DISEASE CONTROL PRIORITIES • THIRD EDITION

# Major Infectious Diseases

# **DISEASE CONTROL PRIORITIES • THIRD EDITION**

# **Series Editors**

Dean T. Jamison

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# **DISEASE CONTROL PRIORITIES**

Budgets constrain choices. Policy analysis helps decision makers achieve the greatest value from limited available resources. In 1993, the World Bank published *Disease Control Priorities in Developing Countries* (*DCP1*), an attempt to systematically assess the cost-effectiveness (value for money) of interventions that would address the major sources of disease burden in low- and middle-income countries. The World Bank's 1993 *World Development Report* on health drew heavily on *DCP1*'s findings to conclude that specific interventions against noncommunicable diseases were cost-effective, even in environments in which substantial burdens of infection and undernutrition persisted.

*DCP2*, published in 2006, updated and extended *DCP1* in several aspects, including explicit consideration of the implications for health systems of expanded intervention coverage. One way that health systems expand intervention coverage is through selected platforms that deliver interventions that require similar logistics but deliver interventions from different packages of conceptually related interventions, for example, against cardiovascular disease. Platforms often provide a more natural unit for investment than do individual interventions. Analysis of the costs of packages and platforms—and of the health improvements they can generate in given epidemiological environments—can help to guide health system investments and development.

DCP3 differs importantly from DCP1 and DCP2 by extending and consolidating the concepts of platforms and packages and by offering explicit consideration of the financial risk protection objective of health systems. In populations lacking access to health insurance or prepaid care, medical expenses that are high relative to income can be impoverishing. Where incomes are low, seemingly inexpensive medical procedures can have catastrophic financial effects. DCP3 offers an approach to explicitly include financial protection as well as the distribution across income groups of financial and health outcomes resulting from policies (for example, public finance) to increase intervention uptake. The task in all of the DCP volumes has been to combine the available science about interventions implemented in very specific locales and under very specific conditions with informed judgment to reach reasonable conclusions about the impact of intervention mixes in diverse environments. DCP3's broad aim is to delineate essential intervention packages and their related delivery platforms to assist decision makers in allocating often tightly constrained budgets so that health system objectives are maximally achieved.

*DCP3*'s nine volumes are being published throughout 2015–18 in an environment in which serious discussion continues about quantifying the sustainable development goal (SDG) for health. *DCP3*'s analyses are well-placed to assist in choosing the means to attain the health SDG and assessing the related costs. Only when these volumes, and the analytic efforts on which they are based, are completed will we be able to explore SDG-related and other broad policy conclusions and generalizations. The final *DCP3* volume will report those conclusions. Each individual volume will provide valuable, specific policy analyses on the full range of interventions, packages, and policies relevant to its health topic.

More than 500 individuals and multiple institutions have contributed to *DCP3*. We convey our acknowledgments elsewhere in this volume. Here we express our particular gratitude to the Bill & Melinda Gates Foundation for its sustained financial support, to the InterAcademy Medical Panel (and its U.S. affiliate, the Institute of Medicine of the National Academy of Sciences), and to World Bank Publications. Each played a critical role in this effort.

Dean T. Jamison Rachel Nugent Hellen Gelband Susan Horton Prabhat Jha Ramanan Laxminarayan Charles N. Mock DISEASE CONTROL PRIORITIES • THIRD EDITION

# Major Infectious Diseases

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# **Contents**

Foreword	xiii	
Abbreviations		$x\nu$

 Major Infectious Diseases: Key Messages from Disease Control Priorities, Third Edition 1

King K. Holmes, Stefano Bertozzi, Barry R. Bloom, Prabhat Jha, Hellen Gelband, Lisa M. DeMaria, and Susan Horton

### PART 1 HIV/AIDS AND OTHER SEXUALLY TRANSMITTED INFECTIONS

2. Global Mortality and Morbidity of HIV/AIDS 29

Kristen Danforth, Reuben Granich, Danielle Wiedeman, Sanjiv Baxi, and Nancy Padian

3. HIV/AIDS Comorbidities: Impact on Cancer, Noncommunicable Diseases, and Reproductive Health 45

Corey Casper, Heidi Crane, Manoj Menon, and Deborah Money

4. HIV Care Continuum in Adults and Children: Cost-Effectiveness Considerations 67

Katherine Harripersaud, Margaret McNairy, Saeed Ahmed, Elaine J. Abrams, Harsha Thirumurthy, and Wafaa M. El-Sadr

5. Effectiveness and Cost-Effectiveness of Treatment as Prevention for HIV 91

Charles B. Holmes, Timothy B. Hallett, Rochelle P. Walensky, Till Bärnighausen, Yogan Pillay, and Myron S. Cohen

6. Prevention of Mother-to-Child Transmission of HIV and Syphilis 113

Grace John-Stewart, Rosanna W. Peeling, Carol Levin, Patricia J. Garcia, David Mabey, and John Kinuthia

## 7. Cost-Effectiveness of Interventions to Prevent HIV Acquisition 137

Geoff P. Garnett, Shari Krishnaratne, Kate L. Harris, Timothy B. Hallett, Michael Santos, Joanne E. Enstone, Bernadette Hensen, Gina Dallabetta, Paul Revill, Simon A. J. Gregson, and James R. Hargreaves

#### 8. Tailoring the Local HIV/AIDS Response to Local HIV/AIDS Epidemics 157

David Wilson and Jessica Taaffe

### Improving the Efficiency of the HIV/AIDS Policy Response: A Guide to Resource Allocation Modeling 179

James G. Kahn, Lori A. Bollinger, John Stover, and Elliot Marseille

#### 10. Sexually Transmitted Infections: Impact and Cost-Effectiveness of Prevention

203

Harrell W. Chesson, Philippe Mayaud, and Sevgi O. Aral

#### PART 2 TUBERCULOSIS

#### 11. Tuberculosis 233

Barry R. Bloom, Rifat Atun, Ted Cohen, Christopher Dye, Hamish Fraser, Gabriela B. Gomez, Gwen Knight, Megan Murray, Edward Nardell, Eric Rubin, Joshua Salomon, Anna Vassall, Grigory Volchenkov, Richard White, Douglas Wilson, and Prashant Yadav

#### PART 3 OTHER MAJOR INFECTIONS

#### 12. Malaria Elimination and Eradication 315

Rima Shretta, Jenny Liu, Chris Cotter, Justin Cohen, Charlotte Dolenz, Kudzai Makomva, Gretchen Newby, Didier Ménard, Allison Phillips, Allison Tatarsky, Roly Gosling, and Richard Feachem

#### 13. Malaria Control 347

Fabrizio Tediosi, Christian Lengeler, Marcia Castro, Rima Shretta, Carol Levin, Tim Wells, and Marcel Tanner

#### 14. Febrile Illness in Adolescents and Adults 365

John A. Crump, Paul N. Newton, Sarah J. Baird, and Yoel Lubell

# Cost-Effectiveness of Strategies for the Diagnosis and Treatment of Febrile Illness in Children 385

Joseph B. Babigumira, Hellen Gelband, and Louis P. Garrison Jr.

#### 16. Viral Hepatitis 401

Stefan Z. Wiktor

# 17. An Investment Case for Ending Neglected Tropical Diseases 411

Christopher Fitzpatrick, Uzoma Nwankwo, Edeltraud Lenk, Sake J. de Vlas, and Donald A. P. Bundy

# 18. Drug-Resistant Infections 433

Molly Miller-Petrie, Suraj Pant, and Ramanan Laxminarayan

DCP3 Series Acknowledgments 449
Volume and Series Editors 451
Contributors 453
Advisory Committee to the Editors 457
Reviewers 459
Index 461

# **Foreword**

Since the publication of the second edition of Disease Control Priorities in 2006, we have experienced some of the most substantial progress in infectious disease—caused mortality and morbidity. The number of annual deaths attributable to human immunodeficiency virus/acquired immune deficiency syndrome (HIV/AIDS) declined 50 percent between 2004 and 2015, thanks to an unprecedented expansion of life-saving antiretroviral therapy to over 18 million people (UNAIDS 2016); since 2006, mother-to-child transmission of HIV has been reduced to low levels, even in generalized epidemic settings (AVERT 2017). Similarly, fewer children and adults die from malaria, diarrheal diseases, and lower respiratory infections. Two infectious diseases are close to eradication: polio and dracunculiasis (Guinea worm disease).

This third edition of the *Disease Control Priorities* (*DCP3*) comes at a pivotal moment for infectious disease control and research. Its chapters clearly demonstrate that, despite the remarkable progress, infectious diseases remain a major threat to health worldwide—particularly in South Asia and Sub-Saharan Africa—but that an increasing range of highly cost-effective interventions is available.

As this volume amply illustrates, innovations in the prevention, diagnosis, and treatment of infectious diseases have been impressive. They include preexposure prophylaxis (PrEP) to prevent HIV infection, new forms of computer-based education for clinicians to manage sexually transmitted infections, HPV vaccines to prevent cervical cancer, and a cure for hepatitis C. The new attention to viral hepatitis in this volume is most welcome, as greatly improved control is now technically feasible—although the history of tuberculosis illustrates that a cure alone is insufficient to bring a disease under control. Much of the progress is due to political and technical leadership, greatly increased funding,

and improved delivery of interventions through health systems and other sectors. Community engagement is the key to success in many cases; a community buy-in to very simple, non-technological prevention mechanisms was instrumental in the sharp decline in dracunculiasis cases, from 130,000 in 2000 to only 22 in 2015 and 0 cases at the time of writing in 2017. However, dogs, which act as alternative hosts for the worm, present a threat to total eradication and remind us of the importance of a "One Health" approach.

At the same time, several epidemics and new pathogens have emerged, including the swine flu (H1N1) pandemic; the Middle East Respiratory Syndrome (MERS); the largest Ebola outbreak ever known in the West African region where it had never caused an outbreak; and an epidemic of Zika and associated neurological disorders. In particular, the collective failure to respond to the Ebola outbreak in a timely and coordinated fashion before it spiraled out of control—infecting over 28,000 people and causing over 11,000 deaths—was a wake-up call for the world. The disastrous impact of the Ebola epidemic prompted an urgent rethinking of how governments, nongovernmental organizations, and international organizations can better work to contain emerging disease threats in an increasingly interconnected world.

It is, however, noteworthy that almost as many people in the three Ebola-affected West African countries died from the disease's disruption on increasing mortality from HIV/AIDS, tuberculosis, and malaria as from Ebola itself (Parpia and others 2016). These three diseases, as well as diarrheal diseases and lower respiratory infections, continue to exact a heavy burden, particularly in Sub-Saharan Africa, where infectious diseases remain the leading cause of death. In 2015, over 1.8 million people worldwide died from tuberculosis (including

0.4 million among people with HIV) (WHO 2017); 1.1 million people from AIDS (UNAIDS 2016); and an estimated 429,000 people died from malaria (WHO 2016). In spite of real achievements in improved access to HIV treatment, over 2 million new infections occur each year, with hardly any decline in new infections over the past five years, and several subpopulations continue to be heavily affected. A critical review of current HIV strategies may be needed to achieve the United Nations goals of ending the AIDS epidemic. Lower respiratory infections remain a major persistent cause of death in children.

Many of these infectious diseases have sophisticated vaccines, diagnostics, and therapeutics available, but political, economic, and social factors limit the extent to which populations can benefit. Furthermore, in a world of growing resistance to antimicrobials and drug-resistant infections, we need to continue to develop innovations in biomedicine. We also need to improve incentives for rational antibiotic use, antimicrobial stewardship, and increased acceptance of the importance of prevention to avoid infection.

The global health agenda is an increasingly crowded space, and the cost-effectiveness of interventions is under growing scrutiny. While there is more information than ever regarding the cost-effectiveness of different interventions in a growing spectrum of contexts, hard choices remain in terms of allocating scarce funding to infectious diseases, especially in light of the complexities of fragile health systems, comorbidities with other infections and NCDs, structural factors that can undermine disease prevention, and treatment programs. One particularly valuable facet of DCP3 is that it demonstrates that some of the most effective steps we can take to reduce the burden of infections are not necessarily expensive, as exemplified by the low cost per disability-adjusted life year averted of condoms for female sex workers or insecticide-treated bednets. Often, the key is not just more, but smarter, investment, for example, better integration of services, strong community engagement, and targeted interventions based on the population most in need in specific locations. In addition to cost-effectiveness, key questions are

whether people will accept and use the interventions, whether the interventions are affordable and work in various parts of the real world, and what the best way is to deliver them.

If we are to reach the ambitious targets under the Sustainable Development Goals, we must focus not only on delivery of innovation but also on "innovation of delivery." One example might be new systems of community-based treatment for tuberculosis to minimize transmission in health care settings. *DCP3* helps us to think about improving health care delivery models through its unique focus on packages of interventions, and on the interrelationships among different kinds of interventions, at both the policy level and in terms of the outcomes across populations.

DCP3 is to be lauded for its focus on equity, recognizing that cost-effective intervention is not cost-effective if the financial burden falls on the poor. With this DCP3 volume on major infectious diseases, we have a highly pragmatic addition to the literature that will help policy makers across the world make smarter decisions to improve health sustainably and equitably in the ongoing fight against infectious disease threats, old and new.

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# **Abbreviations**

ACTs artemisinin-combination therapies

ADCs AIDS-defining cancers AEM AIDS Epidemic Model

AIDS acquired immune deficiency syndrome

AIM AIDS Impact Model ANC antenatal clinic

ART antiretroviral treatment

BCC behavior change communication

CBT community-based testing

CDC Centers for Disease Control and Prevention (U.S.)

CHMI controlled human malaria infection

CHW community health worker CI confidence interval

CQ chloroquine
CRP C-reactive protein
CSF cerebrospinal fluid
CVD cardiovascular disease

DAA direct-acting antiviral DALYs disability-adjusted life years

DBS dried blood spots

DCP2 Disease Control Priorities in Developing Countries, second edition

DCP3 Disease Control Priorities (third edition)
DDT dichloro-diphenyl-trichloroethane
DMPPT Decision Makers' Program Planning Tool

EID early infant diagnostic

ELISA enzyme-linked immunosorbent assay

EMOD Epidemiological Modeling

EMTCT elimination of mother-to-child transmission

EPP Epidemic Projection Package

FBC facility-based care
FI febrile illness
FSW female sex worker

G6PD glucose-6-phosphate dehydrogenase

GBD Global Burden of Disease
GDP gross domestic product
GHD Global Health Decisions
GMAP Global Malaria Action Plan

GMEP Global Malaria Eradication Program

GRADE Grading of Recommendations, Assessment, Development and Evaluation

GRNE Global Resource Needs Estimates

GTS Global Technical Strategy

HAI health care-associated infections HAT Human African trypanosomiasis

HBC home-based care HBV hepatitis B virus

HCC hepatocellular carcinoma

HCV hepatitis C virus

HDL high-density lipoprotein cholesterol

HHV-8 human herpes virus 8 HICs high-income countries

HIV/AIDS human immunodeficiency virus/acquired immune deficiency syndrome

HIVAM HIV-associated malignancies
HIV human immunodeficiency virus

HPV human papillomavirus

HRQL health and health-related quality of life

HSV herpes simplex virus
HTC HIV testing and counseling
HTS HIV/AIDS testing service

ICER incremental cost-effectiveness ratio

IDU injecting drug user

IMAI Integrated Management of Adolescent and Adult Illness

IMCI Integrated Management of Childhood Illness

IRS indoor residual spraying ITN insecticide-treated net

IVM integrated vector management
JEV Japanese encephalitis virus

KS Kaposi sarcoma

LDL low-density lipoprotein cholesterol LGBT lesbian, gay, bisexual, and transgender

LICs low-income countries

LLIN long-lasting insecticide-treated nets LMICs low- and middle-income countries

LTFU loss to follow-up

MCH maternal and child health
MDA mass drug administration
MDG Millennium Development Goal
MDR-TB multidrug-resistant tuberculosis
MMC medical male circumcision
MERS Middle East respiratory syndrome

MCC mobile clinic care

MRSA Methicillin-resistant Staphylococcus aureus

MSM men who have sex with men MTCT mother-to-child transmission

NADCs non–AIDS-defining cancer NBS National Bureau of Statistics

NCC noncommunicable chronic comorbidities

NCDs noncommunicable diseases

NECT nifurtimox-eflornithine combination therapy

NHL non-Hodgkin lymphoma

NNRTI non-nucleoside reverse transcriptase inhibitors NRTI nucleoside reverse transcriptase inhibitors

NSP needle and syringe program NTDs neglected tropical diseases

NT-NMFI nontreatable nonmalaria febrile illness

OOP out of pocket

PCR polymerase chain reaction

PEPFAR President's Emergency Plan for AIDS Relief (United States)

PMI President's Malaria Initiative

PMTCT prevention of mother-to-child transmission

POC point-of-care POCT point-of-care test

PPT periodic presumptive treatment PrEP preexposure prophylaxis

PROMISE Promoting Maternal-Infant Survival Everywhere

QALY quality-adjusted life year

RNM Resource Needs Model

RBM Roll Back Malaria (Partnership)

RPR rapid plasma reagin

RCT randomized controlled trial RDTs rapid diagnostic tests

SAC school-age children

SARS severe acute respiratory syndrome SDG Sustainable Development Goal

SMART Strategies for Management of Antiretroviral Therapy

STDs sexually transmitted diseases
SP sulfadoxine/pyrimethamine
STIs sexually transmitted infections
STH soil-transmitted helminthiases

SW sex workers

TasP treatment as prevention

TB tuberculosis

TCPs target candidate profiles

T-NMFI treatable nonmalaria febrile illness

UI uncertainty interval

UNAIDS Joint United Nations Programme on HIV/AIDS

UN United Nations

VCT voluntary counseling and testing

VIMT vaccines that interrupt malaria transmission VMMC voluntary male medical circumcision

WHO World Health Organization
WWR What Works Reviews

YLDs years lived with disability

YLLs years of life lost