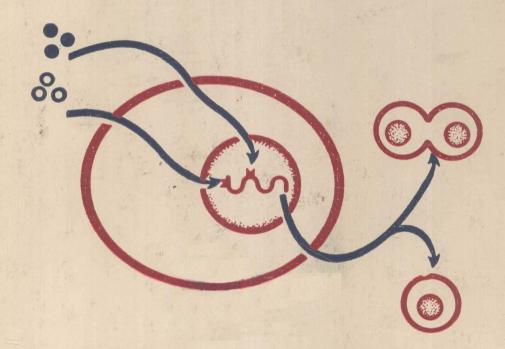
PATHOPHYSIOLOGY

The Biologic Basis for Disease in Adults and Children



Kathryn L. McCance Sue E. Huether

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Developmental Editor: Penny Rudolph Project Manager: Karen Edwards Production Editor: Gail Brower Designer: Elizabeth Fett

Manufacturing Supervisor: Karen Lewis

SECOND EDITION

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Printed in the United States of America Composition by The Clarinda Company Printing/binding by Von Hoffmann Press, Inc.

Mosby – Year Book, Inc. 11830 Westline Industrial Drive St. Louis, Missouri 63146

Library of Congress Cataloging in Publication Data

McCance, Kathryn L.

Pathophysiology: the biologic basis for diseases in adults and children / Kathryn L. McCance, Sue E. Huether. —2nd ed.

p. cm.

Includes bibliographical references and index.

ISBN 0-8016-6902-2

1. Physiology, Pathological. 2. Nursing. I. Huether, Sue E. II. Title.

[DNLM: 1. Disease—nurses' instruction. 2. Pathology—nurses' instruction. 3. Physiology—nurses' instruction. QZ 4 M478p 1993] RB113.M35 1993

616.07—dc20 DNLM/DLC

for Library of Congress

93-31475

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To Kallie, Margot, and their parents,
Beth and Greg,
their aunt and uncles,
Anne, Eric, and Mark,
and their grandfather,
Papa John
. . . all Wolfers
"Here's lookin at all the kids!"

To the many wonderful teachers and students who have taught me through the years.

SEH

Preface

Spectacular advances in modern biology are revolutionizing our understanding of pathophysiology. Although the knowledge explosion—so overwhelming—is comprehensive and progressing rapidly, we have made every effort to stay current. Each chapter has been updated and in several instances revised extensively. Yet, we have retained our initial three broad objectives in offering this second edition:

- We have continued to provide comprehensive presentations of underlying principles common to all disease processes.
- We have retained, updated, and consistently revised coverage of disease processes in all age groups.
- We have paid scrupulous attention to making this edition an even more effective learning tool.

Specifically, we have added new two-color illustrations and updated and revised first edition illustrations appropriately. Algorithms have been added, and many have been revised to facilitate ease of learning.

ORGANIZATION OF THE BOOK

The basic organization has not changed from the first edition. The book is divided into two parts: *Part One*, *Central Concepts of Pathophysiology: Cells and Tis-*

sues, and Part Two, Pathophysiologic Alterations: Organs and Systems.

Part One: Central Concepts of Pathophysiology: Cells and Tissues

PART ONE: CENTRAL CONCEPTS OF PATHOPHYSIOLOGY: CELLS AND TISSUES

The Cell
Genes and Gene-Environment Interaction
Mechanisms of Self-Defense
Cellular Proliferation—Cancer

As before, Part One begins with an indepth study of the cell, its workings, and other processes related to cells, but now has the addition of new areas of study:

- · Completely revised chapters on cancer
- Cell signaling and cell communication processes
- Cytokines and their biologic activities
- Updated information on the identification of the genome and gene therapy
- · Gender and life expectancy and cellular aging
- · Mechanisms of immune defense

- Significantly revised content on AIDS and psychoneuroimmunologic regulation
- · Concepts of stress, coping, and illness

Part One lays the foundation for organ system diseases and disorders that are presented in Part Two. Physical laws and principles, for example, fluid movement, energy expenditures, and electrical conductances, are emphasized to increase student awareness of how these laws and principles can be applied to understanding mechanisms and complications of disease. Cells of the immune system and inflammatory responses are activated with every type of trauma or disease. Understanding these mechanisms of protection and healing provides a powerful source of knowledge that can be generalized to disorders of many organ systems. Once the normal responses are understood, students can more easily comprehend the consequences of both immune deficiencies or hyperactive responses.

Knowledge of the rapidly expanding information on the genetic basis of disease and gene-environment interaction is critical to understanding developmental alterations, cancer, cardiovascular disease, diabetes, cystic fibrosis, degenerative neural diseases, and other common disorders. Therefore both cellular and population genetics have been updated to provide the student the necessary knowledge base for appreciating the importance of risk factors, diagnostic procedures, and treatments that are influenced by genetics.

Part Two: Pathophysiologic Alterations: Organs and Systems

PART TWO: PATHOPHYSIOLOGIC ALTERATIONS: ORGANS AND SYSTEMS

The Neurologic System

The Endocrine System

The Reproductive System

The Hematologic System

The Cardiovascular and Lymphatic Systems

The Pulmonary System

The Renal and Urologic Systems

The Digestive System

The Musculoskeletal System

The Integumentary System

Part Two is still a survey of specific diseases from the traditional systems organization but adds new information and some strategic changes in pedagogy. Because student and faculty response to the format of this text has been overwhelmingly enthusiastic, each system unit still begins with a chapter on the normal anatomy and physiology of the system. We have paid scrupulous at-

tention to updating these chapters, knowing that normal function *directly* informs the discussion of the specific diseases that follow. These chapters have been extensively edited to include new diagnostic tests and updated information on the normal aging process unique to each system.

Throughout Part Two updated or new content has been added, including:

- Reorganization of the alterations of neurologic function chapter to include new information on head trauma and diffuse axonal injury
- Neurologic complications of AIDS and a new section on HIV infection in children
- Expansion of content on sexual dysfunction and cancer of the reproductive organs
- New information on tuberculosis and lung cancer
- New algorithms on conditions such as multiple organ system dysfunction, septic shock, disseminated intravascular coagulation, asthma, adult respiratory distress syndrome, and glomerulonephritis
- More tables to increase clarity of content on topics such as croup syndromes, glomerulonephritis, pneumonia in children, cancer of the GI tract, and hepatitis
- Application of cytokine function from Part One to pathophysiologic processes in Part Two, including the role of colony-stimulating factor (CSF) in hematopoiesis, leukemia, and leukopenia
- Pathologic effects of interleukins and tumor necrosis factor
- Cellular mechanisms of atherosclerosis leading to myocardial infarction and Kawasaki disease in children

Many features of the first edition have been retained for most of the disorders, including:

- Epidemiology and risk factors
- Pathogenesis and clinical manifestations
- · Evaluation and treatment
- Clinical commentaries (listed on p. xvii)
- Use of physical laws to aid students in understanding pathophysiologic alterations
- Extensive integration of cancer research data

New features that enhance the second edition include revised historical openers with new art, new two-color illustrations, figure titles in boldface type, updated and easily retrieved content on aging, detailed chapter outlines with page numbers, updated and revised key terms, and complete updated summary reviews. The reference lists have been divided into cited references and references for additional reading to aid further study.

Although major emphasis is devoted to pathophysiology and clinical manifestations, significant attention is also paid to epidemiology and risk factors. Throughout the book, we have continued to extensively cross-reference to fully integrate the two parts of the book. Rather than delete controversial material, for example

the role of estrogens in breast cancer, we have summarized theoretical positions and identified gaps in current understanding. Sensitivity to gender differences in the development and manifestations of various diseases and disorders has been included as appropriate.

LIFESPAN CONTENT

Aging Content

Beginning in Chapter 2, theories of aging are presented. Reference is made to this content throughout Part Two as the normal changes associated with aging are presented in the normal anatomy and physiology chapter of each system unit. The pathophysiology of diseases associated with advanced age is discussed in the alterations chapters. In this way, we attempt to make it clear to students that there are physiologic changes related to aging that are normal and healthy, and that old age should not be considered synonymous with sickness and death.

Pediatric Content

In the same fashion, content related to alterations in children is firmly rooted in Part One of the book, and frequent reference is made to cellular content during the discussion of specific diseases of childhood presented in the systems units in Part Two. Material in Part One includes considerations related to childhood, such as congenital immune deficiencies, genetic disorders, familial diseases, and cancer in children. We take the opportunity in the systems chapters on children in Part Two to summarize, when appropriate, the differences between diseases in adults and children, such as differences in tumors.

ADDITIONAL LEARNING AIDS

Study Guide and Workbook

A most exciting addition to the second edition is a new Study Guide and Workbook for students written by Dr. Clayton Parkinson. Dr. Parkinson has designed the study guide to help students learn pathophysiology by reinforcing key concepts and integrating workbook-like exercises. To this end, he has designed the anatomy and physiology chapters to have review objectives keyed to the text and keyed to practice examinations with answers. The pathophysiology chapters have prerequisite objectives with review assignments referenced to the text. The pathophysiology objectives are in boldface type and are keyed to study assignments and have an abbreviated outline narrative with summary charts. A practice examination with answers and a case study follow the narrative for application of concepts.

Instructor's Resource Manual and Test Bank

The instructor's manual has been significantly revised and updated. The revisions include updated chapter objectives; new sections on difficult concepts, with suggested teaching and learning approaches; and integrative content questions (i.e., critical thinking exercises). The integrative questions are designed to reinforce the application of physical laws and principles for problem solving. For example, students are asked to apply the concepts of osmotic gradient to the clinical problem of dehydration and to Starling's forces for understanding edema formation in an individual with hypertension.

A 1200-item test bank is available in the instructor's manual. Questions include multiple choice, true/false, matching, and short answer. Answers and page references to the text are provided.

Transparency Acetates

A new package of transparency acetates is available. This includes 150 transparencies, primarily of disorders. A legend booklet is included, and the acetates have been 3-hole punched so they can be stored in a binder.

With the revisions, updated material, and new student learning aids, we believe this new edition offers a *state* of the art and science approach to learning pathophys-

iology.

Kathryn L. McCance Sue E. Huether

Acknowledgments

Many people worked on this book! Most significant was Sue Meeks, who generated the manuscript. Sue's astute and scrupulous approach kept us consistently on target. Thanks again, Sue, for your energy and remarkable attention to detail.

Again, we thank our colleagues and friends at the University of Utah College of Nursing. Special thanks to Debbie Bachan and Molly Berigan-Spira for their critical assistance with many details related to the revision and transmittal of the manuscript. Thanks to Mary Youngkin and Jeanne La Ber for their generous and skilled attention as reference librarians in Eccles Health Sciences Library. In addition, retrieving and securing hundreds of references was facilitated by Cheryl Mansen. Thank you, Cheryl. We acknowledge the librarians at the National Library of Medicine for their very helpful assistance in retrieving illustrations for the historical unit openers.

To our outstanding editors—thank you. Sally Schrefer, our editor, inherited us part way through revision. Sally got us focused and kept us on track and managed

the entire project with "grit and grace." To Sally, thank you, and we'll have you in cowboy boots yet!

Penny Rudolph was our developmental editor. Penny is quick, thorough, and gently persistent. Thank you, Penny, for managing such detail completely and confidently. To Brian Dennison, thanks for all the administrative help and kindness, especially when we lost or forgot things.

Gail Brower was our chief production editor. Gail's ability to orchestrate thousands of details easily and accurately was astonishing. Thank you, Gail.

Again, we had the great fortune of having two extremely talented artists—Donald O'Connor and George Wassilchenko. Thanks, not only from us, but from many faculty and students who have written to tell us how the art program made the book!

Reviewers for this edition included faculty, students, clinicians, and investigators. Many comments and suggestions were essential and notably consistent—saving us invaluable time and energy. To all of the reviewers, thank you.

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