Syphilis • HIV • Chlamydia • Herpes
HPV • Gonorrhea • LGV • Syphilis •
HIV • Chlamydia • Herpes • HPV •
Gonorrhea • LGV • Syphilis • HIV •
Chlamydia • Herpes • HPV • Gonorrhea
LGV • Syphilis • HIV • Chlamydia • Herpes
Syphilis • HIV • Chlamydia • Herpes

Canadian Guidelines on Sexually Transmitted Infections

2006

Canadian STI Guidelines · Canadian STI Guidelines · Canadian STI STI Guidelines · Canadian STI Guidelines ·





Our mission is to promote and protect the health of Canadians through leadership, partnership, innovation and action in public health.

Public Health Agency of Canada

Revised edition of the 1998 Canadian STD Guidelines.

This publication can be made available in alternative formats upon request, and can also be found on the Internet at the following address: www.publichealth.gc.ca/sti

Disponible en français sous le titre : Lignes directrices canadiennes sur les infections transmissibles sexuellement édition 2006

Correspondence:

Sexual Health and Sexually Transmitted Infections Section Community Acquired Infections Division Infectious Disease and Emergency Preparedness Branch Public Health Agency of Canada Ottawa, Ontario K1A 0K9

Fax: (613) 957-0381

Email: PHAC_Web_Mail@phac-aspc.gc.ca

© HER MAJESTY THE QUEEN IN RIGHT OF CANADA (2006)

Catalogue number: HP40-1/2006E

ISBN 0-662-42798-X

Canadian Guidelines on Sexually Transmitted Infections 2006 Edition was coordinated by the Expert Working Group on Canadian Guidelines for Sexually Transmitted Infections.

Chair

Tom Wong, MD, MPH, FRCPC, Director, Community Acquired Infections Division, Infectious Disease and Emergency Preparedness Branch, Public Health Agency of Canada

Section Chairs

Primary Care and Sexually Transmitted Infections Marc Steben, MD, FCFP, médecin-conseil, Direction risques biologiques, environnementaux et occupationnels, Institut national de santé publique du Québec et clinique des maladies de la vulve, Hôpital Notre-Dame, Centre hospitalier de l'Université de Montréal

Laboratory Diagnosis of Sexually Transmitted Infections

Max Chernesky, PhD, Professor Emeritus, McMaster University

Management and Treatment of Specific Syndromes Mark Yudin, MD, MSc, FRCSC, Assistant Professor, University of Toronto; Deputy Head, Department of Obstetrics and Gynecology, St. Michael's Hospital

Management and Treatment of Specific Infections Barbara Romanowski, MD, FRCPC, Clinical Professor of Medicine, Division of Infectious Diseases, Faculty of Medicine and Dentistry, University of Alberta

Specific Populations

Rhonda Kropp, BScN, MPH, Senior Public Health Analyst, Sexual Health and STI Section, Community Acquired Infections Division, Public Health Agency of Canada

Members

Joanne Embree, MD, FRCPC, Departments of Medical Microbiology and Pediatrics and Child Health, University of Manitoba

William Fisher, PhD, Professor, Departments of Psychology and Obstetrics and Gynaecology, University of Western Ontario

Janice Mann, MD, Head, Knowledge Development and Research, Sexual Health and STI Section, Community Acquired Infections Division, Public Health Agency of Canada

Lai-King Ng, PhD, Director of Bacteriology and Enteric Diseases Program, National Microbiology Laboratory, Public Health Agency of Canada

David Patrick, MD, MHSc, FRCPC, Associate Professor, UBC Healthcare & Epidemiology; Director, Epidemiology Services, British Columbia Centre for Disease Control

Michael Rekart, MD, DTM&H, MHSc, Director, HIV/ AIDS Control, British Columbia Centre for Disease Control

Cathy Sevigny, RN, BScN, Program Consultant, Sexual Health and STI Section, Community Acquired Infections Division, Public Health Agency of Canada Ameeta Singh, MD, BMBS, MSc, FRCPC, Infectious Diseases Medical Consultant, Alberta Health and Wellness; Clinical Associate Professor, Department

of Medicine, University of Alberta; Medical Director, Capital Health STD Centre

Secretariat

Allison Ringrose, BHSc, Program Officer, Sexual Health and STI Section, Community Acquired Infections Division, Public Health Agency of Canada

Guidelines presented in this document reflect the views of the Expert Working Group on Canadian Guidelines for Sexually Transmitted Infections. They should be construed not as rules but rather as recommendations.

PREFACE

In March 2003, the Community Acquired Infections Division, Public Health Agency of Canada (PHAC) (then part of Health Canada), brought together an Expert Working Group (EWG) on sexually transmitted infections (STIs) from across Canada to begin planning the revision of the 1998 Canadian STD Guidelines. STI experts from the fields of medicine, nursing, laboratory, public health and research voluntarily participated as authors, reviewers and EWG members in an effort to develop updated, evidence-based recommendations for the prevention, diagnosis, treatment and management of STIs in Canada. The content of the Canadian Guidelines on Sexually Transmitted Infections 2006 Edition reflects emerging issues and highlights changes in the STI literature since the release of the 1998 guidelines.

These guidelines were created as a resource for clinical and public health professionals — especially nurses and physicians — for the prevention and management of STIs across a diverse patient population, including neonates, children, adolescents and adults.

While this document addresses key issues related to the prevention, diagnosis, treatment and management of the most common STIs, it is beyond the scope of these guidelines to provide comprehensive recommendations for the treatment and management of HIV and viral hepatitis C. When confronted with these infections, either as a primary infection or a co-infection, we suggest that you refer to alternate resources (see below for suggestions), including colleagues experienced in the area.

- Strader DB, Wright T, Thomas DL, Seeff LB. AASLD practice guideline: diagnosis, management, and treatment of hepatitis C. *Hepatology* 2004;39:1147–1171.
- U.S. Department of Health and Human Services, Panel on Clinical Practices for Treatment of HIV Infection. Guidelines for the Use of Antiretroviral Agents in HIV-1-infected Adults and Adolescents. Available at: AIDSinfo.nih.gov/ ContentFiles/AdultandAdolescentGL.pdf. Accessed February 6, 2006.

The EWG and PHAC acknowledge that the advice and recommendations set out in this statement are based upon the best current available scientific knowledge and medical practices, and they are disseminating this document to clinical and public health professionals for information purposes. Persons administering or using drugs, vaccines or other products should also be aware of the contents of the individual product monograph(s) for those products, or other similarly approved standards or instructions for use provided by the licensed manufacturer(s). Recommendations for use and other information set out in these guidelines may differ from that set out in product monograph(s) or other similarly approved standards or instructions for use. Manufacturers have sought approval and

Preface

provided evidence as to the safety and efficacy of their products only when used in accordance with the product monograph(s) or other similarly approved standards or instructions for use.

Practitioners should report adverse drug reactions to the Canadian Adverse Drug Reaction Monitoring Program (CADRMP). For specifications and standards of reporting, consult Health Canada's CADRMP guidelines.

While these guidelines have been based on current evidence and clinical practice, the prevention, diagnosis, treatment and management of STIs is an evolving field. The EWG and PHAC, in producing these recommendations, will regularly update this information. Readers are encouraged to consult the STIs page of the PHAC website for the latest chapter update(s).

iv Preface

ACKNOWLEDGMENTS

With the assistance of the Sexual Health and Sexually Transmitted Infections Section, Community Acquired Infections Division, Public Health Agency of Canada:

Editor-in-Chief: Dr. Tom Wong

Associate Editors: Rhonda Kropp, Dr. Janice Mann

Production Manager: Barbara Jones

Production Coordinator: Robert Lerch

Production Assistant: Linda Gardiner

Many health professionals from across Canada volunteered their time to author chapters for these guidelines, and their participation is acknowledged by chapter:

Fred Y. Aoki, MD, Professor of Medicine, Medical Microbiology and Pharmacology & Therapeutics Member, Section of Adult Infectious Diseases, Faculty of Medicine, University of Manitoba, Author: Genital Herpes Simplex Virus (HSV) Infections; Dr. Max Chernesky, PhD, Professor Emeritus, McMaster University, Author: Laboratory Diagnosis of Sexually Transmitted Infections; François Coultée, Clincial Researcher, Laboratoire de Virologie Moléculaire, Centre de Recherche Centre Hospitalier de l'université de Montréal Hôpital Notre-Dame, Co-Author: Genital Human Papillomavirus (HPV) Infections; Laurent Delorme, MD CSPQ FRCP(C), médecin microbiologiste infectiologue, Hôpital Charles-LeMoyne, Co-Author: Genital Ulcer Disease (GUD); Francisco Diaz-Mitoma, MD, PhD, FRCPC, Professor and Chief, Division of Virology, Children's Hospital of Eastern Ontario, University of Ottawa, Co-Author: Genital Ulcer Disease (GUD); Alex Ferenczy, MD, Professor of Pathology and Obstetrics & Gynecology, McGill University and the Sir Mortimer B. Davis-Jewish General Hospital Montreal, Co-Author: Genital Human Papillomavirus (HPV) Infections; William A. Fisher, PhD, Professor, Departments of Psychology and Obstetrics and Gynaecology, University of Western Ontario, Author: Primary Care and Sexually Transmitted Infections; Sarah Forgie, MD FRCPC, Assistant Professor, Pediatrics, Division of Infectious Diseases, University of Alberta, Associate Director, Infection Control, Stollery Children's Hospital and University of Alberta Hospital, Co-Author: Sexual Abuse in Peripubertal and Prepubertal Children, Sexual Assault in Postpubertal Adolescents and Adults; Eduardo L. Franco, MPH, DrPH, James McGill Professor of Epidemiology and Oncology Director, Division of Cancer Epidemiology, McGill University, Co-Author: Genital Human Papillomavirus (HPV) Infections; Deana Funaro, MD, FRCPC dermatology, Clinical professor at Notre-Dame Hospital and Ste-Justine Hospital, University of Montreal, Co-Author: Genital Ulcer Disease (GUD); David Hasse,

MD, Professor, Department of Medicine, Division of Infectious Diseases, Dalhousie University, Author: Human Immunodeficiency Virus (HIV) Infections; Rhonda Kropp, BScN, MPH, Senior Public Health Analyst, Sexual Health and STI Section, Community Acquired Infections Division, Public Health Agency of Canada, Author: Men Who Have Sex With Men (MSM)/Women Who Have Sex With Women (WSW), Substance Use, Co-Author: Lymphogranuloma Venereum (LGV); Annie-Claude Labbé, MD, FRCPC, Department of Microbiology, Hôpital Maisonneuve-Rosemont Montreal, Co-Author: Genital Ulcer Disease (GUD); Janice Mann, MD, Head, Knowledge Development and Research, Sexual Health and STI Section, Community Acquired Infections Division, Public Health Agency of Canada, Author: Inmates and Offenders, Lymphogranuloma Venereum (LGV); Lynette J. Margesson, MD, FRCPC, Assistant Professor of Obstetrics and Gynecology and of Medicine (Dermatology), Dartmouth Medical School, Co-Author: Genital Ulcer Disease (GUD); Deborah M. Money, MD, FRCSC, Associate Professor, University of British Columbia, B.C. Women's Hospital, Co-Author: Pregnancy; Gina Ogilvie, MD, MSc, University of British Columbia, Vancouver, Author: Urethritis; Ron Read, MD, Head, Infectious Diseases, Department of Medicine, Microbiology and Infectious Diseases, University of Calgary/Calgary Health Region, Author: Hepatitis B Virus Infections; Michael L. Rekart, MD, DTM&H, MHSc, Director, HIV/AIDS Control, British Columbia Centre for Disease Control, Author: Sex Workers; Barbara Romanowski, MD, FRCPC, Clinical Professor of Medicine, Division of Infectious Diseases, Faculty of Medicine and Dentistry, University of Alberta, Author: Ectoparasitic Infestations, Gonococcal Infections; Allan Ronald, MD, Distinguished Professor Emeritus, University of Manitoba, Author: Chancroid; Shelly Sarwal, MD, MSc, FRCP(C), Medical Officer of Health, Nova Scotia Department of Health, Author: Vaginal Discharge; Cathy Sevigny, RN, BScN, Program Consultant, Sexual Health and STI Section, Community Acquired Infections Division, Public Health Agency of Canada, Author: Primary Care and Sexually Transmitted Infections; Rita Shahin, MD, Toronto Public Health, Author: Travellers; Ameeta Singh, BMBS, MSc, FRCPC, Infectious Diseases Medical Consultant, Alberta Health and Wellness Clinical Associate Professor, Department of Medicine, University of Alberta, Medical Director, Capital Health STD Centre, Author: Sexual Abuse in Peripubertal and Prepubertal Children, Sexual Assault in Postpubertal Adolescents and Adults, Syphilis; Marc Steben, MD, médecinconseil, Direction risques biologiques, environnementaux et occupationnels, Institut national de santé publique du Québec et clinique des maladies de la vulve, Hôpital Notre-Dame, Centre hospitalier de l'Université de Montréal, Author: Genital Human Papillomavirus (HPV) Infections, Genital Ulcer Disease (GUD), Primary Care and Sexually Transmitted Infections; Bruno Turmel, MD, médecin-conseil, Direction générale de la santé publique, Ministère de la Santé et des Services sociaux du Québec, Author: Epididymitis, Prostatitis, Sexually Transmitted Intestinal and Enteric Infections; Julie van Schalkwyk, MD, MSc, FRCSC, Assistant Clinical Professor, Department of Obstetrics and Gynecology, University of British Columbia, Author: Pregnancy; Tom Wong, MD, MPH, FRCPC, Director Community Acquired Infections Division Infectious Disease and Emergency Preparedness

Branch, Author: Chlamydial Infections, Immigrants and Refugees; Mark H. Yudin, MD, MSc, FRCSC, Assistant Professor, University of Toronto, Deputy Head, Department of Obstetrics and Gynecology, St. Michael's Hospital, Author: Pelvic Inflammatory Disease (PID), Co-Author: Urethritis.

The following people are thanked for volunteering their time as external reviewers for the guideline chapters:

Robert Brunham, MD, Director of Medical and Academic Affairs, BC Centre for Disease Control, Director, UBC Centre for Disease Control, Professor of Medicine, Division of Infectious Diseases, University of British Columbia; Susan Comay, MD, Medical Director, Sexual Assault Service, BC Women's Hospital; Curtis Cooper, MD, FRCPC, University of Ottawa; Francisco Diaz-Mitoma, MD, PhD, FRCPC, Professor and Chief, Division of Virology, Children's Hospital of Eastern Ontario, University of Ottawa; Harold Dion, MD, CCFP, FCFP, Clinique médicale l'Actuel, Chair of the Board, Quebec College of Family Physicians; Shelia Dunn, MD, CCFP(EM), University of Toronto, Sunnybrook and Women's College Health Sciences Centre; Alex Ferenczy, MD, Professor of Pathology and Obstetrics & Gynecology, McGill University, Sir Mortimer B. Davis-Jewish General Hospital; David Fisman, MD, MPH, Visiting Scholar Center for Health and Wellbeing, Woodrow Wilson School, Princeton University; Jennifer Geduld, Manager, HIV/ AIDS Surveillance Section, CIDPC, Surveillance and Risk Assessment Division, HIV/AIDS Epidemiology and Surveillance, Public Health Agency of Canada; Mary Gordon, MD, City of Ottawa, Public Health, Sexual Health Centre; Kevin Gough, MD, FRCPC, MEd, St. Michael's Hospital, University of Toronto; Andree Gruslin, MD, FRCS, Post Graduation Program Director, Assistant Professor, MFM Department of Obstetrics and Gynecology, University of Ottawa; Hunter Handsfield, MD, Professor of Medicine, University of Washington, Center for AIDS and Sexually Transmitted Diseases; Sandra Hooper, RN(EC), MScN, Nurse Practitioner Sexual Health Centre, Ottawa Public Health; Robbi Howlett, MASc, PhD (candidate), Manager, Ontario Cervical Screening Program, Division of Preventive Oncology, Cancer Care Ontario; Gaya Jayaraman, PhD, MPH, Manager, HIV Drug Resistance and Field Surveillance Section, Surveillance and Risk Assessment Division, Public Health Agency of Canada; Hugh D. Jones, MD, Dip Ven, STD/AIDS Control, Clinic Physician; Fadel Kane, MD, MSc, HIV/AIDS Programs and Policy Division, Public Health Agency of Canada; Sari Kives, MD, University of Toronto, St. Michael's Hospital; Claude Laberge, MD, Service de lutte contre les ITSS, Direction générale de la santé publique, Ministère de la santé et des services sociaux du Québec; Gilles Lambert, MD, Institut national de santé publique du Québec; Debbie Lindsay, MD, University of Manitoba, Child Protection Centre, Health Sciences Centre; Noni MacDonald, MD, MSc, FRCP, Dalhousie University, IWK Health Centre; Louisa MacKenzie, MD, FRCPC, DTMH, Calgary Refugee Health Program, Margaret Chisholm Resettlement Centre; Lorette Madore, RN, DPHN, BN, Supervisor, Clinical Services, Healthy Sexuality and Risk Reduction Program, Ottawa Public Health/Santé publique

d'Ottawa; Nathalie Mondain, PhD, Groupe de Recherche Interdisciplinaire en Santé (GRIS), University of Montreal; Deborah Money, MD, FRCSC, Associate Professor, University of British Columbia, BC Women's Hospital; Carolyn A. Montgomery, MB, STD/AIDS Control, Clinic Physician; Curtis Nickel, MD, Professor of Urology, Queen's University; Gina Ogilvie, MD, MSc, University of British Columbia; Caroline Paquet, SF, MSc, Professeure, Baccalauréat en pratique sage-femme, Département de Chimie-biologie, Université du Québec à Trois-Rivières; Gordon Phaneuf, Director of Strategic Initiatives, Child Welfare League of Canada; Raphael Saginur, MD, FRCPC, Ottawa Hospital and University of Ottawa; John Sellors, MD, Senior Medical Advisor, Reproductive Health, PATH; Alberto Severini, MD, National Microbiology Laboratory, Public Health Agency of Canada; Stephen Shafran, MD, FRCPC, Professor and Director, Division of Infectious Diseases, Department of Medicine, University of Alberta; Rita Shahin, MD, Toronto Public Health; Brenna Shearer-Hood, MSA(HSA), BMR (OT) Cancer Care Manitoba; Jonathan M Smith, CSC Epidemiologist, Tuberculosis Prevention and Control, Community Acquired Infections Division, Public Health Agency of Canada; Gwen Stephens, MD, FRCPC, University of British Columbia, Department of Pathology & Laboratory Medicine; Jill Tinmouth, MD, PhD, Sunnybrook and Women's Health Sciences Centre, University of Toronto; Baldwin Toye, MD, FRCPC, Head, Division of Microbiology, Ottawa Hospital, University of Ottawa; Sharonie Valin, MD, CCFP, MHSc, North York General Hospital, Women's College Hospital, Bay Centre for Birth Control; Heidi Wood, PhD, Head, Diagnostics, Zoonotic Diseases and Chlamydia Section, National Microbiology Laboratory, Public Health Agency of Canada.

TABLE OF CONTENTS

Pr	reface	iii
Acknowledgments Introduction Primary Care and Sexually Transmitted Infections 1. Assessing the Reason for a Consultation 2. Knowing about STI Risk Factors and Epidemiology 3. Performing a Brief Patient History and STI Risk Assessment 4. Providing Patient-Centred Education and Counselling 5. Performing a Physical Examination 6. Selecting Appropriate Screening/Testing 7. Diagnosing by Syndrome or by Organism and Post-test Counselling 8. Treating 9. Reporting to Public Health and Partner Notification 10. Managing Co-morbidity and Associated Risks 11. Following up Laboratory Diagnosis of Sexually Transmitted Infections Collection and Transportation of Specimens Laboratory Testing Methods 3	v	
ln	troduction	1
Pr	imary Care and Sexually Transmitted Infections	7
	Assessing the Reason for a Consultation	8
	2. Knowing about STI Risk Factors and Epidemiology	9
	3. Performing a Brief Patient History and STI Risk Assessment	12
	4. Providing Patient-Centred Education and Counselling	15
	5. Performing a Physical Examination	19
	6. Selecting Appropriate Screening/Testing	20
	7. Diagnosing by Syndrome or by Organism and Post-test Counselling	21
	8. Treating	22
	9. Reporting to Public Health and Partner Notification	22
	10. Managing Co-morbidity and Associated Risks	28
	11. Following up	28
La	aboratory Diagnosis of Sexually Transmitted Infections	30
	Collection and Transportation of Specimens	30
	Laboratory Testing Methods	34
	Laboratory Diagnosis of Specific Infections	35
M	anagement and Treatment of Specific Syndromes	42
	Syndromic Management of Sexually Transmitted Infections	42
	Epididymitis	53
	Genital Ulcer Disease (GUD)	59
	Pelvic Inflammatory Disease (PID)	71
	Prostatitis	80
	Sexually Transmitted Intestinal and Enteric Infections	92
	Urethritis	98
	Vaginal Discharge (Bacterial vaginosis, Vulvovaginal Candidiasis, Trichomoniasis)	106

Table of Contents ix

Management and Treatment of Specific Infections	122
Chancroid	122
Chlamydial Infections	126
Ectoparasitic Infestations (Pubic Lice, Scabies)	140
Genital Herpes Simplex Virus (HSV) Infections	145
Genital Human Papillomavirus (HPV) Infections	160
Gonococcal Infections	174
Hepatitis B Virus Infections	189
Human Immunodeficiency Virus (HIV) Infections	198
Lymphogranuloma Venereum (LGV)	223
Syphilis	232
Specific Populations	248
Immigrants and Refugees	248
Inmates and Offenders	255
Men Who Have Sex with Men (MSM)/ Women Who Have Sex With Women (WSW)	262
Pregnancy	273
Sexual Abuse in Peripubertal and Prepubertal Children	292
Sexual Assault in Postpubertal Adolescents and Adults	305
Sex Workers	315
Substance Use	319
Travellers	330
Appendices	
A: Patient Counselling Guide on Condom Use	334
B: How to Use a Male Condom/How to Use a Female Condom	337
C: Resources and Reference Tools for Health Professionals	339
D: Provincial and Territorial Directors of STI Control	341
E: Provincial Laboratories	343
F: Forensic Evidence, Services and Laboratories	345
G: Referral Centres for STIs in Peripubertal and Prepubertal Children	349
H: Tanner Scale of Sexual Maturity	352
Index	354

x Table of Contents

INTRODUCTION

The Process Underlying the Creation of the Canadian Guidelines on Sexually Transmitted Infections, 2006 Edition

The process used to create the *Canadian Guidelines on Sexually Transmitted Infections 2006 Edition* was developed by the 14-member expert working group (EWG) (chaired by Dr. Tom Wong from the Public Health Agency of Canada [PHAC]) and by the Sexual Health and Sexually Transmitted Infections (STI) Section, PHAC. Chapters were written by STI experts from across Canada on a voluntary basis. To facilitate the evidence-based revision, PHAC conducted literature reviews on all chapters and provided additional literature assistance as requested by the authors during chapter writing. Each of the 27 chapters underwent a minimum of four rounds of blinded expert review, three within the EWG and one with at least two external reviewers. Final approval of each chapter by the EWG was required before the chapter was considered complete. In order to ensure the integrity and impartiality of the process and the recommendations in the final document, all EWG members and chapter authors have signed a conflict of interest and disclosure form.

This edition has been enhanced to include references throughout each chapter, as well as level of recommendation and quality of evidence indicators for the treatment recommendations. The indicators used reflect a combination of the methodologies from the U.S. Preventive Services Task Force and the Canadian Task Force on Preventive Health Care and have been modified and simplified for use in these guidelines as outlined in Tables 1 and 2.

Table 1. Levels of recommendation (Modified from Harris RP, et al.¹)

Recommendation: A	Strongly recommends that clinicians routinely provide the treatment to eligible patients. Good evidence that the treatment improves important health outcomes and concludes that benefits substantially outweigh harms.
Recommendation: B	Recommends that clinicians routinely provide the treatment to eligible patients. At least fair evidence that the treatment improves important health outcomes and concludes that benefits outweigh harms.
Recommendation: C	No recommendation for or against routine provision of the treatment. At least fair evidence that the treatment can improve health outcomes but concludes that the balance of the benefits and harms is too close to justify a general recommendation .
Recommendation: D	Recommends against routinely providing the treatment to asymptomatic patients. At least fair evidence that the treatment is ineffective or that harms outweigh benefits.
Recommendation: I	Evidence is insufficient to recommend for or against routinely providing the treatment. Evidence that the treatment is effective is lacking , of poor quality or conflicting , and the balance of benefits and harms cannot be determined.

Table 2. Quality of evidence (Modified from Harris RP, et al¹ and Gross PA, et al²)

1	Evidence from at least one properly randomized, controlled trial.
II	Evidence from at least one well-designed clinical trial without randomization, from cohort or case-control analytic studies (preferably from more than one centre), from multiple time-series studies or from dramatic results in uncontrolled experiments.
Ш	Evidence from opinions of respected authorities based on clinical experience, descriptive studies or reports of expert committees.

NEW TERMINOLOGY AND CHAPTERS

The Canadian Guidelines on Sexually Transmitted Infections 2006 Edition reflects the change in terminology from sexually transmitted disease (STD) to STI, which has been adopted to encompass both symptomatic and asymptomatic patient presentation. This shift helps legitimize the need for thorough patient assessment and screening of those with identified risk, regardless of symptomatology.

Each chapter belongs to one of five sections: Primary Care and Sexually Transmitted Infections, Laboratory Diagnosis of Sexually Transmitted Infections, Management and Treatment of Specific Syndromes, Management and Treatment of Specific Infections and Specific Populations.

The Primary Prevention of STD and Clinical Approach to the Diagnosis and Management of STD chapters from the 1998 guidelines have been combined into one chapter for the current revision, titled Primary Care and Sexually Transmitted Infections. Chapters from the 1998 guidelines that have been incorporated into other sections of the current revision include Cervicitis, Persons with Repeated STD and Youth and Street Youth

New chapters have been added to the Management and Treatment of Specific Infections section (Chancroid, Lymphogranuloma Venereum) and to the Specific Populations section (Immigrants and Refugees, Inmates and Offenders, Sex Workers, Men Who Have Sex with Men/Women Who Have Sex with Women and Substance Use) of this edition.

NEED TO STRENGTHEN PREVENTION

In Canada, there are three nationally reportable STIs: chlamydia, gonorrhea and infectious syphilis. Since 1997, there has been a steady increase in the rates of all three infections. This phenomenon is not unique to Canada; other countries, including the U.S. and the U.K., have reported similar trends.^{3,4} Targeted enhanced surveillance and research are required to determine the factors that may be playing a role in these trends. Some of the possible factors may include the following:

- Nucleic acid amplification tests (NAATs) have been introduced.
- Some people may have developed safer-sex burnout.
- There have been innovations in HIV therapy (e.g., highly active antiretroviral therapy), leading to related treatment optimism.
- Youth awareness of risks and knowledge of risk reduction behaviours remain less than optimal.⁵
- Sex is occurring at an early age, with a high rate of serially monogamous relationships.
- Sex is continuing later in life.

- The transmission risks of STIs associated with sexual activity (vaginal, anal and oral) are not well understood by the public.
- "Party drugs," such as ecstasy and crystal meth, are being increasingly linked to unsafe sexual behaviours.⁶
- Anonymous partnering venues, such as the Internet, are expanding.

By being aware of trends in STIs, risk factors and affected populations, primary care providers and public health practitioners can be strategically placed to apply relevant and complementary individual and community-based education and patient services.

The prevention and control of STIs cannot be approached with a narrow focus. The appropriate medical management of identified cases of STIs is but one piece of the puzzle. Both primary and secondary prevention activities are paramount to reducing the incidence (newly acquired infections) and prevalence (number of cases) of STIs. Primary prevention aims to prevent exposure by identifying at-risk individuals and performing thorough assessments, patient-centred counselling and education. Secondary prevention involves reducing the prevalence of STIs through the detection of infections in at-risk populations, counselling, conducting partner notification and treating infected individuals and contacts in a timely manner, thus preventing and/or limiting further spread.

Both the burden of disease and potential complications associated with STIs are relevant and significant considerations for health professionals and decision makers. The presence of an acute infection can increase the risk of co-infection: for example, an ulcer from an infection such as syphilis can significantly increase the risk of acquiring and transmitting an HIV infection. The sequelae for women from untreated infections such as chlamydia and gonorrhea can include pelvic inflammatory disease, chronic pelvic pain, ectopic pregnancy and infertility. In recent years, there has also been increasing evidence to support the role of persistent human papillomavirus (HPV) infections in cervical dysplasia and carcinoma.

As we strive to attend to the physiological needs of patients, we must also be prepared to attend to their psychological needs as well. Chronic viral STIs can have long-standing negative impacts on a patient's psychosocial well-being. The many potential impacts and sequelae of STIs highlight the need for strengthened prevention efforts.

FUTURE DEVELOPMENTS

As within many areas in the health sector, innovation and development are part of the growing body of knowledge and tools used in the prevention, treatment and management of disease and infection. We recommend consulting a variety of mechanisms/sources to maintain and enhance your clinical practice.

Two future developments with significant potential for impact on the field of STIs are the upcoming HPV and HSV vaccines. The latest data on these two developments are outlined below. As these are evolving areas of inquiry, please consult the STI section of the PHAC website for the latest available information.

HPV vaccine

Preliminary data on virus-like particle vaccines for HPV prevention demonstrate positive results in terms of both safety and short-term efficacy. As of 2005, two candidate vaccines are well into phase 3 trials. Both candidate vaccines include protection against HPV-16 and HPV-18, which cause 70% of cervical cancers.⁸ One of the candidate products also includes protection against HPV-6 and HPV-11 antigens, which cause 90% of external genital warts.⁹ Therapeutic vaccines have also been studied, but the initial results have not been favourable.

HSV vaccine

Preliminary data about a viral glycoprotein-based vaccine against HSV type 2 has shown good results in terms of safety. It provides short-term protection for HSV type 1-negative women but no protection has been found in men.¹⁰ Therapeutic vaccines have also been studied, but results to date have demonstrated a lack of effect compared to placebo.

REFERENCES

- 1. Harris RP, Hefland M, Woolf SH, et al. Current methods of the US Preventive Services Task Force: a review of the process. *Am J Prev Med* 2001;20(3 suppl):21–35.
- 2. Gross PA, Barrett TL, Dellinger EP, et al. Purpose of quality standards for infectious diseases. Infectious Diseases Society of America. *Clin Infect Dis* 1994:18:421.
- Centers for Disease Control and Prevention. Trends in reportable sexually transmitted diseases in the United States, 2004. In: 2004 STD Surveillance Report. Atlanta, GA: Centers for Disease Control and Prevention; 2005.
 Available at: www.cdc.gov/std/stats/default.htm. Accessed January 17, 2006.
- 4. Health Protection Agency Centre for Infections. *Mapping the Issues HIV and other Sexually Transmitted Infections in the United Kingdom: 2005.*London: Health Protection Agency Centre for Infections; 2005. Available

- at: www.hpa.org.uk/hpa/publications/hiv_sti_2005/default.htm. Accessed January 17, 2006.
- Council of Ministers of Education Canada. Canadian Youth, Sexual Health and HIV/AIDS Study 2002: Factors Influencing Knowledge, Attitudes and Behaviours. Toronto, ON: Council of Ministers of Education Canada; 2003. Available at: www.cmec.ca/publications/aids/. Accessed January 17, 2006.
- 6. Buchacz K, McFarland W, Kellogg TA, et al. Amphetamine use is associated with increased HIV incidence among men who have sex with men in San Francisco. *AIDS* 2005;19:1423–1424.
- 7. World Health Organization. *Preventing and Treating Sexually Transmitted and Reproductive Tract Infections*. Geneva, Switzerland: World Health Organization; 2006. Available at: www.who.int/hiv/topics/sti/prev/en/print.html. Accessed January 17, 2006.
- 8. Bosch FX, de Sanjose S. Chapter 1: Human papillomavirus and cervical cancer—burden and assessment of causality. *J Natl Cancer Inst Monogr* 2003;31:3–13.
- 9. von Krogh G. Management of anogenital warts (condylomata acuminata). *Eur J Dermatol* 2001;11:598–603.
- 10. Stanberry LR, Spruance SL, Cunningham AL, et al. Glycoprotein-D-adjuvant vaccine to prevent genital herpes. *N Engl J Med* 2002;347:1652–1661.

PRIMARY CARE AND SEXUALLY TRANSMITTED INFECTIONS

PREVENTION, DIAGNOSIS AND CLINICAL MANAGEMENT OF SEXUALLY TRANSMITTED INFECTIONS IN THE PRIMARY CARE SETTING

It is important for practitioners to recognize that sexually transmitted infection (STI) risks will vary from person to person and should be viewed as dynamic across the lifespan.

- Only through proper assessment can a patient's risk for STIs be identified.
- Assumptions and inferences about patient STI risk may prove inaccurate.
- Sexually inactive individuals can be made aware of STI risks in the course of routine care.

Primary care providers can incorporate STI primary and secondary prevention in the course of routine patient care by doing the following:

- · Assessing and discussing STI risk.
- Informing patients about signs and symptoms of STIs (and lack thereof).
- · Helping patients recognize and minimize STI risk.
- · Offering patient-centred counselling.
- Offering hepatitis A virus (HAV) and hepatitis B virus (HBV) immunization when indicated.
- Offering STI screening and testing.
- Appropriately treating, following up and counselling infected patients and their partners.

This chapter provides an overview of best practices for the prevention and clinical management of STIs in primary care settings. It includes recommendations for the assessment, counselling, screening, diagnosis and management of STIs, including partner notification and public health reporting.

Effective prevention and management of STIs requires the following elements on the part of the health care practitioner:

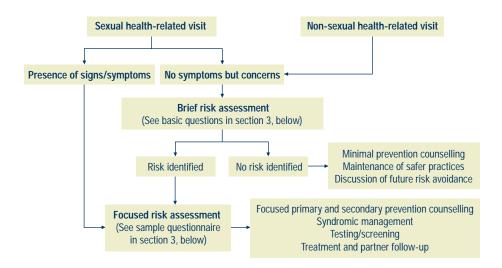
- 1. Assessing the reason for a consultation.
- 2. Knowing about STI risk factors and epidemiology.
- 3. Performing a brief patient history and STI risk assessment.
- 4. Providing patient-centred education and counselling.
- 5. Performing a physical examination.
- 6. Selecting appropriate screening/testing.
- 7. Diagnosing by syndrome or by organism and post-test counselling.
- 8. Treating.
- 9. Reporting to public health and partner notification.
- 10. Managing co-morbidity and associated risks.
- 11. Following up.

Each of these elements is outlined in more detail below.

1. Assessing the Reason for a Consultation

Patients may seek medical attention for issues unrelated to sexual health, but they may be at risk for STIs and benefit from interventions to address identified risk factors. For example, consultation for contraception often has implications for STI prevention counselling and STI screening; management of contraception and management of STI risk are closely related. When patients present for contraceptive advice, it can be an ideal time to assess and discuss STI risk. The type of STI risk a patient may encounter also has implications for appropriate contraceptive choice.

Figure 1. STI risk assessment in primary care settings



In some cases, patients may consult to inquire about signs or symptoms related to a possible STI, to request STI testing or to discuss prevention issues. Identifying a person who has STI concerns, who is at risk for an STI or who has an STI provides an opportunity for discussing barriers to risk reduction and means to overcome them.

2. Knowing about STI Risk Factors and Epidemiology

Identifying the index of suspicion of STI infection in a patient requires the health care practitioner to understand the epidemiologic trends of STIs, as well as the risk factors associated with STI transmission and infection.

Summarized in Table 1 are the key epidemiologic trends for bacterial and viral STIs in Canada, as well as patient risk factors for STIs.

Table 1. Epidemiology of STIs in Canada

Table 1. Epidemiology of 5113 in Gandad			
Infection	How common in clinical practice?	Trends in incidence	Most affected
Chlamydia	 Most commonly diagnosed and reported bacterial STI Cases reported in Canada in 2002: 56,241 Cases reported in Canada in 2004: 63,000 (preliminary data)* 	Steadily increasing in Canada since 1997	 Young women aged 15–24 Young men aged 20–29
Gonorrhea	 Second most commonly diagnosed and reported bacterial STI Cases reported in Canada in 2002: 7,367 Cases reported in Canada in 2004: 9,200 (preliminary data)* 	 From 1997–2004 (preliminary data),* rate has increased by approximately 94% Quinolone resistance has increased from <1% in the early 1990s to 6.2% in 2004 (national rate)[†] 	 Males account for approximately ²/₃ of reported cases Increase in MSM Young men aged 20–29 Young women aged 15–24
Infectious syphilis	 Previously rare in Canada Cases reported in Canada in 2002: 463 Cases reported in Canada in 2004: 1,112 (preliminary data)* 	Dramatic national increases noted since 1997 related to regional outbreaks across Canada	 MSM (HIVpositive and negative) aged 30–39 Sex workers and their clients Acquisition in endemic regions

Table 1. Epidemiology of STIs in Canada (continued)

Infection	How common in clinical practice?	Trends in incidence	Most affected
Chancroid	Exceedingly rare in Canada	Stable	Acquisition in endemic regions
Granuloma inguinale	Exceedingly rare in Canada	• Stable	Acquisition in endemic regions
Lympho- granuloma venereum	Previously rare in Canada	Unknown Recent outbreaks in Canada have resulted in the development and implementation of an enhanced surveillance system	MSM Acquisition in endemic regions
Human papillomavirus	Very common: 70% of the adult population will have had at least one genital HPV infection over their lifetime	 True incidence not known, as HPV is not a reportable disease 	Adolescents and young adult women and men (but affects women and men of all ages)
Genital herpes (HSV-1 and -2)	• Common	 True incidence not known, as HSV is not a reportable disease Seroprevalence studies indicate rates of at least 20% 	Very common in both adolescents and adult men and women Women are more affected than men
HIV	 Rare in general practice 2,529 cases reported in Canada in 2004 	• 20% rise in number of HIV positive test reports in Canada (2000–2004)	 MSM Acquisition in endemic regions Injection drug users Young women aged 15–19

Table 1. Epidemiology of STIs in Canada (continued)

Infection	How common in clinical practice?	Trends in incidence	Most affected
Hepatitis B	 Low to moderate in general practice and varies in different populations Approximately 700 acute cases per year in Canada 	 Acute hepatitis B is twice as high for men than for women Peak incidence rates are found in the 30–39 age group 	 Infants born to HbsAg positive mothers Injection drug users who share equipment Persons with multiple sex partners Acquisition in endemic regions Sexual and household contacts of an acute or chronic carrier

HbsAg = hepatitis B surface antigen
MSM = men who have sex with men

HPV = human papillomavirus STI = sexually transmitted infection

HSV = herpes simplex virus

Note: For up-to-date epidemiologic information, consult the Public Health Agency of Canada website:

- · www.publichealth.gc.ca/sti
- www.phac-aspc.gc.ca/publicat/aids-sida/haic-vsac1204/index.html
- www.phac-aspc.gc.ca/publicat/ccdr-rmtc/05vol31/31s2/index.html

STI risk factors

The following STI risk factors are associated with increased incidence of STIs:

- Sexual contact with person(s) with a known STI.
- Sexually active youth under 25 years of age.
- A new sexual partner or more than two sexual partners in the past year.
- Serially monogamous individuals who have one partner at present but who have had a series of one-partner relationships over time.
- No contraception or **sole** use of non-barrier methods of contraception (i.e., oral contraceptives, Depo Provera, intrauterine device).
- Injection drug use.
- Other substance use, such as alcohol or chemicals (pot, cocaine, ecstasy, crystal meth), especially if associated with having sex.
- Any individual who is engaging in unsafe sexual practices (i.e., unprotected sex, oral, genital or anal; sex with blood exchange, including sadomasochism; sharing sex toys).
- Sex workers and their clients.
- "Survival sex": exchanging sex for money, drugs, shelter or food.
- · Street involvement, homelessness.
- Anonymous sexual partnering (i.e., Internet, bathhouse, rave party).
- Victims of sexual assault/abuse.
- Previous STL

Preliminary data is subject to change; does not include Nunavut. Surveillance and Epidemiology Section, Community Acquired Infections Division, Public Health Agency of Canada, unpublished data, 2006.
 National Microbiology Laboratory, Public Health Agency of Canada, unpublished data, 2005.

3. Performing a Brief Patient History and STI Risk Assessment General principles

- Information should be requested in a simple, non-judgmental manner, using language understandable to the patient.
- History should enquire about the following:
 - Genital symptoms associated with STIs (discharge, dysuria, abdominal pain, testicular pain, rashes, lesions).
 - Systemic symptoms associated with STIs (fever, weight loss, lymphadenopathy).
 - Personal risk factors and prevention (condom use, vaccination against hepatitis B and, in the case of individuals at risk, hepatitis A).
 - Patient's knowledge of increased risk of STIs.
 - Other pertinent elements of a general history, such as relevant drug treatments, allergies and follow-up of previous problems.
- A brief risk assessment should aim to quickly identify or rule out major risk factors associated with STIs. Use of an STI risk assessment script such as the following may be helpful in rapidly assessing risk:
 - "Part of my job is to assess sexual and reproductive health issues. Of course, everything we talk about is completely confidential. If it is OK with you,
 I would like to ask you a few questions in this area.
 - Are you sexually active now, or have you been sexually active? This
 includes oral sex or anal sex, not just vaginal sex.
 - Do you have any symptoms that might make you think that you have an STI? (Do you have any sores on or around your genitals? Does it hurt or burn when you pee? Have you noticed an unusual discharge from your penis, vagina or anus? Do you have pain during sex?)
 - What are you doing to avoid pregnancy? (Do you or your partner use any type of birth control?)
 - What are you doing to avoid STIs including HIV?
 - Do you have any concerns about sexual or relationship violence or abuse?
 - Have you or your partner(s) used injection or other drugs (e.g., crystal meth)?"
 - For women also ask:
 - "When was your last menstrual period?
 - When was your last Pap test?"

Performing a focused risk assessment

Any patient whose current or past history identifies a potential risk factor for STIs should have a more detailed history completed. The focused STI risk assessment questionnaire (Table 2) is intended to serve as a practical guide to assist clinicians in further evaluating an individual patient's risk factors and behaviours, as well as guiding counselling and testing recommendations.

Table 2. STI risk assessment questionnaire¹

Category and elements	Important questions to guide your assessment	
Relationship Present situation	Do you have a regular sexual partner?If yes, how long have you been with this person?	
Identify concerns	Do you have any concerns about your relationship?If yes, what are they? (e.g., violence, abuse, coercion)	
Sexual risk behaviour Number of partners	 When was your last sexual contact? Was that contact with your regular partner or with a different partner? How many different sexual partners have you had in the past 2 months? In the past year? 	
Sexual preference, orientation	Are your partners, men, women or both?	
Sexual activities	 Do you perform oral sex (i.e., Do you kiss your partner on the genitals or anus)? Do you receive oral sex? Do you have intercourse (i.e., Do you penetrate your partners in the vagina or anus [bum]? Or do your partners penetrate your vagina or anus [bum])? 	
Personal risk evaluation	 Have any of your sexual encounters been with people from a country other than Canada? If yes, where and when? How do you meet your sexual partners (when travelling, bathhouse, Internet)? Do you use condoms, all the time, some of the time, never? What influences your choice to use protection or not? If you had to rate your risk for STI, would you say that you are at no risk, low risk, medium risk or high risk? Why? 	

Table 2. STI risk assessment questionnaire¹ (continued)

Category and elements	Important questions to guide your assessment
STI history Previous STI screening	Have you ever been tested for STI/HIV? If yes, what was your last screening date?
Previous STI	Have you ever had an STI in the past? If yes, what and when?
Current concern	When was your sexual contact of concern?If symptomatic, how long have you had the symptoms that you are experiencing?
Reproductive health history Contraception	• Do you and/or your partner use contraception? If yes, what? Any problems? If no, is there a reason?
Known reproductive problems	Have you had any reproductive health problems? If yes, when? What?
Pap test	Have you ever had an abnormal Pap test? If yes, when? Result if known.
Pregnancy	 Have you ever been pregnant? If yes, how many times? What was/were the outcome(s) (number of live births, abortions, miscarriages)?
Substance use Share equipment for injection	 Do you use alcohol? Drugs? If yes, frequency and type? If injection drug use, have you ever shared equipment? If yes, what was your last sharing date?
Sex under influence	 Have you had sex while intoxicated? If yes, how often? Have you had sex while under the influence of alcohol or other substances? What were the consequences? Do you feel that you need help because of your substance use?
Percutaneous risk other than drug injection	Do you have tattoos or piercings? If yes, were they done using sterile equipment (i.e., professionally)?

Table 2. STI risk assessment questionnaire¹ (continued)

Category and elements	Important questions to guide your assessment
Psychosocial history Sex trade worker or client	 Have you ever traded sex for money, drugs or shelter? Have you ever paid for sex? If yes, frequency, duration and last event.
Abuse	 Have you ever been forced to have sex? If yes, when and by whom? Have you ever been sexually abused? Have you ever been physically or mentally abused? If yes, when and by whom?
Housing	Do you have a home? If no, where do you sleep?Do you live with anyone?

STI=sexually transmitted infection

4. Providing Patient-Centred Education and Counselling

On completing the risk assessment, a number of topics may be identified where sexual health- or STI-related education may be indicated for a given patient. Below are a number of common counselling topics and recommendations for information to share with patients, as well as some tips on how to approach sexual health education/counselling using a patient-centred approach.

Common counselling topics

Serial monogamy

It is important for practitioners to recognize and address the issue of serial monogamy. Serial monogamy consists of a series of faithful, monogamous relationships, one after the other. Although they may "feel safe" and "look safe," serially monogamous relationships, with known and committed partners, do not themselves provide adequate protection from STIs. Consistent condom use and STI testing followed by *mutual* monogamy are far safer strategies than relying on a serially monogamous partners' apparent safety.

For youth contemplating initiation of sexual activity

Many youth will ask for contraceptive information prior to becoming sexually active. Many young women will begin using oral contraception for cycle control as opposed to contraceptive reasons. Both represent excellent opportunities to counsel on safer-sex practices.

- When discussing non-barrier contraceptive options, discussion of safer sex and condom use should occur.
- Promote partner testing prior to becoming sexually active for partners who have already been sexually active.
- · Let patients know the benefits of preventive behaviour.

Contraceptive advice

Oral contraceptive prescription is commonly associated with cessation of condom use. It has been documented that prescription of oral contraception is very often associated with the offset of barrier method use and increased incidence of STIs.² Individuals in relationships very often move on from initial barrier protection to oral contraception without the benefit of STI testing. Clinicians need to counsel about alternatives to this risky pattern (e.g., testing before cessation of condom use), particularly when prescribing oral contraceptives.

Plan and motivate prevention and risk-reduction strategies

Acceptance of sexuality

- Individuals must accept the fact that they are or might be sexually active before
 they can plan for STI prevention. Primary care providers, by their actions, can
 show understanding of patient sexuality by initiating a non-judgmental, twoway dialogue that will help individuals examine the choices they make related
 to their sexuality. Examining these choices can be useful in helping patients
 to proactively plan for risk reduction measures appropriate to their specific
 situation.
- Provide easy-to-apply information:
 - Challenge patients to plan if and how they will discuss STI preventive actions with their partners, or take STI preventive actions unilaterally (e.g., put on a condom), and how they will practice safer sex consistently.
 - Assess whether patients know where they can comfortably obtain condoms in their community, if they know how to use condoms correctly, if they are aware of the signs of STIs and if they know how to seek testing and treatment if needed.

Planning prevention

- Individuals who take STI preventive action need to engage in a number of advance preparations, such as buying condoms, seeking STI/HIV testing and talking about STIs with their health care provider(s). Primary care providers can discuss setting and maintaining personal limits with their patients and identify the most "user-friendly" local STI prevention resources available.
- Health care practitioners can help patients to plan for prevention by openly
 discussing safer sex using a continuum approach (i.e., masturbation/mutual
 masturbation, low risk; oral sex, level of risk varies between HIV and other STIs;
 unprotected vaginal or anal intercourse, high risk for STIs and HIV). This can be
 useful in helping patients understand the risks associated with various activities,

make informed choices about the initiation and maintenance of STI preventive actions and deal with possible partner resistance.

- Provide easy-to-apply information:
 - Discuss limiting alcohol or drug intake prior to sexual activity, as they
 decrease inhibitions and could affect decision-making and negotiation skills.
 - Reinforce that it is *not* possible to assess the chances that a partner has
 an STI on the basis of knowing the partner's sexual history, being in a close
 relationship with a partner or being monogamous with a partner who has a
 sexual history and who has not been tested.
 - It is important to tell patients that we do not and cannot routinely test for all STIs (e.g., human papillomavirus [HPV], herpes simplex virus [HSV]), so even if they or their partner's tests are all negative they may still have an asymptomatic STI.

Safer-sex counselling

Safer-sex counselling as a primary or secondary prevention strategy should include the following at minimum:³

- · STI modes of transmission.
- Risks of various sexual activities (oral, genital, rectal).
- Barrier-method options and availability (male condom, female condom, dental dam).
- Harm-reduction counselling: determining which prevention measures are
 appropriate and realistic to implement, given the patient's personal sexual
 situation(s) (e.g., if practicing receptive anal intercourse, always use a condom
 and extra lubrication, and avoid use of spermicidal condoms).

Statements related to the fact that effective safer-sex practice requires negotiation and is something that should be discussed with partners may be approached by stating: "If you or your partner(s) have ever had another sexual partner, there are a number of options open to you for safer sex. Always using a condom, or getting tested for STI/HIV with your partner followed by mutual monogamy are a few of these options. Do you think any of these might work for you and your partner?"

Proper use of condoms

Reasons for condom failure are most often the result of improper or inconsistent use. For counselling guidelines and instructions on use, see *Appendix A* and *B*.

Efficacy of condoms in STI prevention

- Although latex and polyurethane condoms are effective in preventing the majority of STIs, including HIV, HBV, chlamydia and gonorrhea, they do not provide complete protection against HPV or HSV infection.
- Natural skin condoms may be permeable to HBV and HIV.

Discussing alternatives

- An allergy to latex may be an issue for some patients; male or female polyurethane condoms can offer needed protection in these patients.
- The female condom (a polyurethane vaginal pouch) is commercially available
 and represents an alternative to male condoms or in persons who have a latex
 allergy for both STI and pregnancy prevention. Female condoms are available in
 most drug stores and are more expensive than male condoms, approximately
 \$3.00 each. For instructions on use of a female condom see Appendix B.

Female condom use for anal intercourse

Some individuals are using the female condom for anal intercourse, although the manufacturer does not provide recommendations for use in this way. What limited studies have been done on the use of female condoms for anal intercourse have found that there tends to be a higher incidence of rectal bleeding and condom slippage in comparison to the male condom.⁴

These studies concluded that modifications, training and research on the clinical significance of safety outcomes are needed for the use of female condoms with anal sex, and redesign of the female condom could increase acceptability and use by men who have sex with men (MSM) and address possible safety concerns.^{4,5}

Warning re: nonoxynol-9

Spermicidal lubricated condoms are coated with a lubricant containing nonoxynol-9 (N-9), which may provide added protection against pregnancy. N-9 may increase the risk of infection/transmission of HIV and STIs by causing disruptions and lesions in the genital/anal mucosal lining.⁶ N-9 should not be recommended as an effective means of HIV or STI prevention. The best STI and HIV barrier is a latex or polyurethane condom *without* N-9.

- N-9 should never be used rectally. Even low doses used infrequently can cause
 massive disruption of the rectal mucosal lining, which is likely to increase the risk
 of infection by HIV and other STIs.
- If N-9 is used as an aid to contraception, its benefit should be carefully
 considered in light of the increased risk of genital lesions and the resulting
 potential for an increased risk of HIV transmission.

Motivational interviewing techniques

Motivational interviewing is an intervention strategy that has been used to promote primary and secondary prevention of STIs. Motivational interviewing strategies are well researched clinician-implemented intervention techniques that may be helpful in encouraging patients to practice safer sexual behaviour.⁷⁻⁹ **Motivational interviewing strategies can be used to enhance safer-sex practices and condom use among patients who may require focused counselling.**^{8,9} Table 3 provides an example of a motivational interviewing script.

Table 3. Motivational interviewing script

(Adapted from techniques suggested in Rollnicks, et al.)9

"Let me ask you a couple of questions about condoms..."

Health care provider asks:

Q1. "On a scale of 1 to 10, where 1 is "not at all important" and 10 is "very important," **how important is it** to you to...**always use condoms**?

If patient responds with a score of 8 or more, proceed to Q3.

If patient responds with a score of 7 or less, ask: "Why did you say X and not **lower?**" (This paradoxical question challenges patients to come up with reasons why it **is** important to use condoms.)

- Q2. "What would it take or what would have to happen for it to become more important to you to use condoms?" (Patients are the world's foremost experts in what it would take to change their views, and they will tell the clinician what it would take to make condom use more important to them personally. Health care provider and patient can then discuss these responses.)
- Q3. "On a scale of 1 to 10, how *confident* are you that you (or you and your partner) could always use condoms?"

If patient responds with a score of 8 or more, ask about and explore possible barriers that could occur and how patient might deal with them.

If patient responds with a score of 7 or less, ask: "Why did you say X and not **lower**?" (This paradoxical question prompts patients to think about their strengths in managing condom use.)

Q4. "What would it take or what would have to happen for you to become more confident that you (or you and your partner) could always use condoms?" (Patients again are the world's foremost experts in what it would take to change their behaviour, and they will tell the clinician what it would take to do so. Patient and health care provider can use this as a context for problem solving around condom use.)

5. Performing a Physical Examination

Physical examination may be embarrassing for some patients. Therefore, physicians should develop a trusting environment:

- Some patients may feel more comfortable having an assistant of the same gender present.
- All patients should be assured that confidentiality will be maintained at all times.

Table 4. Components of a physical examination

Components common to both sexes

- General assessment
- Search for systemic signs of STIs, such as weight loss, fever, enlarged lymph nodes (palpate inquinal lymph nodes)
- Inspect mucocutaneous regions, including pharynx
- Inspect external genitalia for cutaneous lesions, inflammation, genital discharge and anatomical irregularities
- · Perform a perianal inspection
- Consider anoscopy (or, if unavailable, digital rectal examination) if patient has practiced receptive anal intercourse and has rectal symptoms
 - For prepubertal females and males, see *Sexual Abuse in Peripubertal and Prepubertal Children* chapter

Components specific to adolescent and adult males

- · Palpate scrotal contents with attention to the epididymis
- When foreskin is present, retract it to inspect the glans
- Have the patient or examiner "milk" the urethra to make any discharge more apparent

Components specific to adolescent and adult females

- · Be sure to separate labia so as to adequately visualize vaginal orifice
- Perform an illuminated speculum examination to visualize the cervix and vaginal walls and to evaluate endocervical and vaginal discharges. Obtain specimens as indicated in the *Laboratory Diagnosis of Sexually Transmitted Infections* chapter.
- Perform a bimanual pelvic examination to detect uterine or adnexal masses or tenderness
- In certain circumstances, such as primary genital herpes or vaginitis, speculum and bimanual examination may be deferred until the acute symptoms have subsided

6. Selecting Appropriate Screening/Testing

- Selecting the appropriate laboratory tests for patients is a crucial step in the diagnosis and management of STIs. The selection of appropriate laboratory tests and biologic samples and specimens should be based on patient history, risk factors and findings on physical examination.
- Be aware of the "I have been tested" syndrome. There are two dimensions to this syndrome:
 - The false sense of security that individuals at risk may develop after multiple STI screenings with repeat negative results. These individuals may develop a sense that "it can never happen to me." This can be a focus for counselling. (See Providing Patient-Centred Education and Counselling, above.)

- The individual who has had some form of medical attention (i.e., a physical, been in a hospital, Pap smear, given blood) and thinks they have been tested for STIs. This is an educational opportunity.
- Simply asking a patient if he or she has been screened for STIs is not enough.
 There is a need to be infection-specific and clarify for the individual that routine blood work at an annual exam does not include syphilis or HIV testing, that a pelvic examination does not include testing for chlamydia and gonorrhea and that a routine urine for culture and sensitivity (C&S) does not screen for chlamydia, etc.

7. Diagnosing by Syndrome or by Organism and Post-test Counselling

- The results of microbiologic testing are not immediately available in most offices.
- When particular symptoms and signs are present, a syndromic diagnosis may be made and treatment and post-test counselling provided. (See Syndromic Management of Sexually Transmitted Infections chapter for a summary table.)
- When microbiologic results are available, treatment and counselling should be directed at specific pathogens; see appropriate chapter(s).

Post-test counselling

Post-test counselling is an integral part of management of the individual with a newly diagnosed STI and should include, at a minimum, the following:³

- · Organism- or syndrome-specific advice.
- Safer-sex practices that can remove or reduce the risk of transmitting the STI to a partner or reduce the risk of re-infection in the patient.
- Treatment information and issues that differ as a function of whether the infection is bacterial (curable) versus viral (manageable).
- Case reporting requirements to local public health unit.
- Partner notification either via the index case, the physician or a public health official, and the implications of partners not being tested or treated.

Post-test prevention counselling can also be a very important educational opportunity for individuals who have presented with STI concerns but tested negative for STIs.

Motivational interviewing strategies, as discussed above, can be effective in promoting risk-reduction behaviour change for patients who have tested positive for an STI.⁷⁻⁹ The difference in motivational interviewing as a primary or secondary prevention strategy is simply in the wording. For example, the health care provider may begin by saying, "I ask all of my patients who are dealing with a sexually transmitted infection a couple of questions. Could you tell me how important it is for *you now* to always use condoms (or always carry out another relevant STI-prevention/harm-reduction strategy)?" (Follow the motivational-interviewing script in Table 3, above.)

8. Treating

Treatment can be curative in the case of bacterial, fungal and parasitic infections or palliative/suppressive in the case of viral STIs. For more specific discussion about particular issues, see *Syndromic Management of Sexually Transmitted Infections* chapter or infection-specific chapters.

Free treatment is available for index cases and their contacts with bacterial STIs in all provinces and territories in Canada.

Patients, whether symptomatic or not, should be told not to share their medications with partners and to complete the full course of their prescribed medication, even if their signs and symptoms resolve before they finish their medication. Patients should also be advised that if vomiting occurs more than 1 hour postadministration, a repeat dose is not required.

Patients diagnosed with a bacterial STI or trichomonal infection should be advised that they and their partners should abstain from unprotected intercourse until 7 days after treatment of both partners is complete (e.g., 7 days after single-dose therapy).

9. Reporting to Public Health and Partner Notification

STI reporting requirements and confidentiality

Patients should be advised of the provincial/territorial public health acts and the *Child Protection Act*, which supersede physician/patient confidentiality and require release of personal information without patient consent for all reportable STIs and in cases where child abuse is suspected.

Those working in agencies receiving personal information are bound by ethical, legal and professional obligations to protect the confidentiality of this information. Patients need to be informed that the information will be reported to authorities only as required by law as noted above but will otherwise remain confidential. This is often a crucial concern for young people who come forward for STI care.

Confidentiality applies to all persons, including infected persons, sexual/needlesharing partners, all youth who are competent to understand their infection and care, and people who may be involved in cases of child sexual abuse.

Partner notification

Rationale

Partner notification is a secondary prevention process through which sexual partners and other contacts exposed to an STI are identified, located, assessed, counselled, screened and treated. Partner notification not only produces a public health benefit (e.g., disease surveillance and control) but dramatically reduces the risk of reinfection for the original patient.

While partner notification is sometimes construed as an exercise in societal vs. individual rights, its aim is clearly to assist people in honouring the individual rights of their partners to know they have been put at risk and to make informed decisions regarding their health and in some instances their life.

A review of the evidence supports several recommendations related to the partnernotification process.¹⁰ There is good evidence to show that partner notification can be an effective means of finding at-risk and infected persons and that health care provider referral generally ensures that more partners are notified and medically evaluated.^{10,11}

Who performs partner notification?

Partner notification may be done by the patient, health care providers or public health authorities. Often, more than one strategy may be used to notify different partners of the same infected person.

- Self- or patient referral: the infected person accepts full responsibility for informing partners of the possibility of exposure to an STI and for referring them to appropriate services.
- Health care provider/public health referral: with the consent of the infected person, the health care provider takes responsibility for confidentially notifying partners of the possibility of their exposure to an STI (without ever naming the index case).
- Contract referral: the health care provider negotiates a time frame with the infected person (usually 24–48 hours) to inform his or her partners of their exposure and to refer them to appropriate services.¹¹

Under certain circumstances (i.e., apparently monogamous relationships) the partner may deduce who the index case is by the process of elimination. The health care provider is still required to maintain confidentiality related to the index case, and no information related to the index case can be released to the partner.

If the index case does not wish to notify partners, or if partners have not come forward:

- Explore impediments/barriers to partner notification (see below).
- · If needed, report to public health authorities.

Barriers to partner notification

- Actual or feared physical or emotional abuse that may result from partner
 notification (e.g., conjugal violence): health care provider/public health referral
 may be the best option in these cases so as to protect the index case. If there
 is a threat to patient safety, public health officials should be notified of this
 so that proper safety precautions are taken to protect the index case. Safety
 always trumps the notification process.
- Fear of losing a partner due to a STI diagnosis (blame/guilt): discuss the
 asymptomatic nature of STIs and the benefits of asymptomatic partner(s)
 knowing that they may be infected.
- Feared legal procedures: cases need to be advised that their identity is protected at all times, and unless their records are subpoenaed, no information can be released.
- Fear of re-victimization on the part of sex crime victims: health care provider/ public health referral may be the best option for notification of partners in these cases.
- Anonymous partnering is a significant barrier to partner notification: wherever possible, encourage patient referral.

Note:

Actual or suspected child sexual abuse must be reported to your local child protection agency. The *Child Protection Act* supersedes all other acts and requires health professionals to release the names of any named contacts of a minor to the Children's Aid Society for further investigation.

Novel partner-notification practices

With changing trends in STI rates and transmission, research is being conducted to look at the feasibility of alternative methods of partner notification. One such method is the use of expedited patient-initiated treatment of sex partners. The index case is given medication, together with safety information and contraindications, to give to partners for presumptive treatment without assessment to reduce gonorrhea or chlamydia reinfections and to increase the proportion of partners treated. Although still controversial, this method may be beneficial in high-risk and hard-to-reach populations.^{11,12}

Practice points to maximize partner notification

- Request a notification form for STIs from the local public health unit or call the communicable disease reporting line for assistance.
- Develop a notification plan, including which partners will be notified by whom.
- See Table 5 for recommendations on partners to notify and the recommended trace back period for reportable and non-reportable STIs.

Table 5. Partner notification reference chart

Infection/syndrome	Reportable	Trace- back period	Who to notify/ evaluate	Special considerations
Chlamydia (LGV and non-LGV serovars)	Yes	60 days	SP/NB	If no sexual partner(s) in the last 60 days, trace back to last
Gonorrhea	Yes	60 days	SP/NB	sexual partner Partner notification
Chancroid	Yes	14 days	SP	is not required in most provinces and
Non-gonococcal urethritis	No	60 days	SP	territories as a public health measure but is highly recommended
Mucopurulent cervicitis	No	60 days	SP	for NGU, MPC, PID and epididymitis
Pelvic inflammatory disease	No	60 days	SP	, ,
Epididymitis	No	60 days	SP	
Primary syphilis	Yes	3 months	SP/NB	
Secondary syphilis	Yes	6 months	SP/NB	
Early latent syphilis	Yes	1 year	SP/NB	
Late latent syphilis/ stage undetermined	Yes	Variable	SP/NB/ CMC	
Genital herpes	In some jurisdictions	Current/ future	SP/NB	Partner notification is not required as a public health measure but is highly recommended

Table 5. Partner notification reference chart (continued)

Infection/syndrome	Reportable	Trace- back period	Who to notify/ evaluate	Special considerations
Trichomoniasis	In some jurisdictions	Current	SP	No need to test partners; treat as for index case
Human papillomavirus	No	Current/ future	SP	Partner notification is not required as a public health measure. Patients should be encouraged to notify their sexual partners, but there is no proof that this will lower the risk to the partner
Acute hepatitis B	Yes	Variable	SP/NSP/ HC/ NB/CMC	 All unvaccinated/ non-immune contacts should be notified. May benefit from PEP¹³ Newborns must receive HBIG and vaccine postnatally¹³
Chronic hepatitis B	Yes	Variable	SP/NSP/ HC/ NB/CMC	 All unvaccinated/ non-immune contacts should be notified. May benefit from PEP¹³ Newborns must receive HBIG and vaccine postnatally¹³

Table 5. Partner notification reference chart (continued)

Infection/syndrome	Reportable	Trace- back period	Who to notify/ evaluate	Special considerations
HIV/AIDS	Yes	Variable	SP/NSP/ NB/CMC	Start with recent sexual and needle-sharing partners; outer limit is onset of risk behaviour or to last known negative test PEP may be considered by health care providers for individuals who have been or are highly suspected to have been in contact with HIV. PEP, if considered, should be initiated within 72 hours. Please consult with an expert in HIV.

CMC = children of maternal case

HBIG = hepatitis B immune globulin

HC = household contacts

LGV = lymphogranuloma venereum

MPC = mucopurulent cervicitis

NB = newborns of infected mothers

NGU = non-gonococcal urethritis

NSP = needle-sharing partners PEP = post-exposure prophylaxis

PID = pelvic inflammatory disease

SP = sexual partners

10. Managing Co-morbidity and Associated Risks

Many STIs are transmitted in the context of other medical and social challenges. Recurrent exposure and infection are likely unless underlying issues are dealt with. Specific management for conditions such as drug addiction and mental health conditions must be integrated into the overall multidisciplinary health care plan.

When counselling and testing for STIs, it is important to include HIV pre-test counselling and offer testing. Being infected with an STI (including syphilis, genital herpes, chlamydia, gonorrhea and trichomoniasis) increases the risk of transmission and acquisition of HIV. HIV-infected individuals may be less responsive to STI treatment and require special monitoring post-treatment to ensure treatment effectiveness and to prevent long-term complications arising from inadequately treated STIs.

For individuals diagnosed with chronic viral hepatitis — either HBV or hepatitis C virus (HCV) — co-infection with HIV affects the choice of treatment, the response to treatment and individual patient outcomes. These patients should be referred to a specialist for treatment and management recommendations. Testing for viral hepatitis B and HIV in any chronically infected patient is required to ensure proper management of the infection. In addition, for those infected with HCV, ensuring vaccination against HAV and HBV is essential to prevent co-infection which can further assault the liver, complicate treatment options and compromise response to treatment and patient prognosis.¹⁴

If lymphogranuloma venereum (LGV) is suspected and linked to a current outbreak in Canada, it is also important to test for HCV, because there is a high rate of LGV-HCV co-infection.

11. Following up

Ideally, follow-up should be conducted by the same health care provider to ensure resolution of symptoms, follow-up testing as indicated and follow-through on partner notification to reduce the likelihood of reinfection. Where this is not possible, patients should be directed to the appropriate community resources, counselled on when to get follow-up (especially if tests were done) and advised of indicators of treatment failure. (See infection-specific chapters for follow-up recommendations.)

For individuals identified at ongoing risk for STIs, recommend screening for gonorrhea, chlamydia, syphilis and HIV at 3-month intervals and reinforce safer sexual practices.

RESOURCES

For a list of provincial and territorial STI contacts see *Appendix D* and for a list of current sexual health/STI/safer-sex resources to assist in counselling and assessing risk see *Appendix C*.

REFERENCES

- Public Health Agency of Canada. Terry's case. In: STD Self Learning Module. Available at: www.phac-aspc.gc.ca/slm-maa/terry/index.html. Accessed December 19, 2005.
- MacDonald NE, Wells GA, Fisher WA, et al. High-risk STD/HIV behavior among college students. *JAMA* 1990;263:3155–3159.
- Canadian Medical Association. Counselling Guidelines for HIV Testing. Ottawa, ON: Canadian Medical Association: 1995.
- 4. Renzi C, Tabet SR, Stucky JA, et al. Safety and acceptability of the Reality condom for anal sex among men who have sex with men. *AIDS* 2003;17:727–731.
- Gross M, Buchbinder SP, Holte S, Celum CL, Koblin BA, Douglas JM Jr. Use of reality "female condoms" for anal sex by US men who have sex with men. HIVNET Vaccine Preparedness Study Protocol Team. *Am J Public Health* 1999:89:1739–1741.
- Nonoxynol-9 and the risk of HIV transmission. HIV/AIDS Epi Update
 April 2003. Ottawa, ON: Public Health Agency of Canada; 2003. Available
 at: www.phac-aspc.gc.ca/publicat/epiu-aepi/hiv-vih/nonoxynol_e.html.
 Accessed December 19, 2005.
- Fisher JD, Cornman DH, Osborn CY, Amico KR, Fisher WA, Friedland G. Clinician-initiated HIV risk reduction intervention for HIV+ persons: formative research, acceptability, and fidelity of the Options Project. *J Acquir Immune Deficienc Syndr* 2004;37(suppl 2):S78–S87.
- 8. Fisher JD, Fisher WA, Cornman DH, Amico RK, Bryan A, Friedland GH. Clinician-delivered intervention during routine clinical care reduces unprotected sexual behavior among HIV-infected patients. *J Acquir Immune Deficienc Syndr* 2006;41:44–52.
- 9. Rollnick S, Mason P, Butler C. *Health Behavior Change. A Guide for Practitioners.* Edinburgh: Churchill Livingstone; 1999.
- Program Operations Guidelines for STD Prevention. Atlanta, GA: Centers for Disease Control and Prevention; 2005. Available at: www.cdc.gov/std/ program/partner/ TOC-PGpartner.htm. Accessed December 19, 2005.
- Mathews C, Coetzee N, Zwarenstein M, et al. Strategies for partner notification for sexually transmitted diseases. *Cochrane Database Syst Rev* 2001; 4:CD002843
- 12. Golden MR, Whittington WL, Handsfield HH, et al. Effect of expedited treatment of sex partners on recurrent or persistent gonorrhea or chlamydial infection. *N Engl J Med* 2005;352:676–685.
- 13. Canadian Immunization Guide. 6th ed. Ottawa, ON: Health Canada; 2002.
- 14. Sherman M, Bain V, Villeneuve JP, et al. Management of viral hepatitis: a Canadian consensus conference 2003/2004. Ottawa, ON: Health Canada; 2004. Available at: www.phac-aspc.gc.ca/hepc/hepatitis_c/pdf/ccc_04/pdf/consensus_04.pdf. Accessed December 19, 2005.

LABORATORY DIAGNOSIS OF SEXUALLY TRANSMITTED INFECTIONS

A. COLLECTION AND TRANSPORTATION OF SPECIMENS¹

General Principles

- Swabs, transport systems and types of tests used may vary depending on the agent sought and techniques used by the laboratory.
- Contact the laboratory to obtain further information, especially concerning transport requirements, turn-around time and interpretation of results. See Appendix E for a listing of local contact information.
- Laboratories may use a variety of commercial specimen-collection devices. Follow the instructions provided by the manufacturer.
- All specimen-collection and handling procedures should be performed while wearing appropriate protective clothing and following recommended universal precautions.
- Contamination from indigenous commensal flora should be avoided to ensure a representative sampling of organisms involved in the infectious process.
- Adequate volumes of each liquid specimen should be collected.
- Each specimen container should be labelled with the patient's name and identification number, the source of the specimen and the date and time of collection.
- All specimen containers should be leak-proof and transported within a sealable, leak-proof plastic bag that has a separate compartment for paperwork.
- Sexually transmitted pathogens are usually fastidious and fragile; cultures and techniques that detect viable organisms may give false-negative results unless storage and transport conditions are optimal.
- Storage recommendations need to be observed, and transport must be as rapid as possible for the recovery of infectious organisms, with excesses of temperature avoided.

Specimens

For most sexually transmitted infections (STIs), specimens will be collected by health care providers to be packaged and delivered to diagnostic laboratories. There is an effort to produce commercial point-of-care testing kits for in-office testing, but there are none that are approved and validated at this time. Self-collection of urine, vaginal and lesion swabs is currently being evaluated for home collection, but these strategies lack appropriate evaluation, especially for transportation conditions.

1. Cervix

- After inserting a speculum to view the cervix, remove overlying vaginal secretions and cervical exudate.
- Insert a sterile swab 1–2 cm into the endocervical canal, rotate 180° and withdraw for collection of columnar epithelial cells for diagnosis of *Chlamydia trachomatis* and *Neisseria gonorrhoeae*. The choice of swab should be based on the type of testing being done; consult with the laboratory providing the service.
- Obtain a specimen for N gonorrhoeae before taking a specimen for C trachomatis.
- If a culture is to be performed for *N gonorrhoeae*, directly inoculate the culture tube or plate, or place the swab in the transport medium. Alternatively, place the swab in a nucleic acid amplification transport tube.
- Exocervical samples are better for herpes simplex virus (HSV) and human papillomavirus (HPV).
- Vaginal swabs may be submitted for culture from women who are menstruating or have had a hysterectomy.

Notes:

- Cervical specimens should not be taken from prepubertal girls, since STIs in this
 age group involve the vagina, not the cervix. See Sexual Abuse in Peripubertal
 and Prepubertal Children chapter for more information.
- Obtaining several specimens from the cervix does not usually produce discomfort and may be required to perform various tests.

2. Lesions (vesicles or ulcers)

- a) Vesicles
- Fluid can be obtained by lifting the top of the vesicle and swabbing the lesion.
- An alternative method is to clean the vesicle with a disinfectant and, after drying, piercing into the fluid with a syringe, collecting fluid, capping, sealing the plunger and transporting to the laboratory.

b) Ulcers

- Warn the patient that specimen collection may be painful.
- Swab the lesion bed for culture, polymerase chain reaction (PCR) or direct examination for HSV.
- For direct examination, obtain cellular material by firm swabbing or gentle scraping from the base of the lesion.
- For culture, use the swab and viral transport medium supplied with the collection kit.
- For the detection of *Treponema pallidum*, contact the laboratory to determine the availability of dark-field microscopy or direct fluorescent antibody (DFA) testing. Where available, collect a specimen as follows:
 - Remove scabs or overlying debris.
 - Cleanse the lesion with sterile saline without preservatives and dry the area.

- Abrade the lesion with a dry sterile gauze pad to provoke slight bleeding and exudation of tissue fluid.
- As oozing occurs, wipe away the first few drops and await the appearance of relatively clear serous exudate. It is sometimes necessary to apply pressure at the base of the lesion to express tissue fluid.
- Collect fluid into a capillary tube, small-bore syringe or directly onto a slide for DFA testing.
- Seal the tube, cap the syringe or immobilize the plunger before transportation.
- Store at 4°C before transportation and deliver to the laboratory within 24 hours.
- For Haemophilus ducreyi, a special medium is required for culture. Obtain a swab from the base of a lesion, avoiding pus, and place into a transport tube.

3. Pharynx

- Swab the posterior pharynx and the tonsillar crypts.
- Use the swab to directly inoculate the appropriate culture medium, or place it in a transport medium.
- · For infants, obtain a nasopharyngeal aspirate.

Notes:

- There are promising data on the performance of non-culture tests using pharyngeal specimens.
- Smears of pharyngeal swabs are of no value in detecting pharyngeal N gonorrhoeae and are not recommended.

4. Rectum

- For blind swabbing, insert 2–3 cm into the anal canal, press laterally to avoid fecal material and, in the case of *C trachomatis* or *N gonorrhoeae*, to obtain columnar epithelial cells.
- If there is visible fecal contamination, discard the swab and obtain another specimen.
- With unlubricated anoscopy using only tap water, fecal contamination can be avoided and specimens can be collected under direct visualization.

Notes:

- Specimens may be obtained blindly or through an anoscope. The latter is preferred for symptomatic patients.
- There are promising data needing confirmation on the use of rectal swabs for N gonorrhoeae and C trachomatis in nucleic acid amplification tests (NAATs), but at this time neither use is approved.

5. Urethra

- Warn the patient that specimen collection may be painful, that the next urination may be painful, and that increasing fluid intake may help to decrease urine concentration and therefore discomfort.
- Ideally, the patient should not have voided for at least 2 hours, as voiding reduces the amount of exudate and may decrease the ability to detect organisms.
- Use a thin, dry swab with a flexible wire shaft. Moistening the swab with water before insertion may help reduce discomfort.
- Introduce the swab slowly (3–4 cm in males; 1–2 cm in females), rotate slowly and withdraw gently.
- The swab can be used to prepare a smear by slowly unrolling the secretions onto a slide; then, directly inoculate the appropriate culture medium or place the swab in a transport medium.
- If a NAAT is used, follow the manufacturer's instructions.

Notes:

- "Milking" the penis three or four times from the base to the glans enhances the ability to detect otherwise inapparent urethral discharge.
- In prepubertal boys and girls, collection of an intraurethral specimen is not recommended; obtain first-void urine specimens for NAATs or a meatal specimen using a thin swab with a flexible wire shaft.

6. Urine (first-void)

- The patient should not have voided for at least 2 hours, but having done so does not preclude testing.
- Provide the patient with a leak-proof container.
- Ask the patient to collect only the first 10–20 mL of urine² into the container and to cap it tightly.

Note:

Most commercial NAATs for *C trachomatis* and *N gonorrhoeae* are approved for urine testing and are recommended for detecting these organisms in asymptomatic men or women, women without a cervix or those who wish to avoid pelvic examination. A first-void urine (FVU) may be collected at any time and may also be termed a first-catch urine (FCU).

7. Vagina

- · Collect pooled vaginal secretions, if present.
- When no vaginal secretions are present, swab the vaginal wall in the posterior fornix to prepare a smear, or place the swab in a transport medium.
- Wet-mount and Gram-stain smears are useful in the diagnosis of microbial vulvovaginitis, candidiasis, bacterial vaginosis, trichomoniasis or desquamative inflammatory vaginitis.

- Collection of vaginal specimens from youth and adults is usually done as part of a speculum examination.
- In prepubertal girls, vaginal-wash specimens are most preferred and patient acceptable. If not possible, use swabs moistened with water. See *Sexual Abuse in Peripubertal and Prepubertal Children* chapter for more information.
- In very young children, use very thin swabs.

Note:

In the past, vaginal specimens were not recommended for the diagnosis of STIs, except in the management of vulvovaginitis, bacterial vaginosis and child sexual abuse. More recent data show that NAATs for *C trachomatis*, *N gonorrhoeae* and *Trichomonas vaginalis* may identify as many or more infected women using vaginal swabs than cervical swabs, urethral swabs or urine.³ Check with your laboratory to see if this is an option.

8. Warts and Other HPV Infections

- Scrape the exocervix for superficial epithelial cells.
- Cytobrushes, other collecting devices or swabs can be used to collect cells from the squamo-columnar junction of the cervix.
- Currently commercial and non-commercial assays with specific collection devices are available for HPV DNA detection. Consult with your laboratory.

Note:

Urine samples have not been shown to be as accurate as cervical samples for detecting high-risk HPV.⁴

B. LABORATORY TESTING METHODS

STIs may be diagnosed in the laboratory using (a) culture, (b) microscopy, (c) antigen detection, (d) nucleic acid detection, (e) serology and (f) surrogate markers. The sensitivity and specificity of these different approaches vary according to specimen type and organism assayed. The number of false positives or negatives will be influenced by the prevalence of infection in the population being sampled. NAATs are the most sensitive methods, and culture the most specific. Antigen detection, nucleic acid hybridization, culture and microscopy are less sensitive but may be effective for certain types of patients and specimen types. Since not all diagnostic laboratories perform the same tests, clinical conditions and specimen types should be discussed before collecting the specimen. In some situations, serology is very useful (e.g., syphilis), but in others (e.g., non-LGV *C trachomatis*) it is of no use. Surrogate markers such as leukocyte esterase strip tests, pH or amines point-of-care tests may provide useful screening for some conditions, but are generally insensitive and not very specific.^{5,6}

C. LABORATORY DIAGNOSIS OF SPECIFIC INFECTIONS

1. Chlamydia trachomatis

- Results are highly dependent on the type of test available, ⁷ appropriate specimen collection, ⁸ storage, transport and laboratory expertise.
- Contact the laboratory for specific instructions before submitting specimens, and read and follow test-kit instructions regarding specimen collection, storage and transport.
- NAATs are the most sensitive and specific and should be used whenever possible for urine, urethral, and cervical specimens; blood and mucous can affect NAAT performance.⁹
- Non-invasive specimens such as urine can be used in NAATs, making testing more acceptable to patients.¹⁰
- Both C trachomatis and N gonorrhoeae can be detected from a single specimen in some NAATs.¹¹
- Because successful treatment rates are high, a test of cure is not usually performed.
- Other assays, such as nucleic acid hybridization and antigen detection, may be used, but they are less sensitive and specific, and positives may need to be confirmed.¹²
- C trachomatis IgM serology is useful for diagnosing C trachomatis pneumonia in infants under 3 months of age.¹³
- Serology is not useful for the diagnosis of acute genital chlamydial infections (non-LGV only).
- Culture is the preferred method for medico-legal purposes, but NAATs may be suitable, provided that positive results are confirmed using a different set of primers, which may not be readily available in most labs.
- Strains of lymphogranuloma venereum (LGV) have emerged in Europe and North America, mainly in rectal samples (RS) of men who have sex with men (MSM). Existing NAATs are not cleared by the U.S. Food and Drug Administration or Health Canada for use on rectal or oropharyngeal samples, but will record positives that need to be confirmed as LGV by restriction fragment length polymorphism (RFLP) or sequencing techniques. Samples can also be cultured undiluted and at a dilution of 1:10 (to dilute fecal toxicity) using shell vials with and without centrifugation. LGV grows readily to high levels of elementary bodies without centrifugation, while non-LGV strains require centrifugation. As with NAATs, positive cultures need to be confirmed as LGV by RFLP or sequencing. A NAAT or culture can also be used on other samples in the diagnosis of LGV such as bubo aspirates; urine; or rectal, vaginal or urethral swabs. Emphasis should be placed on clinical samples for a definitive diagnosis; however, serology such as microimmunofluorescence (MIF) may be helpful in supporting the diagnosis. For more information on specimen collection and available tests, please contact your local laboratory (See Lymphogranuloma Venereum chapter for more information on specimen collection and testing by stage of infection).

2. Neisseria gonorrhoeae

- The presence of Gram-negative diplococci inside polymorphonuclear leukocytes (PMNs) is highly predictive for the direct microscopic examination of smears; their presence outside PMNs is not, and confirmation by culture is required.
- The sensitivity and specificity of the Gram stain depends on the type of specimen.¹⁴ Urethral specimens from young adult males have a sensitivity and specificity of 95%; endocervical specimens from adult females have a sensitivity of 45–65% and a specificity of 90%.
- Culture for N gonorrhoeae is required for the determination of antimicrobial susceptibility in cases of sexual abuse/assault, as well as in cases of treatment failure.
- Successful culture of specimens requires proper collection and transportation of appropriate specimens or immediate inoculation of medium.¹⁵ Consult with your laboratory.
- NAATs are approved for cervical and urethral swabs and urine; some NAATs
 are also approved for vaginal swabs.¹¹ Urine and vaginal swabs are convenient
 specimens for women without a cervix, and urine may be most convenient for
 those who may not readily submit to a pelvic examination.
- Urine is a preferred specimen for men if a NAAT is performed.
- A NAAT is not recommended as a test of cure.
- NAAT can be used to detect reinfection, after waiting for at least 2 weeks after completion of therapy.
- For medico-legal purposes, a positive result obtained from NAATs should be confirmed using a different set of primers.
- Serology is not available.

3. Haemophilus ducreyi (chancroid)

- Because H ducreyi is rare in Canada, consult with your laboratory.
- Culture is the current method of choice, using two media in a biplate.16
- Specimens of choice are a calcium alginate or cotton swab from the base of the ulcer or an aspirate if buboes are present.
- There are no useful serologic tests to diagnose H ducreyi. Gram stain with Gram-negative coccobacilli in a "school of fish" pattern may be useful.
- If a NAAT is available, a second ulcer swab should be collected into an appropriate transport medium.

4. Herpes simplex virus

- NAATs are being used increasingly for cerebrospinal fluid, vesicle fluid or ulcer swabs.¹⁷ Consult with your laboratory.
- NAATs approach sensitivities and specificities of 100%, with rapid turn-around of results.
- Cultures are easy to perform and can yield positive results within 24 hours from primary or first-episode genital herpes.

- Other methods, such as antigen detection and Tzanck smear cytology, lack accuracy.
- For neonates, gently rub conjunctiva, insert separate swab into mouth (and gently rub around the lips), external ear canal, umbilicus, axillae and groin.
 Specimens should be collected 24–48 hours after birth.
- Type-specific antibody assays are commercially available and may be useful in some clinical situations (though availability in Canada is currently limited):
 (a) patients with apparent first-episode genital herpes with a negative culture or NAAT;
 (b) identification of a seropositive pregnant woman with no history of herpes;
 (c) counselling HSV serologically discordant couples.¹⁸

5. Treponema pallidum (syphilis)

- Consult with your laboratory on tests available.
- When lesions are present in primary, secondary or early congenital syphilis, clear serous fluid should be collected for dark-field microscopy, enabling observation of morphology and movement of the spirochetes (not reliable for oral or rectal lesions).¹⁹
- Other non-serological methods involve direct fluorescent antibody tests or NAATs. The latter are very sensitive and specific.²⁰
- In cases where pregnant women are suspected of having syphilis, sections of placenta should be collected at birth and sent for DFA testing.
- Serological diagnosis involves initial screening of sera by non-treponemal tests such
 as the Venereal Disease Research Laboratory (VDRL), rapid plasma reagin (RPR),
 toluidine red unheated serum test (TRUST) or the reagin screening test (RST).
- Sera positive in non-treponemal tests are retested by treponemal assays such as the *Treponema pallidum* particle agglutination (TP-PA) test, fluorescent treponemal antibody absorption (FTA-ABS) test and microhemagglutination for *Treponema pallidum* (MHA-TP).²¹ Several enzyme immunoassays (EIA) have been developed commercially to detect IgG or IgM to specific *T pallidum* antigens and are useful in HIV co-infected patients. See *Syphilis* chapter for information on cerebrospinal fliud examination.

6. Human Immunodeficiency Virus

- HIV diagnostic laboratories in Canada are instructed to use only tests approved by Health Canada.
- Sera are initially screened by EIA and may detect antibodies by 3 weeks after infection, but can take up to 6 months.²²
- All positives are confirmed using a different EIA or Western blot.
- Qualitative PCR is used to detect small amounts of nucleic acid in babies born to HIV-infected mothers.
- Quantitative PCR (viral load testing) is used to monitor HIV-positive patients prior to and during antiretroviral therapy.²³
- Genotyping is used to detect the development of drug resistance in selected patients, enabling physicians to choose appropriate antiretroviral drug combinations.²⁴

7. Human papillomavirus

- Liquid-based cytology has increased the accuracy of Pap testing, and the HPV signal amplification hybrid capture assay (Digene) can be performed on the same or a separate cervical sample.²⁵
- The presence of high-risk HPV in patients with atypical squamous cells of undetermined significance (ASCUS) may enable recommendation for immediate colposcopy.²⁶
- Microscopy, culture and antigen detection have no proven utility for the diagnosis of HPV infections.
- NAATs and serology are only for epidemiological purposes at the present time.
- Consult with your laboratory concerning HPV testing, as few laboratories are currently providing this service in Canada.

8. Hepatitis B virus

- Patients acutely infected with HBV will have positive EIA results for hepatitis
 B surface antigen (HBsAg) and/or anti-hepatitis B core (anti-HBc) IgM tests
 performed on sera.
- Most patients (90%) develop immunity within 6 months of infection, lose their HBsAg and have it replaced by anti-HBc IgG and anti-hepatitis B surface antibodies (anti-HBs).²⁷
- Patients chronically infected will demonstrate HBsAg persistence for 6 months or more.
- The presence of hepatitis B e antigen (HBeAg) in acutely or chronically infected individuals indicates greater infectivity for contacts and for babies born to positive mothers.²⁸ These antigens may eventually be replaced by antibodies (anti-HBe).
- Quantitative PCR assays to detect viral DNA are available to monitor response to treatment.^{29,30}

9. Hepatitis A virus

- The presence of hepatitis A virus (HAV) IgM antibodies, which may be present for 3 months, is diagnostic of acute infection.³¹
- HAV IgG antibody testing can demonstrate immunity.

10. Trichomonas vaginalis

- The vaginal pH is >4.5, and the whiff test is usually negative (the withdrawn speculum does not have an abnormal odour).³²
- Because of the low sensitivity of direct microscopy, culture may be used, where available, to isolate the parasite from urethral swabs, urine sediments, prostate fluid and vaginal specimens.³³

11. Candida albicans

- The vaginal pH is normal (<4.5), and the whiff test is negative.34
- Wet-mount preparation with 10% KOH shows budding yeast and/or branching pseudohyphae.

12. Bacterial vaginosis

- The vaginal pH is >4.5, and the whiff test is positive.³⁵
- Gram stain demonstrates a shift in vaginal flora, with a decrease in large Grampositive rods (lactobacilli) and an increase in small Gram-variable coccobacilli and clue cells (vaginal epithelial cells covered with numerous coccobacilli).

REFERENCES

- Chernesky MA. Laboratory services for sexually transmitted diseases: overview and recent developments. In: Holmes KK, Sparling P, Mardh PA, et al, eds. Sexually Transmitted Diseases. 3rd ed. New York, NY: McGraw Hill; 1999: 1281–1294.
- 2. Chernesky M, Jang D, Chong S, Sellors J, Mahony J. Impact of urine collection order on the ability of assays to identify Chlamydia trachomatis infections in men. *Sex Transm Dis* 2003;30:345–347.
- Schachter J, McCormack WM, Chernesky MA, et al. Vaginal swabs are appropriate specimens for diagnosis of genital tract infection with Chlamydia trachomatis. J Clin Microbiol 2003;41:3784–3789.
- Sellors J, Lorincz AT, Mahony JB, et al. Comparison of self-collected vaginal, vulvar and urine samples with physician-collected cervical samples for human papillomavirus testing to detect high-grade squamous intraepithelial lesions. CMAJ 2000;163:513–518.
- O'Brien SF, Bell TA, Farrow JA. Use of a leukocyte esterase dipstick to detect Chlamydia trachomatis and Neisseria gonorrhoeae urethritis in asymptomatic adolescent male detainees. Am J Public Health 1988;78:1583–1584.
- Hedin G, Abrahamsson G, Dahlberg E. Urethritis associated with Chlamydia trachomatis: comparison of leukocyte esterase dipstick test of first-voided urine and methylene blue-stained urethral smear as predictors of chlamydial infection. APMIS 2001;109:595–600.
- 7. Van Dyck E, Ieven M, Pattyn S, Van Damme L, Laga M. Detection of Chlamydia trachomatis and Neisseria gonorrhoeae by enzyme immunoassay, culture, and three nucleic acid amplification tests. *J Clin Microbiol* 2001;39:1751–1756.
- Shafer M, Moncada J, Boyer CB, Betsinger K, Flinn SD, Schachter J.
 Comparing first-void urine specimens, self-collected vaginal swabs, and
 endocervical specimens to detect Chlamydia trachomatis and Neisseria
 gonorrhoeae by a nucleic acid amplification test. *J Clin Microbiol*2003;43:4395–4399.
- 9. Chernesky MA. The laboratory diagnosis of Chlamydia trachomatis infections. *Can J Infect Dis Med Microbiol* 2005;16:39–44.

- Serlin M, Shafer MA, Tebb K, et al. What sexually transmitted disease screening method does the adolescent prefer? Adolescents' attitudes toward first-void urine, self-collected vaginal swab, and pelvic examination. *Arch Pediatr Adolesc Med* 2002;156:588–591.
- Gaydos CA, Quinn TC, Willis D, et al. Performance of the APTIMA Combo 2 assay for detection of Chlamydia trachomatis and Neisseria gonorrhoeae in female urine and endocervical swab specimens. *J Clin Microbiol* 2003;41:304–309.
- Clarke LM, Sierra MF, Daidone BJ, Lopez N, Covino JM, McCormack WM. Comparison of the Syva MicroTrak enzyme immunoassay and Gen-Probe PACE 2 with cell culture for diagnosis of cervical Chlamydia trachomatis infection in a high-prevalence female population. *J Clin Microbiol* 1993; 31:968–971.
- 13. Mahony JB, Chernesky MA, Bromberg K, Schachter J. Accuracy of an IgM immunoassay for the diagnosis of chlamydial infections in infants and adults. *J Clin Microbiol* 1986;24:731–735.
- 14. Ng LK, Martin IE. The laboratory diagnosis of Neisseria gonorrhoeae. *Can J Infect Dis Med Microbiol* 2005;16:15–25.
- 15. Whittington W, Ison C, Thompson S. Gonorrhoea. In: Morse S, ed. *Atlas of Sexually Transmitted Diseases and AIDS*. 2nd ed. London: Mosby-Wolfe; 1996: 99–117.
- 16. Alfa M. The laboratory diagnosis of Haemophilus ducreyi. *Can J Infect Dis Med Microbiol* 2005;16:31–34.
- 17. Singh A, Preiksaitis J, Romanowski B. The laboratory diagnosis of herpes simplex virus infections. *Can J Infect Dis Med Microbiol* 2005;16:92-98.
- 18. Ashley RL. Sorting out the new HSV type specific antibody tests. Sex Transm Infec 2001;77:232–237.
- 19. Ratnam S. The laboratory diagnosis of syphilis. *Can J Infect Dis Med Microbiol* 2005;16:45–51.
- Wicher K, Hororitz HW, Wicher V. Laboratory methods of diagnosis of syphilis for the beginning of the third millennium. *Microbes Infect* 1999;1:1035–1049.
- Stoll BJ, Lee FK, Larsen S, et al. Clinical and serologic evaluation of neonates for congenital syphilis: a continuing diagnostic dilemma. *J Infect Dis* 1993; 167:1093–1099.
- 22. Fearon M. The laboratory diagnosis of HIV infections. *Can J Infect Dis Med Microbiol* 2005;16:26–30.
- Phillips KA, Bayer R, Chen JL. New Centers for Disease Control and Prevention's guidelines on HIV counseling and testing for the general population and pregnant women. *J Acquir Immune Defic Syndr* 2003; 32:182–191.
- Hirsch MS, Brun-Vezinet F, Clotet B, et al. Antiretroviral drug resistance testing in adults infected with human immunodeficiency virus type 1: 2003 recommendations of an international AIDS society–USA Panel. *Clin Infect Dis* 2003;37:113–128.

- Coutlee F, Rouleau D, Ferenczy A, Franco E. The laboratory diagnosis of genital human papillomavirus infections. Can J Infect Dis Med Microbiol 2005;16:83–91.
- Wright TC Jr, Cox JT, Massad LS, Twiggs LB, Wilkinson EJ; ASCCP-Sponsored Consensus Conference. 2001 Consensus Guidelines for the management of women with cervical cytological abnormalities. *JAMA* 2002:287:2120–2129.
- 27. Krajden M, McNabb S, Petric M. The laboratory diagnosis of hepatitis B virus. Can J Infect Dis Med Microbiol 2005;16:65–72.
- Okada K, Kamiyama I, Inomata M, Imai M, Miyakawa Y. e antigen and anti-e in the serum of asymptomatic carrier mothers as indicators of positive and negative transmission of hepatitis B virus to their infants. N Engl J Med 1976;294:746–749.
- 29. Chu CJ, Hussain M, Lok AS. Quantitative serum HBV DNA levels during different stages of chronic hepatitis B infection. *Hepatology* 2002;36:1408–1415.
- Lok AS, Zoulim F, Locarnini S, et al. Monitoring drug resistance in chronic hepatitis B virus (HBV)- infected patients during lamivudine therapy: evaluation of performance of INNO-LiPA HBV DR assay. *J Clin Microbiol* 2002;40:3729–3734.
- 31. Chernesky MA, Gretch D, Mushahwar IK, Swenson PD, Yarbough PO. Laboratory diagnosis of hepatitis viruses. *Cumitech* 1998;Nov:18A.
- 32. Garber GE. The laboratory diagnosis of Trichomonas vaginalis. *Can J Infect Dis Med Microbiol* 2005;16:35–38.
- 33. Beal C, Goldsmith R, Kotby M, et al. The plastic envelope method, a simplified technique for culture diagnosis of trichomoniasis. *J Clin Microbiol* 1992;30:2265–2268.
- 34. Hillier S, Arko R. Vaginal infections. In: Morse S, ed. *Atlas of Sexually Transmitted Diseases and AIDS.* 2nd ed. London: Mosby-Wolfe; 1996: 149–158.
- 35. Money D. The laboratory diagnosis of bacterial vaginosis. *Can J Infect Dis Med Microbiol* 2005;16:77–79.

MANAGEMENT AND TREATMENT OF SPECIFIC SYNDROMES

SYNDROMIC MANAGEMENT OF SEXUALLY TRANSMITTED INFECTIONS

Diagnosis of a syndrome according to standard criteria predicts the likelihood that a specific pathogen or pathogens is/are present and thus facilitates initiation of appropriate empiric treatment at the first visit rather than deferring treatment until there is microbiological confirmation. In the context of variable access to laboratory testing and variable rates of follow-up, the syndromic approach takes on greater relevance in controlling transmission and negative sequelae. See Table 1, below, for the management of sexually transmitted infection (STI) syndromes.

While the syndromic approach is an important tool in the control of STIs and their sequelae, management by syndrome alone is inadequate because infections with important pathogens such as *Chlamydia trachomatis* and *Neisseria gonorrhoeae* may be present without any symptoms or findings. Although an infection may be suspected because of disease in a partner or the presence of another STI, the infection may be diagnosed only by using a specific laboratory test. Thus, in managing STIs, diagnosis by syndrome and laboratory diagnosis by testing for specific organisms are both important and complementary. Consult the chapters of the *Management and Treatment of Specific Infections* section for details on the diagnosis, treatment and management of specific infections.

Table 1. Syndromic approach to the management of sexually transmitted infections

(Patients may present with more than one STI; this table provides an outline of investigations and relevant chapters where more in-depth information can be found. In many cases, screening for other STIs should be carried out.)

Syndrome	Signs and symptoms	Etiology	Specimens and testing	Microscopy results and clinical findings	Next steps/ special considerations
Asymptomatic and at risk for STIs (see Primary Care and Sexually Transmitted Infections	None	Neisseria gonorrhoeae Chlamydia trachomatis Treponema pallidum Herpes simplex virus type 1 or 2 Human papilloma- virus HIV Viral hepatitis	First-catch urine Urethral swab Cervical swab for: C trachomatis N gonorrhoeae		If testing is done by methods other than NAAT and sexual contact occurred <1 week prior to testing, tests may be falsely negative
chapter).			Serology for: Syphilis HIV		Typical window period for syphilis is 6 weeks Typical window period for HIV is 3 months
			Hepatitis A (particularly with oral-anal contact)		If non-immune for hepatitis A and B, consider immunization
			Hepatitis B (if no history of vaccine) Hepatitis C (particularly in IDU)		For chronic viral hepatitis, consult a colleague experienced in this area
		Pap testing if indicated (as per local or provincial/ territorial recommendations)	An abnormal Pap test result (e.g., ASCUS, LSILS) is not diagnostic of HPV	Follow up as per recommendations of province/territory	

ASCUS = atypical squamous cells of undetermined significance bHCG = beta human chorionic gonadotropin DFA = direct fluorescent antibody EIA = enzyme immunoassay FTA-ABS = fluorescent treponemal antibody absorbed HPF = high-power field

HPV = human papillomavirus
HSV = herpes simplex virus
IDU = injection drug use
LGV = lymphogranuloma venereum
LSIL = low-grade squamous
intraepithelial lesions
MHA-TP = microhemagglutinationTreponema pallidum
MSM = men who have sex with men

NAAT = nucleic acid amplification test PMN = polymorphonuclear leukocytes RPR = rapid plasma reagin RUQ = right upper quadrant STI = sexually transmitted infection VDRL = venereal disease research laboratory

Table 1. Syndromic approach to the management of sexually transmitted infections (continued)

Syndrome	Signs and symptoms	Etiology	Specimens and testing	Microscopy results and clinical findings	Next steps/ special considerations
Urethritis	Urethral discharge Burning on urination Irritation in the distal urethra or meatus	gonorrhoeae C trachomatis Trichomonas	causes: for Gram stain and culture for gonorrhoeae gonorrhae (NAAT may also be used where available) and Trichomonas vaginalis Trichomonas vaginalis First-catch urine Herpes for C trachomatis simplex (NAAT) prices (NAAT) Virus Mycoplasma genitalium Ureaplasma urealyticum	Presence of ≥5 PMNs per HPF and absence of Gram-negative diplococci (likely Non-Gonococcal Urethritis)	See urethritis treatment flow chart in the <i>Urethritis</i> chapter for treatment and management recommendations
	Meatal erythema	Herpes simplex virus Mycoplasma genitalium Ureaplasma		Presence of ≥5 PMNs per HPF AND Gram-negative intracellular or extracellular diplococci OR Gram-negative intracellular diplococci alone.	See Gonococcal Infections chapter for treatment recommendations
				Presence of Gram-negative extracellular diplococci alone requires further testing.	See Table 5 in Gonococcal Infections chapter
				Where microscopy results are not immediately available	Treat as per recommendations for chlamydia and gonorrhea.
					If patient treated for gonorrhea and chlamydia and symptoms persist consider other causes or resistance in the case of gonorrhea (see Gonococcal Infections chapter)

OF SPECIFIC SYNDROMES

Table 1. Syndromic approach to the management of sexually transmitted infections (continued)

Syndrome	Signs and symptoms	Etiology	Specimens and testing	Microscopy results and clinical findings	Next steps/ special considerations
Cervicitis (females)	(females) lent cervical discharge Cervical friability gor Vaginal discharge Strawberry cervix Strawberry cervix	causes: for N g Cul gonorrhoeae C trachomatis Trichomonas vaginalis HSV Sw les	Cervical swab for Gram stain, N gonorrhoeae culture and C trachomatis (NAAT or culture)	On Gram stain, presence of ≥20 PMNs per HPF with mucopuru- lent discharge and/or cervical friability	See Chlamydial Infections chapter for treatment recommendations unless gonorrhea is suspected; then, see Gonococcal Infec- tions chapter; although not a sensitive test, Gram stain may be helpful in diagnosing mucopurulent cervi- citis and gonorrhea in symptomatic females
			Swab of cervical lesions for HSV		If HSV is suspected or detected see <i>Genital</i> <i>Herpes Simplex Virus</i> <i>Infections</i> chapter for treatment recommendations
			Vaginal swab for wet mount	Trichomonads	See Vaginal Discharge chapter for treatment recommendations
				Where microscopy results are not immediately available	Treat as per recommendations for chlamydia and gonorrhea

Table 1. Syndromic approach to the management of sexually transmitted infections (continued)

Syndrome	Signs and symptoms	Etiology	Specimens and testing	Microscopy results and clinical findings	Next steps/ special considerations
Genital ulcer disease	ulcer (erosive or pustular) Herpe Vesicles simple Papules Inguinal lymphade- nopathy comm comm Herpe Vericles simple virus 1 C trace matis	Most common: Herpes simplex virus 1 or 2 T pallidum C trachomatis (LGV serovars L1,	Routine: Swab of lesion for culture (herpes)	Herpes Painful lesions Grouped vesicles Erythematous base Fever and malaise	Consider genital herpes and empiric treatment for either primary or suspected recurrent infection (see <i>Genital Herpes Simplex Virus Infections</i> chapter for treatment recommendations)
			Swab of serous fluid from lesion for dark-field microscopy or DFA for syphilis. Check with laboratory re: availability and Serology for syphilis to include both non-treponemal (RPR/VDRL/EIA) and treponemal-specific tests (MHA-TP and FTA-ABS)	Syphilis Non-painful lesions Indurated with serous exudate Single lesion in over 70% of cases	Consider primary syphilis. Empiric treatment should be considered if follow-up is uncertain
			Non-routine: If indicated through patient history Swab of lesion for non-LGV <i>C trachomatis</i> for culture (MSM, travel) or consider serology for <i>C trachomatis</i>	If initial <i>C trachomatis</i> testing is positive, serovarspecific testing is required to confirm a diagnosis of LGV. See <i>Lymphogranuloma Venereum</i> chapter	If LGV is suspected, treat empirically according to the recommendations in Lymphogranuloma Venereum chapter
			Consider testing for chancroid and granuloma ingui- nale (link to travel); consult laboratory for availability		See <i>Genital Ulcer Disease</i> chapter for treatment recommendations

OF SPECIFIC SYNDROMES

Table 1. Syndromic approach to the management of sexually transmitted infections (continued)

Syndrome	Signs and symptoms	Etiology	Specimens and testing	Microscopy results and clinical findings	Next steps/ special considerations
Epididymitis	Unilateral testicular pain/ swelling May have erythema and edema of the over- lying skin With or without urethral discharge	Most common (varies with age): C trachomatis N gonorrhoeae Coliforms Pseudomonads	First-catch urine for NAAT (<i>C trachomatis</i>); may be used for gonorrhea where available Midstream urine for culture and sensitivity (enteric organisms, coliforms) Urethral swab for Gram stain and gonorrhea culture	Palpable swelling of the epididymis Gram stain: presence of ≥5 PMNs per HPF and/or Gram-negative intracellular diplococci Gram stain: absence of PMNs and Gram-negative intracellular diplococci	For empiric treatment recommendations, see Epididymitis chapter See Epididymitis chapter for treatment recommendations for epididymitis likely caused by chlamydial or gonococcal infections See Epididymitis chapter for treatment of organisms other than chlamydia or gonorrhea
			Doppler ultrasound if testicular torsion is suspected		If symptom onset is rapid, testicular torsion needs to be consid- ered, as this is a surgical emergency

Table 1. Syndromic approach to the management of sexually transmitted infections (continued)

Syndrome	Signs and symptoms	Etiology	Specimens and testing	Microscopy results and clinical findings	Next steps/ special considerations
Pelvic inflamma- tory disease	Lower abdominal pain Deep dyspareunia Abnormal bleeding Fever	C trachomatis N gonorrhoeae Genital-tract mycoplasms Other aerobic or anaerobic bacterial species	Cervical swab for Gram stain and gonorrhea culture Cervical swab for <i>C trachomatis</i> (NAAT or culture) Vaginal swab for culture, Gram stain, PH test, whiff test and wet mount Urine ± serum bHCG to rule out ectopic pregnancy Other serological tests: ESR C-reactive protein	On bimanual exam: Cervical motion tenderness Adenexal tenderness Adenexal masses Other findings: RUQ pain Cervicitis Fever	For empiric treatment recommendations and definitive diagnostic criteria, see <i>Pelvic Inflammatory Disease</i> chapter Ensure treatment is appropriate to results of clinical findings and Gram stain, wet mount, PH test and whiff test, see <i>Pelvic Inflammatory Disease</i> chapter

OF SPECIFIC SYNDROME

Table 1. Syndromic approach to the management of sexually transmitted infections (continued)

Syndrome	Signs and symptoms	Etiology	Specimens and testing	Microscopy results and clinical findings	Next steps/ special considerations
Vaginal discharge and low risk for STIs (for risk factors see <i>Primary</i> Care and Sexually Transmitted Infections chapter)	discharge discharge common: pH test and Gra and low risk for STIs (for risk factors see Primary Care and Sexually Transmitted Infections discharge common: pH test and Gra stain stain vaginosis vaginals vaginosis vaginals vaginosis vaginal vaginals richamon. Trichomoniasis vulvar	Vaginal swab for wet mount/amine	On examination: Watery white/ grey copious discharge On microscopy: Predominance of Gram-negative curved bacilli and coccobacilli and presence of clue cells, vaginal pH > 4.5, whiff test positive	Treat for bacterial vaginosis. See <i>Vaginal Discharge</i> chapter for recommendations	
				On examination: Clumpy white, curdy discharge On microscopy: Budding yeast, pseudohyphea and, if able to test, vaginal pH < 4.5, whiff test negative	Treat for candidiasis. See <i>Vaginal</i> <i>Discharge</i> chapter for recommendations
		On examination: Frothy white or yellow discharge On microscopy: Motile flagellated protozoa (tricho- monads) and, if able to test, vaginal pH >4.5, whiff test negative	Treat for trichomo- niasis.See Vaginal Discharge chapter for recommendations Treat sexual partner(s)		
				For low-risk individuals where no testing/microscopy is available or follow-up is not assured, treat according to clinical picture	

Table 1. Syndromic approach to the management of sexually transmitted infections (continued)

Syndrome	Signs and symptoms	Etiology	Specimens and testing	Microscopy results and clinical findings	Next steps/ special considerations
Vaginal discharge and high risk for STIs (for risk factors see <i>Primary</i> Care and Sexually Transmitted Infections chapter)	Vaginal discharge Vaginal odour Vaginal/ vulvar pruritis Vaginal/ vulvar erythema Dysuria	Most common: Bacterial vaginosis Vulvovaginal candidiasis Trichomo- niasis	As above, plus cervical swab for gonorrhea culture Cervical swab for <i>C trachomatis</i> (NAAT or culture) For women without a cervix, see <i>Gonococcal Infections</i> and <i>Chlamydial Infections</i> chapters for specimen collection recommendations	As above	As above For high-risk individuals where no testing/microscopy is available or follow-up is not assured, treat for bacterial vaginosis, Vulvovaginal candidiasis, trichomonas, chlamydia and gonorrhea

OF SPECIFIC SYNDROMES

Table 1. Syndromic approach to the management of sexually transmitted infections (continued)

mechons		7			
Syndrome	Signs and symptoms	Etiology	Specimens and testing	Microscopy results and clinical findings	Next steps/ special considerations
Intestinal syndromes: Proctitis Proctocolitis Enteritis	Varies according to specific syndrome: Mucopuru- lent rectal discharge Anorectal pain Constipation Bloody stools Diarrhea	ording according to specific to specific syndrome: opuru- rectal gonorrhoeae harge C trachomatis (LGV and non-LGV seripation dy Herpes	according to specific syndrome: N togonorrhoeae C trachomatis (LGV and non-LGV serovars) T pallidum Herpes Herpes T according lection should be adapted to the clinical presentation and patient history By anoscopic exam routinely obtain: Rectal swab for gonorrhea culture and chlamydia culture or NAAT (NAAT is not	On examination: Mucopurulent and/or bloody rectal discharge	Treat for gonorrhea and chlamydia as per the recommendations in the Sexually Transmitted Intestinal and Enteric Infections chapter If LGV is suspected, treat empirically as per the Lymphogranuloma Venereum chapter
	Diarrhea Nausea Abdominal pain/cramps Bloating Fever Salmonella spp. Shigella spp. Giardia lamblia	moeba specimens at this time) spylo- er spp. If chlamydia- positive: send for LGV serovar testing; see Lymphogranuloma Venereum chapter lif lesions are present:	On examination: Anal lesion	If syphilis is suspected and follow-up is not assured, treat empirically as per the <i>Syphilis</i> chapter If HSV is suspected, see <i>Genital Herpes Simplex Virus Infections</i> chapter to determine whether treatment is warranted	
		Syphilis serology Swab for herpes culture Stool for culture and ova and parasites	History and symptoms suggestive of enteric pathogens	See Sexually Transmit- ted Intestinal and Enteric Infections chapter for possible causative organisms	

Table 1. Syndromic approach to the management of sexually transmitted infections (continued)

Syndrome	Signs and symptoms	Etiology	Specimens and testing	Microscopy results and clinical findings	Next steps/ special considerations
Papular genital lesions	Growths in anal/genital region or on mucous membranes Multiple and/or polymorphic	Human papilloma- virus <i>Molluscum</i> <i>contagiosum</i> Skin tags	Visual examination and anal and/or vaginal exam as required by history/findings Pap testing if indicated as per local or provincial/territorial recommendations	Multiple or single cauliflower- like lesions (condyloma accuminata)	Treat as per the recommendations in the <i>Genital Human Papillomavirus Infections</i> chapter
	Asym-metrical	Carcinoma Normal variations		Internally anal/vaginal or cervical	Refer to a specialist for consultation and treatment
	inflammatory May be accompanied by: Pruritis Bleeding/obstruction, depending on location (i.e., urethra or vagina)			Flat, asymmetric lesions (condy- loma lata)	Sign of secondary syphilis; see <i>Syphilis</i> chapter for treatment recommendations
				Round, flat, umbilicated pap- ule (<i>Molluscum</i> contagiosum)	May heal spontane- ously with or without treatment. Can be treated with liquid nitrogen
				Symmetrical papular genital lesions Coronal sulcus (pearly penile papules) Vestibular papillae (micropapillomatis labialis)	Normal findings; no need for treatment
				Chronic lesion, ulceration or irregular pig- mentation (may be indicative of cancerous lesion)	Refer to a specialist for consultation and treatment

EPIDIDYMITIS

Definition

- Epididymitis can be defined as inflammation of the epididymis manifested by a relative acute onset of unilateral testicular pain and swelling often with tenderness of the epididymis and vas deferens and occasionally with erythema and edema of the overlying skin.
- The term epididymo-orchitis is primarily used when inflammation occurs in both the epididymis and the testes together.¹

Etiology²

- Before tests for detecting Chlamydia trachomatis were available, the cause of
 most cases of acute epididymitis was unknown. Subsequent studies have shown
 that epididymitis is primarily an infective condition.
- In men less than 35 years of age, sexually transmitted infection (STI) accounts for ²/₃ of epididymitis (47% *Chlamydia trachomatis* and 20% *Neisseria* gonorrhoeae). In men over 35 years of age, 75% of cases can be attributed to coliforms or pseudomonas. Isolation of *Chlamydia trachomatis* or *Neisseria* gonorrhoeae is unusual.
- The determination of the possible etiologic agent should always be based on the evaluation of the risk of the individual having acquired a sexually transmitted agent.
- In children and young adults, it is important to consider non-infectious causes
 of scrotal swelling, such as trauma, torsion of the testicle and tumour. Torsion of
 the testicle, which has a high risk of testicular infarction if treatment is delayed,
 is a surgical emergency and should be suspected when the onset of scrotal pain
 is sudden.

Table 1. Microbial etiology and predisposing factors in acute epididymitis3

Age group	Etiology and predisposing factors
Prepubertal children	 Usual etiology: coliforms, <i>P aeruginosa</i> Unusual etiology: hematogenous spread from primary infected site Predisposing factors: underlying genitourinary pathology
Men under 35	 Usual etiology: <i>C trachomatis, N gonorrhoeae</i> Unusual etiology: coliforms or <i>P aeruginosa, Mycobacterium tuberculosis</i> Predisposing factors: sexually transmitted urethritis
Men over 35	 Usual etiology: coliforms or <i>P aeruginosa</i> Unusual etiology: <i>N gonorrhoeae, C trachomatis, Mycobacterium tuberculosis</i> Predisposing factors: underlying structural pathology or chronic bacterial prostatitis

Epidemiology

- Accurate data on acute epididymitis are lacking. Therefore, the incidence of this
 condition in the general population is unknown. In a large retrospective study,
 49% of cases were in those 20–29 years old, with 70% of cases in those aged
 20–39 years.⁴
- In adolescents with epididymitis, sexual behaviour should be ascertained as the cause may be a sexually transmitted infection.
- Coliforms may be a frequent cause of acute epididymitis in sexually active men in all age groups who practice unprotected insertive anal intercourse.

Prevention

- At the time of diagnosis of a suspected sexually acquired epididymitis, safer sex practices should be reviewed.
- Appropriate information should be provided concerning the level of protection provided by barrier methods such as male condoms.
- Patient and contact(s) should abstain from unprotected intercourse until treatment of both patient and contact(s) is complete, or for 7 days in the case of single dose treatment of partners.

Manifestations5,6

- Patients with acute epididymitis usually present with unilateral testicular pain and tenderness.
- The onset of pain is generally gradual.
- In sexually transmitted epididymitis, symptoms of urethritis or a urethral discharge may be present. However, urethritis is often asymptomatic.
- Testicular torsion should be considered in all cases, as it is a surgical emergency. Torsion is more likely if onset of pain is sudden and the pain is severe. Torsion is more frequent in men less than 20 years of age, but can occur at any age.

Signs of acute epididymitis may include any of the following:

- Tenderness to palpation on the affected side.
- Palpable swelling of the epididymis.
- Urethral discharge.
- Hydrocele.
- Erythema and/or edema of the scrotum on the affected side.
- Fever.

Diagnosis5

- If diagnosis is questionable, a specialist should be consulted immediately because in the case of testicular torsion, testicular viability may be compromised.
- Evaluation for epididymitis should include the following:
 - Urethral swab for Gram stain.
 - Collection of specimens for identification of *N gonorrhoeae* and *C trachomatis* (intraurethral exudate or urine according to available laboratory techniques).
 - Microscopy and culture of mid-stream urine.
- If it can be arranged without delay, a Doppler ultrasound may be useful to help differentiate epididymitis from testicular torsion.
- There is no role for epididymal aspiration in routine clinical practice. It may be useful in recurrent infection that fails to respond to therapy or in patients with suspected abscess formation.

Management and Treatment

See Table 2, below, for published treatment recommendations for acute Epididymitis.

Table 2. Recommended regimens for the treatment of acute epididymitis⁵⁻¹⁰

Epididymitis most likely caused by chlamydial or gonococcal infections	Doxycycline 100 mg PO bid for 10–14 days [A-I] PLUS Ceftriaxone 250 mg IM in a single dose [A-I] OR Ciprofloxacin 500 mg PO in a single dose [A-I]
	(unless not recommended due to quinolone resistance*)
Epididymitis most likely caused by enteric organisms	Ofloxacin 200 mg PO bid for 14 days [A-I]

- * Quinolones are not recommended if the case or contact are from, or are epidemiologically linked to any area with rates of quinolone-resistant *N gonorrhoeae* > 3 5%
 - Asia
 - Pacific Islands (including Hawaii)
 - India
 - Israel
 - Australia
 - United Kingdom
 - Regions of the United States (check with the U.S. Centers for Disease Control and Prevention for rates of quinolone resistance by geographic area)
 - Men who have sex with men with contact or epidemiologically linked to the United States
 - Areas in Canada experiencing high rates of quinolone resistance; current numbers provided by the National Microbiology Laboratory place Quebec, Ontario, Alberta and British Columbia above the 3% threshold for quinolone resistance. Please check with your local public health officials to learn about quinolone resistance in your area. In Alberta all ciprofloxacin resistant cases in 2004–05 were in MSM or linked to travel outside of Alberta, therefore ciprofloxacin remains a recommended agent for the treatment of gonorrhea in Alberta except in these situations (source: Alberta Health and Wellness STD Services). For data on national quinolone resistance in Canada, please visit the Public Health Agency of Canada website (www.phac-aspc.gc.ca). For more information see *Gonococcal Infections* chapter.

Consideration for Other STIs

- Depending on sexual history, gonococcal and/or chlamydial infections should be considered as the etiology of acute epididymitis in all sexually active men with acute epididymitis, especially those under the age of 35.
- Consideration for testing for other STIs, including HIV, should be made according to the patient's sexual history and the presence of risk factors for specific infections.

Reporting and Partner Notification

- Patients with conditions that are reportable according to provincial and territorial laws and regulations should be reported to the local public health authority.
- Local public health authorities are available to assist with partner notification and help with appropriate referral for clinical evaluation, testing, treatment and health education.
- When treatment is indicated for the index case, and they are presumed to have sexually acquired epididymitis, all sexual partners within 60 days prior to symptom onset or date of diagnosis where asymptomatic should be clinically evaluated and treated with an appropriate regimen.

Follow-up

Follow-up should be arranged to evaluate the response to treatment. If a
recommended regimen has been given and correctly taken, symptoms and
signs have disappeared and there is no re-exposure to an untreated sexual
partner, then repeat diagnostic testing for N gonorrhoeae and C trachomatis
is not routinely recommended.

Special Considerations

- Rare causes of clinical sterile acute epididymitis include amiodarone therapy, vasculitis, polyarteritis nodosa, Behçet disease and Henoch-Schönlein purpura and a proportion of cases remains idiopathic.
- A condition described as "chronic epididymitis" has been recently characterized
 in the literature. 11 Although defined by the author as the presence of "symptoms
 of discomfort and/or pain at least 3 months in duration in the scrotum, testicle
 or epididymis localized to one or each epididymis on clinical examination," there
 is no clear natural history of the condition. The authors conclude that further
 studies on the epidemiology, etiology and pathogenesis of this condition are
 needed.

References

- Hagley M. Epididymo-orchitis and epididymitis: a review of causes and management of unusual forms. *Int J STD AIDS* 2003;14:372–378.
- 2. Luzzi GA, O'Brien TS. Acute epididymitis. BJU Int 2001;87:747–755.
- 3. Berger E. Acute epididymitis. In: Holmes KK, Sparling PF, Mardh PA, et al, eds. Sexually Transmitted Diseases. 3rd ed. New York, NY: McGraw Hill; 1999: 847–858.
- 4. Mittemeyer BT, Lennox KW, Borski AA. Epididymitis; a review of 610 cases. *J Urol* 1966;95:390–392.
- 5. Sexually transmitted diseases treatment guidelines 2002. Centers for Disease Control and Prevention. *MMWR Recomm Rep* 2002;51(RR-6):1–78.
- 6. UK national guidelines on sexually transmitted infections and closely related conditions. Introduction. *Sex Transm Infect* 1999;75(suppl 1):S2–3.
- 7. Epididymitis in youth and adults. In: *Canadian STD Guidelines 1998 Edition*. Ottawa, ON: Health Canada; 1998: 100–102.
- 8. Hoosen AA, O'Farrell N, van den Ende J. Microbiology of acute epididymitis in a developing community. *Genitourin Med* 1993;69:361–363.
- 9. Melekos MD, Asbach HW. Epididymitis: aspects concerning etiology and treatment. *J Urol* 1987;138:83–86.
- 10. Weidner W, Schiefer HG, Garbe C. Acute nongonococcal epididymitis. Aetiological and therapeutic aspects. *Drugs* 1987;34(suppl 1):111–117.
- Nickel JC, Siemens DR, Nickel KR, Downey J. The patient with chronic epididymitis: characterization of an enigmatic syndrome. *J Urol* 2002; 167:1701–1704.

GENITAL ULCER DISEASE (GUD)

Etiology

Definition

 Ulcerative, erosive, pustular or vesicular genital lesion(s), with or without regional lymphadenopathy, caused by a number of sexually transmitted infections (STIs) and non-STI-related conditions.

STIs

- For most young, sexually active patients with genital ulcer disease (GUD), etiology is related to an STI. Most often it is due to herpes simplex virus type 1 or 2 (HSV-1 or HSV-2), causing genital herpes.¹ More than one etiology may be found if a careful evaluation is conducted.² Other STI causes of GUD are as follows:
 - Treponema pallidum spp., causing primary syphilis.
 - Haemophilus ducreyi, causing chancroid.
 - *Chlamydia trachomatis* serotype L1, 2 or 3, causing lymphogranuloma venereum (LGV).
 - Klebsiella granulomatis, causing granuloma inquinale (donovanosis).

Non-STI-related infections or conditions

- Non-STI-related infections or conditions causing GUD may also be seen (see Differential diagnosis, below).
- Even after a complete diagnostic evaluation, at least 25% of patients with GUD have no laboratory-confirmed diagnosis.³

Epidemiology

- The cause of GUD can be related to a number of factors, such as geographical
 area where sexual intercourse has taken place; socioeconomic factors; gender
 of sexual partners; number of partners; HIV status and local prevalence; drug
 use; commercial sex; and circumcision.⁴
- GUD constitutes at most 5% of visits to physicians for a possible STI.5
- About 70–80% of genital ulcers are due to HSV-1 or HSV-2.
- Genital ulcers in sexually active persons can be associated with two or more pathogens.²
- Women and men with GUD are at increased risk of acquiring and transmitting HIV.⁶
- Syphilis and LGV are rare causes of GUD in Canada, but should be considered
 in persons having sex while travelling to endemic areas or among men who
 have sex with men (MSM). When identified, the potential for a localized discrete
 outbreak exists. Rarely, granuloma inguinale and chancroid should also be
 considered.

- Syphilis incidence is increasing in Canada, with regional outbreaks of infectious syphilis occurring in recent years, including Vancouver, the Yukon, Calgary, Edmonton, Toronto, Ottawa, Montreal and Halifax.⁷⁻⁹
- Chancroid has been sporadically associated with focal urban epidemics in North America, particularly among cocaine users. Sex workers are the usual reservoir.
- Rectal LGV outbreaks are now occurring among MSM in Europe, with recent reports of cases in North America. Co-infection with HIV and hepatitis C virus are seen at a high rate, 10-11 including in Canada. 12
- HIV infection increases the transmission of STI genital ulcers, and the reverse is also true.¹³

Risk factors

- The following are risk factors for STI-related GUD:¹⁴
 - Sexual contact with:
 - MSM.
 - A person with GUD.
 - A new partner.
 - A partner who is from or has travelled to an endemic area.
 - Sex workers and their clients.
 - An anonymous sexual contact (e.g., from the Internet, bathhouse, rave/circuit party).
 - A partner or index case who is HIV-positive.
 - Travel to endemic areas.
 - Living in region(s) in Canada experiencing outbreaks (e.g., syphilis).
 - Previous genital lesions or STI.
 - Drug use by self and/or partner.

Prevention

- Sexual activity of any mucosal type oral, anal or genital can be associated
 with sexually transmitted ulcers. Patients presenting with concerns about
 STIs and/or birth control should be given information on the efficacy of barrier
 methods in preventing STI/HIV transmission and provided safer-sex counselling
 (see Primary Care and Sexually Transmitted Infections chapter).
- Identify barriers to prevention practices and the means to overcome them (see *Primary Care and Sexually Transmitted Infections* chapter).
- In the case of bacterial GUD caused by an STI, patients and contacts should abstain from unprotected intercourse until treatment of both partners is complete.
 For genital herpes, see Genital Herpes Simplex Virus Infections chapter.

Manifestations

- Diagnosis is often inadequate when based solely on history and physical examination, because of the lack of sensitivity and specificity of lesion(s), even in so-called "classic" cases.³
- Concurrent infection with HIV can change the clinical features of genital ulcers; the therapeutic regimen may also be different.

Table 1. Manifestations

Table 1. Manifestations			
STI	Site	Appearance	Other signs/ symptoms
Herpes simplex virus ¹⁵	 For both sexes, anywhere in the "boxer short" area Men: glans, prepuce, penile shaft, anus, rectum (for MSM) Women: cervix, vulva, vagina, perineum, legs and buttocks 	 Grouped vesicles evolving toward superficial circular ulcers on an erythematous base Smooth margin and base Enlarged, nonfluctuant and tender inguinal lymph nodes most common in primary infection 	 Ulcers usually painful and/or pruritic Genital pain Constitutional symptoms, such as fever, malaise and pharyngitis, are common with primary infection
Primary syphilis (see <i>Syphilis</i> chapter)	At site of inoculation, although most individuals with syphilis fail to notice primary chancre ¹⁶	 Papule evolving to a painless chancre Indurated with serous exudates Single ulcer in 70% of cases Smooth margin and base 	Firm, enlarged, non-fluctuant, non-tender lymphadeno- pathy is common
Chancroid	At site of inoculation	 Single or multiple necrotizing and painful ulcers Two or more in 50% of cases 	Often painful swelling and suppuration of regional lymph nodes, with erythema and edema of overlying skin

Table 1. Manifestations (continued)

STI	Site	Appearance	Other signs/ symptoms
Lymphogranuloma venereum ¹⁷	At site of inoculation	 Self-limited single painless papule, which may ulcerate, followed some weeks later by tender inguinal and/or femoral lymphadenopathy, mostly unilateral, and/or proctocolitis. Recent outbreaks in MSM have been characterized primarily by proctocolitis If not treated, fibrosis can lead to fistulas and strictures and/or obstruction of the lymphatic drainage, causing elephantiasis 	• Signs/ symptoms of urethritis
Granuloma inguinale	At site of inoculation	 Single or multiple progressive ulcerative lesions Highly vascular (beefy red appearance) Bleeds easily on contact Two or more in 50% of cases Hypertrophic, necrotic and sclerotic variants Relapse can occur 6–18 months after apparently effective therapy 	• Painless

MSM = men who have sex with men

Diagnosis

Table 2. Diagnostic features of STI-related GUD

Disease	% of STI-related GUD	Incubation period
Herpes (recurrent genital herpes more frequent than primary genital herpes)	95%	2–7 days for primary genital herpes
Primary syphilis	>1%	3–90 days
Chancroid	<1%	5–14 days
Lymphogranuloma venereum	<1%	3–30 days
Granuloma inguinale	<1%	1–180 days

GUD = genital ulcer disease STI = sexually transmitted infection

Differential diagnosis

Table 3. Infectious, non-STI-related causes of genital ulcers18

Fungal	Viral	Bacterial
CandidaDeep fungi (rare)	 Cytomegalovirus (rare) Varicella or herpes zoster virus (rare) Epstein-Barr virus (rare) 	 Staphylococcus spp. Streptococcus spp. Salmonella spp. Pseudomonas spp. Mycobacteria Parasite (e.g., scabies)

Table 4. Non-infectious skin and mucosal conditions and diseases¹⁹

Bullous dermatoses	Non-bullous dermatoses	Malignancy
 Non-autoimmune Contact dermatitis Erythema multiforme (almost always HSV-related) Toxic epidermolysis Auto-immune Pemphigus Cicatricial pemphigoid 	 Non-specific vulvitis/balanitis Aphthae or aphthous ulcers, aphthosis Lichen planus, erosive lichen planus Lichen sclerosus Behçet disease Pyoderma gangrenosum Fixed drug eruption Lupus erythematosus Crohn's disease Vasculitis 	Squamous-cell carcinoma Vulvar intraepithelial neoplasia Less common: Extramammary Paget's disease Basal-cell carcinoma Lymphoma/leukemia Histiocytosis X

HSV = herpes simplex virus

- Other causes of ulcerative lesions of the skin and mucosa:
 - Trauma (less common)
 - Idiopathic: 12–51% of genital ulcers have no definite cause in research settings. Referral to an expert when no etiology is found may diminish this fraction.⁴

Specimen collection and laboratory diagnosis

- The minimum testing for all cases of GUD should include a viral identification test for HSV and a syphilis serology.
- Inform laboratory in advance when special procedures need to be followed. Consultation with an experienced colleague may be warranted.
- Biopsies, cultures, smears, and serology should be ordered as appropriate for evaluation of all vulvar ulcers.

Herpes simplex virus

- See Genital Herpes Simplex Virus Infections chapter.
- Herpes testing is important for all lesions, initial and recurrent, even in classic
 cases, because of false-positive clinical diagnosis. Retesting following a positive
 test is almost always of limited value. Typing is important to aid in the discussion
 of the natural history, help assess partners and help discuss preventative
 agendas.

Viral identification

- Viral identification by either viral culture or nucleic acid amplification test (NAAT), or, if not available, by antigen test.
- Culture should be carried out on at least three unroofed pustules/vesicles
 or wet ulcers unless HSV infection has been previously confirmed by a
 laboratory test. The specimen must be transported in a special viral transport
 medium.
- NAATs are considered superior, but their availability is limited (see Laboratory Diagnosis of Sexually Transmitted Infections chapter).
- Type-specific serology
 - In the presence of a potential case of genital herpes and two negative viral identification tests, or if there is difficulty organizing testing when lesions are present or lesions are rare, type-specific serology can help confirm possible genital herpes cases.²⁰ If both HSV-1 and HSV-2 serology are negative 12 weeks after the first manifestation, genital herpes is not likely.
 - It should be noted that the availability of type-specific serology is limited in Canada.

T pallidum

- · See Syphilis chapter.
- Identification: dark-field examination or direct fluorescent antibody test on swab from ulcers. Contact your local laboratory regarding these tests, as they are not widely available.
- Serology
 - Syphilis serology should include a non-treponemal test (e.g., rapid plasma reagin [RPR], Venereal Disease Research Laboratory [VDRL]) or treponemalspecific enzyme immunoassay (EIA). As treponemal tests are far more sensitive in primary syphilis than non-treponemal tests, many authorities advocate proceeding directly to treponemal tests when primary syphilis is suspected.
 - If non-treponemal syphilis serology is found, positive confirmation by treponemal-specific test (e.g., *Treponema pallidum* particle agglutination [TP-PA], microhemagglutination for *Treponema pallidum* [MHA-TP] or fluorescent treponemal antibody absorption [FTA-ABS]) should be sought if not already ordered (see *Syphilis* chapter).
 - Serologic tests should be repeated 2–4 weeks after the original negative test if syphilis is a possibility.
 - Dark-field examination or fluorescent antibody for T pallidum of lesions, if available.

Other causes

 If history, risk factors and physical findings warrant testing for other less common causes of GUD, special laboratory tests may be needed to properly assess the etiology of ulcerative disease. Consider testing for chancroid, LGV and granuloma inguinale.

- H ducreyi (chancroid)
 - See Chancroid chapter.
 - Bacterial culture on specific culture medium (special arrangement to be made in advance).
 - NAAT where available (e.g., polymerase chain reaction [PCR]).
 - Gram stain may also be useful (see *Laboratory Diagnosis of Sexually Transmitted Infections* chapter).
- C trachomatis serovar L1, L2 or L3 (LGV)
 - See Lymphogranuloma Venereum chapter
 - Identification of C trachomatis by culture, NAAT or serology, followed by confirmation of LGV serovars through DNA sequencing or restriction fragment length polymorphism (RFLP).
- Klebsiella granulomatis (granuloma inguinale)
 - Identification of dark-staining Donovan bodies on crushed or biopsy specimen.

Caution

- Except for genital herpes, most Canadian clinicians have limited experience with STI-related genital ulcers. Early referral to a colleague experienced in this area should be considered, particularly if the case involves the following:
 - Travel.
 - MSM.
 - HIV-infected individuals.
 - Immunocompromised patients.
 - Systemic disease.
- Atypical and/or non-healing lesions may require a biopsy and should be referred to a colleague experienced in this area.²¹

Management²²

If test results are not yet available

- · Treatment considerations:
 - Empiric treatment for chancroid, LGV and syphilis should be discussed with a local expert or public health official only if follow-up is uncertain and if risk factors for these diseases are present.
 - Treatment at the time of presentation should be considered for genital herpes for almost all cases of GUD, especially if the symptoms are typical.
- See Chancroid, Lymphogranuloma Venereum and Syphilis chapters for more information.

If results are available for RPR, VDRL, TP-PA, MHA-TP/dark-field examination/fluorescent antibody test

Positive (motile corkscrew spirochetes present): treat for syphilis (see Syphilis chapter).

- Dark-field examinations, fluorescent antibody tests and tests for HSV infection and *H ducreyi* are negative or not performed: treat as syphilis if there is a recent history of contact with infectious syphilis or clinical suspicion is strong and follow-up cannot be ensured.
- · Otherwise:
 - Consider therapy for HSV if laboratory tests are negative and presentation is typical of HSV infection (see *Genital Herpes Simplex Virus Infections* chapter).
 - Treat for chancroid if presentation suggests chancroid (see Chancroid chapter).

Treatment²³

- For treatment recommendations for syphilis, HSV, chancroid and LGV, see appropriate chapters.
- Treatment of ulcerative STIs in HIV co-infected patients may represent a challenge.²⁴ See relevant chapters on treatment of specific infections, or, if not experienced in this area, consult an experienced colleague.

Granuloma inguinale^{3,25-29}

- · Preferred:
 - Doxycycline 100 mg P0 bid for 21 days (based on studies of older preparations of tetracyclines) [C-III].
 - Trimethoprim-sulfamethoxazole double strength PO bid for 21 days [C-III].
- · Alternatives:
 - Ciprofloxacin 750 mg PO bid for 21 days [C-III].
 - Erythromycin 500 mg qid for 21 days [C-III].
 - Azithromycin 500 mg daily or 1 g weekly for a minimum of 21 days [C-III].

Consideration for Other STIs

- See Primary Care and Sexually Transmitted Infections chapter.
- Obtain specimen(s) for the diagnosis of chlamydial and gonococcal infections and other STIs when appropriate (including LGV, chancroid and granuloma inquinale if there has been travel to regions where these infections are endemic).
- HIV testing and counselling are recommended (see Human Immunodeficiency Virus Infections chapter). Patients with syphilis, LGV and chancroid are at especially high risk for concurrent HIV infection.³ Timing of HIV testing is important, as genital ulceration is a marker for HIV risk. Baseline testing at the initial visit and repeat HIV testing in 12 weeks should be considered.
- Immunization against hepatitis B in those with no immunity against this virus is also recommended (see *Hepatitis B Virus Infections* chapter).

Reporting and Partner Notification

- Conditions that are reportable according to provincial and territorial laws and regulations must be reported to the local public health authority (see chapters of specific infections for reporting requirements).
- Partner notification is vitally important for the rare bacterial ulcerative conditions discussed in this section in order to prevent an outbreak.
- When treatment is indicated for a diagnosis of syphilis, chancroid, LGV or granuloma inguinale, all partners who have had sexual contact with the index case should be located, clinically evaluated and treated appropriately.³ For more information on partner notification and treatment by infection, see *Chancroid*, *Lymphogranuloma Venereum* and *Syphilis* chapters.
- Local public health authorities are available to assist with partner notification and appropriate referral for clinical evaluation, testing, treatment and health education.

Follow-up

- A follow-up visit should be arranged for re-evaluation.
 - For chancroid and granuloma inguinale, if the patient is compliant with the prescribed treatment, symptoms resolve and there is no risk of reexposure to an untreated partner, repeat diagnostic testing is not routinely recommended.
 - For LGV, see Lymphogranuloma Venereum chapter.
 - For genital HSV infection, no test of cure is necessary.
 - For syphilis, see Syphilis chapter.
- Timing for HIV testing should be considered at this stage. Most patients
 presenting with an acute genital ulcer will be too early in the window to have
 reactive serology related to an HIV infection.

Special Considerations

Children

- Sexual abuse must be considered when GUD is found in children beyond the
 neonatal period. Consultation with a colleague experienced in such cases
 should be sought (see Sexual Abuse in Peripubertal and Prepubertal Children
 chapter).
- Reporting sexual abuse:
 Sexual abuse of children must be reported to the local child protection agency.
 Local public health authorities may be helpful in evaluating both the source of the infection and potential transmission in the community.
- Whenever possible, it is strongly recommended that the child be evaluated at or in conjunction with a referral centre (see Appendix F and G).

References

- Mertz KJ, Trees D, Levine WC, et al. Etiology of genital ulcers and prevalence of human immunodeficiency virus infection in 10 US cities. The Genital Ulcer Disease Surveillance Group. J Infect Dis 1998;178:1795–1798.
- DiCarlo RP, Martin DH. The clinical diagnosis of genital ulcer disease in men. Clin Infect Dis 1997;25:292–298.
- 3. Sexually transmitted disease treatment guidelines 2002. Centers for Disease Control and Prevention. *MMWR Morb Mortal Wkly Rep* 2002;51(RR-6):1–78.
- Ballard R. Genital ulcer adenopathy syndrome. In: Holmes KK, Sparling PF, Mardh PA, et al, eds. Sexually Transmitted Diseases. 3rd ed. New York, NY: McGraw Hill; 1999: 887–892.
- Piot P, Meheus A. Genital ulcerations. In: Taylor-Robinson D, ed. *Clinical Problems in Sexually Transmitted Diseases*. Boston, MA: Martinus Nyhoff; 1985:207.
- 6. Celum CL. The interaction between herpes simplex virus and human immunodeficiency virus. *Herpes* 2004;11(suppl 1):36A-45A.
- Sexual Health and Sexually Transmitted Infections Section, Centre for Infectious Disease Prevention and Control, Public Health Agency of Canada. Reported cases and rates of notifiable STI from January 1 to June 30, 2004, and January 1 to June 30, 2003. Ottawa, ON: Public Health Agency of Canada; 2004. Available at: www.phac-aspc.gc.ca/std-mts/stdcases-casmts/index. html. Accessed January 18, 2005.
- Sarwal S, Shahin R, Ackery J-A, Wong T. Infectious syphilis in MSM, Toronto, 2002: outbreak investigation. Paper presented at: Annual Meeting of the International Society for STD Research; July 2003; Ottawa, ON. Abstract 0686.
- Shahin R, Sarwal S, Ackery J-A, Wong T. Infectious syphilis in MSM, Toronto, 2002: public health interventions. Paper presented at: Annual Meeting of the International Society for STD Research; July 2003; Ottawa, ON. Abstract 0685.
- Nieuwenhuis RF, Ossewaarde JM, Gotz HM, et al. Resurgence of lymphogranuloma venereum in Western Europe: an outbreak of Chlamydia trachomatis serovar L2 proctitis in the Netherlands among men who have sex with men. Clin Infect Dis 2004;39:996–1003.
- 11. Centers for Disease Control and Prevention. Lymphogranuloma venereum among men who have sex with men Netherlands, 2003–2004. *MMWR Morb Mortal Wkly Rep* 2004;53:985–988.
- 12. Kropp RY, Wong T, the Canadian LGV Working Group. Emergence of lymphogranuloma venereum in Canada. *CMAJ* 2005;172:1674–1676.
- 13. Wasserheit JN. Epidemiological synergy. Interrelationships between human immunodeficiency virus infection and other sexually transmitted diseases. Sex Transm Dis 1992;19:61–77.
- 14. Agence de développement de réseaux locaux de services de santé et de services sociaux. Direction de santé publique. Campagne provinciale de prévention de la syphilis "Je suis Phil". 1. La syphilis, état de situation et caractéristiques. Quebec, QC: Direction de santé publique; 2004.

- 15. Corey L, Holmes KK. Clinical course of genital herpes simplex virus infections: current concepts in diagnosis, therapy, and prevention. *Ann Intern Med* 1983;48:973–983.
- 16. Singh AE, Romanowski B. Syphilis: review with emphasis on clinical, epidemiologic and some biologic features. *Clin Microbiol Rev* 1999;12:187–209.
- 17. Mabey D, Peeling RW. Lymphogranuloma venereum. *Sex Transm Infect* 2002;78:90–92.
- 18. Leibowitch M, Staughton R, Neill S, Barton S, Marwood R. *An Atlas of Vulval Disease: A Combined Dermatological, Gynaecological and Venereological Approach.* London: Martin Dunitz; 1995.
- Lynch PJ, Edwards L. Genital Dermatology. Oxford: Churchill Livingstone; 1994.
- Wald A, Ashley-Morrow R. Serological testing for herpes simplex virus (HSV)-1 and HSV-2 infection. Clin Infect Dis 2002;35(suppl 2):S173–S182.
- 21. Black MM, McKay M, Braude P. *Obstetric and Gynecologic Dermatology*. London: Mosby-Wolfe; 1995.
- 22. Canadian STD Guidelines 1998 edition. Ottawa ON: Health Canada; 1998.
- 23. *Guidelines for the Management of Sexually Transmitted Infections*. Geneva, Switzerland: World Health Organization; 2001.
- 24. Wu JJ, Huang DB, Pang KR, Tyring SK. Selected sexually transmitted diseases and their relationship to HIV. *Clin Dermatol* 2004;22:499–508.
- 25. Association for Genitourinary Medicine and the Medical Society for the Study of Venereal Diseases, Clinical Effectiveness Group. 2001 National Guideline for the Management of Donovanosis (Granuloma Inguinale). British Association for Sexual Health and HIV website. Available at: www.bashh.org/guidelines/2002/donovanosis_0901b.pdf. Accessed September 22, 2005.
- 26. Greenblatt RB, Barfield WE, Dienst RB, West RM. Terramycin in the treatment of granuloma inguinale. *J Vener Dis* Inf 1951;32:113–115.
- Lal S, Garg BR. Further evidence of the efficacy of co-trimoxazole in the donovanosis. Br J Vener Dis 1980;56:412–413.
- 28. Robinson HM, Cohen MM. Treatment of granuloma inguinale with erythromycin. *J Invest Dermatol* 1953;20:407–409.
- 29. Bowden FJ, Mein J, Plunkett C, Bastian I. Pilot study of azithromycin in the treatment of genital donovanosis. *Genitourin Med* 1996;72:17–19.

PELVIC INFLAMMATORY DISEASE (PID)

Etiology

- There are multiple causes of lower abdominal pain in women, including gynecologic disease or dysfunction (complications of pregnancy, acute infections, endometriosis, adnexal disorders, menstrual disorders), as well as gastrointestinal (appendicitis, gastroenteritis, inflammatory bowel disease), genitourinary (cystitis, pyelonephritis, nephrolithiasis), musculoskeletal and neurologic causes.
- The most common infectious cause of lower abdominal pain in women is pelvic inflammatory disease (PID).¹
- PID is a polymicrobial infection with multiple microbial etiologies.
- Most cases of PID are associated with more than one organism.
- Pathogens can be categorized as sexually transmitted or endogenous organisms.

Table 1. Microbial causes

Sexually transmitted organisms	 Chlamydia trachomatis Neisseria gonorrhoeae Viruses and protozoa (rare) Herpes simplex virus Trichomonas vaginalis
Endogenous organisms	 Genital-tract mycoplasmas Mycoplasma genitalium Mycoplasma hominis Ureaplasma urealyticum
Anaerobic bacteria	 Bacteroides spp. Peptostreptococcus spp. Prevotella spp.
Facultative (aerobic) bacteria	Escherichia coliGardnerella vaginalisHaemophilus influenzaeStreptococcus spp.

Definition

 PID is an infection of the female upper genital tract involving any combination of the endometrium, fallopian tubes, pelvic peritoneum and contiguous structures.

Epidemiology

- · PID is a very significant public health problem.
- Up to ²/₃ of cases go unrecognized, and underreporting is common.
- There are approximately 100,000 cases of symptomatic PID annually in Canada, although PID is not nationally reportable, so exact numbers are unknown.
- It is estimated that 10–15% of women of reproductive age have had one episode of PID.²
- In recent years, hospitalization rates for PID have declined (118/100,000 women in 1995 and 55/100,000 women in 2001, data from Health Canada) because increasing numbers of patients are treated as outpatients, but the number of patient visits to physician offices for PID has remained stable.
- The incidence of long-term sequelae of PID (tubal factor infertility, ectopic pregnancy, chronic pelvic pain) is directly related to the number of episodes of PID.³
- In jurisdictions with long-standing chlamydia control programs, PID rates and ectopic pregnancy rates have declined.

Prevention

- At the community level, health promotion and education programs are essential to promote screening for sexually transmitted infections (STIs).
- Health care providers must assume responsibility for primary prevention activities, such as risk-reduction counselling and patient education.
- At the time of diagnosis of infection, health care providers should reinforce prevention and safer-sex practices. They should also identify barriers to prevention practices and ways to overcome them.
- Patients and contacts must be counselled to abstain from unprotected sexual contact until treatment of both partners is complete.

Manifestations and Diagnosis

- Abdominal pain may be a clinical feature of many disorders, and the symptoms
 of PID may overlap with other gynecologic disorders or disorders of the
 gastrointestinal, urinary and musculoskeletal systems.
- There is no single historical, physical or laboratory finding that is both sensitive and specific for a diagnosis of PID.⁴
- Only 1/3 of women with acute PID have a temperature above 38°C.5
- Common findings on physical examination of patients with acute PID include bilateral lower abdominal, uterine, adnexal and cervical motion tenderness, but these findings may be present with a variety of other conditions as well.
- The clinical diagnosis of PID is imprecise, and clinicians must have a high index of suspicion.

Table 2. Criteria for diagnosis

Minimum diagnostic criteria	Additional diagnostic criteria	Definitive diagnostic criteria
 Lower abdominal tenderness Adnexal tenderness Cervical motion tenderness 	 Oral temperature >38.3°C Presence of white blood cells on saline microscopy of vaginal secretions/wet mount Elevated erythrocyte sedimentation rate Elevated C-reactive protein Laboratory documentation of cervical infection with Neisseria gonorrhoeae or Chlamydia trachomatis 	 Endometrial biopsy with histopathologic evidence of endometritis (at least 1 plasma cell per x120 field and at least 5 neutrophils per x400 field) Transvaginal sonography or other imaging techniques showing thickened fluid-filled tubes, with or without free pelvic fluid or tubo-ovarian complex Gold standard: Laparoscopy demonstrating abnormalities consistent with PID, such as fallopian tube erythema and/or mucopurulent exudates

PID = pelvic inflammatory disease

Physical examination and specimen collection

- A complete abdominal and pelvic examination should be performed in any patient with lower abdominal pain.
- Pelvic examination should include speculum and bimanual examinations.
- The external genital area, vagina and cervix must all be inspected.
- Stat serum beta HCG to rule out ectopic pregnancy.
- With the aid of a speculum, endocervical swabs should be obtained for diagnostic tests for Neisseria gonorrhoeae and Chlamydia trachomatis.
- Cervical lesions should be sampled with swabs for diagnostic tests for herpes simplex virus, if suspected.
- Vaginal swabs should be obtained for culture; pH testing; amine odour whiff
 testing; normal saline and potassium hydroxide wet preparations; and Gram
 stain. Clinical assessment for bacterial vaginosis includes three of four Amsel
 criteria (vaginal discharge, elevated pH, amine odour whiff test and clue cells on
 microscopy).⁶ An aerobic and anaerobic culture may assist with the detection of
 unusual vaginal pathogens, such as Group A streptococcus.

Laboratory diagnosis

- Negative laboratory results do not rule out a diagnosis of PID.
- A normal ultrasound study does not rule out a diagnosis of PID.
- Ultrasound may aid in the diagnosis, especially if tubo-ovarian abscess is suspected.

- A pregnancy test can be helpful to exclude ectopic pregnancy from the differential diagnosis.
- Detection of Gram-negative intracellular diplococci on a stained smear of endocervical secretions; positive results of a diagnostic test for N gonorrhoeae or C trachomatis; or both.
- Detection of *N gonorrhoeae* or *C trachomatis* may be enhanced by using nucleic acid amplification tests, such as ligase chain reaction or polymerase chain reaction.
- Other tests that may be helpful in the diagnosis of acute PID include complete blood count, erythrocyte sedimentation rate, C-reactive protein and endometrial biopsy.

Management

- · Early diagnosis and treatment are crucial to the maintenance of fertility.
- Antibiotic therapy can be administered orally or parenterally, and in inpatient or outpatient settings.
- Data suggest that efficacy and long-term complication rates are not significantly different between parenteral and oral therapy or inpatient and outpatient treatment.⁷
- Individuals treated as outpatients need careful follow-up and should be re-evaluated 2–3 days after therapy is initiated.
- If no clinical improvement has occurred, hospital admission for parenteral therapy, observation and consideration for laparoscopy is required; consultation with colleagues experienced in the care of these patients should be considered.

Table 3. Criteria for hospitalization

- Surgical emergencies such as appendicitis cannot be excluded.
- The patient is pregnant.
- The patient does not respond clinically to oral antimicrobial therapy.
- The patient is unable to follow or tolerate an outpatient oral regimen.
- The patient has severe illness, nausea and vomiting, or high fever.
- The patient has a tubo-ovarian abscess.

Consider hospitalization for observed oral or parenteral therapy in the following cases:

- HIV infection
- Youth/adolescents (particularly if compliance is an issue)

Treatment

- Goals of treatment are to control the acute infection and to prevent long-term sequelae such as infertility, ectopic pregnancy and chronic pelvic pain.
- Treatment regimens must provide empiric broad-spectrum coverage of likely etiologic pathogens and take into account the polymicrobial nature of PID.
- Treatment regimens must provide coverage for N gonorrhoeae, C trachomatis, anaerobic bacteria, Gram-negative facultative bacteria and streptococci.
- Discontinuation of parenteral therapy may be considered 24 hours after a patient improves clinically.⁸
- Oral step-down therapy should then begin and continue for a total of 14 days of treatment.⁸
- If recovery does not occur, other differential diagnoses must be entertained and a laparoscopy considered.

Table 4. Recommended parenteral treatment regimens

D 00 [4 1]	
Regimen A ^o [A-I]	 Cefotetan 2 g IV every 12 hours PLUS doxycycline 100 mg IV or PO every 12 hours Cefoxitin 2 g IV every 6 hours PLUS doxycycline 100 mg IV or PO every 12 hours Parenteral therapy may be discontinued 24 hours after a patient improves clinically, and oral therapy with doxycycline (100 mg bid) should continue for a total of 14 days Most authorities recommend administering doxycycline in oral form even in hospitalized patients, because IV administration is painful and more costly, and because oral and IV administration provide similar bioavailability
Regimen B [A-I]	 Clindamycin 900 mg IV every 8 hours PLUS Gentamicin* loading dose IV or IM (2 mg/kg of body weight), followed by a maintenance dose (1.5 mg/kg) every 8 hours. Once-daily dosing may be substituted (5mg/kg of body weight IV every 24 hours) Parenteral therapy may be discontinued 24 hours after a patient improves clinically, and oral therapy with doxycycline (100 mg bid) or clindamycin (450 mg PO qid) should continue for a total of 14 days

Table 4. Recommended parenteral treatment regimens (continued)

Alternative regimens¹⁰ [A-II]

- Ofloxacin 400 mg IV every 12 hours \pm metronidazole 500 mg IV every 8 hours

OR

- Levofloxacin 500 mg IV once daily \pm metronidazole 500 mg IV every 8 hours

OR

 Ampicillin/sulbactam 3 g IV every 6 hours PLUS doxycycline 100 mg IV or PO every 12 hours

OR

- Ciprofloxacin 200 mg IV every 12 hours PLUS doxycycline 100 mg IV or PO every 12 hours PLUS metronidazole 500 mg IV every 8 hours
 - Because ciprofloxacin has poor coverage against C trachomatis, it is recommended that doxycycline be added routinely
 - Because of concerns regarding the anaerobic coverage of both quinolones, metronidazole should be included with each regimen

Notes

The use of ofloxacin, ciprofloxacin, levofloxacin, and doxycycline is contraindicated for pregnant and lactating women. Pregnant women should not be treated with quinolones or tetracyclines.

Table 5. Recommended outpatient treatment regimens

Regimen A ¹¹ [A-II]	 Ofloxacin 400 mg P0 bid for 14 days ± metronidazole 500 mg P0 bid for 14 days [A-I] OR Levofloxacin 500 mg P0 qd ± metronidazole 500 mg P0 bid for 14 days [B-II] Metronidazole is added to provide anaerobic coverage Preliminary data suggest that oral levofloxacin is as effective as oral ofloxacin, with the advantage of once-daily dosing⁹
Regimen B ¹² [A-II]	 Ceftriaxone 250 mg IM qd PLUS doxycycline 100 mg PO bid for 14 days OR Cefoxitin 2 g IM PLUS probenecid 1 g PO in a single dose concurrently once PLUS doxycycline 100 mg PO bid for 14 days OR Other parenteral third-generation cephalosporin (e.g., ceftizoxime or cefotaxime) PLUS doxycycline 100 mg PO bid for 14 days Many authorities recommend the addition of metronidazole 500 mg PO bid for 14 days to this regimen for additional anaerobic coverage and the treatment of bacterial vaginosis [B-III]

^{*} These recommendations apply for those patients with normal renal function; gentamicin dosage should be adjusted in cases of renal impairment. Renal function and gentamicin levels should be monitored during treatment.

Consideration for Other STIs

- Individuals infected with one STI are at risk of concurrent infection with one or more other STIs.
- Following a diagnosis of PID, testing and counselling should be performed for other infections, including HIV and syphilis.
- Immunization against hepatitis B is recommended if not already immune.

Reporting and Partner Notification

- Patients with conditions that are notifiable according to provincial and territorial laws and regulations should be reported to local public health authorities.
- The management of women with PID is considered inadequate unless their sexual partners are also evaluated and treated.
- Evaluation should occur if there was sexual contact with the patient during the 60 days prior to symptom onset or date of diagnosis where asymptomatic.
- After evaluation, sexual partners should be treated empirically with regimens effective against both gonorrhea and chlamydia.
- Local public health authorities are available to assist with partner notification and appropriate referral for clinical evaluation, testing, treatment and health education when the causative organism is identified as a reportable STI.

Follow-up

- Pain and tenderness resulting from acute PID should begin to resolve within 48–72 hours of initiating antibiotics.¹³
- If no improvement is observed, further work-up is essential.
- Individuals treated as outpatients need careful follow-up and should be re-evaluated 2–3 days after treatment is initiated.
- If no clinical improvement has occurred, hospital admission for parenteral therapy and observation is required.
- Following a diagnosis of PID, patients should be informed that they are at risk of both short-term consequences such as Fitz-Hugh-Curtis syndrome (perihepatitis) and tubo-ovarian abscess, and long-term sequelae, including infertility, ectopic pregnancy and chronic pelvic pain.

Special Considerations

Pregnancy

- PID is uncommon in pregnancy, especially after the first trimester.
- Pregnant patients with suspected PID should be hospitalized for evaluation and treatment with parenteral therapy because of an increased risk of adverse outcomes for both the mother and the pregnancy.
- There is a large differential diagnosis of acute abdominal pain in pregnancy, and consultation with an expert should be sought.

HIV infection

- HIV-positive women with PID may represent a subgroup of patients with a more difficult clinical course.
- Some studies have suggested that HIV-positive women with PID have longer hospital stays and are at higher risk for the development of tubo-ovarian abscesses and are more likely to require surgical intervention.^{14,15}
- These women should be followed closely and managed aggressively, and consideration should be given to hospitalization.
- Consultation with a colleague experienced in HIV care is recommended.

Adolescents

 Consideration should be given to hospitalization for adolescents with suspected PID if poor compliance is expected to be an issue.

Patients with an intrauterine contraceptive device in situ

 In patients with an intrauterine device (IUD) in situ, the device should not be removed until after therapy has been initiated and at least two doses of antibiotics have been given.

References

- 1. Eschenbach DA. Epidemiology and diagnosis of acute pelvic inflammatory disease. *Obstet Gynecol* 1980;55(suppl 5):142S–152S.
- 2. Aral SO, Mosher WD, Cates W Jr. Self-reported pelvic inflammatory disease in the United States, 1988. *JAMA* 1991;266:2570–2573.
- 3. Westrom L, Joesoef MJ, Reynolds G, Hagdu A, Thompson SE. Pelvic inflammatory disease and fertility. A cohort study of 1,844 women with laparoscopically verified disease and 657 control women with normal laparoscopic results. *Sex Transm Dis* 1992;19:185–192.
- Kahn JG, Walker CK, Washington AE, Landers DV, Sweet RL. Diagnosing pelvic inflammatory disease: a comprehensive analysis and considerations for developing a new model. *JAMA* 1991;266:2594–2604.
- Wolner-Hanssen P. Diagnosis of pelvic inflammatory disease. In: Landers DV, Sweet RL, eds. *Pelvic Inflammatory Disease*. New York, NY: Springer-Verlag; 1997:60–75.

- Amsel R, Totten PA, Spiegel CA, Chen KC, Eschenbach D, Holmes KK. Nonspecific vaginitis. Diagnostic criteria and microbial and epidemiologic associations. Am J Med 1983;74:14–22.
- Ness RB, Soper DE, Holley RL, et al. Effectiveness of inpatient and outpatient treatment strategies for women with pelvic inflammatory disease: results from the Pelvic Inflammatory Disease Evaluation and Clinical Health (PEACH) Randomized Trial. Am J Obstet Gynecol 2002;186:929–937.
- 8. Walker CK, Kahn JG, Washington AE, Peterson HB, Sweet RL. Pelvic inflammatory disease: meta-analysis of antimicrobial regimen efficacy. *J Infect Dis* 1993;168:969–978.
- Sweet RL, Schachter J, Landers DV, Ohm-Smith M, Robbie MO. Treatment of hospitalized patients with acute pelvic inflammatory disease: comparison of cefotetan plus doxycycline and cefoxitin plus doxycycline. *Am J Obstet Gynecol* 1988;158:736–741.
- 10. Matsuda S. Clinical study of levofloxacin (LVFX) on the infectious diseases in the field of obstetrics and gynecology. *Chemotherapy* 1992;40:311–323.
- Peipert JF, Sweet RL, Walker CK, Kahn J, Reilly-Gauvin K. Evaluation of ofloxacin in the treatment of laparoscopically documented acute pelvic inflammatory disease (salpingitis). *Infect Dis Obstet Gynecol* 1999;7:138–144.
- 12. Walker CK, Workowski KA, Washington AE, Soper D, Sweet RL. Anaerobes in pelvic inflammatory disease: implications for the Centers for Disease Control and Prevention's guidelines for treatment of sexually transmitted diseases. *Clin Infect Dis* 1999;28(suppl 1):S29–S36.
- 13. Cunningham FG, Hauth JC, Strong JD, et al. Evaluation of tetracycline or penicillin and ampicillin for treatment of acute pelvic inflammatory disease. *N Engl J Med* 1977;296:1380–1383.
- Korn AP, Landers DV, Green JR, Sweet RL. Pelvic inflammatory disease in human immunodeficiency virus-infected women. *Obstet Gynecol* 1993;82: 765–768.
- 15. Barbosa D, Macasaet M, Brockmann S, Sierra MF, Xia Z, Duerr A. Pelvic inflammatory disease and human immunodeficiency virus infection. *Obstet Gynecol* 1997;89:65–70.

PROSTATITIS

Prostatitis is generally not considered a sexually transmitted infection (STI). It is included here to assist health care providers in the management of men who present with urogenital symptoms.

Definition

Providing a global definition of prostatitis is difficult because each prostatitis syndrome has its own features. One definition, provided by J.N. Krieger, is as follows: "Prostatitis is the diagnosis given to a large group of men who present with a variety of complaints referable to the lower genital tract and perineum."

In 1995, a classification for prostatitis syndromes was first proposed by the U.S. National Institute of Health, National Institute of Diabetes and Digestive and Kidney Diseases (NIH-NIDDK), it was subsequently published in 1998. A consensus meeting of the National Institutes of Health Chronic Prostatitis Collaborative Research Network held in March 2002 reconfirmed the urology research community's approval of this classification system. ² Table 1 compares the NIH-NIDDK classification system with the traditional classification system.

Table 1. NIH-NIDDK classification of prostatitis syndromes

NIH-NIDDK classification	Traditional classification	Features
Category I: acute bacterial prostatitis	Acute bacterial prostatitis	Acute bacterial infection of the prostate gland
Category II: chronic bacterial prostatitis	Chronic bacterial prostatitis	Chronic infection of the prostate characterized by recurrent urinary tract infections
Category III: chronic prostatitis / chronic pelvic pain syndrome (CP/CPPS)		Symptoms of discomfort or pain in the pelvic region for at least 3 months in the absence of uropathogenic bacteria cultured by standard techniques
Category IIIA: inflammatory chronic pelvic pain syndrome	Chronic non-bacterial prostatitis	Significant number of leuko- cytes in EPS, VB3 or semen

OF SPECIFIC SYNDROMES

Table 1. NIH-NIDDK classification of prostatitis syndromes (continued)

NIH-NIDDK classification	Traditional classification	Features
Category IIIB: non- inflammatory chronic pelvic pain syndrome	Prostatodynia	No evidence of significant leukocytes found in EPS, VB3 or semen
Category IV: asymptomatic inflammatory prostatitis	None	Leukocytes in EPS, VB3, semen or prostate tissue during evaluation for other disorders in men without symptoms of prostatitis

EPS = expressed prostatic secretions specimen (see Diagnosis section, below) VB3 = voided bladder 3 specimen (see Diagnosis section, below)

There are three major differences between the traditional and the NIH-NIDDK approaches to the classification of prostatitis syndrome:³

- The new clinical classification includes a systematic evaluation of specific symptoms characteristic of prostatitis, usually done using the NIH — Chronic Prostatitis Symptom Index (see Table 2). This symptom index is meant to be evaluative rather than discriminative in focus, it is not meant to be used as a screening or diagnostic tool. Rather, it is meant to provide a valid index of symptom severity and impact on quality of life for men with chronic prostatitis.
- The difference between inflammatory and non-inflammatory CP/CPPS is substantially different from the distinction between the traditional approach of nonbacterial prostatitis and prostatodynia.
- The new concepts have provided a critical framework for the development of research into the causes, evaluation and treatment of prostatitis syndromes.

Table 2. NIH-Chronic Prostatitis Symptom Index (NIH-CPSI)⁴

PAIN OR DISCOMFORT	
In the last week, have you experienced any pain or discomfort in the following areas?	Area between rectum and testicles (perineum) b. Testicles c. Tip of the penis (not related to urination) d. Below your waist, in your pubic or bladder area
2. In the last week, have you experienced:	a. Pain or burning during urination? □(1) □(0) b. Pain or discomfort during or after sexual climax (ejaculation)? □(1) □(0)
3. How often have you had pain or discomfort in any of these areas over the last week?	□ (0) Never □ (3) Often □ (1) Rarely □ (4) Usually □ (2) Sometimes □ (5) Always
4. Which number best describes your average pain or discomfort on the days that you had it, over the last week?	□ □ □ □ □ □ □ □ □ □ □ □ □ □ □ □ □ □ □
URINATION	
5. How often have you had a sensation of not emptying your bladder completely after you finished urinating over the last week?	☐ (0) Not at all ☐ (3) About half the time ☐ (1) Less than 1 time in 5 ☐ (4) More than half the time ☐ (5) Almost always
6. How often have you had to urinate again less than 2 hours after you finished urinating over the last week?	☐ (0) Not at all ☐ (3) About half the time ☐ (1) Less than 1 time in 5 ☐ (4) More than half the time ☐ (5) Almost always

MANAGEMENT AND TREATMI
OF SPECIFIC SYNDROMES

Table 2. NIH-Chronic Prostatitis Symptom Index (NIH-CPSI)⁴ (continued)

IMPACT OF SYMPTOMS		
7. How much have your symptoms kept you from doing the kind of things you would usually do over the last week?	☐ (0) None ☐ (1) Only a little	☐ (2) Some ☐ (3) A lot
8. How much did you think about your symptoms over the last week?	☐ (0) None ☐ (1) Only	a little (2) Some
QUALITY OF LIFE		
9. If you were to spend the rest of your life with your symptoms just the way they have been during the last week, how would you feel about that?		☐ (6) Terrible
Scoring the NIH-Chronic Prostatitis Symptom Index Domains		
Pain: tota	l of items 1a, 1b, 1c, 1d, 2a, 2b, 3	and 4 =(0-21)
Urinary symptoms: tota	total of items 5 and 6 =(0-10)	
Quality of life impact: total	total of items 7, 8 and 9 =(0–12)	

Etiology

Table 3 presents recent understanding of the etiologic agents involved in the different prostatitis syndromes.⁵

Table 3. Etiologic agents of the different prostatitis syndrome

Table 3. Ethologic agents of the different prostatitis syndrome		
Prostatitis syndrome	Etiologic agents	
Category I: acute bacterial prostatitis	Most frequent: Escherichia coli followed by species of Proteus, Providentia	
	Less common: Klebsiella, Pseudomonas, Serratia and Enterobacter	
	Minor importance: Enterococci	
	Role of Gram-positive bacteria is debated but believed to rarely cause bacterial prostatitis	
Category II: chronic bacterial prostatitis	Predominant agents are the same as for Category I	
Category IIIA: inflammatory chronic pelvic pain syndrome	Cause not known Infection with <i>Chlamydia trachomatis, Mycoplasma hominus, Ureaplasma urealyticum, Trichomonas vaginalis</i> or a viral agent may cause this type of prostatitis syndrome, but most studies do not support this view	
Category III B: non-inflammatory chronic pelvic pain syndrome	Cause not known Suggested explanations for this syndrome include a dyssynergia between bladder detrusor and internal sphincter muscles (stress prostatitis) or "pelvic floor tension myalgia"	
Category IV: asymptomatic inflammatory prostatitis	Cause not known	

Epidemiology

By some estimates, up to 50% of men experience symptoms of prostatitis at some time in their lives. Many men remain symptomatic for prolonged periods.¹

Table 4 summarizes some epidemiological characteristics as well as relative frequency of prostatitis syndromes.

Table 4. Epidemiological characteristics of prostatitis syndromes⁶

Prostatitis syndrome	Typical presentation	Approximate percent of all prostatitis syndromes
Category I: acute bacterial prostatitis	Acute illness	1–5%
Category II: chronic bacterial prostatitis	Recurrent urinary tract infection	5–10%
Category IIIA: inflammatory chronic pelvic pain syndrome	Discomfort or pain in the pelvic region for at least 3 months	40-65%
Category IIIB: non-inflammatory chronic pelvic pain syndrome	Discomfort or pain in the pelvic region for at least 3 months	20-40%
Category IV: asymptomatic inflammatory prostatitis	Asymptomatic. Discovered during evaluation for other disorders in men without symptoms of prostatitis.	Unknown

Manifestations⁵

Table 5. Main clinical features of the different prostatitis syndromes

Prostatitis syndrome	Clinical presentation
Category I: acute bacterial prostatitis	 Typically presents with fever, chills and pain in the low back, rectum or perineum, accompanied in most cases by irritative or obstructive genitourinary symptoms On digital rectal examination, the prostate is warm, firm, swollen and exquisitely tender Prostatic massage should be avoided because it is painful and may cause bacteremia

Table 5. Main clinical features of the different prostatitis syndromes (continued)

Prostatitis syndrome	Clinical presentation
Category II: chronic bacterial prostatitis	 Often presents as relapsing urinary tract infections, even after appropriate antibiotic treatment Symptoms vary from dysuria or other voiding complaints, to ejaculatory pain, hemospermia or pelvic or genital pain Some patients may be asymptomatic Urogenital physical examination is generally unremarkable
Category IIIA: inflammatory chronic pelvic pain syndrome	 Symptoms similar to those of Category II Typically does not cause cystitis-like dysuria Chronic pelvic pains (perineal, testicular, penile, lower abdominal and ejaculatory) are most prominent symptoms Urogenital physical examination is generally unremarkable
Category IIIB: noninflammatory chronic pelvic pain syndrome	 Symptoms similar to those of Category II Typically does not cause cystitis-like dysuria Chronic pelvic pains (perineal, testicular, penile, lower abdominal and ejaculatory) are most prominent symptoms Common complaints include dysuria, hesitancy, interrupted or pulsed flow, diminution in stream size or force, and dribbling Symptoms may be exacerbated by sexual activity Urogenital physical examination is generally unremarkable
Category IV: asymptomatic inflammatory prostatitis	Asymptomatic

Diagnosis⁴

- The gold-standard test for a diagnosis of bacterial prostatitis would be a prostatic biopsy, but this is rarely indicated.
- Examination of expressed prostatic secretions has been the definite test for differentiating the prostatitis syndromes. The procedure is referred to as the "four-glass" localization test (see Table 6).

- Unfortunately, the prostatic localization test has not been properly validated and its limitations are significant. Very few urologists routinely use this test and some suggest it should be confined to research trials.⁵
- A simpler, "two-glass" pre- and post-massage screening test consisting of a
 urine specimen taken before and after a prostatic massage could be as sensitive
 and specific as the "four-glass" test⁶⁻¹⁰ (same interpretation as Table 6 below for
 the "four-glass" test: pre-massage specimen is the same as voided bladder 2
 specimen [VB2] and the post-massage specimen is the same as voided bladder 3
 specimen [VB3]).
- Avoid voided bladder 1 specimen (VB1) in patients with no clinical urethritis and expressed prostatic secretions specimen (EPS), which is difficult to obtain and deal with.

Table 6. Localization cultures ("four-glass" test) for diagnosis of prostatitis syndromes

Technique

- Ensure that the patient has a full bladder at the start of the procedure
- · Retract the foreskin of uncircumcised men throughout the procedure
- · Cleanse the glans penis with soap and water or povidone-iodine
- · Collect first 10 mL of voided urine (VB1)
- Discard next 100 mL urine voided, then collect a 10 mL midstream urine specimen (VB2)
- Massage prostate and collect any expressed prostatic secretions (EPS)
- Collect first 10 mL urine voided after prostatic massage (VB3)
- Make sure all specimens are taken immediately to the laboratory for quantitative culture

Interpretation

- All specimens yield less than 10³ colony-forming units / mL: negative test for bacterial prostatitis
- VB3 or EPS yields a colony count of one or more log(s) greater than the VB1: chronic bacterial prostatitis
- VB1 yields a colony count greater than other specimens: urethritis or specimen contamination
- All specimens yield at least 10³ colony-forming units / mL: not interpretable. In this case, treat the patient for 2 to 3 days with an antibiotic that does not penetrate the prostate but will sterilize bladder urine (such as ampicillin or nitrofurantoin), then repeat procedure

EPS = expressed prostatic secretions specimen

VB1 = voided bladder 1 specimen

VB2 = voided bladder 2 specimen

VB3 = voided bladder 3 specimen

Management and Treatment⁵

Table 7 summarizes the suggested antibiotic regimens for treating acute bacterial prostatitis (Category I) and chronic bacterial prostatitis (Category II).

- Acute bacterial prostatitis responds promptly to most antibiotics.
- Treatment of acute bacterial prostatitis should be for at least 3–4 weeks with an appropriate antimicrobial with excellent tissue penetration in order to avoid complications such as prostatic abscess or chronic bacterial prostatitis.
- Available data do not allow for the recommendation of a specific fluoroquinolone, but only norfloxacin, ciprofloxacin, or ofloxacin are at present approved for the treatment of bacterial prostatitis.
- Most patients with acute prostatitis can be managed with oral antibiotics, though some patients may require IV treatment. If IV treatment is needed, ampicillin/gentamicin is recommended, although both trimethoprim-sulfa and ciprofloxacin may also be given IV (see Table 7). There are other beta-lactam antibiotic regimens that can be used, but listing them is beyond the scope of these guidelines. When IV antibiotic treatment is needed, switch promptly to oral antibiotics when the patient has clinically improved.
- Chronic bacterial prostatitis requires at least 4–6 weeks of appropriate antibiotic therapy.
- For relapse of chronic bacterial prostatitis, a treatment of 3 months may be advisable.
- If there is no response to antibiotic treatment, consider referral for evaluation.

MANAGEMENT AND TREATM
OF SPECIFIC SYNDROMES

Table 7. Potential regimens for empiric therapy of bacterial prostatitis

Prostatitis syndrome	Antibiotic regimen	Duration
Category I: acute bacterial prostatitis	In some cases, treatment may be given intravenously for the first few days.	
	 Trimethoprim-sulfamethoxazole 160/800 mg PO bid* [C-II] OR 	4 weeks
	Ofloxacin 400 mg PO bid [A-I] OR	4 weeks
	• Ciprofloxacin 500 mg PO bid [A-I] OR	4 weeks
	 Ampicillin 1 g IV every 6 hours PLUS gentamicin 5 mg/kg lean body weight IV/day [A-I][†] 	4 weeks
Category II: chronic bacterial prostatitis	Trimethoprim-sulfamethoxazole 160/800 mg PO bid* [C-II] OR	6–12 weeks
	Ofloxacin 400 mg PO bid [A-I] OR	6–12 weeks
	Ciprofloxacin 500 mg PO bid [A-I] OR	6–12 weeks
	Doxycycline 100 mg P0 bid* [I-III]	6–12 weeks

^{*} Not an approved indication by the U.S. Food and Drug Administration.

[†] This is the recommended gentamicin dose for patients with normal renal function; needs to be adjusted with renal impairment. Renal function and gentamicin levels should be monitored during therapy. Antibiotics should be stepped down promptly to oral therapy when the patient has clinically improved.

Table 8. Treatment regimens for non-bacterial prostatitis and chronic pelvic pain syndrome (Category IIIA and Category IIIB)

Treatment for Category IIIA (inflammatory non-bacterial prostatitis) is not well defined

- Occasional successes have been reported with antibiotic therapy.
- A single 4-week course of antibiotic therapy with an appropriate agent may be defensible.
- · Repeated or prolonged antibiotic courses should be avoided.
- Other measures have been suggested, but few are evidence-based (NSAIDs, alpha-blockers, finasteride [Proscar], allopurinol, nutritional supplements, lifestyle changes and prostatic massage).
- Persistent or severe voiding symptoms, especially in older patients, should be evaluated for interstitial cystitis or carcinoma of the bladder.¹¹

Treatment for *Category IIIB* (non-inflammatory chronic pelvic pain syndrome) is even more empirical than for Category IIIA

• In addition to those listed for Category IIIA, suggested approaches include muscle relaxants, analgesics, alpha-blockers, physiotherapy, neuromodulators, biofeedback, sitz baths, relaxation exercises and psychotherapy.

Consideration for Other STIs

- Evaluation for possible sexually transmitted infections should be made when appropriate, especially in younger sexually active patients, and patients with primarily urethral symptomatology or urethral discharge.
- When investigation reveals a VB1 specimen colony count greater than all other specimens (see Diagnosis section, above), consider urethritis as a possible diagnosis and investigate appropriately.

Reporting and Partner Notification

- Because prostatitis syndromes are not typically caused by a sexually transmitted pathogen, sexual partners of patients with prostatitis do not usually require evaluation or treatment.
- When investigation reveals a condition that is notifiable according to provincial and territorial laws and regulations, patients should be reported to the local public health authority.

Follow-up

 Appropriate follow-up should be arranged depending on the proven or presumed diagnosis or on the need to further investigate certain patients according to clinical presentation.

References

- Krieger JN. Prostatitis syndromes. In: Holmes KK, Sparling PF, Mardh PA, et al, eds. Sexually Transmitted Diseases. 3rd ed. New York, NY: McGraw Hill; 1999: 859–871.
- Nickel JC. Classification and diagnosis of prostatitis; a gold standard? Andrologia 2003;35:160–167.
- 3. Krieger JN, Wiedner W. Prostatitis: ancient history and new horizons. *World J Urol* 2003;21:51–53.
- Litwin MS, McNaughton-Collins M, Fowler FJ Jr, et al. The National Institutes of Health chronic prostatitis symptom index: development and validation of a new outcome measure. Chronic Prostatitis Collaborative Research Network. *J Urol* 1999;162:369–375.
- 5. Lipsky BA. Prostatitis and urinary tract infection in men: what's new: what's true? *Am J Med* 1999:106:327–334.
- Nickel JC. The Pre and Post Massage Test (PPMT): a simple screen for prostatitis. *Tech Urol* 1997;3:38–43.
- Nickel JC, Wang Y, Shoskes D, Propert K. Validation of the Pre and Post Massage Test (PPMT) for the evaluation of the patient with chronic prostatitis/ chronic pelvic pain syndrome (CP/CPPS). Chronic Prostatitis Collaborative Research Network. J Urol 2005;173(suppl 4):29.
- 8. Weidner W, Ebner H. Cytological analysis of urine after prostatic massage (VB3): a new technique for discriminating diagnosis of prostatitis. In: Brunner H, Krause W, Rothaug CF, Weidner E, eds. *Chronic Prostatitis*. Stuttgart: Schattauer; 1985: 141–151.
- Ludwig M, Schroeder-Printzen I, Ludecke G, Weidner W. Comparison of expressed prostatic secretions with urine after prostatic massage — a means to diagnose chronic prostatitis/inflammatory chronic pelvic pain syndrome. *Urology* 2000;55:175–177.
- 10. Seiler D, Zbinden R, Hauri D, John H. Four-glass or two-glass test for chronic prostatitis. *Urologe A* 2003;42:238–242.
- 11. Nickel JC. Cytologic evaluation of urine is important in evaluation of chronic prostatitis. *Urology* 2002:60;225–227.

SEXUALLY TRANSMITTED INTESTINAL AND ENTERIC INFECTIONS

Etiology¹

- Sexually transmitted intestinal syndromes involve a wide variety of pathogens at different sites of the gastrointestinal tract.
- The diversity of sexually transmissible pathogens responsible for intestinal disease remains a challenge for the clinician.
- Polymicrobial infection often occurs, causing an overlap of symptoms.
- Infections of the anus and rectum are often sexually transmitted and typically occur in men and women who engage in unprotected receptive anal intercourse.
- Sexually transmitted infections (STIs) must always be considered, but trauma and foreign bodies may result in findings suggestive of proctitis or proctocolitis.
- Some anorectal infections in women are secondary to the contiguous spread of the pathogens from the genitalia.
- Infections with pathogens traditionally associated with food- or water-borne acquisition are known to occur via sexual transmission, most often via the fecal-oral route.
- Infections are often more severe in persons infected with HIV, and the list of potential causes is greater.
- In persons with advanced HIV infection, consider cryptosporidium and microsporidium.

Definitions

- Proctitis: Inflammation limited to the rectal mucosa, not extending beyond 10–12 cm of the anal verge. Transmission of the involved pathogens is usually due to direct inoculation into the rectum during anal intercourse.
- Proctocolitis: Inflammation of the rectal mucosa and of the colon extending above 10–12 cm of the anal verge; generally has an infectious etiology different from proctitis. Transmission is usually fecal-oral.
- Enteritis: Inflammation of the duodenum, jejunum and/or ileum. Transmission is usually fecal-oral.

Table 1 lists the pathogens involved in the common sexually transmitted gastrointestinal syndromes and their modes of acquisition.

Table 1. Common sexually transmitted gastrointestinal syndromes¹

Syndrome	Pathogen(s)	Mode of acquisition
Proctitis	 Neisseria gonorrhoeae Chlamydia trachomatis(LGV and non-LGV serovars) Treponema pallidum Herpes simplex virus 	Receptive anal intercourse in the majority of cases
Proctocolitis	 Entamoeba histolytica Campylobacter species Salmonella species Shigella species C trachomatis (LGV serovars) 	Direct or indirect fecal-oral contact
Enteritis	Giardia lamblia	Direct or indirect fecal-oral contact

LGV = lymphogranuloma venereum

Epidemiology²

- Sexual practices of individuals often involve direct or indirect contact with the rectal mucosal membranes (i.e., sharing sex toys).
- Sexually transmitted intestinal syndromes occur commonly in men who have sex with men who engage in unprotected anal intercourse or oral-anal and oral-genital sexual activities.
- Heterosexual men and women can also be at risk for acquiring enteric infections by oral-anal sexual activities.
- Women can acquire sexually transmitted anorectal pathogens by unprotected anal intercourse.
- Unprotected anal intercourse is being reported more frequently among several subpopulations, such as sexually active adolescents and street youth.

Prevention

- Since anal intercourse is the main mode of sexual transmission for pathogens that cause proctitis, clinicians should identify barriers to prevention practices and discuss means to overcome them.
- Since oral-anal sexual activities are the main mode of acquisition for sexually transmitted proctocolitis and enteritis, the risks of fecal-oral contamination should be discussed, particularly with sex workers and men who have sex with men.

Manifestations

- Typical presenting symptoms of the different sexually transmitted intestinal syndromes are listed in Table 2.
- · Asymptomatic infections are also prevalent.
- Clinicians should routinely inquire about specific sexual activities, regardless
 of the patient's reported sexual preference (see Primary Care and Sexually
 Transmitted Infections chapter).

Table 2. Possible symptoms of sexually transmitted intestinal syndromes

Syndrome	List of possible symptoms
Proctitis	Anorectal painTenesmusConstipationHematochezia (bloody stools)Mucopurulent discharge
Proctocolitis	Proctitis symptomsDiarrheaCrampsAbdominal painFever
Enteritis	DiarrheaCrampsBloatingNausea

Diagnosis

- If a symptomatic patient reports any anorectal sexual activities, anoscopic evaluation should be a routine part of the physical examination.
- Specimen collection should be adapted to the clinical presentation and history, including possible exposure to lymphogranuloma venereum (LGV) (see Lymphogranuloma Venereum chapter). For example, in some cases of enteric infections, evaluation for sexually transmitted pathogens might not be relevant.
- Anoscopic examination for proctitis:
 - Obtain rectal swabs for culture, preferably under direct vision through an anoscope, for appropriate diagnostic testing for *N gonorrhoeae*, *C trachomatis* (further testing is required for positive cultures to differentiate between Chlamydia and LGV infections), and herpes simplex virus.

- A specimen from the lesions should also be collected for a diagnostic test for HSV.
- Syphilis serology should also be performed in all patients (see *Syphilis* chapter).
- Although nucleic acid amplification tests (NAATs) are available for detection
 of gonococcal and chlamydial infections in urogenital specimens, they have
 not been extensively studied for rectal specimens.
- If indicated by clinical presentation and/or history, collect stool specimen for culture for enteric pathogens and examination for ova and parasites.

Management and Treatment

- Treatment of sexually transmitted intestinal infections should be based on physical findings.
- A high index of suspicion concerning the different etiological agents should be maintained by the clinician.
- Most often, treatment of suspected proctitis will be empirical and should not await test results.

Table 3. Recommended treatment regimens according to suspected or proven diagnosis²

Suspected or proven diagnosis	Recommended treatment regimens*
If an anorectal exudate is found on examination, treat for proctitis due to <i>N gonorrhoeae</i> [†] and <i>C trachomatis</i> (see <i>Gonococcal Infections</i> chapter, and <i>Chlamydial Infections</i> chapter, for alternative treatment recommendations; see <i>Lymphogranuloma Venereum</i> chapter for treatment recommendations for LGV serovars of <i>C trachomatis</i>)	 Cefixime 400 mg PO in a single dose [A-I] OR Ciprofloxacin 500 mg PO in a single dose (unless not recommended due to quinolone resistance: see Gonococcal Infections chapter) [A-I] OR Ofloxacin 400 mg PO in a single dose (unless not recommended due to quinolone resistance: see Gonococcal Infections chapter) [A-I] PLUS Doxycycline 100 mg PO twice a day for 7–10 days [A-I] OR Azithromycin 1 g PO in a single dose if poor compliance is expected [A-I]
If patient is suspected or proven to have HSV infection	Treat with antiviral regimens according to genital HSV infection recommendations (see <i>Herpes Simplex Virus Infections</i> chapter)

Table 3. Recommended treatment regimens according to suspected or proven diagnosis² (continued)

Suspected or proven diagnosis	Recommended treatment regimens*
If patient is suspected or proven to have <i>T pallidum</i> infection	 Benzathine penicillin 2.4 million units IM in a single dose (primary and secondary syphilis) [A-I] OR Treat according to syphilis treatment recommendations for other suspected stages of syphilis or in HIV-infected individuals (see <i>Syphilis</i> chapter)
If patient is suspected or proven to have an enteric pathogen other than those listed above	Treat according to the specific pathogen management and treatment recommendations

HSV = herpes simplex virus

LGV = lymphogranuloma venereum

Consideration for Other STIs

- Proctitis is associated with specific high-risk sexual activities; therefore, patients presenting with symptoms should be evaluated for other STIs.
- Counselling and testing for HIV are recommended.
- Screening for hepatitis B markers may be considered in certain high-risk individuals before considering immunization.
- Immunization against hepatitis A and B is recommended.
- Serologic testing for syphilis should be strongly considered in all individuals presenting with proctitis.

Reporting and Partner Notification

- Patients with conditions that are notifiable according to provincial and territorial laws and regulations should be reported to the local public health authority.
- When treatment for proctitis is indicated, all partners who have had sexual
 contact with the index case within 60 days prior to onset of symptoms or date
 of diagnosis where asymptomatic should be located, clinically evaluated and
 treated with the same regimen as the index case.
- Local public health authorities are available to assist with partner notification and help with appropriate referral for clinical evaluation, testing, treatment and health education.

^{*} For references associated with the treatment recommendations, see *Chlamydial Infections, Gonococcal Infections, Genital Herpes Simplex Virus Infections* and *Lymphogranuloma Venereum* chapters.

[†] Other broad-spectrum quinolones may be effective but not recommended as first line agents because of their cost.

Follow-up

- Follow-up should be arranged for every patient. If a recommended treatment
 regimen has been given and properly taken, symptoms and signs have
 disappeared and there has been no re-exposure to any untreated partner, then
 repeat diagnostic testing for N gonorrhoeae and C trachomatis is not routinely
 recommended.
- In cases of confirmed syphilis, appropriate serological follow-up according to syphilis recommendations should be carried out.

Special Considerations

Despite movement toward more social consciousness and awareness of STIs
and diversity in sexual practices, real and perceived prejudice on the part of
some clinicians against anorectal activities may contribute to a reluctance to
seek medical care or to disclose sexual behaviours.

References

- 1. Rompalo AM. Diagnosis and treatment of sexually acquired proctitis and proctocolitis: an update. *Clin Infect Dis* 1999;28(suppl 1):S84–90.
- 2. Sexually transmitted diseases treatment guidelines 2002. Centers for Disease Control and Prevention. *MMWR Recomm Rep* 2002;51(RR-6):1–78.

URETHRITIS

Etiology¹

- · Important causes to consider:
 - Neisseria gonorrhoeae
 - Chlamydia trachomatis
- · Other possible causes:
 - Trichomonas vaginalis²
 - Herpes simplex virus³
 - Mycoplasma genitalium^{4,5}
 - Ureaplasma urealyticum¹
- Other, less common considerations include the following:
 - Adenovirus^{6,7}
 - Candida albicans⁸

Definition

- Clinical syndrome:
 - Inflammation of the urethra, with or without urethral discharge.
 - Discharge, if present, can be mucoid, mucopurulent or purulent.
 - May also be manifested by dysuria, urethral pruritis or meatal erythema.
- Microscopic definition: presence of ≥5 polymorphonuclear leukocytes (PMNs) per oil immersion field (x1000) in five non-adjacent, randomly selected fields on a smear.⁹
- Non-gonococcal urethritis (NGU) refers to urethritis not caused by N gonorrhoeae.

Epidemiology

• Limited data are available on the incidence or prevalence of urethritis.

Natural history

- Symptoms of gonococcal urethritis develop 2–6 days after acquisition.
- Symptoms of NGU develop 1–5 weeks after acquisition (usually at 2–3 weeks).
- Up to 25% of infections, especially NGU, can be asymptomatic.¹⁰

Prevention

- Use clinical evaluation as an opportunity to review safer sexual practices, explore barriers to adopting these practices and problem-solve to overcome such barriers in the future.
- · Advise on consistent condom use.
- Advise patient to abstain from unprotected intercourse until 7 days after initiating treatment.

Manifestations

- · Urethral discharge.
- · Dysuria.
- · Urethral itching or meatal erythema.
- Often asymptomatic.
- Although urinary frequency, hematuria and urgency can, on rare occasions, occur with urethritis, the presence of any of these symptoms requires more extensive evaluation.

Diagnosis

Specimen collection

- Discharge: obtain sample by having patient milk penis three to four times from base to glans.¹¹
- Endourethral swab: pass swab 2 cm into the urethra, rotate and remove for Gram stain and testing.
- Urine sample: obtain first 10–20 mL of first-catch urine, any time of day, but preferably after having not voided for at least 2 hours.¹²

Laboratory diagnosis

- Testing for both gonorrhea and chlamydia is recommended (see Chlamydial Infections and Gonococcal Infections chapters for more information on testing).
- Obtain the following:
 - Gram stain of discharge or endourethral specimen for PMNs and Gramnegative diplococci (if available).
 - If nucleic acid amplification tests (NAATs) are available:
 - NAAT of urine for *C trachomatis*^{13,14} and culture of endourethral swab for *N gonorrhoeae*.
 - If NAAT is unavailable:
 - Direct fluorescent antibody (DFA), enzyme immunoassay (EIA), or culture for *C trachomatis*¹⁴ and culture of endourethral swab for *N gonorrhoeae*.
- Although NAATs for gonorrhea may be considered in cases where transport
 and storage conditions are not conducive to maintaining the viability of
 N gonorrhoeae or obtaining a swab is not possible, culture is the preferred
 method, because it allows for antimicrobial susceptibility testing.

Caution:

- Presence of the following symptoms suggest an alternative diagnosis:
 - Hematuria.
 - Fever, chills.
 - Frequency, nocturia, urgency.
 - Perineal pain, scrotal masses.
 - Difficulties initiating and maintaining stream.
 - Lymphadenopathy.

Management and Treatment (see Figure 2)

- Gonococcal urethritis: Cefixime 400 mg PO in a single dose PLUS EITHER doxycycline 100 mg PO bid for 7 days¹⁵ [A-I] OR azithromycin 1 g PO in a single dose if poor compliance is expected [A-I].
- Non-gonococcal urethritis: doxycycline 100 mg PO bid for 7 days^{16–18} [A-I] OR azithromycin 1 g PO in a single dose if poor compliance is expected [A-I].
- Alternative regimens are available for gonococcal infections/chlamydial infections (see *Gonococcal Infections and Chlamydial Infections* chapters).
- Single-dose regimens offer improved compliance and are especially useful in certain populations such as street youth, but they are also the most expensive.
- Resolution of symptoms can take up to 7 days after therapy has been completed.
- Patients should abstain from unprotected intercourse until 7 days after initiating treatment.
- Asymptomatic infections in men are common and should be treated.

Consideration for Other STIs

- Obtain serology for syphilis.
- Review immunization status for hepatitis B; offer vaccination if the patient is not protected and testing if the patient is at high risk.
- · Offer HIV testing and counselling.
- In men who have sex with men, consider hepatitis A vaccine.

Reporting and Partner Notification

- Urethritis caused by certain agents (e.g., C trachomatis, N gonorrhoeae) is a
 notifiable communicable disease for provinces and territories. All conditions and
 diseases that are notifiable should be reported to public health departments in
 accordance with local regulations and laws.
- All sexual partners of the index case from 60 days prior to symptom onset or date of diagnosis where asymptomatic should be identified and receive a clinical evaluation, including appropriate screening tests and appropriate prophylactic treatment, regardless of findings on clinical examination.
- Where possible, encourage the use of public health authorities or treating physician to conduct contact tracing in order to increase the number of partners contacted.¹⁹

Follow-up

- If treatment is taken and symptoms resolve, test of cure is not routinely recommended.
- If symptoms persist or recur after completed therapy (1 week after initiation of therapy), the patient should be re-evaluated.

- Symptoms alone are not sufficient for retreatment in the absence of laboratory findings or clinical signs.
- If a test of cure is indicated and a NAAT is being used for follow-up testing, testing should not be conducted until 3 weeks after treatment to avoid a false positive.

Special Considerations

Recurrent or persistent urethritis

- · Often a difficult problem.
- Must reconfirm the presence of urethritis using smear and Gram stain.
- Critical to differentiate urethritis from functional complaints.
- Important to inform patient at the start of care for recurrent urethritis that it can be a difficult clinical problem to address, but that symptoms often resolve.
- If there is a microbiologically or clinically documented failure with persistent urethritis, consider the following:
 - Re-exposure to untreated partner.
 - Infection acquired from new partner.
 - Medication not taken correctly/not completed.
 - Infection with other pathogens.
 - Presence of resistant organisms. 20
 - Other causes (e.g., urinary tract infection, prostatitis, phimosis, chemical irritation, urethral strictures, tumours).
- Consider:
 - Repeat specimens (urine and endourethral) for Gram stain, culture and NAAT for N gonorrhoeae and C trachomatis.
 - Endourethral swab or urine for T vaginalis. 2,21
 - Endourethral swab or urine for herpes simplex virus culture, although usually associated with lesion.^{3,22}
 - Endourethral specimen or first-void urine for culture for *U urealyticum* and *M genitalium*⁵ (usually at specialized laboratory).
 - Urology or infectious diseases consultation if unresolved.
 - Determine whether other underlying etiologies, such as anxiety, contribute to symptoms.

Children with urethritis

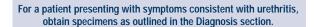
- Sexual abuse must be considered if there are symptoms of unexplained pyuria in prepubertal boys or young males who are not sexually active (see Sexual Abuse in Peripubertal and Prepubertal Children chapter).
- Practitioners need to follow provincial guidelines for reporting any suspected cases of child sexual abuse to appropriate authorities.
- Young men and women with urethritis may be erroneously diagnosed with urinary tract infections.

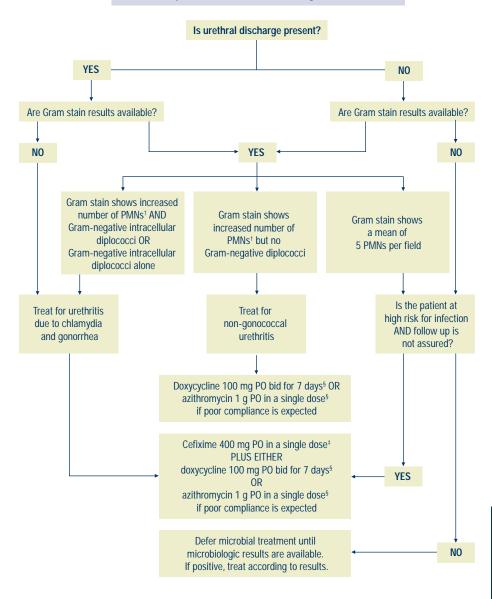
- In addition to symptoms present in adults, children with urethritis can also demonstrate the following:
 - Abdominal pain.
 - Unwillingness to void.
 - Enuresis.
- For treatment regimens in children, see Gonococcal Infections and Chlamydial Infections chapters.
- · Repeat testing should be offered to all children.

Urethritis in women

- Urethritis caused by *N gonorrhoeae* and *C trachomatis* in women can occur without cervicitis.
- Dysuria and urinary frequency may be symptoms of urethritis and thus may mimic cystitis.
- Specimens for *C trachomatis* and *N gonorrhoeae* in women should be obtained from both urine and endocervical specimens.

Figure 2. Urethritis treatment* flow chart





PMN = polymorphonuclear leukocytes

- * Treatment flow chart only. Specimens to be collected and sent for laboratory testing as outlined in the Diagnosis section.
- † A mean of \geq 5 PMNs per field (x 1000) in five non-adjacent fields.
- [‡] For alternative regimens, see *Gonococcal Infections* chapter.
- § For alternative regimens, see *Chlamydial Infections* chapter.

References

- McKee KT Jr, Jenkins PR, Garner R, et al. Features of urethritis in a cohort of male soldiers. Clin Infect Dis 2000;30:736–741.
- Wendel KA, Erbelding EJ, Gaydos CA, Rompalo AM. Use of urine polymerase chain reaction to define the prevalence and clinical presentation of Trichomonas vaginalis in men attending an STD clinic. Sex Transm Infect 2003;79:151–153.
- 3. Madeb R, Nativ O, Benilevi D, Feldman PA, Halachmi S, Srugo I. Need for diagnostic screening of herpes simplex virus in patients with nongonococcal urethritis. *Clin Infect Dis* 2000;30:982–983.
- Mena L, Wang X, Mroczkowski TF, Martin DH. Mycoplasma genitalium infections in asymptomatic men and men with urethritis attending a sexually transmitted diseases clinic in New Orleans. Clin Infect Dis 2002;35:1167–1173.
- 5. Dupin N, Bijaoui G, Schwarzinger M, et al. Detection and quantification of Mycoplasma genitalium in male patients with urethritis. *Clin Infect Dis* 2003;37:602–605.
- 6. Bradshaw CS, Denham IM, Fairley CK. Characteristics of adenovirus associated urethritis. *Sex Transm Infect* 2002;78:445–447.
- Azariah S, Reid M. Adenovirus and non-gonococcal urethritis. *Int J STD AIDS* 2000;11:548–550.
- Varela JA, Otero L, Garcia MJ, et al. Trends in the prevalence of pathogens causing urethritis in Asturias, Spain, 1989–2000. Sex Transm Dis 2003;30: 280–283.
- Swartz SL, Kraus SJ, Herrmann KL, Stargel MD, Brown WJ, Allen SD. Diagnosis and etiology of nongonococcal urethritis. *J Infect Dis* 1978;138: 445–454.
- 10. Grosskurth H, Mayaud P, Mosha F, et al. Asymptomatic gonorrhea and chlamydial infection in rural Tanzanian men. *BMJ* 1996;312;277–280.
- 11. Martin DH, Bowie WR. Management of STD syndromes in men. In: Holmes KK, Sparling P, Mardh PA, et al, eds. *Sexually Transmitted Diseases*. 3rd ed. New York, NY: McGraw Hill; 1999: 833–845.
- 12. Simmons PD. Evaluation of the early morning smear investigation. *Br J Vener Dis* 1978;54:128–129.
- 13. Burstein GR, Zenilman JM. Nongonococcal urethritis a new paradigm. *Clin Infect Dis* 1999;28(suppl 1):S66–73.
- Centers for Disease Control and Prevention (CDC). Screening tests to detect Chlamydia trachomatis and Neisseria gonorrhoeae infections — 2002. MMWR Recomm Rep 2002;51(RR-15): 1–27.
- Handsfield HH, McCormack WM, Hook EW 3rd, et al. A comparison of singledose cefixime with ceftriaxone as treatment for uncomplicated gonorrhea. The Gonorrhea Treatment Study Group. N Engl J Med 1991;325:1337–1341.
- 16. Stamm WE, Hicks CB, Martin DH, et al. Azithromycin for empirical treatment of the nongonococcal urethritis syndrome in men. A randomized double-blind study. *JAMA* 1995;274:545–549.

- 17. Steingrimsson O, Olafsson JH, Thorarinsson H, Ryan RW, Johnson RB, Tilton RC. Single dose azithromycin treatment of gonorrhea and infections caused by C. trachomatis and U. urealyticum in men. *Sex Transm Dis* 1994; 21:43–46.
- 18. Lau CY, Qureshi AK. Azithromycin versus doxycycline for genital chlamydial infections. A meta-analysis of randomized clinical trials. *Sex Transm Dis* 2002;29:497–502.
- Macke BA, Maher JE. Partner notification in the United States: an evidencebased review. Am J Prev Med 1999;17:230–242.
- Public Health Agency of Canada. Interim Statement on the Treatment of Gonorrhea in Canada. Ottawa, ON: Public Health Agency of Canada; November 2004. Available at: www.phac-aspc.gc.ca/std-mts/pdf/ is-gonorrhea-2004_e.pdf. Accessed March 1, 2005.
- 21. Borchardt KA, al-Haraci S, Maida N. Prevalence of Trichomonas vaginalis in a male sexually transmitted disease clinic population by interview, wet mount microscopy and the InPouch TV test. *Genitourin Med* 1995;71:405–406.
- 22. Lautenschlager S. Eichmann A. Urethritis: an underestimated clinical variant of genital herpes in men? *J Am Acad Dermatol* 2002;46:307–308.

VAGINAL DISCHARGE (BACTERIAL VAGINOSIS, VULVOVAGINAL CANDIDIASIS, TRICHOMONIASIS)

Etiology

- The three infections most commonly associated with vaginal discharge in adult women are:
 - Bacterial vaginosis (BV)
 - Vulvovaginal candidiasis (VVC)
 - Trichomoniasis
- On occasion, vaginal discharge may be seen in cervicitis caused by *Neisseria* gonorrhoeae or *Chlamydia trachomatis*.
- Non-infectious causes of vaginal discharge include the following:
 - Excessive physiologic secretions
 - Desquamative inflammatory vaginitis
 - Atrophic vaginitis (scant discharge)
 - Foreign bodies
- Non-infectious causes of vulvovaginal pruritis without discharge should also be considered:
 - Irritant or allergic dermatitis (e.g., latex, soaps, perfumes)
 - Skin disorders, such as the following:
 - Lichen sclerosus (may increase the risk of vulvar cancer)
 - Squamous cell hyperplasia
 - Lichen planus
 - Psoriasis

Bacterial vaginosis

- · Most common cause of vaginal discharge.
- Characterized by an overgrowth of genital tract organisms (e.g., *Gardenerella*, *Prevotella*, *Mobiluncus* spp.) and a depletion of lactobacilli.
- · Not usually considered sexually transmitted.

Vulvovaginal candidiasis

- Approximately 90% of cases caused by Candida albicans; remainder caused by other Candida spp. (e.g., C glabrata) or Saccharomyces cerevisiae.
- Not usually considered sexually transmitted.

Trichomoniasis

- · Caused by Trichomonas vaginalis, a protozoa.
- Sexually transmitted.

Epidemiology

 Vaginal complaints are common in primary care and are among the most common reasons for gynecological consultation.

Bacterial vaginosis

- Prevalence has been estimated at 10–30% of pregnant women and 10% of family practice patients.^{1,2}
- BV during pregnancy is associated with premature rupture of the membranes, chorioamnionitis, preterm labour, preterm birth and post-cesarean delivery endometritis.³
- The presence of BV during an invasive procedure, such as placement of an intrauterine device (IUD), endometrial biopsy or uterine curettage, has been associated with post-procedure pelvic inflammatory disease and vaginal cuff cellulitis.^{4,5}
- Presence of BV is associated with increased acquisition of HIV.^{6,7}

Vulvovaginal candidiasis

- Approximately 75% of women will experience at least one episode of VVC during their lifetime, and 5–10% will have more than one episode.⁸
- The incidence of recurrent VVC (four or more symptomatic episodes of VVC a year) has been estimated at 5% of women of reproductive age.⁸
- Among HIV-positive women, lower CD4 counts and high viral loads are associated with persistent Candida colonization and an increased incidence of VVC.⁹⁻¹²

Trichomoniasis

- The prevalence of trichomoniasis has not been well determined. In one study in a U.S. sexually transmitted infection (STI) clinic, the prevalence was estimated to range from 10–35%; however, these data are not likely to be generalizable.¹³ Among men attending STI clinics, the prevalence has been estimated at 3–20%.¹³
- Trichomoniasis is associated with an increased risk of HIV acquisition and transmission in women. ¹³⁻¹⁵

Prevention

- Predisposing factors for BV and VVC are listed in Table 1.
- Trichomoniasis is sexually transmitted and can be prevented by practicing safer-sex.

Manifestations and Diagnosis

- The symptoms and signs associated with these infections are not specific (see Table 1).
- Definitive diagnosis is based on laboratory testing. 16

Table 1. Diagnostic features and laboratory diagnosis

	Bacterial vaginosis	Candidiasis	Trichomoniasis
Sexual transmission	Not usually considered sexually transmitted	Not usually considered sexually transmitted	Sexually transmitted
Predisposing factors	 Often absent More common if sexually active New sexual partner IUD use 	 Often absent More common if sexually active Current or recent antibiotic use Pregnancy Corticosteroids Poorly controlled diabetes Immunocompromised 	Multiple partners
Symptoms	 Vaginal discharge Fishy odour 50% asymptomatic	Vaginal dischargeItchExternal dysuriaSuperficial dyspareuniaUp to 20% asymptomatic	 Vaginal discharge Itch Dysuria 10–50% asymptomatic
Signs	White or grey, thin, copious discharge	White, clumpy, curdy dischargeErythema and edema of vagina and vulva	 Off-white or yellow, frothy discharge Erythema of vulva and cervix ("strawberry cervix")
Vaginal pH	• >4.5	• <4.5	• >4.5
Wet mount	• PMNs • Clue cells*	Budding yeastPseudohyphae	• Motile flagellated protozoa (38–82% sensitivity)†

MANAGEMENT AND TREATMES
OF SPECIFIC SYNDROMES

Table 1. Diagnostic features and laboratory diagnosis (continued)

	Bacterial vaginosis	Candidiasis	Trichomoniasis
Gram stain	 Clue cells* Decreased normal flora Predominant Gram-negative curved bacilli and coccobacilli 	PMNsBudding yeastPseudohyphae	• PMNs • Trichomonads
Whiff test	• Positive	Negative	 Negative
Preferred treatment (see Tables 3–9)	Metronidazole Clindamycin	Antifungals	Metronidazole Treat partner

IUD = intrauterine device

PMN = polymorphonuclear leukocytes

Specimen collection

- Speculum examination.
- · Rule out cervicitis.
- Collect a sample of the discharge from the vaginal wall for microscopy (if microscopy is not available on-site, see Figure 1 for syndromic management).
- Although not a sensitive test, Gram stain may be helpful in diagnosing mucopurulent cervicitis (MPC) and gonorrhea in symptomatic females.
- A negative wet mount does not rule out an infectious cause of vaginitis.
- Culture is rarely needed in acute cases of vaginitis.

^{*}Clue cells are vaginal epithelial cells covered with numerous coccobacilli.

[†]Culture is more sensitive than microscopy for *T vaginalis*.

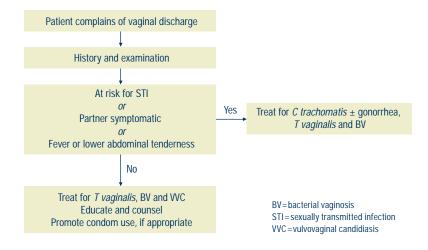
Table 2. Specimen collection

Test	Procedure	Normal result
pH test	Use narrow-range pH paper	pH ≤4.5
Wet mount	 Place a drop of vaginal discharge on a slide; mix with a drop of 0.9% saline*; apply a cover slip; examine immediately under a microscope at low and high power Examine for leukocytes, clue cells, lactobacilli, yeast and trichomonads 	Epithelial cells and rare white blood cells
Whiff test/ KOH slide (optional)	 Place a drop of discharge on a slide; mix with a drop of 10% KOH; an amine (fishy) odour after applying the KOH is a positive test; apply a cover slip; examine under a microscope at low and high power Examine for yeast 	Negative
Gram stain		Predominantly large Gram-positive bacilli

^{*} While KOH destroys cellular debris and allows one to more clearly detect yeast cells and hyphae, it also destroys the epithelial cells in clue cells needed to diagnose BV and lyses trichomonads. Therefore, for vaginitis, saline is necessary.

Figure 1. Syndromic management of vaginal discharge

For situations where on-site microscopy is not available, the World Health Organization has developed an algorithm for management of vaginal discharge.¹⁷



Consideration for Other STIs

- In a case of trichomoniasis, other STIs must be considered. If appropriate, based on the patient's and partner's risk factors (and immunization status in the case of hepatitis B), specimens can be taken for the following:
 - Gonorrhea and chlamydia
 - Syphilis
 - HIV
 - Hepatitis B

Bacterial Vaginosis

Management and Treatment

Table 3. Treatment of bacterial vaginosis

Asymptomatic	Symptomatic	
Treatment is unnecessary except in cases of: • High-risk pregnancy (history of preterm delivery) • Prior to IUD insertion • Prior to gynecologic surgery, therapeutic abortion or upper tract instrumentation	 Preferred Metronidazole 500 mg PO bid for 7 days Metronidazole gel 0.75%, one applicator (5 g) once a day intravaginally for 5 days Clindamycin cream 2%, one applicator (5 g) intravaginally once a day for 7 days Alternatives Metronidazole 2 g PO in a single dose Clindamycin 300 mg PO bid for 7 days 	
 For therapy with metronidazole, a 7 day oral course and a 5 day course of gel are equally efficacious (cure rate 75–85%).^{18–20} A single oral dose also has a cure rate of 85% but a higher relapse rate at 1 month (35–50% vs. 20–33%) [A-I]²¹ In one study, clindamycin cream was equivalent to both metronidazole regimens (cure rate of 75–86%) [A-I]²⁰ 		

IUD = intrauterine device

Notes:

- Patients should not drink alcohol during and for 24 hours after oral therapy with metronidazole because of a possible disulfiram (antabuse) reaction.
- $\bullet\,$ Clindamycin cream is oil-based and may cause latex condoms or diaphragms to fail.

Recurrent bacterial vaginosis

- 15–30% of patients develop a recurrence in the first 1–3 months after treatment.²²
- · Reconfirm diagnosis.

Table 4. Treatment of recurrent bacterial vaginosis

- Metronidazole 500 mg PO bid for 10-14 days [B-III]^{22,23}
- Metronidazole gel 0.75%, one applicator (5 g) once a day intravaginally for 10 days, followed by suppressive therapy of metronidazole gel twice a week for 4–6 months [B-III]²⁴

Note:

 Patients should not drink alcohol during and for 24 hours after oral therapy with metronidazole because of a possible disulfiram (antabuse) reaction.

Reporting and Partner Notification

- Bacterial vaginosis is not a reportable disease.
- Treatment of male sexual partners is not indicated and does not prevent recurrence.

Follow-up

• No follow-up is necessary unless the patient is pregnant or symptoms recur.

Special Considerations

Pregnancy

- BV during pregnancy is associated with premature rupture of the membranes, chorioamnionitis, preterm labour, preterm birth and post-cesarean delivery endometritis.
- Routine screening for BV during pregnancy is not recommended, although
 evidence is available to support screening and treatment at 12–16 weeks in highrisk pregnancies (see *Pregnancy* chapter). However, symptomatic women should
 be tested and treated.
- Treatment of asymptomatic BV in women with a previous preterm birth may reduce the risk of preterm prelabour rupture of the membranes and low birth weight [B-I].^{25,26} Treat with oral antibiotics: oral metronidazole and clindamycin are not contraindicated during pregnancy or breastfeeding.^{26–31} Topical antibiotics have no effect on preterm birth, though topical clindamycin treatment has been associated with adverse outcomes in the newborn when used in pregnancy (see *Pregnancy* chapter).
- Testing should be repeated after 1 month to ensure that therapy was effective.

HIV

• The same therapy is recommended for HIV-positive as for HIV-negative patients.

Vulvovaginal Candidiasis

Management and Treatment

Uncomplicated vulvovaginal candidiasis

Table 5. Treatment of uncomplicated vulvovaginal candidiasis

Asymptomatic	Symptomatic	
Treatment is unnecessary	• Intravaginal, over-the-counter azole ovules and creams (e.g., clotrimazole, miconazole)	
	Fluconazole 150 mg PO in a single dose. Contraindicated in pregnancy	
 Topical and oral azoles are equally effective [A-I].³² Efficacy estimated at 80–90%³² In most cases, expect resolution of symptoms in 2–3 days 		

Note:

· Oil-based ovules and creams may cause latex condoms or diaphragms to fail.

Complicated vulvovaginal candidiasis

 Defined as recurrent VVC, severe VVC, a non-albicans species or occurring in a compromised host.

Recurrent VVC (RVVC)

- Four or more episodes of VVC in a 12 month period.
- Confirm the diagnosis of RVVC by obtaining a vaginal culture and full identification of the isolated species, which should be used to guide therapy. Non-albicans Candida species are found in 10–20% of patients with RVVC.³³ Conventional antifungal therapy is not as effective against some of these species (see Table 8).
- Treatment requires induction, usually followed by a 6-month maintenance regimen (see Table 6).
- For patients prone to RVVC who require a course of antibiotics, prophylactic
 topical or oral azoles, such as fluconazole 150 mg PO, can be given at the start
 of the antibiotic course and once a week during the duration of the course
 [B-III].8

Table 6. Treatment of recurrent vulvovaginal candidiasis

Induction treatment

- Fluconazole 150 mg PO once every 72 hours for three doses [A-I].³⁴ Efficacy 92%. *Contraindicated in pregnancy*
- Topical azole for 10-14 days [B-II]35-38
- Boric acid 300–600 mg gelatin capsule intravaginally once a day for 14 days [B-II].^{39,40}
 Less mucosal irritation experienced when 300 mg used.⁴⁰ Efficacy approximately 80%.⁴⁰
 Contraindicated in pregnancy

Notes:

- Each individual episode of RVVC caused by C albicans usually responds to a course of oral or topical azoles, with a longer course usually more effective than a shorter one.²⁶
- Without maintenance therapy, VVC recurs in 50% of patients within 3 months.
- · Start maintenance therapy as soon as initial treatment has been completed.

Maintenance treatment

- Fluconazole 150 mg PO once a week [A-I].³⁴ Recurrence occurred in 10% while receiving therapy
- Ketoconazole 100 mg PO once a day [A-I].⁴¹ Recurrence occurred in 5% while receiving therapy. Patients receiving long-term ketoconazole should be monitored for hepatotoxicity (incidence 1 in 12,000)
- Itraconazole 200–400 mg PO once a month [A-I]. 42,43 Recurrence occurred in 36% while receiving therapy43
- Clotrimazole 500 mg intravaginally once a month [A-I]44
- Boric acid 300 mg capsule intravaginally for 5 days each month beginning the first day of the menstrual cycle [B-II].⁴⁰ Recurrence occurred in 30% while receiving therapy⁴⁰

Notes:

- Duration of maintenance therapy is a minimum of 6 months. After 6 months, discontinue therapy and observe.
- Relapse rate is high, with approximately 60% of women relapsing within 1–2 months of discontinuing maintenance therapy.^{8,36}
- · If recurrence occurs, treat the episode and then reintroduce a maintenance regimen.
- · Fluconazole and boric acid are contraindicated in pregnancy.
- · Oil-based ovules and creams may cause latex condoms or diaphragms to fail.

RVVC = recurrent vulvovaginal candidiasis

VVC = vulvovaginal candidiasis

Severe VVC

• Extensive vulvar erythema, edema, excoriation or fissure formation.

Table 7. Treatment of severe vulvovaginal candidiasis

- Fluconazole 150 mg PO once every 72 hours for two doses [A-I]. 33 Contraindicated in pregnancy
- Topical azole for 10-14 days [B-III] 8,35,37,38

Note:

· Oil-based ovules and creams may cause latex condoms or diaphragms to fail.

Non-albicans VVC

 Most commonly due to C glabrata, which is 10- to 100-fold less susceptible to azoles than C albicans.⁸

Table 8. Treatment of non-albicans vulvovaginal candidiasis

Initial treatment

- Boric acid 600 mg capsule intravaginally once a day for 14 days [B-II]. 38,39,45,46 Efficacy 64–81%. Vaginal burning reported in <10%
- Flucytosine cream 5 g intravaginally once a day for 14 days [B-II]. 46,47 Efficacy 90%
- Amphotericin B 50 mg suppository intravaginally once a day for 14 days [*B-III*].⁴⁸ Efficacy 80% (10 patients). Mild external irritation reported in 10%
- Flucytosine 1 g PLUS amphotericin B 100 mg (combined in a lubricating jelly) administered intravaginally once a day for 14 days [*B-III*].^{49,50} Efficacy 100% (4 patients)

If symptoms recur

 Retreat with boric acid 600 mg capsule intravaginally once a day for 14 days followed by: alternate-day boric acid for several weeks or 100,000 units of nystatin vaginal suppositories once a day for 3–6 months [B-III]⁸

Note:

No safety data available for long-term use of boric acid.⁵¹

Compromised host

- Corticosteroids, uncontrolled diabetes.
- *C glabrata* and other non-*albicans* species are isolated more frequently in women with diabetes than in those without.
- Treat with a longer (10–14 day) course of an intravaginal azole [B-III] or boric acid 600 mg capsule intravaginally once a day for 14 days [B-II].^{37,38}

Reporting and Partner Notification

- · Vulvovaginal candidiasis is not a reportable disease.
- Routine screening and treatment of male partners is not indicated.⁵²⁻⁵⁴ However, male sexual partners should be treated if *Candida* balanitis is present. Use a topical azole cream twice a day for 7 days.

Follow-up

- No follow-up necessary unless symptoms persist or recur.
- Consider culture and sensitivity of yeast if not responding to appropriate therapy or if infection recurs.

Special Considerations

Pregnancy

 Only topical azoles are recommended for treatment of vulvovaginal candidiasis during pregnancy. Treatment for 7 days may be necessary.⁵⁵

HIV

- The treatment of candidiasis is the same in HIV-positive as it is in HIV-negative individuals.
- Vaginal candidiasis is often recurrent and more severe in HIV-positive women and, in some cases, will require more aggressive and long-term therapy.

Trichomoniasis

Management and Treatment

Table 9. Treatment of trichomoniasis

- Metronidazole 2 g PO in a single dose [A-I]⁵⁶
- Metronidazole 500 mg PO bid for 7 days [A-I]⁵⁶
- Efficacy 82–88% for both regimens; increases to 95% if partner also treated⁵⁶
- · Intravaginal metronidazole gel is not effective

Note:

 Patients should not drink alcohol during and for 24 hours after oral therapy with metronidazole because of a possible disulfiram (antabuse) reaction.

Reporting and Partner Notification

- Trichomoniasis is a reportable disease in some jurisdictions.
- Partners should be treated for trichomoniasis, regardless of symptoms (it is not necessary to screen partners for trichomonas). The majority of men infected with *T vaginalis* are asymptomatic, but some may have mild urethritis. Treat sexual partners with the same therapy as recommended for the case.

Follow-up

- No follow-up necessary unless symptoms recur; usually due to reinfection.
- Prevalence of metronidazole-resistant T vaginalis estimated at 5%. Usually responds to high-dose metronidazole.⁵⁷

Special Considerations

Pregnancy

- Trichomoniasis may be associated with premature rupture of the membranes, preterm birth and low birth weight.
- Symptomatic pregnant women should be treated with metronidazole 2 g PO in a single dose for symptom relief [A-I]. An alternative treatment is metronidazole 500 mg PO bid for 7 days [A-I]. It is not known whether treatment will improve pregnancy outcomes.^{58,59}
- It is not recommended that asymptomatic pregnant women be treated [D-I].60
- Metronidazole is not contraindicated during pregnancy or breastfeeding. ^{26–31}

HIV

• The same therapy is recommended for HIV-positive as for HIV-negative patients.

The Use of Live Lactobacilli to Restore Normal Vaginal Flora

- Lactobacilli preparations are commonly used in the treatment of BV and VVC. One small randomized trial in healthy women showed that the use of oral Lactobacilli was safe and resulted in increased vaginal Lactobacilli and decreased yeast as compared to the placebo group.⁶¹ However, in a more recent, well-conducted randomized, controlled trial of 278 women, oral and vaginal L rhamnosus was ineffective in the prevention of post-antibiotic VVC.⁶²
- Two randomized, controlled trials have studied the use of a topical
 L acidophilus—low dose estriol combination, one in the management of BV,
 the other for several infections (BV, VVC, trichomoniasis).^{63,64} Both showed
 a statistically significant greater reduction in symptoms and microscopic
 restoration of normal flora than the placebo group.

References

- 1. Mead PB. Epidemiology of bacterial vaginosis. *Am J Obstet Gynecol* 1993;169(2 Pt 2):446–449.
- 2. Schmidt H, Hansen JG. Bacterial vaginosis in a family practice population. *Acta Obstet Gynecol Scand* 2000;79:999–1005.
- 3. Leitich H, Bodner-Adler B, Brunbauer M, Kaider A, Egarter C, Husslein P. Bacterial vaginosis as a risk factor for preterm delivery: a meta-analysis. *Am J Obstet Gynecol* 2003;189:139–147.
- Soper DE, Bump RC, Hurt WG. Bacterial vaginosis and trichomoniasis vaginitis are risk factors for cuff cellulitis after abdominal hysterectomy. *Am J Obstet Gynecol* 1990;163:1016–1021.
- 5. Penney GC, Thompson M, Norman J, et al. A randomised comparison of strategies for reducing infective complications of induced abortion. *Br J Obstet Gynaecol* 1998;105:599–604.
- Taha TE, Hoover DR, Dallabetta GA, et al. Bacterial vaginosis and disturbances of vaginal flora: association with increased acquisition of HIV. AIDS 1998;12:1699–1706.
- Martin HL, Richardson BA, Nyange PM, et al. Vaginal lactobacilli, microbial flora, and risk of human immunodeficiency virus type 1 and sexually transmitted disease acquisition. *J Infect Dis* 1999;180:1863–1838.
- 8. Sobel JD. Management of patients with recurrent vulvovaginal candidiasis. *Drugs* 2003;63:1059–1066.
- 9. Duerr A, Heilig C, Meikle S, et al. Incident and persistent vulvovaginal candidiasis among human immunodeficiency virus-infected women: risk factors and severity. *Obstet Gynaecol* 2003;101:548–556.
- 10. McClelland R, Lavreys L, Katingima C, et al. Contribution of HIV-1 infection to acquisition of sexually transmitted disease: a 10-year prospective study. *J Infect Dis* 2005;191:333–338.
- 11. Ohmit S, Sobel J, Schuman P, et al. Longitudinal study of mucosal Candida species colonization and candidiasis among human immunodeficiency virus (HIV)-seropositive and at-risk HIV-seronegative women. *J Infect Dis* 2003;188:118–127.
- Schuman P, Sobel J, Ohmit S, et al. Mucosal candidal colonization and candidiasis in women with or at risk for human immunodeficiency virus infection. Clin Infect Dis 1998;27:1161–1167.
- 13. Sorvillo F, Smith L, Kerndt P, Ash L. Trichomonas vaginalis, HIV, and African-Americans. *Emerg Infect Dis* 2001;7:927–932.
- 14. Fleming DT, Wasserheit JN. From epidemiological synergy to public health policy and practice: the contribution of other sexually transmitted diseases to sexual transmission of HIV infection. *Sex Transm Infect* 1999;75:3–17.
- 15. Schwebke J. Update of trichomoniasis. Sex Transm Infect 2002;78:378–379.
- Anderson MR, Klink K, Cohrssen A. Evaluation of vaginal complaints. *JAMA* 2004;291:1368–1379.
- 17. Guidelines for the management of sexually transmitted infections. Geneva, Switzerland; World Health Organization: 2003.

- Lugo-Miro V, Green M, Mazur L. Comparison of different metronidazole therapeutic regimens for bacterial vaginosis. A meta-analysis. *JAMA* 1992;268:92–95.
- 19. Hanson JM, McGregor JA, Hillier SL, et al. Metronidazole for bacterial vaginosis. A comparison of vaginal gel vs. oral therapy. *J Reprod Med* 2000:45:889–896.
- 20. Ferris DS, Litaker MS, Woodward L, Mathis D, Hendrich J. Treatment of bacterial vaginosis: a comparison of oral metronidazole, metronidazole vaginal gel, and clindamycin vaginal cream. *J Fam Pract* 1995;41:443–449.
- 21. Swedberg J, Steiner JF, Deiss F, Steiner S, Driggers DA. Comparison of single-dose vs one-week course of metronidazole for symptomatic bacterial vaginosis. *JAMA* 1985;254:1046–1049.
- 22. Sobel JD. Vaginitis. N Engl J Med 1997;337:1896–1903.
- 23. Sobel J. Bacterial vaginosis. *Annu Rev Med* 2000;51:349–356.
- 24. Alfonsi GA, Shlay JC, Parker S, Neher JO. What is the best approach for managing recurrent bacterial vaginosis? *J Fam Pract* 2004;53:650–652.
- 25. Guise JM, Mahon SM, Aickin M, Helfand M, Peipert JF, Westhoff C. Screening for bacterial vaginosis in pregnancy. *Am J Prev Med* 2001;20(suppl 3):62–72.
- 26. McDonald H, Brocklehurst P, Parsons J, Vigneswaran R. Antibiotics for treating bacterial vaginosis in pregnancy. *Cochrane Database Syst Rev* 2003;2: CD000262.
- 27. Piper JM, Mitchel EF, Ray WA. Prenatal use of metronidazole and birth defects: no association. *Obstet Gynecol* 1993;82:348–352.
- 28. Czeizel A, Rockenbauer M. A population based case-control teratologic study of oral metronidazole. *Br J Obstet Gynaecol* 1998;105:322–327.
- 29. Burtin P, Taddio A, Ariburnu O, Einarson TR, Koren G. Safety of metronidazole in pregnancy: a meta-analysis. *Am J Obstet Gynecol* 1995;172(2 Pt 1):525–529.
- Caro-Paton T, Carvajal A, Martin de Diego I, et al. Is metronidazole teratogenic? A meta-analysis. Br J Clin Pharmacol 1997;44:179–182.
- 31. Passmore CM, McElnay JC, Rainey EA, D'Arcy PF. Metronidazole excretion in human milk and its effect on the suckling neonate. *Br J Clin Pharmacol* 1988;26:45–51.
- Watson MC, Grimshaw JM, Bond CM, Mollison J, Ludbrook A. Oral versus intra-vaginal imidazole and triazole anti-fungal treatment of uncomplicated vulvovaginal candidiasis (thrush). *Cochrane Database Syst Rev* 2001;4: CD002845.
- 33. Sobel JD, Kapernick PS, Zervos M, et al. Treatment of complicated Candida vaginitis: comparison of single and sequential doses of fluconazole. *Am J Obstet Gynecol* 2001;185:363–369.
- 34. Sobel JD, Wiesenfeld HC, Martens M, et al. Maintenance fluconazole therapy for recurrent vulvovaginal candidiasis. *N Engl J Med* 2004;351:876–883.
- Sobel JD, Faro S, Force RW, et al. Vulvovaginal candidiasis: epidemiologic, diagnostic, and therapeutic considerations. Am J Obstet Gynecol 1998;178:203–211.

- Sobel JD, Brooker D, Stein GE, et al. Single oral dose fluconazole compared with conventional clotrimazole topical therapy of Candida vaginitis. Fluconazole Vaginitis Study Group. Am J Obstet Gynecol 1995;172(4 Pt 1):1263–1268.
- 37. Rex JH, Walsh TJ, Sobel JD, et al. Practice guidelines for the treatment of candidiasis. *Clin Infect Dis* 2000;30:662–678.
- 38. Pappas PG, Rex JH, Sobel JD, et al. Guidelines for treatment of candidiasis. *Clin Infect Dis* 2004;38:161–189.
- 39. Sobel JD, Chaim W. Treatment of Torulopsis glabrata vaginitis: retrospective review of boric acid therapy. *Clin Infect Dis* 1997;24:649–652.
- 40. Guaschino S, De Seta F, Sartore A, et al. Efficacy of maintenance therapy with topical boric acid in comparison with oral itraconazole in the treatment of recurrent vulvovaginal candidiasis. *Am J Obstet Gynecol* 2001;184:598–602.
- 41. Sobel JD. Recurrent vulvovaginal candidiasis. A prospective study of the efficacy of maintenance ketoconazole therapy. *N Engl J Med* 1986;315: 1455–1458.
- 42. Creatsas GC, Charalambidis VM, Zagotzidou EH, Anthopoulou HN, Michailidis DC, Aravantinos DI. Chronic or recurrent vaginal candidosis: short-term treatment and prophylaxis with itraconazole. *Clin Ther* 1993;15:662–671.
- 43. Spinillo A, Colonna L, Piazzi G, Baltaro F, Monaco A, Ferrari A. Managing recurrent vulvovaginal candidiasis. Intermittent prevention with itraconazole. *J Reprod Med* 1997;42:83–87.
- 44. Roth AC, Milsom I, Forssman L, Wahlen P. Intermittent prophylactic treatment of recurrent vaginal candidiasis by postmenstrual application of a 500 mg clotrimazole vaginal tablet. *Genitourin Med* 1990;66:357–360.
- 45. Jovanovic R, Congema E, Nguyen HT. Antifungal agents vs. boric acid for treating chronic mycotic vulvovaginitis. *J Reprod Med* 1991;36:593–597.
- 46. Sobel JD, Chaim W, Nagappan V, Leaman D. Treatment of vaginitis caused by Candida glabrata: use of topical boric acid and flucytosine. *Am J Obstet Gynecol* 2003;189:1297–1300.
- 47. Horowitz BJ. Topical flucytosine therapy for chronic recurrent Candida tropicalis infections. *J Reprod Med* 1986;31:821–824.
- 48. Phillips AJ. Treatment of non-albicans Candida vaginitis with amphotericin B vaginal suppositories. *Am J Obstet Gynecol* 2005;192:2009–2012.
- 49. White DJ, Habib AR, Vanthuyne A, Langford S, Symonds M. Combined topical flucytosine and amphotericin B for refractory vaginal Candida glabrata infections. *Sex Transm Infect* 2001;77:212–213.
- 50. Shann S, Wilson J. Treatment of Candida glabrata using topical amphotericin B and flucytosine. *Sex Transm Infect* 2003;79:265–266.
- 51. Fidel PL Jr, Vazquez JA, Sobel JP. Candida glabrata: review of epidemiology, pathogenesis, and clinical disease with comparison to C. albicans. *Clin Microbiol Rev* 1999;12:80–96.
- 52. Buch A, Skytte Christensen E. Treatment of vaginal candidosis with natamycin and effect of treating the partner at the same time. *Acta Obstet Gynecol Scand* 1982;61:393–396.

- 53. Bisschop MP, Merkus JM, Scheygrond H, van Cutsem J. Co-treatment of the male partner in vaginal candidosis: a double-blind randomized control study. *Br J Obstet Gynaecol* 1986;93:79–81.
- 54. Calderon-Marquez J. Itraconazole in the treatment of vaginal candidosis and the effect of treatment of the sexual partner. *Rev Infect Dis* 1987;9(suppl 1): S143–S145.
- 55. Young GL, Jewell D. Topical treatment for vaginal candidiasis (thrush) in pregnancy. *Cochrane Database Syst Rev* 2001;4:CD000225.
- 56. Forna F, Gulmezoglu AM. Interventions for treating trichomoniasis in women. *Cochrane Database Syst Rev* 2003;2:CD000218.
- 57. Schmid G, Narcisi E, Mosure D, Secor WE, Higgins J, Moreno H. Prevalence of metronidazole-resistant *Trichomonas vaginalis* in a gynecology clinic. *J Reprod Med* 2001:46:545–549.
- 58. Kigozi GG, Brahmbhat H, Wabwire-Mangen F, et al. Treatment of Trichomonas in pregnancy and adverse outcomes of pregnancy: a subanalysis of a randomized trial in Rakai, Uganda. *Am J Obstet Gynecol* 2003;189:1398–1400.
- 59. Gulmezoglu A. Interventions for trichomoniasis in pregnancy. *Cochrane Database Syst Rev* 2002;3:CD000220.
- 60. Klebanoff MA, Carey JC, Hauth JC, et al. Failure of metronidazole to prevent preterm delivery among pregnant women with asymptomatic Trichomonas vaginalis infection. *N Engl J Med* 2001;345:487–493.
- 61. Reid G, Charbonneau D, Erb J, et al. Oral use of Lactobacillus rhamnosus GR-1 and *L. fermentum* RC-14 significantly alters vaginal flora: randomized, placebo-controlled trial in 64 healthy women. *FEMS Immunol Med Microbiol* 2003;35;131–134.
- 62. Pirotta M, Gunn J, Chondros P, et al. Effect of lactobacillus in preventing post-antibiotic vulvovaginal candidiasis: a randomised controlled trial. *BMJ* 2004;329:548.
- 63. Parent D, Bossens M, Bayot D, et al. Therapy of bacterial vaginosis using exogenously applied Lactobacilli acidophili and a low dose of estriol: a placebo-controlled multicentric clinical trial. *Arzneimettelforschung* 1996;46:68–73.
- 64. Ozkinay E, Terek MC, Yayci M, et al. The effectiveness of live lactobacilli in combination with low dose oestriol (Gynoflor) to restore the vaginal flora after treatment of vaginal infections. *BJOG* 2005;112:234–240.

MANAGEMENT AND TREATMENT OF SPECIFIC INFECTIONS

CHANCROID

Etiology

• Genital ulcer disease (GUD) due to *Haemophilus ducreyi* or chancroid. *H ducreyi* is a fastidious Gram-negative rod.

Epidemiology

- Chancroid has been widespread in areas of the world where sexually transmitted infection (STI) control is inadequate. Vulnerable females (particularly sex workers with limited access to care) who have multiple partners in spite of genital ulceration are the usual reservoir. Chancroid can only remain endemic in this context.^{1,2}
- Reintroduction into societies in which chancroid has been eliminated occasionally occurs with travel. Clusters can occur around an index case (has been described in Canada).¹
- It is readily eliminated with control activities directed toward sex workers, treatment of men with genital ulcers and enhanced attention to STI-control efforts.
- Chancroid is transmitted only by individuals with ulcerations; no latent reservoir
 of transmissible chancroid without active disease is known.
- The attack rate following intercourse with contacts who have not used protection is substantial (probably >50% of exposed men or women); incubation period is 5–14 days.
- In endemic areas, as many as 10% of chancroid patients may have concomitant herpes simplex virus (HSV) infection. *Treponema pallidum* may also co-exist with H ducrevi.
- Chancroid gained significance as an important STI when its role in the transmission of HIV became apparent during the 1980s.³
 - Accelerated increases in HIV prevalence have occurred in societies in which chancroid was endemic.
 - The risk of HIV transmission increases by 10–50-fold following sexual exposure to an individual with concomitant *H ducreyi* and HIV infection.^{2,3}
 As a result, extensive research has been directed toward *H ducreyi* and chancroid.⁴
- Control can be achieved in most societies with limited infrastructure and resources.²
 - Has been essentially eliminated during the past decade from many areas of the world in which it was previously endemic, including much of eastern and southern Africa.²
 - Importation into other countries where it has already been eliminated will likely occur with reduced frequency.

Prevention

- Conventional STI-control measures are very effective: reducing the number of partners, the promotion and use of condoms for all high-risk sexual activities and early diagnosis in countries where chancroid is endemic.
- Female sex workers need to be trained to recognize genital ulceration and should have access to medical care.
- In an outbreak, microbiological diagnosis, enhanced education of sex workers and clients, and syndromic treatment of ulcers have together been very successful at limiting spread and eliminating H ducreyi infection locally.²
- Male circumcision also reduces susceptibility to *H ducreyi* infection; chancroid has been shown not to spread in populations where all men are circumcised.

Manifestations

- A papule develops following exposure, and this rapidly progresses to one or more pustular lesions. These rupture to form painful, purulent, shallow ulcers with a granulomatous base that readily bleeds.
 - In males, lesions occur on the prepuce, coronal sulcus and shaft of the penis.
 - In females, lesions can occur widely on the external genitalia but are rarely seen in the vagina or on the cervix.
- Multiple ulcers are common, particularly in women.
- Painful inguinal lymphadenitis occurs in 30% of patients, and lymph nodes may suppurate, become fluctuant and spontaneously rupture.
- Chancroid can mimic other genital ulcer diseases, particularly syphilis; however, chancroid lesions are usually painful, and classic primary syphilis chancres are generally painless.
- Chancroid rarely spreads from the genital tract and does not cause systemic disease.⁵

Diagnosis

- Clinical etiologic diagnosis is frequently erroneous; in Canada, careful etiologic investigation of an ulcer should be carried out, since chancroid is not known to be endemic.
 - Should include, wherever possible, culture for H ducreyi using specialized culture or transport media; these vary by location (check with your local laboratory for more information).
 - Other causes of GUD should be ruled out by performing either a dark field analysis or direct fluorescent antibody test for *T pallidum* for primary syphilis and a culture for HSV.
 - There are no useful serologic tests for the diagnosis of *H ducreyi*. Gram stain with Gram-negative coccobacilli in a "school of fish" pattern may be useful.
- Culture for *H ducreyi* requires specialized media.⁴ In Kenya, the use of both gonococcal and Mueller Hinton agar facilitated the growth of most strains in prospective studies. Specimens should be collected from the base of ulcers into thioglycolate hemin-based transport media, as this can permit bacterial survival

- (2–3 days at 4°C) while the medium is being prepared.⁴ *H ducreyi* grow optimally at 32°C in a humid atmosphere containing 5% carbon dioxide.
- Nucleic acid amplification tests (NAATs) including a multiplex polymerase chain reaction (M-PCR) technique that identifies H ducreyi, T pallidum and HSV can be used but are not available in most laboratories.

Management

- Syndromic management is used globally for the immediate treatment of GUD
 at first contact with the health care system; it has been particularly effective at
 controlling both syphilis and chancroid. Intermittent, careful investigation should
 be performed in most societies to determine which microbial etiologies require
 syndromic management.
- Outbreak investigation and control should be routine in all countries in which
 syphilis and chancroid have been "eliminated." A rapid-response mode should
 be available to immediately address the appearance of either of these ulcerative
 diseases, with strategies to achieve effective re-establishment of regions "free"
 of both H ducreyi and T pallidum.
- All patients diagnosed with chancroid should undergo testing to rule out co-infection with other STIs, including HIV.

Treatment

- Syndromic treatment for chancroid consists of a single dose of 500 mg of ciprofloxacin, which has a cure rate of >90% [A-I].⁶
- A 1-week course of erythromycin, 500 mg tid, also provides an excellent cure rate of >90% but is associated with poorer compliance [A-I].
- Another macrolide, azithromycin, has cured over 90% of patients when prescribed as a single oral 1 g dose [A-I].⁷⁻⁹
- Ceftriaxone 250 mg IM has been successful, but failures have commonly occurred in HIV co-infected individuals [A-I].^{7,9,10}
- Treatment failures should be carefully evaluated with regard to both the etiology and the possible co-existence of other pathogens. Buboes should be aspirated or incised to relieve pain and prevent spontaneous rupture.

Reporting and Partner Notification

All individuals who had sexual exposure to the index patient during the 2 weeks
prior to the date of initial symptoms should be treated epidemiologically with a
quinolone or another antibacterial known to be effective for index case(s).

Consideration for Other STIs

- Patients suspected of having chancroid should also be considered for the following STIs:
 - Lymphogranuloma venereum
 - HSV
 - Syphilis
 - Donovanosis (granuloma inquinale)

- All patients with presumed chancroid should also be tested for syphilis and HIV infection at presentation and 3 months later. Patients should also be tested appropriately for gonorrhea.
- Immunization for hepatitis B should be offered to non-immune patients.
- The opportunity to provide safer-sex counselling should not be missed.

Follow-up

Repeat diagnostic testing for the detection of H ducreyi is not routinely indicated
if a recommended treatment is given and taken AND symptoms and signs
disappear AND there is no re-exposure to an untreated partner.

References

- Hammond GW, Slutchuk M, Scatliff J, Sherman E, Wilt JC, Ronald AR. Epidemiologic, clinical, and laboratory therapeutic features of an urban outbreak of chancroid in North America. Rev Infect Dis 1980;2:867–879.
- 2. Steen R. Eradicating chancroid. Bull World Health Organ 2001;79:818–826.
- 3. Cameron DW, Simonsen JN, D'Costa LJ, et al. Female to male transmission of human immunodeficiency virus type 1: risk factors for seroconversion in men. *Lancet* 1989;2:403–407.
- 4. Spinola SM, Bauer ME, Munson RS Jr. Immunopathogenesis of Haemophilus ducreyi infection (chancroid). *Infect Immun* 2002;70:1667–1676.
- 5. Trees DL, Morse SA. Chancroid and Haemophilus ducreyi: an update. *Clin Microbiol Rev* 1995;8:357–375.
- Malonza IM, Tyndall MW, Ndinya-Achola JO, et al. A randomized, double-blind, placebo-controlled trial of single-dose ciprofloxacin versus erythromycin for the treatment of chancroid in Nairobi, Kenya. *J Infect Dis* 1999;180:1886–1893.
- Roest RW, van der Meijden WI; European Branch of the International Union against Sexually Transmitted Infection and the European Office of the World Health Organization. European guideline for the management of tropical genito-ulcerative diseases. *Int J STD AIDS* 2001;12(suppl 3):78–83.
- 8. Tyndall MW, Agoki E, Plummer FA, Malisa W, Ndinya-Achola JO, Ronald AR. Single dose azithromycin for the treatment of chancroid: a randomized comparison with erythromycin. *Sex Transm Dis* 1994;21:231–234.
- Martin DH, Sargent SJ, Wendel GD Jr, McCormack WM, Spier NA, Johnson RB. Comparison of azithromycin and ceftriaxone for the treatment of chancroid. *Clin Infect Dis* 1995;21:409–414.
- Tyndall M, Malisa M, Plummer FA, Ombetti J, Ndinya-Achola JO, Ronald AR. Ceftriaxone no longer predictably cures chancroid in Kenya. *J Infect Dis* 1993;167:469–471.

CHLAMYDIAL INFECTIONS

(For Lymphogranuloma venereum, see Genital Ulcer Disease and Lymphogranuloma Venereum chapters)

Etiology

· Caused by Chlamydia trachomatis serovars D to K.

Epidemiology

- Reported rate in Canada and elsewhere has been increasing since 1997.¹
- According to preliminary data, approximately 63,000 cases were reported in 2004 (197 per 100,000 population). (Preliminary data is subject to change; does not include Nunavut.)²
- Sexually active youth and young adults are disproportionately represented in the case reports for Chlamydia. The reported rate in 2004 was highest in youth/young adults 15 to 24 years of age, accounting for approximately ²/₃ of the national reported cases.²
- Chlamydia is underdiagnosed because the majority of infected individuals are asymptomatic.³⁻⁸
- Underscreening is a gap in high-risk males and females. Males, the forgotten reservoir, have infrequent health-maintenance visits. 9-11
- The usual incubation period from time of exposure to onset of infection is 2–3 weeks, but can be as long as 6 weeks.
- In the absence of treatment, infection persists for many months.
- Individuals infected with Neisseria gonorrhoeae are often co-infected with C trachomatis.^{12,13}
- · Risk factors:
 - Sexual contact with a chlamydia-infected person.
 - A new sexual partner or more than two sexual partners in the past year.
 - Previous sexually transmitted infections (STIs).
 - Vulnerable populations (e.g., injection drug users, incarcerated individuals, sex trade workers, street youth etc.) (see Specific Populations section).

Prevention

- Infection and its sequelae can be prevented by:
 - Consistent practice of safer sex (see Primary Care and Sexually Transmitted Infections chapter).
 - Identifying barriers to prevention practices and the means to overcome them.
 - Increased acceptance of testing by using a non-invasive urine-based nucleic acid amplification test (NAAT).

- Screening of at-risk groups (as per risk factors listed above).
 - Sexually active females under 25 years of age: evidence is insufficient for or against screening asymptomatic young males, though males with any risk factors (as listed above) should be screened.^{7,8,10,14-21}
 - Pregnant women: all pregnant women should be screened at the first prenatal visit. For those who are positive or who are at high risk for reinfection, rescreening in the third trimester is indicated.²²⁻²⁸
- Repeat screening of individuals with chlamydia infection after 6 months. 23,29-32
 - To prevent reinfection, partners need to be assessed, tested, treated and counselled.
 - Patients and contacts should abstain from unprotected intercourse until treatment of both partners is complete (i.e., after completion of a multipledose treatment or for 7 days after single-dose therapy).

Manifestations

Table 1. Symptoms and signs³³

Females	Males	Neonates and infants
 Most often asymptomatic Vaginal discharge Dysuria Lower abdominal pain Abnormal vaginal bleeding Dyspareunia Conjunctivitis Proctitis (commonly asymptomatic) 	 Often asymptomatic Urethral discharge Urethral itch Dysuria Testicular pain Conjunctivitis Proctitis (commonly asymptomatic) 	 Conjunctivitis in neonates Pneumonia in infants 6 months of age

Table 2. Major sequelae

Females	Males
Pelvic inflammatory diseaseEctopic pregnancyInfertilityChronic pelvic painReiter syndrome	Epididymo-orchitisReiter syndrome

Diagnosis

Laboratory diagnosis

(See Laboratory Diagnosis of Sexually Transmitted Infections chapter)

- Results are highly dependent on the type of test available; specimen collection
 and transport; and laboratory expertise. Consult with your local laboratory
 regarding available tests and their test performance.
- NAATs (e.g., polymerase chain reaction [PCR], transcription-mediated amplification [TMA]) are more sensitive and specific than culture, enzyme immunoassay (EIA) and direct fluorescent antibody assay (DFA). For non-medico-legal purposes, NAATs should be used whenever possible for urine, urethral or cervical specimens. Blood and mucus interfere with NAAT performance and can result in false-negative results: therefore, culture is recommended in such situations. NAATs have not been approved for use in vaginal specimens outside of a research setting. Culture is recommended for throat and rectal specimens, since NAATs have not been adequately evaluated on these specimens.
- Due to its non-invasive nature a urine-based NAAT is ideal for screening
 asymptomatic persons when a pelvic examination is not warranted for other
 reasons. However, a physical examination remains essential, and more invasive
 specimens may be needed for diagnostic purposes in symptomatic individuals.
- Postexposure testing with a NAAT can be done as soon as desired, since it is not necessary to wait for 48 hours after exposure to collect samples as in the case of cultures.
- Both chlamydia and gonorrhea can be detected from a single specimen by some NAATs.
- Culture has been the preferred method for medico-legal purposes. A NAAT may
 be suitable, provided that positive results are confirmed by a different set of
 primers, but it may not be available in most laboratories.
- C trachomatis IgM serology is useful for diagnosing C trachomatis pneumonia in infants under 3 months of age.
- Serology is not useful for the diagnosis of acute genital chlamydial infections.

Specimen collection

- Potential specimen sites:
 - Cervix in pubertal or older females for NAAT.
 - If the cervix has been surgically removed:
 - Send urine for NAAT.
 - Send urethral swab for culture.
 - Send rectal swab for culture.
 - Send vaginal swab for culture.

- Urethral swab in males for NAAT (preferably not having voided for at least 2 hours, but this does not preclude testing).
- Urine NAAT, vaginal/rectal swab for culture in prepubertal girls.
- Urine NAAT for females and males of any age.
 - · Any time of day.
 - Initial 10–20 mL of the urine stream (not mid-stream).
 - Preferably not having voided for at least 2 hours, but this does not preclude testing.
- Endometrial or fimbrial biopsy specimens for NAAT in women undergoing laparoscopy for investigation of pelvic inflammatory disease.
- Conjunctival swab for culture, EIA or DFA.
- Nasopharyngeal aspirate for culture in infants <6 months of age.
- NAATs are not approved for use with rectal or oropharyngeal samples.
- For information on specimen transport, see *Laboratory Diagnosis of Sexually Transmitted Infections* chapter.

Management

- Evaluation should be appropriate for the presenting symptoms, signs and sexual history.
- Treatment for chlamydia is indicated for the following:
 - A positive chlamydia test.
 - Diagnosis of a syndrome compatible with a chlamydial infection, without waiting for the test results of *C trachomatis*.
 - Diagnosis of chlamydial infection in a sexual partner.
 - Empirical co-treatment when a diagnosis of N gonorrhoeae is made without waiting for test results of C trachomatis due to the significant probability of co-infection (20–42%)^{12,13} and the possibility of false-negative results, especially with non-NAAT methods.

Treatment

- Efficacy and use-effectiveness studies evaluating single-dose azithromycin and a 7-day course of doxycycline have demonstrated similarly high cure rates; azithromycin is much more expensive.³⁴⁻⁴³
- Ofloxacin has an efficacy similar to doxycycline and azithromycin, but it is more expensive and must be taken as a multiple-dose course.^{44–52}
- Erythromycin is associated with significantly higher gastrointestinal side effects than other regimens.⁵²⁻⁵⁶
- Drug resistance is rare but may become an emerging issue. ^{57,58}
- In the absence of a contraindication, the following treatment options are recommended.

Adults (non-pregnant and non-lactating): urethral, endocervical, rectal, conjunctival infection

(For pelvic inflammatory disease, see *Pelvic Inflammatory Disease* chapter; for epididymitis, see *Epididymitis* chapter.)

Table 3. Adults (non-pregnant and non-lactating): urethral, endocervical, rectal, conjunctival infection

Preferred	Alternative
 Doxycycline 100 mg P0 bid for 7 days [A-I] OR Azithromycin 1 g P0 in a single dose if poor compliance is expected* [A-I] 	 Ofloxacin 300 mg PO bid for 7 days [B-II] OR Erythromycin 2 g/day PO in divided doses for 7 days[†][B-II] OR Erythromycin 1g/day PO in divided doses for 14 days[†][B-I]

^{*} If vomiting occurs more than 1 hour post-administration, a repeat dose is not required.

Children

- Topical therapy alone for conjunctivitis is NOT adequate and is unnecessary when systemic treatment is used.
- The use of erythromycin in infants under 6 weeks of age has been associated with infantile hypertrophic pyloric stenosis (IHPS). ⁵⁹⁻⁶² The risk of IHPS with other macrolides (e.g., azithromycin, clarithromycin) is unknown. The risks and benefits of using erythromycin in such infants must be explained to parents. When erythromycin is used in such infants, it is important to monitor for signs and symptoms of IHPS. IHPS following erythromycin use should be reported to the Canadian Adverse Drug Reaction Monitoring Program at 1-866-234-2345.
- The need to treat infants under 6 weeks for *C trachomatis* can be avoided by screening pregnant women and treating before delivery.
- Doxycycline is contraindicated in children under 9 years of age.
- Quinolones have been associated with articular damage in young animals. Such
 joint changes have not been clearly attributable to quinolone use in children. Its
 safety in children has not been established. Quinolones should not be used in
 prepubertal patients. Experience in pubertal patients under 18 years of age
 is limited.

[†] Erythromycin dosages refer to erythromycin base. Equivalent dosages of other formulations may be substituted (with the exception of the estolate formulation which is contraindicated in pregnancy). If erythromycin has been used for treatment, a test of cure should be performed 3–4 weeks after completion of therapy.

Table 4. Children

First week of life	>1 week to 1 month	>1 month to <9 years	9-18 years
Infants ≤ 2000 g • Erythromycin 20 mg/kg/day PO in divided doses for at least 14 days*† [B-II] Infants > 2000 g • Erythromycin 30 mg/kg/day PO in divided doses for at least 14 days*† [B-II]	• Erythromycin 40 mg/kg/day PO in divided doses for at least 14 days*† [B-II]	 Azithromycin 12–15 mg/kg (max. 1 g) PO in a single dose [B-II] Alternatives Erythromycin 40 mg/kg/day PO in divided doses (max. 500 mg qid for 7 days or 250 mg qid for 14 days)*† [B-II] OR Sulfamethoxazole 75 mg/kg/day PO in divided doses (max. 1 g bid) for 10 days† [B-II] 	Preferred Doxycycline Smg/kg/day PO in divided doses (max. 100 mg bid) for 7 days [A-I] OR Azithromycin 12–15 mg/kg (max. 1 g) PO in a single dose if poor compliance is expected [A-I] Alternatives Erythromycin 40 mg/kg/day PO in divided doses (max. 500 mg qid for 7 days or 250 mg qid for 14 days)*† [B-I] OR Sulfamethoxazole 75 mg/kg/day PO in divided doses (max. 1 g bid) for 10 days† [B-II]

^{*} Erythromycin dosages refer to the use of erythromycin base. Equivalent dosages of other formulations may be substituted (with the exception of the estolate formulation, which is contraindicated in pregnancy).

[†] If erythromycin or sulfamethoxazole has been used for treatment, repeat testing after completion of therapy is advisable.

[•] Neonates born to infected mothers must be tested for *C trachomatis*. Neonates should be treated if their test results are positive. They should be closely monitored for signs of chlamydial infection (e.g., conjunctivitis, pneumonitis). Prophylaxis is not recommended unless follow-up cannot be guaranteed.

[•] Test of cure should be performed 3–4 weeks after the completion of treatment in all prepubertal children.

Pregnant women and nursing mothers: urethral, endocervical, rectal infection

- Clinical trials comparing amoxicillin, erythromycin and azithromycin have demonstrated similar microbiological and clinical cure, but maternal gastrointestinal side effects are more common with erythromycin.⁶³⁻⁷¹
- To date, there are limited data collected on azithromycin in pregnancy, but it is considered to be safe in this context by many experts.^{64-66,68-70}
- Doxycycline and quinolones are contraindicated in pregnancy and in lactating women.
- Clindamycin requires dosing three to four times a day for 10–14 days and does
 not offer any advantage. In addition, it is even more expensive than azithromycin
 and is thus not being listed as an option.
- Data on neonatal outcomes are limited.

Table 5. Pregnant women and nursing mothers: urethral, endocervical, rectal infection

• Amoxicillin 500 mg PO tid for 7 days* [B-I]

OR

Erythromycin 2 g/day PO in divided doses for 7 days*† [B-I]

OR

- Erythromycin 1g/day PO in divided doses for 14 days*[†] [B-I]
- Azithromycin 1 g PO in a single dose, if poor compliance is expected[‡] [B-I]
- * If erythromycin or amoxicillin has been used for treatment in nursing mothers, test of cure should be performed 3–4 weeks after the completion of treatment.
- † Erythromycin dosage refers to the use of erythromycin base. Equivalent dosages of other formulations may be substituted (with the exception of the estolate formulation, which is contraindicated in pregnancy). Gastrointestinal side effects are more severe with erythromycin than with amoxicillin.
- ‡ If vomiting occurs more than 1 hour post-administration, a repeat dose is not required.

Note: Test of cure should be performed 3-4 weeks after the completion of treatment in all pregnant women.

Considerations for Other STIs

- See Primary Care and Sexually Transmitted Infections chapter.
- Obtain specimen(s) for the diagnosis of *N gonorrhoeae*.
- Obtain a blood sample for serologic testing for syphilis (see Syphilis chapter).
- HIV testing and counselling are recommended (see *Human Immunodeficiency Virus Infections* chapter).
- Immunization against hepatitis B is recommended in non-immune nonimmunized individuals (see Hepatitis B Virus Infections chapter).

Reporting and Partner Notification

- *C trachomatis* infections must be reported by laboratories and physicians to local public health authorities in all provinces and territories.
- All partners who have had sexual contact with the index case within 60 days
 prior to symptom onset or date of diagnosis where asymptomatic should be
 tested and treated. If there was no partner during this period, then the last
 partner should be tested and treated.
- Parents of infected neonates (i.e., mother and her sexual partner[s]) and persons
 implicated in sexual abuse cases must be located, clinically evaluated and
 treated.
- Local public health authorities are available to assist with partner notification and help with appropriate referral for clinical evaluation, testing, treatment and health education. If resources for local public health authority support are limited, priority for partner notification should be directed toward youth/young adults
 years of age.

Follow-up

- Test of cure for C trachomatis is not routinely indicated if a recommended treatment is taken AND symptoms and signs disappear AND there is no re-exposure to an untreated partner except:
 - Where compliance is suboptimal.
 - If an alternative treatment regimen has been used.
 - In all prepubertal children.
 - In all pregnant women.
- Test of cure using a NAAT, if needed, should be performed at 3–4 weeks after the completion of effective treatment to avoid false-positive results due to the presence of non-viable organisms.
- Repeat testing in all individuals with *C trachomatis* infection is recommended 6 months post-treatment, as reinfection risk is high.
- In patients with apparent treatment failure, possibilities include the following:
 - Failure to take medication correctly or to finish course of therapy.
 - Re-exposure to an untreated partner.
 - Infection acquired from a new partner.
 - A false-positive result.
 - Rarely, resistance is an issue.
- In patients with persistent symptoms, infection with other pathogens and a non-infective etiology should also be considered.

Special Considerations

Children

- Neonates born to infected mothers MUST be tested for C trachomatis. Neonates should be treated if test results are positive. They should be closely monitored for signs of chlamydial infection (e.g., conjunctivitis, pneumonitis). Prophylaxis is not recommended unless follow-up cannot be guaranteed.
- Sexual abuse must be considered when genital, rectal or pharyngeal chlamydial
 infection is diagnosed in any prepubertal child, although perinatally acquired
 C trachomatis can persist in an infant for up to 3 years. Consultation with a
 colleague experienced in such cases should be sought. Siblings and other
 children possibly at risk must also be evaluated.
- Sexual abuse of children must be reported to the local child protection agency (see Sexual Abuse in Peripubertal and Prepubertal Children chapter).
- Follow-up cultures for "test of cure" are indicated approximately 3–4 weeks after completion of therapy in prepubertal children.

References

- Patrick DM, Wong T, Jordan R. Sexually transmitted infections in Canada: recent resurgence threatens national goals. *Can J Hum Sexuality* 2000;9: 149–165.
- Surveillance and Epidemiology Section, Community Acquired Infections Division, Public Health Agency of Canada, unpublished data, 2006.
- 3. Farley TA, Cohen DA, Elkins W. Asymptomatic sexually transmitted diseases: the case for screening. *Prev Med* 2003;36:502–509.
- 4. Stamm WE, Koutsky LA, Benedetti JK, Jourden JL, Brunham RC, Holmes KK. Chlamydia trachomatis urethral infections in men. Prevalence, risk factors, and clinical manifestations. *Ann Intern Med* 1984;100:47–51.
- 5. Stamm WE. Expanding efforts to prevent chlamydial infection. *N Engl J Med* 1998;339:768–770.
- 6. Gaydos CA, Howell MR, Pare B, et al. Chlamydia trachomatis infections in female military recruits. *N Engl J Med* 1998;339:739–744.
- Marrazzo JM, White CL, Krekeler B, et al. Community-based urine screening for Chlamydia trachomatis with a ligase chain reaction assay. *Ann Intern Med* 1997;127:796–803.
- 8. Marrazzo JM, Whittington WL, Celum CL, et al. Urine-based screening for Chlamydia trachomatis in men attending sexually transmitted disease clinics. Sex Transm Dis 2001;28:219–225.
- 9. Chen MY, Donovan B. Screening for genital Chlamydia trachomatis infection: are men the forgotten reservoir? *Med J Aust* 2003;179:124–125.
- Andersen B, Olesen F, Moller JK, Ostergaard L. Population-based strategies for outreach screening of urogenital Chlamydia trachomatis infections: a randomized, controlled trial. *J Infect Dis* 2002;185:252–258.

- Ginocchio RH, Veenstra DL, Connell FA, Marrazzo JM. The clinical and economic consequences of screening young men for genital chlamydial infection. Sex Transm Dis 2003;30:99–106.
- 12. Creighton S, Tenant-Flowers M, Taylor CB, Miller R, Low N. Co-infection with gonorrhoea and chlamydia: how much is there and what does it mean? *Int J STD AIDS* 2003;14:109–113.
- Lyss SB, Kamb ML, Peterman TA, et al; Project RESPECT Study Group. Chlamydia trachomatis among patients infected with and treated for Neisseria gonorrhoeae in sexually transmitted disease clinics in the United States. *Ann Intern Med* 2003;139:178–185.
- 14. Braverman PK, Biro FM, Brunner RL, Gilchrist MJ, Rauh JL. Screening asymptomatic adolescent males for chlamydia. *J Adolesc Health Care* 1990;11:141–144.
- Chernesky MA, Jang D, Lee H, et al. Diagnosis of Chlamydia trachomatis infections in men and women by testing first-void urine by ligase chain reaction. *J Clin Microbiol* 1994;32:2682–2685.
- 16. LaMontagne DS, Fine DN, Marrazzo JM. Chlamydia trachomatis infection in asymptomatic men. *Am J Prev Med* 2003;24:36–42.
- Marrazzo JM, Celum CL, Hillis SD, Fine D, DeLisle S, Handsfield HH.
 Performance and cost-effectiveness of selective screening criteria for
 Chlamydia trachomatis infection in women. Implications for a national
 Chlamydia control strategy. Sex Transm Dis 1997;24:131–141.
- 18. Moncada J, Schachter J, Shafer MA, et al. Detection of Chlamydia trachomatis in first catch urine samples from symptomatic and asymptomatic males. *Sex Transm Dis* 1994;21:8–12.
- Domeika M, Bassiri M, Mardh PA. Diagnosis of genital Chlamydia trachomatis infections in asymptomatic males by testing urine by PCR. *J Clin Microbiol* 1994;32:2350–2352.
- 20. Anestad G, Berdal BP, Scheel O, et al. Screening urine samples by leukocyte esterase test and ligase chain reaction for chlamydial infections among asymptomatic men. *J Clin Microbiol* 1995;33:2483–2484.
- 21. Ciemins EL, Kent CK, Flood J, Klausner JD. Evaluation of chlamydia and gonorrhea screening criteria: San Francisco sexually transmitted disease clinic: 1997 to 1998. *Sex Transm Dis* 2000;27:165–167.
- 22. Davies HD, Wang EE. Periodic health examination, 1996 update: 2. Screening for chlamydial infections. Canadian Task Force on the Periodic Health Examination. *CMAJ* 1996;154:1631–1644.
- 23. Sexually transmitted diseases treatment guidelines 2002. Centers for Disease Control and Prevention. *MMWR Recomm Rep* 2002;51(RR-6):1–78.
- 24. Cohen I, Veille JC, Calkins B. Improved pregnancy outcome following successful treatment of chlamydial infection. *JAMA* 1990;263:3160–3163.
- Ryan GM Jr, Abdella TN, McNeeley SG, Baselski VS, Drummond DE. Chlamydia trachomatis infection in pregnancy and effect of treatment on outcome. Am J Obstet Gynecol 1990;162:34–39.

- Black-Payne C, Ahrabi MM, Bocchini JA Jr, Ridenour CR, Brouillette RM.
 Treatment of Chlamydia trachomatis identified with Chlamydiazyme during pregnancy. Impact on perinatal complications and infants. *J Reprod Med* 1990:35:362–367.
- 27. Schachter J, Sweet RL, Grossman M, Landers D, Robbie M, Bishop E. Experience with the routine use of erythromycin for chlamydial infections in pregnancy. *N Engl J Med* 1986;314:276–279.
- 28. McMillan JA, Weiner LB, Lamberson HV, et al. Efficacy of maternal screening and therapy in the prevention of chlamydia infection of the newborn. *Infection* 1985;13:263–266.
- Whittington WL, Kent C, Kissinger P, et al. Determinants of persistent and recurrent Chlamydia trachomatis infection in young women: results of a multicenter cohort study. Sex Transm Dis 2001;28:117–123.
- 30. Schillinger JA, Kissinger P, Calvet H, et al. Patient-delivered partner treatment with azithromycin to prevent repeated Chlamydia trachomatis infection among women: a randomized, controlled trial. *Sex Transm Dis* 2003;30:49–56.
- 31. Gunn RA, Fitzgerald S, Aral SO. Sexually transmitted disease clinic clients at risk for subsequent gonorrhea and chlamydia infections: possible "core" transmitters. Sex Transm Dis 2000;27:343–349.
- 32. Rietmeijer CA, Van Bemmelen R, Judson FN, Douglas JM Jr. Incidence and repeat infection rates of Chlamydia trachomatis among male and female patients in an STD clinic: implications for screening and rescreening. *Sex Transm Dis* 2002;29:65–72.
- Korenromp EL, Sudaryo MK, de Vlas SJ, et al. What proportion of episodes of gonorrhoea and chlamydia becomes symptomatic? *Int J STD AIDS* 2002;13:91–101.
- 34. Hillis SD, Coles FB, Litchfield B, et al. Doxycycline and azithromycin for prevention of chlamydial persistence or recurrence one month after treatment in women. A use-effectiveness study in public health settings. *Sex Transm Dis* 1998;25:5–11.
- 35. Hammerschlag MR, Golden NH, Oh MK, et al. Single dose of azithromycin for the treatment of genital chlamydial infections in adolescents. *J Pediatr* 1993;122:961–965.
- Johnson RB. The role of azalide antibiotics in the treatment of Chlamydia.
 Am J Obstet Gynecol 1991;164(6 Pt 2):1794–1796.
- 37. Marra F, Marra C, Patrick DM. Cost-effectiveness analysis of azithromycin for Chlamydia trachomatis infection in women: a Canadian perspective. *Can J Infect Dis* 1997;8:202–208.
- Martin DH, Mroczkowski TF, Dalu ZA, et al. A controlled trial of a single dose of azithromycin for the treatment of chlamydial urethritis and cervicitis. The Azithromycin for Chlamydial Infections Study Group. N Engl J Med 1992;327:921–925.
- 39. Nilsen A, Halsos A, Johansen A, et al. A double blind study of single dose azithromycin and doxycycline in the treatment of chlamydial urethritis in males. *Genitourin Med* 1992;68:325–327.

- 40. Nuovo J, Melnikow J, Paliescheskey M, King J, Mowers R. Cost-effectiveness analysis of five different antibiotic regimens for the treatment of uncomplicated Chlamydia trachomatis cervicitis. *J Am Board Fam Pract* 1995;8:7–16.
- Ossewaarde JM, Plantema FHF, Rieffe M, Nawrocki RP, De Vries A, van Loon AM. Efficacy of single-dose azithromycin versus doxycycline in the treatment of cervical infections caused by Chlamydia trachomatis. *Eur J Clin Microbiol Infect Dis* 1992;11:693–697.
- Thorpe EM Jr, Stamm WE, Hook EW 3rd, et al. Chlamydial cervicitis and urethritis: single dose treatment compared with doxycycline for seven days in community based practises. *Genitourin Med* 1996;72:93–97.
- Lau CY, Qureshi AK. Azithromycin versus doxycycline for genital chlamydial infections: a meta-analysis of randomized clinical trials. Sex Transm Dis 2002:29:497–502.
- 44. Judson FN, Beals BS, Tack KJ. Clinical experience with ofloxacin in sexually transmitted disease. *Infection* 1986;14(suppl 4):S309–S310.
- 45. Fransen L, Avonts D, Piot P. Treatment of genital chlamydial infection with ofloxacin. *Infection* 1986;14(suppl 4):S318–S320.
- Batteiger BE, Jones RB, White A. Efficacy and safety of ofloxacin in the treatment of nongonococcal sexually transmitted disease. Am J Med 1989;87(6C):75S-77S.
- Nayagam AT, Ridgway GL, Oriel JD. Efficacy of ofloxacin in the treatment of non-gonococcal urethritis in men and genital infections caused by Chlamydia trachomatis in men and women. *J Antimicrob Chemother* 1988;22(suppl C): 155–158.
- 48. Maiti H, Chowdhury FH, Richmond SJ, et al. Ofloxacin in the treatment of uncomplicated gonorrhea and chlamydial genital infection. *Clin Ther* 1991:13:441–447.
- Faro S, Martens MG, Maccato M, Hammill HA, Roberts S, Riddle G. Effectiveness of ofloxacin in the treatment of Chlamydia trachomatis and Neisseria gonorrhoeae cervical infection. *Am J Obstet Gynecol* 1991;164 (5 Pt 2):1380–1383.
- 50. Hooton TM, Batteiger BE, Judson FN, Spruance SL, Stamm WE. Ofloxacin versus doxycycline for treatment of cervical infection with Chlamydia trachomatis. *Antimicrob Agents Chemother* 1992;36:1144–1146.
- Kitchen VS, Donegan C, Ward H, Thomas B, Harris JR, Taylor-Robinson D. Comparison of ofloxacin with doxycycline in the treatment of non-gonococcal urethritis and cervical chlamydial infection. *J Antimicrob Chemother* 1990;26(suppl D):99–105.
- 52. Mogabgab WJ, Holmes B, Murray M, Beville R, Lutz FB, Tack KJ. Randomized comparison of ofloxacin and doxycycline for chlamydia and ureaplasma urethritis and cervicitis. *Chemotherapy* 1990;36:70–76.
- 53. Linnemann CC Jr, Heaton CL, Ritchey M. Treatment of Chlamydia trachomatis infections: comparison of 1- and 2-g doses of erythromycin daily for seven days. *Sex Transm Dis* 1987;14:102–106.

- 54. Cramers M, Kaspersen P, From E, Moller BR. Pivampicillin compared with erythromycin for treating women with genital Chlamydia trachomatis infection. *Genitourin Med* 1988;64:247–248.
- 55. Scheibel JH, Kristensen JK, Hentzer B, et al. Treatment of chlamydial urethritis in men and Chlamydia trachomatis-positive female partners: comparison of erythromycin and tetracycline in treatment courses of one week. *Sex Transm Dis* 1982;9:128–131.
- 56. Bowie WR, Manzon LM, Borrie-Hume CJ, Fawcett A, Jones HD. Efficacy of treatment regimens for lower urogenital Chlamydia trachomatis infection in women. *Am J Obstet Gynecol* 1982;142:125–129.
- 57. Somani J, Bhullar VB, Workowski KA, Farshy CE, Black CM. Multiple drugresistant Chlamydia trachomatis associated with clinical treatment failure. *J Infect Dis* 2000:181:1421–1427.
- 58. Misyurina OY, Chipitsyna EV, Finashutina YP, et al. Mutations in a 23S rRNA gene of Chlamydia trachomatis associated with resistance to macrolides. Antimicrob Agents Chemother 2004;48:1347–1349.
- Sorensen HT, Skriver MV, Pedersen L, Larsen H, Ebbesen F, Schonheyder HC. Risk of infantile hypertrophic pyloric stenosis after maternal postnatal use of macrolides. Scand J Infect Dis 2003;35:104–106.
- Cooper WO, Griffin MR, Arbogast P, Hickson GB, Gautam S, Ray WA. Very early exposure to erythromycin and infantile hypertrophic pyloric stenosis. *Arch Pediatr Adolesc Med* 2002;156:647–650.
- 61. Mahon BE, Rosenman MB, Kleiman MB. Maternal and infant use of erythromycin and other macrolide antibiotics as risk factors for infantile hypertrophic pyloric stenosis. *J Pediatr* 2001;139:380–384.
- 62. Honein MA, Paulozzi LJ, Himelright IM, et al. Infantile hypertrophic pyloric stenosis after pertussis prophylaxis with erythromcyin: a case review and cohort study. *Lancet* 1999;354:2101–2105.
- 63. Magat AH, Alger LS, Nagey DA, Hatch V, Lovchik JC. Double-blind randomized study comparing amoxicillin and erythromycin for the treatment of Chlamydia trachomatis in pregnancy. *Obstet Gynecol* 1993;81(5 Pt 1):745–749.
- 64. Kacmar J, Cheh E, Montagno A, Peipert JF. A randomized trial of azithromycin versus amoxicillin for the treatment of Chlamydia trachomatis in pregnancy. *Infect Dis Obstet Gynecol* 2001;9:197–202.
- Wehbeh HA, Ruggeirio RM, Shahem S, Lopez G, Ali Y. Single-dose azithromycin for Chlamydia in pregnant women. *J Reprod Med* 1998;43: 509–514.
- 66. Adair CD, Gunter M, Stovall TG, McElroy G, Veille JC, Ernest JM. Chlamydia in pregnancy: a randomized trial of azithromycin and erythromycin. *Obstet Gynecol* 1998;91:165–168.
- 67. Alary M, Joly JR, Moutquin JM, et al. Randomised comparison of amoxycillin and erythromycin in treatment of genital chlamydial infection in pregnancy. *Lancet* 1994;344:1461–1465.
- 68. Bush MR, Rosa C. Azithromycin and erythromycin in the treatment of cervical chlamydial infection during pregnancy. *Obstet Gynecol* 1994;84:61–63.

- 69. Genc MR. Treatment of genital Chlamydia trachomatis infection in pregnancy. Best Pract Res Clin Obstet Gynaecol 2002;16:913–922.
- Jacobson GF, Autry AM, Kirby RS, Liverman EM, Motley RU. A randomized controlled trial comparing amoxicillin and azithromycin for the treatment of Chlamydia trachomatis in pregnancy. *Am J Obstet Gynecol* 2001;184: 1352–1354.
- 71. Silverman NS, Sullivan M, Hochman M, Womack M, Jungkind DL. A randomized, prospective trial comparing amoxicillin and erythromycin for the treatment of Chlamydia trachomatis in pregnancy. *Am J Obstet Gynecol* 1994;170:829–831.

ECTOPARASITIC INFESTATIONS (PUBIC LICE, SCABIES)

Pubic Lice

Etiology/Epidemiology

- · Caused by Phthirus pubis (crab louse).
- · Humans are the only reservoir.
- Shorter life span off host (24 hours) than head lice (several days).
- Usually present in pubic hair, but may also be found in chest, armpits, eyelashes
 or facial hair.
- Transmission occurs through intimate sexual and non-sexual contact.¹

Prevention

- Patients presenting with concerns about sexually transmitted infections (STIs) and/or prevention of pregnancy should be provided with instructions and encouragement about the consistent practice of safer-sex.
- At the time of diagnosis, review and monitor prevention practices.
- Identify barriers to prevention practices and the means to overcome them.
- See Primary Care and Sexually Transmitted Infections chapter.

Manifestations²

- Itching, scratching, erythema, skin irritation and inflammation, all as a reaction to the louse bite.
- Small blue spots can appear where the louse has bitten.
- Extensive infestation can be associated with mild fever and malaise.
- Scratching can lead to a secondary bacterial skin infection.

Diagnosis

- Based on history and index of suspicion.
- Careful examination for adult lice and eggs (nits). Look for an area of scabs with nits in the hair; scabs may be adult lice. Nits attach to hair and are not loose and flaky.

Specimen collection and laboratory diagnosis

If necessary, submit nits or scabs in a container for microscopic examination.

Management

- Clothes, bedding and fomites: washing in hot water (50°C) or dry cleaning kills all stages of lice. Alternatively, place in plastic bags for 1 week.
- Vacuum mattresses.
- Sexual partner(s) within the last month should be treated.
- May re-treat after 1 week if no clinical improvement. Pruritus may be controlled with antihistamines such as hydroxyzine or diphenhydramine, as well as mild topical corticosteroids.²

Treatment

- Wash the affected area and apply pediculocide formulation (cream, lotion or shampoo) according to package instructions.
 - Permethrin 1% cream [A-I]

OR

- 0.33% pyrethrin-piperonyl butoxide shampoo [A-I]

OR

- lindane 1% shampoo [A-I].^{2,3}
- May repeat in 3–7 days.

Special Considerations

- Pediculosis of the eyelashes should not be treated with permethrin, pyrethrin
 or lindane.² Recommended treatment: occlusive ophthalmic ointment to the
 eyelidmargins bid for 10 days.
- Gamma benzene hexachloride (lindane) can cause neurotoxicity. Instructions
 for use must be carefully followed to minimize risk of toxicity.³ Contraindicated
 in children <2 years of age, in pregnancy, in lactating women or in patients with
 extensive dermatitis.
- Permethrin cream has efficacy similar to lindane 1%, with less toxicity and cure rates greater than 80%.³
- Pruritus may persist for several days or weeks after treatment.
- In patients with excoriated or damaged skin, consider dose modification to compensate for increased absorption of topical agents.

(See below for Consideration for Other STIs, Reporting and Partner Notification, and Follow-up.)

Scables

Etiology/Epidemiology

- Caused by Sarcoptes scabiei.
- Incubation period is 3 weeks, but reinfestation provokes immediate symptoms (1–3 days).¹
- Transmission:
 - Often non-sexual, through close person-to-person contact (e.g., in families and institutions).⁴
 - May be via shared personal articles (clothes, bedding).
 - Sexual transmission does occur; usually need more than brief contact.

Prevention

- Patients presenting with concerns about STIs and/or prevention of pregnancy should be provided with instructions and encouragement about the consistent practice of safer-sex.
- At the time of diagnosis, review and monitor prevention practices.
- Identify barriers to prevention practices and the means to overcome them.
- See Primary Care and Sexually Transmitted Infections chapter.

Manifestations

- Intense nocturnal itching.
- · Burrows under the skin.
- Lesions affecting hands (finger webs, sides of digits), flexor surfaces of the wrists, axillae, waist, nipple areola, periumbilical area and male genitalia.
- Papules or nodules, which result from itching, often affect the genital area.
- Pyoderma of the penis.
- HIV-infected patients may present atypically with crusted or "exaggerated" scabies called Norwegian scabies.⁶

Diagnosis

- Based on history, index of suspicion and examination.
- Diagnosis is often difficult and therefore delayed.

Specimen collection and laboratory diagnosis

- If necessary, take a skin scraping of a burrow to remove the mite or ova for microscopic examination.¹
- Burrow ink test: apply fountain pen ink or a washable marker to outside
 of burrow, wipe skin (with alcohol). Burrows will retain the ink and may be
 visualized.²

Management

- Clothes, bedding and fomites: washing in hot water (50°C) or dry cleaning kills all stages of the organism. Alternatively, place in plastic bags for 3 days to 1 week.¹
- · Vacuum mattresses.
- All household contacts and recent sexual partner(s) in the last month should be treated.
- Pruritus may persist for several weeks. Pruritus may be controlled with antihistamines and mild topical corticosteroids.

Treatment

- Permethrin 5% cream [A-I].2,3,7
 - Apply to the body from the neck down; leave for 8–14 hours; shower and wear clean clothes.

OR

- Gamma benzene hexachloride (lindane) 1% cream or lotion [A-I]. 2,3,7,8
 - Apply to the body from the neck down; leave for 8 hours; shower and wear clean clothes.
 - More potential for toxicity than permethrin.
 - Contraindicated in children <2 years of age, in pregnancy, in lactating women or in patients with extensive dermatitis.
- · Alternatives:
 - Crotamiton 10% cream [A-I] (less effective than permethrin or lindane).^{7,9}
 This product is available through the Health Canada Special Drug Access Program.
 - Apply nightly for two nights and wash off thoroughly 24 hours after last application.

OR

- Sulphur 5% in petroleum [A-I] (less effective than permethrin or lindane).^{7,9}
- Apply nightly for three nights and wash off thoroughly 24 hours after last application.

Special Considerations

- In pregnancy, permethrin is the only agent that should be used.²
- Gamma benzene hexachloride (lindane) can cause neurotoxicity. Instructions
 for use must be carefully followed to minimize risk of toxicity.³ Contraindicated
 in children <2 years of age, in pregnancy, in lactating women or in patients with
 extensive dermatitis.
- In patients with excoriated or damaged skin, consider dose modification to compensate for increased absorption of topical agents.

Consideration for Other STIs

- See Primary Care and Sexually Transmitted Infections chapter.
- Obtain a specimen for the diagnosis of Chlamydia trachomatis and Neisseria gonorrhoeae.
- Obtain a blood sample for serologic testing for syphilis (see *Syphilis* chapter).
- HIV counselling and testing are recommended (see Human Immunodeficiency Virus Infections chapter).
- Immunization against hepatitis B is recommended, unless already immune (see Hepatitis B Virus Infections chapter).

Reporting and Partner Notification

- Pubic lice and scabies are not reportable to local public health authorities.
- · Partner notification of ectoparasitic infestations is not required.

Follow-up

· Follow-up only if clinically necessary.

References

- 1. Chosidow O. Scabies and pediculosis. *Lancet* 2000;355:819–826.
- 2. Wendel K, Rompalo A. Scabies and pediculosis pubis: an update of treatment regimens and general review. *Clin Infect Dis* 2002;35(suppl 2):S146–S151.
- 3. Roos TC, Alam M, Roos S, Merk HF, Bickers DR. Pharmacotherapy of ectoparasitic infections. *Drugs* 2001;61:1067–1088.
- Hogan DJ, Schachner L, Tanglertsampan C. Diagnosis and treatment of childhood scabies and pediculosis. *Pediatr Clin North Am* 1991;38:941–957.
- 5. Burkhart CG, Burkhart CN, Burkhart KM. An epidemiologic and therapeutic reassessment of scabies. *Cutis* 2000;65:233–240.
- 6. Orkin M. Scabies in AIDS. Semin Dermatol 1993:12:9-14.
- 7. Scott GR. European guideline for the management of scabies. *Int J STD AIDS* 2001;12(suppl 3):58–61.
- 8. Chouela EN, Abeldano AM, Pellerano G, et al. Equivalent therapeutic efficacy and safety of ivermectin and lindane in the treatment of human scabies. *Arch Dermatol* 1999;135:651–655.
- 9. Morgon-Glenn PD. Scabies. *Pediatr Rev* 2001;22:322–323.

GENITAL HERPES SIMPLEX VIRUS (HSV) INFECTIONS

Etiology

Herpes simplex virus (HSV) types 1 and 2.1

Epidemiology

- The annual incidence in Canada of genital herpes due to HSV-1 and -2 infection is not known (for a review of HSV-1/HSV-2 prevalence and incidence studies worldwide, see Smith and Robinson 2002²). In the United States, it is estimated that about 1,640,000 HSV-2 seroconversions occur yearly (730,000 men and 910,000 women, or 8.4 per 1,000 persons).³
- Based on the change in prevalence of the serum antibody to HSV-2, HSV-2 increased 30% between 1976 and 1994, from 16.4–21.9% in Americans aged 12 years and older.⁴
- In British Columbia in 1999, the seroprevalence of HSV-2 antibody in leftover serum submitted for antenatal testing revealed a prevalence of 17.3%, ranging from 7.1% in women 15–19 years old to 28.2% in those 40–44 years.⁵
- In attendees at an Alberta sexually transmitted infection (STI) clinic in 1994 and 1995, the seroprevalence of HSV-1 and -2 in leftover sera was 56% and 19%, respectively.⁶
- The incidence and prevalence of HSV-1 genital infection is increasing globally, with marked variation between countries.⁷
- In Norway, a recent study found that 90% of genital initial infections were due to HSV-1.8
- In Nova Scotia, 58.1% of 1,790 HSV isolates from genital lesion cultures in women were HSV-1; in men, 36.7% of 468 isolates were HSV-1.9
- Females are at higher risk of acquiring genital herpes from a male partner than males are from a female partner. Studies have found that among discordant heterosexual couples with a source partner who had symptomatic recurrent genital HSV-2 infection, the annual transmission rates were 11–17% in couples with male source partners and 3–4% in couples with female source partners.^{10,11}
- In one study, transmission in 70% of patients appeared to result from sexual contact during periods of asymptomatic virus shedding.¹¹
- Pre-existing seropositivity to HSV-1 reduced the likelihood of acquiring symptomatic genital HSV-2 disease in women by 55–74%,^{11,12} although others have not observed such a protective effect.^{10,13}

Natural history

- The incubation period averages 6 days.¹
- Of new HSV-2 infections diagnosed by seroconversion, approximately 60% are asymptomatic and 40% symptomatic. Of the symptomatic cohort, about 80% present with typical genital symptoms and signs, while 20% have atypical presentations, including nonlesional HSV-2 infections such as genital pain

- or urethritis, aseptic meningitis and cervicitis, which are well-recognized complications of first episodes of genital HSV infection.¹
- No intervention, including early initiation of antiviral therapy, prevents the development of latent sacral sensory ganglion infection.¹⁴
- Recurrences tend to occur in tissues innervated by sacral sensory nerves.
- Recurrences may be preceded by warning signs (prodromal symptoms) a few minutes to several days before lesions appear, such as focal burning, itching (most common), tingling or vague discomfort.¹⁵
- Recurrences may be associated with the menstrual cycle, emotional stress, illness (especially with fever), sexual intercourse, surgery and certain medication
 — so-called "trigger factors." 15
- Initial mean recurrence rates are greater in persons with genital HSV-2 infection than in those with HSV-1: 4% and 1% per year, respectively, with marked interindividual variation.¹⁶
- The average recurrence rate decreases over time by around 0.8 outbreaks per year, every year (no matter how high the initial outbreak rate was). However, approximately 25% of patients reported more recurrences in year 5 than year 1, evidence again of the substantial interindividual differences in recurrence rates.¹⁷
- Asymptomatic shedding of HSV can be demonstrated by virus identification through culture or polymerase chain reaction (PCR). HSV DNA can be detected four to five times more frequently by PCR than by culture.^{18,19} However, identification of virus by PCR may not be synonymous with infectivity. The following data pertain to shedding demonstrated by isolation of infectious virus:
 - Asymptomatic shedding prevalence is greater in women with HSV-2 genital infection than with HSV-1 (55% vs 29% during a median follow-up of 105 days).¹⁸ A similar difference may exist in men.¹⁹
 - Asymptomatic shedding of HSV-2 is as common in persons with symptomatic genital infection (while in between outbreaks) as in those with asymptomatic genital infection.¹⁸⁻²⁰
 - Asymptomatic shedding occurs on an average of 2% of days for a mean duration of 1.5 days.^{18,19} HSV has been isolated from vulva, cervicovaginal and rectal sites in women²⁰ and from penile and perianal skin, urethra and urine in men.¹⁹

Prevention

- Patients presenting with concerns about STIs and/or prevention of pregnancy provide clinicians with an important opportunity for instruction and encouragement about consistent safer-sex practices. Given the increase in HSV-1 genital infection, likely due to orogenital sex (perhaps as an alternative to genital intercourse), patients need also to be advised of the inherent risk of genital herpes from such an activity.²¹
- At the time of diagnosis of an STI, review and monitor prevention practices.
- Identify barriers to prevention and the means to overcome them.
- Condom use reduces transmission of genital HSV-2 from infected men to women by 50% and may reduce transmission from infected women to men

- to a similar degree.²² However, condom effectiveness is greatly limited by non-use and may also be limited because of the location of lesions and the risk of transmission during orogenital sex. Other safer-sex practices should be discussed.
- Valacyclovir 500 mg ingested daily by a patient with genital HSV-2 infection has been shown to reduce transmission to a susceptible heterosexual partner by 48%. The effect of condoms and suppressive valacyclovir may be additive.¹⁰
- Immunization with a glycoprotein D-adjuvanted vaccine has been demonstrated to protect against acquisition of genital HSV disease in women who were seronegative for both HSV-1 and -2, but not for those who were seropositive for HSV-1.²³ It had no protective efficacy in men, regardless of serostatus. Protection against genital HSV disease was 74%, and protection against infection (seroconversion plus symptomatic infection) was 46%. Practitioners should be aware that such a vaccine may become available for use in the next 5–10 years.

Manifestations

• A diagnostic lesion is a cluster of vesicles on an erythematous background.

Initial symptomatic episodes

- Primary
 - First clinically evident episode in an HSV-antibody-negative individual.
 - Five characteristics:¹
 - Extensive painful vesiculoulcerative genital lesions, including exocervix.
 - Systemic symptoms in 58–62% (fever, myalgia).
 - Tender lymphadenopathy in 80%.
 - Complications: 16–26% develop aseptic meningitis, and 10% to 28% develop extragenital lesions.
 - Protracted course: mean 16.5 (men) to 22.7 (women) days to resolve.
- Non-primary¹
 - First clinically evident episode in a person who, by testing, is demonstrated to have pre-existing heterologous antibody. Generally the range and severity of symptoms and signs of even the most severe cases are less marked than in those with severe primary infection. This has been attributed to a mitigating effect of pre-existing heterologous immunity in attenuating the severity of disease.
 - Compared to primary genital herpes, non-primary infections are characterized by the following:
 - · Less extensive genital lesions.
 - Systemic symptoms in only 16%.
 - Complications uncommon: meningitis in 1% and extragenital lesions in 8%.
 - Duration less prolonged: mean 15.5 days.

Recurrent disease^{1,24}

- The first clinically evident episode in a person with pre-existing homologous antibody (i.e., culture of HSV-2 from a first outbreak in an individual with demonstratable HSV-2 antibody) may sometimes be confused with a primary infection.²⁴ This is because overlap occurs in the frequency of local symptoms, fever and size of genital lesions between those with recently acquired genital herpes and those, who, by serologic testing, are determined to have acquired infection remotely but are now experiencing a first outbreak.²⁴
- In one study, almost 10% of patients judged to have a first-episode of genital herpes had serologic evidence of remotely acquired HSV-2 infection, indicating that clinical differentiation of primary genital infection and previously acquired infection can be difficult.
- Thus, typing of the virus isolate and type-specific serologic testing are required
 to differentiate between the two entities: primary/non-primary infection vs. a
 first lesion due to reactivation of a (long) latent infection acquired previously
 (see Diagnosis section, below).

Characteristics of recurrent disease

- Due to reactivation of latent sacral sensory ganglion infection.
- Typically, localized small painful genital lesions (mean lesion area 10% of that in primary genital herpes).¹
- Systemic symptoms in 5-12%.
- Prodromal symptoms in 43–53%, for an average of 1.2–1.5 days.
- Mean duration of lesion 9.3-10.6 days.

Asymptomatic shedding

See Natural history section, above.

Diagnosis

Specimen collection and laboratory diagnosis

- Culture is the most common method currently used in public health laboratories in Canada to confirm the clinical diagnosis of HSV. It is sensitive (70% from ulcers, 94% from vesicles) and permits identification of HSV type.²⁵
- PCR is four times more sensitive than HSV culture and is 100% specific.²⁶
 However, at this time, PCR assays have not yet replaced culture for routine
 diagnosis of genital herpes in public health laboratories in Canada.
- The Tzanck smear demonstrating diagnostic multinucleated giant cell is 40–68% as sensitive as culture, while direct fluorescent antibody has a sensitivity of 56% compared to culture.^{25,27} Neither test can thus be relied on for laboratory confirmation of diagnosis.
- The antibody response to primary infection is characterized by early appearance
 of IgM, followed subsequently by IgG antibody. IgM antibody usually wanes
 within a few months of infection;²⁸ therefore, the presence of IgM antibody is an
 indirect indication of "recent" infection.

- A primary infection is confirmed by demonstrating an absence of HSV antibody in the acute-phase sample and the presence of antibody in the convalescent blood sample (i.e., seroconversion).
- Most individuals seroconvert within 3–6 weeks; by 12 weeks, more than 70% will have seroconverted.^{29,30}
- The advent of testing for type-specific antibody will allow practitioners to establish a diagnosis of primary infection and determine whether the infection is due to HSV-1 or -2. Such information will also permit practitioners to counsel individuals with genital herpes and their partners. Type-specific antibody is best detected by Western blot analysis, although new commercial enzyme immunoassays with improved sensitivity and specificity are available.³¹ Enzyme immunoassay test results need not be routinely confirmed by Western blot analysis. At this time, type-specific HSV antibody assays are available only in a few laboratories in Canada (see Special Considerations section, below).
- During recurrent genital HSV infection, no consistent HSV antibody changes occur. Specifically, IgM appears inconsistently, and IgM titres also do not change between acute and convalescent samples.³²
- Detection of HSV-2 antibody is considered to be accurate for detecting silent genital HSV-2 infection, but detecting HSV-1 antibody is not useful in the same way, because asymptomatic HSV-1 orolabial infection is common.³¹

Management

- Counselling is an important component in management. Genital HSV infection is not curable, but its somatic and psychological morbidity can be ameliorated by sensitive, empathetic, knowledgeable counselling. Thus, all patients who have genital HSV infections and their sexual partner(s) can likely benefit from learning about the chronic aspects of the disease after the acute illness subsides. Explain the natural history of the disease, with emphasis on the potential for recurrent episodes, asymptomatic shedding and sexual transmission. Advise patients that antiviral therapy for recurrent episodes may shorten the duration of lesions, and suppressive antiviral therapy can ameliorate or prevent recurrent outbreaks, with one drug having been demonstrated to reduce transmission.¹⁰
- The most common psychological patient concerns include the following:
 - Fear of transmission.
 - Fear of being judged or rejected by partner.
 - Loneliness, depression and low self-esteem.
 - Anxiety concerning potential effect on childbearing.
- Patients must inform their sex partner(s) that they have genital herpes. It may be
 useful to have the partner receive counselling concurrently for information and
 possible serologic testing for HSV-1 and/or -2 antibody.
- Type-specific serologic testing for HSV-1 and/or -2 antibody can demonstrate
 whether couples are discordant or concordant for HSV-1 and/or -2 infection.
 Such information will be useful in counselling couples about the risk of
 transmission of genital herpes infection.

- It should be emphasized that most transmission of genital herpes occurs in the context of asymptomatic shedding.¹¹ Therefore, emphasizing the use of condoms and suppressive antiviral drug therapy is important for reducing the risk of transmission.
- Transmission of genital herpes is decreased by the following:
 - Avoidance of contacts with lesions during obvious periods of viral shedding (prodrome to re-epithelialization) from lesions. Advise patients that they should abstain from sexual activity from the onset of prodromal symptoms until the lesions have completely healed.
 - Condom use (see Prevention section, above).22
 - Daily suppressive antiviral therapy, which reduces recurrent lesions, asymptomatic viral shedding and transmission.¹⁰
- Assess patients with genital herpes for other STIs and treat as needed.³³
- Discuss the risk of neonatal infection with all patients, including men. Women
 who have genital herpes should be advised to inform the health care providers
 who care for them during pregnancy about their HSV infection.
- Genital herpes increases the risk of acquisition of HIV twofold.³⁴

Treatment35

First episode

- Treatment is recommended for clinically important symptoms.
- Analgesia and laxatives may be required. Urinary retention may be an indication for hospitalization.

Table 1. Treatment for first episode

- For severe primary disease, IV acyclovir 5 mg/kg infused over 60 minutes every 8 hours [A-I] is optimal, with conversion to oral therapy when substantial improvement has occurred.³⁶
- Oral acyclovir 200 mg five times per day for 5–10 days [A-I] $^{\rm 37}$

OR

• Famciclovir 250 mg tid for 5 days [A-I]^{38,39}

OR

- Valacyclovir 1000 mg bid for 10 days [A-I]40
- Acyclovir 400 mg tid for 7–10 days is recommended by the U.S. Centers for Disease Control [A-III]²⁴

Notes:

- · Oral acyclovir, famciclovir and valacyclovir are comparably efficacious.
- Acyclovir has been initiated as late as 5–7 days after onset of symptoms with benefit [A-I]³⁷; famciclovir has been initiated
 only in patients with symptoms of fewer than 5 days' duration [A-I] and valacyclovir in those with fewer than 72 hours of
 symptoms [A-I].
- Topical acyclovir does not alleviate systemic symptoms and should not be used [A-I].³⁷

Recurrent lesions35

Table 2. Treatment for recurrent episodes

- Valacyclovir 500 mg bid OR 1 g qd for 3 days [B-I] 41

OR

• Famciclovir 125 mg bid for 5 days [B-I] 42

ΛR

- Acyclovir 200 mg 5 times/day for 5 days [C-I] 43
- A shorter course of acyclovir 800 mg tid for 2 days appears as efficacious as the approved 5-day regimen [B-I]⁴⁴

Notes:

- Valacyclovir, famciclovir and acyclovir are approved for treatment of recurrent genital herpes lesions.
- To be effective, these drugs need to be started as early as possible during the development of a recurrent lesion preferably fewer than 6 hours (famciclovir) [*B-I*] to 12 hours (valacyclovir) [*B-I*] after the first symptoms appear. Patient-initiated therapy at the onset of prodromal symptoms has been proven effective in a Canadian study.⁴² To achieve this end, patients should have medication on hand and be provided with specific information on when to initiate therapy.

Suppressive therapy35

- Suppressive therapy is intended for patients with frequently recurring genital herpes, generally for those with recurrences at least every 2 months or 6 times per year. In such patients, suppressive therapy is preferred to episode therapy⁴⁵ and improves quality of life.⁴⁶
- For individuals with fewer than 6 recurrences per year or one every 2 months, episode therapy is recommended (see above). However, suppressive therapy will probably be efficacious and may be considered on a case-by-case basis.

Table 3. Suppressive therapy for non-pregnant patients

- Acyclovir 200 mg tid to five times daily OR 400 mg bid [A-I]^{47–59}
 OR
- Famciclovir 250 mg bid [A-I]60,61

ΛR

• Valacyclovir 500 mg qd [A-I] (for patients with nine or fewer recurrences per year) OR 1000 mg qd [A-I]^{57,62} (for patients with more than nine recurrences per year).

Notes:

- · Acyclovir, famciclovir and valacyclovir are approved for suppressive therapy in Canada.
- Safety and efficacy data suggest that acyclovir and valacyclovir can be administered for up to 1 year [A-I] based on controlled trials^{47–59,62} whereas famciclovir has been evaluated only for up to 4 months' [A-I] administration.^{60,61}

Table 4. Suppressive therapy for pregnant patients

- Acyclovir 200 mg gid [A-I] 63,64 OR 400 mg tid [A-I] 65,66
- Both regimens have been evaluated and shown to be efficacious in reducing recurrent disease and the need for cesarean section.
- Both regimens require initiation of suppression with acyclovir 400 mg tid at 36 weeks with termination at parturition [A-I] 65,66

Notes:

- There have been no studies of sufficient power to adequately assess whether suppressive antiviral drug therapy in pregnancy reduces maternal-to-child transmission or neonatal herpes per se.
- Acyclovir safety and efficacy have been evaluated in limited numbers of pregnant women [A-III].^{63,65}
- Suppressive acyclovir has been demonstrated to reduce recurrence rates, as well as asymptomatic shedding, and thereby
 obviate the need for cesarean section to prevent neonatal herpes [A-I].⁶³⁻⁶⁶
- · Use of acyclovir suppression does not eliminate the need to observe the neonate carefully for possible HSV infection.

Table 5. Therapy for neonatal herpes

 Acyclovir 45–60 mg/kg/day IV in three equal 8-hourly infusions, each over 60 minutes for 14–21 days [A-I]⁶⁷

Note:

• Consultation with a colleague experienced in this area should be sought.

Consideration for Other STIs

- Having HSV can increase the risk of acquiring and transmitting HIV. This
 increased risk needs to be explained; HIV testing with pre- and post-test
 counselling should be offered.
- Genital ulcers can also be caused by syphilis, chancroid or lymphogranuloma venereum, and testing for these should be considered.
- Testing for other STIs, including chlamydia and gonorrhea, should be considered.
- Immunization for hepatitis B may be indicated.
- See Primary Care and Sexually Transmitted Infections chapter.

Reporting and Partner Notification

- At the time of publication, genital HSV infections were reportable by physicians
 to local public health authorities in New Brunswick, Nova Scotia, Prince
 Edward Island and Newfoundland. Neonatal HSV infections are reportable in
 some provinces only. Whether cases are to be reported on suspicion or after
 laboratory confirmation also varies.
- Partner notification is not required as a public health measure, in part because of the following:
 - Most disease presents as recurrences.

- It is difficult to assess whether a contact has ever had a primary genital infection.
- Patients with genital herpes should be encouraged to inform their sexual partner(s) from the preceding 60 days prior to symptom onset or date of diagnosis where asymptomatic to make them aware of the risk of infection, if uninfected, and to aid diagnosis in a partner if the disease does arise.

Follow-up

- Follow-up cultures are not indicated, except when there are unusual recurrent symptoms or to determine in vitro susceptibility when resistance is suspected as a cause of therapeutic failure.
- Supportive counselling is an important component of managing patients with genital herpes.

Special Considerations

Neonatal herpes^{68,69}

- Recent epidemiologic work on risk factors for neonatal herpes⁶⁸ has demonstrated that the greatest risk factor for neonatal HSV infection is new maternal genital HSV-1 or -2 infection without a fully developed maternal immune response by the time of delivery, resulting in an infant born without homologous transplacental HSV type-specific antibody. Four of nine such infants developed neonatal HSV infection. On the other hand, infants delivered vaginally by women with reactivation of genital herpes with genital lesions or asymptomatic HSV genital virus shedding at parturition had a 2% risk of infection (2 of 92 cases). Cesarean delivery was shown definitively to protect against neonatal transmission of HSV. Thus, the opportunity for preventing neonatal HSV relates more to obviating maternal genital infection late in pregnancy than to identifying women with known genital HSV infection. That is, there is reason for reassurance of pregnant women with a history of genital herpes.
- Incidence in Canada for 2000–2003 inclusive is 5.85 per 100,000 live births;
 62.5% of these infections were attributed to HSV-1.⁷⁰ From 55–80% are due to HSV-2.^{71–74}
- Intrauterine infection accounts for 5% of neonatal HSV infection, and postnatal infection (usually HSV-1) for 15%.⁷²⁻⁷⁴
- Clinically, neonatal infection is classified as skin-eye-mouth (SEM), central
 nervous system (CNS) or disseminated infection. Mortality is 0%, 15% and
 47%, respectively, and abnormal development at 1 year is 2%, 70% and 25%,
 respectively.^{71,72,74} However, overlap occurs, and up to 30% of babies with SEM
 initially will progress to CNS disease as well.
- In the Canadian study, 63.8% of cases had localized (SEM) disease, while 34.5% had infection that disseminated to the CNS or other organs.
- Vesicular skin lesions may not be observed in 17% with SEM, 32% with CNS and 39% of neonates with disseminated disease.

- · Risk of neonatal infection:
 - Is up to 50% if the mother has primary genital HSV infection with lesions at parturition.⁷³ In approximately 70% of cases the mother has no history of genital herpes.^{72,74}
 - Is from 2–8% when vaginal delivery occurs and the mother has a recurrent genital lesion or has asymptomatic genital HSV shedding at parturition.^{68,75}
- Median incubation period is 4 days, with a range of 1–28 days.^{71,72,74}
- Most neonatal herpes begins after a seemingly healthy neonate has left hospital.
- Acyclovir oral therapy suppresses recurrent genital disease and asymptomatic shedding and thereby has been shown to reduce the need for cesarean delivery (see Treatment section, above).

Laboratories offering HSV type-specific serum antibody testing

- Alberta Provincial Laboratory for Public Health, Edmonton, Alberta (implementation anticipated in 2005).
- National Microbiology Laboratory, Public Health Agency of Canada, Winnipeg, Manitoba.
- Regional Virology & Chlamydia Laboratory, Hamilton, Ontario.
- Children's Hospital of Eastern Ontario Laboratory, Ottawa, Ontario.
- · Warnex Inc., Montreal, Quebec.

References

- Corey L, Adams HG, Brown ZA, Holmes KK. Genital herpes simplex virus infections: clinical manifestations, course, and complications. *Ann Intern Med* 1983;98:958–972.
- Smith JS, Robinson NJ. Age-specific prevalence of infection with herpes simplex virus types 2 and 1: a global review. *J Infect Dis* 2002;186(suppl 1): \$3-28
- 3. Armstrong GL, Schillinger J, Markowitz L, et al. Incidence of herpes simplex virus type 2 infection in the United States. *Am J Epidemiol* 2001;153:912–920.
- 4. Fleming DT, McGuillan GM, Johnson RE, et al. Herpes simplex virus type 2 in the United States, 1976 to 1994. *N Engl J Med* 1997;337:1105–1111.
- Patrick DM, Dawar M, Cook DA, Krajden M, Ng HC, Rekart ML. Antenatal seroprevalence of Herpes simplex virus type 2 (HSV-2) in Canadian women: HSV-2 prevalence increases throughout the reproductive years. Sex Transm Dis 2001;28:424–428.
- 6. Singh AE, Romanowski B, Wong T, et al. Herpes simplex virus seroprevalence and risk factors in 2 Canadian sexually transmitted disease clinics. *Sex Transm Dis* 2005;32:95–100.
- Lafferty WE, Downey L, Celum C, Wald A. Herpes simplex virus type 1 as a cause of genital herpes: impact surveillance and prevention. *J Infect Dis* 2000;181:1454–1457.
- 8. Nilsen A, Myrmel H. Changing trends in genital herpes simplex virus infection in Bergen, Norway. *Acta Obstet Gynecol Scand* 2000;79:693–696.

- Forward KR, Lee SHS. Predominance of herpes simplex virus type 1 from patients with genital herpes in Nova Scotia. Can J Infect Dis 2003;14:94–96.
- 10. Corey L, Wald A, Patel R, et al. Once-daily valacyclovir to reduce the risk of transmission of genital herpes. *N Engl J Med* 2004;350:11–20.
- 11. Mertz GJ, Benedetti J, Ashley R, Selke SA, Corey L. Risk factors for the sexual transmission of genital herpes. *Ann Intern Med* 1992;116:197–202.
- 12. Bryson Y, Dillon M, Bernstein DI, Radolf J, Zakowski P, Garratty E. Risk of acquisition of genital herpes simplex virus type 2 in sex partners of persons with genital herpes: a prospective couple study. *J Infect Dis* 1993;167:942–946.
- 13. Langenberg AG, Corey L, Ashley RL, Leong WP, Straus SE. A prospective study of new infections with herpes simplex virus type 1 and type 2. Chiron HSV Vaccine Study Group. *N Engl J Med* 1999;341:432–1438.
- 14. Corey L, Fife KH, Beneditti JK, et al. Intravenous acyclovir for the treatment of primary genital herpes. *Ann Intern Med* 1983;98:914–921.
- 15. Sacks SL. *The Truth about Herpes*. 4th ed. Vancouver, BC: Gordon Soules Book Publishers: 1997.
- Lafferty WE, Coombs RW, Benedetti J, Critchlow C, Corey L. Recurrences after oral and genital herpes simplex virus infection. Influence of site of infection and viral type. N Engl J Med 1987;316:1444–1449.
- Benedetti JK, Zeh J, Corey L. Clinical reactivation of genital herpes simplex virus infection decreases in frequency over time. *Ann Intern Med* 1999; 131:14–20.
- Wald A, Zeh J, Selke S, et al. Reactivation of genital herpes simplex type 2 infection in asymptomatic seropositive persons. N Engl J Med 2000;342: 844–850.
- Wald A, Zeh J, Selke S, Warren T, Ashley R, Corey L. Genital shedding of herpes simplex virus among men. J Infect Dis 2002;186(suppl 1):S34–S39.
- Wald A, Zeh J, Selke S, Ashley R, Corey L. Virologic characteristics of subclinical and symptomatic genital herpes infections. *N Engl J Med* 1995;333:770–775.
- 21. Cowan FM, Copas A, Johnson AM, Ashley R, Corey L, Mindel A. Herpes simplex virus type 1 infection: a sexually transmitted infection of adolescence? Sex Transm Infect 2002;78:346–348.
- 22. Wald AM, Langenberg AG, Link K, et al. Effect of condoms on reducing the transmission of herpes simplex virus type 2 from men to women. *JAMA* 2001;285:3100–3106.
- 23. Stanberry LR, Spruance SL, Cunningham AL, et al. Glycoprotein-D-adjuvant vaccine to prevent genital herpes. *N Engl J Med* 2002;347:1652–1661.
- 24. Ashley-Morrow R, Krantz E, Wald A. Time course of seroconversion by HerpeSelect ELISA after acquisition of genital herpes simplex virus type 1 (HSV-1) or HSV-2. *Sex Transm Dis* 2003;30:310–314.
- 25. Corey L, Holmes KK. Genital herpes simplex virus infections: current concepts in diagnosis, therapy and prevention. *Ann Intern Med* 1983;98:973–983.

- Wald A, Huang M-L, Carrell D, Selke S, Corey L. Polymerase chain reaction for detection of herpes simplex virus (HSV) DNA on mucosal surfaces: comparison with HSV isolation in cell culture. *J Infect Dis* 2003;188:1345–1351.
- 27. Soloman AR, Rasmussen JE, Varani J, Pierson CL. The Tzanck smear in the diagnosis of cutaneous herpes simplex. *JAMA* 1984;251:633–635.
- 28. Kohl S, Adam E, Matson DO, Kaufman RH, Dreesman GR. Kinetics of human antibody responses to primary genital herpes simplex virus infection. *Intervirology* 1982;18:164–168.
- Lopez C, Arvin AM, Ashley R. Immunity to herpesvirus infections in humans.
 In: Roizman B, Whitley RJ, Lopez C, eds. *The Human Herpesviruses*. New York, NY: Raven Press; 1993.
- Ashley RL, Eagleton M, Pfeiffer N. Ability of a rapid serology test to detect seroconversion to herpes simplex virus type 2 glycoprotein G soon after infection. J Clin Microbiol 1999;37:1632–1633.
- 31. Ashley RL. Progress and pitfalls in serological testing for genital herpes. *Herpes* 1994;1:49–51.
- 32. Diamond C, Selke S, Ashley R, Benedetti J, Corey L. Clinical course of patients with serologic evidence of recurrent genital herpes presenting with signs and symptoms of first episode disease. *Sex Transm Dis* 1999;26:221–225.
- 33. Sexually transmitted diseases treatment guidelines 2002. Centers for Disease Control and Prevention. *MMWR Recomm Rep* 2002;51(RR-6):1–78.
- 34. Wald A, Link K. Risk of human immunodeficiency virus infection in herpes simplex virus type 2-seropositive persons: a meta-analysis. *J Infect Dis* 2002;185:45–52.
- 35. Aoki FY. Contemporary antiviral drug regimens for the prevention and treatment of orolabial and anogenital herpes simplex virus infection in the normal host: four approved indications and 13 off-label uses. *Can J Infect Dis.* 2003;14:17–27.
- Corey L, Benedetti J, Critchlow CW, et al. Treatment of primary first-episode genital herpes simplex virus infections with acyclovir: results of topical, intravenous and oral therapy. *J Antimicrob Chemother* 1983;12(suppl B):79–88.
- Mertz GJ, Critchlow CW, Benedetti J, et al. Double-blind placebo-controlled trial of oral acyclovir in first-episode genital herpes simplex virus infection. *JAMA* 1984;252:1147–1151.
- 38. Murphy SM, Ruck F, Kitchin VS. Oral famciclovir (FCV) a new antiherpes agent: comparative study with acyclovir in clinic initiated treatment of first episode genital herpes (FGH) [abstract]. Presented at: European Academy of Dermatology and Venereology/Triaena Congress. 1991; Athens, Greece.
- 39. Loveless M, Harris JRW, Sacks SL. Famciclovir in the management of first-episode genital herpes. *Infect Dis Clin Pract* 1997;6(suppl 1):S12–S16.
- 40. Fife KH, Barbarash RA, Rudolph T, Degregorio B, Roth R. Valaciclovir versus acyclovir in the treatment of first-episode genital herpes infection. Results of an international, multicenter, double-blind, randomized clinical trial. The Valaciclovir International Herpes Simplex Virus Study Group. Sex Transm Dis 1997;24:481–486.

- Spruance SL, Tyring SK, DeGregorio B, Miller C, Beutner K. A large-scale, placebo-controlled, dose-ranging trial of peroral valaciclovir for episodic treatment of recurrent herpes genitalis. Valaciclovir HSV Study Group. *Arch Intern Med* 1996:156:1729–1735.
- Sacks SL, Aoki FY, Diaz-Mitoma F, Sellors J, Shafran SD. Patient-initiated, twice-daily oral famciclovir for early recurrent genital herpes. A randomized, double-blind, multicenter trial. Canadian Famciclovir Study Group. *JAMA* 1996;276:44–49.
- 43. Tyring SK, Douglas JM Jr, Corey L, Spruance SL, Esmann J. A randomized, placebo-controlled comparison of oral valacyclovir and acyclovir in immunocompetent patients with recurrent genital herpes infections. The Valacyclovir International Study Group. *Arch Dermatol* 1998;134:185–191.
- 44. Wald A, Carrell D, Remington M, Kexel E, Zeh J, Corey L. Two-day regimen of acyclovir for treatment of recurrent genital herpes simplex virus type 2 infection. *Clin Infect Dis* 2002;34:944–948.
- 45. Romanowski B, Marina RB, Roberts JN, Valtrex HS230017 Study Group. Patients' preference of valacyclovir once-daily suppressive therapy versus twice-daily episodic therapy for recurrent genital herpes: a randomized study. *Sex Transm Dis* 2003;30:226–231.
- 46. Patel R, Tyring S, Strand A, Price MJ, Grant DM. Impact of suppressive antiviral therapy on the health related quality of life of patients with recurrent genital herpes infection. *Sex Transm Infect* 1999;75:398–402.
- 47. Sacks SL, Fox R, Levendusky P, et al. Chronic suppression for six months compared with intermittent lesional therapy of recurrent genital herpes using oral acyclovir: effects on lesions and nonlesional prodromes. *Sex Transm Dis* 1988;15:58–62.
- 48. Thin RN, Jeffries DJ, Taylor PK, et al. Recurrent genital herpes suppressed by oral acyclovir: a multicentre double blind trial. *J Antimicrob Chemother* 1985;16:219–226.
- 49. Mindel A, Weller IV, Faherty A, et al. Prophylactic oral acyclovir in recurrent genital herpes. *Lancet* 1984;2:57–59.
- 50. Kinghorn GR, Jeavons M, Rowland M, et al. Acyclovir prophylaxis of recurrent genital herpes: randomized placebo controlled crossover study. *Genitourin Med* 1985;61:387–390.
- 51. Halsos AM, Salo AP, Lassus A, et al. Oral acyclovir suppression of recurrent genital herpes: a double-blind, placebo-controlled, crossover study. *Acta Derm Venereol* 1985;65:59–63.
- 52. Blom I, Bäck O, Egelrud T, et al. Long-term oral acyclovir treatment prevents recurrent genital herpes. *Dermatologica* 1986;173:220–223.
- Mertz GJ, Jones CC, Mills J, et al. Long-term acyclovir suppression of frequently recurring genital herpes simplex virus infection. A multicenter double-blind trial. *JAMA* 1988;260:201–206.
- 54. Baker DA, Blythe JG, Kaufman R, Hale R, Portnoy J. One-year suppression of frequent recurrences of genital herpes with oral acyclovir. *Obstet Gynecol* 1989;73:84–87.

- Kroon S, Petersen CS, Andersen LP, Rasmussen LP, Vestergaard BF. Oral acyclovir suppressive therapy in severe recurrent genital herpes. A doubleblind, placebo-controlled cross-over study. *Dan Med Bull* 1989;36:298–300.
- 56. Mostow SR, Mayfield JL, Marr JJ, Drucker JL. Suppression of recurrent genital herpes by single daily dosages of acyclovir. *Am J Med* 1988;85(2A):30–33.
- 57. Reitano M, Tyring S, Lang W, et al. Valaciclovir for the suppression of recurrent genital herpes simplex virus infection: a large-scale dose range-finding study. International Valaciclovir HSV Study Group. *J Infect Dis* 1998;178:603–610.
- Douglas JM, Critchlow C, Benedetti J, et al. A double-blind study of oral acyclovir for suppression of recurrences of genital herpes simplex virus infection. N Engl J Med 1984;310:1551–1556.
- 59. Strauss SE, Takiff HE, Seidlin M, et al. Suppression of frequently recurring genital herpes. A placebo-controlled double-blind trial of oral acyclovir. *N Engl J Med* 1984;310:1545–1550.
- Mertz GJ, Loveless MO, Levin MJ, et al. Oral famciclovir for suppression of recurrent genital herpes simplex virus infection in women. A multicenter, double-blind, placebo-controlled trial. Collaborative Famciclovir Genital Herpes Research Group. Arch Intern Med 1997;157:343–349.
- Diaz-Mitoma F, Sibbald GR, Shafran SD, Boon R, Saltzman RL. Oral famciclovir for the suppression of recurrent genital herpes: a randomized controlled trial. Collaborative Famciclovir Genital Herpes Research Group. *JAMA* 1998:280:887–892.
- 62. Patel R, Bodworth NJ, Woolley P, et al. Valaciclovir for the suppression of recurrent genital HSV infection: a placebo controlled study of once daily therapy. International Valaciclovir HSV Study Group. *Genitourin Med* 1997;73:105–109.
- 63. Stray-Pedersen B. Acyclovir in late pregnancy to prevent neonatal herpes simplex. *Lancet* 1990;336:756.
- 64. Braig S, Luton D, and Sibony O, et al. Acyclovir prophylaxis in late pregnancy prevents recurrent genital herpes and viral shedding. *Eur J Obstet Gynecol Reprod Biol* 2001;96:55–58.
- 65. Scott LL, Sanchez PJ, Jackson GL, Zeray F, Wendel GD Jr. Acyclovir suppression to prevent cesarean delivery after first-episode genital herpes. *Obstet Gynecol* 1996;87:69–73.
- Watts DH, Brown ZA, Money D, et al. A double-blind, randomized, placebocontrolled trial of acyclovir in late pregnancy for the reduction of herpes simplex virus shedding and caesarean delivery. *Am J Obstet Gynecol* 2003;188:836–843.
- Kimberlin DW, Lin CY, Jacobs RF, et al. Safety and efficacy of high-dose intravenous acyclovir in the management of neonatal herpes simplex virus infections. *Pediatrics* 2001;108:230–238.
- Brown ZA, Wald A, Morrow RA, Selke S, Zeh J, Corey L. Effect of serologic status and cesarean delivery on transmission rates of herpes simplex virus from mother to infant. *JAMA* 2003;289:203–209.

- 69. Brown ZA, Selke S, Zeh J, et al. The acquisition of herpes simplex virus during pregnancy. *N Engl J Med* 1997;337:509–515.
- Kropp RY, Wong T, Cormier L, Ringrose A, Embree J, Steben M, Canadian Paediatric Surveillance Program (CPSP). Epidemiology of neonatal herpes simplex virus infections in Canada. Presented at the International Society for STD Research (ISSTDR) conference 2005, Amsterdam.
- 71. Whitley RJ, Corey L, Arvin A, et al. Changing presentation of herpes simplex virus infection in neonates. *J Infect Dis* 1988;158:109–116.
- 72. Kimberlin DW, Lin CY, Jacobs RF, et al. Natural history of neonatal herpes simplex virus infections in the acyclovir era. *Pediatrics* 2001;108:223–229.
- 73. Enright AM, Prober CG. Neonatal herpes infection: diagnosis, treatment and prevention. *Semin Neonatol* 2002;7:283–291.
- 74. Koskiniemi M, Happonen JM, Jarvenpaa AL, Pettay O, Vaheri A. Neonatal herpes simplex virus infection: a report of 43 patients. *Pediatr Infect Dis* 1989;8:30–35.
- 75. Prober CG, Sullender WM, Yasukawa LL, Au DS, Yeager AS, Arvin AM. Low risk of herpes simplex virus infections in neonates exposed to the virus at the time of vaginal delivery to mothers with recurrent genital herpes simplex virus infections. *N Engl J Med* 1987;316:240–244.

GENITAL HUMAN PAPILLOMAVIRUS (HPV) INFECTIONS

This chapter covers the prevention, diagnosis and treatment of human papillomavirus infection. For complete information on the prevention, diagnosis and treatment of cervical cancer, other sources should be used.

Etiology

Definition

Human papillomavirus (HPV) causes skin or mucosal infections and has a strong
affinity for the moist mucosa of the anal, genital and aerodigestive tracts.

Etiology

More than 130 HPV types have been classified on the basis of DNA sequence,
 40 of which can infect the anogenital epithelium. HPV types are classified as high- or low-risk based on the strength of their association with cervical cancer.

Table 1. HPV types

Association with cervical cancer ¹	Genotypes	Most likely clinical conditions
Low-risk	Most common: 6 and 1140, 42, 43, 44, 54, 61, 70, 72, 81 and CP6108	Condylomata acuminata
Probable high-risk	• 26, 53 and 66	Precancerous or cancerous lesions
High-risk	 Most common: 16 and 18 31, 33, 35, 39, 45, 51, 52, 56, 58, 59, 68, 73 and 82 	Precancerous or cancerous lesions

Epidemiology

- HPV is one of the most common sexually transmitted infections (STIs).²
- The incubation period for exophytic warts is 1–8 months.
- 70% of the adult population will have had at least one genital HPV infection over their lifetime.³
- Canadian HPV prevalence studies show that HPV infection is very common and that wide variability exists between different populations:
 - In young women, prevalence is reaching 29%.^{4,5}
 - In a community health centre in Manitoba where 73% of participants were
 years, 33% of women were found to be HPV-positive.⁶
 - In women aged 15–49, attending for routine cervical cancer screening in Ontario, the prevalence of high-risk HPV was found to be 12.7%.^{7,8}

- In women aged 13–79, attending for routine cervical cancer screening in Nunavut the prevalence of high-risk HPV was found to be 25.7%.9
- HPV infections are often acquired early (15–19 years of age),¹⁰ and the majority (>80%) of these infections clear spontaneously within 18 months.¹¹
- HPV infections usually occur in adolescents and young adults, but affect both women and men of all ages.
- Non-oncogenic or low-risk HPV, which can be expressed as exophytic warts, is associated with a low risk for cancer.
- Clinically visible external genital warts (EGWs) (with low-risk HPV) were noted in ~1% of sexually active adults (aged 15–49) in a U.S. population.¹²
- Thirteen high-risk HPV types have been confirmed in the International Agency for Research on Cancer monograph on cervical cancer screening as necessary factors in the etiology of cervical cancer, while other HPV types have been implicated in skin and oral-pharyngeal cancers, as well as with cancers of the anus and penis.¹³
- The average time from acquiring a high-risk genotype of HPV to the detection of cervical cancer is 20 years.¹⁴
- Infection with one HPV genotype does not protect against infection with other types.^{15,16}
- Simultaneous infection with multiple types of HPV has been reported in 5–30% of women with HPV.¹⁷
- Symptomatic perinatal transmission is infrequent and is usually clinically apparent within 2 years. When it occurs, it is associated with anogenital and vocal-cord lesions in the newborn.¹⁸

Prevention

- While condoms may not reliably prevent sexual transmission of HPV, they may
 protect against the HPV types of genital warts,¹⁹ some co-factors of cervical
 dysplasia and invasive cervical cancer; in addition, they effectively prevent
 transmission of bacterial STIs.
- Counsel patients with HPV infection about risk reduction, including the following:
 - Natural history of the disease, with emphasis on the differences between HPV genotypes and their potential manifestations.
 - Potential for recurrent episodes.
 - Potential for sexual transmission.
- There are conflicting epidemiologic data on risk factors and co-factors for HPV infection. The only factor that emerges consistently is lifetime number of sex partners. Putative co-factors for cervical cancer include the following:
 - Smoking tobacco and exposure to tobacco smoke.
 - Long-term use of oral contraceptives (>5 years).
 - Higher number of pregnancies.
 - Other STIs (e.g., Chlamydia trachomatis, herpes simplex virus-2, HIV).
 - Inadequate diet (especially low antioxidant intake).

- Immunosuppression (e.g., HIV/AIDS, organ transplant and immunosuppressive drug therapy).
- Multiple sex partners, sexual intercourse at an early age and sexual intercourse with those infected with HPV.
- Genetic susceptibility: polymorphisms in certain cell regulatory genes, such as p53.

Information about HPV²⁰⁻²³

- Inform women that regular cervical screening for dysplasia and/or HPV infection is effective in reducing rates of cervical cancer.^{24–26}
- Counselling for patients with HPV and/or abnormal cervical screening results should include the following:
 - Explanation of the natural history of the disease, with emphasis on the differences between types of HPV and their causal associations (i.e., lowrisk types are associated with anogenital warts, and high-risk types are associated with cervical cancer).
 - Discussion of the risk of recurrence.
 - Reduction of the impact of risk and co-factors for progression to dysplasia.
 - Encouragement of patients to examine themselves and seek medical attention if lesions appear.
 - Reassurance that the virus is common, and that it is virtually impossible to determine when or from whom they acquired the virus.
 - Reassurance that the risk of cervical cancer is quite low and that most HPV infections will resolve and clear.
 - Reassurance that only persistent infection with high-risk HPV types may progress to precancerous and cancerous lesions.

Diagnosis

- Most anogenital HPV infections are asymptomatic and subclinical. Of those clinically apparent lesions, most will be asymptomatic.
- The most frequent sites of anogenital HPV infection in females are the cervix, vagina, vulva or anus.
- The most frequent sites of anogenital HPV infection in males are the anus or penis.
- Multiple sites are often involved (e.g., cervix, vagina, vulva etc.).
- The natural history is of fluctuation in size and number of warts and, in most cases, eventual clearance.
- Warts can increase in size and number with pregnancy.
- Intraepithelial lesions on a Pap smear usually indicate cervical involvement. These are classified as one of the following:
 - Low-grade squamous intraepithelial lesions (LSILs): under the old classification system, these were known as condyloma of the cervix, mild to moderate dysplasia or cervical intraepithelial neoplasia (CIN) 1 or CIN2.

- High-grade squamous intraepithelial lesions (HSILs): under the old classification system, these were known as severe dysplasia, CIN3 or in situ neoplasia.
- Invasive carcinoma.

External genital warts27

- Most EGWs are caused by low-risk HPV infections.
- Typical EGWs present as exophytic fronds or cauliflower-like to papular growths
 on anogenital skin and/or mucous membranes called condylomata acuminata.
 They are frequently multiple, asymmetric and polymorphic. They occasionally
 cause bleeding, pruritus and local discharge.
- Less frequent manifestations of EGWs are slightly elevated lesions, papular
 or macular lesions with or without keratinization and/or brown/grey/bluish
 pigmentation, also known as bowenoid papulosis, or warty vulvar intraepithelial
 neoplasia.

Table 2. Non-HPV lesions to consider in a differential diagnosis

Normal variations	 In both sexes: sebaceous glands In women: vestibular papillae, also known as micropapillomatosis labialis In men: pearly penile papules on the coronal sulcus
Pathologic entities	 Infections Secondary syphilis with condylomata lata Molluscum contagiosum Diseases of the skin and mucosa Intradermal nevi Skin tags Seborrheic keratoses Cancer Intraepithelial neoplasia

Note:

This table does not include manifestations, which are listed above.

Specimen collection and laboratory diagnosis

Cervical cytology (Pap smear or Pap test)

- Two different methods can be used to screen for cervical cancer and its
 precursors: a glass slide fixed with Cytospray (conventional) or liquid-based
 cytology (LBC). Access to LBC is limited to only a small number of jurisdictions
 in Canada at present.
 - LBC for women with an ordinary risk of cervical cancer is more sensitive than the conventional glass-slide smear and produces a lower rate of unusable samples.²⁸

- Regular cervical screening is important for all women who are, or have ever been, sexually active. Some North American guidelines recommend starting within 3 years of initiation of penetrative sexual activity,²⁹ but European guidelines recommend starting at 25 years of age.^{30,31}
- Provincial and territorial guidelines for cervical cytology vary across Canada.
- Cervical Cancer Prevention Network guidelines recommend annual Pap smears until two sequential normal Pap smears are obtained, then every 3 years if normal in immunocompetent individuals.³²
- Immunocompromised persons, especially those who are HIV-positive, require special attention. Please refer to a local expert for optimal management.
- Cervical cancer is more frequent in women who have not had cervical screening at regular intervals^{24,25,33} and women who are HIV-positive.³⁴
- Many women who develop cervical cancer have had inadequate cytology on previous smears.³⁵
- The best specimen collection device is the extended-tip spatula combined with the Cytobrush.³⁶
- Results are reported in some jurisdictions using Bethesda 2001 terminology,³⁷ but this varies by province and territory.

HPV typing

- A meta-analysis of the available literature concluded that HPV DNA testing is better than repeat cytology in women who have atypical squamous cells of undetermined significance (ASCUS) on Pap smears.³⁸ The Pan Canadian Forum on Cervical Screening has recommended HPV DNA testing for this indication.³⁹
- Co-testing using LBC and HPV DNA testing is approved in the U.S. for primary screening, but no such recommendation exists in Canada.
- HPV typing is not useful for EGWs, which are most likely caused by low-risk non-oncogenic types,² or in women with LSILs or HSILs, because of the high prevalence of oncogenic types in such cases.⁴⁰
- Access to HPV DNA tests in Canada is limited to a small number of jurisdictions at present.

Colposcopy

- Colposcopy should be performed for the following:
 - Clinically visible growths, warts or suspicious findings on the cervix.
 - Abnormal cervical screening test results, including the following:
 - Repeat ASCUS (especially if HPV detection test is positive)
 - ASCUS cannot exclude high-grade lesion
 - LSILs
 - HSILs
 - · Atypical glandular cells
 - · Invasive carcinoma
 - Positive high-risk HPV detection twice in a 6–12 month period, even in the presence of normal cytology.

 Routine colposcopy for women with EGWs is not likely to be beneficial unless other criteria (see above) are present.⁴¹

Aceto-whitening or aceto-acid testing

- A solution of 5% acetic acid applied to the genital skin or the cervix for 1–3 minutes may lead to whitening of HPV-infected epithelium; however, this test has a high false-positive rate in both female and male patients.
- This test is never recommended for screening of external anogenital warts or subclinical lesions, even for partners of persons with an abnormal Pap smear or EGWs.
- This test should be reserved as an adjunct to colposcopy to increase the visibility of subclinical lesions.

Anoscopy

- · Anoscopy should be considered in patients with anal warts.
- Anal cancer is being studied with anal Pap and viral testing as a screening method. Patients with positive results are then managed following clinical evaluation done by high-resolution anoscopy. This may be particularly important for HIV-positive patients.

Urethroscopy

• Urethroscopy can be considered for patients with extensive urethral warts not amenable to other forms of therapy.

Caution

Atypical and/or non-healing warts

- Suspect neoplasia if any of the following are present:
 - Pigmented lesions
 - Bleeding
 - Persistent ulceration
 - Persistent pruritus
 - Recalcitrant lesions
- Patients with suspicious lesions may require a biopsy; refer to a colleague experienced in this area.

Management

- · No therapy guarantees eradication of HPV.
- Cell-mediated immunity will eradicate most HPV infections over time in teens and young adults.
- Warts often have a high persistence/recurrence rate, but more than 90% of patients with EGWs experience complete clearance within 2 years, with or without treatment. However, disappearance of warts is not synonymous with HPV eradication.

 Clearance of cervical lesions approaches 90–95%. Successful therapy for cervical abnormalities is often followed by clearance of HPV. HPV testing is being used to help detect residual high-grade disease and recurrent high-grade cervical lesions.⁴²

Treatment

EGWs in males and females

- New lesions, with all available treatments, can occur at sites that may have been treated. They can also occur at different sites at a rate of 20–30%.⁴³
- All treatments are associated with local skin reactions that can best be addressed by decreasing the intensity of the treatment.
- Rates of efficacy are difficult to determine because of a lack of uniformity in clinical trials.

Table 3. Patient-applied treatments

Treatment	Recurrence rate	Safety issues	Comments
Imiquimod [A-I] • Self-applied three times a week (with at least 1 day between applications) for up to 16 weeks • Should be washed off after 6–8 hours	• Recurrence rates are lower (10%) than with any other therapeutic modality ⁴⁴	Should <i>not</i> be used in pregnancy	Mechanism of action is through immune modulation
Podofilox/podophyllotoxin 0.5% solution [A-I] • Applied to warts (but not the contiguous skin) every 12 hours for 3 days of each week (4 days off) ⁴⁵ • Can be repeated for up to 6 weeks only, with the total dose per day not to exceed 0.5 mL	 Recurrence rates are high (60%) More efficacious, stable and associated with fewer side effects than podophyllin (see Table 4) 	Should not be used in pregnancy Should not be used for the treatment of cervical, meatal, vaginal or anal warts	 For self-application under the direction of a physician Available under two brand names in Canada: Wartec and Condyline

Note:

There has been no study comparing these two treatment options.

Table 4. Office-based treatments

Treatment	Recurrence rate	Safety issues	Comments
Cryotherapy [A-I] 46-48 • Liquid nitrogen, carbon dioxide (dry ice or Histofreeze), or nitrous oxide using cryoprobes • Provide sufficient freezing with a rim of 1–2 mm around the lesion	Good response rates	 Safe for use in pregnancy Aggressive treatment of genital warts can leave scarring 	Destruction of the skin is usually limited to the epidermis
Podophyllin 10–25% [A-I] • Should be applied to the wart and not contiguous skin, and must be washed off in 1–4 hours • May be repeated once or twice at weekly intervals, the total dose not to exceed 1–2 mL per visit		 Should not be used in pregnancy; fetal death has been reported Should not be used for the treatment of cervical, meatal, vaginal or anal warts 	 Should be discarded for a better option, such as patient-based therapies Should be used only if other therapies cannot be used Should never be left to selfapplication
		 Frequent local reactions such as erythema, tissue edema, pain, burning, itching, tenderness or bullous reactions are often reported Systemic toxicity has also been reported 	

Table 4. Office-based treatments

Treatment	Recurrence rate	Safety issues	Comments
Bi- or trichloracetic acid [A-I] ^{47,48} • Repeated weekly for 6–8 weeks • 50–80% solutions in 70% alcohol are most effective • Does not need to be washed off		 Safe for use in pregnancy Caustic and may produce blisters and ulcerations 	Healthy skin should be protected with Vaseline, 2% Xylocaine ointment or eutectic mixture of lidocaine and prilocaine cream
Electro-fulguration, CO ₂ laser ablation, excision ⁴⁹	Good response rates	Poor depth control may cause excess damage and scarring	These treatment options are done for more extensive genital, perineal or anal warts

Note:

Topical analgesia with lidocaine or eutectic mixture of lidocaine and prilocaine cream can be used for reduction of pain with office-based therapies.

Extensive, large or resistant external lesions, and internal lesions including vaginal, cervical, anal, urethral and meatal warts

- Patients should be referred to a colleague experienced in this area. CO₂ laser, trichloracetic acid, electroexcision, scissor excision and fulguration may require local or general anesthesia. Low rates of complications are expected if performed by an experienced physician.
- Patients with HIV infection often present with extensive anogenital warts respond poorly to treatment.
- · The following treatments are not recommended:
 - Interferon beta (Intron-A)
 - Dinitrochlorobenzene sensitization
 - Cidofovir
 - Retinoic acid
 - Application immunotherapy with autogenous vaccines
 - 5% 5-fluorouracil cream

Male partners of women with abnormal Pap smears

 Since abnormal Pap smears most often represent the reactivation of an oncogenic latent strain, there is no clinical follow-up required for asymptomatic male partners.
 Previously, these men were subjected to aceto-whitening of the genital area and treatment for subclinical lesions. There are no data to support this [D-III].⁴¹

Subclinical lesions

Lesions may be visible only after examination or application of aceto-whitening.
No specific management is recommended or necessary for subclinical lesions
of the external anogenital skin, as neither recurrences of clinical warts nor
transmission to partners is affected [D-III].

Consideration for Other STIs

- See Primary Care and Sexually Transmitted Infections chapter.
- In patients with condylomata acuminata, an abnormal cervical smear and STI risk factors, obtain specimen(s) for the diagnosis of chlamydial and gonococcal infections.
- HIV testing and counselling are recommended (see *Human Immunodeficiency Virus Infections* chapter).
- Immunization against hepatitis B is recommended (see *Hepatitis B Virus Infections* chapter).
- Consider obtaining a blood sample for serologic testing for syphilis (see *Syphilis* chapter), especially in the presence of condylomata lata.

Reporting and Partner Notification

- HPV is not a reportable infection in Canada.
- "Standard" partner notification recommendations that apply for other STIs are not useful in reducing transmission of HPV.
- Patients should be encouraged to inform their sex partner(s) that they have or
 have had genital warts or an abnormal Pap smear, but there is no proof that this
 will lower the risk to the partner.
- Treatment or referral of asymptomatic partners is not indicated.⁴¹

Follow-up

- Once genital warts are healed, conduct routine follow-up of women with cervical screening, with or without HPV DNA testing, as recommended by provincial/ territorial guidelines.
- Loss to follow-up treatment after abnormal cervical cytology is a significant issue, with rates as high as 40% in some jurisdictions.^{50–52}

Special Considerations

Patients with HIV

 Patients with HIV infection require special care. Conjoint follow-up with an experienced colleague may be indicated.

Children and pregnant patients

- Refer to a colleague experienced in this area, since the psychological aspects and management can be difficult.
- Consider the possibility of sexual abuse when genital warts are present in a child older than 18 months, and particularly in a child older than 2 years of age (see Sexual Abuse in Peripubertal and Prepubertal Children chapter).
- Cesarean section is not recommended unless warts obstruct the birth canal.⁴¹ Approximately 50% of cases of condyloma associated with pregnancy spontaneously regress in the first 3 months after delivery.

References

- Muñoz N, Bosch FX, de Sanjosé S, et al. Epidemiologic classification of human papillomavirus types associated with cervical cancer. N Engl J Med 2003:348:518–527.
- Franco EL, Duarte-Franco E, Ferenczy A. Cervical cancer: epidemiology, prevention and the role of human papillomavirus infection. *CMAJ* 2001; 164:1017–1025.
- 3. Koutsky LA, Galloway DA, Holmes KK. Epidemiology of genital human papillomavirus infection. *Epidemiol Rev* 1988;10:122–163.
- 4. Richardson H, Franco E, Pintos J, Bergeron J, Arella M, Tellier P. Determinants of low-risk and high-risk cervical human papillomavirus infections in Montreal University students. *Sex Transm Dis* 2000;27:79–86.
- Richardson H, Kelsall G, Tellier P, et al. The natural history of type-specific human papillomavirus infections in female university students. *Cancer Epidemiol Biomarkers Prev* 2003;12:485–490.
- Young T, McNichol P, Beauvais J. Factors associated with human papillomavirus infection detected by polymerase chain reaction among urban Canadian aboriginal and non-aboriginal women. Sex Transm Dis 1997;24:293–298.
- Sellors JW, Mahony JB, Kaczorowski J, et al. Prevalence and predictors of human papillomavirus infection in women in Ontario, Canada. Survey of HPV in Ontario Women (SHOW) Group. CMAJ 2000;163:503–508.
- Sellors JW, Karwalajtys TL, Kaczorowski JA, et al. Prevalence of infection with carcinogenic human papillomavirus among older women. *CMAJ* 2002; 167:871–873.
- 9. Healey SM, Aronson KJ, Mao Y, et al. Oncogenic human papillomavirus infection and cervical lesions in aboriginal women of Nunavut, Canada. *Sex Transm Dis* 2001;28:694–700.
- 10. Burk RD, Kelly P, Feldman J, et al. Declining prevalence of cervicovaginal human papillomavirus infection with age is independent of other risk factors. Sex Transm Dis 1996;23:333–341.
- 11. Ho GY, Bierman R, Beardsley L, Chang CJ, Burk RD. Natural history of cervicovaginal papillomavirus infection in young women. *N Engl J Med* 1998;338:423–428.

- Jay N, Moscicki AB. Human papillomavirus infections in women with HIV disease: prevalence, risk, and management. AIDS Read 2000;10:659–668.
- International Agency for Research on Cancer. Cervix Cancer Screening. IARC Handbooks of Cancer Prevention, vol. 10. Oxford: Oxford University Press, 2005.
- 14. Myers ER, McCrory DC, Nanda K, Bastian L, Matchar DB. Mathematical model for the natural history of human papillomavirus infection and cervical carcinogenesis. *Am J Epidemiol* 2000;151:1158–1171.
- 15. Thomas KK, Hughes JP, Kuypers JM, et al. Concurrent and sequential acquisition of different genital human papillomavirus types. *J Infect Dis* 2000;182:1097–1102.
- Liaw KL, Hildesheim A, Burk RD, et al. A prospective study of human papillomavirus (HPV) type 16 DNA detection by polymerase chain reaction and its association with acquisition and persistence of other HPV types. J Infect Dis 2001;183:8–15.
- 17. Rousseau MC, Pereira JS, Prado JC, Villa LL, Rohan TE, Franco EL. Cervical coinfection with human papillomavirus (HPV) types as a predictor of acquisition and persistence of HPV infection. *J Infect Dis* 2001;184:1508–1517.
- 18. Syrjänen S. HPV infections in children. *Papillomavirus Rep* 2003;14:93–109.
- 19. Manhart LE, Koutsky LA. Do condoms prevent genital HPV infection, external genital warts, or cervical neoplasia? A meta-analysis. *Sex Transm Dis* 2002:29:725–735.
- Koutsky LA, Holmes KK, Critchlow CW, et al. A cohort study of the risk of cervical intraepithelial neoplasia grade 2 or 3 in relation to papillomavirus infection. N Engl J Med 1992; 327:1272–1278.
- International Agency for Research on Cancer Working Group. Human papillomaviruses (HPV). *IARC Monographs* 1995;64.
- Schlecht NF, Kulaga S, Robitaille J, et al. Persistent human papillomavirus infection as a predictor of cervical intraepithelial neoplasia. *JAMA* 2001;286:3106–3114.
- 23. Moscicki AB, Hills N, Shiboski S, et al. Risks for incident human papillomavirus infection and low-grade squamous intraepithelial lesion development in young females. *JAMA* 2001;285:2995–3002.
- 24. Sigurdsson K. The Icelandic and Nordic cervical screening programs: trends in incidence and mortality rates through 1995. *Acta Obstet Gynecol Scand* 1999;78:478–485.
- Nieminen P, Kallio M, Anttila A, Hakama M. Organised vs. spontaneous Pap-smear screening for cervical cancer: a case-control study. *Int J Cancer* 1999;83:55–58.
- Parkin DM, Nguyen-Dinh X, Day NE. The impact of cervical screening on the incidence of cervical cancer in England and Wales. *Br J Obstet Gynaecol* 1985;92:150–157.
- 27. Van Ranst MA, Tachezy R, Delius H, Burk RD. Taxonomy of the human papillomaviruses. *Papillomavirus Rep* 1993;4:61.

- Noorani HZ, Brown A, Skidmore B, Stuart GCE. Liquid-based cytology and human papillomavirus testing in cervical cancer screening. Ottawa, ON: Canadian Coordinating Office for Health Technology Assessment; 2003. Technology Report No. 40.
- ACOG Committee on Practice Bulletins. ACOG Practice Bulletin: clinical management guidelines for obstetrician-gynecologists. Number 45, August 2003. Cervical cytology screening (replaces committee opinion 152, March 1995). Obstet Gynecol 2003;102:417–427.
- 30. Sasieni P, Adams J, Cuzick J. Benefits of cervical screening at different ages: evidence from the UK audit of screening histories. *Br J Cancer* 2003;89:88–93.
- 31. Anttila A, Ronco G, Clifford G, et al. Cervical cancer screening programmes and policies in 18 European countries. *Br J Cancer* 2004;91:935–941.
- 32. Cervical Cancer Prevention Network. *Programmatic Guidelines for Screening for Cancer of the Cervix in Canada*. Ottawa, ON: Health Canada and the Society of Obstetricians and Gynecologists of Canada; 1998.
- 33. Health Canada. Cervical Cancer Screening in Canada: 1998 Surveillance Report. Ottawa, ON: Health Canada; 2002.
- 34. Hawes SE, Critchlow CW, Faye Niang MA, et al. Increased risk of high-grade cervical squamous intraepithelial lesions and invasive cervical cancer among African women with human immunodeficiency virus type 1 and 2 infections. *J Infect Dis* 2003:188:555–563.
- 35. Paterson ME, Peel KR, Joslin CA. Cervical smear histories of 500 women with invasive cancer in Yorkshire. *BMJ* 1984;289:896–898.
- Martin-Hirsch P, Lilford R, Jarvis G, Kitchener HC. Efficacy of cervical-smear collection devices: a systematic review and meta-analysis. *Lancet* 1999; 354:1763–1770.
- NCI Bethesda System 2001. 2001 terminology. Available at: http://bethesda2001.cancer.gov/terminology.html. Accessed January 5, 2006.
- Arbyn M, Buntinx F, Van Ranst M, Paraskevaidis E, Martin-Hirsch P, Dillner J. Virologic versus cytologic triage of women with equivocal Pap smears: a meta-analysis of the accuracy to detect high-grade intraepithelial neoplasia. J Natl Cancer Inst 2004;96:280–293.
- 39. Stuart G, Taylor G, Bancej CM, et al. Report of the 2003 Pan-Canadian Forum on Cervical Cancer Prevention and Control. *J Obstet Gynecol Can* 2004;26:1004–1014.
- Wright TC Jr, Cox JT, Massad LS, Twiggs LB, Wilkinson EJ; ASCCP-Sponsored Consensus Conference. 2001 consensus guidelines for the management of women with cervical cytological abnormalities. *JAMA* 2002;287:2120–2129.
- 41. Sexually transmitted diseases treatment guidelines 2002. Centers for Disease Control and Prevention. *MMWR Recomm Rep* 2002;51(RR-6):1–78.
- 42. Chao A, Lin CT, Hsueh S, et al. Usefulness of human papillomavirus testing in the follow-up of patients with high-grade cervical intraepithelial neoplasia after conization. *Am J Obstet Gynecol* 2004;190:1046–1051.

- 43. von Krogh G, Lacey CJ, Gross G, Barrasso R, Schneider A. European course on HPV associated pathology: guidelines for primary care physicians for the diagnosis and management of anogenital warts. *Sex Transm Infect* 2000:76:162–168.
- 44. Tyring SK, Arany I, Stanley MA, et al. A randomized, controlled, molecular study of condylomata acuminata clearance during treatment with imiquimod. *J Infect Dis* 1998:178:551–555.
- 45. Kirby P, Dunne A, King DH, Corey L. Double-blind randomized clinical trial of self-administered podofilox solution vehicle in the treatment of genital warts. *Am J Med* 1990; 88:465–470.
- 46. Simmons PD, Langlet F, Thin RN. Cryotherapy versus electrocautery in the treatment of genital warts. *Br J Vener Dis* 1981;57:273–274.
- 47. Godley MJ, Bradbeer CS, Gellan M, Thin RN. Cryotherapy compared with trichloroacetic acid in treating genital warts. *Genitourin Med* 1987;63:390–392.
- 48. Abdullah AN, Walzman M, Wade A. Treatment of external genital warts comparing cryotherapy (liquid nitrogen) and trichloroacetic acid. *Sex Transm Dis* 1993;20:344–345.
- 49. Gross GE, Barasso R, eds. *Human Papillomavirus Infection: A Clinical Atlas.* Wiesbaden: Ullstein Mosby; 1997.
- 50. Sarfati D, Cox B, Jones RW, Sopoaga T, Rimeme C, Paul C. National audit of women with abnormal cervical smears in New Zealand. *Aust N Z J Obstet Gynaecol* 2003;43:152–156.
- 51. Peterson NB, Han J, Freund KM. Inadequate follow-up for abnormal Pap smears in an urban population. *J Natl Med Assoc* 2003;95:825–832.
- 52. Gage JC, Ferreccio C, Gonzales M, Arroyo R, Huivin M, Robles SC. Follow-up care of women with an abnormal cytology in a low-resource setting. *Cancer Detect Prev* 2003;27: 466–471.

GONOCOCCAL INFECTIONS

Etiology

· Caused by Neisseria gonorrhoeae.

Epidemiology

- Preliminary data show that there were approximately 9,200 reported cases of gonorrhea in 2004. Most affected are males 20–24 years of age (reported rate of 127.6/100,000) and females 15–19 years (reported rate of 126.7/100,000).¹ (Preliminary data — is subject to change; does not include Nunavut.)
- There has been a gradual but steady increase in gonococcal infections since 1998. It appears that a network of people with high-transmission activities play a key role in current prevalence levels. Case finding and partner notification are critical strategies for controlling this infection.
- The proportion of penicillin-resistant organisms is >1% in most areas of Canada and may reach 15% or higher in certain urban and rural areas.²
 - Numbers of isolates resistant to tetracyclines or a combination of penicillin
 and tetracyclines are high, and these antimicrobial agents should *not* be
 considered in the treatment of gonorrhea.
 - Quinolone resistance in Canada has been steadily increasing, from 1% in the late 1990s to a rate of 6.2% in 2004.²⁻⁴ This rate reflects samples which have been submitted by individual provinces and territories to the National Microbiology Laboratory (NML). The current rate reported by the NML may not truly reflect the national picture as the submission of samples from individual provinces and territories is voluntary and not standardized across the country. The shift from culture to NAATs has also created difficulty in providing an accurate picture for resistance across Canada as the availability of samples for resistance testing is becoming increasingly limited.
 - Quinolone resistance in certain regions of Canada is significantly higher than the national rate. Please check with your local public health officials to learn about quinolone resistance in your area.
 - Continued monitoring for antimicrobial resistance is important for ensuring high cure rates for this treatable infection.^{5,6}
- HIV transmission is enhanced in people with concomitant gonococcal infections.⁷
- · People at risk:
 - Those who have had contact with a person with proven infection or a compatible syndrome.
 - Those who have had unprotected sex with a partner originating from an area with high endemicity (there is also a higher risk of resistance in this population).
 - Travellers to an endemic country who have had unprotected sex with a resident of that area (there is also a higher risk of resistance in this population).
 - Sex workers and their sexual partners.

- Sexually active youth <25 years of age with multiple partners.
- Street-involved youth.
- Men who have unprotected sex with men.
- Previous gonorrhea and other STI.

Prevention

- Patients presenting with concerns about sexually transmitted infections (STIs) and/or prevention of pregnancy should be provided with instructions and encouragement about the consistent practice of safer-sex.
- At the time of diagnosis, review and monitor prevention practices.
- Identify barriers to prevention practices and the means to overcome them.
- See Primary Care and Sexually Transmitted Infections chapter.
- Provide counselling for the prevention of reproductive sequelae.
- Patients and contacts should abstain from unprotected intercourse until treatment of both partners is complete (i.e., 7 days after single-dose therapy).

Manifestations

Table 1. Manifestations

Neonates and infants	Children	Youth and adults		
mants		Females	Males	Females and males
 Ophthalmia Neonatal amniotic fluid infection Disseminated gonococcal infection 	 Urethritis Vaginitis Conjunctivitis Pharyngeal infection Proctitis Disseminated gonococcal infection 	 Cervicitis Pelvic inflammatory disease Urethritis Perihepatitis Bartholinitis 	Urethritis Epididymitis	 Pharyngeal infection Conjunctivitis Proctitis Disseminated gonococcal infection: arthritis, dermatitis, endocarditis, meningitis

Table 2. Symptoms of genital tract infection with N gonorrhoeae8-10

Neonates	Females	Males
Conjunctivitis Sepsis	 Vaginal discharge Dysuria Abnormal vaginal bleeding Lower abdominal pain Rectal pain and discharge if proctitis (see Sexually Transmitted Intestinal and Enteric Infections chapter) Deep dyspareunia 	 Urethral discharge Dysuria Urethral itch Testicular pain, swelling or symptoms of epididymitis Rectal pain and discharge if proctitis (see Sexually Transmitted Intestinal and Enteric Infections chapter)

Notes:

- Usual incubation period, 2-7 days.
- Many patients are asymptomatic or have symptoms not recognized to be due to N gonorrhoeae.
- Contacts are also likely to be asymptomatic.
- · Long-term carriage occurs.

Table 3. Major sequelae

Females	Males
 Pelvic inflammatory disease Infertility Ectopic pregnancy Chronic pelvic pain Reiter syndrome Disseminated gonococcal infection 	Epididymo-orchitisReiter syndromeInfertility (rare)Disseminated gonococcal infection

Diagnosis¹¹

Laboratory diagnosis

- Cultures obtained less than 48 hours after exposure may be negative.
- If possible, culture is the recommended method, because it allows for antimicrobial susceptibility testing. It is recognized that nucleic acid amplification tests (NAATs)* are the only available method in some jurisdictions. NAATs may be most useful when patients resist pelvic examination or urethral swabbing.¹² In these situations, urine NAATs should be used.

- Culture is especially important in the following cases:
 - Sexual abuse of children (rectal, pharyngeal, vaginal).[†]
 - Sexual assault.[†]
 - Treatment failure.
 - Evaluation of pelvic inflammatory disease (PID).
 - Infection acquired overseas or in areas with recognized antimicrobial resistance.
- Antimicrobial susceptibility testing for all isolates is suggested and is required for all isolates from positive (test of cure) follow-up cultures and treatment failures.
- Non-culture tests are an ideal method when transport and storage conditions
 are not conducive to maintaining the viability of N gonorrhoeae¹³ (see Laboratory
 Diagnosis of Sexually Transmitted Infections chapter).
- NAATs may be considered, but measures should be taken to ensure continued surveillance for antimicrobial resistance. If these tests are used for a test of cure, specimen collection should be delayed for 2–3 weeks post-treatment.¹⁴

Notes:

- * NAATs include polymerase chain reaction, ligase chain reaction, transcription mediated assay and strand displacement amplification.
- [†] When NAATs are used, two different primers should be used in the laboratory (see *Laboratory Diagnosis of Sexually Transmitted Infections* chapter).

Specimen collection^{11,13}

Routine specimen sites

- Urethra in young and adult males, with/without meatal discharge (see *Laboratory Diagnosis of Sexually Transmitted Infections* chapter).
 - For prepubertal boys, see Laboratory Diagnosis of Sexually Transmitted Infections and Sexual Abuse in Peripubertal and Prepubertal Children chapters.
- Cervix in young and adult females (see *Laboratory Diagnosis of Sexually Transmitted Infections* chapter).
- Rectum in females and in men who have sex with men (see *Laboratory Diagnosis* of *Sexually Transmitted Infections* chapter).
 - Colonization can occur without anal intercourse.¹⁵
- Vagina in prepubertal girls (see Laboratory Diagnosis of Sexually Transmitted Infections and Sexual Abuse in Peripubertal and Prepubertal Children chapters).
- Pharynx in those with a history of oral-genital contact (see *Laboratory Diagnosis* of *Sexually Transmitted Infections* chapter).
- Urine (first 10–20 mL) for NAAT if culture is not available, patient is resistant to
 pelvic examination or urethral swabbing, or problems exist with storage and
 transport of specimen.

Table 4. Specimen collection

Site/specimen	Test	Comments	
Urethra (intraurethral) (young and adult	Gram stain (for intracellular diplococci) (symptomatic men only)	Generally diagnostic of gonorrhea	
males)	• Culture	Confirmation and antimicrobial susceptibility testing	
	Non-culture test (NAAT)	In cases where culture not practical (does not provide antibiotic susceptibility)	
Endocervix/ urethra (young and adult females)	Gram stain (for intracellular diplococci)	Sensitivity lower than in male urethral specimens and not routinely recommended	
Terridies)	• Culture	Confirmation and antimicrobial susceptibility testing	
	Non-culture test (NAAT)	In cases where culture not practical (does not provide antibiotic susceptibility)	
Vagina	• Culture	Confirmation and antimicrobial susceptibility testing	
	Non-culture test (NAAT)	In cases where culture not practical (does not provide antibiotic susceptibility)	
Pharynx/ conjunctiva/ rectum	 Culture (Gram stain and non-culture tests not suitable for these sites) NAATs are not approved in Canada for oropharyngeal or rectal use. For conjunctiva and rectum, refer to package insert 	Confirmation and antimicrobial susceptibility testing	
Urine (males and females)	Non-culture test (NAAT)	Should not be used in cases of treatment failure when antimicrobial susceptibility data are critical	

Table 4. Specimen collection (continued)

Site/specimen	Test	Comments
Disseminated infection	Genital testingBlood cultureGram stain and culture of skin lesionSynovial fluid if arthritis	

NAAT=nucleic acid amplification test

Notes

- Specimens should be taken for the diagnosis of both gonococcal and chlamydial infections (see Laboratory Diagnosis of Sexually Transmitted Infections chapter).
- All suspected treatment failures must be investigated with a culture to ensure the availability of antimicrobial susceptibility data

Other sites

- If the cervix has been surgically removed, urine and vaginal swabs are convenient specimens; specimens can also be collected from the rectum and urethra.
- Self-obtained vaginal swabs may be suitable for women who refuse pelvic examination.
- Women undergoing laparoscopy for investigation of PID should have intraabdominal specimens taken (i.e., fallopian tube, cul de sac fluid etc.).
- Urethra in women with urethral syndrome.
- Blood and synovial fluid (in blood culture bottle) in disseminated disease.
 Synovial fluid should also be examined by Gram stain.
- Epididymal aspirate in men with epididymitis may be considered.
- Conjunctiva for ocular infection.

Note:

For further information on specimen transport, see *Laboratory Diagnosis of Sexually Transmitted Infections* chapter.

Transport

- Contact the laboratory for specific instructions regarding the preferred method of specimen transport to ensure pathogen survival for purposes of culture.
- Transport of gonococcal specimens for culture should be at ambient temperature, *not* 4°C as recommended for other organisms.

Management

- Management choices should be based on the site of infection and laboratory results.
- A diagnosis of gonorrhea should be confirmed by the identification of N gonorrhoeae by culture, or if culture is not available, by NAATs. All confirmed or suspected cases must be treated.

Table 5. Management: test results available

Gram stain	 Treat for gonococcal and chlamydial infection if Gram-negative intracellular diplococci observed The presence of Gram-negative diplococci outside PMNs is an equivocal finding that must be confirmed by culture The presence of PMNs without diplococci does not indicate or exclude gonococcal infection
Culture test	Treat all positives
NAATs	A positive test is diagnostic of gonorrhea, and the patient should be treated

NAAT=nucleic acid amplification test PMN=polymorphonuclear leukocyte

Table 6. Management: test results unavailable

Urethral/cervical mucopurulent discharge observed	Treat for N gonorrhoeae and C trachomatis
No urethral/cervical mucopurulent discharge	 Defer therapy until smear/culture/NAAT results available OR Treat for <i>N gonorrhoeae</i> and <i>C trachomatis</i> if follow-up uncertain and history and symptoms suggestive, or if partner is infected

NAAT=nucleic acid amplification test

Treatment

- All patients treated for gonorrhea should also be treated for chlamydial infection, unless a chlamydia test result is available and negative.
- Directly observed therapy with single-dose regimens is desirable if poor compliance is expected.
- For PID, see Pelvic Inflammatory Disease chapter.
- For epididymitis, see *Epididymitis* chapter.

Youth and adults

Table 7: Urethral, endocervical, rectal, pharyngeal infection (except in pregnant women and nursing mothers)^{16–22}

Alternative ONLY if use of quinolones not recommended and cephalosporin allergy OR immediate/anaphylactic penicillin allergy²³ • Cefixime 400 mg PO in a single dose^{†B} [A-I] Azithromycin 2 g PO in a single dose[¶] [A-I] OR Ciprofloxacin 500 mg PO in a single dose^{‡§} Spectinomycin 2 g IM in a single dose^b (unless not recommended due to quinolone (available only through Special Access resistance) [A-I] Program [SAP]) [A-I] OR • Ofloxacin 400 mg PO in a single dose^{‡§} (unless not recommended due to quinolone resistance) [A-I] OR • Ceftriaxone 125 mg IM in a single dose^{†¥B}[A-I]

All regimens should be followed by empiric treatment for chlamydial and non-gonococcal infections (see *Chlamydial Infections* and *Urethritis* chapters)

- * Other broad-spectrum quinolones are effective but not recommended as first-line agents because of their cost.
- † Cefixime and ceftriaxone should not be given to persons with a cephalosporin allergy or a history of immediate and/or anaphylactic reactions to penicillins.
- ‡ Contraindicated in pregnant and lactating women.
 - § Quinolones are not recommended if the case or contact are from, or are epidemiologically linked to, any area with rates of quinolone-resistant N aonorrhoeae > 3-5%:
 - Asia
 - · Pacific Islands (including Hawaii)
 - India
 - Israel
 - Australia
 - · United Kingdom
 - Regions of the United States (check with the U.S. Centers for Disease Control and Prevention for rates of quinolone resistance by geographic area)
 - · MSM with contact or epidemiologically linked to the United States
 - Areas in Canada experiencing high rates of quinolone resistance; current numbers provided by the National Microbiology
 Laboratory place Quebec, Ontario, Alberta and British Columbia above the 3% threshold for quinolone resistance (see
 Epidemiology section, above, for an explanation on national and regional quinolone resistance rates). Please check with
 your local public health officials to learn about quinolone resistance in your area. In Alberta all ciprofloxacin resistant
 cases in 2004–05 were in MSM or linked to travel outside of Alberta, therefore ciprofloxacin remains a recommended
 agent for the treatment of gonorrhea in Alberta except in these situations (source: Alberta Health and Wellness STD
 Services). For data on national quinolone resistance in Canada, please visit the Public Health Agency of Canada website
 (www.phac-aspc.gc.ca).
 - ¥ The preferred diluent for ceftriaxone is 1% lidocaine without epinephrine (0.9 mL/250 mg, 0.45 mL/125 mg) to reduce discomfort.
 - ¶ Associated with a significant incidence of gastrointestinal adverse effects. Taking medication with food may minimize adverse effects. Antiemetics may be needed.
- P Not effective for pharyngeal infection. Test of cure is recommended.
- ß Cefixime is preferred over ceftriaxone as a factor of cost and ease of administration.

Table 8. Urethral, endocervical, rectal or pharyngeal infection in pregnant women and nursing mothers²⁴⁻²⁶

Preferred	Alternatives
• Cefixime 400 mg PO in a single dose* [A-I]	 Ceftriaxone 125 mg IM in a single dose *† [A-I] OR Spectinomycin 2 g IM in a single dose[‡] (available only through SAP) [A-I]

All regimens should be followed by empiric treatment for chlamydial and non-gonococcal infections (see *Chlamydial Infections* and *Urethritis* chapters)

SAP=Special Access Program

- * Cefixime and ceftriaxone should not be given to persons with a cephalosporin allergy or a history of immediate and/or anaphylactic reactions to penicillins.
- † The preferred diluent for ceftriaxone is 1% lidocaine without epinephrine (0.9 mL/250 mg, 0.45 mL/125 mg) to reduce discomfort.
- ‡ Not effective for pharyngeal infection. Test of cure is recommended.

Table 9. Gonococcal ophthalmia/disseminated infection (arthritis, meningitis)

Preferred initial therapy

Ceftriaxone 2 g/day IV/IM AND doxycycline/azithromycin while awaiting consultation* [A-II]

- · Consultation with a colleague experienced in this area is essential
- Hospitalization is necessary for meningitis and may be necessary for other disseminated infections
- The preferred diluent for IM ceftriaxone is 1% lidocaine without epinephrine (0.9 mL/250 mg, 0.45 mL/125 mg) to reduce discomfort.

Children under 9 years of age^{8,27}

Table 10. Urethral, vaginal, rectal, pharyngeal infection

Preferred	Alternative
 Cefixime 8 mg/kg PO in a single dose (maximum 400 mg)*[†] [A-II] OR Ceftriaxone 125 mg IM in a single dose^{†‡} [A-II] 	Spectinomycin 40 mg/kg IM (maximum 2 g) in a single dose [A-II]

- All regimens should be followed by treatment for chlamydial infection. See *Chlamydial Infections* chapter for treatment recommendations for children under 9 years of age.
- * Oral therapies are preferred in children. Recommendations for the use of cefixime are based on data showing efficacy in the treatment of infections caused by organisms similar to *N gonorrhoeae*. Because there is limited experience with the use of cefixime in children with gonococcal infections, antimicrobial susceptibility must be ascertained and a follow-up culture ensured. If follow-up cannot be ensured, use ceftriaxone 125 mg IM in place of cefixime.
- † Cefixime and ceftriaxone should not be given to persons with a cephalosporin allergy or a history of immediate and/or anaphylactic reactions to penicillins.
- ‡ The preferred diluent for ceftriaxone is 1% lidocaine without epinephrine (0.9 mL/250 mg, 0.45 mL/125 mg) to reduce discomfort.

Table 11. Disseminated infection

Infection	Preferred treatment		
Arthritis	Ceftriaxone 50 mg/kg IV/IM in a single daily dose for 7 days* [A-III]		
Meningitis, endocarditis	Ceftriaxone 25 mg/kg IV/IM every 12 hours for 10–14 days for meningitis, 28 days for endocarditis* [A-III]		
Gonococcal ophthalmia beyond neonatal period	Ceftriaxone 50 mg/kg IV/IM in a single dose (maximum 1 g)* [A-III]		
Hospitalization and consultation with a colleggue experienced in this area is assential			

Hospitalization and consultation with a colleague experienced in this area is essential

^{*} The preferred diluent for IM ceftriaxone is 1% lidocaine without epinephrine (0.9 mL/250 mg, 0.45 mL/125 mg) to reduce discomfort.

Neonatal infection

Ophthalmia neonatorum

- Hospitalize and institute appropriate infection-control precautions until 24 hours of effective therapy completed.
- Culture eye discharge, blood (cerebrospinal fluid only if evidence of systemic disease).
- Irrigate eyes immediately with sterile normal saline and at least hourly as long as necessary to eliminate discharge.
- Start ceftriaxone 100 mg/kg IV or IM single-dose therapy [A-II].
- Consult with a colleague experienced in this area as soon as possible.

Table 12. Neonates born to women infected with gonorrhea

Recommended therapy (must also include therapy for chlamydia for 14 days unless the mother's tests are negative)

Ceftriaxone 125 mg IM in a single dose AND erythromycin in the following dosage schedule*† [A-III]:

- If ≤7 days old and ≤2000 g: erythromycin 20 mg/kg/day PO in divided doses† [A-III]
- If ≤7 days old and >2000 g: erythromycin 30 mg/kg/day PO in divided doses[†] [A-III]
- If >7 days of age: erythromycin 40 mg/kg/day PO in divided doses† [A-III]
- * The preferred diluent for ceftriaxone is 1% lidocaine without epinephrine (0.9 mL/250 mg, 0.45 mL/125 mg) to reduce discomfort.
- † Erythromycin dosages refer to erythromycin base. Equivalent dosages of other formulations may be substituted. The use of erythromycin in infants under 6 weeks of age has been associated with infantile hypertrophic pyloric stenosis (IHPS). 28-31 The risk of IHPS with other macrolides (e.g., azithromycin) is unknown. The risks and benefits of using erythromycin in such infants must be explained to parents. When erythromycin is used, it is important to monitor for symptoms and signs of IHPS. IHPS following erythromycin use should be reported to the Canadian Adverse Drug Reaction Monitoring Program at 1-866-234-2345.

Consideration for Other STIs

- See Primary Care and Sexually Transmitted Infections chapter.
- Obtain a specimen for the diagnosis of chlamydial infection.
- Obtain a blood sample for serologic testing of syphilis (see *Syphilis* chapter).
- HIV counselling and testing are recommended (see *Human Immunodeficiency Virus Infections* chapter).
- Immunization against hepatitis B is recommended, if not already immune (see *Hepatitis B Virus Infections* chapter).

Reporting and Partner Notification

- With the changing epidemiology of N gonorrhoeae, case finding and partner notification are critical strategies for maintaining control of gonococcal infections in Canada.
- Gonococcal infections are reportable in all provinces and territories.
- Positive culture and non-culture tests must be reported to the local public health authorities.
- All partners who have had sexual contact with the index case within at least 60 days prior to symptom onset or date of diagnosis where asymptomatic; parents of infected neonates (i.e., mother and her sexual partner), and persons implicated in sexual abuse cases must be located, clinically evaluated and treated.
- Since co-infections are common, persons treated for gonococcal infections should also be treated for *C trachomatis*, unless concurrent tests for chlamydia are negative.
- Local public health authorities are available to assist with partner notification and with appropriate referral for clinical evaluation, testing, treatment and health education.

Follow-up

- Repeat screening of individuals with gonorrhea after 6 months is recommended.
- Follow-up testing by culture *must* be completed if any of the following exist:
 - Treatment failure has occurred previously.
 - Antimicrobial resistance to therapy is documented.
 - Compliance is uncertain.
 - There is re-exposure to an untreated partner.
 - There is concern over a false-positive non-culture test result.
 - Infection occurs during pregnancy.
 - PID or disseminated gonococcal infection is diagnosed.
 - Patient is a child.

Notes:

- Follow-up cultures for test of cure are indicated approximately 4–5 days after the completion of therapy. These should include reculturing of all positive sites.
- NAATs are not recommended for test of cure. However, if this is the only choice, tests should not be done for 3 weeks after treatment to avoid false-positive results due to the presence of non-viable organisms.

Special Considerations

Children

- Neonates born to infected mothers must be tested and treated.
- Sexual abuse must be considered when genital, rectal or pharyngeal gonorrhea
 is diagnosed in any child after the neonatal period. Consultation with a
 colleague experienced in such cases should be sought. Siblings and other
 children possibly at risk must also be evaluated.
- Sexual abuse of children must be reported to the local child protection agency.
- Local public health authorities may be helpful in evaluating the source of infection and spread to others. See Sexual Abuse in Peripubertal and Prepubertal Children chapter.

Notes:

- Follow-up cultures for test of cure are indicated approximately 4–5 days after the completion of therapy. These should include reculturing of all positive sites.
- NAAT is not recommended for test of cure. However if this is the only choice, tests should not be done for 3 weeks after treatment to avoid false-positive results due to the presence of non-viable organisms.

References

- 1. Unpublished data. Surveillance and Epidemiology Section, Community Acquired Infections Division, Public Health Agency of Canada, 2006.
- 2. Mann J, Kropp R, Wong T, et al. Gonorrhea treatment guidelines in Canada: 2004 update. *CMAJ* 2004;171:1345–1346.
- Sarwal S, Wong T, Sevigny C, Ng LK. Increasing incidence of ciprofloxacin resistant Neisseria gonorrhoeae infection in Canada. CMAJ 2003;168:872–873.
- 4. National Microbiology Laboratory, Public Health Agency of Canada, unpublished data, 2005.
- Tapsall JW, Limnios EA, Shultz TR. Continuing evolution of the pattern of quinolone resistance in Neisseria gonorrhoeae isolated in Sydney, Australia. Sex Transm Dis 1998;25:415–417.
- Ng LK, Sawatzky P, Martin IE, Booth S. Characterization of ciprofloxacin resistance in Neisseria gonorrhoeae isolates in Canada. Sex Transm Dis 2002;29:780–788.
- Laga M, Manoka A, Kivuvu M, et al. Nonulcerative sexually transmitted diseases as risk factors for HIV-1 transmission in women; results from a cohort study. AIDS 1993;7:95–102.
- 8. Sung L, MacDonald NE. Gonorrhea: a pediatric perspective. *Pediatr Rev* 1998;19:13–22.
- Korenromp EL, Sudaryo MK, de Vlas SJ, et al. What proportion of episodes of gonorrhea and chlamydia become symptomatic? *Int J STD AIDS* 2002; 13:91–101.

- Mehta SD, Rothman RE, Kelen GD, Quinn TC, Zenilman JM. Clinical aspects of diagnosis of gonorrhea and chlamydia infection in an acute care setting. Clin Infect Dis 2001;32:655–659.
- 11. Johnson RE, Newhall WJ, Papp JR, et al. Screening tests to detect Chlamydia trachomatis and Neisseria gonorrhoeae infections 2002. *MMWR Recomm Rep* 2002;51(RR-15):1–38.
- 12. Davies PO, Low N, Ison CA. The role of effective diagnosis for the control of gonorrhoea in high prevalence populations. *Int J STD AIDS* 1998;9:435–443.
- Koumans EH, Johnson RE, Knapp JS, St. Louis ME. Laboratory testing for Neisseria gonorrhoeae by recently introduced nonculture tests: a performance review with clinical and public health considerations. *Clin Infect Dis* 1998;27:1171–1180.
- Bachmann LH, Desmond RA, Stephens J, Hughes A, Hook EW 3rd. Duration of persistence of gonococcal DNA detected by ligase chain reaction in men and women following recommended therapy for uncomplicated gonorrhea. *J Clin Microbiol* 2002;40:3596–3601.
- 15. McCormack WM, Stumacher RJ, Johnson K, Donner A. Clinical spectrum of gonococcal infections in women. *Lancet* 1977;1:1182–1185.
- Burstein GR, Berman SM, Blumer JL, Moran JS. Ciprofloxacin for the treatment of uncomplicated gonorrhea infection in adolescents: does the benefit outweigh the risk? Clin Infect Dis 2002;35(suppl 2):S191–S199.
- 17. Dan M, Poch F, Sheinberg B. High prevalence of high-level ciprofloxacin resistance in Neisseria gonorrhoeae in Tel Aviv, Israel: correlation with response to therapy. *Antimicrob Agents Chemother* 2002;46:1671–1673.
- Aplasca de los Reyes MR, Pato-Mesola V, Klausner JD, et al. A randomized trial of ciprofloxacin versus cefixime for treatment of gonorrhea after rapid emergence of gonococcal ciprofloxacin resistance in the Philippines. *Clin Infect Dis* 2001;32:1313–1318.
- 19. Jones RB, Schwebke J, Thorpe EM Jr, Dalu ZA, Leone P, Johnson RB. Randomized trial of trovafloxacin and ofloxacin for single dose therapy of gonorrhea. Trovafloxacin Gonorrhea Study Group. *Am J Med* 1998;104:28–32.
- Stoner BP, Douglas JM Jr, Martin DH, et al. Single-dose gatifloxacin compared with ofloxacin for the treatment of uncomplicated gonorrhea: a randomized, double-blind, multicenter trial. Sex Transm Dis 2001;28:136–142.
- 21. Robinson AJ, Ridgway GL. Concurrent gonococcal and chlamydial infection: how best to treat. *Drugs* 2000;59:801–813.
- 22. Tapsall J. Current concepts in the management of gonorrhoea. *Expert Opin Pharmacother* 2002;3:147–157.
- Handsfield HH, Dalu ZA, Martin DH, Douglas JM Jr, McCarty JM, Schlossberg D. Multicenter trial of single dose azithromycin vs ceftriaxone in the treatment of uncomplicated gonorrhea. Azithromycin Gonorrhea Study Group. Sex Transm Dis 1994;21:107–111.
- Ramus RM, Sheffield JS, Mayfield JA, Wendel GD Jr. A randomized trial that compared oral cefixime and intramuscular ceftriaxone for the treatment of gonorrhea in pregnancy. Am J Obstet Gynecol 2001;185:629–632.

- 25. Donders GG. Treatment of sexually transmitted bacterial diseases in pregnant women. *Drugs* 2000;59:477–485.
- 26. Brocklehurst P. Update on the treatment of sexually transmitted infections in pregnancy 1. *Int J STD AIDS* 1999;10:571–578.
- 27. American Academy of Pediatrics. Committee on Child Abuse and Neglect. Gonorrhea in prepubertal children. *Pediatrics* 1998;101(1 Pt 1):134–135.
- Sorensen HT, Skriver MV, Pedersen L, Larsen H, Ebbesen F, Schonheyder HC. Risk of infantile hypertrophic pyloric stenosis after maternal postnatal use of macrolides. *Scand J Infect Dis* 2003;35:104–106.
- 29. Cooper WO, Griffin MR, Arbogast P, Hickson GB, Gautam S, Ray WA. Very early exposure to erythromycin and infantile hypertrophic pyloric stenosis. *Arch Pediatr Adolesc Med* 2002;156:647–650.
- Mahon BE, Rosenman MB, Kleiman MB. Maternal and infant use of erythromycin and other macrolide antibiotics as risk factors for infantile hypertrophic pyloric stenosis. *J Pediatr* 2001;139:380–384.
- 31. Honein MA, Paulozzi LJ, Himelright IM, et al. Infantile hypertrophic pyloric stenosis after pertussis prophylaxis with erythromcyin: a case review and cohort study. *Lancet* 1999;354:2101–2105.

HEPATITIS B VIRUS INFECTIONS

Etiology

 Hepatitis B is a viral disease characterized by infection of the liver by the hepatitis B virus (HBV), a small DNA virus of the family Hepadnaviridae. The virus occurs worldwide, with greatest prevalence in the developing world.

Epidemiology

- Most common cause of sexually transmitted hepatitis.
- Incubation period ranges from days after percutaneous exposure to 4–8 weeks after mucous membrane exposure.
- Incidence of acute hepatitis B in Canada is estimated to be 2.3 per 100,000.1
 - Incidence of acute hepatitis B in men is twice as high as in women (3.0/100,000 vs. 1.5/100,000, respectively).
 - Peak incidence rates are found in those aged 30–39 (6.1/100,000).
- Prevalence of hepatitis B in Canada is estimated to be 0.5–1.0%.²
- Prevalence of chronic hepatitis B varies in different populations:
 - Immigrants: 7.4%³
 - Inuit: 6.9%⁴
 - First Nations: 0.3%⁵
 - Sexually transmitted infection (STI) clinic patients: 0.3%6
- · Routes of transmission:
 - Percutaneous, principally injection drug users.
 - Sexual: anal > vaginal > oral.
 - Horizontal: household contacts.
 - Vertical: mother to neonate.
- Risk factors for acquisition:⁷
 - Injection drug use (IDU): 34%
 - Multiple heterosexual sex partners: 24%
 - Men who have sex with men (MSM): 7.3%
 - Sex with HBV-infected individuals: 12%
 - Hepatitis B carrier in family: 2.4%
- Prior to donor screening, blood and blood products were important sources of infection in Canada and may still be in countries where the quality of the blood supply is questionable.
- Populations at the highest risk include the following:
 - Infants born to hepatitis B surface antigen (HBsAg)-positive mothers.
 - Injection drug users who share drug injection/preparation equipment.
 - Those with multiple sex partners.
 - Those born in or having sexual contact in areas of high endemicity.
 - Sexual and household contacts of an acute case or chronic carrier.
 - Health care workers and others with occupational blood exposure.
 - Those who are incarcerated or institutionalized.
 - Those infected with HIV or hepatitis C virus (HCV).
 - Those with a previous STI.

Prevention

Primary prevention

- Counselling/education regarding risk behaviours.
- Harm-reduction strategies (needle exchanges, etc.).
- Hepatitis B vaccination (pre-exposure prophylaxis).
 - A school-based universal hepatitis B immunization program aimed at children aged 9–13 was implemented in all provinces and territories in the early 1990s.
 - An infant universal hepatitis B vaccination program is run in some provinces and territories, in addition to the school-based preadolescent immunization program.
 - Hepatitis B immunization should be routinely offered to the following risk groups (if not previously immunized):8
 - Children from HBV-endemic areas who may be exposed to HBV via extended family or the community.
 - Populations or communities in which HBV is highly endemic.
 - Residents and staff of institutions for the mentally or developmentally challenged.
 - · Sex workers.
 - · Hemodialysis patients.
 - People with hemophilia and others receiving repeated infusions of blood or blood products.
 - Household and sexual contacts of acute HBV cases and HBV carriers.
 - · Pregnant women.
 - Injection drug users.
 - · Staff and inmates of correctional facilities.
 - · Travellers to HBV-endemic areas.
 - Those who have recently acquired an STI.
 - Those whose regular sex partner is HBsAg-positive.
 - Those with multiple sex partners.
 - MSM.
 - Those with occupational risk (e.g., health care workers and emergency service workers who may be exposed to blood, blood products or body fluids that may contain the virus).
 - Children in childcare settings in which there is an HBV-infected child.
 - · People who are HIV-positive.
 - Sexual partners of any of those listed above.
 - Offer hepatitis B vaccine to all those in the above categories who do not show evidence of immunity [A-I] or do not have proof of immunization, and refer those showing evidence of chronic hepatitis B carriage for consideration for treatment with available agents [A-I].^{9,10} Some authorities suggest that preimmunization screening is not cost-effective in low-risk populations, particularly adolescents, and recommend immunization without screening tests;¹¹ with each passing year after the initiation of universal school-based immunization, screening will become increasingly cost-effective as the proportion of those not immunized diminishes.

Secondary prevention (post-exposure prophylaxis)

- Hepatitis B immune globulin (HBIG) can be given to recipients of percutaneous (needlestick) or mucosal exposure up to 7 days after exposure and to sexual contacts within 14 days of exposure (ideally within 48 hours), followed by hepatitis B vaccine.⁸
- For infants born to HBV-infected mothers, the first dose of hepatitis B vaccine should be administered within 12 hours of birth and HBIG immediately after birth (efficacy decreases sharply after 48 hours).⁸
 - See Figure 1 for an algorithm on the approach to a sexual (penile-anal, penile-vaginal or oral-genital) or percutaneous/mucosal exposure to a known hepatitis B carrier or a high-risk source.
 - Postimmunization screening for the antibody to hepatitis B surface antigen (anti-HBs) is generally not recommended, except for the following:⁸
 - · Infants born to infected mothers.
 - Sexual partners and household contacts of chronic carriers.
 - Those immunized for occupational exposure.
 - Those who are immunocompromised (i.e., lose their response).
 - · Hemodialysis patients.
 - · Pregnant women.

Manifestations and diagnosis

- Although HBV is hepatotropic and the liver is the sole site of infection, viremia may lead to clinical manifestations related to immune complex formation.
- All patients being assessed for STIs should be asked about their vaccination history, risk history, previous icteric illness and previous hepatitis testing.
- Acute hepatitis B infection is often not clinically apparent, with 50–70% of adult cases being asymptomatic. Symptomatic cases may be non-specific (fatigue, nausea, vomiting, anorexia, rash, arthralgia). A smaller proportion of cases are icteric; these can be clinically indistinguishable from other viral or toxic causes of hepatitis.
- Chronic hepatitis B can be detected by persistence of HBsAg, may or
 may not be associated with elevations in hepatic transaminases and is
 generally asymptomatic until clinical signs of cirrhosis, portal hypertension or
 hepatocellular carcinoma supervene.
- Hepatitis serologic testing can be done for a number of potential indications:
 - To diagnose acute infection in symptomatic persons.
 - To detect chronic infection in asymptomatic persons.
 - As a preimmunization screen to identify non-immune persons who may benefit from hepatitis B vaccination.
- See Table 1 for serologic markers for hepatitis B.

Table 1. Serologic markers for hepatitis B

Stage	HBsAg	НВеАд	Anti-HBc IgM	Anti-HBc IgG/total	Hepatitis B viral DNA	Anti HBs
Acute (early)	+	+	+	+	+	-
Acute (resolving)	+	-	+	+	-	-
Chronic	+	+/-	-	+	+/-	-
Resolved	-	-	-	+	-	+/-*
Vaccinated	-	-	-	-	-	+*

anti-HBc = antibody to hepatitis B core antigen

 $anti-HBs = antibody \ to \ hepatitis \ B \ surface \ antigen$

HBeAg = hepatitis B early antigen HBsAg = hepatitis B surface antigen

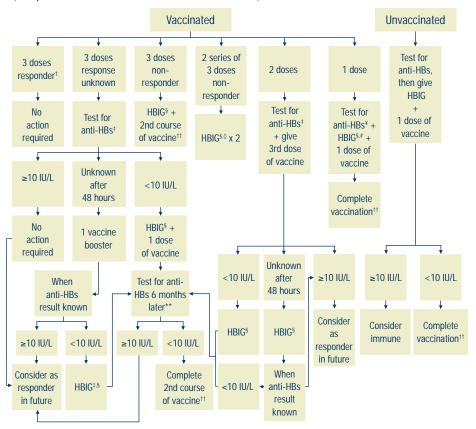
- The choice of serologic testing for suspected acute or chronic cases is
 determined by the clinical situation and should be supplemented by the addition
 of liver function testing and hepatic transaminases. For those who are HBsAgpositive, who may be in the window period before development of anti-HBs
 and anti-HBc antibodies, a positive anti-HBc IgM confirms that this is due to
 early infection.
- There is controversy surrounding the need to prescreen high-risk individuals before vaccination, as well as the optimal choice of serologic tests for screening. For those at high risk and for whom follow-up cannot be ensured, it is prudent to give the first dose of vaccine on the initial visit after drawing blood for screening serology.
- Evaluating the status of a high-risk person should not delay immunization.

^{*} In some patients, anti-HBs may decline over time and become undetectable.

Management

Figure 1. Management of sexual/percutaneous/mucosal exposure to infected (HBsAg-positive) or high-risk* source

(adapted from Canadian Immunization Guide)8

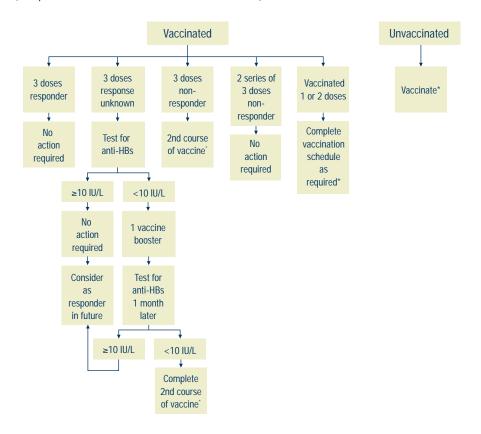


anti-HBs = antibody to hepatitis B surface antigen HBIG = hepatitis B immune globulin

- * A known source is high-risk if the person comes from a highly endemic region for HBV, has sexual relations with multiple partners, has a partner who is infected with HBV or is at high risk of being so, is in close family contact with an infected person, uses injection drugs or received blood or blood products prior to 1970. Wherever possible, the source should be tested. In the case of an unknown source, background circumstances may provide some indication of the degree of risk (e.g., syringe found in the street, attendance at an STI, detoxification or well-baby clinic).
- † Responder known to have ≥10 IU/L anti-HBs. No measures are required if the person has developed an immunity following an infection
- ‡ Anti-HBs titre should be determined as soon as possible to avoid needless administration of HBIG and because efficacy is unknown if given after 7 days for percutaneous/mucosal exposures and up to 14 days for sexual exposures.
- § The administration of HBIG can be omitted if the high-risk source can be tested within 48 hours and the result is negative. In that case, see Figure 2.
- ♦ The second dose of HBIG should be given 1 month after the first.
- ¥ This test does not change the continuation of vaccination, but may reassure the exposed individual about the immediate risk of becoming infected.
- # If it is possible to quickly obtain an anti-HBs titre confirming ≥10 IU/L, administration of HBIG should be omitted.
- ** Determination of anti-HBs titre should be delayed for 6 months to allow HBIG antibodies to wane.
- ††Test for anti-HBs 1–6 months after the course of vaccine.

Figure 2. Management of sexual/percutaneous/mucosal exposure to uninfected (HBsAg-negative) or low-risk source

(adapted from Canadian Immunization Guide)8



anti-HBs = antibody to hepatitis B surface antigen

^{*}Test for anti-HBs 1–6 months after the course of vaccine.

Treatment

- A discussion of the treatment of clinical hepatitis B is beyond the scope of these guidelines. Any patient known to have chronic hepatitis B should be referred to an expert for further management. Those wishing further details on initial workup of the patient with chronic hepatitis B are referred to Management of Viral Hepatitis: A Canadian Consensus Conference 2003/2004¹² and The Management of Chronic Viral Hepatitis: A Canadian Consensus Conference 2004.¹³ Some brief comments can be made:
 - There is no indication for antiviral intervention in acute hepatitis B.
 - Acute cases of hepatitis B should abstain from sexual contact or practice safer-sex until partners and/or relevant contacts have been appropriately screened and/or immunized.
 - In the case of chronic active hepatitis B, there are data to support the efficacy of interferon- α , lamivudine, famciclovir, adefovir, fibavirin and other agents under study. In Canada, most patients are managed with interferon- α and/or lamivudine (3TC) as primary therapeutic modalities [A-I].

Consideration for Other STIs

- Any patient with hepatitis B infection believed to have been acquired sexually should be considered to be at risk for other STIs, including HIV, and should be offered testing for gonorrhea, chlamydia, syphilis and HIV.
- Any patient with hepatitis B infection believed to have been acquired parenterally should be considered to be at risk for HIV and HCV, and should be offered testing for both.
- Concurrent HIV and hepatitis B infection can lead to more rapid progression
 of liver damage and is more likely to lead to chronic infection and impaired
 hepatic function, which may limit the therapeutic options for treatment of the
 HIV co-infection.¹⁷

Reporting, Partner Notification and Follow-up

- Acute hepatitis B is a reportable infection in all Canadian jurisdictions.
- Partner notification/contact tracing is essential to identify those at risk of acquiring hepatitis B, both to clarify their immune status and to provide vaccine protection to the non-immune. Contacts include the following:
 - Sexual and percutaneous exposures during the period of infectivity.
 - Children of hepatitis B-infected mothers who did not receive HBIG and vaccine at birth.
 - Those living in the household of the index case.

Special Considerations

- Pregnant women with no history of hepatitis B immunization should be screened at their initial prenatal visit for HBsAg. A pregnant woman who has no markers of acute or chronic HBV infection but who is at high risk of acquiring HBV should be offered vaccine at the first opportunity and tested for antibody response. Pregnancy is not a contraindication to immunization. If testing has not been done during pregnancy, it should be done at the time of delivery. Repeat testing before delivery may be considered in uninfected and non-immunized women with continuing high-risk behaviour. Infants born to HBsAg-positive women should receive postexposure prophylaxis.
- Children adopted from areas or family situations in which there is a high prevalence of HBV infection should be screened for HBsAg, and if they are positive, household contacts should be immunized before adoption.

References

- Zou S, Zhang J, Tepper M, et al. Enhanced surveillance of acute hepatitis B and acute Hepatitis C in four health regions in Canada 1998–1999. Can J Infect Dis 2001;12:345–350.
- Sherman M. Update 5: June 1996. The epidemiology of hepatitis B in Canada. The Hepatitis Information Network Web site. Available at: www.hepnet.com/ update5.html. Accessed January 9, 2006.
- 3. Delage G, Montplaisir S, Remy-Prince S, Pierri E. Prevalence of hepatitis B virus infection in pregnant women in the Montreal area. *CMAJ* 1986;134: 897–901.
- 4. Baikie M, Ratnam S, Bryant DG, et al. Epidemiologic features of hepatitis B virus infection in Northern Labrador. *CMAJ* 1989;141:791–795.
- Martin JD, Mathias RG. HIV and hepatitis B surveillance in First Nations alcohol and drug treatment centers in British Columbia, Canada. *Int J Circumpolar Health* 1998;57(suppl 1):280–284.
- Romanowski B, Campbell P. Sero-epidemiologic study to determine the prevalence and risk of hepatitis B in a Canadian heterosexual sexually transmitted disease clinic population. Can J Public Health 1994;85:205–207.
- 7. Zhang J, Zou S, Giulivi A. Viral hepatitis and blood-borne pathogens in Canada. Hepatitis B in Canada. *Canadian Commun Dis Rep* 2001;2753:10–12.
- 8. Canadian Immunization Guide. 6th ed. Ottawa, ON: Health Canada; 2002.
- Brook MG, Karayiannis P, Thomas HC. Which patients with chronic hepatitis B will respond to alpha interferon therapy? A statistical analysis of predictive factors. *Hepatology* 1989;10:761–763.
- 10. Nevens F, Main J, Honkoop P, et al. Lamivudine therapy for chronic hepatitis B: a six-month randomized dose-ranging study. *Gastroenterology* 1997;113:1258–1263.
- 11. Sexually transmitted diseases treatment guidelines 2002. Centers for Disease Control and Prevention. *MMWR Recomm Rep* 2002;51(RR-6):1–78.

- 12. Canadian Consensus Conference on the Management of Viral Hepatitis 2003/2004. Ottawa, ON: Health Canada and Correctional Service Canada; 2004. Available at: www.phac-aspc.gc.ca/hepc/hepatitis_c/pdf/ccc_04/index. html. Accessed: 10 January 2006.
- 13. Sherman M, Bain V, Villenueve J-P, et al. The management of chronic viral hepatitis: a Canadian consensus conference 2004. *Can J Infect Dis Med Microbiol* 2004;15:313–326.
- 14. Main J, Brown JL, Howells C, et al. A double-blind, placebo-controlled study to assess the effect of famciclovir on virus replication in patients with chronic hepatitis B virus infection. *J Vir Hepatitis* 1996;3:211–215.
- 15. Tsiang M, Rooney JF, Toole JJ, Gibbs CS. Biphasic clearance kinetics of hepatitis B virus from patients during adefovir dopivoxil therapy. *Hepatology* 1999:29:1863–1869.
- Cotonat T, Quiroga JA, Lopez-Alcorocho JM, et al. Pilot study of combination therapy with ribavirin and interferon alfa for the retreatment of chronic hepatitis B e antibody-positive patients. *Hepatology* 2000;31:502–506.
- 17. Rockstroh JK. Management of hepatitis B and C in HIV co-infected patients. *J Acquir Immune Defic Syndr* 2003;34 (suppl 1):S59–S65.

HUMAN IMMUNODEFICIENCY VIRUS INFECTIONS

Etiology^{1,2}

- The human immunodeficiency virus (HIV) has been shown to be the causative agent of acquired immunodeficiency syndrome (AIDS).
- Infection with HIV results in the progressive destruction of CD4+ T lymphocytes.
 These cells are crucial to the normal function of the human immune system.
- Persons with HIV infection and subsequent immune suppression are, therefore, at risk of developing a variety of clinical AIDS-defining conditions, including opportunistic infections (e.g., *Pneumocystis jiroveci* [formerly *Pneumocystis carinii*] pneumonia, disseminated *Mycobacterium avium* complex [MAC] disease), primary neurologic disease (e.g., AIDS dementia) and malignancy (e.g., lymphoma, Kaposi sarcoma) (see Table 3 for AIDS-defining conditions).

Epidemiology^{3,4}

- The HIV/AIDS epidemic is a complex one, with differing rates of infection in specific at-risk populations. The number of Canadians living with HIV infection continues to increase. There has been a 20% rise in the number of positive HIV test reports in Canada in the last 5 years (2000–2004).
- In 2004, men who have sex with men (MSM) still represented the largest number and proportion of positive HIV test reports; however, the heterosexual exposure category represents a growing number and proportion of positive HIV tests, surpassing injection drug use (IDU) as the second largest exposure category.
- Persons migrating to Canada from countries where HIV is endemic also represent an increasing proportion of the positive HIV test reports in the last 3 years. These reports are included in the heterosexual exposure category.
- Women represent an increasing proportion of those with positive HIV test reports, as well as reported AIDS cases in Canada. Over 25% of the positive HIV test reports in 2004 were in women, compared to less than 10% prior to 1995. The largest rise in this group is seen among those aged 15–19 years. Heterosexual exposure and IDU are the two major risk behaviours for HIV infection in women.
- Aboriginal peoples make up a growing percentage of positive HIV test reports and reported AIDS cases. IDU continues to be a key mode of HIV transmission in the Aboriginal community. Nearly 50% of all positive HIV test reports among Aboriginal Canadians were in women (less than 20% of positive HIV test reports among caucasian Canadians were in women). Aboriginal peoples test positive for HIV at a younger age compared to non-Aboriginal persons.^{4,5}
- Canadians of African ancestry also make up a growing percentage of positive HIV reports and reported AIDS cases. Heterosexual exposure accounts for more than 80% of positive HIV test reports in this group. Approximately 50% of positive HIV test reports in this group are in women.
- Approximately 30% of people living with HIV infection are unaware of their HIV status. These persons representing the "hidden epidemic" are particularly important, because they have not yet taken advantage of services for clinical

- assessment, counselling and therapy. They present for medical attention later in the course of their illness and may unknowingly continue to transmit the infection.
- Although the limited data available suggest that HIV prevalence is currently low among Canadian youth, sexual risk behaviour and sexually transmitted infection (STI) data clearly indicate that the potential for HIV transmission remains significant among young Canadians. Data from targeted studies show that street-involved youth, youth who inject drugs and young MSM are particularly vulnerable to HIV infection.
- Rates of HIV infection in Canadian provincial and federal prisons appear to be
 much higher than in the general population. It is likely that most HIV-positive
 inmates were engaged in high-risk behaviour prior to imprisonment; however,
 there is evidence to indicate that some inmates continue to engage in highrisk behaviour after incarceration, including needle-sharing, tattooing and
 unprotected sex. There is great potential for HIV transmission among inmates,
 with possible transmission later to the spouses/partners of those released.⁶
- In Canada, blood donors have been screened and tested for HIV infection since 1985. This has resulted in a marked decline in the proportion of transfusionassociated HIV infections. The current estimated risk of infection from blood and blood products is exceedingly low in Canada (approximately one per million units of blood).
- The risk of acquiring HIV infection from a single sexual contact with an HIV-infected person is variable; risk increases with number of exposures and higher viral load in the source person.⁷⁻⁹ While oral sex is a lower-risk activity than unprotected anal or vaginal intercourse, repeated exposures may increase the risk.⁴
- Sexual transmission (infectiousness or susceptibility) of HIV is enhanced by the presence of other STIs,^{10–12} including ulcerative genital infections (e.g., syphilis, genital herpes) and non-ulcerative genital infections (e.g., chlamydia, gonorrhea, trichomoniasis).^{13–17} Bacterial vaginosis, although not strictly considered an STI, may also increase sexual transmission of HIV.^{18–21}
- The median time from acquiring HIV infection to the diagnosis of AIDS now exceeds 10 years. There has been a marked decline in the number of persons diagnosed with AIDS in Canada. The use of highly active antiretroviral therapy (HAART) is the major factor responsible for this decline.
- The use of HAART has dramatically changed the face of the HIV epidemic.²² The increased lifespan of persons living with this chronic disease may be leading to a more relaxed attitude and less caution in persons at risk of transmitting and acquiring this infection.²³⁻²⁵
- The success of HAART in transforming HIV infection into a chronic disease has increased the total burden of care. This has resulted in an increased incidence of adverse effects from therapy and greater difficulty with long-term adherence to HAART.
- Widespread use of HAART, including issues of non-adherence, has also increased the potential for transmission of drug-resistant virus.

Prevention

- Persons presenting with concerns about HIV infection provide an important
 opportunity for education and encouragement for the consistent practice of risk
 reduction. These practices include sexual abstinence, reduced number of sexual
 partners, proper use of barrier methods and risk reduction with IDU.
- Persons with known risk behaviour(s) should be offered HIV testing, counselling and diagnosis.
- At the time of diagnostic testing for HIV, review and monitor prevention practices.
- Identify barriers to prevention practices and the means to overcome them.
- Discuss the potential use of HAART to not only improve prognosis, but also reduce infectiousness.²⁶
- Discuss prompt treatment of any STI to reduce the risk of transmitting or acquiring HIV.²⁷⁻³¹

Pre- and Post-Test Counselling32

- Counselling should be age-appropriate and individualized to the person being tested.
- Testing should be done only after informed consent has been obtained.

Pre-test counselling

- · Clarify:
 - The confidentiality of HIV testing, reporting and record handling.
 - The testing options available (i.e., nominal, non-nominal, anonymous) (see Laboratory diagnosis, below).
 - That the test is for antibodies to HIV, not a direct test for the HIV virus or for AIDS.
 - That the majority of persons produce detectable antibodies within 3 months.
 - That an initial positive screening test is automatically followed by a confirmatory test (same blood sample) to rule out a false-positive test. This may mean a delay in the availability of the test result.
 - That the results should not be provided to the patient until confirmatory test results are available.
 - That the test results should be provided in person.
 - That returning for results is preferred, as it provides an opportunity to provide proper post-test counselling.
 - That a negative test may mean the person is not infected, or that it is too soon to detect antibodies.
 - That a positive test means the person is infected with HIV and is infectious to others through unprotected sexual contact, blood, breast milk or tissue/organ donation.

- That an indeterminate confirmatory test result means that testing should be repeated in 3 months or additional testing performed (e.g., qualitative HIV polymerase chain reaction [PCR], serum p24 antigen; please consult your local laboratory regarding test availability).
- That HIV is not casually transmitted through sweat, saliva, urine, feces or tears (unless there is visible blood present in any of these).
- That transmission risks are as follows:
 - Unprotected sexual contact: anal sex (high risk), vaginal sex (high risk), oral sex (low risk).
 - · Direct blood-to-blood contact.
 - Sharing needles or syringes (including IDU, tattooing, piercing with shared/unclean equipment).
 - Transmission from mother to child during pregnancy, at birth or via breast milk.
 - Receiving blood or blood products in Canada before November 1985 (elsewhere risk will vary depending on testing of donated blood).

· Discuss:

- Specific risk behaviours, sexual and otherwise.
- Availability of therapy to decrease the risk of mother-to-child transmission if the person is pregnant (decreased by ≥80%).
- Whether future testing will be necessary.
- Risk-reduction behaviours (see Primary Care and Sexually Transmitted Infections chapter):
 - Practice sexual abstinence (will eliminate risk).
 - Ensure consistent use of latex or polyurethane condoms.
 - · Avoid casual/anonymous/unprotected sex.
 - Avoid sharing needles, syringes or other IDU equipment.

· Explore:

- Psychological implications of testing.
- Coping mechanisms to deal with either result; availability of support systems (personal, community, medical).

Explain:

- The need to return for test results and schedule a post-test counselling visit.
- Public health notification for follow-up if the test is positive and the patient fails to return for results.
- Post-test counselling procedures.
- Partner notification and reporting requirements for HIV infection (depends on jurisdiction and availability of anonymous testing).
- With a positive result, the need for full clinical and laboratory assessments and for discussion regarding antiretroviral therapy and prophylaxis for opportunistic infections.

Post-test counselling33,34

- If the test result is negative:
 - Interpret as:
 - No infection or "window period" with infection, but no detectable antibodies. Retesting may be required 3 months after last potential exposure to allow for detection of an antibody response. Retesting 6 months after last potential exposure may be required for those presenting with late clinical signs and symptoms of HIV infection or in persons with an impaired immune response.
 - In the case of sexual assault (see Sexual Abuse in Peripubertal and Prepubertal Children and Sexual Assault in Postpubertal Adolescents and Adults chapters) and occupational exposure (see Occupational transmission, below) baseline testing should be performed, followed by additional testing at 6 weeks, 12 weeks and 6 months.
 - Reinforce risk reduction:
 - Avoid high-risk behaviours.
 - · Avoid needle/syringe sharing.
 - · Use lubricated latex or polyurethane condoms with sexual activity.
- If the test result is **positive**:
 - Interpret as:
 - Infected with HIV, not diagnostic of AIDS.
 - Explain that a confirmatory test to rule out a false-positive test has been performed.
 - Consider a first priority:
 - Dealing with the issues important to the infected person.
 - Discussing coping and support systems.
 - Discussing and assisting in the partner-notification process (by the infected person or the local public health unit).
 - Providing specific guidance about avoiding HIV transmission:
 - Protect others from sexual secretions, blood and other bodily fluids.
 - Avoid donating blood, organs, tissue, sperm or breast milk.
 - Be aware of infectivity (reinforce mechanisms of transmission, including high- and low-risk behaviours).
 - Discuss disclosure issues:
 - Persons with HIV infection should be informed of the medico-legal requirement to disclose their HIV status to a potential sexual or druginjecting partner. This is particularly important if they will be engaging in high-risk behaviour(s).³⁵⁻³⁷
 - Persons with HIV infection should inform their family physician and consider informing other health care providers (e.g., dentist).
 - Disclosure in the workplace is usually not mandatory but should be individualized (e.g., where the person with HIV infection has direct patient-care responsibilities).
 - Disclosure to friends or family is not essential but might be considered if there is potential for a positive outcome (e.g., positive family support).

- · Discuss benefits of treatment and follow-up.
- Deal with soon:
 - Further medical support, immune testing, HIV viral load testing, CD4 count and counselling are required.
 - Discuss use of laboratory testing to make therapeutic decisions.
- Discuss medical care:
 - Screen for hepatitis B virus (HBV) infection and immunity (see *Hepatitis B Virus Infections* chapter). Screen for hepatitis A virus (HAV) immunity in injection drug users, MSM, individuals with chronic liver disease and hemophiliacs.
 - · Screen for hepatitis C virus (HCV) infection.
 - Screen for syphilis and other STIs.
 - · Screen for tuberculosis.
 - Refer where required (e.g., HIV specialist).
 - Discuss health-enhancing lifestyle modifications, empowerment.
 - Discuss issues of confidentiality in the health care system, community, at school or at work.
 - Discuss avoidance of activities that increase transmission risk of toxoplasmosis and enteric pathogens.

Transmission

 Transmission of HIV infection occurs essentially through specific exposure to blood and/or body fluids from an HIV-infected person. The most concerning types of exposure include sexual exposure, parenteral blood exposure through IDU or blood transfusion, perinatal mother-to-child transmission and occupational exposure in the health care setting. Strategies for prevention should be aimed at risk reduction in these areas. A high viral load in the infected person increases the potential for transmission.³⁸

Sexual transmission

- This is the major route of HIV transmission.³⁹
- Sexual activities can be divided according to risk.⁴⁰ This ranges from touching
 and hugging, which carry no risk, to penile–anal and penile–vaginal intercourse
 without a condom, which carries a high risk. Providers must be aware of and
 counsel patients according to the implications that specific behaviours can have
 on the transmission of other blood-borne pathogens and STIs.
- Persons should be counselled that:
 - Only sexual abstinence and "no-risk" activities are guaranteed to prevent transmission.
 - Low-risk activities are preferable to high-risk activities.
 - Male and female condoms made of latex or polyurethane are an effective barrier for preventing HIV transmission. Correct and consistent use of condoms can reduce but not eliminate the risk of HIV transmission.⁴¹⁻⁴⁴

- The presence of another STI in either the source or the exposed person, particularly ulcerative lesions such as syphilis or genital herpes, increases the potential for sexual transmission of HIV.
- Infected individuals should be strongly urged to inform past, present and future sexual partners about their known HIV-positive status.
- Ongoing counselling and discussion about sexual behaviour is appropriate.

Parenteral transmission

- Risk of parenteral HIV transmission can be divided according to risk.⁴⁰ This
 ranges from the use of sterilized injection equipment, which is considered
 no-risk, to the use of shared needles, which is considered high-risk. Providers
 should be aware of and counsel patients according to the implications that
 specific behaviours can have on the transmission of other blood-borne
 pathogens.
- Active injection drug users should be encouraged to discontinue drug use by using addiction-treatment services and should be counselled on the health risks associated with IDU.
- If the individual is not ready, willing or able to discontinue IDU, harm-reduction strategies should be stressed, including not sharing injecting equipment and adopting safer modes of drug use.
- Access to sterile injecting equipment, such as needle-exchange programs, should be discussed and encouraged.

Perinatal mother-to-child transmission

- The HIV prevalence rate among pregnant women is approximately 3–5/10,000 in Canada.
- Transmission of HIV infection from the HIV-positive mother to her infant may occur in utero, during childbirth or after childbirth through breastfeeding.
 Preventing this mode of transmission can, therefore, be achieved by identifying HIV-infected women who are pregnant and using strategies to minimize the risk of mother-to-child transmission.⁴⁵
- Antiretroviral therapy can dramatically reduce perinatal transmission of HIV.
- In all Canadian provinces and territories, HIV testing of pregnant women remains
 the choice of the woman. Guidelines and/or recommendations for HIV testing
 of pregnant women have been developed in each province and territory to
 encourage informed decision-making.
- All pregnant women should be offered confidential HIV testing and counselling as part of routine prenatal care.
- In some provinces and territories (Alberta, Newfoundland and Labrador, Northwest Territories, Nunavut), an "opt-out" policy treats HIV screening as a routine prenatal screening test. The pregnant woman is informed that testing will be done, but consent is implied unless she specifically refuses.⁴

- Women who present in labour who have not had prenatal HIV testing or who
 have been engaging in high-risk behaviour after initial negative prenatal HIV
 testing should be offered expedited or rapid HIV testing.⁴⁵
- HIV-positive women of childbearing age should be counselled about the risk of
 mother-to-child transmission. They should also be given complete information
 regarding contraceptive and reproductive options, as well as the availability
 of therapy to decrease the risk of transmission to the child (see *Pregnancy*chapter).
- In North America, breastfeeding is contraindicated for HIV-infected mothers.

Occupational transmission46

- Transmission of HIV infection in the workplace (occupational exposure) is
 primarily concerned with the potential for transmission from patient to health
 care personnel. The potential for transmission from health care personnel to
 patient and from one health care person to another is not within the scope of
 this section.
- Occupational exposure to HIV infection may occur in several instances:
 - Percutaneous injury with a sharp object potentially contaminated with blood or other bodily fluid.
 - Mucous membrane exposure to blood or other bodily fluid.
 - Skin exposure to blood or other bodily fluid.
- The average risk of HIV transmission after a percutaneous exposure to HIV-infected blood has been estimated to be approximately 0.3% (3/1,000), and after a mucous membrane exposure, approximately 0.09% (0.9/1,000).^{47,48} Although episodes of HIV transmission after non-intact skin exposure have been documented, the average risk for transmission by this route has not been precisely quantified but is estimated to be less than the risk for mucous membrane exposures.^{49,50} The risk for transmission after exposure to fluids or tissues other than HIV-infected blood also has not been quantified, but is probably considerably lower than for blood exposures.⁵¹
- The decision to initiate postexposure prophylaxis (PEP) for HIV infection is based on clinical judgment and should be a joint decision with the exposed health care worker.
- The choice of no PEP vs. a two- or three-drug PEP regimen is based on the index of suspicion after evaluating the following:
 - Source of exposure: the potential for HIV infection (e.g., high-risk activity or HIV-positive source).
 - Type of exposure: the potential for transmission of HIV infection (e.g., hollowbore needle visibly contaminated with source patient's blood).^{52,53}
- PEP should be initiated as soon as possible, as it may be less effective if initiated more than 72 hours after exposure.

Diagnosis

The diagnosis of HIV infection is based primarily on a positive serologic test.
Persons with HIV infection may be totally asymptomatic. Therefore, serologic
testing is recommended when there is a high index of suspicion (e.g., high-risk
behaviour and/or suspicious clinical symptoms and signs). Persons may also
present with specific opportunistic infections or other conditions indicative of
underlying immunosuppression.

Risk behaviours

- Multiple sex partners.
- Unprotected sexual activity (i.e., no barrier [condom] protection).
- Sex with an HIV-infected partner.
- · Receptive anal/vaginal intercourse.
- Sharing of IDU equipment.
- Acquisition of other STIs, such as HBV or syphilis.

Clinical diagnosis

- The time from initial HIV infection to clinical disease is highly variable, with a median time of approximately 10 years. However some HIV-infected persons experience more rapid progression of disease.
- The person with HIV infection may experience several stages:
 - Primary or acute HIV infection.
 - Chronic asymptomatic HIV infection.
 - Chronic symptomatic HIV infection.

Primary/acute HIV infection

- This is the period from initial infection to development of the full serum antibody profile (i.e., seroconversion).⁵⁴⁻⁵⁶
 - High levels of viral replication and plasma viremia.
 - Shedding from mucosal sites.
 - No detectable antibody.
 - Depressed CD4 count.
- Although some patients in this stage are asymptomatic, up to 90% may be symptomatic (i.e., the acute retroviral syndrome).⁵⁷ Symptoms generally appear 2–4 weeks after initial infection and are often nonspecific or mild. They are usually self-limited, lasting 1–2 weeks, but may last several months.
- The spectrum of symptoms may include an acute mononucleosis-like illness, fever and skin rash. Meningoencephalitis or aseptic meningitis may occur. Less commonly, AIDS-defining conditions such as *Pneumocystis jirovecii* (formerly carinii) pneumonia or oroesophageal candidiasis may occur.

Table 1. Symptoms of acute HIV infection

Symptoms	Frequency
Fever (mean temperature 39.4°C [102.9°F])	>80%
Arthralgia or myalgia, rash, lymphadenopathy, sore throat, fatigue, headache	40-80%
Oral ulcers and/or genital ulcers, >5 kg weight loss, nausea, vomiting or diarrhea	10-40%

- If initial HIV serologic tests are negative or indeterminate, additional testing can be considered. Please consult appropriate resources or colleagues experienced in this area.
- A high index of suspicion in the person with a nonspecific febrile illness and a history of high-risk behaviour is key to making the diagnosis.
- Although the treatment of primary or acute HIV infection is considered optional
 at this time, these persons may be highly infectious.⁵⁸ Detection of primary
 HIV infection provides an opportunity for counselling and preventing further
 transmission.

Chronic asymptomatic HIV infection

- This is the stage where viral replication and plasma viremia are more controlled by the immune response. There is a balance between ongoing viral replication and the host immune response represented by the level of CD4+ T cells.
 - Many persons fall into this category.
 - Generalized lymphadenopathy is frequently present.
 - Thrombocytopenia may be present.

Chronic symptomatic HIV infection

 This is the stage where viral replication depletes the CD4+ T cells to the level of profound immunosuppression.⁵⁹ See Table 2 for signs and symptoms.

Table 2. Signs and symptoms of chronic symptomatic HIV infection

- Oral hairy leukoplakia
- Unexplained fever (>2 weeks)
- · Fatigue or lethargy
- Unexplained weight loss (>10% body weight)
- Chronic diarrhea (>3 weeks)
- Unexplained lymphadenopathy (usually generalized)
- · Cervical dysplasia
- · Dyspnea and dry cough
- · Loss of vision
- Recurrent or chronic mucocutaneous candidiasis (oral, esophageal, vaginal)
- Dysphagia (esophageal candidiasis)
- Red/purple nodular skin or mucosal lesions (Kaposi sarcoma)
- · Encephalopathy
- · Herpes zoster, especially if severe, multidermatomal or disseminated
- · Increased frequency or severity of mucocutaneous herpes simplex virus infection
- Unexplained "anemia of chronic disease"

Table 3. AIDS-defining conditions^{60,61}

(Require concurrent positive HIV serology to be diagnostic of AIDS)

- · Bacterial pneumonia, recurrent
- · Candidiasis (esophageal, bronchi, trachea or lungs)
- · Cervical cancer, invasive
- Coccidioidomycosis (disseminated or extrapulmonary)
- Cryptococcosis (extrapulmonary)
- Cryptosporidiosis (chronic intestinal)
- Cytomegalovirus disease (other than liver, spleen, nodes)
- · Cytomegalovirus retinitis (with loss of vision)
- Encephalopathy, HIV-related (dementia)
- Herpes simplex virus (chronic ulcers or bronchitis, pneumonitis or esophagitis)
- · Isosporiasis, chronic intestinal
- · Kaposi sarcoma
- Lymphoma (Burkitt, immunoblastic, primary in brain)
- Mycobacterium avium complex or M kansasii (disseminated or extrapulmonary)
- Mycobacterium of other species (disseminated or extrapulmonary)
- Mycobacterium tuberculosis (pulmonary, disseminated or extrapulmonary)
- · Pneumocystis jirovecii (formerly carinii) pneumonia
- Progressive multifocal leukoencephalopathy
- · Salmonella septicemia, recurrent
- · Toxoplasmosis of brain
- · Wasting syndrome due to HIV

Laboratory diagnosis — HIV antibody testing

- Any physician or qualified health care provider may order an HIV test.
 (Check with your local laboratory for the availability of these tests.)
- Testing should be carried out only with the informed consent of the person being tested.
- HIV antibody testing should be offered to any person with identified risk behaviour who has clinical or laboratory clues suggestive of HIV infection, or who requests it.
- Explain clearly the nature of the test, and provide appropriate pre- and post-test counselling.
- Point-of-care rapid tests for HIV antibodies are now more widely available.
 All reactive screening tests using these kits require confirmatory testing (e.g., Western blot analysis).⁶²
- CD4 count and viral load testing should not be used as screening or diagnostic tests.
- p24 antigen testing, although occasionally useful in diagnosis of primary or acute infection, is insensitive for screening purposes.
- There are three options for HIV testing and reporting in Canada: nominal, nonnominal or anonymous. The use and availability of these options varies across the provinces and territories. Your local public health authority will provide information on the options available in your region.⁴
 - Nominal testing: the HIV test is ordered using the name of the person being tested.
 - Non-nominal testing: the HIV test is ordered using a code or the initials of the person being tested. Only the person ordering the test knows the identity of the person being tested and is able to link the result to that person's health care record.
 - Anonymous testing: the HIV test is ordered using a unique non-identifying code. The person(s) ordering the test and providing the result (usually by telephone) do not know the identity of the person being tested. Only the person being tested knows the code, so the test result is not linked to that person's health care record. Although anonymous testing may encourage more persons to have testing, it is not available in all provinces and territories.
- A positive enzyme immunoassay (EIA) screening test requires confirmatory testing (e.g., Western blot analysis) using the same specimen.
- Repeat all initial positive HIV serologic tests using a second blood specimen to rule out laboratory error and confirm the diagnosis.

Management, Treatment and Follow-up^{63,64}

 This is an increasingly complex area, with rapid changes in optimal therapy as new research becomes available. Recommendations for a given person should be made in collaboration with a colleague experienced in HIV/AIDS. Your local public health authority will have a listing of these.

Guiding principles

- Asymptomatic infected persons are usually followed at 3–6-month intervals
 if untreated.
- The follow-up interval may vary if antiretroviral therapy is provided or if the person is symptomatic.
- Routine monitoring of CD4 lymphocyte count and plasma HIV RNA viral load are key in assessing effectiveness of antiretroviral therapy.^{65,66}

First visit after positive HIV test

- Conduct a medical history and complete physical examination, including genital and anal inspection.
- Order laboratory tests, including complete blood count with white cell
 differential, CD4 count, viral load, liver function tests, creatinine kinase,
 blood glucose, amylase and lipase. Screen for HBV infection and immunity
 (see Hepatitis B Virus Infections chapter). Screen for HAV immunity in injection
 drug users, MSM and individuals with chronic liver disease and hemophilia.
 Screen for HCV infection. Screen for toxoplasma (IgG) and syphilis. Tests
 for other STIs, such as gonorrhea and chlamydia, should also be considered
 (see Consideration for Other STIs, below).
- For women, cervical screening for dysplasia and/or human papillomavirus (HPV) infection is recommended if not performed within the last 6–12 months. The anal Pap smear for men with a history of anal receptive intercourse and/or a history of anal warts is available only in certain centres.
- Baseline fasting lipids and glucose would be appropriate if considering starting antiretroviral therapy.
- Tuberculin skin testing is essential. A negative test may not rule out latent or active tuberculosis.⁶⁷
 - If past exposure to Mycobacterium tuberculosis is indicated (induration
 ≥5 mm in diameter), the person should be assessed for active tuberculosis.
 - If active tuberculosis is excluded and the person has not previously received therapy to prevent or treat tuberculosis, isoniazid 300 mg once daily for 9–12 months is highly effective in preventing the development of active tuberculosis. Rifampin 600 mg daily or rifabutin 300 mg daily can be used for isoniazid-resistant strains or when isoniazid toxicity precludes isoniazid use.⁶⁸
 - Consultation with a colleague experienced in this area should be sought.

- Immunization (e.g., HAV, HBV) should be discussed according to current guidelines.^{69,70} Generally, there is no contraindication to the use of inactivated or component vaccines in HIV-positive persons. The routine childhood immunization schedule should be completed if indicated. Pneumococcal immunization (boosted once only after 5 years) and annual influenza immunization are recommended.
- Influenza and pneumococcal immunization have been associated with transient increases in plasma viral load levels. However, this does not appear to have any significant impact on disease progression, and the benefits are generally felt to outweigh the risks.
- Smoking cessation is an important issue, particularly in persons with other cardiovascular risk factors who will be starting antiretroviral therapy.

Follow-up visits

- Conduct a clinical assessment, including assessment for cardiovascular disease, lipodystrophy, lactic acidosis and diabetes mellitus.
- Conducting an annual anal inspection for the presence of HPV lesions, particularly in MSM, is encouraged.^{71,72}
- Take the opportunity for risk-reduction counselling. Sexual and drug-use history should be discussed at each visit.
- If the patient is on therapy, assess adverse effects and adherence.
- CD4 counts and viral load testing should be performed every 3–6 months. Other laboratory tests, including complete blood count with white cell differential, liver function tests, creatinine kinase, amylase, lipase, fasting lipids and blood glucose should also be performed every 3–6 months, depending on drug therapy.
- There are two components to drug treatment: antiretroviral therapy and drugs to prevent or treat opportunistic infections.

Antiretroviral therapy73

- This is a rapidly evolving field, and any decision on specific therapy for a given
 person should be made in collaboration with a colleague experienced in
 HIV/AIDS. Therapy should be individualized and based on factors such as
 efficacy, tolerability, potential adverse effects, convenience and drug-drug
 interactions. Specific details and recommendations for antiretroviral drug
 therapy are beyond the scope of this chapter.
- The antiretroviral drug classes licensed in Canada so far include the following:
 - Nucleoside reverse transcriptase inhibitors (NRTIs): e.g., zidovudine (AZT), lamivudine (3TC) and stavudine (d4T).
 - Nucleotide reverse transcriptase inhibitor (NtRTI): tenofovir.
 - Non-nucleoside reverse transcriptase inhibitors (NNRTIs): e.g., efavirenz and nevirapine.
 - Protease inhibitors (PIs): e.g., nelfinavir, saguinavir, ritonavir and atazanavir.
 - Fusion inhibitor: enfuvirtide/T20.

- Other types of investigational antiretroviral drugs are presently in development and clinical trials. Immune-based therapy to boost CD4 counts is still in clinical trials.
- Recommendations for antiretroviral therapy are based on clinical status, CD4
 count, viral load and patient willingness to undertake therapy (see Table 4). It
 must be recognized that prolonged therapy is limited by drug toxicity, adherence
 issues, drug resistance and cost.
- Therapy, when indicated, includes at least three agents (e.g., two NRTIs plus one NNRTI or PI).
- The goal of therapy is to suppress viral replication to the point where plasma HIV RNA is undetectable, with minimal patient toxicity.
- Monotherapy and dual therapy must be avoided, as this has been associated with the emergence of drug resistance.
- Persons must be instructed to take medication regularly, as missed doses and under-dosing may promote drug resistance.
- · Significant drug-drug interactions may occur with some antiretroviral drugs.
- Alteration of HAART is usually indicated if there is a failure to achieve or maintain control of viral replication or there is unacceptable toxicity. Resistance testing (genotyping or phenotyping) may be valuable in the selection of the initial or subsequent regimens.

Table 4. Guidelines for starting antiretroviral therapy for the person with chronic HIV infection

Clinical status	CD4 count	Viral load	Therapy
AIDS-defining illness or severe HIV symptoms	Any	Any	Yes
Asymptomatic	<0.2 x 10°/L (<200/μL)	Any	Yes
Asymptomatic	0.2-0.35 x 10°/L (200-350/μL)	Any	Offer
Asymptomatic	>0.35 x 10 ⁹ /L (>350/ μ L)	≥100,000 copies/mL	Defer or consider
Asymptomatic	>0.35 x 10°/L (>350/µL)	<100,000 copies/mL	Defer

Prevention of opportunistic infections74

- HIV-infected persons are at increased risk of specific opportunistic infections, depending on their CD4 count.
- It is safe to discontinue prophylactic therapy once CD4 count has increased and remained above a certain level for 3–6 months.

Table 5. Prophylactic therapy for opportunistic infections

CD4 count	Opportunistic infection	Prophylactic therapy
<0.2 x 10 ⁹ /L (<200 cells/ <i>μ</i> L)	Pneumocystis jiroveci (formerly carinii) pneumonia	 Preferred: trimethoprim-sulfamethoxazole PO once daily or three times per week Alternate: dapsone PO once daily, atovaquone PO once daily, aerosolized pentamidine once monthly Also indicated with oral candidiasis or prior <i>P jiroveci</i>, regardless of CD4 count
<0.1 x 10 ⁹ /L (<100 cells/ <i>µ</i> L)	Toxoplasma gondi	Same drugs as <i>P jiroveci</i> , except for aerosolized pentamidine
<0.05 x 10°/L (<50 cells/μL)	Mycobacterium avium complex	 Preferred: azithromycin PO once weekly Alternate: clarithromycin PO twice daily, rifabutin PO once daily

- Cytomegalovirus disease:
 - Present guidelines do not recommend primary prophylaxis for cytomegalovirus (CMV) disease. However, persons with CD4 count <0.05 x 10°/L (<50 cells/μL) are at highest risk of CMV disease. These persons should be aware of the symptoms of CMV disease, in particular CMV retinitis (e.g., visual distortions, floaters). A regular 4–6-monthly funduscopic examination performed by an ophthalmologist may be helpful in early detection of CMV retinitis.
- · Other infections:
 - Treatment and prevention of bacterial, viral, parasitic and fungal infections must be individualized and response to therapy monitored.
 - In many instances, long-term suppressive therapy is required.

Consideration for Other STIs

- Persons with risk behaviours for HIV infection should be offered testing for other STIs.
 - Testing from appropriate sites for chlamydia and gonorrhea.
 - Serologic tests for syphilis.
 - Screening for HBV infection and immunity (see Hepatitis B Virus Infections chapter); screening for HAV immunity in injection drug users, MSM, individuals with chronic liver disease and hemophilia; screening for HCV infection.
 - Type-specific herpes simplex virus (HSV) serology (HSV-2 infection): if available this may be useful in identifying persons who are potentially more at risk of acquiring or transmitting HIV infection. The increased risk of acquisition or transmission appears to be primarily during symptomatic genital HSV (active genital ulcers).⁷⁵⁻⁷⁹ However, asymptomatic genital HSV may also be an important factor in HIV acquisition or transmission. Episodes of acute genital HSV have been shown to increase mucosal shedding and plasma levels of HIV.⁸⁰⁻⁸³ Antiviral therapy and suppression of genital HSV reactivation may be an important strategy in minimizing HIV transmission in association with genital HSV infection.^{84,85} If genital ulcers are present, see the *Genital Ulcer Disease* chapter for testing recommendations.
- Offer immunization for HBV and HAV if non-immune as per current guidelines.⁶⁹

Reporting and Partner Notification

- HIV infection is reportable in all provinces and territories; such reporting may be nominal or non-nominal, depending on the jurisdiction.
- AIDS is reportable by physicians to local public health authorities in all provinces and territories.
- Partner notification must be undertaken in all cases of AIDS and HIV infection.
- Local public health authorities are available to assist with partner notification and help with appropriate referral for clinical evaluation, testing, treatment and health education. The treating physician is responsible for ensuring that partner notification is initiated.
- All children born to mothers who are or may be HIV-infected must be evaluated (see *Pregnancy* chapter).
- All HIV-positive persons who have previously received or donated blood should be reported in confidence to the local Canadian Blood Services.

Special Considerations

- The increased risk of cervical cancer in HIV-infected women is related to the degree of immunosuppression.⁸⁶ Pap smears should be performed at baseline and 6 months later, with subsequent Pap smears at least annually depending on the results of initial smears.^{74,87}
- Anal HPV infection with subsequent epithelial changes of anal cancer and its
 precancerous lesions have been detected in HIV-infected persons, even in the
 absence of anal intercourse. These changes may be seen despite the use of
 HAART and immune restoration.^{71,72}
- In some centres, anal Pap smears and HPV detection are performed on a regular basis in HIV-positive MSM. Colposcopy and biopsy is performed if indicated.
 Aggressive therapy of high-grade lesions is warranted.
- It is important to ensure access to psychological counselling for all HIV-infected persons as needed.
- It may be appropriate to provide non-occupational PEP to persons in certain situations (e.g., sexual assault) on a case-by-case basis.⁸⁸
- Some persons may develop acute symptoms, such as fever, arthralgia, myalgia, lymphadenopathy, worsening liver disease or encephalopathy within the first few weeks of starting HAART. This "immune reconstitution syndrome" is associated with the improved immune response to pre-existing co-infection (e.g., with HCV or mycobacterium avium complex [MAC]).
- All persons on HAART have the potential to develop a number of adverse
 effects. These include direct drug-related toxicity (e.g., pancreatitis, peripheral
 neuropathy, body-fat maldistribution [lipodystrophy] or metabolic abnormalities
 such as hyperglycemia or hyperlipidemia). Lactic acidosis and liver dysfunction
 may be more frequent with specific drugs.
- Many persons are also at increased risk of cardiovascular disease related to family history, risk factors such as smoking and drug-induced hyperlipidemia.
- Other issues, such as osteopenia, osteoporosis and hypogonadism, may also become problematic.
- Persons with HIV co-infection may experience a more rapid progression of HCV infection and HBV infection. HBV or HCV co-infection is a risk factor for severe hepatotoxicity during HAART.⁸⁹⁻⁹³
- HIV co-infection may alter the natural history of syphilis and neurosyphilis, including response to therapy.⁹⁴⁻⁹⁷
- Therapeutic drug monitoring is being used to assess therapeutic drug levels in some persons who are adherent but fail an appropriate regimen. This is not yet universally available.
- Discussion of sexual and other risk behaviours should be routinely performed at each visit. The medico-legal implications of infection transmission without disclosure should be reinforced. Referral to local public health authorities should be made in cases where risk behaviours are not being voluntarily controlled.

References

- The Relationship Between the Human Immunodeficiency Virus and the Acquired Immunodeficiency Syndrome. Bethesda, MD: National Institute of Allergy and Infectious Diseases; 1995. Available at: www.niaid.nih.gov/ Publications/hivaids/hivaids.htm. Accessed January 19, 2006.
- The Evidence that HIV Causes AIDS. Bethesda, MD: National Institute of Allergy and Infectious Diseases; 2003. Available at: www.niaid.nih.gov/ Factsheets/evidhiv.htm. Accessed January 19, 2006.
- HIV and AIDS in Canada: Surveillance Report to December 31, 2004. Ottawa, ON: Surveillance and Risk Assessment Division, Centre for Infectious Disease Prevention and Control, Public Health Agency of Canada; 2005. Available at: www.phac-aspc.gc.ca/publicat/aids-sida/haic-vsac1204/index.html. Accessed January 19, 2006.
- HIV/AIDS Epi Updates 2005. Ottawa, ON: Surveillance and Risk Assessment Division, Centre for Infectious Disease Prevention and Control, Public Health Agency of Canada; 2005. Available at: www.phac-aspc.gc.ca/ publicat/epiu-aepi/epi-05/index.html. Accessed January 19, 2006.
- Understanding the HIV/AIDS Epidemic among Aboriginal Peoples in Canada: The Community at a Glance. HIV/AIDS Epi Notes. December 2004. Ottawa, ON: Surveillance and Risk Assessment Division, Centre for Infectious Disease Prevention and Control, Public Health Agency of Canada; 2004. Available at: www.phac-aspc.gc.ca/publicat/epiu-aepi/epi-note/index.html. Accessed January 19, 2006.
- 6. HIV/AIDS and Hepatitis C in Prisons: The Facts. Montreal, QC: Canadian HIV/ AIDS Legal Network; 2004. Available at: www.aidslaw.ca/Maincontent/issues/ prisons/e-revinfo-pa1.pdf. Accessed January 19, 2006.
- Cameron DW, Simonsen JN, D'Costa LJ, et al. Female to male transmission of human immunodeficiency virus type 1: risk factors for seroconversion in men. *Lancet* 1989;2:403–407.
- 8. Quinn TC, Wawer MJ, Sewankambo N, et al. Viral load and heterosexual transmission of human immunodeficiency virus type 1. Rakai Project Study Group. *N Engl J Med* 2000;342:921–929.
- 9. Wawer MJ, Gray RH, Sewankambo NK, et al. Rates of HIV-1 transmission per coital act, by stage of HIV-1 infection, in Rakai, Uganda. *J Infect Dis* 2005;191:1403–1409.
- Rottingen JA, Cameron DW, Garnett GP. A systematic review of the epidemiologic interactions between classic sexually transmitted diseases and HIV: how much really is known? Sex Transm Dis 2001;28:579–597.
- 11. Fleming DT, Wasserheit JN. From epidemiological synergy to public health policy and practice: the contribution of other sexually transmitted diseases to sexual transmission of HIV infection. *Sex Transm Infect* 1999;75:3–17.
- 12. Galvin SR, Cohen MS. The role of sexually transmitted diseases in HIV transmission. *Nat Rev Microbiol* 2004;2:33–42.
- 13. Mabey D. Interactions between HIV infection and other sexually transmitted diseases. *Trop Med Int Health* 2000;5:A32–36.

- 14. Stamm WE, Handsfield HH, Rompalo AM, Ashley RL, Roberts PL, Corey L. The association between genital ulcer disease and acquisition of HIV infection in homosexual men. *JAMA* 1988;260:1429–1433.
- 15. Hook EW 3rd, Cannon RO, Nahmias AJ, et al. Herpes simplex virus infection as a risk factor for human immunodeficiency virus infection in heterosexuals. *J Infect Dis* 1992;165:251–255.
- Guenthner PC, Secor WE, Dezzutti CS. Trichomonas vaginalis-induced epithelial monolayer disruption and human immunodeficiency virus type 1 (HIV-1) replication: implications for the sexual transmission of HIV-1. *Infect Immun* 2005;73:4155–4160.
- 17. Kreiss J, Willerford DM, Hensel M, et al. Association between cervical inflammation and cervical shedding of human immunodeficiency virus DNA. *J Infect Dis* 1994:170:1597–1601.
- 18. Sewankambo N, Gray RH, Wawer MJ, et al. HIV-1 infection associated with abnormal vaginal flora morphology and bacterial vaginosis. *Lancet* 1997;350:546–550.
- 19. Martin HL, Richardson BA, Nyange PM, et al. Vaginal lactobacilli, microbial flora, and risk of human immunodeficiency virus type 1 and sexually transmitted disease acquisition. *J Infect Dis* 1999;180:1863–1868.
- Cu-Uvin S, Hogan JW, Caliendo AM, et al. Association between bacterial vaginosis and expression of human immunodeficiency virus type 1 RNA in the female genital tract. *Clin Infect Dis* 2001;33:894–896.
- Sha BE, Zariffard MR, Wang QJ, et al. Female genital-tract HIV load correlates inversely with Lactobacillus species but positively with bacterial vaginosis and Mycoplasma hominis. *J Infect Dis* 2005;191:25–32.
- 22. Egger M, May M, Chene G, et al. Prognosis of HIV-1-infected persons starting highly active antiretroviral therapy: a collaborative analysis of prospective studies. *Lancet* 2002;360:119–129.
- 23. Katz MH, Schwarcz SK, Kellogg TA, et al. Impact of highly active antiretroviral treatment on HIV seroincidence among men who have sex with men in San Francisco. *Am J Public Health* 2002;92:388–394.
- Chen SY, Gibson S, Katz MH, et al. Continuing increases in sexual risk behavior and sexually transmitted diseases among men who have sex with men: San Francisco, California, 1999–2001, USA. Am J Public Health 2002;92:1387–1388.
- 25. Wolitski RJ, Valdiserri RO, Denning PH, Levine WC. Are we headed for a resurgence of the HIV epidemic among men who have sex with men? *Am J Public Health* 2001;91:883–888.
- Vernazza PL, Troiani L, Flepp MJ, et al. Potent antiretroviral treatment of HIVinfection results in suppression of the seminal shedding of HIV. The Swiss HIV Cohort Study. AIDS 2000;14:117–121.
- Cohen MS, Hoffman IF, Royce RA, et al. Reduction of concentration of HIV-1 in semen after treatment of urethritis: implications for prevention of sexual transmission of HIV-1. AIDSCAP Malawi Research Group. *Lancet* 1997;349:1868–1873.

- Centers for Disease Control and Prevention. HIV prevention through early detection and treatment of other sexually transmitted diseases — United States. Recommendations of the Advisory Committee for HIV and STD Prevention. MMWR Recomm Rep 1998;47(RR-12):1–24.
- 29. Wang CC, McClelland RS, Reilly M, et al. The effect of treatment of vaginal infections on shedding of human immunodeficiency virus type 1. *J Infect Dis* 2001;183:1017–1022.
- 30. McClelland RS, Wang CC, Mandaliya K, et al. Treatment of cervicitis is associated with decreased cervical shedding of HIV-1. *AIDS* 2001;15:105–110.
- 31. Harwell JI, Flanigan TP, Mitty JA, et al. Directly observed antiretroviral therapy to reduce genital tract and plasma HIV-1 RNA in women with poor adherence. *AIDS* 2003;17:1990–1993.
- 32. Centers for Disease Control and Prevention. Revised guidelines for HIV counseling, testing, and referral. *MMWR Recomm Rep* 2001;50(RR-19):1–57.
- 33. Horsburgh CR Jr, Ou CY, Jason J, et al. Duration of human immunodeficiency virus infection before detection of antibody. *Lancet* 1989;2:637–640.
- Busch MP, Lee LL, Satten GA, et al. Time course of detection viral and serologic markers preceding human immunodeficiency virus type 1 seroconversion: implications for screening of blood and tissue donors. *Transfusion* 1995;35:91–97.
- 35. After Cuerrier: Canadian Criminal Law and the Non-Disclosure of HIV-Positive Status. Montreal, QC: Canadian HIV/AIDS Legal Network; 1999. Available at: www.aidslaw.ca/Maincontent/issues/criminallaw/finalreports/cuerrier/tofc.htm. Accessed January 19, 2006.
- Elliott R, Betteridge G. HIV-positive person who did not disclose status convicted of attempted aggravated assault. Can HIV Aids Policy Law Rev 2003;8:50–53
- 37. Persons who fail to disclose their HIV/AIDS status: conclusions reached by an expert working group. *Can Commun Dis Rep* 2005;31:53–61.
- 38. Lee TH, Sakahara N, Fiebig E, Busch MP, O'Brien TR, Herman SA. Correlation of HIV-1 RNA levels in plasma and heterosexual transmission of HIV-1 from infected transfusion recipients. *J Acquir Immune Defic Syndr Hum Retrovirol* 1996;12:427–428.
- Royce RA, Sena A, Cates W Jr, Cohen MS. Sexual transmission of HIV. N Engl J Med 1997:336:1072–1078.
- HIV Transmission: Guidelines for Assessing Risk. A Resource for Educators, Counsellors, and Health Care Providers. 5th ed. Ottawa, ON: Canadian AIDS Society; 2004.
- Centers for Disease Control and Prevention (CDC). Update: barrier protection against HIV infection and other sexually transmitted diseases. MMWR Morb Mortal Wkly Rep 1993:42:589–591, 597.
- 42. Fact Sheet for Public Health Personnel 2002: Male Latex Condoms and Sexually Transmitted Diseases. Atlanta, GA: Centers for Disease Control and Prevention; 2002. Available at: www.cdcnpin.org/FactSheets/condom.pdf. Accessed January 19, 2006.

- 43. Carey RF, Herman WA, Retta SM, Rinaldi JE, Herman BA, Athey TW. Effectiveness of latex condoms as a barrier to human immunodeficiency virus-sized particles under conditions of simulated use. *Sex Transm Dis* 1992;19:230–234.
- 44. Minnis AM, Padian NS. Effectiveness of female controlled barrier methods in preventing sexually transmitted infections and HIV: current evidence and future research directions. *Sex Transm Inf* 2005;81:193–200.
- 45. Centers for Disease Control and Prevention (CDC). Revised recommendations for HIV screening of pregnant women. *MMWR Recomm Rep* 2001;50(RR-19): 63–85
- Panlilio AL, Cardo DM, Grohskopf LA, Heneine W, Ross CS; US Public Health Service. Updated U.S. Public Health Service guidelines for the management of occupational exposures to HIV and recommendations for postexposure prophylaxis. MMWR Recomm Rep 2005;54(RR-9):1–17.
- 47. Bell DM. Occupational risk of human immunodeficiency virus infection in health care workers: an overview. *Am J Med* 1997;102(5B):9–15.
- 48. Ippolito G, Puro V, De Carli G. The risk of occupational human immunodeficiency virus in health care workers. Italian Multicenter Study. The Italian Study Group on Occupational Risk of HIV Infection. *Arch Int Med* 1993;153:1451–1458.
- 49. Centers for Disease Control (CDC). Update: human immunodeficiency virus infections in health-care workers exposed to blood of infected patients. *MMWR Morb Mortal Wkly Rep* 1987;36:285–289.
- Fahey BJ, Koziol DE, Banks SM, Henderson DK. Frequency of nonparenteral occupational exposures to blood and body fluids before and after universal precautions training. Am J Med 1991;90:145–153.
- 51. Henderson DK, Fahey BJ, Willy M, et al. Risk for occupational transmission of human immunodeficiency virus type 1 (HIV-1) associated with clinical exposures. A prospective evaluation. *Ann Intern Med* 1990;113:740–746.
- 52. Cardo DM, Culver DH, Ciesielski CA, et al. A case-control study of HIV seroconversion in health care workers after percutaneous exposure. Centers for Disease Control and Prevention Needlestick Surveillance Group. *N Engl J Med* 1997;337:1485–1490.
- 53. Mast ST, Woolwine JD, Gerberding JL. Efficacy of gloves in reducing blood volumes transferred during simulated needlestick injury. *J Infect Dis* 1993;168:1589–1592.
- 54. Tindall B, Barker S, Donovan B, et al. Characterization of the acute clinical illness associated with human immunodeficiency virus infection. *Arch Intern Med* 1988;148:945–949.
- 55. Schacker T, Collier AC, Hughes J, Shea T, Corey L. Clinical and epidemiologic features of primary HIV infection. *Ann Intern Med* 1996;125:257–264.
- 56. Quinn TC. Acute primary HIV infection. JAMA 1997;278:58-62.
- 57. Kahn JO, Walker BD. Acute human immunodeficiency virus type 1 infection. *N Engl J Med* 1998;339:33–39.

- 58. Pilcher CD, Tien HC, Eron JJ Jr, et al; Quest Study; Duke-UNC-Emory Acute HIV Consortium. Brief but efficient: acute HIV infection and the sexual transmission of HIV. *J Infect Dis* 2004;189:1785–1792.
- 59. Moylett EH, Shearer WT. HIV: clinical manifestations. *J Allergy Clin Immunol* 2002;110:3–16.
- Health and Welfare Canada. Revision of the CDC surveillance case definition for acquired immunodeficiency syndrome. CDWR 1987;13-38:169–177.
- 61. Revision of the surveillance case definition for AIDS in Canada. *CCDR* 1993:19-15:116–117.
- 62. Branson BM. Point-of-care rapid tests for HIV antibody. *J Lab Med* 2003;27:288–295.
- 63. Aberg JA, Gallant JE, Anderson J, et al; *HIV Med*icine Association of the Infectious Diseases Society of America. Primary care guidelines for the management of persons infected with human immunodeficiency virus: recommendations of the *HIV Med*icine Association of the Infectious Diseases Society of America. *Clin Infect Dis* 2004;39:609–629.
- Hecht FM, Wilson IB, Wu AW, Cook RL, Turner BJ. Optimizing care for persons with HIV infection. Society of General Internal Medicine AIDS Task Force. *Ann Intern Med* 1999;131:136–143.
- 65. Mellors JW, Munoz A, Giorgi JV, et al. Plasma viral load and CD4+ lymphocytes as prognostic markers of HIV-1 infection. *Ann Intern Med* 1997;126:946–954.
- Tarwater PM, Gallant JE, Mellors JW, et al. Prognostic value of plasma HIV RNA among highly active antiretroviral therapy users. AIDS 2004; 18:2419–2423.
- 67. Long R, Houston S, Hershfield E; Canadian Tuberculosis Committee of the Centre for Infectious Disease Prevention and Control, Population and Public Health Branch, Health Canada. Recommendations for screening and prevention of tuberculosis in patients with HIV and for screening for HIV in patients with tuberculosis and their contacts. *CMAJ* 2003:169:789–791.
- Hoeppner V, Marciniuk D, Hershfield E. Treatment of tuberculosis disease and infection. In: Long R, ed. *Canadian Tuberculosis Standards*. 5th ed. Ottawa, ON: Canadian Lung Association and Health Canada; 2000:83–109.
- 69. Canadian Immunization Guide. 6th ed. Ottawa, ON: Health Canada; 2002.
- Centers for Disease Control and Prevention (CDC). Recommended adult immunization schedule — United States, October 2005–September 2006. Available at: www.cdc.gov/nip/recs/adult-schedule.htm. Accessed January 19, 2006.
- 71. Piketty C, Darragh TM, Da Costa M, et al. High prevalence of anal human papillomavirus infection and anal cancer precursors among HIV-infected persons in the absence of anal intercourse. *Ann Intern Med* 2003;138:453–459.
- 72. Piketty C, Darragh TM, Heard I, et al. High prevalence of anal squamous intraepithelial lesions in HIV-positive men despite the use of highly active antiretroviral therapy. *Sex Transm Dis* 2004;31:96–99.

- Guidelines for the Use of Antiretroviral Agents in HIV-1-Infected Adults and Adolescents. Rockville, MD: AIDSinfo, Department of Health and Human Services; 2005. Available at: aidsinfo.nih.gov/ContentFiles/ AdultandAdolescentGL.pdf. Accessed January 19, 2006.
- Kaplan JE, Masur H, Holmes KK; USPHS; Infectious Disease Society of America. Guidelines for preventing opportunistic infections among HIV-infected persons—2002. Recommendations of the U.S. Public Health Service and the Infectious Disease Society of America. MMWR Recomm Rep 2002;51 (RR-8):1–52.
- 75. Wald A, Link K. Risk of human immunodeficiency virus infection in herpes simplex virus type 2-seropositive persons: a meta-analysis. *J Infect Dis* 2002;185:45–52.
- Reynolds SJ, Risbud AR, Shepherd ME, et al. Recent herpes simplex virus type 2 infection and the risk of human immunodeficiency virus type 1 acquisition in India. *J Infect Dis* 2003;187:1513–1521.
- 77. Serwadda D, Gray RH, Sewankambo NK, et al. Human immunodeficiency virus acquisition associated with genital ulcer disease and herpes simplex virus type 2 infection: a nested case-control study in Rakai, Uganda. *J Infect Dis* 2003;188:1492–1497.
- 78. Renzi C, Douglas JM Jr, Foster M, et al. Herpes simplex virus type 2 infection as a risk factor for human immunodeficiency virus acquisition in men who have sex with men. *J Infect Dis* 2003:187:19–25.
- 79. Celum CL. The interaction between herpes simplex virus and human immunodeficiency virus. *Herpes* 2004;11(suppl 1):36A–45A.
- Schacker T, Ryncarz AJ, Goddard J, Diem K, Shaughnessy M, Corey L. Frequent recovery of HIV-1 from genital herpes simplex virus lesions in HIV-1-infected men. *JAMA* 1998;280:61–66.
- 81. Golden MP, Kim S, Hammer SM, et al. Activation of human immunodeficiency virus by herpes simplex virus. *J Infect Dis* 1992;166:494–499.
- 82. Mole L, Ripich S, Margolis D, Holodniy M. The impact of active herpes simplex virus infection on human immunodeficiency virus load. *J Infect Dis* 1997;176:766–770.
- 83. Schacker T, Zeh J, Hu H, Shaughnessy M, Corey L. Changes in plasma human immunodeficiency virus type 1 RNA associated with herpes simplex virus reactivation and suppression. *J Infect Dis* 2002;186:1718–1725.
- 84. Grosskurth H, Gray R, Hayes R, Mabey D, Wawer M. Control of sexually transmitted diseases for HIV-1 prevention: understanding the implications of the Mwanza and Rakai trials. *Lancet* 2000;355:1981–1987.
- 85. Celum CL, Robinson NJ, Cohen MS. Potential effect of HIV type 1 antiretroviral and herpes simplex virus type 2 antiviral therapy on transmission and acquisition of HIV type 1 infection. *J Infect Dis* 2005;191(suppl 1):S107–S114.
- 86. Schafer A, Friedmann W, Mielke M, Schwartlander B, Koch MA. The increased frequency of cervical dysplasia-neoplasia in women infected with the human immunodeficiency virus is related to the degree of immunosuppression.

 Am J Obstet Gynecol 1991;164:593–599.

- 87. Duerr A, Kieke B, Warren D, et al; HER study group. Human papillomavirus-associated cervical cytologic abnormalities among women with or at risk of infection with human immunodeficiency virus. *Am J Obstet Gynecol* 2001;184:584–590.
- 88. Smith DK, Grohskopf LA, Black RJ, et al; U.S. Department of Health and Human Services. Antiretroviral postexposure prophylaxis after sexual, injection-drug use, or other nonoccupational exposure to HIV in the United States: recommendations from the U.S. Department of Health and Human Services. MMWR Recomm Rep 2005;54(RR-2):1–20.
- 89. Graham CS, Baden LR, Yu E, et al. Influence of human immunodeficiency virus infection on the course of hepatitis C virus infection: a meta-analysis. *Clin Infect Dis* 2001;33:562–569.
- Ragni MV, Belle SH. Impact of human immunodeficiency virus infection on progression to end-stage liver disease in individuals with hemophilia and hepatitis C virus infection. *J Infect Dis* 2001;183:1112–1155.
- 91. Puoti M, Airoldi M, Bruno R, et al. Hepatitis B virus co-infection in human immunodeficiency virus-infected subjects. *AIDS Rev* 2002;4:27–35.
- Sheng WH, Chen MY, Hsieh SM, et al. Impact of chronic hepatitis B virus (HBV) infection on outcomes of patients infected with HIV in an area where HBV infection is hyperendemic. Clin Infect Dis 2004;38:1471–1417.
- 93. Mathews G, Bhagani S. The epidemiology and natural history of HIV/HBV and HIV/HCV co-infections. *J HIV Ther* 2003;8:77–84.
- 94. Johns DR, Tierney M, Felsenstein D. Alteration in the natural history of neurosyphilis by concurrent infection with the human immunodeficiency virus. *N Engl J Med* 1987;316:1569–1572.
- 95. Hutchinson CM, Hook EW 3rd, Shepherd M, Verley J, Rompalo AM. Altered clinical presentation of early syphilis in patients with human immunodeficiency virus infection. *Ann Intern Med* 1994;121:94–100.
- 96. Gordon SM, Eaton ME, George R, et al. The response of symptomatic neurosyphilis to high-dose intravenous penicillin G in patients with human immunodeficiency virus infection. *N Engl J Med* 1994;331:1469–1473.
- 97. Malone JL, Wallace MR, Hendrick BB, et al. Syphilis and neurosyphilis in a human immunodeficiency virus type-1 seropositive population: evidence for frequent serologic relapse after therapy. *Am J Med* 1995;99:55–63.

LYMPHOGRANULOMA VENEREUM (LGV)

Etiology

- · Caused by Chlamydia trachomatis, serovars L1, L2, L3.
- LGV can be transmitted through vaginal, anal or oral sexual contact.

Epidemiology

- In general, an uncommonly reported sexually transmitted infection (STI) in Canada.
- Endemic in parts of Africa, Asia, South America and the Caribbean,¹ thought to account for 2–10% of genital ulcer disease in areas of India and Africa.²
- A relatively rare disease in industrialized countries; until recently, the majority of cases were acquired in endemic areas.
- However recent outbreaks in men who have sex with men (MSM) starting in the Netherlands in 2003,³ with reports of cases in Belgium,⁴ France,⁵ Germany, Sweden,⁴ the U.K.,⁶ the U.S., ^{7,8} and Canada.⁹
- LGV is not nationally notifiable in the U.S. or Canada. Since the issuing of LGV alerts, cases have started to surface in the U.S., 7,8 and in Canada. 9
- Recent outbreaks among MSM have been associated with concurrent HIV, other STIs, hepatitis C and participation in casual sex gatherings such as "leather scene" parties and high-risk activities such as "fisting."^{3,4}
- LGV may enhance the transmission and acquisition of HIV, other STIs and blood-borne pathogens.
- Nationally notifiable C trachomatis is not broken down into LGV and non-LGV serovars. As such, the national LGV rate is unknown; however, a national enhanced surveillance system was initiated in February 2005 by the Public Health Agency of Canada in partnership with provincial and territorial public health departments.

Prevention

- Condoms or other barrier methods¹⁰ for vaginal, anal and oral sex.
- Extragenital inoculation is possible;¹ therefore, unprotected oral sex is not a safer-sex activity for the prevention of LGV.
- Minimize or avoid sexual activities associated with mucosal damage: for example, fisting, which could facilitate transmission.¹¹ Avoid sharing sex toys and clean toys prior to use.
- See Primary Care and Sexually Transmitted Infections chapter.

Manifestations

- Unlike other *C trachomatis* serovars (A-K), LGV strains are more invasive, preferentially affecting the lymph tissue.³
- Commonly divided into three stages (see Table 1).1

Table 1. Manifestations

Primary LGV	 Incubation period of 3–30 days. Small (1–6 mm), painless papule at site of inoculation (vulva, vagina, penis, rectum, oral cavity, occasionally cervix) that may ulcerate. Self-limited and may go unnoticed in up to 50% of people.¹
Secondary LGV	 Begins within 2–6 weeks of primary lesion.² Often accompanied by significant systemic symptoms, such as low-grade fever, chills, malaise, myalgias, arthralgias; occasionally accompanied by arthritis, pneumonitis or hepatitis/perihepatitis; rarely associated with cardiac involvement, aseptic meningitis and ocular inflammatory disease.² Abscesses and draining sinuses are possible (< 1/3 of patients). Involves the lymph nodes and/or anus and rectum.
Secondary LGV causing lymphadenopathy	 Inguinal/femoral is the most common form and is characterized by painful inguinal and/or femoral lymphadenopathy (unilateral in ½ to ²/₃ of cases), referred to as buboes. "Groove sign": inguinal nodes above and femoral nodes below the inguinal ligament (once considered pathognomonic for LGV). Other lymphadenopathy may occur depending on site of inoculation (e.g., cervical lymphadenopathy following inoculation during oral sex).
Secondary LGV causing anorectal symptoms	 Characterized by acute hemorrhagic proctitis. Symptoms of proctocolitis. Bloody, purulent or mucous discharge from the anus, as well as constipation, are common presenting symptoms.^{3,9,10,12}
Tertiary LGV (chronic LGV occurring in 10–20% of untreated cases)	 More common in females than males. Chronic inflammatory lesions lead to scarring. Lymphatic obstruction causing genital elephantiasis.^{1,2,13} Genital and rectal strictures and fistulae. Possible extensive destruction of genitalia (esthiomene).

Diagnosis

• The diagnosis of LGV is not always straightforward. The symptoms and signs of LGV significantly overlap with other STIs, other infections, drug reactions and malignancies. The diagnosis is often based on the history and clinical picture and is supported by laboratory testing, although in Canada confirmatory testing for LGV is now readily available in some laboratories (see Laboratory testing, below). For surveillance purposes, only cases positive by LGV confirmatory tests are considered confirmed cases.9 It may be appropriate, however, for less strict clinical, epidemiologic and laboratory criteria to be used for the clinical management of cases and contacts.

Diagnostic procedures

- · Anoscopy/sigmoidoscopy/proctoscopy
 - Pattern similar to ulcerative colitis.
 - Granular or ulcerative proctitis.
- · Bubo aspiration
 - Buboes in LGV usually contain a small amount of milky fluid.
 - May require injection of 2-5 mL of sterile saline for aspiration.
 - Buboes should be aspirated through healthy skin.

Laboratory testing

- Routine tests for C trachomatis may be positive in patients with LGV, but generally do not include typing to distinguish LGV serovars from non-LGV serovars. Definitive diagnosis of LGV requires serovar-specific (confirmatory) testing using DNA sequencing or restriction fragment length polymorphism (RFLP). Clinicians will therefore need to request that testing be done for LGV specifically, as most laboratories will not automatically perform serovar typing.
- The availability and type of testing for LGV varies by laboratory. Some local
 laboratories are able to do confirmatory testing for LGV, while others will need
 to involve the National Microbiology Laboratory (NML) via their provincial
 laboratory. Please check with your local laboratory for more information on
 how to collect and transport specimens. Where possible, suspected cases of
 LGV should have both swab and sera samples submitted for laboratory testing.
 Serology and confirmatory testing (DNA sequencing and RFLP) are available at
 the NML.

Table 2. Laboratory testing

Type of test	Test specifics	Differentiate between LGV and non-LGV		
		serovars		
Tests for <i>C trachomatis</i> (not specific to LGV serovars)				
Culture	Culture for <i>C trachomatis</i>	 No Positive specimens may be sent for RFLP or DNA sequencing to identify LGV serovars 		
NAAT	PCR, LCR, TMA and SDA	No Positive specimens may be sent for RFLP or DNA sequencing to identify LGV serovars		
Serology	Testing modalities vary by laboratory: • MIF test for <i>C trachomatis</i> : high-titre (titre ≥1:256) • CF test for <i>C trachomatis</i> : positive (titre ≥1:64) - MIF is a more specific test for LGV than CF - Cross-reactivity may be an issue with CF	 No Because of the invasive nature of LGV, serology titres are in general significantly higher in LGV vs. non-LGV <i>C trachomatis</i> infections High-titre serology is suggestive of LGV infection but is not definitive; low-titre serology does not eliminate possibility of past or current LGV infection 		
LGV-specific te	ests (confirmatory)			
DNA sequencing	Definitively identifies LGV serovars	 Yes Samples that test positive for <i>C trachomatis</i> with NAAT or culture can be sent for DNA sequencing* 		
RFLP	Definitively identifies LGV serovars	 Yes Samples that test positive for <i>C trachomatis</i> with NAAT or culture can be sent for RFLP testing* 		

CF = complement fixation

LGV = lymphogranuloma venereum

NAAT = nucleic acid amplification test

RFLP = restriction fragment length polymorphism

TMA = transcription-mediated amplification

LCR = ligase chain reaction

MIF = microimmunofluorescence

PCR = polymerase chain reaction

SDA = strand displacement amplification

^{*} For laboratories sending samples to NML for confirmatory testing (DNA sequencing or RFLP), please note that it is the original sample that must be submitted to NML. These samples will be tested by PCR for *omp*1, and this PCR product is what must be sent for sequencing by the NML.

Specimen collection

• Table 3 describes types of specimens that may be collected for the laboratory tests described above, for the diagnosis of LGV by stage of infection.

Table 3. Specimen collection

Stage of infection	Sample type	Tests	Comments
Primary	Swab of lesion	Culture or NAAT	Because the invasive nature of LGV has not yet manifested in the primary stage of the infection, serology at this stage is unlikely to be helpful
Secondary and tertiary	Bubo aspirate	Culture or NAAT	Identification of <i>C trachomatis</i> in bubo fluid is highly suggestive of LGV, even prior to or without identification of LGV serovars
	Rectal, vaginal, oropharyngeal, or urethral swab	Culture or NAAT	NAAT is not officially approved in Canada for use with rectal or oropharyngeal swabs. Repeat testing is advised to confirm a positive test
	Urine	NAAT	
	Serology	MIF test	See Table 2
		CF test	

CF = complement fixation

LGV = lymphogranuloma venereum

MIF = microimmunofluorescence

NAAT = nucleic acid amplification test

- For samples being sent to the NML, the following storage and shipping recommendations apply:
 - Dry swabs should be stored and shipped frozen.
 - Swabs stored in chlamydia transport media should be kept frozen at -80°C if culture will be done, or at -20°C if culture will not be done.
 - Urine samples should be stored and shipped frozen.
 - See Laboratory Diagnosis of Sexually Transmitted Infections chapter for more information on collecting and shipping specimens.

Management

- Treatment with appropriate antibiotic regimen (see Treatment section, below).
- Aspiration of buboes may help symptomatically; however, incision/drainage or excision of nodes is not helpful and may delay healing.

Treatment

 Suspected cases should be treated empirically for LGV while awaiting test results.

Table 4. Treatment of lymphogranuloma venereum

First line	Doxycycline 100 mg PO bid for 21 days [B-II]
Alternative	• Erythromycin 500 mg PO qid for 21 days* [C-III]
Possible	Azithromycin 1g PO once weekly for 3 weeks† [C-III]

^{*} Erythromycin dosage refers to the use of erythromycin base. Equivalent dosages of other formulations may be substituted (with the exception of the estolate formulation, which is contraindicated in pregnancy); erythromycin (NOT the estolate formulation) should be used in pregnancy.

Treatment of partners

- Sexual partners from the last 60 days prior to symptom onset or date of diagnosis where asymptomatic should be contacted, tested and treated empirically (regardless of whether signs/symptoms are present) as follows:
 - Azithromycin 1g PO in a single dose [C-III]
 OR
 - Doxycycline 100 mg PO bid for 7 days [C-III]
- Should test results confirm an LGV infection, treat as recommended for cases above.

[†] While some experts believe azithromycin to be effective in the treatment of LGV, clinical data are lacking.

Consideration for Other STIs

- Because of rates of co-infection, testing for HIV, syphilis, HSV, gonorrhea, hepatitis B and hepatitis C is recommended in patients with LGV (see chapters on individual infections for more information on testing).
- Testing for chancroid and donovanosis (granuloma inguinale) should also be considered in patients with LGV, especially if there has been travel to regions where these infections are endemic.
- Immunization for hepatitis B should be offered to non-immune patients (see *Hepatitis B Virus Infections* chapter for more information).
- The opportunity to provide safer-sex counselling should not be missed.

Reporting and Partner Notification

- An enhanced surveillance system was initiated by the Public Health Agency of Canada, in partnership with the provinces and territories, in February 2005.
 - LGV should be reported by local public health authorities to the appropriate regional and provincial/territorial authorities, who have, in turn, agreed to report LGV to the Sexual Health and STI Section of the Public Health Agency of Canada.
 - Case definitions for national enhanced surveillance as of August 2005 are as follows.⁹
- Any sexual partners from the last 60 days prior to symptom onset or date
 of diagnosis where asymptomatic should be contacted, tested and treated
 (see Treatment section).

Table 5. Case definitions

Probable case	Positive result on culture, NAAT or serologic testing for C trachomatis plus the presence of proctitis OR inguinal or femoral lymphadenopathy OR a sexual partner with LGV
Confirmed case	Presence of <i>C trachomatis</i> serotype L1, L2, L3 confirmed by DNA sequencing or RFLP

LGV = lymphogranuloma venereum NAAT = nucleic acid amplification test RFLP = restriction fragment length polymorphism

Follow-up

- Patients should be followed until chlamydial tests are negative (test of cure) and the patient has clinically recovered.³ Serology should not be used to monitor treatment response, as the duration of antibody response has not been defined.
 - Test of cure should be performed at 3–4 weeks after the completion of effective treatment to avoid false-positive results due to the presence of nonviable organisms (especially if using NAAT).
- Surgery may be required to repair genital/rectal damage of tertiary LGV.

Special Considerations

- Based on limited data, HIV appears to have little effect on the clinical presentation, although atypical presentations in HIV-positive patients have been rarely reported.¹⁴
- Disease duration may be prolonged in HIV-positive patients.¹⁴
- In pregnancy, erythromycin (non-estolate preparations) should be used for the treatment of LGV.

References

- Mabey D, Peeling RW. Lymphogranuloma venereum. Sex Transm Infect 2002;78:90–92.
- Roest RW, van der Meijden WI; European Branch of the International Union Against Sexually Transmitted Infection and the European Office of the World Health Organization. European guideline for the management of tropical genitor-ulcerative diseases. *Int J STD AIDS* 2001;12(suppl 3):78–83.
- 3. Nieuwenhuis RF, Ossewaarde JM, Götz HM, et al. Resurgence of Lymphogranuloma venereum in Western Europe: an outbreak of *Chlamydia trachomatis* serovar I2 proctitis in The Netherlands among men who have sex with men. *Clin Infect Dis 2004*;39:996–1003.
- Centers for Disease Control and Prevention. Lymphogranuloma venereum among men who have sex with men — Netherlands, 2003–2004. MMWR Morb Mortal Wkly Rep 2004;53:985–988.
- Institut de veille sanitaire. Emergence de la Lymphogranulomatose vénérienne rectale en France: cas estimés au 31 mars 2004. Synthèse réalisée le 1^{er} juin 2004. Available at: www.invs.sante.fr/presse/2004/le_point_sur/lgv_160604/. Accessed February 14, 2006.
- Health Protection Agency. Enhanced surveillance of lymphogranuloma venereum (LGV) in England. CDR Weekly 2004;14:3. Available at: www.hpa.org. uk/cdr/archives/2004/cdr4104.pdf. Accessed February 14, 2006.
- Lymphogranuloma venereum USA (California). ProMED-mail. December 22, 2004. Archive number:20041222.3376. Available at: www.promedmail.org. Accessed February 14, 2006.
- Lymphogranuloma venereum USA (Texas). ProMED-mail. December 24, 2004. Archive number: 20041224.3397. Available at: www.promedmail.org. Accessed February 14, 2006.

- Kropp RY, Wong T; Canadian LGV Working Group. Emergence of lymphogranuloma venereum in Canada. CMAJ 2005;172:1674–1676.
- Weir E. Lymphogranuloma venereum in the differential diagnosis of proctitis. *CMAJ* 2005:172:185
- Gotz HM, van Doornum G, Niesters HG, den Hollander JG, Thio HB, de Zwart O. A cluster of acute hepatitis C virus infection among men who have sex with men — results from contact tracing and public health implications. *AIDS* 2005; 19:969–974.
- 12. Goens JL, Schwartz RA, DeWolf K. Mucocutaneous manifestations of chancroid, lymphogranuloma venereum and granuloma inguinale. *Am Fam Physician* 1994;49:415–418, 423–425.
- 13. Aggarwal K, Jain VK, Gupta S. Bilateral groove sign with penoscrotal elephantiasis. *Sex Transm Infect* 2002;78:458.
- 14. Czelusta A, Yen-Moore A, Van der Straten M, Carrasco D, Styring K. An overview of sexually transmitted diseases. Part III. Sexually transmitted diseases in HIV-infected patients. *J Am Acad Dermatol* 2000;43:409–432.

SYPHILIS

Etiology

- Caused by Treponema pallidum subsp. pallidum.
- T pallidum subsp. pallidum causes venereal syphilis, T pallidum subsp. endemicum causes endemic syphilis (bejel), T pallidum subsp. pertenue causes yaws and T carateum causes pinta.

Epidemiology

- Infectious syphilis (primary, secondary and early latent stages) is the least common of the three nationally reportable sexually transmitted infections (STIs).
- After achieving rates of 0.4–0.6/100,000 from 1994–2000, rates of infectious syphilis rose in 2002 to 1.5/100,000, and preliminary figures for 2004 show projected rates of 3.9/100,000.^{1,2} (Preliminary data is subject to change; does not include Nunavut.)
- Most affected are males 30–39 years (14.5 per 100,000 population in 2004).¹
 (Preliminary data is subject to change; does not include Nunavut.)
- The rate of infectious syphilis is increasing in both males and females, but more so in males. In recent years, localized outbreaks of infectious syphilis have been reported in a number of locations worldwide^{3,4} and in Canada, including Vancouver, Yukon, Calgary, Edmonton, Toronto, Ottawa, Montreal and Halifax.^{2,5-7}
- Most of the outbreaks have been related to the sex trade and in men who have sex with men (MSM), but some have been in heterosexual persons not fitting into one of these categories. Some large outbreaks among MSM have been associated with the acquisition of anonymous sex partners through the Internet.⁸
- Syphilis, as with other STIs, increases the risk of acquisition and transmission
 of HIV.

Transmission

- The primary mode of transmission is by vaginal, anal and oral sexual contact.
- Kissing, sharing of needles and injection equipment, blood transfusion and accidental inoculation have rarely been reported as routes of transmission.
- Primary, secondary and early latent stages are considered infectious, with an
 estimated risk of transmission per partner of around 60%.¹⁰ Early latent syphilis is
 considered infectious because of the 25% chance of relapse to secondary stage.¹¹
- The majority of infants with congenital syphilis are infected in utero, but they
 can also be infected by contact with an active genital lesion at the time of
 delivery; the risk of transmission is much greater when the mother has untreated
 primary, secondary or early latent syphilis in pregnancy than if she has late latent
 syphilis.¹²

Prevention

• Results of reactive syphilis tests in a pregnant mother and any treatment history should be provided to the primary caregiver of the newborn infant.

Manifestations

Table 1. Manifestations9

Stage	Clinical manifestations	Incubation period
Primary	Chancre, regional lymphadenopathy	3 weeks (3–90 days)
Secondary	Rash, fever, malaise, lymphadenopathy, mucus lesions, condyloma lata, alopecia, meningitis, headaches, uveitis, retinitis	2–12 weeks (2 weeks– 6 months)
Latent	Asymptomatic	Early: <1 year Late: ≥1 year
Tertiary Cardiovascular syphilis	Aortic aneurysm, aortic regurgitation, coronary artery ostial stenosis	10-30 years
Neurosyphilis	Ranges from asymptomatic to symptomatic with headaches, vertigo, personality changes, dementia, ataxia, presence of Argyll Robertson pupil	<2 years– 20 years
Gumma	Tissue destruction of any organ; manifestations depend on site involved	1–46 years (most cases 15 years)
Congenital Early	Fulminant disseminated infection, mucocutaneous lesions, osteochondritis, anemia, hepatosplenomegaly, neurosyphilis	Onset <2 years
Late	Interstitial keratitis, lymphadenopathy, hepatosplenomegaly, bone involvement, anemia, Hutchinson's teeth, neurosyphilis	Persistence >2 years after birth

Diagnosis

Risk factors

A diagnosis of syphilis should be considered in the following individuals:

- Those who have had contact with a known case of syphilis.
- MSM.
- · Sex workers.
- Those with street involvement.
- Injection drug users.
- Those with multiple sex partners.

- Those with a history of syphilis, HIV and other STIs.
- Those originating from or having sex with an individual from a country with a
 high prevalence of syphilis; it should be noted that screening for syphilis (using
 a non-treponemal test) is routinely performed in all immigration applicants to
 Canada who are older than 15 years.
- Sexual partners of any of the above.

Symptoms and signs

- Current or past history of lesions or rash (see Manifestations, above).
- A high proportion of individuals fail to recall a primary chancre.⁹
- Signs and symptoms may be modified in the presence of HIV co-infection.¹³

Special considerations in pregnant women

- Given the resurgence of syphilis in Canada, universal screening of pregnant women continues to be important and remains the standard of care in most jurisdictions.
- Screening should ideally be performed in the first trimester and repeated later in pregnancy in women at high risk of acquiring syphilis (see Risk factors, above).

Laboratory diagnosis

- The interpretation of syphilis serology should be made in conjunction with a colleague experienced in this area (see Table 2).
- Every attempt should be made to obtain and document prior history of treatment for syphilis and prior serologic results in order to avoid unnecessary retreatment.

Specimen collection

- Dark-field microscopy, direct or indirect fluorescent antibody tests (DFA/IFA) or
 polymerase chain reaction (PCR) (for more information on available tests, please
 contact your local laboratory). To visualize *T pallidum* from chancres of primary
 syphilis and some lesions of secondary syphilis (e.g., condyloma lata).
- Dark-field microscopy and DFA/IFA are not reliable for oral/rectal lesions, as non-pathogenic treponemes may be present.
- PCR is available only at specialized laboratories, including the National Microbiology Laboratory.

Serology

Screening for syphilis has traditionally involved the use of non-treponemal tests
 (NTT) such as rapid plasma reagin (RPR), followed by confirmatory treponemal
 tests if the NTT is reactive. However, in patients with suspected primary syphilis or
 late latent syphilis, the NTT may be non-reactive, and it is then appropriate to add
 a treponemal test to the initial screen or, in the case of primary syphilis, to repeat
 the NTT after 2–4 weeks. In regions experiencing outbreaks of syphilis, it may be
 appropriate to screen at baseline with both non-treponemal and treponemal tests.

Table 2. Guide to interpretation of serologic tests for syphilis

Table 2. Guide to interpretation of serologic tests for syphilis			
Test results on blood or serum			
Non- treponemal test: RPR/VDRL	Treponemal test: TP-PA	Treponemal test: FTA-ABS	Most likely condition
NR	NR	R	Primary syphilis with compatible history/clinical findings
R (dilutions can vary)	R	R	 Infectious syphilis (primary, secondary, early latent), especially if titre >1:8 OR Old treated syphilis (especially if titre <1:8) OR Follow-up of treated syphilis OR In persons from endemic countries, yaws (e.g., Caribbean), pinta (e.g., Central America) or bejel
NR	R	R	 Usually treated syphilis OR Late latent of unknown duration if no history of confirmed treatment OR In persons from endemic countries, yaws (e.g., Caribbean), pinta (e.g., Central America) or bejel OR Early infection (primary syphilis)
R	NR	NR	Biological false positive* (repeat in 3–4 weeks)

FTA-ABS = fluorescent treponemal antibody absorbed

NR = non-reactive

R = reactive

RPR = rapid plasma reagin

TP-PA = *T pallidum* particle agglutination

VDRL = Venereal Disease Research Laboratory

^{*} Some causes of false-positive serologic tests for syphilis include certain collagen-vascular diseases, pregnancy, injection drug use, etc.

- The introduction of treponemal tests for IgG/IgM antibodies, such as the treponemal enzyme immunoassay (EIA), may provide a more sensitive screening test for syphilis.
- Non-treponemal tests include RPR, Venereal Disease Research Laboratory (VDRL) and the toluidine red unheated serum test (TRUST).
- Non-treponemal antibody titres usually correlate with disease activity and are used to monitor response to treatment and assess for reinfection.
- Treponemal tests include the T pallidum particle agglutination (TP-PA), fluorescent treponemal antibody absorbed (FTA-ABS) and EIA to detect IgG and/or IgM antibodies.
- Treponemal tests usually remain reactive for life regardless of treatment, although 15–25% will serorevert if the patient is treated during the primary stage.

Cerebrospinal fluid

- Criteria for cerebrospinal fluid (CSF) examination include the following:
 - Presence of neurologic or ophthalmic symptoms or signs.
 - Congenital syphilis.
 - Previously treated patients who fail to achieve an adequate serologic response to treatment.
 - Tertiary syphilis.¹⁴
 - HIV patients with neurologic symptoms or signs, late latent syphilis, RPR ≥1:32 dilutions, CD4 <350 cells/μL or treated syphilis with suboptimal decline in VDRL/RPR titre; some experts recommend CSF examination in all cases.¹⁵
 - Some experts recommend CSF examination in all patients with RPR ≥1:32 dilutions.¹⁵
- CSF should be tested for cell count and differential, protein, VDRL and/or FTA-ABS.
- CSF-VDRL is highly specific but insensitive.
- CSF FTA-ABS is highly sensitive but non-specific for neurosyphilis; a negative CSF FTA-ABS helps to exclude a diagnosis of neurosyphilis.^{14,16-18}
- The diagnosis of neurosyphilis is usually made on a combination of reactive serologic results, abnormalities of CSF cell count or protein or a reactive CSF-VDRL with or without clinical manifestations.

Management

Primary and secondary syphilis

- Attempt to obtain material from primary or secondary lesions for dark-field microscopy and/or DFA/IFA for T pallidum.
- Ulcers should also be tested for herpes simplex virus and/or chancroid (if epidemiologically appropriate) and/or lymphogranuloma venereum (if epidemiologically appropriate).

Serology should include both treponemal and non-treponemal tests. Note that
both non-treponemal and treponemal tests may be negative in early primary
syphilis. Serology should be repeated in 2–4 weeks if they are dark-field or
DFA/IFA negative and/or no treatment has been given. If follow-up cannot be
assured, it may be appropriate to treat presumptively for primary syphilis.

Latent syphilis

- Serology: both treponemal and non-treponemal tests; note that a negative non-treponemal test does not rule out the diagnosis of latent syphilis.
- All patients should undergo a physical examination, including neurologic
 examination, to evaluate for the presence of signs of tertiary syphilis. Chest x-ray
 may be appropriate to evaluate for the presence of cardiovascular syphilis
 (e.g., aneurysm of ascending aorta).
- Lumbar puncture may be appropriate (see Cerebrospinal fluid, above).
- Treat as appropriate for stage.

Tertiary syphilis

- Serology: both treponemal and non-treponemal tests; note that a negative non-treponemal test does not rule out the diagnosis of tertiary syphilis.
- All patients with suspected tertiary syphilis should undergo CSF examination.
 - If CSF is not compatible with a central nervous system (CNS) infection, treat as for late latent syphilis.
 - If CSF is compatible with a CNS infection, treat as for neurosyphilis.

Congenital syphilis

- Obtain venous samples from both mother and baby (note that cord blood is not suitable) for serology (treponemal and non-treponemal tests).
 - The interpretation of reactive antibodies in the neonate must take into consideration the maternal history, including stage of syphilis, history of treatment, and syphilis serology results.
- Placenta, neonatal nasal discharge or skin lesions may be examined by darkfield microscopy or DFA/IFA for T pallidum.
- CSF examination should be performed on all infants with suspected congenital syphilis.
- Long-bone x-rays should be performed.

Treatment

- Although regimens containing daily IM procaine penicillin for 10–14 days are
 equally efficacious to regimens containing benzathine penicillin G, the latter are
 preferred because of better adherence with less frequent dosing.
- Benzathine penicillin G is available in Canada only through provincial/territorial sexually transmitted infection services, which obtain the drug from non-Canadian pharmaceutical companies through Health Canada's Special Access Program, as the drug is no longer available in Canada.

Table 3. Treatment

Stage	Preferred treatment	Alternative treatment for penicillin-allergic patients
All non-pregnant adultsPrimarySecondaryEarly latent (<1 year duration)	Benzathine penicillin G 2.4 million units IM as a single dose*19-22 [A-II; A-III for HIV-infected individuals]	 Doxycycline 100 mg PO bid for 14 days^{23,24} [<i>B-III</i>] Alternative agents (to be used in exceptional circumstances):[†] ceftriaxone 1 g IV or IM daily for 10 days^{25,26} [<i>B-III</i>]
Pregnant women Primary Secondary Early latent (<1 year duration)	• Benzathine penicillin G 2.4 million units IM as a single dose* ²⁷ [A-II]	There is no satisfactory alternative to penicillin for the treatment of syphilis in pregnancy; insufficient data exist to recommend ceftriaxone in pregnancy Strongly consider penicillin desensitization, followed by treatment with penicillin [A-III]
 All non-pregnant adults Late latent syphilis Latent syphilis of unknown duration Cardiovascular syphilis and other tertiary syphilis not involving the CNS 	Benzathine penicillin G 2.4 million units IM weekly for 3 doses ^{28,29} [A-II]	 Consider penicillin desensitization Doxycycline 100 mg PO bid for 28 days²⁴ [B-II] Alternative agents (to be used in exceptional circumstances):¹ ceftriaxone 1 g IV or IM daily for 10 days³⁰ [C-III]
 Pregnant women Late latent syphilis Latent syphilis of unknown duration Cardiovascular syphilis and other tertiary syphilis not involving the CNS 	Benzathine penicillin G 2.4 million units IM weekly for 3 doses ³¹ [A-II]	There is no satisfactory alternative to penicillin for the treatment of syphilis in pregnancy; insufficient data exist to recommend ceftriaxone in pregnancy Strongly consider penicillin desensitization, followed by treatment with penicillin [A-III]

Table 3. Treatment (continued)

Stage	Preferred treatment	Alternative treatment for penicillin-allergic patients
All adults • Neurosyphilis	• Penicillin G 3–4 million units IV every 4 hours (16–24 million units/day) for 10–14 days ²⁹ [A-II]	 Strongly consider penicillin desensitization, followed by treatment with penicillin Ceftriaxone 2 g IV/IM per day for 10–14 days^{29,32,33} [B-II]
Congenital syphilis ³⁴	Early (<1 month): crystalline penicillin G 50,000 units/kg IV every 12 hours for the first week of life and every 8 hours thereafter for 10 days of total therapy [A-II]	
	• Late (≥1 month): crystalline penicillin G 50,000 units/kg/ IV every 6 hours for 10–14 days [A-II]	 If no neurologic involvement and normal CSF: benzathine penicillin G 50,000 units/kg IM (max 2.4 million units) weekly for 3 successive weeks [B-II] No data are available to recommend penicillin alternatives in the case of penicillin allergy
Epidemiological treatment of sexual contacts in the preceding 30 days to primary, secondary and early latent syphilis §¥35	Benzathine penicillin G 2.4 million units IM as a single dose [B-II]	See comment below on azithromycin [¥]

CNS = central nervous system

- * Some experts recommend 3 weekly doses (total of 7.2 million units) of benzathine penicillin G in HIV-infected individuals.
- † The efficacy data supporting the use of these agents is limited, and as such they should only be used in exceptional circumstances and when close patient follow-up is assured.
- ‡ Secondary syphilis in late pregnancy (>20 weeks gestation) should be treated with two doses of benzathine penicillin G 2.4 million units given 1 week apart (see note under Pregnancy, below).
- § If sexual contact is unreliable or unable to test, epidemiological treatment should be strongly considered.
- ¥ Azithromycin: in light of recent reports of failure of azithromycin for the treatment of early syphilis³⁶ and the rapid development of azithromycin resistance in *T pallidum*,^{31,38} this agent should not be routinely used as a treatment option for early or incubating syphilis, unless adequate and close follow-up can be ensured, and only in jurisdictions where little to no azithromycin genotypic resistance in *T pallidum* has been demonstrated. It should be noted, however, that at the present time, very limited Canadian data on the prevalence of azithromycin resistance in T pallidum is available, with 1 of 47 specimens between 2000 and 2003 as compared with 4 of 12 specimens from MSM in 2004–2005 collected in Vancouver demonstrating resistance.³⁸

Penicillin desensitization

- Skin testing with the major and minor determinants can reliably identify persons at high risk for penicillin reactions.
- Patients who have a positive skin test to one of the penicillin determinants can be desensitized.

Table 4. Oral desensitization protocol for patients with a positive skin test³⁹

Penicillin V suspension dose number*	Amount† units/mL	Volume administered (mL)	Units	Cumulative dose (units)
1	1,000	0.1	100	100
2	1,000	0.2	200	300
3	1,000	0.4	400	700
4	1,000	0.8	800	1,500
5	1,000	1.6	1,600	3,100
6	1,000	3.2	3,200	6,300
7	1,000	6.4	6,400	12,700
8	10,000	1.2	12,000	24,700
9	10,000	2.4	24,000	48,700
10	10,000	4.8	48,000	96,700
11	80,000	1.0	80,000	176,700
12	80,000	2.0	160,000	336,700
13	80,000	4.0	320,000	656,700
14	80,000	8.0	640,000	1,296,700

^{*} Interval between doses, 15 minutes; elapsed time, 3 hours and 45 minutes; cumulative dose, 1.3 million units.

[†] The specific amount of drug is diluted in approximately 30 mL of water and then administered orally.

- Oral desensitization is preferable to IV desensitization, as it is safer and less costly.
- Desensitization should occur in a hospital setting, as serious allergic reactions, although unlikely, can occur. The whole procedure can usually be completed in 4 hours, after which the first dose of penicillin is given. After administration of the dose, the patient should be observed for at least 1 hour.

Consideration for Other STIs

- All patients with reactive syphilis serology should be tested for HIV, as this
 affects treatment and follow-up.
- Testing for other STIs, including chlamydia and gonorrhea, should be performed.
- Genital ulcers should also be tested for herpes simplex virus and/or chancroid and/or lymphogranuloma venereum, depending on epidemiologic risk.
- Immunization against hepatitis B and/or A may be indicated if not already immune.

Reporting and Partner Notification

- Infectious syphilis (primary, secondary and early latent syphilis) is reportable in all provinces and territories and to the Public Health Agency of Canada.
- Non-infectious syphilis (late latent, cardiovascular and neurosyphilis) may be reportable at the provincial/territorial level but is not reportable to the Public Health Agency of Canada.
- All sexual or perinatal contacts within the following time periods must be located, tested and treated if serology is reactive.

Table 5. Partner notification

Stage of syphilis	Time period	
Primary syphilis	3 months prior to the onset of symptoms	
Secondary syphilis	6 months prior to the onset of symptoms	
Early latent	1 year prior to the diagnosis	
Late latent	Assess marital or other long-term partners and children as appropriate	
Congenital	Assess mother and her sexual partner(s)	
Stage undetermined	Assess/consult with a colleague experienced in syphilis management	

Follow-up

- In the absence of a test of cure, NTTs should be monitored until they are seronegative or at a stable low titre (e.g., 1:4 dilutions).⁴⁰
- See Table 6 for a guide to the monitoring of NTTs.
- See Table 7 for a guide to adequate serologic response (in NTT: e.g., RPR).⁴¹
- Note that the NTT may revert to non-reactive after treatment or remain at a low steady level (sero-fast); repeat testing is not required if the baseline or follow-up NTT becomes non-reactive, except in HIV-infected individuals.
- A rising NTT after treatment may indicate treatment failure or reinfection. If treatment failure is suspected, further investigation, including CSF examination, may be indicated.

Table 6. Monitoring of NTTs

Primary, secondary, early latent	1, 3, 6, 12 months after treatment
Late latent, tertiary	12 and 24 months after treatment
Neurosyphilis	6, 12 and 24 months after treatment
HIV-infected (any stage)	1, 3, 6, 12 and 24 months after treatment and yearly thereafter
Babies born to mothers with reactive syphilis serology*	3 and 6 months after birth; repeat non-treponemal and treponemal tests at 12 and 18 months if tests remain reactive at 6 months
Congenital syphilis*	0, 3, 6, 12 and 18 months after birth

^{*} NTT titres should decline by 3 months of age and be non-reactive by 6 months if the infant was not infected. If the titres are stable or increase after 6–12 months of age, the child should be evaluated (including cerebrospinal fluid examination) and treated as for congenital syphilis. Passively transferred treponemal antibodies can be present in an infant up to 15 months; a reactive treponemal test after 18 months is diagnostic of congenital syphilis.

Table 7. Adequate serologic response

Primary	2-tube* drop at 6 months, 3-tube drop at 12 months, 4-tube drop at 24 months
Secondary	3-tube and 4-tube drop at 6 and 12 months, respectively
Early latent	2-tube drop at 12 months

^{*2-}tube drop=four-fold drop, e.g., change from 1:32 dilutions to 1:8 dilutions.

Patients with neurosyphilis and abnormal CSF examinations should have a lumbar puncture repeated at 6 month intervals after completion of treatment until CSF parameters normalize. CSF pleocytosis is generally the first measure of improvement and should occur over about 6 months.⁴² Elevated protein levels, if present, will begin to decline during the first 6 months but can take up to 2 years to return to normal.⁴³ CSF protein may decline more slowly in patients who are neurologically abnormal compared with those who are neurologically normal.⁴⁴ The CSF-VDRL titre should decline (four-fold within a year) if it is initially high, but it may take years to revert to negative. 42 A persistent, low CSF-VDRL titre after a course of treatment may warrant retreatment, but if CSF pleocytosis and elevated protein levels have resolved and serum VDRL titre has not risen, additional treatment is unlikely to be beneficial.⁴⁵ All CSF laboratory parameters normalize more slowly in patients co-infected with HIV.44 The possibility of treatment failure should be considered if there is clinical progression, increase in RPR/VDRL by ≥2 dilutions or CSF pleocytosis fails to resolve; treatment options for patients with treatment failure should be discussed with a colleague experienced in this area.

Special Considerations

HIV infection

 Persons co-infected with HIV may require a longer course of treatment, as well as closer and longer follow-up.

Pregnancy⁴⁶

- All women newly diagnosed with syphilis during pregnancy should receive treatment appropriate to their stage of disease, with the exception of secondary syphilis in late pregnancy, where despite the administration of the recommended penicillin regimen, as many as 14% will have a fetal death or deliver infants with clinical evidence of congenital syphilis.^{47–49} These cases should therefore be treated with two doses of benzathine penicillin G 2.4 million units 1 week apart, although the effect of this regimen in preventing fetal syphilis is not known.⁴⁶
- Retreatment during pregnancy is not necessary unless there is clinical or serologic evidence of new infection (four-fold rise in a non-treponemal test titre) or history of recent sexual contact with early syphilis.
- Erythromycin is the least effective agent for the treatment of syphilis and does not penetrate the CSF or placental barrier well; it is therefore not recommended in pregnancy.^{50, 51}
- If the mother is >20 weeks gestation, an ultrasound should be performed and she should be managed with an obstetrician/maternal-fetal medicine specialist; if fetal abnormalities are identified, the mother should be hospitalized for treatment and fetal monitoring.⁵²
- All babies should be assessed at delivery by a pediatrician, and if a maternal non-penicillin regimen was used, consideration should be given to treating the baby empirically for congenital syphilis.

Congenital syphilis53

- Infected infants are frequently asymptomatic at birth and may be seronegative if maternal infection occurred late in gestation.
- · Infants should be treated at birth:
 - If symptomatic.
 - If the infant's non-treponemal titre is four-fold (2 tubes) higher than the mother's.
 - If maternal treatment was inadequate, did not contain penicillin, is unknown or occurred in the last month of pregnancy, or if maternal serologic response is inadequate.
 - If adequate follow-up of the infant cannot be ensured.

Jarisch-Herxheimer reaction54

- Patients should be made aware of this possible reaction to treatment, especially with penicillin.
- An acute febrile illness with headache, myalgia, chills, rigours generally occurring within 8–12 hours and resolving within 24 hours.
- Common in early syphilis, but usually not clinically significant unless there is neurologic or ophthalmic involvement or in pregnancy where it may cause fetal distress and premature labour.
- · Not a drug allergy.
- · Can be treated with antipyretics.
- Steroids may be indicated for the management of severe reactions but should be used in consultation with a colleague experienced in this area.

References

- 1. Unpublished data. Surveillance and Epidemiology Section, Community Acquired Infections Division, Public Health Agency of Canada, 2006.
- Public Health Agency of Canada. Reported Infectious Syphilis Cases and Rates in Canada by Province/Territory and Sex, 1993-2002. Available at: www.phac-aspc.gc.ca/std-mts/stddata_pre06_04/tab3-2_e.html. Accessed July 14, 2005.
- Centers for Disease Control and Prevention. Sexually Transmitted Disease Surveillance 2003 Supplement. Syphilis Surveillance Report, December 2004. Atlanta, GA: Centers for Disease Control and Prevention; 2004. Available at: www.cdc.gov/std/Syphilis2003/SyphSurvSupp2003.pdf. Accessed July 14, 2005.
- 4. Righarts AA, Simms I, Wallace L, Solomou M, Fenton KA. Syphilis surveillance and epidemiology in the United Kingdom. *Euro Surveill* 2004;9:21–25.
- 5. Sarwal S, Shahin R, Ackery JA, Wong T. Infectious syphilis in MSM, 2002: outbreak investigation. Paper presented at: Annual Meeting of the International Society for STD Research; July 2003; Ottawa, ON. Abstract 0686.

- Shahin R, Sarwal S, Ackery JA, Wong T. Infectious syphilis in MSM, 2002: public health interventions. Paper presented at: Annual Meeting of the International Society for STD Research; July 2003; Ottawa, ON. Abstract 0685.
- Alberta Health and Wellness. Notifiable diseases. www.health.gov.ab.ca/ regions/require/list.htm. Accessed July 18, 2005.
- Klausner JD, Wolf W, Fischer-Ponce L, Zolt I, Katz MH. Tracing a syphilis outbreak through cyberspace. *JAMA* 2000;284:447–449.
- Singh AE, Romanowski B. Syphilis: review with emphasis on clinical, epidemiologic, and some biologic features. Clin Microbiol Rev 1999;12:187–209.
- Garnett GP, Aral SO, Hoyle DV, Cates W Jr, Anderson RM. The natural history of syphilis. Implications for the transmission dynamics and control of infection. Sex Transm Dis 1997;24:185–200.
- Gjestland T. The Oslo study of untreated syphilis: an epidemiologic investigation of the natural course of syphilis infection based upon a study of the Boeck-Bruusgaard material. Acta Derm Venereol 1955;35(suppl 34):1–368.
- 12. Fiumara NJ. Syphilis in newborn children. *Clin Obstet Gynecol* 1975;18:183–189.
- 13. Rompalo AM, Lawlor J, Seaman P, Quinn TC, Zenilman JM, Hook EW 3rd. Modification of syphilitic genital ulcer manifestations by coexistent HIV infection. *Sex Transm Dis* 2001;28:448–454.
- 14. Golden MR, Marra CM, Holmes KK. Update on syphilis: resurgence of an old problem. *JAMA* 2003;290:1510–1514.
- 15. Marra CM, Maxwell CL, Smith SL, et al. Cerebrospinal fluid abnormalities in patients with syphilis: association with clinical and laboratory features. *J Infect Dis* 2004;189:369–376.
- 16. Hooshmand H, Escobar MR, Kopf SW. Neurosyphilis. A study of 241 patients. *JAMA* 1972;219:726–729.
- Davis LE, Schmitt JW. Clinical significance of cerebrospinal fluid tests for neurosyphilis. *Ann Neurol* 1990;27:211–212.
- 18. Marra CM, Critchlow CW, Hook EW 3rd, Collier AC, Lukehart SA. Cerebrospinal fluid treponemal antibodies in untreated early syphilis. *Arch Neurol* 1995;52:68–72.
- 19. Smith C, Kamp M, Olansky S, Price EV. Benzathine penicillin G in the treatment of syphilis. *Bull World Health Organ* 1956;15:1087–1096.
- 20. Elliot WC. Treatment of primary syphilis. J Am Vener Dis Assoc 1976;3:128-135.
- 21. Idsoe O, Guthrie T, Wilcox RR. Penicillin in the treatment of syphilis. The experience of three decades. *Bull World Health Organ* 1972;47:1–68.
- Rolfs RT, Joesoef MR, Hendershot EF, et al. A randomized trial of enhanced therapy for early syphilis in patients with and without human immunodeficiency virus infection. The Syphilis and HIV Study Group. N Engl J Med 1997;337: 307–314.
- 23. Harshan V, Jayakumar W. Doxycycline in early syphilis: a long term follow up. *Indian J Dermatol* 1982;27:119–124.
- 24. Onoda Y. Therapeutic effect of oral doxycycline on syphilis. *Br J Vener Dis* 1979;55:110–115.

- Hook EW 3rd, Baker-Zander SA, Moskowitz BL, Lukehart SA, Handfield HH. Ceftriaxone therapy for asymptomatic neurosyphilis. Case report and Western blot analysis of serum and CSF IgG response to therapy. Sex Transm Dis 1986;13(suppl 3):185–188.
- 26. Moorthy TT, Lee CT, Lim KB, Tan T. Ceftriaxone for treatment of primary syphilis in men: a preliminary study. *Sex Transm Dis* 1987;14:116–119.
- 27. Alexander JM, Sheffield JS, Sanchez PJ, Mayfield J, Wendel GD Jr. Efficacy of treatment for syphilis in pregnancy. *Obstet Gynecol* 1999;93:5–8.
- 28. Rolfs RT. Treatment of syphilis, 1993. Clin Infect Dis 1995;20(suppl 1):S23-38.
- 29. Augenbraun MH, Rolfs R. Treatment of syphilis, 1998: nonpregnant adults. *Clin Infect Dis* 1999;29(suppl 1):S21–28.
- 30. Augenbraun M, Workowski K. Ceftriaxone therapy for syphilis: report from emerging infections network. *Clin Infect Dis* 1999;29:1337–1338.
- Walker GJ. Antibiotics for syphilis diagnosed during pregnancy. Cochrane Library 2002:3.
- 32. Dowell ME, Ross PG, Musher DM, Cate TR, Baughn RE. Response of latent syphilis or neurosyphilis to ceftriaxone therapy in persons infected with human immunodeficiency virus. *Am J Med* 1992;93:481–488.
- 33. Marra CM, Boutin P, McArthur JC, et al. A pilot study evaluating ceftriaxone and penicillin G as treatment agents for neurosyphilis in human immunodeficiency virus-infected individuals. *Clin Infect Dis* 2000;30:540–544.
- 34. Chang SN, Chung KY, Lee MG, Lee JB. Seroconversion of the serological tests in the newborns to treated syphilitic mothers. *Genitourin Med* 1995;71:68–70.
- Hook EW 3rd, Stephens J, Ennis DM. Azithromycin compared with penicillin G benzathine for treatment of incubating syphilis. *Ann Intern Med* 1999;131: 434–437.
- 36. Lukehart SA, Godornes C, Molini BJ, et al. Macrolide resistance in *Treponema pallidum* in the United States and Ireland. *N Engl J Med* 2004;351:154–158.
- 37. Klausner JD, Mitchel SJ, Lukehart SA, Gordones C, Engelman J, GISP, CDC. Rapid and large increase in azithromycin resistance in syphilis whilst steady low azithromycin resistance in gonorrhea 2000-2004. Paper presented at: Annual Meeting of the International Society for STD Research; July 2005; Amsterdam, the Netherlands. Abstract TO-203.
- 38. Holmes KK. Azithromycin versus penicillin G benzathine for early syphilis. *N Engl J Med* 2005;353:1291–1293.
- 39. Wendel GD Jr, Stark RJ, Jamison RB, Molina RD, Sullivan TJ. Penicillin allergy and desensitization in serious infections during pregnancy. *N Engl J Med* 1985;312:1229–1232.
- Lukehart SA. Serologic testing after therapy for syphilis; is there a test for cure? Ann Intern Med 1991;114:1057–1058.
- Romanowski B, Sutherland R, Fick GH, Mooney D, Love EJ. Serologic response to treatment of infectious syphilis. *Ann Intern Med* 1991;114: 1005–1009.
- 42. Dattner B, Thomas EW, De Mello L. Criteria for the management of neurosyphilis. *Am J Med* 1951;10:463–467.

- 43. Flores JL. Syphilis. A tale of twisted treponemes. *West J Med* 1995;163: 552–559.
- Marra CM, Longstreith WT Jr, Maxwell CL, Lukehart SA. Resolution of serum and cerebrospinal fluid abnormalities after treatment of neurosyphilis. Influence of concomitant human immunodeficiency virus infection. Sex Transm Dis 1996;23:184–189.
- 45. Jordan KG. Modern neurosyphilis a critical analysis. *West J Med* 1988;149:47–57.
- 46. Genc M, Ledger WJ. Syphilis in pregnancy. Sex Transm Infect 2000;76:73–79.
- 47. McFarlin B, Bottoms SF, Dock BS, Isada NB. Epidemic syphilis: maternal factors associated with congenital infection. *Am J Obstet Gynecol* 1994:170:535–540.
- 48. Mascola L, Pelosi R, Alexander CE. Inadequate treatment of syphilis in pregnancy. *Am J Obstet Gynecol* 1984;150:945–947.
- 49. Conover CS, Rend CA, Miller GB Jr, Schmid GP. Congenital syphilis after treatment of maternal syphilis with a penicillin regimen exceeding CDC guidelines. *Infect Dis Obstet Gynecol* 1998;6:134–137.
- 50. Kiefer L, Rubin A, McCoy JB, Foltz EL. The placental transfer of erythromycin. *Am J Obstet Gynecol* 1955;69:174–177.
- 51. Philipson A, Sabath LD, Charles D. Transplacental passage of erythromycin and clindamycin. *N Engl J Med* 1973;288:1219–1221.
- 52. Wendel GD Jr, Sheffield JS, Hollier LM, Hill JB, Ramsey PS, Sanchez PJ. Treatment of syphilis in pregnancy and prevention of congenital syphilis. *Clin Infect Dis* 2002;35(suppl 2):S200–209.
- 53. Sanchez PJ, Wendel GD. Syphilis in pregnancy. Clin Perinatol 1997;24:71–90.
- 54. Brown ST. Adverse reactions in syphilis therapy. *J Am Vener Dis Assoc* 1976;3:172–176.

SPECIFIC POPULATIONS

IMMIGRANTS AND REFUGEES

Definitions

A *legal immigrant* is a person born outside of Canada who has been granted the right to live in Canada permanently by immigration authorities, whereas an *illegal immigrant* has not been granted such a right. A *refugee* is a person outside his/her country of nationality who is unable or unwilling to return because of persecution on account of race, religion, nationality, membership in a particular social group, or political opinion.¹

Epidemiology

Over 5 million Canadians were born outside of the country, and about 250,000 new immigrants arrive in Canada each year,^{2,3} but data on migrant health in Canada are limited. Recent immigrants underuse health services; it has not been established whether this is associated with cultural barriers, language barriers, reduced perceived needs, reduced needs (this *healthy immigrant effect* refers to recent immigrants being healthier than Canadian-born individuals) or socioeconomic barriers (e.g., lack of access to telephone, transportation to clinic etc).⁴⁻⁷ Many countries of origin have much higher rates of sexually transmitted infections (STIs) than Canada.^{8,9}

Prevention

Health care providers must pay special attention to the complex and stressful process that newcomers may have to undergo to integrate into a new society. Potential loss of social support and cultural identity during the transition may prove challenging. Illegal immigrants present even greater challenges because of the underground nature of their existence. They do not have health insurance and may avoid seeking medical attention for fear of being deported.

Clinical and public health services must be sensitized to the following issues in the provision of affordable, comprehensive and culturally/linguistically appropriate sexual health counselling and STI prevention and management services for the immigrant and refugee population:^{10–12}

- Language, cultural and socioeconomic barriers may prevent access to STI and prevention services.
- This population may experience social isolation from loss of social support.
- Understanding social/sexual mixing patterns, health belief systems, practices and taboos is important in STI prevention, diagnosis, management and partner notification.

- Health care providers must be aware of stigma and discrimination so that individuals from high-prevalence countries are not stigmatized.
- Mental health, including post-traumatic stress disorder, can influence behaviours and interaction with the health system.
- Gender power differential and domestic violence can be barriers to prevention and partner notification.
- Patients may have a history of torture and rape.
- Patients may have a limited knowledge of STIs in Canada and other health resources.
- There may be a travel-related risk of patients either carrying an STI from their place of origin or of acquiring an STI when they return home to visit friends/relatives. This population is less likely to seek pre-travel advice or post-travel care.

Evaluation

Assessment

A non-judgmental and culturally sensitive STI risk assessment should be part of a comprehensive approach to the prevention and early detection of STIs. Issues to explore include the following:

- The presence of opposite-sex and same-sex activity.
- The range and frequency of various sexual practices, taking into account cultural and gender context (see *Primary Care and Sexually Transmitted Infections* chapter).
- The patient's history of STIs, including HIV, with awareness of the stigma and discrimination that come with these infections.
- Injection drug use (IDU).
- Suboptimal screening in pregnant women.

Screening

Based on the results of the risk assessment, conventional STI screening in asymptomatic individuals should be considered for those engaging in high-risk practices (see *Primary Care and Sexually Transmitted Infections* chapter):

- Syphilis testing:
 - Syphilis serology (non-treponemal only) is a standard Citizenship and Immigration Canada (CIC) test requirement for immigrant and refugee applicants 15 years of age or over.
 - Possible false-positive syphilis tests should be kept in mind in individuals from areas of the world where pinta, yaws and bejel are prevalent¹³ (see *Syphilis* chapter).

- HIV serology (unless known to be seropositive):
 - Since 2002, HIV serology has been a standard CIC test requirement for immigrant and refugee applicants 15 years of age or over, and for a child with blood/blood product exposure, born to an HIV-infected mother or being considered as an international adoptee.
 - High-risk individuals who have not had a recent HIV antibody test should be counselled and tested accordingly (see *Human Immunodeficiency Virus Infections* chapter).
 - A child should be offered HIV testing unless there is a reason not to do so, especially if it is likely that the child was born to or breastfed by an undiagnosed HIV-infected mother.

At present, HIV and syphilis are the only mandatory STI tests for immigrant/refugee applicants. Some laboratories abroad may have quality-control issues, and some applicants may pay to obtain negative tests in order to facilitate their application.

For individuals with anogenital symptoms, it is important to keep in mind the following when considering appropriate investigation:

- Chancroid and lymphogranuloma venereum (LGV) are common in parts of Africa, Asia, the Caribbean and Latin America (see *Chancroid* and *Lymphogranuloma Venereum* chapters).
- For assessment of genital ulcers, see Genital Ulcer Disease chapter.
- Quinolone-resistant gonorrhea is particularly prevalent in Asia, the Pacific Islands, Australia, Israel, the United Kingdom, parts of the United States and Canada (see *Gonococcal Infections* chapter).

Hepatitis B and C

Currently, hepatitis B and C testing are not required for the immigration and refugee application process in Canada. However, the prevalence of chronic hepatitis B infection in Asia, Africa, Eastern Europe and Latin America is much higher than in Canada. In asymptomatic individuals from high-prevalence regions, hepatitis B screening should be undertaken for *either* hepatitis B surface antigen carriage and the antibody to hepatitis B surface antigen (for immunity) *or* the antibody to hepatitis B core antigen (for past exposure to the virus). Further hepatitis B testing may be performed depending on the results of the screening tests. Household and sexual contacts of a hepatitis B carrier should be assessed. Those who have not been exposed to hepatitis B or previously immunized should receive a three-dose series of the hepatitis B vaccine (see *Canadian Immunization Guide*¹⁴ and *Hepatitis B Virus Infections* chapter).

The prevalence of chronic hepatitis C infection in Asia, Africa and the Mediterranean is much higher than in Canada.¹⁵ Hepatitis C is primarily transmitted parenterally. Recently, transmission of hepatitis C has been increasingly reported in Europe among men who have sex with men who are not injection drug users, in association with fisting, LGV, HIV and other STIs.^{16–21} As with all patients, hepatitis C should be considered in immigrants and refugees with any of the following risk factors:^{19–32}

- · Any history of IDU.
- Receipt of contaminated blood/blood products in some countries, even after 1990, because of inadequate quality control in the laboratory or inadequate blood screening.
- Procedures (e.g., injection, surgery, transfusion, ceremonial rituals, acupuncture) involving sharing of contaminated equipment in parts of the world with high hepatitis C virus (HCV) prevalence.
- · Exposure to hepatitis C in a prison setting.
- · Needlestick or sharp injuries.
- · Non-sterile tattooing and body piercing.
- · Hemodialysis.
- Sharing personal items contaminated with blood from an HCV-infected individual (e.g., razors, nail clippers, toothbrush).
- Sharing intranasal equipment for snorting drugs.
- · Hepatitis B infection.
- HIV infection.
- · Being a child born to a mother infected with HCV.
- Undiagnosed liver disease.

Sexual transmission is usually inefficient, and the risk of hepatitis C is slightly increased in individuals with the following risk factors:

- A sexual partner with HCV.
- Multiple sexual partners.
- HIV and other STI co-infections.
- Practicing anal intercourse.

Specimen Collection and Laboratory Diagnosis

(See Laboratory Diagnosis of Sexually Transmitted Infections chapter. See individual chapters for specific STIs/syndromes)

- Same as for all patients.
- Please note that specimen collection can be affected by a history of female genital mutilation. The genital structure may appear different, and visualization of the cervix may not be possible with a standard-size speculum. Performing a bimanual pelvic examination may also be difficult, especially if the introitus has been sutured.

In many cultures, screening compliance is poor if swabs are used because of the
invasive nature of specimen collection. Some immigrants and refugees have very
limited opportunity to interface with the health system, especially some patients
who may have cultural sensitivities toward health care providers of the opposite
sex. Urine nucleic acid amplification test screening of high-risk individuals can
enhance compliance and patient comfort.

Management and Treatment

(See individual chapters for specific STIs/syndromes)

- Same as for all patients.
- It is important to address sociocultural and economic factors that may affect treatment adherence. Language barriers may contribute to difficulty following instructions on why and how to take medications, practice safer-sex etc. In some cultures, it may be difficult to discuss monogamy or condom use.
- It is important to obtain a history of traditional/herbal medicine to minimize toxicities and drug interactions.

Reporting and Partner Notification

(See individual chapters for specific STIs/syndromes)

- Same as for all patients.
- It is important to address sociocultural factors that may affect partner
 notification. Language barriers may contribute to difficulty understanding the
 importance of partner notification. In some cultures, fear of domestic violence
 can be an issue with partner notification.

Follow-up

- Same as for all patients.
- Patients who receive their first dose of hepatitis B vaccination should be reminded to return to complete the three-dose immunization series.

References

- 1. Talking about refugees and immigrants: a glossary of terms. Canadian Council for Refugees website. Available at: cpj.ca/refugees/Refugee_Basics/index.html? ap=1&x=81013. Accessed January 25, 2006.
- Canada's ethno-cultural portrait: the changing mosaic. 2001 Census: Analysis Series. Ottawa, ON: Statistics Canada; 2003. Cat. No. 96F0030XIE2001008. Available at: www12.statcan.ca/english/census01/products/analytic/ companion/etoimm/pdf/96F0030XIE2001008.pdf. Accessed January 25, 2006.
- 3. Facts and Figures 2002: Immigration Overview. Ottawa, ON: Citizenship and Immigration Canada; 2003. Cat. No. MP43-333/2003E. Available at: www.cic.gc.ca/english/pdf/pub/facts2002.pdf. Accessed January 25, 2006.
- Hyman I. Immigration and Health. Health Policy Working Paper Series. Ottawa, ON: Health Canada: 2001.

- 5. Kinnon D. *Canadian Research on Immigration and Health: An Overview.* Ottawa, ON: Health Canada; 1999.
- 6. Wen SW, Goel V, Williams JI. Utilization of health care services by immigrants and other ethnic/cultural groups in Ontario. *Ethn Health* 1996;1:99–109.
- 7. Vissandjee B, Desmeules M, Cao Z, Abdool S, Kazanjian A. Integrating ethnicity and migration as determinants of Canadian women's health. *BMC Women's Health* 2004;4(suppl 1):S32.
- 8. Gerbase AC, Rowley JT, Heymann DH, Berkley SF, Piot P. Global prevalence and incidence estimates of selected curable STDs. *Sex Transm Infect* 1998;74(suppl 1):S12–S16.
- 9. Fenton KA, Mercer CH, McManus S, et al. Ethnic variations in sexual behaviour in Great Britain and risk of sexually transmitted infections: a probability survey. *Lancet* 2005:365:1246–1255.
- 10. Holt BY, Effler P, Brady W, et al. Planning STI/HIV prevention among refugees and mobile populations: situation assessment of Sudanese refugees. *Disasters* 2003:27:1–15.
- 11. DuPlessis HM, Cora-Bramble D; American Academy of Pediatrics Committee on Community Health Services. Providing care for immigrant, homeless, and migrant children. *Pediatrics* 2005;115:1095–1100.
- 12. Fowler N. Providing primary health care to immigrants and refugees: the North Hamilton experience. *CMAJ* 1998;159:388–391.
- 13. Ratnam S. The laboratory diagnosis of syphilis. *Can J Inf Dis Med Microbiol* 2005;16:45–51.
- Canadian Immunization Guide. 6th ed. Ottawa, ON: Public Health Agency of Canada: 2002.
- Hepatitis C global prevalence (update). Wkly Epidemiol Rec 1999;74: 425–427
- 16. Kropp RY, Wong T; Canadian LGV Working Group. Emergence of lymphogranuloma venereum in Canada. *CMAJ* 2005;172:1674–1676.
- 17. Nieuwenhuis RF, Ossewaarde JM, Gotz HM, et al. Resurgence of lymphogranuloma venereum in Western Europe: an outbreak of Chlamydia trachomatis serovar I2 proctitis in The Netherlands among men who have sex with men. *Clin Infect Dis* 2004:39:996–1003.
- 18. French P, Ison CA, MacDonald N. Lymphogranuloma venereum in the United Kingdom. *Sex Transm Infect* 2005;81:97–98.
- Gotz HM, van Doornum G, Niesters HG, den Hollander JG, Thio HB, de Zwart O. A cluster of acute hepatitis C virus infection among men who have sex with men — results from contact tracing and public health implications. AIDS 2005;19:969–974.
- 20. Ruys TA, den Hollander JG, Beld MG, van der Ende ME, van der Meer JT. Sexual transmission of hepatitis C in homosexual men. *Ned Tijdschr Geneeskd* 2004:148:2309–2312.
- 21. Ghosn J, Pierre-François S, Thibault V, et al. Acute hepatitis C in HIV-infected men who have sex with men. *HIV Med* 2004;5:303–306.

- 22. NIH Consensus Statement on management of hepatitis C: 2002. *NIH Consens State Sci Statements* 2002;19:1–46.
- 23. Recommendations for prevention and control of hepatitis C virus (HCV) infection and HCV-related chronic diseases. Centers for Disease Control and Prevention. *MMWR Recomm Rep* 1998;47(RR-19):1–39.
- 24. Alter MJ, Seeff LB, Bacon BR, Thomas DL, Rigsby MO, Di Bisceglie AM. Testing for hepatitis C virus infection should be routine for persons at increased risk for infection. *Ann Int Med 2004*;141:715–717.
- 25. Sherman M, Bain V, Villeneuve JP, et al. The management of chronic viral hepatitis: a Canadian consensus conference 2004. *Can J Gastroenterol* 2004;18:715–728.
- 26. Mele A, Spada E, Sagliocca L, et al. Risk of parenterally transmitted hepatitis following exposure to surgery or other invasive procedures: results from the hepatitis surveillance system in Italy. *J Hepatol* 2001;35:284–289.
- 27. Montella M, Crispo A, Grimaldi M, Tridente V, Fusco M. Assessment of iatrogenic transmission of HCV in Southern Italy: was the cause the Salk polio vaccination? *J Med Virol* 2003;70:49–50.
- 28. Frank C, Mohamed MK, Strickland GT, et al. The role of parenteral antischistosomal therapy in the spread of hepatitis C in Egypt. *Lancet* 2000;355:887–891.
- 29. Singh S, Kumar J, Singh R, Dwivedi SN. Hepatitis B and C viral infections in Indian kala-azar patients receiving injectable anti-leishmanial drugs: a community-based study. *Int J Infect Dis* 2000;4:203–208.
- 30. Chlabicz S, Grzeszczuk A, Prokopowicz D. Medical procedures and the risk of iatrogenic hepatitis C infection: case-controlled study in north-eastern Poland. *J Hosp Infect* 2004;58:204–209.
- 31. Alter MJ, Kruszon-Moran D, Nainan OV, et al. The prevalence of hepatitis C virus infection in the United States, 1988 through 1994. *N Engl J Med* 1999;341:556–562.
- 32. Sharma AK, Aggarwal OP, Dubey KK. Sexual behavior of drug-users: is it different? *Prev Med* 2002;34:512–515.

INMATES AND OFFENDERS

Background

The responsibility for corrections in Canada is shared by the federal, provincial and territorial governments.¹ In 2001, the average count of adult offenders incarcerated in federal, provincial and territorial facilities was 32,073 (133/100,000). A further 122,870 adult offenders were under community supervision, including probation, conditional sentence and conditional release.² Statistics on juvenile corrections are not routinely collected at the national level,¹ but in 1994–95, 1,095 female young offenders were sentenced to secure custody, 1,795 were placed in open custody and another 6,952 were placed on probation;³ in 1996–97, 10,396 male young offenders were sentenced to secure custody, 11,541 were placed in open custody and 28,395 were placed on probation.⁴

Although Aboriginal people constitute about 3% of the general population in Canada, they represent approximately 15% of the federal offender population.¹ Women made up approximately 3% of the total incarcerated population in Correctional Service Canada (CSC) facilities in 2000–2001.⁵ Canada's incarceration rate is higher than that of many Western European countries, but is much lower than that of the United States.¹

Epidemiology

Inmates in correctional facilities around the world bear a disproportionate burden of illness related to infectious disease compared to the general population. As a result, rates of sexually transmitted infections (STIs), hepatitis B virus (HBV), hepatitis C virus (HCV) and HIV/AIDS are significantly higher among prison inmates. Often, inmates belong to vulnerable populations in which high-risk behaviours for STI infection are present, such as injection drug use (IDU) and unprotected sexual intercourse.⁵ Although many inmates enter correctional facilities already infected, any inmate who engages in risky behaviours in prison is at risk of becoming infected or reinfected with an STI.⁵ As of 2002, CSC estimated that 70% of the inmates who entered prison had self-identified drug- or alcohol-abuse problems.⁶ Although penetrative sexual activity is known to occur in correctional settings,⁷ it is likely underreported because it is often prohibited and may carry stigma.⁶ Nonconsensual sexual activity may also be an issue.⁸ Other practices within the prison setting, such as IDU, tattooing and/or piercing, may contribute to the transmission of infectious diseases as well.⁶

In January 2000, CSC, in collaboration with Health Canada (now the Public Health Agency of Canada), introduced a comprehensive surveillance system to provide more accurate and extensive information about infectious diseases within federal correctional settings — the CSC Infectious Diseases Surveillance System (CSC-IDSS).⁵ The CSC-IDSS is based on aggregate data on testing and test results for blood-borne and sexually transmitted pathogens, and it allows CSC to monitor trends in prevalence among newly admitted and general-population inmates.

According to CSC, a revised system, which includes line-listed risk behaviour and test-outcome data, is currently being implemented to better target harm-reduction programs.

Reported rates of infection in Canadian penitentiaries to 2002 are as follows:5,9,10

- HIV:* The prevalence of HIV among offenders in federal facilities increased steadily between 2000 and 2002, from 1.7% to 2.0%. In 2002, the rate was higher in women (3.7%) than in men (1.9%).
- HCV:* The prevalence of HCV among offenders in federal facilities increased from 2,542 cases (19.7%) in 2000 to 3,173 (25.4%) in 2002. In 2002, the rate was higher in women (33.7%) than in men (25.2%), but between 2000 and 2002, rates decreased for women and increased for men.
- HBV:† HBV prevalence among federal inmates in 2002 was 0.2%. There was a sharp increase from 2000 (0.1%) to 2001 (0.3%), but rates fell in 2002. Most cases identified were in men.
- Chlamydia:[†] There were 53 cases reported in 2002 (0.32% prevalence). The rate is increasing compared to 2000–2001. Over 90% of cases have been diagnosed in men.
- Gonorrhea:[†] There were 20 cases reported in 2002 (0.12% prevalence). The
 rate has increased compared to 2000–2001. About 85% of cases have been
 diagnosed in men.
- Syphilis:[†] There were three cases reported in 2002. The rate has increased compared to 2000–2001.

Notes:

- * Testing uptake levels for HIV and HCV indicate that up to 70% of inmates may remain unscreened for these infections. As a result, reported rates may severely underestimate the true burden of disease within federal correctional facilities.
- Lack of reporting and underdiagnosis of HBV and STIs (including lack of routine screening for STIs) are likely to result in an underestimate of the actual rates of these infections in inmates.

Prevention

Correctional facilities in Canada are recognized as an important focus for public health measures to control STIs, HBV, HCV, HIV/AIDS and other infectious diseases. By its very nature, incarceration may offer one of the best opportunities to access high-risk individuals and provide them with the preventive services, treatment and skills necessary to stay healthy. Interventions are limited by the length of incarceration, but even brief counselling encounters can have a significant impact on risky behaviours. Since most inmates eventually return to the community, harm-reduction efforts within the correctional system can have favourable implications, not only for the inmate population but also for the wider community. For this reason, it is important to coordinate prevention activities with local public health officials and other community-based care groups. Discharge planning for

infected inmates is also an important step in order to optimize the continuation of care for offenders outside the correctional setting.^{5,11}

Components to be considered for STI prevention programs within correctional facilities are similar to those in the community: ¹² education; voluntary testing and counselling; distribution of clean needles or bleach; distribution of condoms; and drug-dependence treatment (including substitution treatment) have all been proven effective in reducing HIV/STI risk in prisons and have been shown to have no unintended negative consequences. ¹² The appropriate care, treatment and support of inmates with STIs helps prevent transmission of these infections. This includes partner notification, as well as testing and treatment of recent sexual contacts.

It is important to include the issues of alcohol and drug use in educational efforts, acknowledging their contribution to heightened risk for STIs and other infections. Harm-reduction education to minimize the negative consequences of risky behaviours and provide alternatives can impact favourably on the transmission of STIs and other infections. CSC currently has a number of health-education and peer-counselling support programs to disseminate information and encourage behaviour modification.

As part of the Canadian Strategy on HIV/AIDS and in partnership with the Public Health Agency of Canada, CSC has implemented several initiatives aimed at preventing and controlling the transmission of infectious diseases (including STIs, HIV, HBV and HCV) within federal correctional facilities.¹⁴ These include confidential, voluntary testing for inmates on admission and throughout their incarceration, as well as pre- and post-test counselling.14 Serological testing and immunization for hepatitis A virus (HAV) and HBV are offered. Educational programs and materials are provided for offenders and staff.14 In 1992, condoms, dental dams and waterbased lubricants were introduced into federal penitentiaries. 5 CSC has also initiated a national drug strategy aimed at controlling the supply of drugs in federal institutions. Its goal is to reduce the demand for drugs among federal offenders by implementing prevention and treatment programs, 6 such as methadone maintenance treatment and substance-use programs.¹⁴ CSC currently provides bleach kits to inmates for cleaning needles and most recently has instituted a pilot project of tattoo parlours in six federal prisons. Currently, CSC does not provide needle-exchange services to inmates, citing its zero-tolerance policy toward drug use and trafficking in prison, as well as concerns about the health of inmates and the security of the institution. Discussions between CSC and the Public Health Agency of Canada about possible collaborative projects in federal correctional facilities are underway.

Evaluation

Health care professionals may be reluctant to ask and offenders may be reluctant to disclose information about their health, especially when issues such as sexual activity, substance use and possible illegal activities are involved. It is essential that the confidential nature of the interaction be stressed so that a true understanding of a patient's risk for STIs and other infections can be gained.

History

A complete sexual history should be taken (see *Primary Care and Sexually Transmitted Infections* chapter).

It is important to be aware that self-identified sexual identity is not an accurate predictor of sexual behaviour.⁸ Although some inmates may consider themselves heterosexual, they may have been involved in sexual activity with members of the same sex (either prior to and/or during incarceration). Therefore, it is essential that a basic sexual history include questions about opposite-sex and same-sex activity. This can be achieved by asking open-ended questions such as: "Do you have sex with men, women or both?" For a more complete discussion of this topic, see *Men Who Have Sex with Men/Women Who Have Sex with Women* chapter.

Patients engaging in practices (both sexual and non-sexual) that are associated with an increased risk for STIs need to be identified. Such practices include the following:

- Receptive and insertive anogenital intercourse.
- Oral-anal intercourse (anilingus/rimming).
- Unprotected sexual activity (oral, anal or genital).
- Sharing of sex toys.
- Receptive manual-anal intercourse (insertion of finger or fist in anus of partner).
- · Substance use accompanying sex.
- · Tattooing.
- IDU and other substance use.

Because of the high prevalence of substance use in correctional settings, a substance-use history should also be taken (see *Substance Use* chapter).

Screening

Voluntary testing offered to new admissions to the correctional system may be one of the best opportunities for screening and identifying prevalent infections among offenders. Non-invasive tests such as nucleic acid amplification tests (NAATs) (e.g., ligase chain reaction [LCR], polymerase chain reaction [PCR]) of urine has made STI screening in correctional facilities more available and acceptable, but anecdotal reports suggest that this has not translated into higher rates of uptake. One possible explanation may be inmates' reluctance to submit urine, as urine

testing is typically associated with testing for drug use, something inmates may be anxious to hide. A detailed explanation of the testing procedure may help to overcome this hurdle.

Whether for a new admission or not, greater use of routine testing for inmates at risk is needed,⁵ especially given the often asymptomatic nature of STIs. This highlights the importance of the sexual history for identifying those at risk (see *Primary Care and Sexually Transmitted Infections* chapter). Collaborations involving corrections, public health officials and evaluators have been shown to facilitate increased STI screening in inmates.¹⁵

Pre- and post-test counselling for positive and negative results is essential and should reflect the primary public health purposes of counselling and testing: to help non-infected individuals initiate and sustain behaviour changes to reduce their risk of infection and to help infected individuals seek health care and avoid infecting others.⁸

Based on the results of the history/risk assessment, the following screening should be considered for inmates and offenders:

- Routine STI screening at all potential sites of infection: chlamydia, gonorrhea, syphilis, HIV and HBV (if not already immunized or known to be immune).
- Testing for herpes simplex virus, if symptoms are present (see Genital Herpes Simplex Virus Infections chapter).
- HCV serology: IDU, tattooing and high-risk sexual practices are known risk behaviours associated with the transmission of HCV in the prison/inmate population.⁶
- For those with identified risk, screen for HAV immunity prior to vaccination.

For specific screening considerations in men who have sex with men and women who have sex with women, see *Men Who Have Sex with Men/Women Who Have Sex with Women* chapter.

Cervical screening for dysplasia and/or human papillomavirus (HPV) infection in female inmates should be considered if there is no evidence of screening with a normal result within the previous year (see *Genital Human Papillomavirus Infections* chapter).

Specimen Collection and Laboratory Diagnosis

In the correctional setting, the rapid turnover and transfer of offenders within institutions, especially at reception and among temporary detainees (those returning from parole), may limit the time available to diagnose and treat an STI. For this reason, rapid point-of-care testing may be especially relevant.

Urine-based testing is generally more acceptable than more invasive urethral or cervical swabs, but its association with drug testing may make inmates reluctant to provide a sample. A discussion of exactly what the urine is being tested for may facilitate acceptance by inmates.

Management and Treatment

In the correctional setting, the rapid turnover and transfer of offenders within institutions, especially at reception and among temporary detainees (those returning from parole), may limit the use of longer-term treatment regimens. In these cases, single-dose therapies for the treatment of STIs may be more appropriate.

Reporting and Partner Notification

According to a CSC infection control directive, CSC physicians or the Chief of Health Services on behalf of a physician must ensure that all diagnosed provincially reportable communicable diseases are reported to the local public health unit or the appropriate public health office.

Partner notification is a major component of STI follow-up. However, inmates who test positive for an STI may be reluctant to disclose information about contacts or behaviours that may be deemed inappropriate, illegal or stigmatized. It is critical to ensure that the partner-notification process is voluntary and non-coercive, preserving confidentiality and trust and respecting the dignity and human rights of the individual.⁸

Follow-up

Inmates who continue to engage in risky behaviour should be encouraged to be screened regularly for STIs. Safer-sex and harm-reduction education and counselling should continue to be emphasized.

If HAV and HBV vaccination have been initiated, the vaccination schedule must be completed as recommended.

As for all women, female inmates should have regular cervical screening for dysplasia and/or HPV infection as appropriate.

It is important that correctional services work in concert with local public health officials to follow up when appropriate with offenders who have been released to the community (i.e., referral/reporting to public health of unmanaged cases and contacts released to or residing in the community).

References

- Basic Facts about Federal Corrections. Ottawa, ON: Correctional Service Canada: 2001.
- Adult correctional services, average counts of offenders in provincial, territorial and federal programs. CANSIM tables 251-0004 and 251-0007. Ottawa, ON: Statistics Canada, 2002.
- Female Young Offenders in Canada: Recent Trends. Ottawa, ON: Correctional Service Canada; 1997. Available at: www.csc-scc.gc.ca/text/rsrch/briefs/b18/ b18e_e.shtml. Accessed March 28, 2005.
- Male Young Offenders in Canada: Recent Trends. Ottawa, ON: Correctional Service Canada; 1998. Available at: www.csc-scc.gc.ca/text/rsrch/briefs/b22/ b22e e.shtml. Accessed March 28, 2005.
- Infectious Diseases Prevention and Control in Canadian Federal Penitentiaries, 2001–01. Ottawa, ON: Correctional Service Canada; 2003. Available at: www. csc-scc.gc.ca/text/pblct/infectiousdiseases/en.pdf. Accessed March 28, 2005.
- 6. Skoretz S, Zaniewski G, Goedhuis NJ. Hepatitis C virus transmission in the prison/inmate population. *Can Commun Dis Rep* 2004;30:141–148.
- 7. *Guidelines on HIV Infection and AIDS in Prisons*. Geneva, Switzerland: World Health Organization; 1993.
- 8. HIV in Prisons. Geneva, Switzerland: World Health Organization; 2001.
- 9. De P, Connor N, Bouchard F, Sutherland D. HIV and hepatitis C virus testing and seropositivity rates in Canadian federal penitentiaries: a critical opportunity for care and prevention. *Can J Infect Dis Med Microbiol* 2004;15:221–225.
- 10. Correctional Service Canada, unpublished data, 2004.
- 11. Grinstead O, Seal DW, Wolitski R, et al. HIV and STD testing in prisons: perspectives of in-prison service providers. *AIDS Educ Prev* 2003;15:547–560.
- 12. WHO, UNAIDS, United Nations Office on Drugs and Crime. *Policy Brief: Reduction of HIV Transmission in Prisons.* Geneva, Switzerland; World Health Organization: 2004.
- 13. MacGowan RJ, Margolis A, Gaiter J, et al. Predictors of risky sex of young men after release from prison. *Int J STD AIDS* 2003;14:519–523.
- 14. Specific Guidelines for Methadone Maintenance Treatment. Ottawa, ON: Correctional Service Canada; 2003. Available at: www.csc-scc.gc.ca/text/pblct/methadone/english/meth_guidelines_e.pdf. Accessed March 28, 2005.
- 15. Jacob-Arriola KR, Braithwaite RL, Kennedy S, et al. A collaborative effort to enhance HIV/STI screening in five county jails. *Public Health Rep* 2001;116:520–529.

MEN WHO HAVE SEX WITH MEN (MSM)/WOMEN WHO HAVE SEX WITH WOMEN (WSW)

Definition

Men who have sex with men (MSM) may have sex with men exclusively, or with both men and women, and may self-identify as gay, bisexual or heterosexual.

Women who have sex with women (WSW) may have sex with women exclusively, or with both women and men, and may self-identify as gay, lesbian, bisexual or heterosexual.

Epidemiology

Following a decline in the prevalence of reportable sexually transmitted infections (STIs) among MSM beginning in the 1980s, the incidence of syphilis, gonorrhea, chlamydia, genital herpes, hepatitis A virus (HAV), hepatitis B virus (HBV) and HIV infections has risen among MSM in Canada and internationally since the mid-1990s.^{1–12} Recent outbreaks of syphilis among MSM have been reported, ^{2,3,13,14} with a large proportion of cases co-infected with HIV. Similarly, recent outbreaks of lymphogranuloma venereum (LGV) have been reported internationally ^{15–20} and in Canada²¹ among MSM, with a high degree of HIV co-infection. Co-infection is of particular concern, given that syphilis and other STIs can increase the likelihood of HIV transmission and acquisition. ^{22–25}

Rising rates of STIs among MSM are associated with increases in unsafe sexual practices, ²⁶ including unprotected anal intercourse (otherwise known as barebacking), ^{12,27-31} an increase in the number of sex partners; ^{1,12} partner-finding on the Internet; ³²⁻³⁷ other anonymous partnering venues (e.g., bathhouses); ^{1,38} recreational and non-recreational drug use; ^{1,27,39-43} and unprotected oral sex. ¹ Rates of unprotected anal intercourse have increased among MSM of all ages, and between HIV serodiscordant partners. ^{28,31,44}

Many explanations have been proposed for the recent increase in risky sexual practices among MSM, including fatigue with safer-sex messages and reduced fear of acquiring HIV due to optimism about new HIV treatments, 45,46 although the correlation with treatment optimism has not been shown consistently. 47 The increase in unsafe sexual practices among HIV-infected MSM has been attributed in part to the increasing proportion of HIV-infected MSM who feel healthy, are living longer and are therefore having sex more often and with more partners. Lack of knowledge of their own and their partners' STI status, including HIV, is also a concern; for example, almost 27% of HIV-positive men in the *Ontario Men's Survey* were unaware of their HIV status. 26

Common recreational drugs used at bathhouses, raves or circuit parties include alcohol, methamphetamine ("crystal meth"), methylenedioxymethamphetamine (MDMA or "ecstasy"), ketamine ("special K"), gamma hydroxybutyrate (GHB), volatile nitrites (poppers) and cocaine (see *Substance Use* chapter). The reduction in inhibition that accompanies the use of these drugs can increase the likelihood of multiple sex partners and unprotected sex, and may be partnered with the use of sildenafil citrate (Viagra), vardenafil (Levitra) or tadalafil (Cialis) to counteract the erectile-dysfunction side effect of some of them. The use of sildenafil among MSM has been linked to an increased risk for multiple sex partners and STI acquisition. 48,49

Sexually transmitted epidemics of enteric infections such as *Salmonella enterica* serotype *typhi* (typhoid fever)⁵⁰ and *Campylobacter jejuni* subsp. *jejuni*,⁵¹ as well as sexual transmission of human herpes virus^{8,52} have been documented among MSM populations in Canada and the United States.

There are very few data on rates of STIs among WSW, although studies have consistently found higher rates of STIs — specifically human papillomavirus (HPV), genital warts, HIV, syphilis and genital ulcer disease — among heterosexual and bisexual women than among women who have sex with women exclusively.^{53–55} Although STI transmission among WSW is strongly correlated with sexual contact with male partners, sexual transmission of HIV, syphilis, HPV, herpes simplex virus types 1 and 2 (HSV-1 and -2), *Trichomonas vaginalis, Chlamydia trachomatis* and HAV have been reported in WSW with no history of a male partner.^{56–61} Higher rates of bacterial vaginosis and hepatitis C virus (HCV) have been reported for WSW than for women with male sex partners only.^{55,62,63} The few studies exploring STI risk behaviours among WSW have demonstrated higher rates of sexual contact with homosexual/bisexual men;^{55,64,65} sex with HIV-infected partners;⁶⁴ injection drug use;^{54,55,64,66} sex for money or drugs;^{54,64,66} and a greater number of recent partners⁶⁴ among WSW compared to exclusively heterosexual women.

Prevention

Prevention counselling with MSM and WSW, as with all sexually active populations, should emphasize personal risk and risk behaviours, as well as the initiation and maintenance of risk-reduction activities with a patient-centred focus. It is important for health care providers to avoid making assumptions about involvement in risky behaviours, including drug use, based on sexual orientation. It is also important that health professionals accurately communicate the risks associated with various sex acts to their sexually active patient, including the risk of transmission via oral sex (i.e., although the risk of STI transmission is lower via oral sex than vaginal or anal sex, many STIs, including syphilis, chlamydia, gonorrhea, herpes and HIV, can be transmitted through unprotected oral sex).

Risk-reduction strategies to include in discussions with MSM and WSW, and all sexually active patients, include the following (see *Primary Care and Sexually Transmitted Infections* and *Human Immunodeficiency Virus Infections* chapters for more information on safer-sex and HIV-specific counselling):

- Avoiding or minimizing unprotected anal, vaginal, oral and anal-oral intercourse; in addition to intercourse, minimize other sexual activities involving exchange of bodily fluids (i.e., sharing of sex toys), which also carry risk for STI transmission.
- Ensuring consistent and correct use of condoms for vaginal intercourse and both insertive and receptive anal intercourse.
- Ensuring use of barrier protection for oral sex.
- Avoiding or minimizing sexual encounters with multiple and anonymous partners, as well as the use of recreational drugs in conjunction with sex.
- Regular testing for STIs if engaging in unprotected or risky sexual activity.
- Negotiating safety in sexual encounters, including disclosure of STI status to
 partners and learning partners' STI status; it should be noted, however, that
 serostatus disclosures may or may not be accurate, and that safer-sex practices
 (i.e., condom use or non-penetrative acts) provide the best protection against
 STIs in a sexual encounter.
- Avoiding the use of products containing nonoxynol-9 (N-9) during intercourse, given the safety and efficacy concerns regarding its use (see *Primary Care and Sexually Transmitted Infections* chapter for more information on N-9). N-9 found on spermicidally lubricated condoms may provide added protection against pregnancy, but it does not effectively protect against infection with HIV or other STIs and may irritate the genital mucosal lining, facilitating their transmission; however, a condom lubricated with N-9 is better than no condom at all.
- Receiving vaccination for both HBV and HAV; this should be offered to all MSM, given their increased risk of infection^{67,68} and poor vaccination coverage;⁶⁹ the first dose can be given while waiting for serological test results (if performed), as immunization is not harmful for previously vaccinated or infected persons (see *Hepatitis B Virus Infections* chapter for more information on HBV vaccination and preimmunization screening).*
- For WSW, undergoing regular cervical screening for dysplasia and/or HPV infection.

Note:

Preimmunization testing for immunity against HAV should be considered for populations with the potential for higher levels of pre-existing immunity (i.e., older Canadians and people from HAV-endemic areas). Routine preimmunization serologic screening for HBsAg, anti-HBs or anti-HBc is recommended for people at high risk of having been infected, but is not practical for universal immunization programs.⁷⁰

Recognizing that MSM and WSW are diverse populations and that reasons for unsafe sexual practices will vary across individuals and subcultures, prevention messages should be tailored to the individual in question and should allow for discussion of realistic safer-sex goals. To be most effective, safer-sex messaging should not be a discussion of sexual risk alone, but one that takes into account the broader context of sexual health influences, including intimacy; sexuality and arousal; drugs and alcohol; mental health, including self-esteem and self-worth; abuse and coercion; and sexual identity.^{71,72} Using a motivational interviewing approach for prevention counselling can be effective in promoting harm-reduction behaviours (see *Primary Care and Sexually Transmitted Infections* chapter for more information on motivational interviewing).

Evaluation

Prior experiences of MSM and WSW with discrimination, homophobia and heterosexism may have an effect on health care–seeking behaviour and disclosure of sexual behaviour in consultations.^{73,74} In every patient encounter, it is important to avoid the assumption of heterosexuality. Taking a basic sexual history for all sexually active patients is important for establishing the following:

- · Presence of opposite-sex and same-sex activity.
- Range and frequency of sexual practices.
- · Level of risk for specific STIs.

Self-identified sexual identity is not an accurate predictor of behaviour;⁷⁵ it is necessary to ask specific questions about the gender of sexual partners when taking a sexual history. Using gender-neutral terms such as "partner" can help to create an environment that is comfortable for disclosure.⁷³ The best approach to obtaining a sexual history is to begin with open-ended, non-judgmental questions regarding broad categories of sexual behaviour and progressing to specific sexual practices.

Asking, "Do you have sex with men, with women, or with both?" may be a useful question during the sexual history to assess gender of sexual partners (see *Primary Care and Sexually Transmitted Infections* chapter for more information on taking a sexual history).

Specific sexual practices that are associated with increased risk of STIs and should be assessed for with all sexually active patients include the following:

- · Receptive (passive) and insertive (active) anogenital intercourse.
- Oral-anal intercourse (anilingus).
- Unprotected sexual activity (oral, anal or genital).
- Sharing of sex toys.
- · Rectal douching in association with receptive anogenital intercourse.
- Receptive manual-anal intercourse (insertion of finger or fist in anus of partner).
- Anonymous partnering and use of anonymous partnering venues (e.g., bathhouses, Internet, raves, circuit parties).

- Substance use accompanying sex.
- Injection drug use (IDU) and other substance use.

Based on results from the risk assessment, the following screening should be considered for men who have had unprotected sex with another man in the preceding year:

- Routine STI screening at all potential sites of infection (chlamydia, gonorrhea, syphilis), HIV serology (unless known to be seropositive) and HBV and HAV serology (if not previously immunized or known to be immune) (see *Hepatitis B Virus Infections* chapter for more information on HBV screening).
- Although asymptomatic screening for HSV and HPV is not currently recommended, new information may alter these recommendations. Studies are ongoing assessing whether screening in certain situations is cost-beneficial.

Assessment for STI symptoms including dysuria, anorectal symptoms (e.g., pain, discharge, bleeding, pruritus), urethral discharge, genital ulcers or lesions, and skin rash should be completed and appropriate diagnostic testing conducted if symptoms are present. In addition to a careful genital and targeted extragenital examination, a physical examination for MSM may include the following (see *Primary Care and Sexually Transmitted Infections* chapter for more information on physical examination):

- Examination of lymph nodes, skin, sclera, oral cavity, pharynx and perianal region.
- Anoscopy or proctoscopy for symptomatic MSM who are the receptive partner for anogenital sex.

Misconceptions about the STI risk and sexual practices of WSW may negatively impact the sexual history and screening performed for this group of women. STI-screening recommendations for WSW should be based on a detailed risk assessment, not on assumptions of low-risk sexual behaviours (see *Primary Care and Sexually Transmitted Infections* chapter). WSW, including those with no history of a male sexual partner, are at risk for cervical abnormalities^{55,58} and should be encouraged to receive regular cervical screening for dysplasia and/or HPV infection.

Specimen Collection and Laboratory Diagnosis

As for all patients, while the choice of STI screening tests is based on the results of the sexual history (as described above), the choice of STI diagnostic tests should be based on the differential diagnosis of the presenting syndrome (e.g., proctitis). The following recommendations apply (see *Laboratory Diagnosis of Sexually Transmitted Infections* chapter for specific information on specimen collection):

- Anorectal gonorrhea and chlamydia cultures, if engaging in unprotected anal intercourse.
- Pharyngeal gonorrhea cultures, if performing unprotected oral sex.
- Laboratory testing for pathogens not usually associated with STIs (i.e., HAV, enteric organisms) but that can cause sexually transmitted proctitis, proctocolitis and enteritis may be indicated based on risk assessment and symptoms (e.g., examination of stool for ova and parasites).

Note:

Although culture remains the recommended test method for assessing pharyngeal or rectal infections, limited studies suggest a potential role for nucleic acid amplification tests for detection of pharyngeal gonorrhea⁷⁶ and rectal chlamydia;⁷⁷ further studies are needed before recommendations can be made.

Management and Treatment

- · Same as for all patients.
- It is important to be aware of the potential stress associated with the "coming out" process and to be knowledgeable about gay- and lesbian-specific support groups and community networks for referral as needed.

Reporting and Partner Notification

- Same as for all patients.
- Anonymous partnering presents a challenge for partner notification, making it difficult, if not impossible, to contact and treat partners who have been exposed to an STI.

Follow-up

- WSW should be encouraged to undergo regular cervical screening for dysplasia and/or HPV infection.
- Patients whose history reveals unsafe sexual behaviours should be encouraged
 to engage in safer-sex and harm-reduction behaviours, and to be screened
 frequently for STIs (at least yearly) (see Primary Care and Sexually Transmitted
 Infections chapter).
- Patients who receive their first dose of HBV or HAV vaccination should be reminded to return to complete the vaccination series (one additional dose for HAV vaccine and two additional doses for HBV vaccine).

References

- 1. Bellis MA, Cook P, Clark P, Syed Q, Hoskins A. Re-emerging syphilis in gay men: a case-control study of behavioural risk factors and HIV status. *J Epidemiol Community Health* 2002;56:235–235.
- Centers for Disease Control and Prevention (CDC). Primary and secondary syphilis among men who have sex with men — New York City, 2001. MMWR Morb Mortal Wkly Rep 2002;51:853–856.
- Centers for Disease Control and Prevention (CDC). Resurgent bacterial sexually transmitted disease among men who have sex with men — King County, Washington, 1997–1999. MMWR Morb Mortal Wkly Rep 1999; 48:773–777.
- 4. Yamey G. San Francisco's HIV infection rate doubles. *BMJ* 2001;322:260.
- 5. Hogg RS, Weber AE, Chan K, et al. Increasing incidence of HIV infections among young gay and bisexual men in Vancouver. *AIDS* 2001;15:1321–1322.
- Fox KK, del Rio C, Holmes KK, et al. Gonorrhea in the HIV era: a reversal in trends among men who have sex with men. Am J Public Health 2001; 91:959–964.
- Berglund T, Fredlund H, Giesecke J. Epidemiology of the reemergence of gonorrhea in Sweden. Sex Transm Dis 2001;28:111–114.
- Catania JA, Osmond D, Stall RD, et al. The continuing HIV epidemic among men who have sex with men. Am J Public Health 2001;91:907–914.
- 9. Ciemins EL, Flood J, Kent CK, et al. Reexamining the prevalence of Chlamydia trachomatis infection among gay men with urethritis: implications for STD policy and HIV prevention activities. *Sex Transm Dis* 2000;27:249–251.
- Geisler WM, Whittington WL, Suchland SJ, Stamm WE. Epidemiology of anorectal chlamydial and gonococcal infections among men having sex with men in Seattle: utilizing serovar and auxotype strain typing. Sex Transm Dis 2002;29:189–195.
- Calzavara L, Burchell AN, Major C, et al; Polaris Study Team. Increases in HIV incidence among men who have sex with men undergoing repeat diagnostic HIV testing in Ontario, Canada. AIDS 2002;16:1655–1661.
- 12. Chen SY, Gibson S, Katz MH, et al. Continuing increases in sexual risk behavior and sexually transmitted diseases among men who have sex with men: San Francisco, Calif., 1999–2001. *Am J Public Health* 2002;92:1387–1388.
- 13. Centers for Disease Control and Prevention (CDC). Outbreak of syphilis among men who have sex with men Southern California, 2000. *MMWR Morb Mortal Wkly Rep* 2001;50:117–120.
- Centers for Disease Control and Prevention (CDC). Primary and secondary syphilis — United States, 2002. MMWR Morb Mortal Wkly Rep 2003;52: 1117–1120.
- 15. Nieuwenhuis RF, Ossewaarde JM, Götz HM, et al. Resurgence of lymphogranuloma venereum in Western Europe: an outbreak of Chlamydia trachomatis Serovar I2 proctitis in The Netherlands among men who have sex with men. *Clin Infect Dis 2004*;39:996–1003.

- Lymphogranuloma venereum among men who have sex with men— Netherlands, 2003–2004. MMWR Morb Mortal Wkly Rep 2004;53:985–988.
- 17. Health Protection Agency. Enhanced surveillance of lymphogranuloma venereum (LGV) in England. CDR Weekly 2004;14:3. Available at: www.hpa.org.uk/cdr/archives/2004/cdr4104.pdf. Accessed February 14, 2006.
- 18. Lymphogranuloma venereum USA (California). ProMED-mail. December 22, 2004. Archive number:20041222.3376. Available at: www.promedmail.org. Accessed February 14, 2006.
- 19. Lymphogranuloma venereum USA (Texas). ProMED-mail. December 24, 2004. Archive number: 20041224.3397. Available at: www.promedmail.org. Accessed February 14, 2006.
- 20. Institut de veille sanitaire. Emergence de la Lymphogranulomatose vénérienne rectale en France : cas estimés au 31 mars 2004. Synthèse réalisée le 1^{er} juin 2004. Available at: www.invs.sante.fr/presse/2004/le_point_sur/lgv_160604/. Accessed February 14, 2006.
- Kropp RY, Wong T; Canadian LGV Working Group. Emergence of lymphogranuloma venereum in Canada. CMAJ 2005;172:1674–1676.
- Centers for Disease Control and Prevention (CDC). HIV prevention through early detection and treatment of sexually transmitted diseases — United States. MMWR Recomm Rep 1998;47(RR-12):1–24.
- 23. Renzi C, Douglas JM Jr, Foster M, et al. Herpes simplex virus type 2 infection as a risk factor for human immunodeficiency virus acquisition in men who have sex with men. *J Infect Dis* 2003;187:19–25.
- Rottingen JA, Cameron DW, Garnett GP. A systematic review of the epidemiologic interactions between classic sexually transmitted diseases and HIV: how much really is known? Sex Transm Dis 2001;28:579–597.
- Fleming DT, Wasserheit JN. From epidemiological synergy to public health policy and practice: the contribution of other sexually transmitted diseases to sexual transmission of HIV infection. Sex Transm Infect 1999;75:3–17.
- 26. Myers T, Allman D, Calzavara L, et al. Ontario Men's Survey Final Report. Available at: www.mens-survey.ca/doc/OMS_Report_web_final%20.pdf. Accessed February 24, 2005.
- 27. Koblin BA, Chesney MA, Husnik MJ, et al. High-risk behaviors among men who have sex with men in 6 US cities: baseline data from the EXPLORE study. *Am J Public Health* 2003;93:926–932.
- 28. Ekstrand ML, Stall RD, Paul JP, Osmond DH, Coates TJ. Gay men report high rates of unprotected anal sex with partners of unknown or discordant HIV status. *AIDS* 1999;13:1525–1533.
- Dufour A, Alary M, Otis J, et al. Risk behaviours and HIV infection among men who have sex with men: baseline characteristics of participants in the Omega Cohort Study, Montreal, Quebec, Canada. Can J Public Health 2000;91: 345–349.
- Halkitis PN, Parsons JT. Intentional unsafe sex (barebacking) among HIV-positive gay men who seek sexual partners on the Internet. AIDS Care 2003;15:367–378.

- 31. Chen SY, Gibson S, Weide D, McFarland W. Unprotected anal intercourse between potentially HIV-serodiscordant men who have sex with men, San Francisco. *J Acquir Immune Defic Syndr* 2003;33:166–170.
- 32. McFarlane M, Bull SS, Rietmeijer CA. The Internet as a newly emerging risk environment for sexually transmitted diseases. *JAMA* 2000;284:443–446.
- 33. Klausner JD, Wolf W, Fischer-Ponce L, Zolt I, Katz MH. Tracing a syphilis outbreak through cyberspace. *JAMA* 2000;284:447–449.
- 34. Rietmeijer CA, Bull SS, McFarlane M. Sex and the Internet. *AIDS* 2001;15: 1433–1434.
- 35. Elford J, Bolding G, Sherr L. Seeking sex on the Internet and sexual risk behaviour among gay men using London gyms. *AIDS* 2001;15:1409–1415.
- Centers for Disease Control and Prevention (CDC). Internet use and early syphilis infection among men who have sex with men San Francisco, California, 1999–2003. MMWR Morb Mortal Wkly Rep 2003;52:1229–1232.
- Benotsch EG, Kalichman S, Cage M. Men who have met sex partners via the Internet: prevalence, predictors, and implications for HIV prevention. *Arch Sex Behav* 2002;31:177–183.
- Sowell RL, Lindsey C, Spicer T. Group sex in gay men: its meaning and HIV prevention implications. J Assoc Nurses AIDS Care 1998;9:59–71.
- 39. Colfax GN, Mansergh G, Guzman R, et al. Drug use and sexual risk behavior among gay and bisexual men who attend circuit parties: a venue-based comparison. *J Acquir Immune Defic Syndr* 2001;28:373–379.
- 40. Stall R, Purcell D. Intertwining epidemics: a review of research on substance use among men who have sex with men and its connection to the AIDS epidemic. *AIDS Behav* 2000;4:181–192.
- 41. Purcell DW, Parsons JT, Halkitis PN, Mizuno Y, Woods WJ. Substance use and sexual transmission risk behavior of HIV-positive men who have sex with men. *J Subst Abuse* 2001;13:185–200.
- 42. Mattison AM, Ross MW, Wolfson T, Franklin D; San Diego HIV Neurobehavioral Research Center Group. Circuit party attendance, club drug use, and unsafe sex in gay men. *J Subst Abuse* 2001;13:119–126.
- 43. McNall M, Remafedi G. Relationship of amphetamine and other substance use to unprotected intercourse among young men who have sex with men. *Arch Pediatr Adolesc Med* 1999;153:1130–1135.
- 44. Whittington WL, Collis T, Dithmer-Schreck D, et al. Sexually transmitted diseases and human immunodeficiency virus discordant partnerships among men who have sex with men. *Clin Infect Dis* 2002;35:1010–1017.
- 45. Vanable PA, Ostrow DG, McKirnan DJ. Viral load and HIV treatment attitudes as correlates of sexual risk behavior among HIV-positive gay men. *J Psychosom Res* 2003;54:263–269.
- 46. Elford J, Bolding G, Maguire M, Sherr L. Combination therapies for HIV and sexual risk behavior among gay men. *J Acquir Immune Defic Syndr* 2000;23:266–271.

- 47. International Collaboration on HIV Optimism. HIV treatments optimism among gay men: an international perspective. *J Acquir Immune Defic Syndr* 2003;32:545–550.
- 48. Sherr L, Bolding G, Maguire M, Elford J. Viagra use and sexual risk behaviour among gay men in London. *AIDS* 2000;14:2051–2053.
- 49. Chu PL, McFarland W, Gibson S, et al. Viagra use in a community-recruited sample of men who have sex with men, San Francisco. *J Acquir Immune Defic Syndr* 2003:33:191–193.
- 50. Reller ME, Olsen SJ, Kressel AB, et al. Sexual transmission of typhoid fever: a multistate outbreak among men who have sex with men. *Clin Infect Dis* 2003;37:141–144.
- 51. Gaudreau C, Michaud S. Cluster of erythromycin- and ciprofloxacin-resistant Campylobacter jejuni subsp. jejuni from 1999 to 2001 in men who have sex with men, Quebec, Canada. *Clin Infect Dis* 2003;37:131–136.
- 52. Diamond C, Thiede H, Perdue T, et al. Seroepidemiology of human herpes virus 8 among men who have sex with men. *Sex Transm Dis* 2001;28:176–183.
- 53. Johnson SR, Smith EM, Guenther SM. Comparison of gynecologic health care problems between lesbians and bisexual women: a survey of 2,345 women. *J Reprod Med* 1987;32:805–811.
- 54. Bevier PJ, Chiasson MA, Heffernan RT, Castro KG. Women at a sexually transmitted disease clinic who reported same-sex contact: their HIV seroprevalence and risk behaviors. *Am J Public Health* 1995;85:1366–1371.
- 55. Fethers K, Marks C, Mindel A, Estcourt CS. Sexually transmitted infections and risk behaviours in women who have sex with women. *Sex Transm Infect* 2000;76:345–349.
- 56. Marrazzo JM, Stine K, Wald A. Prevalence and risk factors for infection with herpes simplex virus type-1 and -2 among lesbians. *Sex Transm Dis* 2003:30:890–895.
- 57. Kwakwa HA, Ghobrial MW. Female-to-female transmission of human immunodeficiency virus. *Clin Infect Dis* 2003;36:e40–41.
- 58. Marrazo JM, Koutsky LA, Stine KL, et al. Genital human papillomavirus infection in women who have sex with women. *J Infect Dis* 1998;178: 1604–1609.
- Kellock D, O'Mahony CP. Sexually acquired metronidazole-resistant trichomoniasis in a lesbian couple. Genitourin Med 1996;72:60–61.
- 60. Campos-Outcalt D, Hurwitz S. Female-to-female transmission of syphilis: a case report. *Sex Transm Dis* 2002;29:119–120.
- 61. Walters MH, Rector WG. Sexual transmission of hepatitis A in lesbians. *JAMA* 1986;256:594.
- 62. Skinner CJ, Stokes J, Kirlew Y, Kavanagh J, Forster GE. A case-controlled study of the sexual health needs of lesbians. *Genitourin Med* 1996;72:277–280.
- 63. Berger BJ, Kolton S, Zenilman JM, Cummings MC, Feldman J, McCormack WM. Bacterial vaginosis in lesbians: a sexually transmitted disease. *Clin Infect Dis* 1995;21:1402–1405.

- 64. Marrazzo JM, Koutsky LA, Handsfield HH. Characteristics of female sexually transmitted disease clinic clients who report same-sex behaviour. *Int J STD AIDS* 2001;12:41–46.
- 65. Kennedy MB, Scarlett MI, Duerr AC, Chu SY. Assessing HIV risk among women who have sex with women: scientific and communication issues. *J Am Med Womens Assoc* 1995;50:235–248.
- 66. Lemp GF, Jones M, Kellogg TA, et al. HIV seroprevalence and risk behaviors among lesbians and bisexual women in San Francisco and Berkeley, California. *Am J Public Health* 1995;85:1549–1552.
- 67. Goldstein ST, Alter MJ, Williams IT, et al. Incidence and risk factors for acute hepatitis B in United States, 1982–1998: implications for vaccination programs. *J Infect Dis* 2002;185:713–719.
- 68. Bell BP, Shapiro CN, Alter MJ, et al. The diverse patterns of hepatitis A epidemiology in the United States implications for vaccination strategies. *J Infect Dis* 1998;178:1579–1584.
- 69. MacKellar DA, Valleroy LA, Secura GM, et al. Two decades after vaccine license: hepatitis B immunization and infection among young men who have sex with men. *Am J Public Health* 2001;91:965–971.
- 70. Canadian Immunization Guide. 6th ed. Ottawa, ON: Health Canada; 2002.
- 71. Stall R, Mills TC, Williamson J, et al. Association of co-occurring psychosocial health problems and increased vulnerability to HIV/AIDS among urban men who have sex with men. *Am J Public Health* 2003;93:939–942.
- Seal DW, Kelly JA, Bloom FR, Stevenson LY, Coley BI, Broyles LA. HIV
 prevention with young men who have sex with men: what young men
 themselves say is needed. Medical College of Wisconsin CITY Project
 Research Team. AIDS Care 2000;12:5–26.
- 73. McNair RP. Lesbian health inequities: a cultural minority issue for health professionals. *Med J Aust* 2003;178:643–645.
- Harrison AE. Primary care of lesbian and gay patients: educating ourselves and our students. Fam Med 1996;28:10–23.
- 75. Richters J, Bergin S, Lubowitz S, Prestage G. Women in contact with Sydney's gay and lesbian community: sexual identity, practice, and HIV risks. *AIDS Care* 2002;14:193–202.
- 76. Page-Shafer K, Graves A, Kent C, Balls JE, Zapitz VM, Klausner JD. Increased sensitivity of DNA amplification testing for the detection of pharyngeal gonorrhea in men who have sex with men. *Clin Infect Dis* 2002;34:173–176.
- 77. Golden MR, Astete SG, Galvan R, et al. Pilot study of COBAS PCR and ligase chain reaction for detection of rectal infections due to Chlamydia trachomatis. *J Clin Microbiol* 2003;41:2174–2175.

PREGNANCY

This chapter will highlight aspects of STI management particular to pregnancy, but details for each condition should be reviewed elsewhere in these guidelines.

A lower threshold of screening for sexually transmitted infections (STIs) should exist in pregnancy, as there are significant potential complications for both the pregnancy outcome (gestational age at delivery and type of delivery) and the health of the newborn, due to the risk of vertical transmission. As such, the following recommendations are presented.

- At the first prenatal visit, all pregnant women should be:
 - Offered HIV counselling and testing.
 - Screened for hepatitis B surface antigen (HBsAg).
 - Screened for *Chlamydia trachomatis* and *Neisseria gonorrhoeae*.
 - Screened for syphilis.
- All pregnant women should be evaluated for STI risk factors prior to and during pregnancy. Risk factors are described in the *Primary Care and Sexually Transmitted Infections* chapter. Any woman with ongoing risk factors for STI acquisition during pregnancy should be considered for rescreening each trimester.
- If an STI is diagnosed in pregnancy, treatment specific to the disease should be initiated, taking the pregnancy into consideration (see below).
- Due to the potential for decreased efficacy of treatments in pregnancy, follow-up after treatment of STIs for both the patient and her sexual partner(s) is important to ensure therapeutic success.

Antimicrobial Therapy in Pregnancy

- Special attention is required to safely treat STIs in pregnancy.
- Always consult with an experienced colleague if you are unclear about a
 medication risk in pregnancy. Data or risks associated with antimicrobials
 are beyond the scope of this document. The Motherisk Clinic at the Hospital
 for Sick Children in Toronto is an excellent resource and can be accessed at
 www.motherisk.org or by calling (416) 813-6780.
- The following is an incomplete list of drugs that are relatively or absolutely contraindicated in pregnancy:
 - Erythromycin estolate
 - Sulfamethoxazole
 - Fluoroguinolones
 - Podophyllin/podophyllotoxin/5-fluoro-uracil/imiquimod (not licensed for use in pregnancy)
 - Doxycycline/tetracycline/minocycline
 - Gamma benzene hexachloride/lindane
 - Interferons
 - Ribavirin

Specific Issues Related to Obstetric and Gynecologic Circumstances STI and pregnancy termination

Women presenting for surgical or medical termination of pregnancy should ideally be screened for STIs prior to termination. When feasible, screening for chlamydia and gonorrhea and subsequent treatment are appropriate pre-procedure. When screening is not feasible, prophylaxis pre-procedure with single-dose azithromycin (1 g PO [A-I]) or doxycycline for *C trachomatis* coverage is recommended.¹ Although bacterial vaginosis (BV) is thought to contribute to postoperative infection, a recent randomized clinical trial of treatment with metronidazole prior to surgery in documented cases of BV did not improve outcome.² Further study is required in this area.

Artificial insemination and STI risk

STI risk with donor insemination is reduced with current Canadian practices of serologic screening for HIV, hepatitis B virus (HBV), hepatitis C virus (HCV) and syphilis. It is recommended that donor semen be stored until repeat donor serology at 6 months is negative for HIV. Initial and repeat screening of donor semen should include *N gonorrhoeae* and *C trachomatis* testing.³ Antibiotic use at the time of embryo transfer to reduce iatrogenic pelvic inflammatory disease from *C trachomatis* has not been studied in a controlled fashion.⁴ However, a recent survey in the U.K. indicates that *C trachomatis* prophylaxis is used in half of embryo transfers in that country.⁵

Chlamydia Trachomatis

There are variable reports in the literature, but no consistent association exists between poor pregnancy outcome (i.e., preterm birth or preterm prolonged rupture of membranes) and *C trachomatis* cervicitis.⁶ Vertical transmission occurs in 50% of infants born vaginally to infected mothers. Vertical transmission can occur with cesarean section where membranes are intact. Of those neonates who acquire infection, at least 20% develop conjunctivitis, and 20% develop pneumonia.^{7,8} Although provincial guidelines may vary, general national recommendations are to screen for *C trachomatis* early in pregnancy. Repeat screening should be performed in the third trimester for women at continuing risk for STI acquisition. (See *Chlamydial Infections* chapter for a full discussion of *C trachomatis* diagnosis and management.)

Treatment

Table 1. Treatment for C trachomatis during pregnancy

• Amoxicillin 500 mg PO tid for 7 days [A-I]

OR

• Erythromycin base 500 mg PO qid for 7 days [A-I]

OR

Azithromycin 1 g PO in a single dose if poor compliance is expected [A-I]

Note:

Doxycycline and quinolones are contraindicated in pregnancy and in lactating women. Erythromycin estolate is contraindicated in pregnancy due to associated hepatotoxicity and cholestatic hepatitis. Amoxicillin and erythromycin are effective; however, compliance with erythromycin may be difficult due to gastrointestinal side effects. Azithromycin appears to be safe and effective. Azithromycin appears to be safe and effective.

Sexual partners should be treated and undergo follow-up testing to ensure cure. Condoms or abstinence are recommended during treatment and until follow-up tests are negative. Repeat polymerase chain reaction (PCR) chlamydial testing may be positive due to the presence of persistent DNA from killed organisms for up to 4 weeks after the completion of treatment. Repeat testing should therefore be by PCR (as it is most sensitive) at 3–4 weeks post-treatment, or by culture if time does not allow for a 3 week waiting period. All pregnant women should be retested following treatment to ensure cure.

Gonococcal Infections

Infection with *N gonorrhoeae* in pregnancy is associated with endometritis, pelvic sepsis, ophthalmia neonatorum and systemic neonatal infection.¹⁴ Although gonococcal infection is relatively uncommon in many clinical practices, it is still suggested that all pregnant women be screened in early pregnancy due to the adverse consequences of an untreated infection.

Those infected should be treated with a recommended or alternate cephalosporin. Women with a penicillin allergy or those who cannot tolerate a cephalosporin should be administered a single 2 g dose of spectinomycin IM. A diagnosis of *N gonorrhoeae* is strongly associated with co-infection of *C trachomatis*. Treatment for both STIs is recommended when *N gonorrhoeae* is diagnosed, unless testing for *C trachomatis* is negative. In pregnant women, a test of cure is recommended. (See *Gonococcal Infections* chapter for a full discussion of *N gonorrhoeae* diagnosis and management.)

Ensure simultaneous treatment of *C trachomatis* when treating *N gonorrhoeae*, unless testing for *C trachomatis* is documented negative (see *Chlamydial Infections* and *Gonococcal Infections* chapters).

Treatment

Table 2. Treatment for N gonorrhoeae during pregnancy

Preferred	Alternative
 Cefixime 400 mg P0 in a single dose [A-I] OR Ceftriaxone 125 mg IM in a single dose [A-I] 	Spectinomycin 2 g IM in a single dose (available only through SAP) [A-I]

SAP = Special Access Program

All sexual partners of patients who have *N gonorrhoeae* infection should be evaluated and treated for both *N gonorrhoeae* and *C trachomatis* infections. Patients and contacts should abstain from unprotected intercourse until treatment of both partners is complete (i.e., after completion of a multiple-dose treatment or for 7 days after single-dose therapy). In pregnancy, a test of cure in both partners is recommended.

Syphilis

Infectious syphilis in pregnancy, defined as primary, secondary or early latent infection (typically the first year after infection), can lead to fetal infection with stillbirth, preterm birth, congenital abnormalities and active disease at delivery. Transmission occurs transplacentally (as early as 14 weeks and throughout pregnancy) or at the time of delivery. Untreated primary or secondary syphilis carries a transmission risk of up to 100%, while early latent infection has a 40% transmission risk. Treated syphilis has a transmission rate of 1.8%. In a small Canadian study, 1 of 98 treated women had a child with congenital syphilis, whereas 4 of 9 women not treated in pregnancy had infants with congenital syphilis.

All women should be screened serologically with a non-treponemal screening test for syphilis at the first prenatal visit (Venereal Disease Research Laboratories [VDRL] or rapid plasma reagin [RPR]). In those considered high-risk, a treponemal test should be added to initial testing, and repeat testing should be performed at both 28 weeks' gestation and delivery. If screening serology is positive, treponemal-specific testing is required to confirm the diagnosis: *Treponema pallidum* immobilization (TPI), fluorescent treponemal antibody absorbed (FTA-ABS) or microhemagglutination-*T pallidum* (MHA-TP) (*T pallidum* particle agglutination [TP-PA] in Quebec). Any woman who delivers a stillborn infant after 20 weeks' gestation should be tested for syphilis.

Biological false-positive results are possible with non-treponemal and treponemal tests in pregnancy, but they are more common with non-treponemal results.

For details on specific tests, see *Syphilis* chapter.

Diagnostic considerations

Pregnant women with confirmed syphilis should be considered infected unless an adequate treatment history is documented and sequential serologic antibody titres have declined. In some cases, titres will not decline to undetectable levels even after successful treatment and may remain positive at low levels, such as 1:1 or 1:2, indefinitely.

Treatment

Penicillin is effective for preventing maternal transmission to the fetus and for treating fetal infection. Treatment during pregnancy should consist of the penicillin regimen appropriate for the presenting stage. Penicillin alternatives have not been proven effective for the treatment of syphilis during pregnancy. Pregnant women who have a history of significant penicillin allergy should be desensitized and then treated with penicillin.

Table 3. Treatment for syphilis during pregnancy

Primary or secondary syphilis	Benzathine penicillin G 2.4 million units IM in a single dose (available only through SAP) [B-II]
Early latent syphilis	Benzathine penicillin G 2.4 million units IM in a single dose (available only through SAP) [B-II]
Late latent syphilis or latent syphilis of unknown duration	Benzathine penicillin G 7.2 million units total, administered as three doses of 2.4 million units IM each at 1 week intervals (available only through SAP) [B-II]

SAP = Special Access Program

In the second half of pregnancy, management and counselling may be facilitated by a sonographic fetal evaluation for congenital syphilis, but this should not delay therapy. Sonographic signs of fetal syphilis (i.e., hepatomegaly, ascites and hydrops) indicate a greater risk for fetal treatment failure; such cases should be managed in consultation with obstetric specialists.²²

Women treated for syphilis during the second half of pregnancy are at risk for premature labour and/or fetal distress if the treatment precipitates the Jarisch-Herxheimer reaction; this includes fever, uterine irritability and contractions. It is estimated to occur in 40% of patients with primary or secondary syphilis, begins on average within 10 hours of treatment and resolves within 24 hours. ²³ These women should be advised to seek obstetric attention after treatment if they notice any contractions or decrease in fetal movements. Some centres admit and conduct fetal monitoring at the time of treatment. Although stillbirth is a rare complication of treatment, concern about this complication should not delay necessary treatment.

All patients who have syphilis should be offered testing for HIV infection. In the case of suspected congenital syphilis, consult a colleague with experience in this area.

Trichomoniasis

Vaginal trichomoniasis has been associated with adverse pregnancy outcomes, particularly premature rupture of the membranes, preterm delivery and low birthweight. However, data have not indicated that treating asymptomatic trichomoniasis during pregnancy lessens the risk of adverse pregnancy outcomes. In fact, treatment of asymptomatic trichomoniasis with metronidazole 2 g x 2 doses has been shown to increase preterm birth in a placebo-controlled trial. For this reason, screening of all pregnant women cannot be recommended. Women who are symptomatic with trichomoniasis, however, should be treated to ameliorate symptoms and minimize the risk of sexual transmission as described below. 95-27 Women may be treated with 2 g of metronidazole orally in a single dose. Marginally better cure rates have been found with 7 day treatment (as per treatment recommendations, below). Multiple studies and meta-analyses have not demonstrated a consistent association between metronidazole use during pregnancy and adverse fetal effects — it is therefore considered safe in pregnancy.

Diagnostic considerations

Diagnosis of vaginal trichomoniasis is usually performed by microscopy of vaginal secretions (wet mount), but this method has a sensitivity of only about 60–70%. Microscopy and culture performed rapidly from time of sample collection is the most sensitive available method of diagnosis.

Treatment

Table 4. Treatment for trichomoniasis during pregnancy

Preferred	Alternative
Metronidazole 2 g PO in a single dose [A-I]	Metronidazole 500 mg PO bid for 7 days [A-I]

Topical therapy is ineffective for cure compared to oral metronidazole (<50% with intravaginal treatment).³¹ Treatment of sexual partner(s) is essential for cure.

Abstinence during treatment is recommended to avoid reinfection. Retesting in pregnancy is necessary only for those who remain symptomatic after treatment.

Bacterial Vaginosis

Bacterial vaginosis in pregnancy has been associated with adverse outcomes, including premature rupture of membranes, preterm labour, preterm birth and postpartum endometritis. There is evidence to support screening and treatment at 12–16 weeks in high-risk pregnancies (i.e., previous preterm labour/delivery or preterm premature rupture of membranes). If the patient is symptomatic or at high risk, test for BV and treat as below. Treatment of BV in

such cases may reduce the risk of prematurity, low birthweight and preterm premature rupture of membranes.^{32–35} In low-risk and asymptomatic women, screening is not recommended, as it has not been shown to affect adverse outcomes in well-designed randomized, controlled trials.^{35,36} If symptoms suggest BV, testing is appropriate, and positive results warrant treatment for symptom resolution.

Treatment

Table 5. Treatment for bacterial vaginosis during pregnancy

Preferred	Alternative
• Metronidazole 500 mg PO bid for 7 days [A-I]	• Clindamycin 300 mg PO bid for 7 days [A-I]

Systemic rather than topical treatment is recommended in pregnancy, as vaginal treatment has not been shown to decrease the risk of adverse pregnancy outcomes. Also, clindamycin topical treatment has been associated with adverse outcomes in the newborn when used in pregnancy. Based on multiple studies, most recently assessed by meta-analysis, the evidence supports the safety and lack of teratogenicity of systemic metronidazole use in pregnancy. Bescreening and re-treating may be advisable in women with high-risk pregnancies (i.e., previous preterm labour, delivery or preterm premature rupture of membranes). It is important to note that clindamycin has been associated with increased risk of pseudomembranous colitis and should be used only when alternatives are not possible.

Vulvovaginal Candidiasis

Vulvovaginal candidiasis is a common occurrence in pregnancy. Management depends on the degree of symptomatology. Often *Candida* is difficult to eradicate in pregnancy, so the primary goal of therapy should be symptom control. To date, only topical "azole" treatments are recommended in pregnancy, and these should be monitored by a physician. Treatment for 7 days may be necessary in pregnancy to achieve resolution of symptoms.⁴⁰ Oral fluconazole is considered teratogenic in animal studies,⁴¹ but in 226 cases of first-trimester exposure in humans there was no increased risk of complications.⁴² There are reports, however, of women with chronic exposure in pregnancy who have had infants with skeletal malformation syndromes suggestive of fluconazole teratogenic effects.^{43,44} Therefore, oral "azoles" are not recommended. The use of intravaginal boric acid is not recommended in pregnancy due to its teratogenic potential in animal studies.⁴⁵

Treatment

Table 6. Treatment options for vulvovaginal candidiasis during pregnancy

Butoconazole [A-I]	2% cream 5 g (butaconazole1-sustained release) in a single intravaginal application
Clotrimazole [A-I]	 1% cream 5 g intravaginally per day for 7–14 days OR 100 mg vaginal tablet, one tablet per day for 7 days OR 100 mg vaginal tablet, two tablets per day for 3 days OR 500 mg vaginal tablet, one tablet in a single application
Miconazole [A-I]	 2% cream 5 g intravaginally per day for 7 days OR 100 mg vaginal suppository, one suppository per day for 7 days OR 200 mg vaginal suppository, one suppository per day for 3 days
Nystatin [A-I]	• 100,000 unit vaginal tablet, 1 tablet per day for 14 days
Terconazole [A-I]	 0.4% cream 5 g intravaginally per day for 7 days 0.8% cream 5 g intravaginally per day for 3 days 0R 80 mg vaginal suppository, one suppository per day for 3 days

Ectoparasitic Infestations

Phthirus pubis

Patients who have *P pubis* (i.e., pubic lice) usually seek medical attention because of pruritus, lice or nits in their pubic hair. Pediculosis pubis is usually transmitted by sexual contact.⁴⁶ Treatment should be given in pregnancy as follows (see also *Ectoparasitic Infestations* chapter).

Treatment

Table 7. Treatment for pubic lice during pregnancy

- Permethrin 1% cream rinse applied to affected areas and washed off after 10 minutes [B-II] OR
- Pyrethrins with piperonyl butoxide applied to the affected area and washed off after 10 minutes [B-III]

Note: Lindane is contraindicated in pregnancy.

Follow-up

Patients should be evaluated after 1 week if symptoms persist. Re-treatment may be necessary if lice or eggs are observed at the hair-skin junction. Patients who do not respond to one of the recommended regimens should be re-treated with an alternative regimen. However, pruritus alone in the absence of persistent organisms warrants symptomatic treatment only.

Sexual partners within the last month should be treated. Patients should avoid sexual contact with their sexual partner(s) until patients and partners have been treated and re-evaluated to rule out persistent disease.

Scabies

The predominant symptom of scabies is pruritus. Sensitization to *Sarcoptes scabiei* must occur before pruritus begins. The first time a person is infected with *S scabiei*, sensitization takes up to several weeks to develop. However, pruritus may occur within 24 hours after a subsequent reinfestation. Scabies in adults often is sexually acquired, although scabies in children is usually not (see *Ectoparasitic Infestations* chapter for more information on transmission). Pruritus may persist for several days or weeks after treatment.^{46–48}

Treatment

Table 8. Treatment for scabies during pregnancy

 Permethrin cream (5%) applied to all affected areas of the body from the neck down and washed off after 8–14 hours [B-II]

Note: Lindane and ivermectin are contraindicated in pregnancy and lactation.

Both sexual and close personal or household contacts within the preceding month should be examined and treated. Re-treat if symptoms persist or recur.

Genital Herpes Simplex Virus Infection

Counselling on the signs and symptoms of herpes simplex virus (HSV), as well as risk reduction behaviours to avoid contracting genital herpes, is important for all women who present for pregnancy care. There is currently no evidence to investigate or treat pregnant women who have no history of genital herpes and whose partners also have no history. However, without past history, these women are at risk of acquiring primary infection in pregnancy. Primary infection in pregnancy is associated with significant vertical transmission rates.

Women without a history of HSV should receive risk-reduction behaviour counselling to avoid contracting HSV. Both HSV-1 and HSV-2 can cause genital lesions, be vertically transmitted and lead to neonatal disease. Diagnosis of genital

herpes can be complicated due to the common phenomenon of asymptomatic or subclinical disease. Diagnosis requires a careful assessment of clinical features, culture or PCR of genital sites and type-specific serology. Neonatal HSV is associated with significant morbidity and mortality, causing cutaneous, central nervous system and disseminated disease, such as pneumonitis and encephalitis.

Primary infection

If the mother is seronegative, she is at risk of primary infection with HSV-1 or -2 in pregnancy. If this occurs during the second half of pregnancy, a vertical transmission rate of 30–50% exists. 49,50 A significant proportion of neonatal herpes cases are born to mothers with no recognized history of genital herpes. 51,52 At present, there is no evidence that routine serotesting in pregnancy will be successful at decreasing the risk of neonatal herpes. However, if a known serosusceptible pregnant woman is known to have a partner with oral or genital herpes, it is prudent to advise abstinence from oral and/or genital sexual contact. In addition, non-pregnancy data would suggest that suppressive therapy in the male partner with genital herpes would decrease the risk of sexual transmission, but this should not replace abstinence or judicious condom use.53

Treatment

Table 9. Treatment for genital HSV during pregnancy

Acyclovir 200 mg five times per day for 5–10 days [A-I]⁵⁴

Primary infection in pregnancy warrants acyclovir treatment and consideration of cesarean section for delivery, especially if infection is in the late third trimester. Such measures reduce, but do not eliminate, the risk of vertical transmission.⁵⁵ See *Genital Herpes Simplex Virus Infections* chapter for more information on treatment.

Recurrent infection

In a woman with prior infection, the risk of vertical transmission is 2–4%. For those who have had an outbreak within the previous year, prophylaxis at 36 weeks' gestation until delivery with acyclovir 400 mg PO tid is recommended [A-I]. ⁵⁴ Transmission can occur at the time of delivery, with or without lesions, due to asymptomatic shedding. Treatment with acyclovir reduces the risk of lesions and the risk of asymptomatic viral shedding, thereby reducing the cesarean section rate. ^{54,56} See *Genital Herpes Simplex Virus Infections* chapter for more information on suppressive therapy.

If genital lesions or prodromal symptoms are present at the time of delivery, cesarean section is recommended.⁵⁶ In the event of ruptured membranes, cesarean section is thought to confer protection, ideally if performed within less than 4 hours ^{57,58}

Genital Warts and Genital Human Papillomavirus Infection

Vertical transmission of genital human papillomavirus (HPV) types 6 and 11 can cause recurrent respiratory papillomatosis (RRP) in infants and children. Symptomatic perinatal transmission is infrequent and is usually clinically apparent within 2 years. When it occurs, it is associated with anogenital and vocal-cord lesions in the newborn. Although maternal HPV prevalence is high, HPV vertical transmission is low, and RRP is rare. ⁵⁹⁻⁶¹ The value of cesarean section for reducing/preventing transmission is unknown. Cesarean section is not recommended for the sole purpose of reducing transmission of HPV to the newborn. If the pelvic outlet is obstructed by warts, or if the warts are significant in number as to cause a bleeding complication with vaginal delivery, cesarean section may be warranted.

Genital warts may proliferate, reappear and become friable in pregnancy. Women should be reassured that this growth usually regresses postpartum. In general, the practice is to defer treatment due to poor response to therapy in pregnancy. If treatment is desired, the following options are appropriate. Weekly treatment may be required.

Treatment

Table 10. Treatment for genital HPV during pregnancy

- TCA trichloroacetic acid (85%) [B-II]
- · Cryotherapy (liquid nitrogen) [B-II]
- CO₂ laser [B-II]
- Surgical excision [B-II]

Note: Imiquimod, podophyllin, podofilox, podophyllotoxin, 5-fluoro-uracil and interferon are contraindicated in pregnancy.

Hepatitis A Virus Infection

Vertical transmission of hepatitis A virus is not described. An infected woman can infect her newborn through the usual fecal/oral routes of transmission. Immunization and/or gammaglobulin treatment in pregnancy is safe and may confer some protection for the newborn.⁶²

If a pregnant woman is infected, consider prophylaxis with vaccine and/or gammaglobulin for household contacts. Household contacts should consider receiving hepatitis A vaccine. If a pregnant woman is a contact of an infected person, there is no contraindication to the use of gammaglobulin or hepatitis A vaccine in pregnancy [B-II].

Hepatitis B Virus Infection

Mothers who are acutely infected with HBV, or are carriers, can transmit the virus to their infant. Transmission appears to occur at time of delivery, but not transplacentally. Depending on the stage of maternal infection, the vertical transmission risk of hepatitis B can be as high as 90% in the absence of intervention at the time of delivery. 63 Ninety-five percent of cases can be prevented with the use of hepatitis B immune globulin (HBIG) and hepatitis B vaccine administered at birth to the neonate, followed by two additional vaccine doses at 1 and 6 months. 64 The first dose of hepatitis B vaccine should be administered within 12 hours of birth and HBIG immediately after birth (efficacy decreases sharply after 48 hours). 65

If a woman is newly identified to be HBsAg-positive in pregnancy, she warrants further investigation. Consideration should be given to testing for HIV, hepatitis B e antigen, hepatitis B core antibody, HBV DNA, hepatitis A IgM and hepatitis C antibodies (anti-HBc IgM and IgG). If she is found to be positive for any of these, an evaluation of liver transaminases and function is warranted (see *Hepatitis B Virus Infections* chapter).

If the mother is infectious at time of delivery, document the diagnosis on prenatal forms and plan for administering HBIG and the first dose of hepatitis B vaccine to the neonate immediately after birth. The second and third dose of the vaccine should be given to the infant 1 and 6 months after the first dose. Special attention is required to complete the three-dose schedule, since long-term exposure is possible and there may be difficulty in reaching the family for the third dose. A follow-up hepatitis B surface antibody at 1–2 months after completion of HBV vaccine series to document adequate immune response is recommended (see *Hepatitis B Virus Infections* chapter and *Canadian Immunization Guide*).³ Breastfeeding is safe if the neonate is treated.

If the mother is a contact of an infected person or is at risk of acquiring hepatitis B, there is no contraindication to HBIG or HBV vaccine in pregnancy [A-I].

Hepatitis C Virus Infection

Approximately 0.8% of the Canadian population is infected with hepatitis C.⁶⁶ Persons with hepatitis C should be referred to health care professionals with experience in the treatment of hepatitis C. Pregnancy does not appear to have an effect on the progression of hepatitis C.

Hepatitis C in pregnancy may be associated with increased rates of cholestasis. 67

The risk of vertical transmission is estimated to be 7.9%.⁶⁸ It is not yet known if cesarean section reduces the vertical transmission of HCV, as it has not been adequately studied to date.⁶⁹

Breastfeeding is considered to be safe unless nipples are cracked or bleeding. Although HCV RNA has been identified in breast milk,⁷⁰ breastfeeding is still considered safe. Assessment of and education to reduce risk behaviour is important in pregnancy.

Current treatments available for HCV infection are contraindicated in pregnancy (i.e., interferon-alpha and ribavirin, combination therapies of pegylated interferon-alpha 2a and 2b plus ribavirin). Although not well studied, interferon-alpha does not appear to have an adverse affect on the human embryo or fetus, but it is associated with increased rates of preterm delivery and intrauterine growth restriction. Animal studies have shown an increased rate of fetal loss.⁷¹ If interferon is to be used in pregnancy, the potential benefits of its use should clearly outweigh possible hazards.⁷²⁻⁷⁴ Because there are no large studies of ribavirin use during human pregnancy and ribavirin is highly teratogenic in animal studies, its use during pregnancy is absolutely contraindicated.⁷⁵ Ribavirin has been given Pregnancy Category X by the U.S. Food and Drug Administration. It is recommended that women and/or their male partners who have received ribavirin as part of a combination treatment for HCV infection both use a highly effective method of birth control to prevent pregnancy during ribavirin therapy and for 6 months afterward.

Canadian guidelines for the management of hepatitis C in pregnancy are detailed elsewhere. 68

Human Immunodeficiency Virus Infection

All women should be offered HIV antibody testing with appropriate counselling and informed consent at their first prenatal visit. A diagnosis of HIV and pregnancy presents a need for complex care and requires consultation with experts in the area as soon as possible. Initiation of antiretroviral therapy in HIV-infected pregnant women is critical for the reduction of vertical transmission; this typically consists of combination antiretroviral therapy, also known as highly active antiretroviral therapy (HAART). Effective suppression of viral load in pregnancy prior to delivery, along with intrapartum and 6 weeks of neonatal antiretroviral therapy, reduces vertical transmission from 25% to less than 1%.76

If the mother is found to be HIV-positive on confirmatory testing (see *Human Immunodeficiency Virus Infections* chapter), consultation should be made with a specialist in HIV pregnancy care. The best care and greatest chance for viral suppression is with early management. If the pregnancy is to be continued, HAART should be initiated either immediately or at 14–18 weeks' gestation, depending on CD4 counts and viral load. Women should be counselled regarding the potential side effects of antiretroviral therapy, the importance of strict compliance and need for close monitoring. At a minimum, monthly complete blood count, aspartate aminotransferase, alanine aminotransferase, amylase, bilirubin, creatinine, serum lactate, glucose, CD4 count and viral load are recommended. Specific guidelines are found elsewhere.⁷⁶

Specific antiretroviral drugs that are contraindicated in pregnancy include the following:

- Efavirenz
- Delavirdine
- Hydroxyurea
- Nevirapine (the initiation of continuous nevirapine in pregnancy is not currently recommended due to its potential toxicities: rash, severe hepatitis, Stevens-Johnson syndrome)

If a woman presents in pregnancy already taking nevirapine and tolerating it well, continuation may be considered. One-time maternal dosing of nevirapine used in the high-risk setting at the time of delivery is still appropriate.

Because of the complexity associated with the use of antiretroviral drugs in pregnancy, all HIV-positive pregnant women should be managed in cooperation with an HIV specialist.

If HIV viral load is undetectable at the time of delivery, vaginal delivery is usually recommended, unless cesarean section is required for obstetric reasons. With a viral load greater than 1,000 copies/mL, a cesarean section is usually recommended to reduce the risk of vertical transmission.⁷⁷⁻⁸¹ Additionally, all infected women should receive IV zidovudine from the onset of labour until delivery or before a cesarean section is started. Breastfeeding is contraindicated, as HIV can be transmitted through breast milk.

Women who are diagnosed HIV-positive late in pregnancy or in labour are at very high risk for perinatal transmission of infection. Further management should be in cooperation with both adult and pediatric HIV specialists, who may recommend one or more of the following: intrapartum prophylaxis options with IV zidovudine, cesarean section, single-dose nevirapine to the woman in labour and single-dose nevirapine to the infant, and 6 weeks of oral antiretroviral therapy to the infant.⁷⁶

Note that these guidelines are under constant revision, and each case should be managed with an expert in the area. For more detailed information, please see the Canadian guidelines for management of HIV-affected pregnancy, labour and delivery, and postpartum period.⁷⁶

References

- Montgomery C, Norman W, Money D, Rekart M. Antibiotic at time of induced abortion. BCMJ 2002:44:367–373.
- Miller L, Thomas K, Hughes JP, Holmes KK, Stout S, Eschenbach DA. Randomised treatment trial of bacterial vaginosis to prevent post-abortion complication. *BJOG* 2004;111:982–988.

- Guidance on the Processing and Distribution of Semen for Assisted Conception Regulations. Ottawa, ON: Health Products and Food Branch, Health Canada; 2004.
- Witkin SS, Linhares IM. Chlamydia trachomatis in subfertile women undergoing uterine instrumentation: an alternative to direct microbial testing or prophylactic antibiotic treatment. *Hum Reprod* 2002;17:1938–1941.
- 5. Sowerby E, Parsons J. Prevention of iatrogenic pelvic infection during in vitro fertilization current practice in the UK. *Hum Fertil (Camb)* 2004;7:135–140.
- 6. Monif GRG, Baker DA. *Infectious Diseases in Obstetrics and Gynecology.* 5th ed. New York, NY: Parthenon Publishing; 2004:323–324.
- Johnson RE, Newhall WJ, Papp JR, et al. Screening to detect Chlamydia trachomatis and Neisseria gonorrhoeae infections—2002. MMWR Recomm Rep 2002;51(RR-15):1–38.
- Schachter J, Grossman M, Sweet RL, Holt J, Jordan C, Bishop E. Prospective study of perinatal transmission of Chlamydia trachomatis. *JAMA* 1986;255:3374–3377.
- Alary M, Joly JR, Moutquin JM, et al. Randomized comparison of amoxicillin and erythromycin in treatment of genital chlamydial infection in pregnancy. *Lancet* 1994;344:1461–1465.
- Brocklehurst P, Rooney G. Interventions for treating genital Chlamydia trachomatis infections in pregnancy. *Cochrane Database Syst Rev* 2000; 2:CD000054.
- 11. Adair CD, Gunter M, Stovall TG, McElvoy G, Veille JC, Ernest JM. Chlamydia in pregnancy: a randomized trial of azithromycin and erythromycin. *Obstet Gynecol* 1998;91:165–168.
- 12. Wehbeh HA, Ruggeirio RM, Shakem S, Lopez G, Ali Y. Single dose azithromycin for Chlamydia in pregnant women. *J Reprod Med* 1998;43:509–514.
- 13. Takahashi S, Hagiwara T, Shiga S, Hirose T, Tsukamoto T. Detection of antimicrobial-treated Chlamydia trachomatis with Amplicor PCR test kit. *J Infect Chemother* 2000;6:211–215.
- 14. Brocklehurst P. Antibiotics for gonorrhoeae in pregnancy. *Cochrane Database Syst Rev* 2002;2:CD000098.
- 15. Ramus RM, Sheffield JS, Mayfield JA, Wendel GD Jr. A randomized trial that compared oral cefixime and intramuscular ceftriaxone for the treatment of gonorrhea in pregnancy. *Am J Obstet Gynecol* 2001;185:629–632.
- 16. Kouri YH, Gonzalez L, Perez M, et al. Effect of penicillin and spectinomycin given for urethritis and cervicitis with Neisseria gonorrhoeae: high prevalence of penicillin-resistant isolates. *Genitourin Med* 1989;65:342–346.
- 17. Creighton S, Tenant-Flowers M, Taylor CB, Miller R, Low N. Co-infection with gonorrhea and chlamydia: how much is there and what does it mean? *Int J STD AIDS* 2003;14:109–113.
- 18. Washington AE, Browner WS, Korenbrot CC. Cost-effectiveness of combined treatment for endocervical gonorrhea. Considering co-infection with chlamydia trachomatis. *JAMA* 1987;257:2056–2060.

- 19. Fiumara NJ, Fleming WL, Downing JG, Good FL. The incidence of prenatal syphilis at the Boston City Hospital. *N Engl J Med* 1952;247:48–52.
- Alexander JM, Sheffield JS, Sanchex PJ, Mayfield J, Wendel GD Jr. Efficacy of treatment for syphilis in pregnancy. *Obstet Gynecol* 1999;93:5–8.
- Jones H, Taylor D, Montgomery CA, et al. Prenatal and congenital syphilis in British Columbia. J Obstet Gynaecol Can 2005;27:467–472.
- 22. Wendel GD Jr, Sheffield JS, Hollier LM, Hill JB, Ramsey PS, Sanchez PJ. Treatment of syphilis in pregnancy and prevention of congenital syphilis. *Clin Infect Dis* 2002;35(suppl 2):S200–209.
- 23. Myles TD, Elam G, Park-Hwang E, Nguyen T. The Jarisch-Herxheimer reaction and fetal monitoring changes in pregnant women treated for syphilis. *Obstet Gynecol* 1998;92:859–864.
- 24. Klebanoff MA, Carey JC, Hauth JC, et al; National Institute of Child Health and Human Development Network of Maternal-Fetal Medicine Units. Failure of metronidazole to prevent preterm delivery among pregnant women with asymptomatic Trichomonas vaginalis infection. N Engl J Med 2001;345: 487–493.
- duBouchet L, McGregor JA, Ismail M, McCormack WM. A pilot study of metronidazole vaginal gel versus oral metronidazole for the treatment of Trichomonas vaginalis vaginitis. Sex Transm Dis 1998;25:176–179.
- Tidwell BH, Lushbaugh WB, Laughlin MD, Cleary JD, Finley RW. A doubleblind placebo-controlled trial of single-dose intravaginal versus single-dose oral metronidazole in the treatment of trichomonal vaginitis. *J Infect Dis* 1994;170:242–246.
- 27. Hager WD, Brown ST, Kraus SJ, Kleris GS, Perkins GJ, Henderson M. Metronidazole for vaginal trichomoniasis. Seven-day vs single-dose regimens. *JAMA* 1980;244:1219–1220.
- 28. Caro-Paton T, Carvajal A, Martin de Diego I, Martin-Arias LH, Alvarez Requejo A, Rodriguez Pinilla E. Is metronidazole teratogenic? A meta-analysis. *Br J Clin Pharmacol* 1997;44:179–182.
- 29. Burtin P, Taddio A, Ariburnu O, Einarson TR, Koren G. Safety of metronidazole in pregnancy: a meta-analysis. *Am J Obstet Gynecol* 1995;172(2 Pt 1):525–529.
- 30. Piper JM, Mitchel EF, Ray WA. Prenatal use of metronidazole and birth defects: no association. *Obstet Gynecol* 1993;82:348–352.
- 31. Antonelli NM, Diehl SJ, Wright JW. A randomized trial of intravaginal nonoxynol 9 versus oral metronidazole in the treatment of vaginal trichomoniasis. *Am J Obstet Gynecol* 2000;182:1008–1010.
- 32. Hauth JC, Goldenberg RL, Andrews WW, DuBard MB, Copper RL. Reduced incidence of preterm delivery with metronidazole and erythromycin in women with bacterial vaginosis. *N Engl J Med* 1995;333:1732–1736.
- 33. Morales WJ, Schorr S, Albritton J. Effect of metronidazole in patients with preterm birth in preceding pregnancy and bacterial vaginosis: a placebocontrolled, double-blind study. *Am J Obstet Gynecol* 1994;171:345–347.

- McDonald HM, O'Loughlin JA, Vigneswaran R, et al. Impact of metronidazole therapy on preterm birth in women with bacterial vaginosis flora (Gardnerella vaginalis): a randomised, placebo controlled trial. *Br J Obstet Gynaecol* 1997:104:1391–1397.
- 35. McDonald H, Brockelhurst P, Parsons J, Vigneswaran R. Antibiotics for treating bacterial vaginosis in pregnancy. *Cochrane Database Syst Rev* 2003;2: CD000262.
- 36. Carey JC, Klebanoff MA, Hauth JC, et al. Metronidazole to prevent preterm delivery in pregnant women with asymptomatic bacterial vaginosis; National Institute of Child Health and Human Development Network of Maternal-Fetal Medicine Units. *N Engl J Med* 2000;342:534–540.
- 37. McGregor JA, French JI, Jones W, et al. Bacterial vaginosis is associated with prematurity and vaginal fluid mucinase and sialidase: results from a controlled trial of topical clindamycin cream. *Am J Obstet Gynecol* 1994;170:1048–1059.
- 38. Joesoef MR, Hillier SL, Wiknjosastro G, et al. Intravaginal clindamycin treatment for bacterial vaginosis: effects on preterm delivery and low birth weight. *Am J Obstet Gynecol* 1995;173:1527–1531.
- 39. Vermeulen GM, Bruinse HW. Prophylactic administration of clindamycin 2% vaginal cream to reduce the incidence of spontaneous preterm birth in women with an increased recurrence risk: a randomised placebo-controlled double-blind trial. *Br J Obstet Gynaecol* 1999;106:652–657.
- 40. Young GL, Jewell D. Topical treatment for vaginal (thrush) in pregnancy. *Cochrane Database Syst Rev* 2001;4:CD000225.
- 41. Menegola E, Broccia ML, DiRenzo F, Giavini E. Antifungal triazoles induce malformations in vitro. *Reprod Toxicol* 2001;15:421–427.
- 42. Mastoiacovo P, Mazzone T, Botto LD, et al. Prospective assessment of pregnancy outcomes after first-trimester exposure to fluconazole. *Am J Obstet Gynecol* 1996;175:1645–1650.
- 43. Aleck XA, Bartley DL. Multiple malformation syndrome following fluconazole use in pregnancy: report of an additional patient. *Am J Med Genet* 1997:72:253–256.
- 44. Pursley TJ, Blomquist IK, Abraham J, Andersen HF, Bartley JA. Fluconazole-induced congenital anomalies in three infants. *Clin Infect Dis* 1996;22:336–340.
- 45. Chapin RE and Ku WW. The reproductive toxicity of boric acid. *Environ Health Perspect* 1994;102(suppl 7):87–91.
- 46. Hart G. Factors associated with pediculosis pubis and scabies. *Genitourin Med* 1992;68:294–295.
- 47. Scott GR. European guideline for the management of scabies. *Int J STD AIDS* 2001;12(suppl 3):58–61.
- 48. Hollier LM, Workowski K. Treatment of sexually transmitted diseases in women. *Obstet Gynecol Clin North Am* 2003;30:751–775.
- Brown ZA, Benedetti J, Ashley R, et al. Neonatal herpes simplex virus infection in relation to asymptomatic maternal infection at the time of labour. N Engl J Med 1991;324:1247–1252.

- 50. Prober CG, Corey L, Brown ZA, et al. The management of pregnancies complicated by genital infections with herpes simplex virus. *Clin Infect Dis* 1992;15:1031–1038.
- 51. Whitley RJ, Corey L, Arvin A, et al. Changing presentation of herpes simplex virus infection in neonates. *J Infect Dis* 1988;158:109–116.
- 52. Kropp RY, Wong T, Burton S, Embree J, Steben M. Neonatal herpes simplex virus infections in Canada; Valacyclovir HSV Transmission Study Group. Int J STD AIDS 2004;15(suppl 1):2.
- 53. Corey L, Wald A, Patel R, et al. Once-daily valacyclovir to reduce the risk of transmission of genital herpes. *N Engl J Med* 2004;350:11–20.
- Watts DH, Brown ZA, Money D, et al. A double-blind, randomized, placebocontrolled trial of acyclovir in late pregnancy for the reduction of herpes simplex virus shedding and cesarean delivery. *Am J Obstet Gynecol* 2003;188:836–843.
- Brown ZA, Wald A, Morrow RA, Selke S, Zeh J, Corey L. Effect of serologic status and cesarean delivery on transmission rates of herpes simplex virus from mother to infant. *JAMA* 2003;289:203–209.
- 56. Sheffield JS, Hooler LM, Hill JB, Stuart GS, Wendel GD. Acyclovir prophylaxis to prevent herpes simplex virus recurrence at delivery: a systematic review. *Obstet Gynecol* 2003;102:1396–1403.
- 57. Amstey MS, Monif GR. Genital herpesvirus infection in pregnancy. *Obstet Gynecol* 1974;44:394–397.
- 58. Nahmias AJ, Josey WE, Naib ZM, Freeman MG, Fernandez RJ, Wheeler JH. Perinatal risk associated with maternal genital herpes simplex virus infection. *Am J Obstet Gynecol* 1971;110:825–837.
- 59. Smith EM, Ritchie JM, Yankowitz J, et al. Human papillomavirus prevalence and types in newborns and parents: concordance and modes of transmission. *Sex Transm Dis* 2004;31:57–62.
- 60. Armstrong LR, Preston EJ, Reichert M, et al. Incidence and prevalence of recurrent respiratory papillomatosis among children in Atlanta and Seattle. *Clin Infect Dis* 2000;31:107–109.
- 61. Watts DH, Koutsky LA, Holmes KK, et al. Low risk of perinatal transmission of human papillomavirus: results from a prospective cohort study. *Am J Obstet Gynecol* 1998;178:365–373.
- 62. Gall SA. Maternal immunization. *Obstet Gynecol Clin North Am* 2003;30: 623–636.
- 63. Sweet RL. Hepatitis B infection in pregnancy. *Obstet Gynecol Rep* 1990;2: 128–139.
- 64. Ip HM, Lelie PN, Wong VC, Kuhns MC, Reesink HW. Prevention of hepatitis B virus carrier state in infants according to maternal serum levels of HBV DNA. *Lancet* 1989;1:406–410.
- 65. Canadian Immunization Guide. 6th ed. Ottawa, ON: Health Canada; 2002.
- 66. Hepatitis C prevention and control: a public health consensus. Ottawa, Canada, October 14–16, 1998. *Can Commun Dis Rep* 1999;25(suppl 2):1–22.

- Locatelli A, Roncaglia N, Arreghini A, Bellini P, Vergani P, Ghidini A. Hepatitis C virus infection is associated with a higher incidence of cholestasis of pregnancy. *Br J Obstet Gynaecol* 1999;106:498–500.
- 68. Boucher M, Gruslin A, Delage G, et al. The reproductive care of women living with hepatitis C infection. *J SOGC* 2000;96:4–29.
- European Paediatric Hepatitis C Virus Network. Effects of mode of delivery and infant feeding on the risk of mother-to-child transmission of hepatitis C virus. European Paediatric Hepatitis C Virus Network. BJOG 2001;108:371–377.
- 70. Kumar RM, Shahul S. Role of breast-feeding in transmission of hepatitis C virus to infants of HCV-infected mothers. *J Hepatol* 1998;29:191–197.
- 71. Hiratsuka M, Minakami H, Koshizuka S, Sato I. Administration of interferonalpha during pregnancy: effects on fetus. *J Perinat Med* 2000;28:372–376.
- 72. Alter MJ, Hadler SC, Judson FN, et al. Risk factors for acute non-A, non-B hepatitis in the United States and association with hepatitis C virus infection. *JAMA* 1990;264:2231–2235.
- 73. Alter MJ, Coleman PJ, Alexander WJ, et al. Importance of heterosexual activity in the transmission of hepatitis B and non-A, non-B hepatitis. *JAMA* 1989;262:1201–1205.
- 74. Dienstag JL. Sexual and perinatal transmission of hepatitis C. *Hepatology* 1997;26(3 suppl 1):66S–70S.
- 75. Morris DJ. Adverse effects and drug interactions of clinical importance with antiviral drugs. *Drug Saf* 1994;10:281–291.
- 76. Burdge D, Money DM, Forbes JC; Canadian HIV Trials Network Working Group on Vertical HIV Transmission. Canadian consensus guidelines for the management of pregnancy, labour and delivery and for postpartum care in HIV-positive pregnant women and their offspring (summary of 2002 quidelines). CMAJ 2003;168:1671–1674.
- 77. Mandelbrot L, Le Chenadec J, Berrebi A, et al. Perinatal HIV-1 transmission: interaction between zidovudine prophylaxis and mode of delivery in the French Perinatal Cohort. *JAMA* 1998;280:55–60.
- 78. Kind C, Rudin C, Siegrist CA, et al. Prevention of vertical HIV transmission: additive protective effect of elective Cesarean section and zidovudine prophylaxis. Swiss Neonatal HIV Study Group. *AIDS* 1998;12:205–210.
- 79. Elective cesarean-section versus vaginal delivery in prevention of vertical HIV-1 transmission: a randomized clinical trial. European Mode of Delivery Collaboration. *Lancet* 1999;353:1035–1039.
- 80. The mode of delivery and vertical transmission of human immunodeficiency virus type 1 a meta-analysis of 15 prospective cohort studies. The International Perinatal HIV Group. *N Engl J Med* 1999;340:977–987.
- 81. Boucher M, Cohen HR, Gruslin A, Money DM, Steben M, Wong T. Mode of delivery for pregnant women infected by the human immunodeficiency virus. *J SOGC* 2001;101:1–3.

SEXUAL ABUSE IN PERIPUBERTAL AND PREPUBERTAL CHILDREN

Background

Canadian Law regarding Age of Consent to Sexual Activity (at the time of publication)

Canadian law is fairly nuanced in respect to defining the points at which sexual activities involving persons under the age of 18 become criminal offences.¹ Depending on the circumstances, any form of touching for a sexual purpose can constitute an offence. Consent is the key factor in determining whether any form of sexual activity is a criminal offence. The law recognizes some minors as having the ability to consent, in some situations. Generally speaking, persons over 14 are recognized as being able to give consent to participate in sexual activities, unless the activities are taking place in a relationship where one participant has some authority over or is in a position of trust in relation to the other person, where there is dependency, or where there is exploitation of one participant by the other. The *Criminal Code* provides a "close in age" exception: a 12 or 13 year old can consent to engage in sexual activity with another person who is less than two years older and with whom there is no relationship of trust, authority, dependency, or exploitation Children under 12 do not have the legal capacity to consent to any form of sexual activity.

Definition

The definition of sexual abuse varies, but involves all sexual acts that the child cannot comprehend, for which the child is not developmentally prepared and/or cannot give consent to, and/or that violates the law.² Activities may range from fondling to penetration. For the purpose of these guidelines, as is relevant to the potential transmission of sexually transmitted infections (STIs), the definition will include complete or partial penetration by a penis of the mouth, anus and/or vagina, although it is noted that contact of the mouth with the external genitalia or anus could potentially transmit herpes simplex virus (HSV) infections.

In addition, for the purpose of these guidelines, peripubertal refers to individuals aged 11–13 and prepubertal to individuals less than 11 years of age.

Epidemiology

It is difficult to accurately estimate the prevalence of sexual abuse due to underreporting. The reported prevalence varies from study to study, depending on a number of factors. This form of abuse affects children of all ages, socioeconomic classes and geographic locations.³ Some studies estimate that approximately 1% of children experience some form of sexual abuse each year, resulting in sexual victimization of 12–25% of girls and 8–10% of boys by age 18.⁴ The perpetrator may be a member of a child's family or a complete stranger, but in either case the abuser is often an adult male (adolescents may be the perpetrators in as many as 20% of all cases). Boys may be abused as often as girls, but they are less likely to report the abuse.

The Canadian Incidence Study of Reported Child Abuse and Neglect⁶ estimated that 135,573 child maltreatment investigations were carried out in Canada in 1998, an annual incidence rate of 21.52 investigations per 1,000 children. Ten percent (15,614 or 2.48 investigations per 1,000 children) of these investigations involved sexual abuse as the primary reason for investigation. Of these, an estimated 2,742 child investigations involved allegations of oral, vaginal or anal sexual activities. Non-parental figures were most often investigated in sexual abuse cases, with non-parental relatives representing 28%, biological fathers 15% and stepfathers 9% of all cases. Seven percent of sexual abuse investigations involved mothers as alleged perpetrators (5% biological mothers and 2% stepmothers). Sixty-eight percent (~9,813 cases) involved female children, with adolescent females aged 12–15 accounting for 21% of investigations and girls 4–7 years accounting for 23%.

Multiple factors affect the risk of transmission of infection with sexual abuse, including the following:⁶⁻⁹

- Prevalence of STIs within the local population.
- Type of sexual activity: the risk of STI transmission with penile-rectal penetration is greater than penile-vaginal penetration, which is greater than penile-oral penetration etc.
- Degree of trauma: injuries to the genital tract are more common in children.
- Sexual maturity of the child: altered susceptibility to STIs due to maturational differences in the genital tract.
- · Lack of use of barrier contraception.
- · Multiple episodes of abuse.

Prevention

Children should be screened throughout childhood, during routine visits to health care providers' offices, for evidence of sexual abuse. Children who may be at higher risk include those with developmental, behavioural and medical problems. Health care providers should also be aware that recognizing and reporting child sexual abuse is the most effective means of preventing further abuse, reactive abuse and pedophilia. 12-15

Evaluation

Sexually abused children may present in many ways. They may present alone or with their parents for evaluation of alleged sexual abuse. They may present at a health care provider's office with an unrelated complaint and then disclose abuse. The health care provider may even suspect abuse during a routine visit, highlighting the need for vigilance, because abuse may present in ways that may be so non-specific that the problem may not be considered. Rectal or genital bleeding, the presence of STIs and developmentally unusual sexual behaviour are some of the more specific signs of sexual abuse.

Victims of sexual assault may be reluctant to disclose that they have been sexually assaulted for a variety of reasons, including being coerced into secrecy, fear of not

being believed or fear of retribution. In some instances, children may not recognize that abuse has taken place.

Assessment and follow-up of children who are victims of sexual abuse should be carried out with great sensitivity and ideally with the direct involvement of experienced teams or services (see *Appendix G*). When direct referral cannot be made (e.g., in remote areas), every effort should be made to consult with the nearest referral centre.

Health care providers who suspect the occurrence or possibility of sexual abuse should inform the parents/guardians in a calm, non-accusatory manner.² Health care providers must be aware of local reporting requirements (see Reporting and Partner Notification, below).

The health care provider's role is not to conduct a legal interview or obtain details of the abuse from the child, but rather to do the following:²⁰

- 1. Take a pertinent medical history.
- 2. Ensure the physical and emotional well-being of the patient.
- 3. Treat or prevent illness or injury.
- 4. Accurately record spontaneous disclosure or volunteered information.
- Obtain and document physical findings consistent with abuse or suspicions of abuse.
- 6. Inform the child and caregivers about the medical outcome of the investigation.
- 7. Assist child protection and law enforcement agencies in their investigation.

History

When a health care provider suspects abuse, he/she must take a pertinent medical history to satisfy the medical needs of the child and generate adequate information to assist child protection agencies.

When direct referral to specialist referral centre is not possible (e.g., in remote areas), several methods may be used when asking young children about abuse.²¹ The child may also spontaneously provide information. If possible, the child should be interviewed alone, although the presence of a non-threatening caregiver may be appropriate. In addition, the parents/guardians may provide a history of behavioural changes that may be relevant to the situation.

Physical exam

The following information is provided as a guide and may be useful when screening for the possibility of sexual abuse. A full evaluation should ideally be performed by a clinician experienced in this area.

Injuries requiring immediate attention should take precedence over any other examination. The physical examination should be explained to the child before it is performed and should not result in additional emotional trauma.

A complete pediatric examination should be performed, with special attention paid to the growth parameters and sexual development of the child using Tanner staging (see *Appendix H*). Injuries and other evidence of abuse should be documented, including bruising, swelling and areas of tenderness. If the abuse has occurred within 72 hours, or if there is bleeding or acute injury, the examination should be performed immediately so that forensic specimens can be collected.² After 72 hours has passed and when no acute injuries are evident, then the evaluation should be performed when convenient for the child and investigative team.

Careful examination of all areas involved in sexual activity should be performed and notes made of any abnormalities. Examination of the genital and rectal areas may be aided by instruments that illuminate and/or magnify the area. In both sexes, the anus should be examined, and in females the hymenal opening should be examined. Digital and speculum examination is not usually necessary and should not be performed in prepubertal children.

Specimen Collection and Laboratory Diagnosis

For prepubertal children, the decision to perform testing should be done on an individual basis. The following situations put the child at higher risk for STIs and are indications for testing:²²

- The child has symptoms or signs of an STI (e.g., vaginal discharge or pain, genital itching or odour, urinary symptoms, genital ulcers or lesions).
- The suspected assailant is known to have an STI or to be at risk for an STI.
- Another child or adult in the household is known to have an STI.
- The prevalence of STIs in the community is high.
- There is evidence of genital, oral or anal penetration.

If testing is warranted, an experienced clinician (ideally one involved with a referral centre) must be consulted; the testing procedures described below are intended as a guide and for information only.

Minimal investigation should include testing for *Neisseria gonorrhoeae* and *Chlamydia trachomatis*, and if genital ulcers are present, for herpes simplex virus and syphilis. The genital organs of female infants, children and adolescents vary significantly from those of adults, influencing the microbiological flora of the genital tract and sampling sites for screening. Sampling sites must be specific for the sexual maturity of the young person. Speculum examinations should not be performed on prepubertal children.

The health care provider may elect to use several techniques, including the use of small swabs (such as urethral or ear, nose and throat swabs) moistened with sterile saline for transhymenal vaginal sampling. Placing the child in the prone knee-chest position allows cultures to be taken without touching the hymen and causing pain and without the child being alarmed by the sight of the swab.²³ Vulvar or vaginal washings are also suitable (see Table 1).

Table 1. Initial visit: prepubertal children

Table 1. Illitial visit. prepubertal ciliuren		
Specimen type by gender	Condition or organism to be detected	
Males and females Urine • First-catch urine (10–20 mL), after not voiding for 2 hours	 A molecular diagnostic test, preferably a NAAT, should be collected for gonorrhea and chlamydia. This test is generally more sensitive than genital culture and may be acceptable for medico-legal purposes if confirmed by a second set of primers or, in some cases, a second test sent to another laboratory Postexposure NAAT testing can be taken at the time of presentation, without needing to wait for 48 hours after exposure 	
Females Vagina, vestibule or discharge (if present) • 1 urethral swab, premoistened with sterile water (to minimize discomfort)* • Vaginal wash¹ technique preferred to multiple vaginal swabs if NAAT used for Chlamydia trachomatis and Neisseria gonorrhoeae	 Gram stain, if available, for abnormal bacterial flora, bacterial vaginosis, candidiasis, gonorrhea should be taken Molecular diagnostic tests, especially NAATs, are more sensitive than culture for <i>C trachomatis</i> and <i>N gonorrhoeae</i> Cultures have been the preferred method for medicolegal purposes, but NAATs may be acceptable if positive results are confirmed by a second set of primers or, in some cases, a second test sent to another laboratory If available, both tests (culture and NAAT) should be taken If available, wet mount and/or culture for <i>T vaginalis</i> should be taken Since culture tests collected <48 hours after exposure may be falsely negative, they should be repeated 1–2 weeks after exposure if prophylaxis is not offered; a postexposure NAAT can be taken at the time of presentation 	

Table 1. Initial visit: prepubertal children (continued)

Specimen type by gender	Condition or organism to be detected
Males Meatus • 1 urethral swab, premoistened in sterile water for meatal specimen; intraurethral specimen not recommended	 Gram stain for gonococcal urethritis should be taken Molecular diagnostic tests, especially NAATs, are more sensitive than culture for <i>C trachomatis</i> and <i>N gonorrhoeae</i> Cultures have been the preferred method for medicolegal purposes, but NAATs may be acceptable if positive results are confirmed by a second set of primers or, in some cases, a second test sent to another laboratory If available, both tests (culture and NAAT) should be taken If available, wet mount and/or culture for <i>T vaginalis</i> should be taken Since culture tests collected <48 hours after exposure may be falsely negative, they should be repeated 1–2 weeks after exposure if prophylaxis is not offered; a postexposure NAAT can be taken at the time of presentation
Pharynx • 1 swab	 N gonorrhoeae culture should be taken Test for C trachomatis by culture if available; note that organisms can be detected in oropharynx from perinatal transmission for up to 6 months following birth No approved NAAT for throat specimens
Rectum - 1–2 swabs	 N gonorrhoeae and C trachomatis culture should be taken; no approved NAATs at present HSV culture should be taken (if inflammation present)
Genital ulcers • 1 swab	 HSV culture should be taken Treponema pallidum direct test should be taken (see Syphilis chapter)

Table 1. Initial visit: prepubertal children (continued)

Specimen type by gender	Condition or organism to be detected
Serologic specimens	Syphilis Consider screening test(s) for syphilis [†] Syphilis tests should be repeated at 12 and 24 weeks after exposure. In some instances (e.g., high-risk assailant; see the <i>Syphilis</i> chapter) and in areas experiencing outbreaks of syphilis, it may be appropriate to repeat tests 2–4 weeks post-assault
	Hepatitis B • If the child is known to be immune to hepatitis B (HBsAb ≥10 IU/L) or HBsAg-positive, then no testing is required • Baseline antibodies to HBsAg should be collected when hepatitis B immune status is unknown
	 HIV Baseline HIV antibody testing should be collected HIV antibody testing should be repeated at 6, 12 and 24 weeks following significant exposures
	 Hepatitis C Baseline HCV antibody is optional, since transmission of HCV is low via sexual contact. It may be considered if the (alleged) perpetrator(s) is/are at high risk for hepatitis C (e.g., known injection drug user[s]) and significant trauma has occurred with the assault If baseline testing has been performed and is negative, HCV antibody testing should be repeated at 12 and 24 weeks following significant exposures

HBsAb = hepatitis B surface antibody HCV = hepatitis C virus

NAAT = nucleic acid amplification test

HBsAg = hepatitis B surface antigen HSV = herpes simplex virus

- * Vaginal specimens can be taken without a speculum in a relaxed child as long as the hymenal ring is not touched. A small swab (e.g., urethral swab), is preferred. Speculum examination is only rarely required, and in prepubertal females requires consultation with a specialist or may require a general anesthetic.
- † Vaginal washes are performed by instilling 1.5–2 mL of sterile, preservative-free normal saline at room temperature into the vagina via a modification of the method described by Pokorny and Stormer. ^{24,25} The tubing from a 25 mm butterfly needle, with the needle and butterfly wings removed, is inserted into the distal end of a No. 8 bladder catheter. This assembly is then attached to a 3 mL syringe by the end of the butterfly tubing. This system allows for aspiration of the vaginal contents without the end of the butterfly tube becoming occluded by the vaginal walls. The normal saline and vaginal discharge fluid are then aspirated from the vagina.
- ‡ Baseline screening for syphilis should be considered in areas with high prevalence or regional outbreaks of syphilis, foreign-born children, parents/family members/perpetrators diagnosed with syphilis and children diagnosed with another STI.²⁶

All specimens for forensic evidence should be collected by professionals experienced in these procedures and should follow established regional/local protocols (see *Appendix F*). It should be noted that most forensic kits do not contain tests for STIs or blood-borne pathogens. They are useful in the identification of semen or other body fluids, forensic DNA analysis, microscopic hair examination, textile damage assessment and examinations involving fibres and other types of trace evidence. These, in turn, may be used to establish that some form of association occurred between the victim and the accused, that sexual contact occurred and/or that the assault was violent or forceful, thereby indicating lack of consent. All isolates and specimens should be retained in case additional or repeated testing is required.

Table 2. Implications of a diagnosis of STIs for a diagnosis of sexual abuse^{2,9}

Incubation period of infection	Probability of abuse	Mother-to-child transmission
Gonorrhea: 2–7 days	Strong; probable if child <1 year	Can be seen in children from 0–6 months of age
Chlamydia: 1–3 weeks, but up to 6 weeks	Probable; strong if child >3 years	Can be seen in children up to 3 years of age
HSV: 2–14 days	Probable	Can be seen in children up to 3 months of age
Trichomoniasis: 1–4 weeks	Strong if child >6 months	Can be seen in children 0-6 months of age
HPV: ≥1 month	Possible; probable if >2 years	Can be seen in children from 0–2 years of age
Syphilis: up to 90 days	Strong	Must be excluded
HIV: up to 6 months, but the majority seroconvert within 4–12 weeks	Possible	Must be excluded
Hepatitis B: up to 3 months	Possible	Must be excluded

HSV = herpes simplex virus

HPV = human papillomavirus

Management and Treatment

Considerations for prophylaxis

- · Offer prophylaxis if:
 - The patient presents within 48 hours after an assault.
 - It is requested by a parent/patient/guardian.
 - The patient is at high risk for an STI (see Specimen Collection and Laboratory Diagnosis, above).
- It should be noted that the efficacy of antibiotic prophylaxis has not been studied in sexual assault; prophylaxis should be as recommended for treatment of specific infections. See chapters on specific infections for more information.

Table 3. Recommended prophylaxis for uncomplicated urogenital infections (See chapters on specific infections for alternate treatment choices and non-genital infections.)

Sexually transmitted infection	Recommended prophylaxis	
Gonorrhea	 <45 kg: cefixime 8 mg/kg PO in a single dose (max 400 mg PO)*† [A-I] >45 kg: cefixime 400 mg PO in a single dose*† [A-II] 	
Chlamydia	 <45 kg: azithromycin 15 mg/kg P0 in a single dose (max 1 g) [A-I] >45 kg: azithromycin 1 g P0 in a single dose [A-I] 	
Trichomoniasis	 Treat only if positive for trichomonas <45 kg: metronidazole 30 mg/kg/day divided every 6–12 hours for 1 week [B-III] >45 kg: metronidazole 2 g PO as a single dose²⁷ [A-I] 	
Syphilis	 Prophylaxis with azithromycin (given for prophylaxis against chlamydia) is no longer considered to be effective against incubating syphilis in light of the recent emergence of syphilis resistant to azithromycin. Prophylaxis with other agents may be considered if the patient is unlikely to return or there is a potentially high-risk source in an area experiencing an outbreak of infectious syphilis (see <i>Syphilis</i> chapter for more information) If the child subsequently has reactive syphilis serology, he/she should be retreated with a recommended treatment for syphilis 	

Table 3. Recommended prophylaxis for uncomplicated urogenital infections (continued)

Sexually transmitted infection	Recommended prophylaxis
Hepatitis B	 Prophylaxis for hepatitis B should be considered in all cases of sexual assault/abuse where the sexual acts have included anal or vaginal penetration or oral-anal contact without a condom, or if condom status is unknown and the source is not immune to hepatitis B (see Table 1). Oral-genital and oral-oral contact do not appear to be significant modes of transmission²⁸ Recommended prophylaxis as outlined in the <i>Canadian Immunization Guide</i>, 2002,²⁹ includes the following: HBIG 0.06 mL/kg IM up to 14 days after exposure A 3-dose course of hepatitis B vaccine at 0, 1 and 6 months following exposure or on an accelerated schedule
Hepatitis C	No PEP available
HIV	 HIV PEP is recommended when the assailant is known to be HIV-infected and significant exposure has occurred (e.g., oral, anal and/or vaginal penetration without a condom or condom status unknown/broken)³⁰ PEP may also be available on a case-by-case basis for other high-risk exposures (e.g., source a known injection drug user, multiple assailants and/or significant injury) and vaginal, anal or oral penetration has occurred Recommendations vary by province, and the decision to offer PEP should be made in conjunction with a pediatric HIV specialist If HIV PEP is used, it should be started as soon as possible — no later than 72 hours after the assault — and continued for 28 days³⁰

HBIG = hepatitis B immune globulin

PEP = postexposure prophylaxis

^{*} Cefixime should not be given to persons with cephalosporin allergy or a history of immediate and/or anaphylactic reactions to penicillins.

[†] Treatment for gonorrhea should be accompanied by treatment for chlamydia unless a NAAT is negative for chlamydia.

• See the Pregnancy section in the *Sexual Assault in Postpubertal Adolescents* and *Adults* chapter if this is a possibility.

Other management issues

- Appropriate referral should be made as necessary and available (e.g., to child protection authorities, sexual assault teams, local police/Royal Canadian Mounted Police, psychological support, local victim support organizations etc.).
- Consideration should be given to assessing other children in the family or setting where the abuse is thought to have occurred, as it is not unusual to find other children who have also been sexually abused.⁵
- If the patient is sexually active, advise of the need to practice safer-sex or abstain from sexual intercourse until infection has been ruled out and/or prophylaxis is complete.
- Offer tetanus toxoid if relevant (e.g., dirty wounds/abrasions sustained outdoors) and the child's immunization schedule is not up to date.

Reporting and Partner Notification

- Every province and territory has statutes in place that require the reporting of child abuse. Although the exact requirements may differ by province/territory, health care professionals should be aware of local reporting requirements and procedures with respect to child abuse and other acts of maltreatment. If reasonable cause to suspect child abuse exists, local child protection services and/or law enforcement agencies must be contacted promptly.
- An individual with a confirmed notifiable STI should be reported to provincial/ territorial authorities as appropriate.
- Partner notification of individuals found to be infected with an STI should follow the recommendations in the relevant chapter.

Follow-up

- Follow-up tests of cure are recommended for all curable STIs identified in peripubertal and prepubertal children. Follow-up will vary depending on the type of test performed and the type and duration of treatment given. In general, nucleic acid amplification tests should be repeated 3–4 weeks after completion of treatment and culture tests 4–5 days after completion of treatment.
- If no prophylaxis was taken, follow-up should be arranged for 7–14 days after the
 original visit to review available laboratory test results and repeat an STI screen
 to detect infections acquired at the time of the assault that were not detected at
 the initial examination.
- If empiric prophylactic therapy was given, follow-up should be arranged at 3-4 weeks.
- Arrange follow-up serologic testing for HIV, hepatitis B and C, and syphilis as required (see Table 1).

- Review mental state and arrange appropriate referral to mental health services if necessary.
- Psychological and social support should also be offered to affected family members.

References

- 1. Canada Criminal Code, R.S., 1985, c. C-46, s.150.1 153.1
- Guidelines for the evaluation of sexual abuse of children: subject review.
 American Academy of Pediatrics, Committee on Child Abuse and Neglect.
 Pediatrics 1999:103:186–191.
- 3. Muram D. The medical evaluation of sexually abused children. *J Pediatr Adolesc Gynecol* 2003;16:5–14.
- 4. Finkelhor D. *Sourcebook on Sexual Abuse*. Beverly Hills, CA: Sage Publications; 1986.
- 5. Trocmé N, MacLaurin B, Fallon B, et al. *Canadian Incidence Study of Reported Child Abuse and Neglect: Final Report*. Ottawa, ON: Public Health Agency of Canada; 2001. Available at: www.phac-aspc.gc.ca/publicat/cisfr-ecirf/pdf/cis e.pdf. Accessed February 1, 2006.
- 6. Hammerschlag MR. The transmissibility of sexually transmitted diseases in sexually abused children. *Child Abuse Negl* 1998;22:623–635.
- 7. Duncan ME, Tibaux G, Pelzer A, et al. First coitus before menarche and risk of sexually transmitted disease. *Lancet* 1990;335:338–340.
- 8. Greenberg J, Magder L, Aral S. Age at first coitus. A marker for risky sexual behavior in women. *Sex Transm Dis* 1992:19:331–334.
- Thomas A, Forster G, Robinson A, Rogstad K; Clinical Effectiveness Group (Association of Genitourinary Medicine and the Medical Society for the Study of Venereal Disease). National guideline for the management of suspected sexually transmitted infections in children and young people. Sex Transm Infect 2002;78:324–331.
- 10. Fallon MA, Eifler K, Niffenegger JP. Preventing and treating sexual abuse in children with disabilities: use of a team model of intervention. *J Paediatr Nurs* 2002;17:363–367.
- 11. Balogh R, Bretherton K, Whibley S, et al. Sexual abuse in children and adolescents with intellectual disability. *J Intellect Disabil Res* 2001; 45(Pt 3):194–201.
- 12. Andrews G, Gould B, Corry J. Child sexual abuse revisited. *Med J Aust* 2002;176:458–459.
- 13. Jankowski MK, Leitenberg H, Henning K, Coffey P. Parental caring as a possible buffer against sexual revictimization in young adult survivors of child sexual abuse. *J Trauma Stress* 2002;15:235–244.
- 14. Lee JK, Jackson HJ, Pattison P, Ward T. Developmental risk factors for sexual offending. *Child Abuse Negl* 2002;26:73–92.
- 15. Bentovim A. Preventing sexually abused young people from becoming abusers, and treating the victimization experiences of young people who offend sexually. *Child Abuse Negl* 2002;26:661–678.

- Krugman RD. Recognition of sexual abuse in children. Pediatr Rev 1986;8:25–30.
- 17. Adams JA, Harper K, Knudson S, Revilla J. Examination findings in legally confirmed child sexual abuse: it's normal to be normal. *Pediatrics* 1994;94:310–317.
- 18. Johnson CF. Child sexual abuse. Lancet 2004;364:462-470.
- 19. Friedrich WN, Grambsch P. Child sexual behaviour inventory: normative and clinical. *Psychol Assess* 1992;4:303–311.
- Protocol for Medical Examination of the Abused Child. Calgary, AB: Alberta Medical Association; 1998.
- Faller KC. Child Sexual Abuse: Intervention and Treatment Issues. Washington, DC: U.S. Department of Health and Human Services; 1993. Available at: nccanch.acf.hhs.gov/pubs/usermanuals/sexabuse/index.cfm. Accessed February 1, 2006.
- 22. Sexually transmitted diseases treatment guidelines, 2002. Centers for Disease Control and Prevention. *MMWR Recomm Rep* 2002;51(RR-6):1–78.
- 23. Adams J, Alderman E, Konop R, et al. Genital complaints in prepubertal girls, 2004. Available at: www.emedicine.com/ped/topic2894.htm. Accessed February 1, 2006.
- 24. Pokorny SF, Stormer J. Atraumatic removal of secretions from the prepubertal vagina. *Am J Obstet Gynecol* 1987;156:581–582.
- Embree JE, Lindsay D, Williams T, Peeling RW, Wood S, Morris M. Acceptability and usefulness of vaginal washes in premenarchal girls as a diagnostic procedure for sexually transmitted diseases. *Pediatr Infect Dis J* 1996;15:662–667.
- 26. Bays J, Chadwick D. The serologic test for syphilis in sexually abused children. *Adolesc Pediatr Gynecol* 1991;4:148–151.
- 27. Forna F, Gulmezoglu AM. Interventions for treating trichomoniasis in women. *Cochrane Database Syst Rev* 2000;3:CD000218.
- 28. Schreeder MT, Thompson SE, Hadler SC, et al. Hepatitis B in homosexual men: prevalence of infection and factors related to transmission. *J Infect Dis* 1982;146:7–15.
- 29. Canadian Immunization Guide, 6th ed. Ottawa, ON: Health Canada; 2002.
- 30. Smith DK, Grohskopf LA, Black RJ, et al; U.S. Department of Health and Human Services. Antiretroviral postexposure prophylaxis after sexual, injection-drug use, or other nonoccupational exposure to HIV in the United States: recommendations from the U.S. Department of Health and Human Services. *MMWR Recomm Rep* 2005;54(RR-2):1–20.

SEXUAL ASSAULT IN POSTPUBERTAL ADOLESCENTS AND ADULTS

Definition

The definition of sexual assault varies but involves all non-consensual sexual acts, ranging from fondling to penetration. For the purpose of these guidelines, as is relevant to the potential transmission of sexually transmitted infections (STIs), the definition will include complete or partial penetration by a penis of the mouth, anus and/or vagina, although it is noted that contact of the mouth with the external genitalia or anus could potentially transmit herpes simplex virus (HSV) infections.

Epidemiology

Both females and males of any age may be affected by sexual assault. Incidence varies by geographic location and appears, in some studies, to have a seasonal distribution, with peaks occurring in the summer.^{1,2} In the majority of assaults, the victims are young females, but 5–6% of assaults are reported among males.³ Assaults by acquaintances have been estimated to occur at least as often as assaults by strangers and may be underreported.⁴

Canadian data show that 16% of all women (1.7 million) have been involved in at least one incident of sexual or physical assault by a date or boyfriend by the age of 16, and 24% of women 18–24 years had been sexually and/or physically assaulted by a date or boyfriend.⁵ According to Canadian crime statistics, male-against-female violence was the most common type of overall violence but the least likely to involve a stranger.⁶ In 76.8% of reported cases, the woman knew her assailant. In 28.9% of reports, the woman was assaulted by her spouse/ex-spouse.

Gonorrhea, chlamydia and trichomoniasis are the most frequent infections identified in women who give a history of sexual assault.⁷⁻⁹ The peak age incidence of sexual assault victims corresponds with the peak age incidence of many STIs, so their presence does not necessarily indicate acquisition as a result of the assault.⁸

Prevention

Most sexual assaults cannot be prevented, but becoming aware of situations that can make sexual assault more likely and taking preventative steps is of primary importance. Such steps can include measures to remain safe (i.e., at home or while driving), and the avoidance of situations whereby a perpetrator may use alcohol or drugs to impair the victim's ability to resist the assault.

Evaluation

Victims may be reluctant to disclose that they have been sexually assaulted for a variety of reasons, including fear of becoming involved in the criminal system; fear of not being believed or fear of retribution; feelings of guilt, shame or self-blame; or a desire to forget the event. Despite this reluctance to disclose events surrounding the assault, these victims may present for medical attention because of concerns about pregnancy, STIs or injury.¹⁰ In addition, they may present with post-traumatic stress, depressive symptoms, alcohol or substance abuse, or self-harm.¹¹

Assessment and follow-up of sexual assault victims should be carried out with great sensitivity and in conjunction with local teams or services experienced in the management of victims of sexual assault.

Documentation

Clear and complete documentation of history, physical examination findings and specimen collections should be made.

History

History taking should include the date, location and time(s) of the assault(s); what is known about the (alleged) perpetrator(s) (e.g., relationship to the victim, known injection drug use etc.); orifice(s) that have been penetrated and condom use; sexual history before and after the assault; past medical history (e.g., gynecological, menstrual and contraceptive history); current medications; immunization history; if a shower or bath was taken after the assault; if clothing was changed; and available support systems for the patient. Extensive interviewing about the details of the assault should be left to law-enforcement agencies, as this may adversely affect the forensic interview.

Physical exam

Injuries requiring immediate attention should take precedence over any other examination. Ideally, the patient should be asked to disrobe completely, and if forensic specimens are to be collected, this should be done while standing on an open sheet (to collect evidence that may fall off). All clothing worn during the assault should be collected in separate labelled plastic bags. The patient should put on a gown so that a complete examination for bruises and other injuries can be performed. All injuries (including those seen on genital examination) should be accurately documented on body-map diagrams. It is important to look for petechial hemorrhages on the palate if there was a history of forced oral penetration. Colposcopy and photography rarely provide any useful information and may produce unnecessary distress.^{7,12}

Specimen Collection and Laboratory Diagnosis

The decision to obtain genital or other specimens for the diagnosis of STIs or blood-borne pathogens (BBPs) should be made on a case-by-case basis. Since baseline diagnostic testing for STIs and BBPs facilitates optimum medical management of the victim, this is strongly recommended whenever possible. It may be appropriate, however, to inform the individual that the results of any test for an STI will become part of his/her medical record, and in the case of a sexual assault could be brought into evidence in a court proceeding.

Wherever possible, baseline screening for common STIs should be performed due to the significant incidence of pre-existing STIs among women who present after sexual assault and the smaller but significant incidence of acquisition of STIs resulting from rape. Baseline testing also facilitates recommended follow-up (e.g., test of cure in pregnant women) if an STI is identified. When it is not possible to screen for all STIs, a minimal investigation should include testing for *Neisseria gonorrhoeae* and *Chlamydia trachomatis*.

Speculum examination should be performed in females, including postpubertal females, whenever possible. If it is not possible to pass a speculum, blind vaginal sampling, together with urethral and/or urine nucleic acid amplification tests (NAATs), are advised.

Wherever possible, the (alleged) perpetrator(s) should also be screened.

All specimens for forensic evidence should be collected by professionals experienced in these procedures and should follow established regional/local protocols (see *Appendix F*). It should be noted that most forensic kits do not contain tests for STIs or BBPs. They are useful in the identification of semen or other body fluids, forensic DNA analysis, microscopic hair examination, textile damage assessment and examinations involving fibres and other types of trace evidence. These, in turn, may be used to establish that some form of association occurred between the victim and the accused, that sexual contact occurred and/or that the assault was violent or forceful, thereby indicating lack of consent. All isolates and specimens should be retained in case additional or repeated testing is required.

Table 1. Initial visit: postpubertal children/adolescents/adults

Sexually transmitted infection	Recommended specimen type
Gonorrhea (see Gonococcal Infections chapter)	 Gram stain (for Gram-negative intracellular diplococci) if available, should be taken Culture from all penetrated (partially or fully) orifice(s) and urethra in males and females should be taken A molecular diagnostic test, preferably a NAAT, should also be performed on specimens collected from the urethra (males), endocervix/urethra (females), urine (males and females), as appropriate. This test is generally more sensitive than genital culture and may be acceptable for medicolegal purposes if confirmed by a second set of primers or, in some cases, a second test sent to another laboratory. Note that a NAAT should not be performed on pharyngeal specimens, and referral to the manufacturer's guidelines is recommended for testing of rectal specimens Since culture tests collected <48 hours after exposure may be falsely negative, they should be repeated in 1–2 weeks after exposure if prophylaxis is not offered; a postexposure NAAT can be taken at the time of presentation
Chlamydia (see <i>Chlamydial</i> <i>Infections</i> chapter)	 Molecular diagnostic tests, especially NAATs, are more sensitive than culture and should be performed whenever possible on urine (males and females), urethral (males) or cervical (females) specimens. Urine testing may make testing more acceptable to some individuals Cultures have been the preferred method for medico-legal purposes, but NAATs may be acceptable if the positive results are confirmed by a second set of primers or, in some cases, a second test sent to another laboratory. NAATs have not been adequately evaluated for throat and rectal specimens If available, both tests (culture and NAAT) should be performed Since culture tests collected <48 hours after exposure may be falsely negative, they should be repeated 1–2 weeks after exposure if prophylaxis is not offered; a postexposure NAAT can be taken at the time of presentation
Trichomoniasis	If available, wet mount and/or culture for <i>Trichomonas vaginalis</i>
Syphilis (see <i>Syphilis</i> chapter)	 A non-treponemal test (e.g., RPR, VDRL) and a treponemal test (e.g., TP-PA) should be performed Both the treponemal and non-treponemal tests should be repeated at 12 and 24 weeks after exposure. In some instances (e.g., a high-risk assailant; see <i>Syphilis</i> chapter) and in areas experiencing outbreaks of syphilis, it may be appropriate to repeat tests 2–4 weeks post-assault

Table 1. Initial visit: postpubertal children/adolescents/adults (continued)

Sexually transmitted infection	Recommended specimen type
Hepatitis B	 If the individual is known to be immune to hepatitis B (HBsAb ≥10 IU/L) or HBsAg-positive, then no testing is required Baseline antibodies to HBsAg should be collected when hepatitis B immune status is unknown
HIV	 Baseline HIV antibody testing should be performed HIV antibody testing should be repeated at 6, 12 and 24 weeks following significant exposures
Hepatitis C	 Baseline HCV antibody is optional, since transmission of HCV is low via sexual contact. Testing may be considered if the (alleged) perpetrator(s) is/are at high risk for hepatitis C (e.g., known injection drug user[s]) and significant trauma has occurred with the assault If baseline testing performed and is negative, HCV antibody testing should be repeated at 12 and 24 weeks following significant exposures

HBsAb = hepatitis B surface antibody
HBsAg = hepatitis B surface antigen
HCV = hepatitis C virus
NAAT = nucleic acid amplification test
RPR = rapid plasma reagin
TP-PA = Treponema pallidum particle agglutination
VDRL = Venereal Disease Research Laboratory

Management and Treatment

Considerations for prophylaxis

- · Offer prophylaxis if:
 - Unsure that the patient will be returning for follow-up.
 - It is known that the assailant is infected with a specific STI.
 - It is requested by the patient/parent/guardian.
 - The patient has signs or symptoms of an STI.
- In addition, it may be appropriate to routinely offer prophylaxis in situations where vaginal, oral or anal penetration has occurred, because most sexual assault victims do not return for follow-up visits.^{8,13,14}
- It should be noted that the efficacy of antibiotic prophylaxis has not been studied in sexual assault; prophylaxis should be as per recommendations for treatment of specific infections (see chapters on specific infections for more information).

Table 2. Recommended prophylaxis for uncomplicated urogenital infections

(See chapters on specific infections for alternative treatment choices and non-genital infections.)

Sexually transmitted infection	Recommended prophylaxis
Gonorrhea	 Non-pregnant adults Cefixime 400 mg PO in a single dose* [A-I] OR Ciprofloxacin 500 mg PO in a single dose¹ (unless not recommended due to quinolone resistance) [A-I] Pregnant adults Cefixime 400 mg PO in a single dose [A-I]
Chlamydia	 Non-pregnant adults Azithromycin 1 g PO in a single dose if poor compliance is expected [A-I] OR Doxycycline 100 mg PO bid for 7 days [A-I] Pregnant adults Amoxicillin 500 mg PO tid for 7 days [B-I] OR Azithromycin 1 g PO in a single dose if poor compliance is expected [B-I]
Trichomonas	 Treat only if test is positive for trichomonas All adults: metronidazole 2 g PO in a single dose¹⁵ [A-I]
Syphilis	 Prophylaxis with azithromycin (given for prophylaxis against chlamydia) is no longer considered to be effective against incubating syphilis in light of the recent emergence of syphilis resistant to azithromycin. Prophylaxis with other agents may be considered if the patient is unlikely to return or there is a potentially high-risk source in an area experiencing an outbreak of infectious syphilis (see <i>Syphilis</i> chapter for more information) If the recipient subsequently has reactive syphilis serology, he/she should be retreated with a recommended treatment agent for syphilis

Table 2. Recommended prophylaxis for uncomplicated urogenital infections (continued)

Sexually transmitted infection	Recommended prophylaxis
Hepatitis B	 Prophylaxis for hepatitis B should be considered in all cases of sexual assault/ abuse where the sexual acts have included anal or vaginal penetration or oral-anal contact without a condom or condom status is unknown and the source is not immune to hepatitis B (see Table 1). Oral-genital and oral-oral contact do not appear to be significant modes of transmission¹⁶ Recommended prophylaxis as outlined in the <i>Canadian Immunization Guide</i>, 2002¹⁷ includes the following: HBIG up to 14 days after exposure A 3-dose course of hepatitis B vaccine at 0, 1 and 6 months following exposure or accelerated schedule as appropriate
Hepatitis C	No PEP available
HIV	 HIV PEP is recommended when the assailant is known to be HIV-infected and significant exposure has occurred (e.g., oral, anal, and/or vaginal penetration without a condom or condom status unknown/broken)¹⁸ PEP may also be available on a case-by-case basis for other high-risk exposures (e.g., source a known injection drug user, multiple assailants and/or significant injury) and vaginal, anal or oral penetration has occurred Recommendations vary by province, and the decision to offer PEP should be made in conjunction with an HIV specialist and/or provincial/territorial/regional protocols If HIV PEP is used, it should be started as soon as possible — no later than 72 hours after exposure — and continued for 28 days¹⁸

HBIG = hepatitis B immune globulin

PEP = postexposure prophylaxis

- * Cefixime and ceftriaxone should not be given to persons with cephalosporin allergy or a history of immediate and/or anaphylactic reactions to penicillins.
- † Quinolones are not recommended if the case or contact are from, or are epidemiologically linked to, any area with rates of quinolone-resistant N gonorrhoeae > 3-5%:
 - Asia

- · Pacific Islands (including Hawaii)
- India

- · Australia
- · United Kingdom
- · Regions of the United States (check with the U.S. Centers for Disease Control and Prevention for rates of quinolone resistance by geographic area)
- · MSM with contact or epidemiologically linked to the United States
- · Areas in Canada experiencing high rates of quinolone resistance; current numbers provided by the National Microbiology Laboratory place Quebec, Ontario, Alberta and British Columbia above the 3% threshold for quinolone resistance. Please check with your local public health officials to learn about quinolone resistance in your area. In Alberta all ciprofloxacin resistant cases in 2004-05 were in MSM or linked to travel outside of Alberta, therefore ciprofloxacin remains a recommended agent for the treatment of gonorrhea in Alberta except in these situations (source: Alberta Health and Wellness STD Services). For data on national quinolone resistance in Canada, please visit the Public Health Agency of Canada website (www.phac-aspc.qc.ca). For more information see *Gonococcal Infections* chapter.

 If pregnancy is a possible result of the assault, the emergency contraceptive pill (ECP) should be considered:¹⁹

Preferred	Alternative
Plan B: levonorgestrel 1.5 mg PO as a single dose	• levonorgestrel 0.75 mg PO bid x 2 doses if a single dose is not likely to be tolerated

- Treatment should be taken as soon as possible, up to 72 hours after exposure (efficacy declines after this, but some benefit may be achieved up to 120 hours after exposure).
- The ECP is more effective and better tolerated than the Yupze method.²⁰
- The ECP is contraindicated if there is evidence of an established pregnancy as confirmed by a positive pregnancy test.
- For the two-dose regimen, Gravol 50 mg given 30 minutes before the second dose of levonorgestrel may prevent vomiting.

Other management issues

- If the patient consents, appropriate referral should be made as necessary and
 as available (e.g., to sexual assault teams, local police/Royal Canadian Mounted
 Police, psychological support, local victim support organizations etc.). Advise of
 the need to practice safer sex or abstain from sexual intercourse until infection
 has been ruled out and/or prophylaxis is complete.
- Offer tetanus toxoid if relevant (e.g., dirty wounds/abrasions sustained outdoors).

Reporting and Partner Notification

- Every province and territory has statutes in place that require the reporting of child abuse. Although the exact requirements may differ by province/territory, health professionals should be aware of local reporting requirements and procedures with respect to child abuse and other acts of maltreatment. If reasonable cause to suspect child abuse exists, local child protection services and/or law enforcement agencies should be contacted.
- An individual with a confirmed notifiable STI should be reported to provincial/ territorial authorities as appropriate.
- Partner notification of individuals found to be infected with an STI should follow the recommendations in the relevant chapter.

Follow-up

If no prophylaxis was taken, follow-up should be arranged for 7–14 days after
the original visit to review available laboratory test results and to repeat an STI
screen to detect infections acquired at the time of the assault that were not
detected at the initial examination.

- Test of cure for specific infections should follow recommendations outlined in the relevant chapters.
- If empiric prophylactic therapy was given, follow-up should be arranged at 3-4 weeks.
- Arrange follow-up serologic testing as required (see Table 1).
- Review mental state and arrange appropriate referral to mental health services if necessary.

References

- 1. Everett RB, Jimerson GK. The rape victim: a review of 117 consecutive cases. *Obstet Gynecol* 1977;50:88–90.
- 2. Michael RP, Zumpe D. Sexual violence in the United States and the role of season. *Am J Psychiatry* 1983;140:883–886.
- 3. Anderson CL. Males as sexual assault victims: multiple levels of trauma. *J Homosex* 1982;7:145–162.
- Schwarcz SK, Whittington WL. Sexual assault and sexually transmitted diseases: detection and management of adults and children. *Rev Infect Dis* 1990;12(suppl 6):S682–690.
- 5. Johnson H. *Dangerous Domains: Violence against Women in Canada.* Toronto, ON: Nelson; 1996.
- 6. Canadian Crime Statistics 2000. Ottawa, ON: Statistics Canada; 2000.
- Estreich S, Forster GE, Robinson A. Sexually transmitted diseases in rape victims. Genitourin Med 1990;66:433–438.
- 8. Jenny C, Hooton TM, Bowers A, et al. Sexually transmitted diseases in victims of rape. *N Engl J Med* 1990;322:713–716.
- Lacey HB. Sexually transmitted diseases and rape: the experience of a sexual assault centre. Int J STD AIDS 1990;1:405–409.
- 10. Mein JK, Palmer CM, Shand MC, et al. Management of acute adult sexual assault. *Med J Aust* 2003;178:226–230.
- 11. Petter LM, Whitehill DL. Management of female sexual assault. *Am Fam Physician* 1998;58:920–926, 929–930.
- 12. Bowyer L, Dalton ME. Female victims of rape and their genital injuries. *Br J Obstet Gynaecol* 1997;104;617–620.
- 13. Forster GE, Pritchard J, Munday PE, Goldmeier D. Incidence of sexually transmitted diseases in rape victims during 1984. *Genitourin Med* 1986;62:267–269.
- 14. Tintinalli JE, Hoelzer M. Clinical findings and legal resolution in sexual assault. *Ann Emerg Med* 1985;14:447–453.
- 15. Forna F, Gulmezoglu AM. Interventions for treating trichomoniasis in women. *Cochrane Database Syst Rev* 2000;3:CD000218.
- 16. Schreeder MT, Thompson SE, Hadler SC, et al. Hepatitis B in homosexual men: prevalence of infection and factors related to transmission. *J Infect Dis* 1982:146:7–15.
- 17. Canadian Immunization Guide, 6th ed. Ottawa, ON: Health Canada; 2002.

- 18. Smith DK, Grohskopf LA, Black RJ, et al; U.S. Department of Health and Human Services. Antiretroviral postexposure prophylaxis after sexual, injection-drug use, or other nonoccupational exposure to HIV in the United States: recommendations from the U.S. Department of Health and Human Services. *MMWR Recomm Rep* 2005;54(RR-2):1–20.
- 19. Dunn S, Guilbert E. Emergency contraception. *SOGC Clinical Practice Guidelines* 2003;131:1–7. Available at: www.sogc.org/guidelines/pdf/ps131.pdf. Accessed February 1, 2006.
- Randomised controlled trial of levonorgestrel versus the Yupze method of combined oral contraceptives for emergency contraception. Task Force on Postovulatory Methods of Fertility Regulation. *Lancet* 1998;352:428–433.

SEX WORKERS

Definition

Sex workers are female, male or transgendered adults or young people who receive money, shelter or goods in exchange for sexual services, either regularly or occasionally, and who may or may not consciously define those activities as income-generating. There are no reliable verbal or visual clues as to whether a patient is a sex worker or not. Where appropriate, patients should be asked whether they have ever received money, shelter or goods in exchange for sexual services.

Epidemiology

Sex workers are vulnerable to sexually transmitted infections (STIs), including HIV, because of the following factors:

- Lack of control (e.g., condom use, refusing clients).
- Lifestyle risks, such as violence, substance use and mobility.¹
- Stigmatization and marginalization.
- · Limited economic options.
- Limited access to health, social and legal services.
- Limited access to information about and the means of prevention.
- · Gender-related differences and inequalities.
- Sexual abuse and exploitation, including trafficking and child prostitution.
- Legislation and policies affecting the rights of sex workers.
- Mental health problems.
- Incarceration.
- Lack of family and social support.

Because of high rates of partner change, sex workers play an important role in the transmission of STIs, especially of those with short periods of infectiousness, such as syphilis and gonorrhea.² Studies from developed and developing countries have shown increased STI and HIV incidence and prevalence among sex workers.²⁻¹⁰ Sex workers tend to use condoms less often with regular partners, even though these individuals are often at high risk for STIs and HIV themselves. Adolescents and children who work in the sex trade are especially vulnerable to STIs due to the cellular immaturity of the female vagina and cervix, as well as an inferior ability to negotiate for safer sex and higher risk of violence and abuse.¹¹

Prevention

Successful STI/HIV prevention focuses on the promotion of safer sexual behaviour through female and male condom availability and correct usage; improved negotiating skills; and supportive policies and laws.¹⁻³ Peer education, outreach work, accessible services, advocacy, community development, program coordination and sex worker involvement are all important prevention principles and strategies.^{1-3,12-15}

Lubricants have been associated with a reduced risk for STIs.¹⁶ Spermicides such as nonoxynol-9 have been linked to enhanced susceptibility to infection and have not been shown to enhance the protective effect of condoms.¹⁷ Hepatitis B vaccination should be available to sex workers, since they are at increased risk for infection.¹⁸ Hepatitis A vaccination should be available to sex workers at high risk, such as male sex workers who engage in oral-anal contact with male customers.

Evaluation

Sex workers presenting for STI care or a routine medical examination should have an STI/HIV history taken and undergo a physical examination focusing on the genital area, including a speculum exam for women and a throat and rectal exam if indicated. Privacy and confidentiality must be assured. STI/HIV evaluation of sex workers cannot always be performed in ideal clinic conditions; it may need to be adapted to less structured settings, such as mobile clinics. In addition to the standard STI/HIV evaluation, it is important to ask about current or past drug use, whether there is a regular partner and whether there is condom use with both customers and partners. 19

Specimen Collection and Laboratory Diagnosis

History, examination and setting should determine specimen collection. With counselling and informed consent, sex workers should receive regular laboratory screening for syphilis, HIV infection (unless known to be HIV-positive), gonorrhea, chlamydia, vaginitis/vaginosis and HPV infection (if available). Pegular cervical screening for dysplasia and/or HPV infection is important. Persons at risk for hepatitis C should be counselled and tested.

Because of the nature of sex work and the social situation of many sex workers, urine-based laboratory testing, rapid point-of-care testing and self-collected specimens are especially relevant.

Management and Treatment

Sex workers should be able to access standard STI and HIV/AIDS management and treatment recommendations.²⁰ Curing a single sex worker of gonorrhea can result in fewer secondary cases and reduce the risk of HIV, thereby saving 120 disability-adjusted life years (DALYs) at a cost below US\$1 per DALY.²¹ Singledose, oral therapies for STIs should be available to sex workers who are unable to complete a longer course of treatment. Epidemiological or syndromic treatment without a full evaluation or laboratory testing may sometimes be necessary.^{1,19}

Education and counselling is a vital component of STI/HIV management for sex workers, as well as for other patients.^{1,19} It is especially important that sex workers know how to use condoms, how to negotiate for safer-sex and why they should use condoms with regular partners. Clinicians need to understand the specific circumstances of risk for each patient and develop an individualized risk reduction plan for him/her.

Reporting and Partner Notification

STI/HIV surveillance is important, and accurate and prompt reporting is the basis of effective STI surveillance and case management. Sex workers and other marginalized populations often rely on publicly funded STI/HIV services, so to facilitate case management and cooperation with patient reporting, trust, respect and confidentiality should be emphasized in these situations.

Sex worker partners (both regular and commercial) need to be notified in a confidential manner and offered treatment in the same manner as non–sex workers. The possibility that partner notification may result in violence toward the index sex worker should be explored and mitigated where possible. In this instance, notification by the health department or a health care worker (keeping the identity of the sex worker anonymous) is often preferable.

Follow-up

Sex workers should be encouraged to have a regular monthly STI evaluation.¹⁹ Children and youth who may have been exploited should be reported to the relevant youth protection agency (see *Sexual Abuse in Peripubertal and Prepubertal Children* chapter). Sex workers with mental health, social service, housing or legal issues need to be referred to appropriate agencies or practitioners.

References

- Sex Work and HIV/AIDS. Geneva, Switzerland: Joint United Nations Programme on HIV/AIDS; 2002.
- Plummer FA, Coutinho RA, Ngugi EN, Moses S. Sex workers and their clients in the epidemiology and control of sexually transmitted diseases. In: Holmes KK, Sparling PF, Mardh P-A, et al, eds. Sexually Transmitted Diseases. 3rd ed. New York, NY: McGraw-Hill; 1999:143–150.
- Ngugi EN, Branigan E, Jackson DJ. Interventions for commercial sex workers and their clients. In: Gibney L, DiClemente RJ, Vermund SH, eds. *Preventing HIV in Developing Countries*. New York, NY: Kluwer Academic/Plenum Publishers; 1999:205–229.
- Roy E, Haley N, LeClerc P, et al. Prevalence of HIV infection and risk behaviours among Montreal street youth. *Int J STD AIDS* 2000;11:241–247.
- Uribe-Salas F, Hernandez-Avila M, Juarez-Figueroa L, Conde-Glez CJ, Uribe-Zuniga P. Risk factors for herpes simplex type 2 infection among female commercial sex workers in Mexico City. *Int J STD AIDS* 1999;10:105–111.
- 6. Tsunoe H, Tanaka M, Nakayama H, et al. High prevalence of Chlamydia trachomatis, Neisseria gonorrhoeae and Mycoplasma genitalium in female commercial sex workers in Japan. *Int J STD AIDS* 2000;11:790–794.

- Desai VK, Kosambiya JK, Thakor HG, Umrigar DD, Khandwala BR, Bhuyan KK. Prevalence of sexually transmitted infections and performance of STI syndromes against aetiological diagnosis, in female sex workers of red light area in Surat, India. Sex Transm Infect 2003;79:111–115.
- 8. Zachariah R, Spielmann MP, Harries AD, Nikhoma W, Chantulo A, Arendt V. Sexually transmitted infections and sexual behaviour among commercial sex workers in a rural district of Malawi. *Int J STD AIDS* 2003;14:185–188.
- Estcourt CS, Marks C, Rohrsheim R, Johnson AM, Donovan B, Mindel A. HIV, sexually transmitted infections, and risk behaviours in male commercial sex workers in Sydney. Sex Transm Infect 2000;76:294–298.
- Poulin C, Alary M, Bernier F, Carbonneau D, Boily MC, Joly JR. Prevalence of Chlamydia trachomatis and Neisseria gonorrhoeae among at-risk women, young sex workers, and street youth attending community organizations in Quebec City, Canada. Sex Transm Dis 2001;28:437–443.
- 11. Willis BM, Levy BS. Child prostitution: global health burden, research needs, and interventions. *Lancet* 2002;359:1417–1422.
- 12. Basuki E, Wolffers I, Deville W, et al. Reasons for not using condoms among female sex workers in Indonesia. *AIDS Educ Prev* 2002;14:102–116.
- 13. Ibbitson M. Out of the sauna: sexual health promotion with "off street" sex workers. *J Epidemiol Community Health* 2002;56:903–904.
- Sanchez J, Campos PE, Courois B, et al. Prevention of sexually transmitted diseases (STDs) in female sex workers: prospective evaluation of condom promotion and strengthened STD services. Sex Transm Dis 2003;30:273–279.
- 15. Morton AN, Wakefield T, Tabrizi SN, Garland SM, Fairley CK. An outreach programme for sexually transmitted infection screening in street sex workers using self-administered samples. *Int J STD AIDS* 1999;10:741–743.
- Rojanapithayakorn W, Goedken J. Lubrication use in condom promotion among commercial sex workers and their clients in Ratchaburi, Thailand. J Med Assoc Thai 1995;78:350–354.
- 17. Forbes A, Heise L. What's up with nonoxynol-9? *Reprod Health Matters* 2000;8:156–159.
- 18. Mak R, Traen A, Claeyssens M, Van Renterghem L, Leroux-Roels G, Van Damme P. Hepatitis B vaccination for sex workers: do outreach programmes perform better? *Sex Transm Infect* 2003;79:157–159.
- 19. Guidelines for the Management of Sexually Transmitted Infections in Female Sex Workers. Geneva, Switzerland: World Health Organization; 2002.
- 20. Guidelines for the Management of Sexually Transmitted Infections. Geneva, Switzerland: World Health Organization; 2001.
- 21. *The Public Health Approach to STD Control.* Geneva, Switzerland: Joint United Nations Programme on HIV/AIDS; 1998.

SUBSTANCE USE

The objective of this chapter is to provide an overview of substance use issues as they pertain to the prevention, management and treatment of sexually transmitted infections (STIs). Additional sources of information^{1,2} can provide a more comprehensive overview of substance use prevention and treatment in general.

Definition

Substance use may be for medicinal or non-medicinal purposes and may be done legally or illegally. It occurs along a continuum from experimental use to harmful use and dependence:³

- No use: the person does not use alcohol or other drugs.
- Experimental use: the person tries a substance out of curiosity and may or may not use the drug again.
- Social or occasional use: the person uses the drug in an amount or frequency that is not harmful (e.g., to health, family, school or work).
- Harmful use: the person experiences negative consequences of drug use (e.g., health, family, school, work or legal problems).
- Dependence: the person is psychologically and/or physically dependent on a drug, which is used excessively, and continues use despite experiencing serious problems.

Epidemiology

- The 2002 Canadian Community Epidemiology Network on Drug Use national report on drug trends in Canada indicates that self-reported alcohol use is rising for both males and females, with an average of 20.2% of Canadians (29.0% male and 11.4% female) reporting frequent heavy alcohol use (five or more drinks on one occasion, 12 or more times a year).⁴
- Cannabis is the most frequently used illicit drug among Canadian youth and adults, with 18.6% of respondents reporting lifetime use; 3.6% report ever using LSD, speed or heroin; and 2.7% report cocaine use.⁴
- There are approximately 50,000 to 100,000 injection drug users in Canada, with concentration in Vancouver, Montreal and Toronto.^{5,6} In 2002, 24% of positive HIV tests reported to the Centre for Infectious Disease Prevention and Control were attributable to injection drug use (IDU).⁷
- Aboriginal Canadians and street youth are at greater risk for and have higher rates of alcohol and illicit substance abuse than other Canadians.⁵
- Although there are few data available on the abuse of solvents in Canada, there
 is particular concern about solvent abuse among Aboriginal youth.⁴

- The use of alcohol and illicit drugs is associated with risky sexual behaviour. Alcohol and illicit drug use, especially the use of crack cocaine⁸⁻¹³ and methamphetamine,^{9,10} are associated with poor and inconsistent condom use,^{9,11,13-19} sex with multiple partners,^{9,10,13-21} early sexual debut,^{20,22} trading sex,^{10,11,14,15,18,19} buying sex,²³ sex with known injection drug users,¹⁹ lower condom use self-efficacy or perceived ability to use condoms,¹⁶ and lower HIV-related knowledge.¹⁶
- Substance use has also been linked to elevated hepatitis C^{24,25} and STI transmission,¹⁹⁻²³ including herpes simplex virus type 2,²¹⁻²⁴ hepatitis B,²⁴ trichomoniasis,^{20,26} syphilis,^{24,27} HIV,^{19,24,27} chlamydia^{20,24,26,27} and gonorrhea.^{20,24,26,27}
- Users of more stigmatized substances, such as injection drugs and crack, are at higher behavioural risk for STIs than users of less stigmatized drugs, such as marijuana.²⁸
- Youth who abuse substances are more likely to engage in risky sexual behaviour and continue these risky behaviours and drug use into adulthood.^{17,29}
- The use of recreational drugs among men who have sex with men (MSM) has risen in recent years and has been linked to increases in risky sexual behaviour and rising STI rates (see Men Who Have Sex with Men/Women Who Have Sex with Women chapter). Sildenafil citrate (Viagra), vardenafil (Levitra) or tadalafil (Cialis) can be used to counteract the erectile dysfunction side effect of some of these drugs, a practice that has been linked to multiple sex partners and STI acquisition.^{37,38}

Prevention

While the elimination of harmful substance use is the ideal approach to preventing substance use reducing the associated STI risk, this can be a difficult if not unattainable goal, especially when substance dependence has already developed. For substance users, substance abstinence should not be used as the exclusive focus of substance use or STI-prevention efforts and should not be a requirement for using STI treatment services. Two prevention strategies are recommended, depending on the patient's placement on the risk continuum:³⁹

- Risk avoidance: to avoid or prevent the adoption of risk behaviours among non-users and low-risk users (e.g., people of legal drinking age who drink at low or moderate levels).
- Harm or risk reduction: to encourage the adoption of acceptable behavioural change, no matter how small, to reduce, if not eliminate, risk (e.g., using clean needles from a needle exchange, cessation of needle sharing).

A harm reduction approach is non-judgmental and takes into account individual needs and a number of potential approaches when discussing realistic personal risk reduction goals. Some potential harm-reduction strategies related to substance use include the following:

- Abstaining from one or more drugs for a limited or open time period.
- Decreasing the frequency and/or amount of a substance used.
- Switching to lower-risk substances and delivery methods (e.g., methadone, cannabis).
- Separating substance use from driving, working and other tasks.
- Creating a safer drug use environment (where, when and with whom; safer purchases/possession; use of needle-exchange programs; and safer injection sites).
- Considering treatment, rehabilitation, detoxification, counselling or support programs.
- Developing a trusting relationship with a health care professional to help monitor physical and mental health.
- Learning about overdose prevention and response.
- · Addressing nutritional needs and ways to improve nutrition.

Harm reduction strategies specific to injection drug users include safer injection practices:40

- Use a new needle and syringe for each injection.
- If sharing cannot be avoided, clean the syringe properly before use:40
 - Fill syringe completely with clean water, and shake vigorously for 30 seconds.
 Squirt water out.
 - Fill syringe with full-strength (undiluted) bleach, leave in for at least
 30 seconds, and shake vigorously. Squirt bleach out. Do this at least twice,
 using fresh bleach each time.
 - Rinse bleach from syringe by repeating the first step at least twice, using clean water each time.
- Avoid sharing vials, cotton and spoons, as well as recapping the needles
 of others.
- Before shooting up, always clean the injection site with a sterile alcohol swab, rubbing alcohol, aftershave lotion (which contains alcohol) or soap and water.
- Sterilize spoons with an alcohol swab or bleach and water before each use.
- Mix drugs using sterile water or, if this is not available, water that has been
 recently boiled. To remove impurities from the mix, it is best to fill the syringe by
 drawing the liquid through a cotton filter (or a piece torn from an alcohol swab).

STI prevention should be discussed within the context of potential influences on sexual behaviour, including substance use, and should also focus on harm reduction (see *Primary Care and Sexually Transmitted Infections* chapter). For substance users with poor condom practices, skill-building around condom use and negotiation may help to improve condom use.⁴¹ A motivational interviewing approach for prevention counselling can help promote harm reduction behaviours (see *Primary Care and Sexually Transmitted Infections* chapter).

Because involvement in illicit drug use is a risk factor for hepatitis A virus (HAV) and hepatitis B virus (HBV) infection, and because vaccination coverage for this population is poor, HAV and HBV vaccination is recommended for injection drug users. HAV vaccination is also recommended for those involved in oral drug use in unsanitary conditions⁴² (see *Hepatitis B Virus Infections* chapter).

- As self-reported HBV infection and immunization status among both injection and non-injection drug users may not be accurate,⁴³ vaccination should be offered to all in these groups.
- To maximize reach in high risk populations beyond primary care settings, immunization for HBV and HAV can be successfully delivered in non-traditional settings (e.g., public health nursing outreach to geographic areas with high rates of substance use).⁴⁴

Note:

According to the *Canadian Immunization Guide*,⁴² pre-immunization testing for immunity against HAV should be considered for populations with the potential for higher levels of pre-existing immunity. Routine pre-immunization serologic screening for HBsAg, anti-HBs or anti-HBc is recommended for people at high risk of infection, but is not practical for universal immunization programs.

Evaluation

- Evaluation of current and past substance use is an important component of STI risk assessment (see *Primary Care and Sexually Transmitted Infections* chapter). Table 1 outlines the six main elements of a substance use history, including sexual risk associated with substance use and possible questions for each element.
- The word use carries no value judgment, but abuse does. Asking about substance use is more likely to lead to an open, honest answer.
- Elicit information on legal drug use, illegal drug use and harmful use of drugs sold for medicinal purposes.
- When assessing substance use as part of the STI risk assessment, use language
 that will be easily understood. Becoming familiar with the terms used in your
 region can help you to effectively communicate. Table 2 provides a quick
 reference for frequently used substances, street names and possible modes
 of delivery.

Table 1. Main elements of taking a substance use history⁴⁵

Main element	Possible questions
Substance/alcohol use	Do you or have you ever used drugs? What drugs do you use? How often do you use drugs? Do you drink alcohol? How often?
Injection drug use and equipment	Have you ever injected drugs? Do you have your own injection equipment? Do you prepare your own drug for injection? Do you use a needle-exchange program? Have you ever shared a needle, syringe, cooker, cotton or water — even once?
Other drug use risks	Do you ever snort drugs? Have you ever shared a snorting straw? Are others present when you inject so that help can be summoned if needed?
Sex under the influence	Have you ever had sex under the influence of alcohol or drugs? If so, have you been more likely to have risky sexual encounters when under the influence, such as having unprotected sex or multiple partners?
Consequences	What effect has drug or alcohol use had on your life? Has your drug or alcohol use caused problems with work? With family? With your health?
Other percutaneous risks	Do you have any body piercings? Any tattoos? Where did you have your piercings or tattoos done?

Table 2. Reference to frequently used substances and their modes of delivery⁴⁶

or delivery									
Substance	Street name	Eat	Freebase*	Inhale	Inject	Oral	Smoke	Sniff/snort	Spray into mouth
Alcohol	Booze, brew, hooch, grog				Some- times	Χ			
Amphetamines	Speed, ice, glass, crystal, crank, bennies, uppers				X	X	X		
Barbiturates	Downers, barbs, blue heavens, yellow jackets, red devils				Some- times	X			
Cannabis	Marijuana, pot, grass, hashish, hash oil, weed	X					X		
Cocaine	Crack, coke, C, blow, flake, snow		X		X		X	X	
LSD/ hallucinogens	Derived from mushrooms (psilocybin), cactus (mescaline), glory seeds, jimson weed. Other examples include LSD (acid), PCP (angel dust), hog				X	X		X	

Table 2. Reference to frequently used substances and their modes of delivery⁴⁶ (continued)

Substance	Street name	Eat	Freebase*	Inhale	Inject	Oral	Smoke	Sniff/snort	Spray into mouth
Narcotic analgesic	Derived from Asian poppy; opium, codeine, morphine, heroin			X	X	X	X		
Ritalin, talwin	T and R				X	X			
Solvents/ aerosols	Glue, gas, sniff			X				X	X
Steroids	Juice, white, stuff, roids				X	Χ			

^{*} Freebase: to use purified cocaine by burning it and inhaling the fumes. Cocaine is "purified" by dissolving it in a heated solvent and separating and drying the precipitate.

Specimen Collection and Laboratory Diagnosis

- · Same as for all patients.
- Given the circumstances often surrounding substance use, urine-based testing, rapid point-of-care testing, self-collected specimens and use of locally based clinics should be considered to improve access to STI testing for this population.

Management and Treatment

- Where patient compliance is a concern, effective single-dose or short-course treatments for STIs are recommended; epidemiologic or syndromic treatment without full evaluation or laboratory testing may sometimes be necessary.
- Integrating STI screening, counselling and treatment into substance treatment and outreach programs has been recommended.^{24,26,47–49} Entry into substance treatment has been linked to a reduction in risky sexual behaviour.⁵⁰
- Be aware of substance use treatment programs and community resources (including safer injection sites, needle-exchange programs and support networks) for referral as needed.

Substance users who are infected with HIV may be at particular risk for serious outcomes. For example, methamphetamine use by people infected with HIV can result in hypertension, hyperthermia, rhabdomyolysis and stroke, and it can produce paranoia, auditory hallucinations and violent behaviour when the user is intoxicated.⁵¹ Fatal interactions between antiretroviral medications (stavudine, saquinavir and ritonavir) and methamphetamine, as well as between ritonavir and ecstasy (MDMA), have been reported.⁵¹

Reporting and Partner Notification

- As with all patients, conditions reportable according to provincial and territorial regulations should be reported to the local public health authority.
- Persons diagnosed with a blood-borne infection such as HIV or infectious syphilis and who share drug using equipment should have their sharing partners notified about possible infection and encouraged to go for testing.
- There are a number of potential reasons substance using patients may not
 accurately report their own substance use or their sexual/injection partners,
 including fear of violence from partner(s), fear of legal repercussions, stigma,
 confidentiality concerns, lack of information on partner(s) and forgetting.
- Repeat prompting and reading back the list of recalled sexual and injection partners can elicit reports of additional sexual and injection partners.⁵²

Follow-up

Patients with substance use problems participating in sexual and/or injection risk behaviours should be encouraged to get regularly screened for STIs, including HIV. Patients whose assessment indicates moderate to severe substance use should be encouraged and facilitated as appropriate to enter a substance treatment/rehabilitation program for follow-up.

References

- Canada's drug strategy. Health Canada website. Available at: www.hc-sc.gc.ca/hecs-sesc/cds/index.htm. Accessed April 5, 2005.
- 2. Canadian Centre on Substance Abuse website. Available at: www.ccsa.ca. Accessed April 5, 2005.
- Health Canada. Straight Facts about Drugs and Drug Abuse. Ottawa, ON: Health Canada; 2000. Available at: www.hc-sc.gc.ca/hecs-sesc/cds/pdf/ straight_facts.pdf. Accessed April 5, 2005.
- Canadian Community Epidemiology Network on Drug Use (CCENDU). 2002 National Report: Drug Trends and the CCENDU Network. Ottawa, ON: Canadian Centre on Substance Abuse; 2003. Available at: www.ccsa.ca/ ccendu/pdf/report_national_2002_e.pdf. Accessed April 5, 2005.
- Canadian Centre on Substance Abuse and Centre for Addiction and Mental Health. Canadian Profile 1999: Alcohol, Tobacco, and Other Drugs. Ottawa, ON: CCSA and CAMH; 1999.
- Single E. A Socio-demographic Profile of Drug Users in Canada. Ottawa, ON: HIV/AIDS Prevention and Community Action Programs of Health Canada; 2000.

- Centre for Infectious Disease Prevention and Control, Health Canada. HIV/ AIDS among injecting drug users in Canada. In: HIV/AIDS Epi Update — May 2004. Ottawa, ON: Health Canada; 2004. Available at: www.phac-aspc.gc.ca/ publicat/epiu-aepi/epi_update_may_04/11_e.html. Accessed April 5, 2005.
- 8. Ross MW, Williams ML. Sexual behavior and illicit drug use. *Annu Rev Sex Res* 2001;12:290–310.
- Wingood GM, DiClemente RJ. The influence of psychosocial factors, alcohol, drug use on African-American women's high-risk sexual behavior. Am J Prev Med 1998:15:54–59.
- Baseman J, Ross M, Williams M. Sale of sex for drugs and drugs for sex: an economic context of sexual risk behavior for STDs. Sex Transm Dis 1999;26:444–449.
- Jones DL, Irwin KL, Inciardi J, et al. The high-risk sexual practices of cracksmoking sex workers recruited from the streets of three American cities. The Multicenter Crack Cocaine and HIV Infection Study Team. Sex Transm Dis 1998;25:187–193.
- 12. Gomez MP, Kimball AM, Orlander H, Bain RM, Fisher LD, Holmes KK. Epidemic crack cocaine use linked with epidemics of genital ulcer disease and heterosexual HIV infection in the Bahamas: evidence of impact of prevention and control measures. *Sex Transm Dis* 2002;29:259–264.
- 13. Wilson TE, Minkoff H, DeHovitz J, Feldman J, Landesman S. The relationship of cocaine use and human immunodeficiency virus serostatus to incident sexually transmitted diseases among women. *Sex Transm Dis* 1998;25:70–75.
- 14. Tyndall MW, Patrick D, Spittal P, Li K, O'Shaughnessy MV, Schechter MT. Risky sexual behaviours among injection drug users with high HIV prevalence: implications for STD control. *Sex Transm Infect* 2002;78(suppl 1):i170–175.
- Moliter F, Ruiz JD, Flynn N, Mikanda JN, Sun RK, Anderson R.
 Methamphetamine use and sexual and injection risk behaviors among out-of-treatment injection drug users. *Am J Drug Alcohol Abuse* 1999;23:475–493.
- Morrison TC, DiClemente RJ, Wingood GM, Collins C. Frequency of alcohol use and its association with STD/HIV-related risk practices, attitudes and knowledge among an African-American community-recruited sample. *Int J STD AIDS* 1998;9:608–612.
- 17. Tapert SF, Aarons GA, Sedlar GR, Brown SA. Adolescent substance use and sexual risk-taking behavior. *J Adolesc Health* 2001;28:181–189.
- 18. Castrucci BC, Martin SL. The association between substance use and risky sexual behaviors among incarcerated adolescents. *Matern Child Health J* 2002;6:43–47.
- Moliter F, Truax SR, Ruiz JD, Sun RK. Association of methamphetamine use during sex with risky sexual behaviors and HIV infection among non-injection drug users. West J Med 1998;168:93–97.
- 20. Novotna L, Wilson TE, Minkoff HL, et al. Predictors and risk-taking consequences of drug use among HIV-infected women. *J Acquir Immune Defic Syndr* Hum Retrovirol 1999;20:502–507.

- 21. Cook RL, Pollock NK, Rao AK, Clark DB. Increased prevalence of herpes simplex virus type 2 among adolescent women with alcohol use disorders. *J Adolesc Health* 2002;30:169–174.
- 22. Ramrakha S, Caspi A, Dickson N, Moffitt TE, Paul C. Psychiatric disorders and risky sexual behaviour in young adulthood: cross sectional study in birth cohort. *BMJ* 2000;321:263–266.
- Sharma AK, Aggarwal OP, Dubey KK. Sexual behavior of drug-users: it is different? Prev Med 2002;34:512–515.
- Hwang LY, Ross MW, Zack C, Bull L, Rickman K, Holleman M. Prevalence of sexually transmitted infections and associated risk factors among populations of drug abusers. Clin Infect Dis 2000;31:920–926.
- 25. Stein MD, Maksad J, Clarke J. Hepatitis C disease among injection drug users: knowledge, perceived risk and willingness to receive treatment. *Drug Alcohol Depend* 2001;61:211–215.
- Bachmann LH, Lewis I, Allen R, et al. Risk and prevalence of treatable sexually transmitted diseases at a Birmingham substance abuse treatment facility. Am J Public Health 2000;90:1615–1618.
- 27. Poulin C, Alary M, Bernier F, Ringuet J, Joly JR. Prevalence of *Chlamydia trachomatis*, *Neisseria gonorrhoeae*, and HIV infection among drug users attending an STD/HIV prevention and needle-exchange program in Quebec City, Canada. *Sex Transm Dis* 1999;26:410–420.
- Flom PL, Friedman SR, Kottiri BJ, et al. Stigmatized drug use, sexual partner concurrency, and other sex risk network and behavior characteristics of 18- to 24-year old youth in a high-risk neighborhood. Sex Transm Dis 2001;28:598–607.
- Staton M, Leukefeld C, Logan TK, et al. Risky sex behavior and substance use among young adults. *Health Soc Work* 1999;24:147–154.
- 30. Bellis MA, Cook P, Clark P, Syed Q, Hoskins A. Re-emerging syphilis in gay men: a case-control study of behavioural risk factors and HIV status. *J Epidemiol Community Health* 2002;56:235–236.
- 31. Koblin BA, Chesney MA, Husnik MJ, et al. High-risk behaviors among men who have sex with men in 6 US cities: baseline data from the EXPLORE study. *Am J Public Health* 2003;93:926–932.
- 32. Colfax GN, Mansergh G, Guzman R, et al. Drug use and sexual risk behavior among gay and bisexual men who attend circuit parties: a venue-based comparison. *J Acquir Immune Defic Syndr* 2001;28:373–379.
- 33. Stall R, Purcell DW. Intertwining epidemics: a review of research on substance use among men who have sex with men and its connection to the AIDS epidemic. *AIDS Behav* 2000;4:181–192.
- 34. Purcell DW, Parsons JT, Halkitis PN, Mizuno Y, Woods WJ. Substance use and sexual transmission risk behavior of HIV-positive men who have sex with men. *J Subst Abuse* 2001;13:185–200.
- 35. Mattison AM, Ross MW, Wolfson T, Franklin D, San Diego HIV Neurobehavioral Research Center Group. Circuit party attendance, club drug use, and unsafe sex in gay men. *J Subst Abuse* 2001;13:119–126.

- 36. McNall M, Remafedi G. Relationship of amphetamine and other substance use to unprotected intercourse among young men who have sex with men. *Arch Pediatr Adolesc Med* 1999;153:1130–1135.
- Sherr L, Bolding G, Maguire M, Elford J. Viagra use and sexual risk behaviour among gay men in London. AIDS 2000;14:2051–2053.
- 38. Chu PL, McFarland W, Gibson S, et al. Viagra use in a community-recruited sample of men who have sex with men, San Francisco. *J Acquir Immune Defic Syndr* 2003;33:191–193.
- 39. Prevention Source BC. *The Workbook on Prevention Concepts*. Available at: www.preventionsource.bc.ca/guides/workbook/sec6.html. Accessed April 5, 2005.
- 40. Canadian AIDS Society. *HIV and HCV Transmission: Guidelines for Assessing Risk*. Ottawa, ON: Canadian AIDS Society; 2004.
- 41. van Empelen P, Schaalma HP, Kok G, Jansen MW. Predicting condom use with casual and steady sex partners among drug users. *Health Educ Res* 2001;16:293–305.
- 42. Canadian Immunization Guide. 6th ed. Ottawa, ON: Health Canada; 2002.
- 43. Kuo I, Mudrick DW, Strathdee SA, Thomas DL, Sherman SG. Poor validity of self-reported hepatitis B virus infection and vaccination status among young drug users. *Clin Infect Dis* 2004;38:587–590.
- 44. Weatherill SA, Buxton JA, Daly PC. Immunization programs in non-traditional settings. *Can J Public Health* 2004;95:133–137.
- 45. Sexual Health and Sexually Transmitted Infections Section, Health Canada. Substance use history. In: Self Learning Module on Sexually Transmitted Diseases. Available at: www.phac-aspc.gc.ca/slm-maa/terry/sixdoors_ e.html#sub. Accessed April 5, 2005.
- 46. Sexual Health and Sexually Transmitted Infections Section, Health Canada. Quick reference on frequently used substances. In: Self Learning Module on Sexually Transmitted Diseases. Available at: www.phac-aspc.gc.ca/slm-maa/ terry/in02_e.html. Accessed April 5, 2005.
- 47. Ross MW, Hwang LY, Leonard L, Teng M, Duncan L. Sexual behavior, STDs and drug use in a crack house population. *Int J STD AIDS* 1999;10:224–230.
- 48. Ross MW, Hwang LY, Zack C, Bull L, Williams ML. Sexual risk behaviours and STIs in drug abuse treatment populations whose drug of choice is crack cocaine. *Int J STD AIDS* 2002;13:769–774.
- 49. Houlding C, Davidson R. Beliefs as predictors of condom use by injecting drug users in treatment. *Health Educ Res* 2003;18:145–155.
- Hoffman JA, Klein H, Clark DC, Boyd FT. The effect of entering drug treatment on involvement in HIV-related risk behaviors. Am J Drug Alcohol Abuse 1998;24:259–284.
- 51. Urbina A, Jones K. Crystal methamphetamine, its analogues, and HIV infection: medical and psychiatric aspects of a new epidemic. *Clin Infect Dis* 2004;38:890–894.
- 52. Brewer DD, Garrett S, Kulasingam S. Forgetting as a cause of incomplete reporting of sexual and drug injection partners. *Sex Transm Dis* 1999;26:166–176.

TRAVELLERS

Definition

There has been a long-standing association between travel, sexual behaviour and the risk of acquiring sexually transmitted infections (STIs). Travellers are defined as people who are journeying temporarily, permanently or episodically for recreational or occupational reasons.¹ Categories of travellers include but are not limited to: tourists, business travellers, military personnel, seamen, truckers, diplomats, college and university students on school break and immigrants visiting their country of origin.²-⁴ Sex tourists are a particular category of travellers whose main intention is to engage in sexual activity when abroad.² They are more likely to engage in sex with sex workers and to have multiple partners when travelling.² Prostitution has developed around tourist resorts in some countries, particularly in Southeast Asia and increasingly in Eastern Europe.².5.6

Epidemiology

In 2002, Canadians took 13 million overnight trips to the United States with an average length of stay of 4 nights and 4.7 million trips overseas with an average length of stay of 15.2 nights. The ten most popular destinations in descending order (excluding the United States) were the United Kingdom, Mexico, France, Cuba, the Dominican Republic, Germany, Italy, the Netherlands, Spain and China.

The risk of acquiring STIs is increased in travellers because travel affords freedom from the normal social constraints of daily life at home, as well as increased time and opportunity for casual sex.8 Studies have shown that 5-50% of travellers engage in casual sex.9 and that 1/3 to over 1/2 of travellers do not consistently use condoms.1 Those at higher risk include males, younger travellers, those who are travelling alone or with friends, those who are single, men who have sex with men (MSM), those planning a long duration of stay, those travelling on business, smokers or those using alcohol or illicit drugs.1 4.5.8 1.112

STIs are among the most common notifiable infections worldwide, with rates being particularly high in developing countries. Chlamydia trachomatis is the most prevalent bacterial STI worldwide. Gonococcal infections remain common worldwide, with the incidence of antibiotic resistance increasing. Antimicrobial susceptibility patterns of Neisseria gonorrhoeae vary worldwide, with a high prevalence of resistance seen in Africa and Asia. For further information on antimicrobial resistance, see Gonococcal Infections chapter.

The World Health Organization estimated that, worldwide, at the end of 2003, there were 38 million adults and children living with HIV — 4.8 million infected in 2003 alone. In Canada, the HIV epidemic is largely due to infection with B subtype viruses. However, travellers may be at risk for transporting non-B subtypes of HIV virus home. In the subtype of HIV virus home.

Prevention

Evidence of the effectiveness of pre-travel interventions is very limited.^{1,8,14} Health care providers should advise travellers to take condoms with them, alert them to the high rates of STIs and reinforce the message that alcohol or illicit drug use lowers inhibitions.^{5,10,11,14} Travellers should be informed that condoms available overseas may be of inferior quality and that hot, humid conditions can decrease the effectiveness of condoms.¹¹ Collaboration between travel clinics and STI programs or clinics is helpful in ensuring appropriate prevention and treatment.¹

Hepatitis B virus (HBV) vaccine is recommended for travellers to areas of HBV endemnicity. Opto-date information on HBV prevalence can be found on the World Health Organization website on International Travel and Health at www.who.int/ith/en or by consulting the 2001 *International Travel Health Guide.* Hepatitis A virus (HAV) vaccination is recommended for MSM, injection drug users and travellers to countries where HAV is endemic to prevent person-to-person transmission of HAV. Combination vaccines for HAV and HBV are useful for patients who require protection against both infections. Chemoprophylatic use of antibiotics for the prevention of STIs while travelling is not recommended.

Prevention efforts should also be targeted at immigrants from HIV-endemic countries who are at increased risk of acquiring HIV infection during visits to their country of origin following immigration to Canada.^{3,6}

Evaluation

Early diagnosis and treatment is key in preventing further spread of STIs, particularly to regular sexual partners at home. A travel and sexual history should be taken. It is important to be aware that self-identified sexual identity is not an accurate predictor of sexual behaviour while travelling. Although some travellers may consider themselves heterosexual, they may have been involved in sexual activities with members of the same sex (either prior to and/or during travel). Therefore it is essential that a sexual history include questions regarding opposite sex and same sex activity. This can be achieved by asking open-ended questions such as: "Do you have sex with men, women or both?"

For a more complete discussion, see *Men Who Have Sex with Men/Women Who Have Sex with Women* and *Primary Care and Sexually Transmitted Infections* chapters.

Practices while travelling (both sexual and non-sexual) that are associated with an increased risk for acquiring STIs should be assessed. These include the following:

- Unprotected oral, vaginal or anal intercourse (receptive and insertive).
- Oral-anal intercourse (anilingus).
- Receptive manual-anal intercourse (insertion of finger or fist in anus of partner).
- · Substance use accompanying sex.
- · Tattooing or body piercing.
- IDU and other drug use.

A substance use history should also be taken.

Travellers who have had unprotected sex with a new partner while travelling should be offered STI screening for chlamydia, gonorrhea, syphilis, HIV and HBV (if unvaccinated). Hepatitis C virus (HCV) testing should be offered if the history reveals drug use, tattooing, body piercing, or other activities where sharing of contaminated equipment may have occurred (see *Immigrants and Refugees* chapter for more information). Health care workers should be aware that travellers may present with STIs rarely seen in Canada, such as chancroid or lymphogranuloma venereum (LGV) (see *Chancroid* and *Lymphogranuloma venereum* chapters). HIV testing should be accompanied by recommended counselling (see *Human Immunodeficiency Virus Infections* chapter).

Specimen Collection and Laboratory Diagnosis

Same as for all patients. See relevant chapters on specific infections.

Management and Treatment

Same as for all patients. See relevant chapters on specific infections.

Reporting and Partner Notification

Same as for all patients. See relevant chapters on specific infections.

Notification of partners abroad may pose a challenge but should be attempted in conjunction with local and provincial departments of health and the Public Health Agency of Canada.

Follow-up

Travellers who engage in high-risk sexual activities when travelling should be encouraged to undergo regular STI screening. Safer-sex and harm reduction counselling should continue to be emphasized. HIV, HBV and HCV testing should be scheduled following the window period, and adherence to safer-sex practices until that time may be indicated to prevent infection of current partners. HAV and HBV vaccination series should be completed if initiated prior to travelling.

References

- Abdullah ASM, Ebrahim SH, Fielding R, Moriskey DE. Sexually transmitted infections in travelers: implications for prevention and control. *Clin Infect Dis* 2004;39:533–538.
- 2. Thomson MM, Najera R. Travel and the introduction of human immunodeficiency virus type 1 non-B subtype genetic forms into Western countries. *Clin Infect Dis* 2001;32:1732–1737.
- Fenton KA, Chinouya M, Davidson O, Copas A; MAYISHA research team. HIV transmission risk among sub-Saharan Africans in London travelling to their countries of origin. AIDS 2001;15:1442–1445.
- Apostolopolous Y, Sonmez S, Yu CH. HIV-risk behaviours of American spring break vacationers: a case of situational disinhibition? *Int J STD AIDS* 2002;13:733–743.
- Memish ZA, Osoba AO. Sexually transmitted diseases and travel. Int J Antimicrob Agents 2003;21:131–134.
- Perrin L, Kaiser L, Yerly S. Travel and the spread of HIV-1 genetic variants. Lancet Infect Dis 2003;3:22–27.
- International Travel 2002. Ottawa, ON: Statistics Canada, Minister of Industry; 2004.
- 8. Cabada MM, Echevarria JI, Seas CR, et al. Sexual behaviour of international travelers visiting Peru. Sex Transm Dis 2002;29:510–513.
- Matteelli A, Carosi G. Sexually transmitted diseases in travelers. Clin Infect Dis 2001;32:1063–1067.
- 10. Ryan ET, Kain KC. Health advice and immunization for travelers. *N Engl J Med* 2000;342:1716–1725.
- 11. Hamlyn E, Dayan L. Sexual health for travellers. *Aust Fam Physician* 2003:32:981–984.
- Arvidson M, Kallings I, Nilsson S, Hellberg D, Mardh PA. Risky behaviour in women with history of casual travel sex. Sex Transm Dis 1997;24:418–421.
- 13. *2004 Report on the Global AIDS Epidemic*. Geneva, Switzerland: Joint United Nations Programme on HIV/AIDS; 2004.
- 14. Thomas RE. Preparing patients to travel abroad safely. Part 4: Reducing risk of accidents, diarrhea and sexually transmitted diseases. *Can Fam Physician* 2000;46:1634–1638.
- 15. Spira AM. A review of combined hepatitis A and hepatitis B vaccination for travelers. *Clin Ther* 2003:25:2337–2351.
- 16. Canadian Immunization Guide. 6th ed. Ottawa, ON: Health Canada; 2002.
- 17. Rose, SR. *International Travel Health Guide*. 12th ed. Northampton, MA: Travel Medicine; 2001.

APPENDIX A: PATIENT COUNSELLING GUIDE ON CONDOM USE

Essential Information on Condoms and Patient Counselling Guide Check the label

- The most common type of condom is the latex condom, but synthetic (polyurethane) condoms also offer protection against unintended pregnancies as well as sexually transmitted infections (STIs), including HIV.
- Natural membrane condoms (also called "sheepskin") are not recommended for use in protection against certain viral diseases such as hepatitis and HIV.
- Novelty condoms, such as "edible condoms," do not offer pregnancy or STI prevention.

Store condoms properly and check them before use

 Condoms should be stored in a cool, dry place out of direct sunlight (i.e., do not store in wallet, in automobile or any place where condoms will be exposed to extreme heat or cold).

Always check the expiry date before using the condom; expired condoms should not be used

- Condoms in damaged packages or those that show obvious signs of age (e.g., those that are brittle, sticky or discoloured) should not be used, because they cannot be relied upon to prevent infection.
- Condoms should be put on before any genital contact to prevent exposure to body fluids that may contain infectious agents. Nonoxynol-9 (N-9) is not recommended as an effective means of HIV or STI prevention. The best STI and HIV barrier is a latex or polyurethane condom without N-9.
 - If N-9 is used as an aid to contraception, its benefit should be carefully considered in light of the increased risk of genital lesions and the resulting potential for an increased risk of HIV transmission.¹

334 Appendix A

Suggestions for Enhancing Adherence to Condom Use for STI Prevention

- Routinely recommend "dual protection" using condoms together with oral contraceptives — for STI prevention and highly effective birth control.
- Prepare a "Prescription Pad Counselling Guide" as follows:²
 If you or your partner has ever had another sexual partner, we strongly recommend that you make one of these safer-sex choices:
 - Use a condom consistently to prevent pregnancy and STIs.
 - Always use a condom for the first 3 months of a sexual relationship with a new partner, and then come in with your partner for STI and HIV testing.
 If your tests are negative, you can quit using condoms, as long as you and your partner feel that you are willing and able to remain monogamous and take appropriate birth control measures.

Barriers to Condom Use and Means to Overcome Them

Table 1. Perceived barrier and proposed intervention strategy

Decreases sexual pleasure or sensation	Often perceived by those who have never used a condom Encourage patient to try the following: Put a drop of water-based lubricant or saliva inside the tip of the condom or on the glans of the penis prior to putting on the condom Try a thinner latex condom Try different brands Try more lubrication
Decreases spontaneity of sexual activity	Encourage incorporation of condom use during foreplay Remind patient that peace of mind may enhance pleasure for self and partner
Embarrassing, juvenile, "unmanly"	Remind patient that it is "manly" to protect oneself and others
Poor fit (too small or too big, slips off, uncomfortable)	Smaller and larger condoms are available
Requires prompt withdrawal after ejaculation	Reinforce the protective nature of prompt withdrawal Suggest substitution of other postcoital sexual activities

Appendix A 335

Table 1. Perceived barrier and proposed intervention strategy (continued)

Fear of breakage may lead to less vigorous sexual activity	With prolonged intercourse, lubricant wears off and condom begins to rub. Have a water- soluble lubricant available to reapply
Non-penetrative sexual activity	 Condoms are advocated for use during fellatio; non-lubricated condoms may prove best for this purpose There are flavoured condoms now on the market, but they should not be confused with edible condoms found in some novelty sex shops Other barriers, such as dental dams or a non-lubricated condom cut down the middle to form a barrier, have been advocated for use during certain forms of non-penetrative sexual activity (e.g., cunnilingus and anilingual sex).

References

- Nonoxynol-9 and the risk of HIV transmission. HIV/AIDS Epi Update
 April 2003. Ottawa, ON: Public Health Agency of Canada; 2003. Available
 at: www.phac-aspc.gc.ca/publicat/epiu-aepi/hiv-vih/nonoxynol_e.html.
 Accessed February 15, 2006.
- 2. Society of Obstetricians and Gynaecologists of Canada Contraception and Sexual Health Initiative: Facilitator's Manual. Ottawa, ON: Society of Obstetricians and Gynaecologists of Canada; 2002.

336 Appendix A

APPENDIX B: HOW TO USE A MALE CONDOM/ HOW TO USE A FEMALE CONDOM

How to Use a Male Condom

It is possible to communicate many of these points to patients clearly in a simple demonstration by putting a condom over two fingers or a model.

- Open the package; handle carefully to avoid damaging the condom.
- A water-based lubricant may be used inside the tip
 of the condom or on the penis to avoid irritation or
 tearing the condom; KY Jelly or a liquid form such as
 Astro-Glide should be used. Petroleum- or oil-based
 lubricants (such as petroleum jelly, cooking oils,
 shortening and lotions) should not be used, because
 they weaken the latex.
- 3. Press the air out of the tip, leaving enough space to hold the semen (about 1 cm).
- 4. Pinching the condom tip, unroll the condom over as much of the hard penis as possible.
- 5. After sex, take the penis out with the condom still on and the penis still hard. Hold the base of the condom firmly so that the semen doesn't spill.
- 6. After use, tie a knot at the open end and dispose of the condom in the garbage (not in the toilet). Do not reuse.

Note:

If a condom breaks, it should be replaced immediately. If ejaculation occurs after condom breakage and there is need for protection against pregnancy, emergency oral contraception should be used.













Appendix B 337

How to Use a Female Condom

Insert the condom into the vagina before sexual contact.

- Open the package, handling carefully to avoid tearing the condom.
- Squeeze the flexible inner ring at the closed end of the sheath.
- 3. Gently insert the inner ring into the vagina.
- 4. Place the index finger on the inside of the condom, and push the inner ring up as far as it will go.
- 5. Be sure the sheath is not twisted. The outer ring should remain on the outside of the vagina.
- 6. Guide the penis into the sheath's opening. Be sure that the penis is not entering on the side, between the vagina wall and the sheath.
- If the condom moves out of place during sex, lubrication can be used either on the inside of the condom or on the penis.
- 8. To remove the condom, twist the outer ring and gently pull the condom out to avoid spilling the semen.
- Dispose of the condom in the garbage (not in the toilet).Do not reuse.

Note:

If the condom is dislodged, twisted or breaks, it should be replaced immediately. If ejaculation occurs after condom failure and there is need for protection against pregnancy, emergency oral contraception should be used.













338 Appendix B

APPENDIX C: RESOURCES AND REFERENCE TOOLS FOR HEALTH PROFESSIONALS

Books

Canadian Guidelines for Sexual Health Education, Health Canada

A resource and reference tool developed by Health Canada in collaboration with sexual health experts to provide the basis for program planners, policy makers, health care professionals, researchers and those working in related fields to build effective sexual health education programs to meet the diverse needs of Canadians. Available in PDF format online at www.phac-aspc.gc.ca/publicat/cgshe-ldnemss/cgshe_index.htm.

HIV/HCV Transmission: Guidelines for Assessing Risk: A Resource for Educators, Counsellors, and Health Care Providers. 4th ed. Canadian AIDS Society

A comprehensive, evidence-based guide outlining the risks associated with various sexual activities, graded from no-risk to high-risk. Available in PDF format online at www.cdnaids.ca.

Sex Sense. Society of Obstetricians and Gynaecologists of Canada

A comprehensive booklet about sexuality and contraception. This booklet covers all contraceptive methods available in Canada and provides fact-based information on protection against sexually transmitted infections. It contains helpful websites and phone numbers for support across Canada. Available to order online at www.sogc.org/sexsense/book.htm.

Sexual and Reproductive Health Counselling Guidelines. Planned Parenthood Federation of Canada

These guidelines can be used as a tool to improve support skills, train staff or provide additional information for patients in a clinical, community or educational setting. Available to order online at www.ppfc.ca.

Appendix C 339

Internet Links

www.aidssida.cpha.ca

The National AIDS Clearinghouse of the Canadian Public Health Association (1565 Carling Avenue, Suite 400, Ottawa, ON, K1Z 8R1) distributes a variety of pamphlets, posters and other safer-sex materials.

www.phac-aspc.gc.ca/std-mts/index.html

The Public Health Agency of Canada Sexual Health and Sexually Transmitted Infections website provides resources on STI support surveillance and targeted research studies, evidence-based national guidelines and policy, and the dissemination and exchange of information.

www.sexualityandu.ca/masexualite.ca

A sexual- and reproductive-health website sponsored by the Society of Obstetricians and Gynaecologists of Canada. It is widely used by teens, parents, adults, teachers and health care professionals to access relevant sexual and reproductive health information.

Note:

If you are not aware of a local source of health promotion material, contact your local public health authority or provincial/territorial director of STI control (see *Appendix D*).

340 Appendix C

APPENDIX D: PROVINCIAL AND TERRITORIAL DIRECTORS OF STI CONTROL

Alberta

Dr. Ameeta Singh Infectious Diseases Medical Consultant Office of Provincial Health Officer 24th Floor, Telus Plaza, North Tower 10025 Jasper Avenue Edmonton, AB T5J 2N3

Tel: 780-427-5263 Fax: 780-427-7683

ameeta.singh@gov.ab.ca

British Columbia

Dr. Michael Rekart, Director Division of STD/AIDS Control BC Centre for Disease Control 655 West 12th Avenue Vancouver, BC V5Z 4R4 Tel: 604-660-6178

Fax: 604-775-0808 michael.rekart@bccdc.ca

Manitoba

Dr. Carole Beaudoin Epidemiologist, Communicable Disease Manitoba Health 4th Floor, 300 Carleton Street Winnipeg, MB R3B 3M9 Tel: 204-788-6786

Fax: 204-948-2040 cabeaudoin@gov.mb.ca

New Brunswick

Dr. Holy Akwar Communicable Disease Epidemiologist Office of the Chief Medical Officer of Health New Brunswick Department of Health and Wellness 2nd Floor, 520 King Street, PO Box 5100

Fredericton, NB E3B 5G8 Tel: 506-453-2323

Fax: 506-453-8702 holy.akwar@gnb.ca

Newfoundland and Labrador

Dr. Faith Stratton Chief Medical Officer of Health Department of Health Building 801, Pleasantville St. John's, NF A1B 4J6 Tel: 709-729-3430 Fax: 709-729-5824 fstratton@mail.gov.nf.ca

Northwest Territories

Dr. André Corriveau Chief Medical Health Officer Department of Health and Social Services Population Health. Health Protection Unit Government of Northwest Territories Yellowknife, NT X1A 2I 9

Tel: 867-920-8646 Fax: 867-873-0442

andre_corriveau@gov.nt.ca

Appendix D

Nova Scotia

Dr. Jeff Scott Office of the Chief Medical Officer of Health PO Box 488 Halifax, NS B3J 2R8 Tel: 902-424-8698 Fax: 902-424-0550

medicalofficerofhealth@gov.ns.ca

Nunavut

Elaine Randell Communicable Disease Consultant Dept of Health & Social Services PO Box 1000, Station 1000 Igaluit, NU X0A 0H0 Tel: 867-975-5775

Fax: 867-979-3190 ERandell@gov.nu.ca

Ontario

STI Medical Director STI/AIDS Sexual Health Unit Ministry of Health and Long-Term Care 8th Floor, 5700 Yonge Street Toronto, ON M2M 4K5 Tel: 416-327-7429

Fax: 416-327-7439

Prince Edward Island

Dr. Lamont Sweet Chief Medical Officer of Health 16 Garfield Street, Box 2000 Charlottetown, PE C1A 7N8 Tel: 902-368-4996

Fax: 902-620-3354 lesweet@ihis.org

Quebec

Mme Lise Guérard Chef de service Service de lutte contre les infections transmissibles sexuellement et par le sang Direction générale de la santé publique Ministère de la Santé et des Services sociaux 201, rue Crémazie Est, RC-03 Montréal, QC H2M 1L2 Tel: 514-873-9892 Fax: 514-873-9997

lise.guerard@msss.gouv.qc.ca

Saskatchewan

Dr. Huiming Yang Deputy Chief Medical Health Officer Communicable Disease Control and Vaccines Population Health Branch Saskatchewan Health 3475 Albert Street Regina SK S4S 6X6 Tel: 306-787-3148 Fax: 306-787-9576 hyang@health.gov.sk.ca

Yukon Territory

Ms. Colleen Hemsley Communicable Disease Officer Health & Social Services Yukon Territorial Government 4 Hospital Road Whitehorse, YT Y1A 3H8 Tel: 867-667-8369

Fax: 867-667-8349

colleen.hemsley@gov.yk.ca

Appendix D

APPENDIX E: PROVINCIAL LABORATORIES

For more information on laboratory diagnosis of sexually transmitted infections, first consult your local facility or your nearest public health laboratory.

Alberta

Provincial Laboratory for Public Health (Microbiology)

Edmonton site:

8440 – 112 Street Edmonton, AB T6G 2J2

Tel: 780-407-7121 Fax: 780-407-8984

Calgary site:

3030 Hospital Drive NW Calgary, AB T2N 4W4 Tel: 403-944-1200 Fax: 403-283-0142

British Columbia

Provincial Laboratory
BC Centre for Disease Control
Laboratory Services
655 12th Avenue West
Vancouver, BC V5Z 4R4
Tel: 604-660-6030

Fax: 604-660-6073

Manitoba

Cadham Provincial Laboratory 750 William Avenue Winnipeg, MB R3E 3J7 Tel: 204-945-6123 Fax: 204-786-4770

New Brunswick

Department of Laboratory Medicine St. John Regional Hospital 400 University Avenue Saint John, NB E2L 4L2 Tel: 506-648-6501

Fax: 506-648-6576

Newfoundland and Labrador

Newfoundland Public Health Laboratories The Leonard A. Miller Centre for Health Sciences 100 Forest Road, PO Box 8800 St. John's, NF A1B 3T2 Tel: 709-777-6555 Fax: 709-737-7070

Nova Scotia

Department of Pathology and Laboratory Medicine Queen Elizabeth II Health Science Centre 5788 University Avenue Halifax, NS B3H 1V8 Tel: 902-473-2231

Ontario

Central Public Health Laboratory 81 Resources Road Etobicoke, ON M9P 3T1 Tel: 416-235-6132

Toll-free: 1-800-640-7221 Fax: 416-235-6103

Fax: 902-473-4432

Hamilton Public Health Laboratory 250 Fennell Avenue West, PO Box 2100

Hamilton, ON L8N 3R5 Tel: 905-385-5379 Fax: 905-385-0083

Kingston Public Health Laboratory 181 Barrie Street, PO Box 240 Kingston, ON K7L 3K2

Tel: 613-548-6630 Fax: 613-548-6636

Appendix E 343

London Public Health Laboratory 850 Highbury Avenue, PO Box 5704, Terminal A

London, ON N6A 4L6 Tel: 519-455-9310 Fax: 519-455-3363

Orillia Public Health Laboratory 750 Memorial Avenue, PO Box 600 Orillia, ON L3V 6K5

Tel: 705-325-7449 Fax: 705-329-6001

Ottawa Public Health Laboratory 2380 Saint Laurent Boulevard Ottawa, ON K1G 6C4

Tel: 613-736-6800 Fax: 613-736-6820

Peterborough Public Health Laboratory 99 Hospital Drive, PO Box 265 Peterborough, ON K9J 6Y8

Tel: 705-743-6811 Fax: 705-745-1257

Sault Sainte-Marie Public Health Laboratory 160 McDougall Street, PO Box 220 Sault Sainte-Marie, ON P6A 3A8

Tel: 705-254-7132 Fax: 705-945-6873

Sudbury Public Health Laboratory 2 - 1300 Paris Street Sudbury, ON P3E 6H3 Tel: 705-564-6917

Fax: 705-564-6918

Thunder Bay Public Health Laboratory 336 South Syndicate Avenue Thunder Bay, ON P7E 1E3 Tel: 807-622-6449

Fax: 807-622-5423

Timmins Public Health Laboratory 67 Wilson Avenue Timmins, ON P4N 2S5 Tel: 705-267-6633

Fax: 705-360-2006

Toronto Public Health Laboratory PO Box 9000, Terminal A Toronto, ON M5W 1R5 Tel: 416-235-6132

Toll-free: 1-800-640-7221 Fax: 416-235-6103

Windsor Public Health Laboratory 3400 Huron Church Road, PO Box 1616 Windsor, ON N9E 4H9

Tel: 519-969-4341 Fax: 519-973-1481

Prince Edward Island

Division of Laboratories Provincial Health Laboratory Queen Elizabeth Hospital Riverside Drive, PO Box 6600 Charlottetown, PE C1A 8T5

Tel: 902-894-2300 Fax: 902-894-2385

Quebec

Institut national de santé publique du Québec

Laboratoire de santé publique du Québec

20045, chemin Sainte-Marie ouest Sainte-Anne-de-Bellevue, QC H9X 3R5

Tel: 514-457-2070 Fax: 514-457-6346

Saskatchewan

Saskatchewan Provincial Laboratory Services Saskatchewan Health 3211 Albert Street Regina, SK S4S 5W6 Tel: 306-787-3131

Fax: 306-787-9122

APPENDIX F: FORENSIC EVIDENCE, SERVICES AND LABORATORIES

Forensic Evidence

- Forensic evidence is invaluable in supporting the testimony of victims of sexual assault.
- The purpose of forensic analysis of specimens is to establish one or more of the following:
 - That there was some form of association between the victim and the accused.
 - That sexual contact occurred.
 - That the assault was violent or forceful, thereby indicating lack of consent.
 - That the victim may have been drugged.
- Types of forensic analyses most useful in sexual assault are as follows:
 - Identification of semen or other bodily fluids.
 - Forensic DNA analysis.
 - Hair examination (suitability for DNA analysis).
 - Textile damage assessment.
 - Examinations involving fibres and other types of trace evidence.
 - Drug screen (including alcohol) in bodily fluids (blood and urine).
- In some situations, it may be impossible to collect certain specimens for forensic
 analysis. The availability of specimens depends on the sex of the perpetrator,
 the nature of the molestation (fondling vs. penetration) and the time between the
 event and the examination. An interval of more than 48 hours or cleansing the
 sexually abused areas will reduce the availability of specimens and the strength
 of forensic evidence.
- When specimens are being collected as forensic evidence with the objective of
 establishing the identification of the perpetrator, certain strict guidelines must
 be followed. This is essential if the information gathered is to be unequivocally
 accepted in court. Particular attention must be paid to the manner of collection,
 the labelling and identification of individual specimens, and obtaining signed
 specific consent forms. For details on the collection of specimens for forensic
 analysis, local police authorities should be consulted (see Forensic Laboratories,
 below).

Collection of specimens

- Physicians should familiarize themselves with the test kit before they need to use it.
- Sexual assault examination kits differ by jurisdiction. An approved sexual assault
 examination kit should be used for the collection of specimens. Local practices
 and the instructions contained within the sexual assault kit should be carefully
 followed.
- An attempt should be made to obtain specimens of seminal fluid (pristine
 material) from all possible sites with sterile cotton swabs. The swabs should then
 be allowed to air dry. The forensic laboratory will examine these specimens for
 the presence of semen and conduct DNA typing.

Appendix F 345

- Any residual fluids from affected areas, such as the vaginal vestibule, should be collected by aspiration. A sterile eye dropper is ideal for this purpose in children.
 - Before aspiration, the area should be moistened with 1–2 mL of sterile saline.
 - Depending on local policies and the availability of appropriate equipment and training, samples can be examined for the presence of motile sperm. A positive finding suggests that the sexual activity occurred less than 6 hours previously. Confirmation of the presence of spermatozoa by the forensic laboratory is essential.
- Demonstration of saliva on the body or clothing of the person who has been abused or assaulted may provide valuable forensic evidence.
 - Samples from the body can be collected with a sterile cotton swab. The swab should be moistened slightly with distilled water and rubbed over the affected area of the body. The specimen should be allowed to dry and then packaged and labelled.
 - If a child or adult is unclear about which area(s) is (are) affected, the common target areas (the neck, breast, belly, genital area, penis, thighs and buttocks) may be swabbed; a separate swab should be used for each area and labelled accordingly.
- Judgment is required in deciding whether these investigations are sensible. It is
 pointless to collect such samples if weeks have elapsed since the incident or if
 the critical areas have since been bathed.
- The body and the clothing worn at the time of the incident may contain trace evidence (foreign material left by the perpetrator). Items commonly encountered include hair from any part of the body, clothing fibres, lubricants, petroleum jelly and lipstick. Any suspicious hair or fibre material found on the body of the person should be removed with forceps, folded in a piece of clean paper and put in a separate, properly labelled envelope. Suspicious material such as lubricants, petroleum jellies and lipstick on the body of the person should be removed using a sterile swab, then packaged and labelled. Each item of clothing worn by the person should be packaged separately and labelled.
- If the assaulted or abused person has reached puberty, the pubic hair should be combed and the comb, as well as any free hair collected, should be folded in a piece of paper or tissue and put in a labelled envelope or placed in a plastic bag and then sealed and labelled. Hairs can be assessed to determine their body area of origin (pubic, scalp or body hair). In addition, the root portions of any hairs may be suitable for DNA analysis.
- Fingernall scrapings/clippings should be collected if there is a possibility that
 the perpetrator was scratched during the incident. The forensic laboratory will
 examine these samples for the presence of blood and foreign DNA. Clippings can
 be collected using clean nail clippers or scissors, folded into a piece of paper or
 tissue and placed into a labelled envelope or container. Fingernail scrapings can
 be collected using a nail scraper and the scraper and debris folded into a piece
 of paper or tissue and placed into a labelled envelope or container.

346 Appendix F

Collection of known samples for DNA analysis

It is essential for DNA typing analysis to collect a known sample from the victim. A blood stain, mouth swab or pulled hair sample can be collected as a known sample from the victim following the instructions provided in the approved sexual assault examination kit. A known blood stain is the preferred sample to be collected from the victim. A known blood stain, mouth swab or pulled hair sample can also be collected using the appropriate consent sample collection kits that are available from the Case Receipt Units of the Royal Canadian Mounted Police Forensic Laboratory Services.

Collection of samples for toxicological analysis

Blood and urine samples should be collected from the victim for toxicological analyses using the blood collection tube and urine jar provided in the sexual assault kit or grey-stoppered blood collection tubes available at the hospital.

Forensic Services

- Investigative and scientific forensic laboratory services to detect evidence of sexual assault and abuse are available throughout Canada.
- Services are supplied by the Royal Canadian Mounted Police and by federal, provincial, regional and local agencies and police forces.
- Current legislation on the abuse of children obliges physicians to notify local child protection agencies of such cases. These local agencies maintain close liaison with police force personnel familiar with the investigation of suspected abuse and with the availability of forensic laboratory services.
- Physicians should not submit specimens for forensic study directly to laboratories. This should be done through police services.
- Physicians wishing to consult scientists on forensic matters may do so by contacting the nearest laboratory.
- Most forensic evaluations do not include tests to detect sexually transmitted infections.

Appendix F 347

Forensic Laboratories

Alberta

General Manager
Forensic Laboratory Services —
Edmonton

Royal Canadian Mounted Police 15707 118th Avenue Edmonton, AB T5V 1B7

Tel: 780-451-7400 Fax: 780-495-6961

British Columbia

General Manager
Forensic Laboratory Services —
Vancouver
Royal Canadian Mounted Police
5201 Heather Street
Vancouver, BC V5Z 3L7

Tel: 604-264-3400 Fax: 604-264-3499

Manitoba

General Manager
Forensic Laboratory Services —
Winnipeg
Royal Canadian Mounted Police
621 Academy Road
Winnipeg, MB R3N 0E7
Tel: 204-983-4267

Tel: 204-983-4267 Fax: 204-983-6399

Nova Scotia

General Manager Forensic Laboratory Services — Halifax

Royal Canadian Mounted Police 3151 Oxford Street, PO Box 8208 Halifax, NS B3K 5L9

Tel: 902-426-8886 Fax: 902-426-5477

Ontario

Chief Scientific Officer
Forensic Laboratory Services —
Ottawa
Royal Canadian Mounted Police
1200 Vanier Parkway, PO Box 8885

Ottawa, ON K1G 3M8 Tel: 613-993-0986 Fax: 613-952-0156

Northern Regional Laboratory of the Centre of Forensic Sciences Suite 500, 70 Foster Drive Sault Sainte-Marie, ON P6A 6V3

Tel: 705-945-6550 Fax: 705-945-6569

Director Centre of Forensic Sciences 25 Grosvenor Street Toronto, ON M7A 2G8 Tel: 416-314-3200

Fax: 416-314-3225

Quebec

Le directeur
Laboratoire de sciences judiciaires et
de médecine légale
1701, rue Parthenais, PO Box 1500
Montreal, QC H2K 3S7

Tel: 514-873-2704 Fax: 514-873-4847

Saskatchewan

General Manager
Forensic Laboratory Services —
Regina
Royal Canadian Mounted Police
6101 Dewdney Avenue West,
PO Box 6500

Regina, SK S4P 3J7 Tel: 306-780-5810 Fax: 306-780-7571

APPENDIX G: REFERRAL CENTRES FOR STIS IN PERIPUBERTAL AND PREPUBERTAL CHILDREN

This list of child and youth abuse treatment centres in Canada is not inclusive; however, it can be used as a reference for obtaining more specific local information.

Alberta

Child Abuse Program Alberta Children's Hospital 1820 Richmond Road Southwest Calgary, AB T2T 5C7

Tel: 403-943-7886

Department of Pediatrics
Stollery Children's Hospital
2C–300 Walter McKenzie Health Centre
University of Alberta
Edmonton, AB T6G 2B7
Tel: 780-407-6370

British Columbia

Child Protection Services Royal Columbian Hospital 330 East Columbia Street New Westminster, BC V3L 3W7

Tel: 604-520-4253

BC Children's Hospital 4480 Oak Street Vancouver, BC V6H 3V4 Tel: 604-875-2345

Sexual Assault Assessment Project Department of Family Practice University of British Columbia 5804 Fairview Avenue Vancouver, BC V6T 1Z3 Tel: 604-822-5431

Suspected Child Abuse and Neglect Team Victoria General Hospital 1 Hospital Way Victoria, BC V8Z 6R5 Tel: 250-727-4212

Manitoba

Child Protection Centre Children's Hospital of Winnipeg Health Sciences Centre 685 William Avenue Winnipeg, MB R3A 1R9 Tel: 204-787-2811

New Brunswick

Child Protection Consultation Team Attn: Social Work Moncton Hospital 135 MacBeath Avenue Moncton, NB E1C 6Z8 Tel: 506-857-5331

Child Protection Team Saint John Regional Hospital PO Box 2100 Saint John, NB E2L 4L2 Tel: 506-648-6811

Newfoundland and Labrador

Protection Team
Janeway Children's Health &
Rehabilitation Centre
300 Prince Phillip Drive
St. John's, NF A1A IR8
Tel: 709-777-6300

Northwest Territories

Department of Health and Social Services Government of the Northwest Territories PO Box 1320 Yellowknife, NT X1A 2L9

Tel: 867-920-3231 Fax: 867-873-0442

Appendix G 349

Nova Scotia

Child Abuse Team IWK Health Centre 5850/5980 University Avenue, PO Box 9700 Halifax, NS B3K 6R8

Tel: 902-470-8888

Nunavut

Director of Child and Family Services
Department of Health and
Social Services
Government of Nunavut
PO Box 1000, Station 1000
Iqaluit, NU XOA 0H0

Tel: 867-975-5750 Fax: 867-975-5705

Ontario

Child Abuse Committee
Brampton Memorial Hospital
20 Lynch Street
Brampton, ON L6W 2Z8
Tel: 905-451-1710

Child Protection Team Hamilton Health Sciences PO Box 2000, Station A Hamilton, ON L8N 3Z5 Tel: 905-521-2100

Child Protection Team Hotel Dieu Hospital 166 Brock Street Kingston, ON K7L 5G2 Tel: 613-544-3310

Gyne/Endo Clinic Children's Hospital of Western Ontario 800 Commissioners Road East London, ON N6A 4G5 Tel: 519-685-8484 Child Abuse Team
Trillium Health Centre
100 Queensway West
Mississauga, ON L5B 1B8
Tel: 905-848-7100, ext. 2548

Child and Youth Protection Children's Hospital of Eastern Ontario 401 Smyth Road Ottawa, ON K1H 8L1 Tel: 613-737-7600

Child Abuse Committee
Blue Water Health
220 North Milton Street
Sarnia, ON N7T 6H6
Tel: 519-464-4500 ext. 8259

Child Abuse Team Shoniker Clinic 2867 Ellesmere Road Scarborough, ON M1E 4B9 Tel: 416-281-7301

Chief of Pediatrics St. Joseph's Care Group 35 North Algoma Street PO Box 3251 Thunder Bay, ON P7B 5G7 Tel: 807-343-2431

Suspected Child Abuse and Neglect Program Hospital for Sick Children 555 University Avenue Toronto, ON M5G 1X8 Tel: 416-813-6275

Child Abuse Team North York General Hospital 4001 Leslie Street Toronto, ON M2K 1E1 Tel: 416-756-6000

350 Appendix G

Quebec

Adolescent Clinic Montreal Children's Hospital 1040 Atwater Street Montreal, QC H3Z 1X3 Tel: 514-934-1934, ext. 24481

Comité de prévention de l'enfance maltraitée

Direction de la protection de la jeunesse Hôpital Maisonneuve-Rosemont 5415, boulevard de l'Assomption Montreal, QC H1T 2M4 Tel: 514-252-3400, ext. 3826

Clinique de pédiatrie socio-juridique Hôpital Sainte-Justine 3175, chemin Côte Ste-Catherine Montreal, QC H3T 1C5 Tel: 514-345-4866 (0–11 years old) Tel: 514-345-4721 (12–18 years old)

Comité de protection de l'enfance Centre hospitalier de l'Université Laval (CHUL) 2705, boulevard Laurier Ste-Foy, QC G1V 4G2 Tel: 418-656-4141

Clinique médico-juridique Centre hospitalier universitaire de l'Estrie Sherbrooke, QC J1H 5N4 Tel: 819-346-1110, ext. 14644

Saskatchewan

Child Abuse Team Regina General Hospital 1440 14th Avenue Regina, SK S4P 0W5 Tel: 306-766-4444

Child and Youth Service Department of Psychiatry Royal University Hospital 103 Hospital Drive Saskatoon, SK S7N 0W8

Tel: 306-655-1000

Yukon

Communicable Disease Officer Yukon Communicable Disease Control 4 Hospital Road Whitehorse, YT Y1A 2C6 Tel: 867-667-8369

Fax: 867-667-8349

Appendix G 351

APPENDIX H: TANNER SCALE OF SEXUAL MATURITY

Sexual maturity ratings have replaced the traditional indicators of growth status such as height, weight and skinfold thickness. Sexual maturity ratings have proven useful in assessing growth and development during adolescence.

Classification of patients may be done as part of a general physical examination and does not require any special procedures.

The scale of development is based on secondary sexual characteristics. The ratings range from stage 1, which represents the prepubertal child, to stage 5, which represents the adult.

Boys: Genital Development

- Stage 1: Preadolescent. Testes, scrotum and penis are about the same size and proportion as in early childhood.
- Stage 2: Enlargement of scrotum and testes. Skin of scrotum reddens and changes in texture. Little or no enlargement of penis.
- Stage 3: Enlargement of penis, at first mainly in length. Further growth of testes and scrotum.
- Stage 4: Increased size of penis, with growth in breadth and development of glans. Testes and scrotum larger. Scrotal skin darkened.
- Stage 5: Genitalia are adult in size and shape.

Girls: Breast Development

- Stage 1: Preadolescent. Elevation of papilla only.
- Stage 2: Breast bud stage. Elevation of breast and papilla as small mound.
 Enlargement of diameter of areola.
- Stage 3: Further enlargement and elevation of the breast and areola, with no separation of their contours.
- Stage 4: Projection of areola and papilla to form a secondary mound above the level of the breast.
- Stage 5: Mature stage. Projection of papilla only, owing to recession of the areola to the general contour of the breast.

352 Appendix H

Both Sexes: Pubic Hair

- Stage 1: Preadolescent. Vellus over pubes is not developed further than that over abdominal wall (i.e., no pubic hair).
- Stage 2: Sparse growth of long, slightly pigmented downy hair, straight or slightly curled, chiefly at base of penis and along labia.
- Stage 3: Hair is considerably darker, coarser and more curled. It spreads sparsely over the junction of pubes.
- Stage 4: Hair is adult in type, but area covered is still considerably smaller than in adult. No spread to medial surface of thighs.
- Stage 5: Hair is adult in quantity and type, with distribution of horizontal (or classic "feminine" in females) pattern. Spread to medial surface of thighs but not up linea alba or elsewhere above base of inverse triangle (spread up linea alba occurs late and is rated Stage 6).

Appendix H 353

INDEX

alopecia 233
amine odour whiff testing 49, 73, 110
aminoglycosides
gentamicin 75, 76, 88, 89
amiodarone 57
amoxicillin. See penicillin
amphotericin B 115
ampicillin/sulbactam penicillin.
See penicillin
amsel criteria 73
anal 11, 12, 16–20, 32, 43
discharge. See discharge
See rectal
See individual syndrome, infection and population chapters for specific
considerations
anemia 208, 233
anilingus (oral-anal intercourse). See oral sex
anogenital 160–163, 165, 168, 250, 258, 265, 266, 283
anoscopy. See sigmoidoscopy and proctoscopy
antibody
IgG 37, 38, 210
IgM 35, 37, 38, 48, 149, 192
Genital Herpes Simplex Virus Infections chapter 145, 147–149, 153, 154
Hepatitis B Virus Infections chapter
191–192
Human Immunodeficiency Virus
Infections chapter 200, 206, 209
Lyphogranuloma Venereum chapter 230
230

	Syphilis chapter 237	azole
	<i>Immigrants and Refugees</i> chapter 250	butoconazole 280
		clotrimazole 113, 114
	Pregnancy chapter 276, 285	fluconazole 113–115, 279
	Sexual Abuse of Peripubertal and Prepubertal Children chapter 298	itraconazole 114
	Sexual Assault in Postpubertal	ketoconazole 114
	Adolescents and Adults chapter 309	miconazole 113, 280
an	itiemetics 181	terconazole 280
an	ntimicrobial resistance (AMR). See drug resistance	В
ар	pendicitis 71, 74	bacteria
Ar	gyll Robertson pupil 233	aerobic 71
ar	thralgia 191, 207, 215, 224	anaerobic 48, 71, 75
ar	thritis 175, 179, 182, 224	facultative 71, 75
ar	tificial insemination 274	bacterial vaginosis (BV) 33, 39
AS	SCUS. See cervical dysplasia	Pelvic Inflammatory Disease chapter
as	septic meningitis. See meningitis	73, 76
as	pirate	Vaginal Discharge chapter 106–112,
	bubo 35, 36, 124, 225, 227, 228,	117 Human Immunodeficiency Virus
	298, 346	Infections chapter 199
	epididymal 65, 179	Men Who Have Sex With Men/
	nasopharyngeal 32, 129	Women Who Have Sex With Women
	vaginal 298, 346	chapter 263
	sault, sexual 11, 36, 177, 202, 215, 93, 305-314, 345-347	Pregnancy chapter 274, 278–279
	symptomatic. See Primary Care and	Sexual Abuse in Peripubertal and Prepubertal Children chapter 296
	exually Transmitted Infections chapter	bare-backing 262
ata	axia 233	barriers. See condoms
ato	ovaquone 213	bartholinitis 175
at	ypical squamous cells of	bathhouses 11, 13, 262, 265
	undetermined significance (ASCUS). See cervical dysplasia	Behçet's disease
27	ithromycin. See macrolide	benzathine penicillin G. See penicillin
	ithromycin resistance. <i>See</i> resistance	bichloracetic acid 168
αZ	and the state of t	bimanual exam 20, 48, 73, 251
		23.1ddi 0./diii 201

cannabis. See substance use

biopsy 66, 165, 215	carcinoma 4, 52, 64, 90, 163, 164
endometrial 73, 107, 129	hepatocellular carcinoma 191
prostatic 86	CD4. See HIV/AIDS
vulvar 64	cefixime. See cephalosporin
bisexual 262, 263	cefotetan. See cephalosporin
bleach kits 257	cefoxitin. See cephalosporin
blood 11, 21, 35, 51, 74, 94, 179, 190,	ceftriaxone. See cephalosporin
201, 203, 251, 346	cephalosporin 76, 275
blood borne pathogens (BBP) 203, 204, 223, 299, 307	cefixime 95, 100, 103, 181–183, 275, 300, 310–311
donors 189, 193, 199, 201, 202, 214	cefotetan 75
See white blood cell	cefoxitin 75, 76
boric acid 114, 115, 279	ceftriaxone 76, 124, 181-184,
bowenoid papulosis 163	238–239, 275, 311
breastfeeding 112, 117, 205, 285, 286	cephalosporin allergy. See allergy
buboes. See lymphadenopathy	cervical cancer. See cervical dysplasia
burrow ink test 142	cervical discharge. See discharge
butoconazole. See azole BV. See bacterial vaginosis	cervical dysplasia 4, 161–163, 208, 210, 259, 266, 316
C	atypical squamous cells of undetermined significance (ASCUS) 38, 43, 164
c-reactive protein 48, 73	cervical intra-epithelial neoplasia
c-section. See cesarean	(CIN) 162, 163
Campylobacter jejuni subsp. Jejuni 51, 93	high-grade squamous intra-epithelial lesions (HSIL) 163, 164
Canadian Adverse Drug Reaction Monitoring Program iv, 130	low-grade squamous intra-epithelial lesions (LSIL) 43, 162, 164
cancer 5, 52, 106, 160–169, 208, 215	cervical friability 45
See carcinoma	cervical mability 43
See neoplasia	strawberry cervix 45, 108
candida (<i>Candida albicans</i>) 39, 63, 98, 106, 107, 113–116	See individual syndrome, infection
candidiasis 208	and population chapters for specific considerations
oroesophageal 206, 208, 213	cervix 20, 25, 31, 34–36, 38, 43, 45
vulvovaginal 33, 49, 50, 106, 107, 110, 113–117, 279–280, 296	cesarean 107, 152–154, 274, 282–283, 284, 286

chancroid (*Haemophilus ducreyi*) 10, 25, 32, 36, 46

Genital Ulcer Disease chapter 59-68

Chancroid chapter 122–125

Lymphogranuloma Venereum chapter **229**

Syphilis chapter 236

Immigrants and Refugees chapter 250

Travellers chapter 332

Child Protection Act 22, 24

chlamydia (*Chlamydia trachomatis*) 9, 21, 25, 28, 35, 43, 45

Chlamydial Infections chapter 126–134

Epididymitis chapter 54, 56

Genital Ulcer Disease chapter 59

Pelvic Inflammatory Disease chapter 71, 73, 75, 76

Prostatitis chapter 84

Sexually Transmitted Intestinal and Enteric Infections chapter 93, 95

Urethritis chapter 98, 103

Vaginal Discharge chapter 106, 110

Gonococcal Infections chapter 179–180, 184

Human Immunodeficiency Virus Infections chapter 199

Lymphogranuloma Venereum chapter 223–230

Inmates and Offenders chapter 259

Men Who Have Sex With Men/ Women Who Have Sex With Women chapter 263, 267

Pregnancy chapter 273, 274-275

Sexual Abuse in Peripubertal and Prepubertal Children chapter 296–297, 299–301 Sexual Assault in Postpubertal Adolescents and Adults chapter 308, 310

Sex Workers chapter 316

Travellers chapter 332

cholestasis 284

Cialis. See tadalafil

CIN. See cervical dysplasia ciprofloxacin. See fluoroquinolone circuit parties 60, 263, 265 circumcision 59, 123

cirrhosis 191

Citizenship and Immigration Canada (CIC) 249–250

clarithromycin. See macrolide

clindamycin 75, 109, 111, 112, 132, 279

clotrimazole. See azole

clue cells 39, 49, 73, 108-110

CO₂ laser ablation 168, 283

cocaine. See substance use

coliforms 47, 53, 54

colposcopy 38, 164, 165, 215, 306

complement fixation (CF) 226–227

condoms 19, 21 111, 147, 161, 201–203, 206, 223, 324–328

See individual population chapters for specific considerations

condyloma 162, 170

condyloma lata (condylomata lata) 52, 163, 233, 234

condylomata acuminata 160, 163 congenital syphilis. *See* syphilis

conjunctivitis 127, 130, 134, 175, 176, 274

constipation 51, 94, 224

dermatitis 64, 106, 141, 143, 175
DFA. See direct fluorescent
antibody assay
diarrhea 51, 94, 207, 208 diplococci 36, 44, 47, 74, 103, 178,
180, 308
direct fluorescent antibody assay (DFA) 31, 32, 37, 46, 65, 99, 123, 129, 234, 236, 237
disability-adjusted life years (DALY) 316
discharge 12, 20, 163 abnormal
anal/rectal 51, 94, 176, 224
cervical 45, 180
eye 184
nasal 237
urethral 33, 44, 47, 55, 98, 99, 103, 127, 176, 180
vaginal 20, 45, 49, 50, 73, 106–118, 127, 176, 296, 298
donovanosis. See granuloma inguinale
doxycycline. See tetracycline
drug resistance. See resistance
dyspareunia 48, 108, 127, 176
dysphagia 208
dysuria 12, 49, 50, 86, 99, 102, 108,
127, 176
E
ectoparasitic infestations
pubic lice (<i>Phthirus pubis</i>)/ pediculosis pubis 140, 141, 144, 280–281
scabies (<i>Sarcoptes scabiel</i>) 63, 142–144, 281
ectopic pregnancy. See pregnancy
edema 53, 55, 61, 108, 116, 167

efavirenz. See non-nucleoside reverse	fimbrial. See biopsy
transcriptase inhibitors	fisting 223, 251, 258, 265, 332
electro-fulguration 168	Fitz-Hugh-Curtis syndrome 77
emergency contraception 312, 337, 338	fluconazole. See azole
endocarditis 175, 183	flucytosine 115
endometrial. See biopsy	fluorescent treponemal antibody
endometriosis 71	absorption (FTA-ABS) test 37, 65,
endometritis 73, 107, 275, 278	235, 236, 276
enfuvirtide/T20. See fusion inhibitor	fluoroquinolone. See quinolones
enteric 47, 203, 263	forensic 295, 299, 306–307, 345–348
See Sexually Transmitted Intestinal	four-glass localization test 86, 87
and Enteric Infections chapter	fusion inhibitor
enteritis 51, 56, 92–94, 267	enfuvirtide/T20 211
enuresis 102	
enzyme immunoassay (EIA) 37, 38, 65, 99, 149, 209, 236	G gamma benzene hexachloride 141, 143,
epididymitis 25, 47, 53-57, 175-176	273
epididymo-orchitis 53, 127, 176	gamma hydroxybutyrate (GHB).
erythema 53, 55, 61, 140, 167	See substance use
fallopian tube 73	gammaglobulin 283
meatal 44, 99	gastroenteritis 71
multiforme 64	gastrointestinal. See diarrhea
vulvar 49, 50, 108, 115	See nausea
erythromycin. See macrolides	See vomiting
excision 168, 228, 283	gay. See men who have sex with men
extragenital inoculation 223	See women who have sex with women
F	genital herpes. See herpes simplex virus
facultative bacteria. See bacteria	genital ulcer disease (GUD) 46, 59–68, 122, 223
famciclovir. See purine nucleoside analogs	genital warts. See Human Papillomavirus
femoral 224, 229	gentamicin. See aminoglycosides
fever 12, 20	giardia lamblia 93
See individual syndrome and infection chapters for specific	

considerations

gonorrhea (*Neisseria gonorrhoeae*) Gonococcal Infections chapter 178-179 31-34, 36, 43, 45 Epididymitis chapter 54, 56 Sexual Abuse in Peripubertal and Prepubertal Children chapter Pelvic Inflammatory Disease chapter 296-297 71, 73, 75 Sexual Assault in Postpubertal Sexually Transmitted Intestinal and Adolescents and Adults chapter 308 Enteric Infections chapter 93, 95 granuloma inquinale 10, 59, 62, 63, 66, Urethritis chapter 99, 102 67, 229 Vaginal Discharge chapter 106, 109, groove sign 224 110 GUD. See genital ulcer disease Chlamydial Infections chapter 126, 128 Gonococcal Infections chapter н 176-186 Human Immunodeficiency Virus Haemophilus ducreyi. See chancroid Infections chapter 199 haemorrhagic proctitis 224 Immigrants and Refugees chapter harm-reduction. See counselling 250 HAV. See hepatitis A virus Men Who Have Sex With Men/ Women Who Have Sex With Women HBIG. See hepatitis B immune globulin chapter 266-267 HBV. See hepatitis B virus Pregnancy chapter 274, 275–276 HCV. See hepatitis C virus Sexual Abuse of Peripubertal Health Canada Special Drug Access and Prepubertal Children chapter Program (SAP) 181, 182, 237, 296-297, 299-301 275-276, 277 Sexual Assault in Postpubertal hematochezia 95 Adolescents and Adults chapter 308, hematuria 99 310-311 hemodialysis 190, 191, 251 Sex Workers chapter 316 Henoch-Schönlein purpura 57 Travellers chapter 330 hepatitis A virus (HAV) 7, 12, 28, 38, 43 gram-negative 36, 74, 75, 99, 103, 109, 122, 123, 180, 308 Hepatitis B Virus Infections chapter 195 gram-positive 39, 84, 110 Human Immunodeficiency Virus Infections chapter 210 gram stain 33, 36, 39, 45 Men Who Have Sex With Men/ Epididymitis chapter 55 Women Who Have Sex With Women Pelvic Inflammatory Disease chapter chapter 264 Pregnancy chapter 283, 284 Urethritis chapter 99, 103 Substance Use chapter 322

360 Index

Travellers chapter 331

vaccination. See vaccine

Vaginal Discharge chapter 109, 110

Chancroid chapter 123

hepatitis B immune globulin (HBIG) 26, Herpes Simplex Virus (HSV) 10, 20, 25, 181, 191, 193, 284, 301, 311 28, 31, 36, 43 hepatitis B virus (HBV) 7, 11, 17, 26, 28, Genital Ulcer Disease chapter 59, 61, 38, 43 63, 64, 66 Hepatitis B Virus Infections chapter Pelvic Inflammatory Disease chapter 189-196 71 Human Immunodeficiency Virus Sexually Transmitted Intestinal and Infections chapter 210, 215 Enteric Infections chapter 93, 95 Immigrants and Refugees chapter Urethritis chapter 98 250-251 Chancroid chapter 122 Pregnancy chapter 274, 284 Genital Herpes Simplex Virus Sexual Abuse of Peripubertal Infections chapter 145-154 and Prepubertal Children chapter Genital Human Papillomavirus 298-299, 300-301 Infections chapter 161 Sexual Assault in Postpubertal Human Immunodeficiency Virus Adolescents and Adults chapter 309, Infections chapter 208, 214 310 Pregnancy chapter 281–282 Travellers chapter 331 Sexual Abuse of Peripubertal and vaccination. See vaccine Prepubertal Children chapter 297, hepatitis C virus (HCV) iii, 28, 43, 60 299 Hepatitis B Virus Infections chapter heterosexual 93, 145, 147, 198, 258, 189 262, 265 Human Immunodeficiency Virus high-grade squamous intra-epithelial Infections chapter 203 lesions (HSIL). See cervical dysplasia Lyphogranuloma Venereum chapter 223 highly active antiretroviral therapy (HAART). See HIV/AIDS *Immigrants and Refugees* chapter 250-251 HIV/AIDS iii, 4, 9, 10, 17, 18, 27, 28, 37, 43 Pregnancy chapter 284–285 CD4 107, 203, 206, 207, 210, 213,

236, 285

highly active antiretroviral therapy (HAART) 199, 212, 215, 285

Genital Ulcer Disease chapter 60, 61, 67

Pelvic Inflammatory Disease chapter

Sexually Transmitted Intestinal and Enteric Infections chapter 92

Vaginal Discharge chapter 107

Chancroid chapter 122, 124

hepatocellular carcinoma. See carcinoma

Appendix C 339

298, 301

309, 311

Sexual Abuse of Peripubertal and Prepubertal Children chapter

Sexual Assault in Postpubertal

Adolescents and Adults chapter

hepatosplenomegaly 233 heroin. See substance use

Hutchinson's teeth 233 Ectoparasitic Infestations chapter 142 Genital Human Papillomavirus hydrocele 55 Infections chapter 164, 165, 168 hydroxyzine 141 Hepatitis B Virus Infections chapter 189-190 Human Immunodeficiency Virus IgG. See antibody Infections chapter 198-215 IgM. See antibody Syphilis chapter 234, 236, 238-239, 243 imiguimod 166, 273, 283 Appendices 334–335, 339–340 immigrants 189, 234, 248-254, 331 See individual population chapters immunization. See vaccine for specific considerations incarceration 189, 199, 255, 315 homophobia 265 incubation period. See individual homosexual. See men who have sex infection chapters for specific with men considerations See women who have sex with infantile hypertrophic pyloric stenosis women (IHPS) 130, 184 HSIL. See cervical dysplasia infertility 72, 127, 176 HSV. See herpes simplex virus inflammatory bowel disease 71 HTLV. See Human T-lymphocyte virus inguinal nodes. See lymph node Human Immunodeficiency Virus (HIV). injection drug use (IDU). See HIV/AIDS See substance use human papillomavirus (HPV) 10, 26, 31, interferon 273, 283 34, 38, 43 alpha 195, 285 vaccine. See vaccine beta 168 warts (anal, genital, oral) 5, 34, Internet 4, 11, 13, 232, 262, 340 160-170, 210, 283 interstitial Genital Human Papillomavirus Infections chapter 160-170 cystitis 90 keratitis 233 Human Immunodeficiency Virus Infections chapter 210, 211, 215 intestinal and enteric infections 92-97. Inmates and Offenders chapter 259 263 Men Who Have Sex With Men/ intradermal nevi 163 Women Who Have Sex With Women intrapartum. See pregnancy chapter 263 intrauterine device (IUD). Pregnancy chapter 283-284 See contraception Sexual Abuse of Peripubertal and itraconazole. See azole Prepubertal Children chapter 299

J	lymph node 20, 123, 224, 266
	inguinal 20, 61, 224
Jarisch-Herxheimer reaction 244, 277	lymphadenopathy 12, 59, 99, 144, 207,
K	208, 233
K	femoral 62, 224, 229
Kaposi sarcoma 198, 208	inguinal 46, 224, 229
ketamine. See substance use	lymphogranuloma venereum (LGV) 25,
ketoconazole. See azole	28, 35
Klebsiella granulomatis. See granuloma inguinale	Genital Ulcer Disease chapter 59, 62, 63, 66
L	Sexually Transmitted Intestinal and Enteric Infections chapter 93–95
laboratory diagnosis 30–39	<i>Immigrants and Refugees</i> chapter 250, 251
See individual syndrome, infection and population chapters for specific considerations	Men Who Have Sex With Men/ Women Who Have Sex With Women chapter 262
laboratory testing methods 34	Travellers chapter 332
lactating 76, 132, 141, 143, 181, 275, 281	'
lactobacilli 39, 106, 110, 117	M
lamivudine (3TC). See nucleoside reverse transcriptase inhibitors	M-PCR. See multiplex polymerase
•	chain reaction
reverse transcriptase inhibitors	chain reaction macrolides 130, 184
reverse transcriptase inhibitors laparoscopy 73, 74, 129, 179	chain reaction
reverse transcriptase inhibitors laparoscopy 73, 74, 129, 179 LCR. See nucleic acid amplification test lesbian. See women who have sex	chain reaction macrolides 130, 184 azithromycin 67, 95, 100, 124, 129, 130–132, 181, 182, 213, 228, 239,
reverse transcriptase inhibitors laparoscopy 73, 74, 129, 179 LCR. See nucleic acid amplification test lesbian. See women who have sex with women	chain reaction macrolides 130, 184 azithromycin 67, 95, 100, 124, 129, 130–132, 181, 182, 213, 228, 239, 274, 300, 310
reverse transcriptase inhibitors laparoscopy 73, 74, 129, 179 LCR. See nucleic acid amplification test lesbian. See women who have sex with women Levitra. See vardenafil	chain reaction macrolides 130, 184 azithromycin 67, 95, 100, 124, 129, 130–132, 181, 182, 213, 228, 239, 274, 300, 310 clarithromycin 130, 213
reverse transcriptase inhibitors laparoscopy 73, 74, 129, 179 LCR. See nucleic acid amplification test lesbian. See women who have sex with women Levitra. See vardenafil LGV. See lymphogranuloma venereum lidocaine 168, 181, 182, 183, 184 ligase chain reaction (LCR). See nucleic	chain reaction macrolides 130, 184 azithromycin 67, 95, 100, 124, 129, 130–132, 181, 182, 213, 228, 239, 274, 300, 310 clarithromycin 130, 213 erythromycin 67, 124, 130–132, 184,
reverse transcriptase inhibitors laparoscopy 73, 74, 129, 179 LCR. See nucleic acid amplification test lesbian. See women who have sex with women Levitra. See vardenafil LGV. See lymphogranuloma venereum lidocaine 168, 181, 182, 183, 184 ligase chain reaction (LCR). See nucleic acid amplification test	chain reaction macrolides 130, 184 azithromycin 67, 95, 100, 124, 129, 130–132, 181, 182, 213, 228, 239, 274, 300, 310 clarithromycin 130, 213 erythromycin 67, 124, 130–132, 184, 228, 243, 273–275
reverse transcriptase inhibitors laparoscopy 73, 74, 129, 179 LCR. See nucleic acid amplification test lesbian. See women who have sex with women Levitra. See vardenafil LGV. See lymphogranuloma venereum lidocaine 168, 181, 182, 183, 184 ligase chain reaction (LCR). See nucleic	chain reaction macrolides 130, 184 azithromycin 67, 95, 100, 124, 129, 130–132, 181, 182, 213, 228, 239, 274, 300, 310 clarithromycin 130, 213 erythromycin 67, 124, 130–132, 184, 228, 243, 273–275 malaise 61, 140, 224, 233
reverse transcriptase inhibitors laparoscopy 73, 74, 129, 179 LCR. See nucleic acid amplification test lesbian. See women who have sex with women Levitra. See vardenafil LGV. See lymphogranuloma venereum lidocaine 168, 181, 182, 183, 184 ligase chain reaction (LCR). See nucleic acid amplification test Lindane. See gamma benzene	chain reaction macrolides 130, 184 azithromycin 67, 95, 100, 124, 129, 130–132, 181, 182, 213, 228, 239, 274, 300, 310 clarithromycin 130, 213 erythromycin 67, 124, 130–132, 184, 228, 243, 273–275 malaise 61, 140, 224, 233 medico-legal 36, 128, 202, 215, 296, 308 men who have sex with men (MSM)
reverse transcriptase inhibitors laparoscopy 73, 74, 129, 179 LCR. See nucleic acid amplification test lesbian. See women who have sex with women Levitra. See vardenafil LGV. See lymphogranuloma venereum lidocaine 168, 181, 182, 183, 184 ligase chain reaction (LCR). See nucleic acid amplification test Lindane. See gamma benzene hexachloride low-grade squamous intra-epithelial lesions (LSIL).	chain reaction macrolides 130, 184 azithromycin 67, 95, 100, 124, 129, 130–132, 181, 182, 213, 228, 239, 274, 300, 310 clarithromycin 130, 213 erythromycin 67, 124, 130–132, 184, 228, 243, 273–275 malaise 61, 140, 224, 233 medico-legal 36, 128, 202, 215, 296, 308 men who have sex with men (MSM) 9–11, 18, 262–272 See individual syndrome, infection and population chapters for specific

ndex 363

uncommon 147

hominis 71, 84

methadone 257, 321	N
methamphetamine (crystal meth). See substance use	NAAT. See nucleic acid amplification test
methylenedioxymethamphetamine	nasopharyngeal aspirate. See aspirate
(MDMA, ecstasy). See substance use	National Microbiology Laboratory (NML)
metronidazole 76, 111, 112, 116, 117,	11, 154, 225, 228, 234
274, 278, 279, 300, 310	nausea 74, 94, 191, 207
miconazole. See azole	needle-exchange 204, 257, 321
microhemagglutination for <i>Treponema</i> pallidum (MHA-TP) 37, 65, 276	Neisseria gonorrhoeae. See gonorrhea neonate
microimmunofluorescence (MIF) 35, 226, 227	Chlamydial Infections chapter 127, 131–134
micropapillomatosis labialis 163	Genital Ulcer Disease chapter 68
microsporidium 92	Gonococcal Infections chapter 184
miscarriage 14	Hepatitis B Virus Infections chapter
Mobiluncus sp. 106	189
molluscum contagiosum 163	Genital Herpes Simplex Virus
monogamy	Infections chapter 37, 152–154
mutual 15, 17	Syphilis chapter 237
serial 15	Pregnancy chapter 273–286
Motherisk 273	neoplasia 163, 165
motivational interviewing 18, 21, 265, 322	See cervical dysplasia
mucopurulent	See vulvar intra-epithelial neoplasia
discharge, cervical 25, 45, 109, 180	neurosyphilis. See syphilis
discharge, rectal 51, 94 discharge, urethral 98, 180	nevirapine. See non-nucleoside reverse transcriptase inhibitors (NNRTIs)
exudates 73	nocturia 99
Mueller Hinton agar 123	non-nucleoside reverse transcriptase inhibitors (NNRTIs)
multiplex polymerase chain reaction 124	efavirenz 211, 286
myalgia 84, 147, 207, 215, 224, 244	nevirapine 211, 286
Mycobacterium avium complex (MAC)	nonoxynol-9 (N-9) 18, 264, 316, 334
198, 208, 213	non-treponemal test (NTT) 37, 65, 237,
Mycobacterium tuberculosis. See tuberculosis	243, 308
Mycoplasma	norfloxacin. See quinolones
genitalium 71, 98	notifiable STI. See reportable STIs

notification	pap test. See papanicoulau
partner 8, 21–28	papanicoulau 12, 14, 43
See individual syndrome, infection	anal 165, 210, 215
and population chapters for specific considerations	cervical 38, 162-165, 169, 215
nucleic acid amplification test (NAAT)	parturition 152-154
31, 38, 43, 95, 124, 128, 176, 180, 226,	pearly penile papules 163
258, 307	pediatric examination 295
ligase chain reaction (LCR) 74, 177,	pediculocide 141
226, 258	pediculosis pubis.
polymerase chain reaction (PCR) 31, 66, 128, 146, 177, 201, 226, 234,	See ectoparasitic infestations
258, 275	pelvic inflammatory disease (PID) 25, 71–78, 107, 127, 129, 175, 274
nucleoside reverse transcriptase inhibitors (NRTIs)	penicillin
lamivudine (3TC) 195, 211	amoxicillin 132, 274, 310
stavudine (d4T) 211, 326	ampicillin 87–89
zidovudine (AZT) 211, 286	ampicillin/sulbactam 76
nucleotide reverse transcriptase	benzathine 96, 237, 238, 243, 277
inhibitor (NtRTI)	crystalline 239
tenofovir 211	procaine 237
nystatin 115	penicillin allergy. See allergy
	penicillin resistance. See resistance
0	pentamidine 213
ocular inflammatory disease 224	PEP. See prophylaxis
ofloxacin. See quinolones	perihepatitis 77, 175, 224
ophthalmia neonatorum 184, 275	perinatal 134, 161, 204, 286, 297
oral contraception. See contraception	permethrin 141, 143, 280, 281
oral sex 11–13, 16, 17, 60, 199, 201,	pharynx 20, 32, 178, 297
224, 263	Phthirus pubis.
oral-anal 43, 93, 258, 264, 265, 301,	See ectoparsitic infestations
311, 316, 332	PID. See pelvic inflammatory disease
oral-genital 93, 177, 191, 301, 311 osteochondritis 233	Pneumocystis jiroveci pneumonia (PCP) 198, 208, 213
OSteochondriis 233	pneumonia 127, 128, 208, 274
P	pneumonitis 131, 134, 208, 224, 282
	podofilox/podophyllotoxin 166, 273
p24 antigen testing 209	podophyllin 167, 283
P aeruginosa 54	

pap smear. See papanicoulau

point of care (POC) tests 30, 34, 209, 259, 316, 325	prostatitis 38, 54, 80–90, 101 prostatodynia 81
polymerase chain reaction (PCR). See nucleic acid amplification test	protease inhibitors
polymorphonuclear leukocytes (PMN) 36, 43, 47, 103, 108, 109, 180	ritonavir 211 saquinavir 211
portal hypertension 191	pruritus 98, 106, 280, 281
postexposure prophylaxis (PEP). See prophylaxis	pseudomembranous colitis 279 pseudomonas 53, 63, 84
pregnancy 12, 14, 18, 273–291	pubic lice (<i>Phthirus pubis</i>).
ectopic 48, 72–75, 127, 176	See ectoparasitic infestations
intrapartum 285	Public Health Agency of Canada 11, 56, 154, 181, 223, 257, 332, 340
termination. See abortion, therapeutic	purine nucleoside analogs
See individual syndrome and	acyclovir 150-154, 282
infection chapters for specific	famciclovir 150, 151, 195
considerations	valacyclovir 147, 150, 151
premature rupture of membranes (PROM) 112, 117, 278	pyoderma 64, 143
prenatal visit 127, 196, 273, 276, 285	pyrethrin-piperonyl butoxide 141, 280
preterm labour 111, 112, 117, 278	Q
prevention	
primary 4, 7, 8, 17, 18, 21	quality of evidence 1, 2
secondary 4, 7, 8, 17, 18, 21, 23	quinolone resistance. <i>See</i> resistance
See individual syndrome, infection and population chapters for specific	quinolones 76, 88, 96, 124, 130, 132, 181, 275
considerations	
	ciprofloxacin 67, 76, 88, 89, 95, 124,
primary care 7–28, 107	181, 310
primary care 7–28, 107 primary syphilis. <i>See</i> syphilis	181, 310 norfloxacin 88
	181, 310
primary syphilis. See syphilis	181, 310 norfloxacin 88 ofloxacin 76, 88, 89, 95, 129, 130, 181
primary syphilis. <i>See</i> syphilis procaine penicillin. <i>See</i> penicillin	181, 310 norfloxacin 88
primary syphilis. <i>See</i> syphilis procaine penicillin. <i>See</i> penicillin proctitis 92–97, 127, 175, 224, 225, 229	181, 310 norfloxacin 88 ofloxacin 76, 88, 89, 95, 129, 130, 181
primary syphilis. <i>See</i> syphilis procaine penicillin. <i>See</i> penicillin proctitis 92–97, 127, 175, 224, 225, 229 proctocolitis 62, 92–97, 224	181, 310 norfloxacin 88 ofloxacin 76, 88, 89, 95, 129, 130, 181
primary syphilis. <i>See</i> syphilis procaine penicillin. <i>See</i> penicillin proctitis 92–97, 127, 175, 224, 225, 229 proctocolitis 62, 92–97, 224 proctoscopy 225, 266	181, 310 norfloxacin 88 ofloxacin 76, 88, 89, 95, 129, 130, 181 R rape 249, 307
primary syphilis. <i>See</i> syphilis procaine penicillin. <i>See</i> penicillin proctitis 92–97, 127, 175, 224, 225, 229 proctocolitis 62, 92–97, 224 proctoscopy 225, 266 prodrome 146, 148, 150, 151, 282	181, 310 norfloxacin 88 ofloxacin 76, 88, 89, 95, 129, 130, 181 R rape 249, 307 rapid plasmid plasma reagin (RPR) 37, 65
primary syphilis. <i>See</i> syphilis procaine penicillin. <i>See</i> penicillin proctitis 92–97, 127, 175, 224, 225, 229 proctocolitis 62, 92–97, 224 proctoscopy 225, 266 prodrome 146, 148, 150, 151, 282 prophylaxis 131, 134, 213, 274, 282,	181, 310 norfloxacin 88 ofloxacin 76, 88, 89, 95, 129, 130, 181 R rape 249, 307 rapid plasmid plasma reagin (RPR) 37, 65 raves 11, 60, 263, 265

reinfection 23, 24, 28, 36, 127, 133, 236, Genital Ulcer Disease chapter 65, 255, 278 67, 68 Sexually Transmitted Intestinal and Reiter syndrome 127, 176 Enteric Infections chapter 95 reportable STIs 3, 9-11, 22, 24, 25, 185, 214, 241, 260 Chancroid chapter 123 See individual syndrome, infection Chlamydial Infections chapter 128 and population chapters for specific Genital Herpes Simplex Virus considerations Infections chapter 148, 149 resistance 37, 199, 212, 101, 129, 153 Hepatitis B Virus Infections chapter antimicrobial 185 191-192 azithromycin 239 Lymphogranuloma Venereum chapter 226 metronidazole 117 Syphilis chapter 234–235 penicillin 174 See individual population chapters quinolone 9, 44, 56, 95, 174, 177, for specific considerations 181, 310 serovar 25, 66, 93, 95, 126, 223, 226 restriction fragment length polymorphism (RFLP) 35, 66, 225, sex toys 11, 93, 223, 258, 265 226, 229 sex workers 9, 11, 60, 93, 122, 315-318, retinitis 208, 233 330 ribavirin 273, 285 sexual abuse. See abuse rifabutin 210, 213 sexual assault. See assault risk assessment 8, 12-15, 249, 259, shedding 266, 322 asymptomatic 146, 150, 152, 154, 282 mucosal 206, 214 S sigmoidoscopy 225 sadomasochism (S&M) 11 sildenafil citrate 263, 320 safer injection sites 321, 325 skin tags 163 SAP. See Health Canada Special Drug spectinomycin 181, 182, 183, 275 **Access Program** spermicide 17, 18, 246, 316 Sarcoptes scabiei. spontaneous abortion. See miscarriage See ectoparasitic infestations stavudine(d4T). See nucleoside reverse scables. See ectoparasitic infestations transcriptase inhibitors (NRTIs) sebaceous glands 163 stigma 249, 255, 326 secondary syphilis. See syphilis stillbirth 276, 277 sepsis 176, 275 strand displacement amplification (SDA) serial monogamy. See monogamy 177, 226 seroconversion 145, 147, 206 strawberry cervix. See cervix serology 34-38, 43 street youth 11, 93, 100, 126, 319

tertiary syphilis 237

tadalafil 263, 320 tattoo 14, 201, 251, 257, 258, 259, 332 teen. See adolescent tenesmus 94 terconazole. See azole tertiary syphilis. See syphilis test of cure 35, 36
See individual syndrome and infection chapters for specific considerations testicular 12, 53, 55, 86, 127, 176 torsion 53, 55
tetracycline 67, 76, 174, 273 doxycycline 67, 75, 76, 89, 95, 100, 103, 129–132, 182, 228, 238, 273, 274, 275, 310 thioglycolate hemin-based transport
media 123 toluidine red unheated serum test (TRUST) 37, 236 topical therapy 112–117, 130, 141, 143, 150, 168, 278, 279
Toxoplasma gondi 213 traditional medicine 252 transcription mediated amplification (TMA) 128, 226 transgendered 315
travel 13, 59, 60, 66, 122, 174, 190, 229, 249 travellers 330–333 Treponema pallidum particle agglutination (TP-PA) test 37, 65, 235, 276, 308 treponemal test 37, 65, 234, 236, 276, 308 trichloracetic acid 168

trichomoniasis (Trichomonas vaginalis) urethral strictures. See strictures 26, 28, 33, 38, 45 urethritis 25, 54, 55, 62, 87, 98-103, 117, Pelvic Inflammatory Disease chapter 146, 175, 297 71 urinary tract infection (UTI) 80, 85, Prostatitis chapter 84 86, 101 Urethritis chapter 98 urine 33-36, 38, 43, 345, 347 Vaginal Discharge chapter 106-111, See individual syndrome, infection 116, 117 and population chapters for specific considerations Human Immunodeficiency Virus Infections chapter 199 U.S. National Institutes of Health-Chronic Prostatitis Symptom Index 81-83 Pregnancy chapter 278 uveitis 233 Sexual Abuse in Peripubertal and Prepubertal Children chapter 299, 300 V Sexual Assault in Postpubertal vaccine Adolescents and Adults chapter 308, hepatitis A virus 7, 12, 28, 43, 210-211 trimethoprim-sulfamethoxazole See individual population chapters (TMP-SMX) 67, 88, 89, 213 for specific considerations tubal factor infertility 72 hepatitis B virus 7, 12, 26, 28, 38, 43, tuberculosis 54, 203, 210 190-196, 210-211 Mycobacterium tuberculosis 208, See individual population chapters 210 for specific considerations two-glass pre and post massage herpes simplex virus (HSV) 5, 147 screening test 87 human papillomavirus (HPV) 5 typhoid fever (Salmonella enterica vaginal 12, 16, 18, 20, 33-36, 38, 39, 45 serotype typhi) 263 discharge. See discharge Tzanck smear 36, 37, 148 See individual syndrome, infection U and population chapters for specific considerations ulcer 4, 31, 36 vaginal bleeding 127, 176 See individual syndrome, infection valacyclovir. and population chapters for specific See purine nucleoside analogs considerations vardenafil 262, 320 Ureaplasma urealyticum 71, 84 Venereal Disease Research Laboratory urethral 33, 35, 36, 38, 43 (VDRL) 37, 65, 235-236, 243, 276,

Index 369

discharge. See discharge

considerations

See individual syndrome, infection

and population chapters for specific

308

vertical transmission 273-286

vesicle 31, 36, 61, 65, 147, 148

vestibular papillae.

See micropapillomatosis labialis

Viagra. See sildenafil citrate

VIN. See vulvar intra-epithelial neoplasia viral hepatitis. See hepatitis A virus

See hepatitis B virus

See hepatitis C virus

viral load 37, 107, 199, 203, 210, 211, 212, 285, 286

viremia 181, 206, 207

vomiting 22, 74, 130, 132, 191, 207, 312

vulvar intra-epithelial neoplasia (VIN) 64, 163

vulvovaginal candidiasis. See candida

VVC. See candida

W

warts. *See* human papillomavirus
WBC. *See* white blood cell
western blot 37, 149, 209
wet-mount 33, 39, 45, 73, 108–110, 278, 296, 297, 308
white blood cell (WBC) 73, 110
window period 43, 68, 182, 202, 332
women who have sex with women
(WSW) 259, 262–267, 331

Υ

young adults 9, 10

See individual syndrome, infection and population chapters for specific considerations

youth 11, 15, 22

See individual syndrome, infection and population chapters for specific considerations

Z

zidovudine (AZT). See nucleoside reverse transcriptase inhibitors (NRTIs)