ELEVENTH Edition

APPLIED THERAPEUTICS

The Clinical Use of Drugs

Caroline S. Zeind Michael G. Carvalho



. Wolters Kluwer

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Caroline S. Zeind, PharmD

Associate Provost for Academic and International Affairs Chief Academic Officer Worcester, Massachusetts and Manchester, New Hampshire Campuses Professor of Pharmacy Practice Academic Affairs MCPHS University Boston, Massachusetts

Michael G. Carvalho, PharmD, BCPP

Assistant Dean of Interprofessional Education Professor and Chair Department of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

• Wolters Kluwer

Philadelphia • Baltimore • New York • London Buenos Aires • Hong Kong • Sydney • Tokyo Acquisitions Editor: Matt Hauber
Development Editor: Andrea Vosburgh
Editorial Coordinator: Annette Ferran
Editorial Assistant: Brooks Phelps
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Dedication

The editors wish to express their sincere thanks to Dr. Mary Anne Koda-Kimble and Dr. Lloyd Young who created Applied Therapeutics, the gold standard textbook used in the teaching of patient-centered drug therapeutics. Their contributions vision and legendary have inspired generations of health profession students, faculty, and clinicians. We are grateful to the past editors and current editorial team and contributors, as well as faculty colleagues and clinicians across the globe for their dedication and tremendous to their patients their communities. They have shaped and inspired the next generation of health care professionals through innovative teaching, mentoring, and dissemination of their scholarship. Finally, we wish to thank our families for their patience and understanding as we embarked on this journey and completed this eleventh edition of the textbook.

Preface

It has been over forty years since the first edition of Applied Therapeutics: The Clinical Use of Drugs was published, and the landscape of health care has changed dramatically. While we have seen tremendous scientific and technologic advancements transform personalized medicine, we also realize the significant challenges that we face within our increasingly complex health care delivery system. More than ever, we are in need of health professionals who are able to think critically and to utilize problem-solving skills to improve patient outcomes. Approximately four decades later, the founding principle for this textbook—a patient-centric, case-based approach to learning—remains the cornerstone of health professions education. Our authors present approximately 900 patient cases that stimulate the reader to integrate and apply therapeutic principles in the context of specific clinical situations. Health profession students and practitioners gain a glimpse into the minds of clinicians as they work to assess and solve therapeutic problems and develop their own critical-thinking and problem-solving skills.

Readers familiar with past editions of the book will notice that the overall design of the book is consistent with the tenth edition, which incorporates a Core Principles section at the beginning of each chapter, providing the most important "take home" information from the chapter. Each Core Principle is mapped to specific cases within the chapter where the principle is discussed in detail. Key references and websites are listed at the end of each chapter, whereas the full reference lists for each chapter have been moved online.

Building upon the excellent foundation of case-based learning provided in previous editions, the eleventh edition has incorporated changes to meet the evolving educational needs of health profession educators and students across the globe. The editors and contributors have utilized the five Institute of Medicine (IOM) core competencies as a broad framework for proposing case studies and questions within the textbook: patient-centered care; interdisciplinary teams; evidence-based practice; quality improvement; and informatics. In addition, the Accreditation Council for Pharmacy Education (ACPE) Accreditation Standards 2016, the Center for the Advancement of Pharmacy Education (CAPE) Educational Outcomes, and the North American Pharmacist Licensure Examination (NAPLEX) revised competency statements have served as road maps for the editorial team and contributors in designing the eleventh edition.

Featuring contributions from more than 200 experienced clinicians, every chapter

has been revised and updated to reflect our ever-changing knowledge of drugs and the application of this knowledge to the individualized therapy of patients. Content within several sections has been extensively reorganized, with new chapters introduced to expand important topics. Among these are seven new chapters within the General Principles, Immunologic Disorders, Rheumatic and Musculoskeletal Diseases, Neurologic Disorders, Psychiatric Disorders & Substance Abuse, and Hematology and Oncology sections. Of particular note are featured new chapters in the General Principles section on Drug Interactions, Pharmacogenomics and Personalized Medicine, and Interprofessional Education and Practice. In addition, a chapter has been redesigned to focus on Care of the Critically Ill Adult, which now complements the chapter on Care of the Critically Ill Child.

Given the importance of incorporating Interprofessional Education (IPE) within the didactic, practicum, and clinical settings, we have added an array of IPE case studies prepared by contributors that are representative of various sections of the textbook. These IPE cases, along with answer guides, will be available for instructors on the textbook's website (see the "Additional Resources" section).

We welcome your feedback as we undertake planning for the next edition. The authors have drawn on information from the literature, current standards, and their own clinical experiences to share the process involved in making sound and thoughtful therapeutic decisions. However, it remains the responsibility of every practitioner to evaluate the appropriateness of a particular opinion in the context of the actual clinical situation, bearing in mind any recent developments in the field. We strongly urge students and practitioners to consult several appropriate information sources when working with new and unfamiliar drugs.

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Acknowledgments

We are deeply indebted to the many dedicated people who have given of themselves to complete the eleventh edition of Applied Therapeutics: The Clinical Use of Drugs. We are grateful to our contributing authors who have worked tirelessly to provide the highest quality of work, while balancing numerous responsibilities as educators, clinicians, and researchers. We are grateful for the exceptional work of our twenty-six section editors, who provided critical feedback necessary both in the organizational structure of the textbook and in the individual editing of chapters; without their dedication and support, this edition would not be possible. In particular, we wish to recognize those returning section editors as they have been a guiding force for the eleventh edition: Drs. Jean M. Nappi, Timothy J. Ives, Marcia L. Buck, Judith L. Beizer, and Myrna Y. Munar. We sincerely thank the past editorial team of Applied Therapeutics, with special thanks to Dr. Brian K. Alldredge and Dr. B. Joseph Guglielmo for their guidance and support and contributions to the eleventh edition. We would also like to thank Facts and Comparisons for allowing us to use their data for the construction of some of our tables.

The team from Wolters Kluwer, Matt Hauber, Andrea Vosburgh, and Annette Ferran, deserve special recognition for their efforts. Their exceptional patience, attention to detail, and guidance have been critical to the success of this project. We sincerely thank Tara Slagle (project management) and Samson Premkumar (production) for their assistance in completing this edition. Most importantly, we wish to acknowledge the love, understanding, and unwavering support of our spouses and families. They selflessly gave to us early mornings, late nights, weekends, and vacation time that we spent writing and editing.

Consistent with past editions, we continue to dedicate our work to our students who inspire us and to the many patients we have been privileged to care for and who have taught us invaluable lessons. We also dedicate the eleventh edition to those clinicians and educators who have served as pioneering leaders and role models in the delivery of patient-centered care using team-based approaches.

Additional Resources

The eleventh edition of Applied Therapeutics: The Clinical Use of Drugs includes additional resources for both instructors and students, available on the book's companion website at http://thepoint.lww.com/AT11e.

STUDENT RESOURCES

Students who have purchased Applied Therapeutics: The Clinical Use of Drugs, Eleventh Edition have access to the following additional resources for each chapter:

A full online reference list for that chapter

INSTRUCTOR RESOURCES

Approved adopting instructors will be given access to the following additional resources:

- Interprofessional Education (IPE) Case Studies and Answer Guides
- Each chapter will map which NAPLEX competency statements are addressed with it, which will be a helpful resource for instructors.

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Section Editors

Michael C. Angelini, PharmD, MA, BCPP

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Judith L. Beizer, PharmD, CGP, FASCP

Clinical Professor Department of Clinical Pharmacy Practice College of Pharmacy & Allied Health Professions St. John's University Jamaica, New York

Marcia L. Buck, PharmD, FCCP, FPPAG

Professor
Department of Pediatrics
School of Medicine
Clinical Coordinator, Pediatrics
Department of Pharmacy
University of Virginia
Charlottesville, Virginia

Michael G. Carvalho, PharmD, BCPP

Assistant Dean of Interprofessional Education Professor and Chair Department of Pharmacy Practice School of Pharmacy—Boston MCPHS University Boston, Massachusetts

Judy W. Cheng, PharmD, MPH, BCPS, FCCP

Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

R. Rebecca Couris, PhD, RPh

Professor of Nutrition Science and Pharmacy Practice Department of Pharmacy Practice, School of Pharmacy–Boston MCPHS University

Steven Gabardi, PharmD, BCPS, FAST, FCCP

Abdominal Organ Transplant Clinical Specialist & Program Director PGY-2 Organ Transplant Pharmacology Residency

Brigham and Women's Hospital

Departments of Transplant Surgery/Pharmacy/Renal Division

Assistant Professor of Medicine

Harvard Medical School

Boston, Massachusetts

Jennifer D. Goldman, BS, PharmD, CDE, BC-ADM, FCCP

Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Christy S. Harris, PharmD, BCPS, BCOP

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Timothy R. Hudd, PharmD, AE-C

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Timothy J. Ives, PharmD, MPH, FCCP, BCPS

Professor Eshelman School of Pharmacy The University of North Carolina at Chapel Hill Chapel Hill, North Carolina

Susan Jacobson, MS, EdD, RPh

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Maria D. Kostka-Rokosz, PharmD

Assistant Dean of Academic Affairs Professor of Pharmacy Practice School of Pharmacy—Boston MCPHS University Boston, Massachusetts

Trisha LaPointe, PharmD, BCPS

Associate Professor of Pharmacy Practice

School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Michele Matthews, PharmD, CPE, BCACP

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

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Susan L. Mayhew, PharmD, BCNSP, FASHP

Professor and Dean Appalachian College of Pharmacy Oakwood, Virginia

William W. McCloskey, BA, BS, PharmD

Professor and Vice-Chair Department of Pharmacy Practice School of Pharmacy-Boston MCPHS University Boston, Massachusetts

Myrna Y. Munar, PharmD

Associate Professor
Department of Pharmacy Practice
College of Pharmacy
Oregon State University
Oregon Health and Science University
Portland, Oregon

Jean M. Nappi, PharmD, FCCP, BCPS AQ-Cardiology

Professor Clinical Pharmacy and Outcome Sciences South Carolina College of Pharmacy Medical University of South Carolina Charleston, South Carolina

Kamala M. Nola, PharmD, MS

Professor and Vice-Chair Department of Pharmacy Practice Lipscomb University College of Pharmacy Nashville, Tennessee

Dorothea C. Rudorf, PharmD, MS

Professor of Pharmacy Practice School of Pharmacy–Boston

Carrie A. Sincak, PharmD, BCPS, FASHP

Assistant Dean for Clinical Affairs and Professor Department of Pharmacy Practice Midwestern University Chicago College of Pharmacy Downers Grove, Illinois

Timothy E. Welty, PharmD, FCCP

Professor Department of Pharmacy Practice University of Kansas School of Pharmacy Lawrence, Kansas

G. Christopher Wood, PharmD, FCCP, FCCM, BCPS

Associate Professor of Clinical Pharmacy University of Tennessee Health Science Center College of Pharmacy Memphis, Tennessee

Kathy Zaiken, PharmD

Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Caroline S. Zeind, PharmD

Associate Provost for Academic and International Affairs
Chief Academic Officer
Worcester, Massachusetts and Manchester, New Hampshire Campuses
Professor of Pharmacy Practice
Academic Affairs
MCPHS University
Boston, Massachusetts



Steven R. Abel, PharmD, FASHP

Professor of Pharmacy Practice Associate Provost for Engagement Purdue University West Lafayette, Indiana

Jessica L. Adams, PharmD, BCPS, AAHIVP

Assistant Professor of Clinical Pharmacy
HIV and Infectious Diseases Specialist
Department of Pharmacy Practice and Pharmacy Administration
Philadelphia College of Pharmacy
University of the Sciences
Philadelphia, Pennsylvania

Brian K. Alldredge, PharmD

Professor and Vice Provost University of California—San Francisco San Francisco, California

Mary G. Amato, PharmD, MPH, BCPS

Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Jaime E. Anderson, PharmD, BCOP

Oncology Clinical Pharmacy Specialist MD Anderson Medical Center University of Texas Houston, Texas

Michael C. Angelini, PharmD, MA, BCPP

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Albert T. Bach, PharmD

Assistant Professor of Pharmacy Practice School of Pharmacy

Jennifer H. Baggs, PharmD, BCPS, BCNSP

Clinical Assistant Professor University of Arizona Tucson, Arizona

David T. Bearden, PharmD

Clinical Professor and Chair
Department of Pharmacy Practice
Clinical Assistant Director
Department of Pharmacy Services
College of Pharmacy
Oregon State University
Oregon Health and Science University
Portland, Oregon

Sandra Benavides, PharmD, FCCP, FPPAG

Professor
Assistant Dean for Programmatic Assessment and Accreditation
Interim Chair
Department of Clinical and Administrative Sciences
Larkin Health Sciences Institute College of Pharmacy

Paul M. Beringer, PharmD, FASHP, FCCP

Associate Professor Department of Clinical Pharmacy University of Southern California Los Angeles, California

Snehal H. Bhatt, PharmD, BCPS

Associate Professor of Pharmacy Practice School of Pharmacy—Boston MCPHS University Clinical Pharmacist Beth Israel Deaconess Medical Center Boston, Massachusetts

Jeff F. Binkley, PharmD, BCNSP, FASHP

Administrative Director of Pharmacy Maury Regional Medical Center and Affiliates Columbia, Tennessee

Marlo Blazer, PharmD, BCOP

Assistant Director Xcenda, an AmerisourceBergen Company Columbus, Ohio

KarenBeth H. Bohan, PharmD, BCPS

Professor and Founding Chair Department of Pharmacy Practice School of Pharmacy and Pharmaceutical Sciences Binghamton University Binghamton, New York

Suzanne G. Bollmeier, PharmD, BCPS, AE-C

Professor of Pharmacy Practice School of Pharmacy—Boston St. Louis College of Pharmacy St. Louis, Missouri

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p. xii

Laura M. Borgelt, PharmD, BCPS

Associate Dean of Administration and Operations Professor Departments of Clinical Pharmacy and Family Medicine University of Colorado Anschutz Medical Campus Skaggs School of Pharmacy Aurora, Colorado

Jolene R. Bostwick, PharmD, BCPS, BCPP

Clinical Associate Professor Department of Clinical, Social, and Administrative Sciences University of Michigan College of Pharmacy Ann Arbor, Michigan

Nicole J. Brandt, PharmD, MBA, CGP, BCPP, FASCP

Executive Director
Peter Lamy Center on Drug Therapy and Aging
Professor
University of Maryland School of Pharmacy
Baltimore, Maryland

Marcia L. Buck, PharmD, FCCP, FPPAG

Professor Department of Pediatrics School of Medicine Clinical Coordinator, Pediatrics Department of Pharmacy University of Virginia Charlottesville, Virginia

Deanna Buehrle, PharmD

Infectious Diseases Clinical Specialist University of Pittsburgh Medical Center Presbyterian Pittsburgh, Pennsylvania

Sara K. Butler, PharmD, BCPS, BOCP

Clinical Pharmacy Specialist, Medical Oncology Barnes-Jewish Hospital Saint Louis, Missouri

Beth Buyea, MHS, PA-C

Assistant Professor Tufts University, School of Medicine Boston, Massachusetts

Charles F. Caley, PharmD, BCCP

Clinical Professor School of Pharmacy University of Connecticut Storrs, Connecticut

Joseph Todd Carter, PharmD

Assistant Professor of Pharmacy Practice Appalachian College of Pharmacy Oakwood, Virginia Primary Care Centers of Eastern Kentucky Hazard, Kentucky

Michael G. Carvalho, PharmD, BCPP

Assistant Dean of Interprofessional Education Professor and Chair Department of Pharmacy Practice School of Pharmacy—Boston MCPHS University Boston, Massachusetts

Jamie J. Cavanaugh, PharmD, CPP, BCPS

Assistant Professor of Clinical Education, Pharmacy Assistant Professor of Medicine University of North Carolina at Chapel Hill Chapel Hill, North Carolina

Michelle L. Ceresia, PharmD, FACVP

Associate Professor of Pharmacy Practice
School of Pharmacy—Boston
MCPHS University
Boston, Massachusetts
Adjunct Associate Professor
Department of Clinical Sciences
Cummings Veterinary School of Medicine at Tufts University
North Grafton, Massachusetts

Laura Chadwick, PharmD

Clinical Specialist in Pharmacogenomics Boston Children's Hospital

Michelle L. Chan, PharmD, BCPS

Clinical Pharmacy Specialist Infectious Diseases Methodist Hospital of Southern California Arcadia, California

Lin H. Chen, MD, FACP, FASTMH

Associate Professor of Medicine Harvard Medical School Boston, Massachusetts Director of the Travel Medicine Center Mount Auburn Hospital Cambridge, Massachusetts

Steven W. Chen, PharmD, FASHP, FNAP

Associate Professor and Chair
Titus Family Department of Clinical Pharmacy
William A. Heeres and Josephine A. Heeres Endowed Chair in Community Pharmacy
University of Southern California School of Pharmacy
Los Angeles, California

Judy W. Cheng, PharmD, MPH, BCPS, FCCP

Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Michael F. Chicella, PharmD, FPPAG

Pharmacy Clinical Manager Children's Hospital of The King's Daughters Norfolk, Virginia

Jennifer W. Chow, PharmD

Director of Professional Development and Education Pediatric Pharmacy Advocacy Group Memphis, Tennessee

Cary R. Chrisman, PharmD

Assistant Professor
Department of Clinical Pharmacy
University of Tennessee College of Pharmacy
Clinical Pharmacist, Department of Pharmacy
Methodist Medical Center
Memphis and Oak Ridge, Tennessee

Edith Claros, PhD, MSN, RN, APHN-BC

Assistant Dean and Associate Professor School of Nursing MCPHS University Worcester, Massachusetts

John D. Cleary, PharmD, FCCP, BCPS

Director of Pharmacy St. Dominic-Jackson Memorial Hospital Schools of Medicine and Pharmacy University of Mississippi Medical Center Jackson, Mississippi

Michelle Condren, PharmD, BCPPS, AE-C, CDE, FPPAG

Professor and Department Chair University of Oklahoma College of Pharmacy University of Oklahoma School of Community Medicine Tulsa, Oklahoma

Amanda H. Corbett, PharmD, BCPS, FCCP

Clinical Associate Professor Eshelman School of Pharmacy and School of Medicine Global Pharmacology Coordinator Institute for Global Health and Infectious Diseases University of North Carolina Chapel Hill, North Carolina

Mackenzie L. Cottrell, PharmD, MS, BCPS, AAHIVP

Research Assistant Professor UNC Eshelman School of Pharmacy University of North Carolina at Chapel Hill Chapel Hill, North Carolina

R. Rebecca Couris, PhD, RPh

Professor of Nutrition Science and Pharmacy Practice Department of Pharmacy Practice, School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Steven J. Crosby, MA, BSP, RPh, FASCP

Assistant Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Jason Cross, PharmD

Associate Professor Pharmacy Practice School of Pharmacy–Worcester/Manchester MCPHS University Worcester, Massachusetts

Sandeep Devabhakthuni, PharmD, BCPS-AQ Cardiology

Assistant Professor of Cardiology/Critical Care University of Maryland School of Pharmacy Baltimore, Maryland

Andrea S. Dickens, PharmD, BCOP

Clinical Pharmacy Specialist MD Anderson Cancer Center University of Texas Houston, Texas

Lisa M. DiGrazia, PharmD, BCPS, BCOP

Director, Medical Affairs Amneal Biosciences Bridgewater, New Jersey

Suzanne Dinsmore, BSP, PharmD, CGP

Assistant Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Betty J. Dong, PharmD, FASHP, FAPHA, FCCP, AAHIVP

Professor of Clinical Pharmacy and Family and Community Medicine Department of Clinical Pharmacy Schools of Pharmacy and Medicine University of California, San Francisco San Francisco, California

Richard H. Drew, PharmD, MS, FCCP

Professor and Vice-Chair of Research and Scholarship Campbell University College of Pharmacy and Health Sciences Buies Creek, North Carolina Associate Professor of Medicine (Infectious Diseases) Duke University School of Medicine Durham, North Carolina

Robert L. Dufresne, PhD, PhD, BCPS, BCPP

INBRE Behavioral Science Coordinator and Professor College of Pharmacy University of Rhode Island Kingston, Rhode Island Psychiatric Pharmacotherapy Specialist PGY-2 Psychiatric Pharmacy Residency Program Director Providence VA Medical Center Providence, Rhode Island

Kaelen C. Dunican, PharmD

Professor of Pharmacy Practice School of Pharmacy–Worcester/Manchester MCPHS University

Brianne L. Dunn, PharmD

Associate Dean for Outcomes Assessment & Accreditation Clinical Associate Professor Department of Clinical Pharmacy and Outcomes Sciences University of South Carolina College of Pharmacy Columbia, South Carolina

Robert E. Dupuis, PharmD, FCCP

Clinical Professor of Pharmacy Eshelman School of Pharmacy University of North Carolina at Chapel Hill Chapel Hill, North Carolina

Cheryl R. Durand, PharmD

Associate Professor of Pharmacy Practice School of Pharmacy–Worcester/Manchester MCPHS University Manchester, New Hampshire

Megan J. Ehret, PharmD, MS, BCPP

Behavior Health Clinical Pharmacy Specialist United States Department of Defense Fort Belvoir Community Hospital Fort Belvoir, Virginia

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p. xiv

Carol Eliadi, EdD, JD, NP-BC

Professor and Dean of Nursing MCPHS University School of Nursing-Worcester, Massachusetts and Manchester, New Hampshire Campuses

Shareen Y. El-Ibiary, PharmD, FCCP, BCPS

Professor of Pharmacy Practice Department of Pharmacy Practice Midwestern University College of Pharmacy–Glendale Glendale, Arizona

Katie Dillinger Ellis, PharmD

Clinical Specialist
Neonatal/Infant Intensive Care
Department of Pharmacy
The Children's Hospital of Philadelphia
Philadelphia, Pennsylvania

Justin C. Ellison, PharmD, BCPP

Clinical Pharmacy Specialist–Mental Health Providence Veterans Affairs Medical Center Providence, Rhode Island

Rachel Elsey, PharmD, BCOP

Clinical Pharmacist Avera Cancer Institute South Dakota State University Sioux Falls, South Dakota

Gregory A. Eschenauer, PharmD, BCPS (AQ-ID)

Clinical Assistant Professor University of Michigan Ann Arbor, Michigan

John Fanikos, MBA, RPh

Executive Director of Pharmacy
Brigham and Women's Hospital
Adjunct Associate Professor of Pharmacy Practice
MCPHS University
Department of Pharmacy Practice, School of Pharmacy–Boston
Boston, Massachusetts

Elizabeth Farrington, PharmD, FCCP, FCCM, FPPAG, BCPS

Pharmacist III-Pediatrics Department of Pharmacy New Hanover Regional Medical Center Wilmington, North Carolina

Erika Felix-Getzik, PharmD

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Jonathan D. Ference, PharmD

Assistant Dean of Assessment and Alumni Affairs Associate Professor of Pharmacy Practice Director of Pharmacy Care Labs Nesbitt School of Pharmacy Wilkes University Wilkes-Barre, Pennsylvania

Kimberly Ference, PharmD

Associate Professor Department of Pharmacy Practice Nesbitt College of Pharmacy and Nursing Wilkes University Wilkes-Barre, Pennsylvania

Victoria F. Ferraresi, PharmD, FASHP, FCSHP

Director of Pharmacy Services Pathways Home Health and Hospice Sunnyvale, California

Joseph W. Ferullo, PharmD

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Christopher K. Finch, PharmD, BCPS, FCCM, FCCP

Director of Pharmacy Methodist University Hospital Associate Professor College of Pharmacy University of Tennessee Memphis, Tennessee

Douglas N. Fish, PharmD, BCPS-AQ ID

Professor and Chair
Department of Clinical Pharmacy
Skaggs School of Pharmacy and Pharmaceutical Science
University of Colorado
Clinical Specialist in Critical Care/Infectious Diseases
University of Colorado Hospital
Aurora, Colorado

Jeffrey J. Fong, PharmD, BCPS

Associate Professor of Pharmacy Practice School of Pharmacy–Worcester/Manchester MCPHS University Worcester, Massachusetts

Andrea S. Franks, PharmD, BCPS

Associate Professor, Clinical Pharmacy and Family Medicine College of Pharmacy and Graduate School Medicine University of Tennessee Health Science Center Knoxville, Tennessee

Kristen N. Gardner, PharmD

Clinical Pharmacy Specialist–Behavioral Health Highline Behavioral Clinic Kaiser Permanente Colorado Denver, Colorado

Virginia L. Ghafoor, PharmD

Pharmacy Specialist—Pain Management University of Minnesota Medical Center Minneapolis, Minnesota

Brooke Gildon, PharmD, BCPPS, BCPS, AE-C

Associate Professor of Pharmacy Practice Southwestern Oklahoma State University College of Pharmacy Weatherford, Oklahoma

Ashley Glode, PharmD, BCOP

Assistant Professor Department of Clinical Pharmacy Skaggs School of Pharmacy and Pharmaceutical Sciences University of Colorado Anschutz Medical Campus Aurora, Colorado

p. xiv

p. xv

Jeffery A. Goad, PharmD, MPH, FAPhA, PCPhA, FCSHP

Professor and Chair Department of Pharmacy Practice School of Pharmacy Chapman University Irvine, California

Jennifer D. Goldman, BS, PharmD, CDE, BC-ADM, FCCP

Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Joel Goldstein, MD

Assistant Clinical Professor Harvard Medical School Division of Child/Adolescent Psychology Cambridge Health Alliance Cambridge, Massachusetts

Luis S. Gonzalez, III, PharmD, BCPS

Manager Clinical Pharmacy Services PGY1 Pharmacy Residency Program Director Conemaugh Memorial Medical Center Johnstown, Pennsylvania

Larry Goodyer, PhD, MRPharmS, BCPS

Professor, School of Pharmacy
De Montfort University
Leicester, United Kingdom
Medical Director
Nomad Travel Stores and Clinic
Bishop's Stortford, United Kingdom

Mary-Kathleen Grams, PharmD, BCGP

Assistant Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Philip Grgurich, PharmD, BCPS

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

B. Joseph Guglielmo, PharmD

Professor and Dean School of Pharmacy University of California, San Francisco San Francisco, California

Karen M. Gunning, PharmD, BCPS, BCACP, FCCP

Professor (Clinical) and Interim Chair of Pharmacotherapy
Adjunct Professor of Family and Preventive Medicine
PGY2 Ambulatory Care Residency Director
Clinical Pharmacist—University of Utah Family Medicine Residency/Sugarhouse Clinic
University of Utah College of Pharmacy and School of Medicine
Salt Lake City, Utah

Mary A. Gutierrez, PharmD, BCPP

Professor of Pharmacy Practice Chapman University School of Pharmacy Irvine, California

Justinne Guyton, PharmD, BCACP

Associate Professor of Pharmacy Practice Site Coordinator PGY2 Ambulatory Care Residency Program St. Louis College of Pharmacy St. Louis, Missouri

Matthew Hafermann, PharmD, BCPS

Medical ICU/Cardiology Clinical Pharmacist Harborview Medical Center PGY1 Pharmacy Residency Coordinator Medicine Clinical Instructor University of Washington School of Pharmacy Seattle, Washington

Jason S. Haney, PharmD, BCPS, BCCCP

Assistant Professor Department of Clinical Pharmacy and Outcome Sciences South Carolina College of Pharmacy

Christy S. Harris, PharmD, BCPS, BCOP

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Mary F. Hebert, PharmD, FCCP

Professor
Department of Pharmacy
Adjunct Professor of Obstetrics and Gynecology
University of Washington
Seattle, Washington

Emily L. Heil, PharmD, BCPS-AQ ID

Assistant Professor Infectious Diseases University of Maryland School of Pharmacy Baltimore, Maryland

Erika L. Hellenbart, PharmD, BCPS

Clinical Assistant Professor University of Illinois at Chicago College of Pharmacy Chicago, Illinois

David W. Henry, PharmD, MS, BCOP, FASHP

Associate Professor and Chair Pharmacy Practice University of Kansas School of Pharmacy Lawrence, Kansas

Christopher M. Herndon, PharmD, BCPS, CPE

Associate Professor Department of Pharmacy Practice School of Pharmacy Southern University Illinois Edwardsville Edwardsville, Illinois

Richard N. Herrier, PharmD, FAPhA

Clinical Professor Department of Pharmacy Practice and Science College of Pharmacy University of Arizona Tucson, Arizona

Karl M. Hess, PharmD, CTH, FCPhA

Vice Chair of Clinical and Administrative Sciences Associate Professor Certificate Coordinator for Medication Therapy Outcomes Keck Graduate Institute Claremont, California

Curtis D. Holt, PharmD

Clinical Professor Department of Surgery University of California, Los Angeles Los Angeles, California

Evan R. Horton, PharmD

Associate Professor of Pharmacy Practice School of Pharmacy–Worcester/Manchester MCPHS University Worcester, Massachusetts

Priscilla P. How, PharmD, BCPS

Assistant Professor
Director of PharmD Program
Department of Pharmacy
Faculty of Science
National University of Singapore
Principal Clinical Pharmacist
Department of Medicine
Division of Nephrology
National University Hospital
Singapore, Republic of Singapore

Molly E. Howard, PharmD, BCPS

Clinical Pharmacy Specialist Central Alabama Veterans Health Care System Montgomery, Alabama

Timothy R. Hudd, PharmD, AE-C

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Bethany Ibach, PharmD, BCPPS

Assistant Professor of Pharmacy Practice School of Pharmacy, Pediatrics Division Texas Tech University Health Sciences Center Abilene, Texas

Gail S. Itokazu, PharmD

Clinical Associate Professor Department of Pharmacy Practice University of Illinois, Chicago Clinical Pharmacist Division of Infectious Diseases John H. Stroger Jr. Hospital of Cook County Chicago, Illinois

Timothy J. Ives, PharmD, MPH, FCCP, CPP

Professor of Pharmacy Adjunct Professor of Medicine Eshelman School of Pharmacy University of North Carolina at Chapel Hill Chapel Hill, North Carolina

Nicole A. Kaiser, RPh, BCOP

Oncology Clinical Pharmacy Specialist Children's Hospital Colorado Aurora, Colorado

James S. Kalus, PharmD, FASHP

Director of Pharmacy Henry Ford Health System Henry Ford Hospital Detroit, Michigan

Marina D. Kaymakcalan, PharmD

Clinical Pharmacy Specialist Dana Farber Cancer Institute Boston, Massachusetts

Michael B. Kays, PharmD, FCCP

Associate Professor Department of Pharmacy Practice Purdue University College of Pharmacy West Lafayette and Indianapolis, Indiana

Jacob K. Kettle, PharmD, BCOP

Oncology Clinical Pharmacy Specialist University of Missouri Health Care Columbia, Missouri

Rory E. Kim, PharmD

Assistant Professor of Clinical Pharmacy University of Southern California School of Pharmacy Los Angeles, California

Lee A. Kral, PharmD, BCPS, CPE

Clinical Pharmacy Specialist, Pain Management Department of Pharmaceutical Care The University of Iowa Hospitals and Clinics Iowa City, Iowa

Donna M. Kraus, PharmD, FAPhA, FPPAG, FCCP

Pediatric Clinical Pharmacist/Associate Professor of Pharmacy Practice Departments of Pharmacy Practice and Pediatrics Colleges of Pharmacy and Medicine University of Illinois at Chicago Chicago, Illinois

Susan A. Krikorian, MS, PharmD

Professor of Pharmacy Practice School of Pharmacy–Boston

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MCPHS University Boston, Massachusetts

Andy Kurtzweil, PharmD, BCOP

Pharmacy Supervisor—Adult Hematology and Oncology/BMT University of Minnesota Health Minneapolis, Minnesota

Benjamin Laliberte, PharmD, BCPS

Clinical Pharmacy Specialist, Cardiology Massachusetts General Hospital Boston, Massachusetts

Jerika T. Lam, PharmD, AAHIVP

Assistant Professor of Pharmacy Practice School of Pharmacy Chapman University Irvine, California

Trisha LaPointe, PharmD, BCPS

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Alan H. Lau, PharmD

Professor Director, International Clinical Pharmacy Education College of Pharmacy University of Illinois at Chicago Chicago, Illinois

Elaine J. Law, PharmD, BCPS

Assistant Clinical Professor of Pharmacy Practice Thomas J. Long School of Pharmacy and Health Sciences University of the Pacific Stockton, California

Kimberly Lenz, PharmD

Clinical Pharmacy Manager Office of Clinical Affairs University of Massachusetts Medical School Quincy, Massachusetts

Russell E. Lewis, PharmD, FCCP

Associate Professor of Medicine, Infectious Diseases Department of Medical and Surgical Services Infectious Diseases Unit, Policlinico S. Orsola-Malpighi University of Bologna Bologna, Italy

Rachel C. Long, PharmD, BCPS

Clinical Staff Pharmacist Carolinas HealthCare System Charlotte, North Carolina

Ann M. Lynch, BSP, PharmD, AE-C

Professor of Pharmacy Practice School of Pharmacy–Worcester/Manchester MCPHS University Worcester, Massachusetts

Matthew R. Machado, PharmD

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Emily Mackler, PharmD, BCOP

Clinical Pharmacist and Project Manager Michigan Oncology Quality Consortium University of Michigan Ann Arbor, Michigan

Daniel R. Malcolm, PharmD, BCPS, BCCCP

Associate Professor and Vice-Chair Clinical and Administrative Services Sullivan University College of Pharmacy Louisville, Kentucky

Shannon F. Manzi, PharmD, NREMT, FPPAG

Director, Clinical Pharmacogenomics Service Manager, Emergency and ICU Pharmacy Services Boston Children's Hospital Boston, Massachusetts

Joel C. Marrs, PharmD, FCCP, FASHP, FNLA, BCPS-AQ Cardiology, BCACP, CLS, ASH-CHC

Associate Professor
Department of Clinical Pharmacy
University of Colorado Anschutz Medical Campus
Skaggs School of Pharmacy and Pharmaceutical Sciences
Clinical Pharmacy Specialist
Department of Pharmacy
Denver Health and Hospital Authority
Aurora, Colorado

John Marshall, PharmD, BCPS, BCCCP, FCCM

Clinical Pharmacy Coordinator–Critical Care Beth Israel Deaconess Medical Center Boston, Massachusetts

Darius L. Mason, PharmD, BCPS, FACN

Clinical Pharmacist Methodist South Hospital Memphis, Tennessee

Susan L. Mayhew, PharmD, BCNSP, FASHP

Professor and Dean Appalachian College of Pharmacy Oakwood, Virginia

James W. McAuley, RPh, PhD, FAPhA

Associate Dean for Academic Affairs and Professor Departments of Pharmacy Practice and Neurology The Ohio State University College of Pharmacy Columbus, Ohio

Sarah E. McBane, PharmD, CDE, BCPS, FCCP, FCPhA, APh

Professor and Chair Department of Pharmacy Practice West Coast University Los Angeles, California

William W. McCloskey, BA, BS, PharmD

Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Chephra McKee, PharmD

Assistant Professor of Pharmacy Practice School of Pharmacy Pediatrics Division Texas Tech University Health Sciences Center Abilene, Texas

Molly G. Minze, PharmD, BCACP

Associate Professor of Pharmacy Practice Ambulatory Care Division School of Pharmacy Texas Tech University Health Sciences Center Abilene, Texas

Amee D. Mistry, PharmD

Associate Professor Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

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Katherine G. Moore, PharmD, BCPS, BCACP

Executive Director of Experiential Education Associate Professor of Pharmacy Practice Presbyterian College School of Pharmacy Clinton, South Carolina

Jill A. Morgan, PharmD, BCPS, BCPPS

Associate Professor and Chair Department of Pharmacy Practice and Science University of Maryland School of Pharmacy Baltimore, Maryland

Anna K. Morin, PharmD

Professor of Pharmacy Practice and Dean School of Pharmacy–Worcester/Manchester MCPHS University Worcester, Massachusetts

Pamela B. Morris, MD, FACC, FAHA, FASPC, FNLA

Director, Seinsheimer Cardiovascular Health Program Co-Director, Women's Heart Care Medical University of South Carolina Charleston, South Carolina

Oussayma Moukhachen, PharmD, BCPS

Assistant Professor Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts Clinical Care Specialist Mount Auburn Hospital Cambridge, Massachusetts

Kelly A. Mullican, PharmD

Primary Care Clinical Pharmacy Specialist Kaiser Permanente–Mid-Atlantic States Washington, District of Columbia

Myrna Y. Munar, PharmD

Associate Professor of Pharmacy College of Pharmacy Oregon State University Oregon Health and Science University Portland, Oregon

Yulia A. Murray, PharmD, BCPS

Assistant Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Milap C. Nahata, MS, PharmD, FCCP, FAPhA, FASHP

Director, Institute of Therapeutic Innovations and Outcomes Professor Emeritus of Pharmacy, Pediatrics, and Internal Medicine Colleges of Pharmacy and Medicine The Ohio State University Columbus, Ohio

Richard S. Nicholas, PharmD, ND, CDE, BCPS, BCACP

Assistant Professor of Pharmacy Practice Appalachian College of Pharmacy Oakwood, Virginia

Stefanie C. Nigro, PharmD, BCACP, BC-ADM

Assistant Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Cindy L. O'Bryant, PharmD, BCOP, FCCP, FHOPA

Professor
Department of Clinical Pharmacy
Skaggs School of Pharmacy and Pharmaceutical Sciences
Clinical Pharmacy Specialist in Oncology
University of Colorado Cancer Center
Aurora, Colorado

Kirsten H. Ohler, PharmD, BCPS, BCPPS

Clinical Assistant Professor of Pharmacy Practice
College of Pharmacy
University of Illinois at Chicago
Clinical Pharmacy Specialist—Neonatal ICU
University of Illinois at Chicago Hospital and Health Sciences System

Julie L. Olenak, PharmD

Assistant Dean of Student Affairs Associate Professor Department of Pharmacy Practice Nesbitt College of Pharmacy and Nursing Wilkes University Wilkes-Barre, Pennsylvania

Jacqueline L. Olin, MS, PharmD, BCPS, CDE, FASHP, FCCP

Professor of Pharmacy School of Pharmacy Wingate University Wingate, North Carolina

Neeta Bahal O'Mara, PharmD, BCPS

Clinical Pharmacist Dialysis Clinic, Inc. North Brunswick, New Jersey

Robert L. Page, II, PharmD, MSPH, FHFSA, FCCP, FASHP, FASCP, CGP, BCPS (AQ-Cards)

Professor Departments of Clinical Pharmacy and Physical Medicine School of Pharmacy and Pharmaceutical Sciences University of Colorado Aurora, Colorado

Louise Parent-Stevens, PharmD, BCPS

Assistant Director of Introductory Pharmacy Practice Experiences Clinical Assistant Professor Department of Pharmacy Practice University of Illinois at Chicago College of Pharmacy Chicago, Illinois

Dhiren K. Patel, PharmD, CDE, BC-ADM, BCACP

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Katherine Tipton Patel, PharmD, BCOP

Clinical Pharmacy Specialist The University of Texas MD Anderson Cancer Center Houston, Texas

Jennifer T. Pham, PharmD, BCPS, BCPPS

Clinical Assistant Professor, Department of Pharmacy Practice University of Illinois at Chicago College of Pharmacy Clinical Pharmacy Specialist, Neonatal Clinical Pharmacist University of Illinois Hospital and Health Sciences System Chicago, Illinois

Jonathan D. Picker, MBChB, PhD

Assistant Professor Harvard Medical School Clinical Geneticist Boston Children's Hospital Boston, Massachusetts

Brian A. Potoski, PharmD, BCPS

Associate Professor
Departments of Pharmacy and Therapeutics
University of Pittsburgh School of Pharmacy
Associate Director, Antibiotic Management Program
University of Pittsburgh Medical Center
Presbyterian University Hospital
Pittsburgh, Pennsylvania

David J. Quan, PharmD, BCPS

Health Sciences Clinical Professor of Pharmacy
Department of Clinical Pharmacy
School of Pharmacy
University of California, San Francisco
Pharmacist Specialist—Solid Organ Transplant
University of California, San Francisco Medical Center
San Francisco, California

Erin C. Raney, PharmD, BCPS, BC-ADM

Professor of Pharmacy Practice Midwestern University College of Pharmacy–Glendale Glendale, Arizona

Valerie Relias, PharmD, BCOP

Clinical Pharmacy Specialist Division of Hematology/Oncology Tufts Medical Center Boston, Massachusetts

Lee A. Robinson, MD

Instructor Department of Psychiatry Harvard Medical School Boston, Massachusetts Associate Training Director Child and Adolescent Psychiatry Fellowship Primary Care Mental Health Integrated Psychiatrist Cambridge Health Alliance Cambridge, Massachusetts

Charmaine Rochester-Eyeguokan, PharmD, BCPS, BCACP, CDE

Associate Professor of Pharmacy Practice and Science University of Maryland School of Pharmacy Baltimore, Maryland

Carol J. Rollins, PharmD, MS, RD, CNSC, BCNSP

Clinical Associate Professor Department of Pharmacy Practice and Science College of Pharmacy The University of Arizona Tucson, Arizona

Melody Ryan, PharmD, MPH, GCP, BCPS

Professor
Department of Pharmacy Practice and Science
College of Pharmacy
University of Kentucky
Lexington, Kentucky

David Schnee, PharmD, BCACP

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Eric F. Schneider, BS Pharm, PharmD

Assistant Dean for Academics Professor School of Pharmacy Wingate University Wingate, North Carolina

Sheila Seed, PharmD, MPH

Professor of Pharmacy Practice School of Pharmacy–Worcester/Manchester MCPHS University Worcester, Massachusetts

Timothy H. Self, PharmD

Professor of Clinical Pharmacy College of Pharmacy University of Tennessee Health Science Center Memphis, Tennessee

Amy Hatfield Seung, PharmD, BCOP

Senior Director of Clinical Development Physician Resource Management/Caret Cary, North Carolina

Nancy L. Shapiro, PharmD, FCCP, BCPS

Operations Coordinator
University of Illinois Hospital and Health Sciences System
Clinical Associate Professor of Pharmacy Practice
Director, PGY2 Ambulatory Care Residency
College of Pharmacy
University of Illinois at Chicago
Chicago, Illinois

Iris Sheinhait, PharmD, MA, RPh

Certified Poison Information Specialist Adjunct Assistant Professor Regional Center for Poison Control Serving Massachusetts and Rhode Island Boston Children's Hospital and MCPHS University Boston, Massachusetts

Greene Shepherd, PharmD, DABAT

Clinical Professor and Vice-Chair Division of Practice Advancement and Clinical Education Director of Professional Education, Asheville Campus Eshelman School of Pharmacy University of North Carolina at Chapel Hill Asheville, North Carolina

Devon A. Sherwood, PharmD, BCPP

Assistant Professor Psychopharmacology College of Pharmacy University of New England Portland, Maine

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Richard J. Silvia, PharmD, BCCP

Associate Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Carrie A. Sincak, PharmD, BCPS, FASHP

Assistant Dean for Clinical Affairs and Professor Department of Pharmacy Practice Midwestern University Chicago College of Pharmacy Downers Grove, Illinois

Harleen Singh, PharmD, BCPS-AQ Cardiology, BCACP

Clinical Associate Professor of Pharmacy Practice Oregon State University Oregon Health and Science University Portland, Oregon

Jessica C. Song, MA, PharmD

Clinical Pharmacy Supervisor PGY1 Pharmacy Residency Coordinator Department of Pharmacy Services Santa Clara Valley Medical Center San Jose, California

Suellyn J. Sorensen, PharmD, BCPS, FASHP

Director Clinical Pharmacy Services St. Vincent Indianapolis Indianapolis, Indiana

Linda M. Spooner, PharmD, BCPS (AQ-ID), FASHP

Professor of Pharmacy Practice School of Pharmacy–Worcester/Manchester MCPHS University Clinical Pharmacy Specialist in Infectious Diseases Saint Vincent Hospital Worcester, Massachusetts

Karyn M. Sullivan, PharmD, MPH

Professor of Pharmacy Practice School of Pharmacy–Worcester/Manchester MCPHS University Worcester, Massachusetts

David J. Taber, PharmD, MS, BCPS

Associate Professor Division of Transplant Surgery College of Medicine Medical University of South Carolina Charleston, South Carolina

Candace Tan, PharmD, BCACP

Clinical Pharmacist Kaiser Permanente Los Angeles, California

Yasar O. Tasnif, PharmD, BCPS, FAST

Associate Professor Cooperative Pharmacy Program University of Texas at Austin and University of Texas, Rio Grande Valley Clinical Pharmacist Specialist Doctor's Hospital at Renaissance–Renaissance Transplant Institute Edinburg, Texas

Daniel J. G. Thirion, BPharm, MSc, PharmD, FCSHP

Professeur Titulaire de Clinique Faculté de Pharmacie Université de Montréal Pharmacien Centre Universitaire de Santé McGill Montréal, Québec, Canada

Angela M. Thompson, PharmD, BCPS

Assistant Professor Department of Clinical Pharmacy Skaggs School of Pharmacy and Pharmaceutical Sciences University of Colorado Aurora, Colorado

Lisa A. Thompson, PharmD, BCOP

Clinical Pharmacy Specialist in Oncology Kaiser Permanente Colorado Lafayette, Colorado

Toyin Tofade, MS, PharmD, BCPS, CPCC

Dean and Professor Howard University College of Pharmacy Washington, District of Columbia

Tran H. Tran, PharmD, BCPS

Associate Professor Midwestern University, Chicago College of Pharmacy Downers Grove, Illinois

Dominick P. Trombetta, PharmD, BCPS, CGP, FASCP

Associate Professor Department of Pharmacy Practice Nesbitt School of Pharmacy Wilkes University Wilkes-Barre, Pennsylvania

Toby C. Trujillo, PharmD, FCCP, FAHAH, BCPS-AQ Cardiology

Associate Professor Department of Clinical Pharmacy Skaggs School of Pharmacy and Pharmaceutical Sciences University of Colorado Aurora, Colorado

Sheila K. Wang, PharmD, BCPS (AQ-ID)

Associate Professor of Pharmacy Practice Chicago College of Pharmacy Midwestern University Downers Grove, Illinois Clinical Pharmacist, Infectious Disease Program Director, Rush University Medical Center Chicago, Illinois

Brian Watson, PharmD, BCPS

Pharmacist University of Maryland Medical System St. Joseph's Medical Center Baltimore, Maryland

Kristin Watson, PharmD, BCPS-AQ Cardiology

Associate Professor, Vice-Chair of Clinical Services University of Maryland School of Pharmacy Baltimore, Maryland

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Lynn Weber, PharmD, BCOP

Clinical Pharmacy Specialist, Oncology/Hematology Pharmacy Residency Coordinator and PGY-1 Residency Director Hennepin County Medical Center Minneapolis, Minnesota

Kellie Jones Weddle, PharmD, BCOP, FCCP, FHOPA

Clinical Professor of Pharmacy Practice College of Pharmacy Purdue University Indianapolis, Indiana

C. Michael White, PharmD, FCP, FCCP

Professor and Head Department of Pharmacy Practice School of Pharmacy University of Connecticut Storrs, Connecticut

Natalie Whitmire, PharmD, BCPS, BCGP

Pharmacist Specialist University of California, San Diego Health

Barbara S. Wiggins, PharmD, BCPS, CLS, AACC, FAHA, FCCP, FNLA

Clinical Pharmacy Specialist–Cardiology Medical University of South Carolina Charleston, South Carolina

Kristine C. Willett, PharmD, FASHP

Associate Professor of Pharmacy Practice School of Pharmacy–Worcester/Manchester MCPHS University Manchester, New Hampshire

Bradley R. Williams, PharmD, CGP

Professor of Clinical Pharmacy and Clinical Gerontology School of Pharmacy University of Southern California Los Angeles, California

Casey B. Williams, PharmD, BCOP, FHOPA

Director, Center for Precision Oncology Director, Department of Molecular and Experimental Medicine Avera Cancer Institute Sioux Falls, South Dakota

Dennis M. Williams, PharmD, BCPS, AE-C

Associate Professor and Vice-Chair for Professional Education and Practice Division of Pharmacotherapy and Experimental Therapeutics Eshelman School of Pharmacy University of North Carolina at Chapel Hill Chapel Hill, North Carolina

Katie A. Won, PharmD, BCOP

Clinical Pharmacist Hennepin County Medical Center Minneapolis, Minnesota

Annie Wong-Beringer, PharmD, FIDSA

Professor of Pharmacy School of Pharmacy University of Southern California Los Angeles, California

Dinesh Yogaratnam, PharmD, BCPS, BCCCP

Assistant Professor of Pharmacy Practice School of Pharmacy–Worcester/Manchester MCPHS University Worcester, Massachusetts

Kathy Zaiken, PharmD

Professor of Pharmacy Practice School of Pharmacy–Boston MCPHS University Boston, Massachusetts

Caroline S. Zeind, PharmD

Associate Provost for Academic and International Affairs Chief Academic Officer Worcester, Massachusetts and Manchester, New Hampshire, Campuses Professor of Pharmacy Practice MCPHS University Boston, Massachusetts

Sara Zhou, PharmD

Certified Poison Information Specialist Adjunct Assistant Professor Regional Center for Poison Control Serving Massachusetts and Rhode Island Boston Children's Hospital and MCPHS University Boston, Massachusetts

Kristin M. Zimmerman, PharmD, CGP, BCACP

Associate Professor Department of Pharmacotherapy & Outcomes Science Virginia Commonwealth University Richmond, Virginia

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SECTION 1 | General Principles

Section Editors: William W. McCloskey and Maria D. Kostka-Rokosz



Medication Therapy Management and Assessment of Therapy

Matthew R. Machado, Amee D. Mistry, and Joseph W. Ferullo

CORE PRINCIPLES	CHAPTER CASES
Medication Therapy Management Services (MTMS) are a service or group of services that optimize therapeutic outcomes for individual patients.	Case 1-5 (Questions 1-4)
A successful MTMS encounter includes medication reconciliation and a comprehensive medication history.	Case 1-1 (Questions 1-3)
Sources of patient information for MTMS include the patient, the electronic health record, the paper chart, and the pharmacy information system.	Case 1-5 (Questions 1, 5), Table 1-1
A careful and complete patient interview should include medical, medication, and social histories, and it must be provided in a culturally sensitive manner.	Case 1-1 (Questions 1-3), Table 1-2, Online Content
A successful MTMS encounter must be well documented following the Problem Oriented Medical Record (POMR). Documentation involves collecting subjective and objective data to identify the primary problem.	Case 1-5 (Question 1), Table 1-4 Case 1-2 (Question 1), Case 1-3 (Question 1), Case 1-4 (Question 1)
The clinician must assess the drug therapy or disease-specific problem and create a treatment plan.	Case 1-5 (Questions 1, 2)
The final step in documenting the MTMS encounter is developing the medication action plan and processing any billing requirements.	Case 1-5 (Questions 1, 2, 4)
Accurate and complete communication of the MTMS encounter to the patient's health care team is vital.	Case 1-5 (Question 3)

This chapter discusses medication therapy management services (MTMS) with a focus on the assessment of drug therapy. The illustrations in this chapter primarily focus on the pharmacist; however, the principles used to assess patient response to drug therapy are of value to all health care providers.

As defined by the American Pharmacists Association, medication therapy management (MTM) is a term used to describe a broad range of health care services provided by pharmacists, the medication experts on the health care team. A consensus definition created by 11 pharmacy associations, adopted by the pharmacy profession in 2004, defines MTM as a service or group of services that optimize therapeutic outcomes for individual patients. Pharmacists provide MTM to help patients get the best benefits from their medications by actively managing drug therapy and by identifying, preventing, and resolving medication-related problems.

MTM has a direct relationship to pharmaceutical care. Pharmaceutical care has been described as the responsible provision of drug therapy to achieve definite outcomes that are intended to improve a patient's quality of life.^{2,3} In fact, MTM has been described as a service provided in the practice of pharmaceutical care.⁴ However, unlike pharmaceutical care, MTM is recognized by payers, has current procedural terminology (CPT) codes specifically for pharmacists, and has several clearly defined interventions. Therefore, MTMS will be the term used to describe the activity of MTM in various patient populations.

With the passage of the Patient Protection and Affordable Care Act and the Health Care and Education Reconciliation Act of 2010, pharmacists have tremendous opportunities in the implementation of health care reform.^{5,6} One of the hallmarks of this law is delivery system reform. As health care delivery systems change, pharmacists have an opportunity to improve overall quality of

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care, to become involved in coordinated health care approaches such as medical home teams and accountable care organizations, and to collaborate to improve care for high-risk patients and those with chronic conditions in primary-care settings. In the United States, there are an estimated 133 million people that have at least one chronic medical condition.⁷ In 2010, 86% of all health care spending was for people with one or more chronic health condition(s).⁸ Pharmacists will have additional opportunities as hospitals will have financial incentives to improve quality, reduce costs, and decrease hospital-acquired conditions.^{5,6}

Both patient self-care and medication reconciliation are critical aspects of any MTMS encounter regardless of the setting (i.e., inpatient, community, ambulatory, or institutional). Patient self-care requires the patient to take responsibility for the illness; however, the help of a professional to structure healthy self-care is important. For example, patients with diabetes who monitor their blood glucose levels regularly and adjust their diet according to the guidelines published from the American Diabetes Association (ADA) would be practicing self-care. Self-care is often the work that the patient performs between visits with the provider. The patient should be involved in his or her own care to ensure the best outcomes.