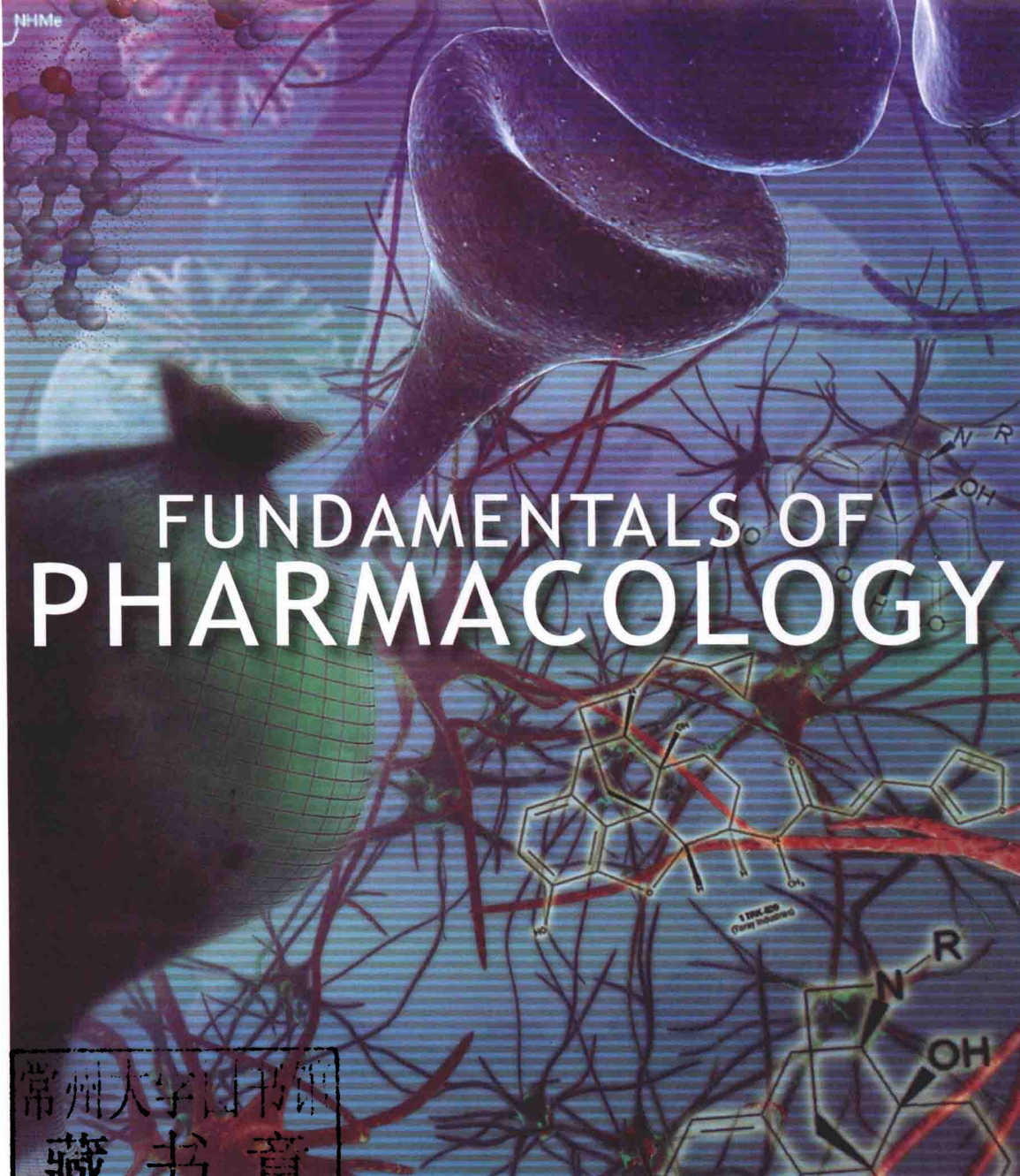




FUNDAMENTALS OF PHARMACOLOGY

SHANE BULLOCK AND ELIZABETH MANIAS 7th EDITION



SHANE BULLOCK AND ELIZABETH MANIAS 7th EDITION

Copyright © Pearson Australia (a division of Pearson Australia Group Pty Ltd) 2014
Pearson Australia
Unit 4, Level 3
14 Aquatic Drive
Frenchs Forest NSW 2086

www.pearson.com.au

The *Copyright Act 1968* of Australia allows a maximum of one chapter or 10% of this book, whichever is the greater, to be copied by any educational institution for its educational purposes provided that that educational institution (or the body that administers it) has given a remuneration notice to Copyright Agency Limited (CAL) under the Act. For details of the CAL licence for educational institutions contact:

Copyright Agency Limited, telephone: (02) 9394 7600, email: info@copyright.com.au

All rights reserved. Except under the conditions described in the *Copyright Act 1968* of Australia and subsequent amendments, no part of this publication may be reproduced, stored in a retrieval system or transmitted in any form or by any means, electronic, mechanical, photocopying, recording or otherwise, without the prior permission of the copyright owner.

Senior Acquisitions Editor: Mandy Shepherd
Senior Project Editor: Rebecca Pomponio
Development Editor: Katie Pittard
Editorial Coordinator: Sophie Attwood
Production Controller: Julie McArthur
Copy Editor: Biotext Pty Ltd
Proofreader: Jane Tyrrell
Senior Copyright and Pictures Editor: Emma Gaulton
Indexer: Garry Cousins
Cover and internal design by Natalie Bowra
Cover illustration © MEDI-MATION/Science Photo Library
Typeset by Midland Typesetters, Australia

Printed in China (GCC/01)

1 2 3 4 5 18 17 16 15 14

National Library of Australia
Cataloguing-in-Publication Data

Author:	Bullock, Shane, author.
Title:	Fundamentals of pharmacology / Shane Bullock, Elizabeth Manias.
Edition:	7th edition.
ISBN:	9781442563100 (paperback).
Notes:	Includes index.
Subjects:	Pharmacology—Study and teaching (Higher). Drugs—Study and teaching (Higher)
Other Authors/Contributors:	Manias, Elizabeth, author.
Dewey Number:	615.1900711

Every effort has been made to trace and acknowledge copyright. However, should any infringement have occurred, the publishers tender their apologies and invite copyright owners to contact them.

FUNDAMENTALS OF PHARMACOLOGY

PREFACE

Fundamentals of Pharmacology is primarily a text for undergraduate and postgraduate students in the health science disciplines, particularly those in nursing. Students of other health disciplines whose roles involve pharmacological therapy (such as pharmacy, podiatry, optometry, paramedic and physiotherapy), as well as those studying basic science, should find much of the material relevant to their studies. Qualified health professionals and pharmaceutical company sales representatives will also find the information useful in their daily roles. Unashamedly, we have written a pharmacology textbook for students of the health professions that does not compromise the scientific basis of the discipline. Many pharmacology texts previously published have been strong on clinical considerations, yet relatively weak in the science of pharmacology.

Our approach

Philosophically, our goal is to empower health professionals through an understanding of the fundamental scientific principles of pharmacology. We believe that, to promote understanding, the effects of drugs on physiological and pathophysiological processes have to be clearly explained. We have included a small amount of chemistry and biochemistry where appropriate in order to facilitate this understanding. With a greater appreciation of the action of drugs and their target tissues, the reader should be able to deduce what adverse effects to expect, as well as the precautions and contraindications to consider.

Furthermore, where possible we have tended to describe the important characteristics of medicine groupings rather than focusing on individual agents, and have used prototypes and common generics as examples. The rationale for this approach is that new medicines are regularly entering the market while older agents are removed. The average practitioner cannot possibly keep up with all these changes. However, if a student knows which grouping a new agent belongs to, the principal characteristics of the medicine can be easily deduced.

This book is primarily designed to establish the foundations in pharmacology. We encourage students to refer to the electronic and hard copy references commonly found in the clinical setting and in hospital wards, such as the *Australian Medicines Handbook*, *MIMS* or *Therapeutic Guidelines*, for more detailed information regarding individual therapeutic agents (e.g. dosage, special precautions and toxicological information).

We hope that you will find this textbook a valuable companion in your pursuit of a fundamental understanding in a most fascinating area of clinical knowledge—pharmacology.

Changes in the seventh edition

This edition reflects the availability of medicines in Australia and New Zealand at the time of publication. Consistent with information currently available to us, we have updated new medicines that have entered the marketplace, as well as those that have been removed since the last edition.

We use the word “medicine” rather than “drug” or “medication” where appropriate. This change was implemented in recognition of the increasing use of the word “medicine” as evidenced by a number of industry websites such as:

- the National Prescribing Service (www.nps.org.au);
- Australian Prescriber (www.australianprescriber.com); and
- the Therapeutic Goods Administration (www.tga.gov.au/industry/pm.htm).

Where appropriate, the therapeutic approaches associated with the management of important clinical conditions, such as cardiovascular disease, diabetes mellitus and psychiatric illness have been brought up to date with current clinical guidelines.

FULL COLOUR FIGURES AND TABLES

This edition is printed in full colour for the first time. Chapter figures are more dynamic, providing the representations of structures and processes with greater depth and vibrancy. Receptors are rendered more often in figures as G-protein-coupled or ion channels rather than basic geometric shapes.

A number of new figures and tables have been included to assist students in visualising difficult pharmacological concepts, the sites of actions of drugs and the range of drug effects expected in a person when particular drug groups are administered.

END-OF-CHAPTER AND END-OF-SECTION FEATURES

The book contains over 800 end-of-chapter questions to assist in the consolidation of learning—all of these have been reviewed.

New and revised integrated case studies appear at the end of sections to assist with making links between theory and practice.

ACKNOWLEDGMENTS

We would like to thank a number of people who have contributed to the development of this textbook, and this edition in particular. Elizabeth wishes to thank her family for their patience and support, and for giving her an appreciation of things beyond the world of medicines. She would also like to thank her colleagues and students, who have provided her with helpful comments about the textbook and made suggestions for improvement.

For Shane the writing of this edition was fuelled by the primary producers situated around his homebase in the Gippsland region of Victoria—yummy cheese, chutney, jam and wine. With respect to the latter indulgence, students are advised to do as I say (see Chapter 24) rather than as I do. He is grateful to the backyard chooks who proved to be a more receptive and attentive audience than other family members when workshopping new ideas for the book.

We would like to thank the team at Pearson Australia for the preparation of this edition. Our thanks to Mandy Sheppard for her support, encouragement and good humour. We are also grateful for access to the expertise of Katie Pittard, Emma Gaulton and Rebecca Pomponio. It is always a pleasure working with you. We thank our copy editor, Anneliese Gillard, and proofreader, Jane Tyrrell, for their valuable advice on contemporary word usage and for picking up on our writing idiosyncrasies.

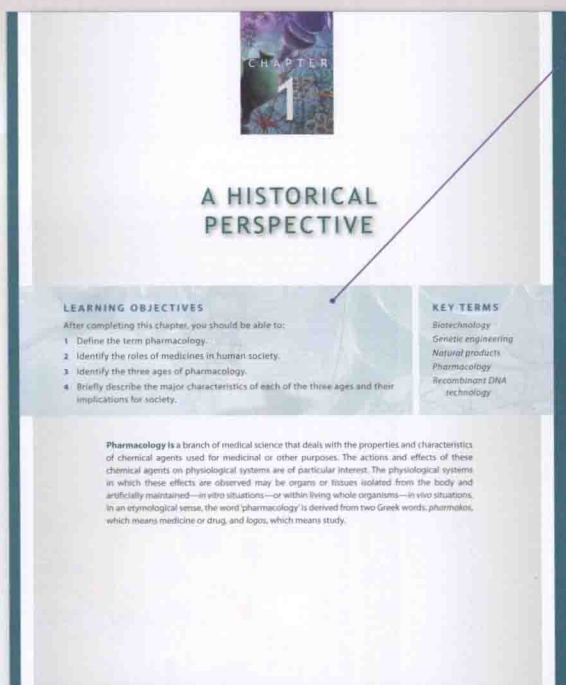
Thanks also to the proposal reviewers:

- Peter Athanasos, Flinders University
- Dr Hemant Mehta, Australian Catholic University
- Rebekkah Middleton, University of Wollongong
- Dr Srinivas Nammi, University of Western Sydney
- Dr Nicole Reinke, James Cook University
- Dr Ross Richards, Charles Sturt University
- Dr Scott Smid, The University of Adelaide
- Dr Jenny Wilkinson, Charles Sturt University

Shane Bullock and Elizabeth Manias

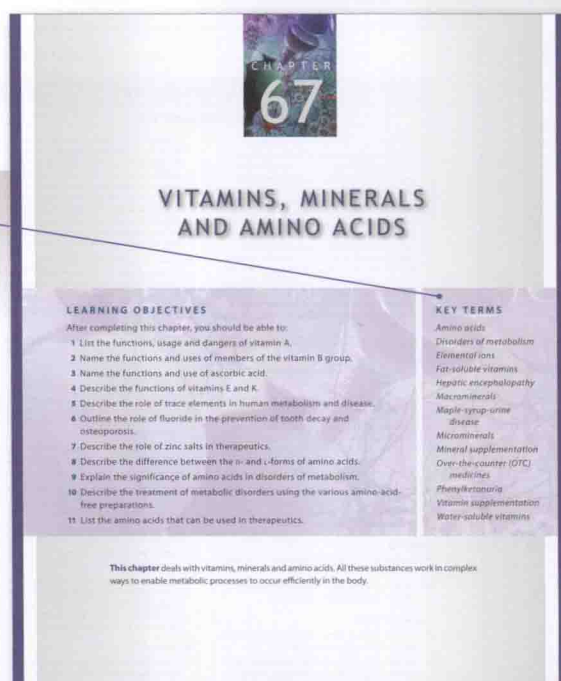
July 2013

FEATURES



Learning Objectives make clear what students will learn in each chapter.

Key Terms introduce students to new terminology and are helpful when revising for exams.



2 What is a chelating agent?

3 Name the agent(s) used in the treatment of poisoning by each of the following substances:

- a cyanide
- b lead
- c mercury
- d pesticides

4 Define the term envenomation.

5 State the three aims of emergency care when someone is bitten or stung by a venomous animal.

6 Your neighbour visits you in an extremely distressed state. Joey, her three-year-old son, has just swallowed an unknown quantity of paracetamol tablets. What would you advise her to do? Why?

7 Mario Malodoro, a 60-year-old farmer, is brought into the emergency department with organophosphate poisoning. How would this form of poisoning be treated?

8 While clearing rubbish in his backyard, 28-year-old Jeffrey Abbott is bitten on the hand by a redback spider. His partner bandages his hand and arm firmly. She then drives him to your clinic, which is only five minutes down the road. Comment on the suitability of this treatment. Describe the management of this type of envenomation.

22 MEDICINE SUMMARY TABLE

FAMILY NAME	GENERIC NAME	TRADE NAME(S)
Emetic	iprécacuan	
Adsorbent	activated charcoal + sorbitol	CarboSorb X CarboSorb XS
Iso-osmotic laxatives	electrolytes + polyethylene glycol + ascorbic acid	CololyTELY- C ClycoPrep OlycoPrep-C Klean Prep Monicol Movicol
Methanol intoxication	ethanol + glucose	
Cyanide antidote	amyl nitrite dicoumat edetate sodium nitrite sodium thiosulfate	
Organophosphate antidotes	atropine sulfate pralidoxime iodide	PAM injection

Australia only
 New Zealand only

Medicine Summary Tables provide a handy list of family names, generic names and trade names for specific medicines. Icons indicate medicines that are only available in Australia or New Zealand. Special considerations are listed where necessary.

Figures illustrate and clarify complex processes, aiding student comprehension.

CHAPTER 27 ADRENERGIC PHARMACOLOGY 279

CLINICAL MANAGEMENT

SYMPATHOMIMETICS

Assessment

- Obtain baseline vital signs for the person. Report any abnormal findings. These include blood pressure and rate, and rhythm of pulse. Assess colour and temperature of the person's extremities (for drug with α_1 effects). Conscious state is assessed to determine cerebral perfusion (this is an important consideration if the medicine is administered intravenously for the purpose of maintaining blood pressure). Determine rate, rhythm and depth of respiration. Assess for wheezing if the medicine is used for asthma. Listen to the heart with a stethoscope for dysrhythmias and palpitations (for drugs with α_1 or β_1 effects). Compare the person's apical beat with the radial rate. A difference indicates irregularity in rhythm. Determine urinary output and assess for bladder distension (for drugs with α_1 effects).
- Assess whether the person has a history of the following:
 - glaucoma or prostatic hypertrophy (for drugs with α_1 effects);
 - cardiovascular, cerebrovascular or circulatory disease, hyperthyroidism (for drugs with α_1 or β_1 effects);
 - diabetes mellitus (for drugs with α_1 or β_1 effects). The sympathomimetic agent may intensify the condition, therefore, leading to elevated blood glucose levels from increased glycogen breakdown. The situation would require further clarification with the prescriber.
- Determine whether the person is taking monoamine oxidase inhibitors, β -blockers or digoxin, as their effects can be either nullified or intensified by the administration of sympathomimetics.

Planning

- The person's vital signs will remain within an acceptable range for the person.
- The person will experience minimal or no adverse effects from the sympathomimetic.

Implementation

- Carefully and regularly monitor the person's vital signs, conscious state and urinary output.
- Sympathomimetic administered intravenously can produce profound effects on vital organs at small

dosages. Their haemodynamic effects should, therefore, be carefully monitored and recorded. Dosages are then titrated according to the person's response. A large central vein should be used for the administration of intravenous sympathomimetics to prevent peripheral necrosis. The use of intravenous sympathomimetics is restricted generally to clinical settings in which close monitoring of venous and arterial pressures, electrocardiogram and urinary output can be performed, such as intensive care or coronary care units.

- Report and record adverse effects of the sympathomimetic, including palpitations, tachycardia (pulse greater than 100 beats/min), tremors or increased glucose levels.
- Regularly monitor the person's urinary output (for drugs with α_1 effects).
- Prolonged use of a sympathomimetic may lead to a diminished clinical effect, which is caused by a regulatory decrease in receptor numbers.

Medicine education

- Drugs with β_1 effects are usually given by inhalation or sublingual. Check the methods for inhalation and nebulisation (refer to Chapter 7, Tables 7.17 and 7.18, for a description of methods).
- Instruct the person on the method of administering solid or flu preparations by nasal spray and drops (refer to Chapter 7, Tables 7.7 and 7.8, for description of methods).
- Instruct the person that nasal sprays used in excess could lead to a rebound nasal congestion. Directions for dosage should be carefully followed.
- Excessive use of bronchodilator inhalers could lead to adverse effects, such as tachycardia and skeletal muscle tremor. If asthma symptoms appear to be getting worse, the doctor should be consulted.
- Instruct the person to read all labels of over-the-counter preparations. Many of these preparations contain sympathomimetics and should not be taken if the person has a history of cardiac disease, diabetes, hypertension or cardiac dysrhythmias.

Evaluation

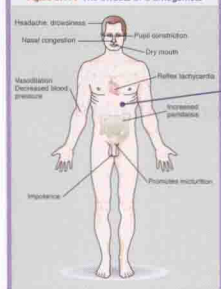
- Examine the person's response to the sympathomimetic for expected and adverse effects. Continue to monitor

260 SECTION VI AUTONOMIC PHARMACOLOGY

Figure 27.1 Adrenergic nerve action

A The branching nerve terminal of a sympathetic postganglionic fibre shows a series of small swellings known as varicosities along the length of each branch. **B** A summary of events involved in adrenergic nerve stimulation. The action potential travels along the axon until it reaches the varicosities of the axon terminals (1). Depolarisation of the membrane of the varicosity causes the release of chemical transmitters, noradrenaline (NA), into the synaptic gap (2). NA diffuses across the gap and mimics with the adrenergic postganglionic receptors, triggering an effector response via a G-protein-coupled second messenger system (3). The transmitter is removed from the synaptic gap by the uptake 1 transporter (4) and is returned to the synaptic vesicles. Any excess transmitter within the terminal not returned to the vesicles is degraded by the mitochondrial enzyme, monoamine oxidase (MAO) (5). Any excess transmitter remaining within the synaptic gap is subject to extraneuronal reuptake (uptake 2). The release of transmitter from the varicosity is also subject to modulation by presynaptic adrenoceptors (enhancement of release by α_1 type, inhibition by α_2). Such control of transmitter release is known as autoregulation.

Clinical Management Tables highlight clinical applications of theory and utilise the clinical decision-making framework in a step-by-step process for care of the person.

Figure 27.14 The effects of α_1 antagonists

erectile dysfunction, but it must be used with **papaverine** or **alfentanil** to be effective. Selective α_1 antagonists are used for control of hypertension. All α_1 antagonists may cause a rapid fall in blood pressure after the first dose. The patient should be advised to take the first dose at bedtime to reduce the consequences of this effect. The dose is then titrated slowly at two-weekly intervals. This hypotensive effect is likely to be more severe in the older person and in the individual who takes diuretics. It is recommended, therefore, that diuretics be withheld for a few days before commencing an α_1 antagonist. Postural hypotension and dizziness may occur and the person is advised to get up gradually from a lying or sitting position. Advise individuals to sit down if they become dizzy.

B ANTAGONIST ACTION

Mechanism of action

Acetazolamide, carvedilol, nadolol, oxprenolol, pindolol, propranolol, sotalol and timolol are non-selective β antagonists or blockers. Atenolol, betaxolol, bisoprolol, carvedilol, nebivolol and metoprolol are relatively β_1 -selective (cardioselective) blocking drugs. Cardioselective

β -blockers were developed to reduce potentially life-threatening reactions, such as bronchospasm, resulting from β_1 receptor blockade. Acetazolamide, oxprenolol and pindolol are partial agonists, and will induce sympathomimetic effects when there is low sympathetic tone. Uniquely, nebivolol produces a therapeutic mild vasodilating effect through an interaction with the nitric oxide synthesis pathway.

Common adverse effects

The effects of β -blockers are shown in Figures 27.15 and 27.16. Common adverse effects include dizziness, lethargy, insomnia and diarrhoea. Contraindications include known hypersensitivity, heart block, severe heart failure, cardiogenic shock and other severe circulatory disorders, bradycardia with a heart rate of less than 45–50 beats per minute, sick sinus syndrome, atrioventricular block, severe hypertension or uncontrolled heart failure. They should also not be used in people with a history of asthma or chronic obstructive pulmonary disease.

Clinical considerations

Application for β_1 antagonists are to be found in the control of cardiac disease, hypertension, migraine prophylaxis, situational anxiety and thyrotoxicosis. In a seemingly counter-intuitive way, metoprolol, bisoprolol and carvedilol have been used judiciously in the management of heart failure (for details see Chapter 50). There are no clinical applications for β_2 antagonists.

Always withheld of β antagonists may accentuate angina or produce rebound hypertension, myocardial infarction or ventricular dysrhythmias. It is, therefore, important that β antagonists be slowly reduced when treatment is to cease. Cardioselective β antagonists may be preferred in conditions such as peripheral vascular disease, Raynaud's syndrome or diabetes mellitus because of their decreased effect on altering glucose metabolism and causing peripheral vasoconstriction. In diabetes, non-selective β antagonists may mask important signs of hypoglycaemia, including tachycardia and tremor, therefore increasing the severity of the condition. However, β_1 selectivity diminishes with higher doses of the medicine.

NON-SELECTIVE ADRENERGIC BLOCKING AGENTS

Mechanism of action

Carvedilol and labetalol non-selectively block both α_1 and α_2 adrenoceptors in the periphery.

Human Models visually illustrate the effects, both positive and negative, of pharmacological agents on the human body. Male and female human models are used to illustrate the effects of pharmacological agents.

Case Studies with Accompanying Questions immerse students in scenarios involving people taking medicines, family members and health professionals. Students are given the opportunity to apply knowledge, practise drug calculations and dosages, and convey their understanding of pharmacological principles and interactions in a variety of clinical settings.

SECTION 1 PHARMACOLOGY WITHIN THE SOCIAL CONTEXT

CHAPTER REVIEW

- Advertising of medicines can affect the medicine management activities of health professionals.
- Advertising can influence the medicinal activities of consumers.
- Over-the-counter preparations are available to consumers without a prescription, and often without supervision of a health professional.
- The generic name of a medicine is the shortened, simplified version of the chemical name.
- The brand name is the trademark used by a pharmaceutical pharmaceutical company to identify the preparation of a particular drug.
- Generic prescribing means that a pharmacist can supply any formulation of a particular medicine.
- Generic substitution means that a pharmacist can supply any formulation of the medicine without referring back to the prescriber.
- Polyprescription, which is a major problem for older people, involves the excessive or inappropriate use of medicines.
- The traditional beliefs and values of a particular culture influence individuals' perceptions and expectations about drug therapy.

FURTHER READING

- Banning M, 2007, *Medication Management in Care of Older People*, Blackwell Publishing, Oxford.
- Cermody D & Mansfield PR, 2010, 'What do medical students think about pharmaceutical promotion?' *Australian Medical Student Journal*, 11, 54–7.
- DeLorme DE & Huh J, 2009, 'Sensors' uncertainty management of direct-to-consumer prescription drug advertising usefulness', *Health Communication*, 24, 494–503.
- Hamilton HJ, Gallagher PJ & O'Mahony D, 2009, 'Inappropriate prescribing and adverse drug events in older people', *BMJ Geriatrics*, 9, 5, 28.
- Perry DE, Patel AA, Cass A, Howard MR, Tchan ML, Brady JR, De Vries J, Rickards BA, Varnold DJ, Hayman NE & Brown AD, 2009, 'Cardiovascular disease risk management for Aboriginal and Torres Strait Islander peoples in primary health care settings: Findings from the Kanyini audit', *Medical Journal of Australia*, 191, 304–9.
- Spartling GK, Mansfield PR, Montgomery BD, Lescchin J, Doust J, Othman N & Vitry AJ, 2010, 'Information from pharmaceutical companies and the quality, quantity, and cost of physicians' prescribing: a systematic review', *Public Library of Science Medicine*, 19, 7(10), e1000352.
- Wessell AM, Nierster PJ, Jenkins RG, Nemeth LS & Ornstein SM, 2008, 'Inappropriate medication use in the elderly', *American Journal of Geriatric Pharmacotherapy*, 6, 21–7.

WEB RESOURCES

- A Brief History of Pharmacology pubs.acs.org/subscribe/journals/mvld/v04/i05/html/05timeline.html
- Australian Bureau of Statistics www.abs.gov.au/AUSSTATS
- What is Pharmacology? www.pharmacology.med.unn.edu/whatispharm.html
- Everybody (Health Consumer Information) www.everybody.co.nz
- Maori Health www.health.govt.nz/our-work/populations/maori-health
- New Zealand Deserves Better: Direct-to-Consumer Advertising (DCA) of Prescription Medicines in New Zealand for Health or Profit? journal.nzma.org.nz/journal/116/1180/564
- Office for Aboriginal and Torres Strait Islander Health www.health.gov.au/ostsihs

CASE STUDY 1

Mrs. JH is a 62-year-old woman who has had rheumatoid arthritis in her hands, hips and knees for about eight years. She is receiving weekly assistance from her local district nursing service because of impaired mobility. For the arthritis, she is taking the non-steroidal anti-inflammatory drug ibuprofen daily and receives intermittent hydrocortisone therapy when the condition worsens.

You are caring for Mrs. JH. She tells you that her eyes have not been the best of late and she is finding it hard to see things out of the corners of her eyes. She is referred to her family doctor. He, in turn, refers her to the local eye clinic where a diagnosis of open-angle glaucoma is made. Mrs. JH is prescribed eye drops containing a miotic agent. This medicine causes pupil constriction and facilitates the drainage of aqueous humour through the canal of Schlemm.

Questions

1. Applying your knowledge of adrenergic and cholinergic pharmacology, which groups of drugs are well suited as miotics?
2. What receptor types are they acting on and how are they affecting the function of these receptors?
3. State three common side-effects associated with each of these drug groups.
4. Would you expect to observe systemic side-effects associated with this therapy? Why?
5. Referring to Chapter 18, explain why Mrs. JH may be predisposed to glaucoma.

CASE STUDY 2

Mr. JT is a 22-year-old man who has been admitted to your hospital emergency department. He has been working as a labourer at a nearby market garden that specialises in growing flowers. He was spraying the crops with the organophosphate insecticide malathion when he collapsed. He was not wearing the appropriate protective clothing. You observe that he is conscious and complains of gastrointestinal cramps and nausea. He vomited a couple of times in the ambulance as he was transported to hospital. You note the following manifestations: profuse sweating, drooling, lacrimation, bradycardia, agitation, muscle twitching and constricted pupils.

Supportive treatment is implemented, which involves respiratory support and the administration of antidotes. His progress is carefully monitored during this critical period. His recovery is without complications. He is discharged from hospital several days later.

Questions

1. Underlying Mr. JT's condition is a change in the level of activity of a division of the autonomic nervous system. Which division is affected and what is the nature of the change?
2. Which type or types of histamine receptor are involved in this condition?
3. Explain the mechanism by which the organophosphate insecticides induce this state.
4. Which clinical drug group do the organophosphate insecticides closely resemble in terms of their action? Why?
5. Which drug group can be used as an antidote to reverse the effects of the insecticide? Why?

CASE STUDY 3

Mr. JJ, aged 68 years, visits the outpatient clinic for a check-up relating to his asthma condition. He has occasional bouts of acute asthma, which is adequately controlled using a salbutamol inhaler. Mr. JJ indicates that he has just been diagnosed with open-angle glaucoma, which is being treated with timolol 0.25% eye drops. He inserts one drop in each eye twice daily. The outpatient nurse ascertains that he has used the eye drops for two days.

Questions

1. To which drug group does timolol belong and how does it act to lower intraocular pressure? You may wish to refer to Chapter 27.
2. To which drug group does salbutamol belong, and how does it act to relieve asthma? You may wish to refer to Chapters 22 and 33.
3. What is the potential problem for Mr. JJ using salbutamol and timolol?

CASE STUDY 4

Ms. RW is a 50-year-old woman who is suffering from sinus bradycardia (a slow heart rate). Recently, she has had some problems maintaining a normal blood pressure. She is given a medicine that acts on the autonomic innervation of the heart and returns her heart rate to normal.

Questions

1. State the divisions involved, the transmitters released, the receptors concerned and the effects associated with autonomic nervous system innervation of the heart.
2. Name the possible cholinergic and/or adrenergic drug groups that could be used to reverse Ms. RW's bradycardia.

Chapter Review summarises the essential information in each chapter, providing a quick revision tool.

REVIEW QUESTIONS

1. What are the major functions of skin?
2. Indicate the major characteristics of each of the following skin layers:
 - a. dermis
 - b. stratum basale
 - c. stratum corneum
3. Outline the major characteristics of each of the following skin preparations:
 - a. lotion
 - b. gel
 - c. rubefacient
 - d. keratolytic
 - e. cream
4. For each of the following drug groups, indicate the skin condition(s) that they are used to treat:
 - a. antimicrobial agents
 - b. corticosteroids
 - c. immunomodulators
 - d. keratolytics
5. Outline the pathophysiology of the following two conditions:
 - a. acne
 - b. psoriasis
6. For each of the following agents used in the management of psoriasis, indicate whether it is directed towards reducing the inflammation or the rate of cell proliferation:
 - a. the corticosteroids
 - b. methotrexate
 - c. PUVA therapy
 - d. cyclosporin
7. Ebony Tinselle is 17 years old and receiving treatment with the retinoid isotretinoin for severe acne. What medicine education would you offer Ebony?
8. Mark Mitchell is a 35-year-old man about to commence methotrexate therapy for psoriasis. What baseline examinations are required for this therapy? What advice should Mark receive regarding his therapy?
9. Two of the four primary school-aged children in Charlotte Austen's family have head lice infestations. Charlotte buys a shampoo containing piperonyl butoxide and a pyrethrin. Briefly outline the treatment approach to eradicate the infestation.
10. Judy Jones, a 35-year-old mother of two young children, is ordered a dithranol preparation to treat her psoriasis. With what medicine education would you provide her for the application of dithranol? What extra care should she take when tending to her children?
11. Martha Bortolli, a 15-year-old student, complains of pimples and blackheads on her face and back. As the health professional who examines Martha, you recommend a benzoyl peroxide cream to treat the acne. How would you advise Martha on the use of the cream?
12. Jack Brown, aged 35 years, begins a course of treatment with minoxidil liquid for the treatment of androgenic alopecia. What counselling would you offer Mr Brown about using this liquid?

Review Questions check that students remember and understand the clinical significance of key chapter content.

Further Reading lists appear at the end of each section and provide information for students wishing to pursue a topic in further detail for assessment or interest.

FURTHER READING

- Burton R, 2006. Pharmacokinetics and pharmacodynamics: the dynamics of drug absorption, distribution and elimination. In: Brunton LL, Lazo JS & Parker KL, eds. *Clinical Pharmacology and Therapeutics*, 11th edn. McGraw-Hill, New York, pp. 1–39.
- Eshkol T, Shleier E, Ben-Zvi Z & Hsieh C, 2011. Drug transport across the placenta. *Current Pharmaceutical Biotechnology*, 12(8): 707–14.
- Hughes CM, Roughhead E & Kersa N, 2008. Improving use of medicines for older people in long-term care: constructing the policy approach of four countries. *Healthcare Policy*, 3(3): e154–e167.
- Jacobs Aggrain E & Chomera L, eds, 2006. *Pharmaceutical Clinical Pharmacology, Informa HealthCare*, London.
- Knoxworth SC, Wong H & Hup CCA, 2011. *Drug Metabolism and Pharmacokinetics Quick Guide*. Springer, New York.
- Koch S, Glash M, Nay R, eds, 2010. *Medication Management in Older Adults: A Concise Guide for Clinicians*. Springer Science and Business Media, New York.
- Le Conte D, McLachlan A & de Cabo R, 2012. Aging, drugs, and drug metabolism. *Journal of Gerontology Series A: Biological Sciences & Medical Sciences*, 67(2): 137–50.
- McCance K & Huether SE, 2009. *Nephropathology*, 6th edn, Elsevier Mosby, Sydney (for age-related and disease-related changes in body structure and function).
- Sassung TM, Troutman SM, Campbell TJ, Preskier HM, Sung H, Bates SE & Figg WD, 2012. Transporter pharmacokinetics: transporter polymorphisms affect normal physiology, diseases, and pharmacotherapy. *Discovery Medicine*, 13(68): 19–34.
- The Royal Australian College of General Practitioners, Australasian Society of Clinical and Experimental Pharmacologists and Toxicologists, Pharmaceutical Society of Australia, 2012. *Australian Medicines Handbook*, AMH Pty Ltd, Adelaide.
- Weiss N, He SM, Li XT, Wang LL & Zhou SF, 2008. Placental drug disposition and its clinical implications. *Current Drug Metabolism*, 9: 106–21.

WEB RESOURCES

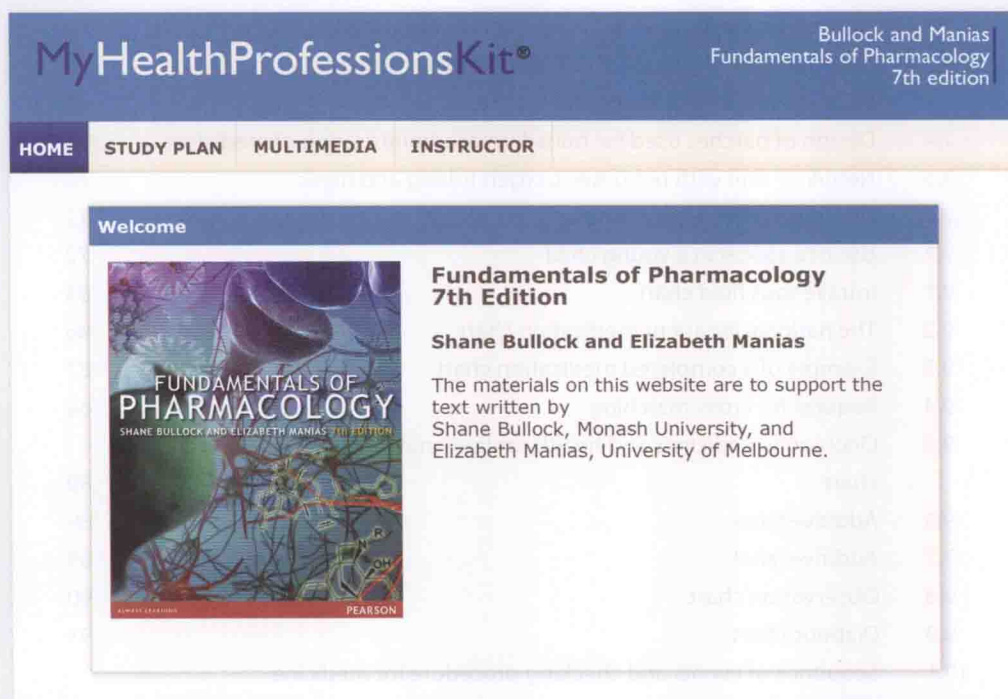
- Australian Government Department of Health and Ageing www.health.gov.au
- Australian Statistics on Medicines www.tga.gov.au/hp/medicines-statistics-2010.htm
- Clinical Trials (US site) www.clinicaltrials.gov
- Health Insite www.healthinsite.gov.au/index.cfm
- Interactive Clinical Pharmacology www.icp.org.au
- Medicines Australia (Pharmaceutical Industry Group) www.medicinesaustralia.com.au
- MedSafe www.medsafe.govt.nz
- NZ Ministry of Health www.moh.govt.nz/medsafe/
- Pharmacokinetics: An Introduction (US site) [www.austlii.edu.au/other/australianlegaljournal/pharmacokinetics/](http://www.austlii.edu.au/au/other/australianlegaljournal/pharmacokinetics/)
- Therapeutic Goods Administration (TGA) www.tga.gov.au/index.htm
- Trials Central: online register of US clinical trials www.trialscentral.org

WEB RESOURCES

- Better Health Channel: Haemorrhoids www.betterhealth.vic.gov.au/1hcv2/bhcarticles.nsf/pages/haemorrhoids
- Centre for Digestive Diseases: Disease Information www.cdd.com.au
- Crohn's and Colitis Australia www.cacca.net.au
- Gastroenterological Society of Australia: Professional Information www.gesa.org.au
- Health Insite: Digestion and Stomach Disorders www.healthinsite.gov.au/topics/Digestion_and_Stomach_Disorders
- Medline Plus: Constipation (US site) www.nlm.nih.gov/medlineplus/constipation.html
- Medline Plus: Nausea and Vomiting (US site) www.nlm.nih.gov/medlineplus/nauseaandvomiting.html
- Nausea and Vomiting During Pregnancy (Canadian site) www.sogc.org/health/pregnancy/nausea_v.htm
- Primary Care Society for Gastroenterology (UK site) www.pcsg.org.uk

Web Resources lists appear at the end of each section and provide links to relevant websites for further study and online research.

TEACHING AND LEARNING PACKAGE



FOR STUDENTS

MyHealthProfessionsKit is an online study tool that will help you understand, revise and master the concepts in the textbook.

MyHealthProfessionsKit gives you access to these study resources:

- multiple-choice revision questions;
- interactive 'drag and drop' revision activities;
- animations demonstrating the mechanisms of action for various medicines;
- glossary flashcards to test your knowledge of key pharmacology terms;
- realistic drug calculation scenarios to give you practice;
- searchable eBook (if you have purchased the MyLab with eBook option).

FOR INSTRUCTORS

Computerised TestBank

Create professional-looking customised printed or online exams in just minutes using Pearson's TestGen software. Build tests from the database of over than 600 true-false and multiple-choice questions, edit questions or add questions of your own.

PowerPoint Slides

Lecture slides pair key points with images from each chapter to facilitate effective lectures and classroom discussions.

Solutions Manual

This manual provides the answers to the end-of-chapter exercises in the text. You have the option of making this manual available to your students.

Digital Media Library

All figures and tables from the textbook are provided in jpeg format.

CONTENTS



	Preface	viii
	Acknowledgments	x
	Features	xi
	Teaching and learning package	xv
	Figures and tables	xvi
SECTION I	PHARMACOLOGY WITHIN THE SOCIAL CONTEXT	1
	1 A historical perspective	2
	2 Sociocultural aspects	8
SECTION II	PHARMACOLOGY WITHIN THE PROFESSIONAL CONTEXT	19
	3 Health professionals and the law	21
	4 Ethical issues	30
	5 The roles and responsibilities of health professionals in medicine adherence, education and advocacy	39
	6 The roles and responsibilities of health professionals in medicine management	44
SECTION III	MEDICINE ADMINISTRATION AND PROFESSIONAL RESPONSIBILITIES	55
	7 Medicine formulations, storage and routes of administration	56
	8 The clinical decision-making process	76
	9 Medicine administration strategies and documentation	82
	10 Medication errors	92
	11 Management of common adverse drug reactions	96
	12 Risk communication: balancing the benefits and risks of drug treatment	113
SECTION IV	GENERAL ASPECTS OF PHARMACOLOGY	125
	13 Drug nomenclature	127
	14 Pharmacokinetics: absorption and distribution	133
	15 Pharmacokinetics: metabolism and excretion	142
	16 Drug interactions	152
	17 Pharmacodynamics	159
	18 Drug development, evaluation and safety	171
	19 Pharmacogenetics	182
	20 Pharmacokinetic factors that modify drug action	190
	21 Paediatric and geriatric pharmacology	195
SECTION V	TOXICOLOGY	209
	22 Poisoning and envenomation	210
	23 The management of acute clinical overdose	218
	24 Contemporary drugs of abuse	227
	25 Drug abuse in sport	239

SECTION VI	AUTONOMIC PHARMACOLOGY	249
	26 General aspects of neuropharmacology	250
	27 Adrenergic pharmacology	258
	28 Cholinergic pharmacology	286
SECTION VII	CHEMICAL MEDIATORS	313
	29 An introduction to chemical mediators	314
	30 Histamine and antihistamines	318
	31 Prostaglandins and serotonin	329
	32 Nitric oxide and the endothelins	337
SECTION VIII	THE MODULATION OF BEHAVIOUR, COGNITION AND MOTOR ACTIVITY	345
	33 General concepts of psychopharmacology	347
	34 Antipsychotic agents	352
	35 Anxiolytics and hypnotics	367
	36 Antidepressants and mood stabilisers	380
	37 Medicines used in neurodegenerative disorders	399
	38 Antiseizure agents and muscle relaxants	418
	39 Central nervous system stimulants	437
SECTION IX	MEDICINES USED TO RELIEVE PAIN AND PRODUCE ANAESTHESIA	447
	40 Narcotic analgesics	448
	41 Non-steroidal anti-inflammatory, antipyretic and analgesic agents	466
	42 Medicines used to treat migraine	488
	43 General anaesthesia	498
	44 Local anaesthesia	508
SECTION X	THE MODULATION OF OXYGENATION AND PERFUSION	521
	45 Medicines used to lower blood lipids	522
	46 Antihypertensive agents	534
	47 Medicines used to promote tissue perfusion	556
	48 Antithrombotic, fibrinolytic and haemostatic agents	572
	49 Diuretics and other renal medicines	594
	50 Medicines used to treat heart failure	606
	51 Medicines used to treat cardiac dysrhythmia	626
	52 Fluid and potassium imbalances	636
	53 Antianaemic agents	651
	54 Medicines used to maintain gas exchange	659
	55 Medicines for upper respiratory tract conditions	681
SECTION XI	THE MODULATION OF GASTROINTESTINAL FUNCTION	697
	56 Upper gastrointestinal tract drugs	698
	57 Lower gastrointestinal tract drugs	713
	58 Antiemetic agents	734

SECTION XII	THE MODULATION OF BODY GROWTH, DEVELOPMENT AND METABOLISM	747
59	Medicines and the pituitary gland	749
60	Medicines and the thyroid	762
61	Medicines and the pancreas	770
62	Medicines and the adrenal cortex	791
63	Medicines and the gonads	801
64	Medicines and bone metabolism	822
65	Hyperuricaemia and gout	833
66	Obesity and its treatment	842
SECTION XIII	NUTRITIONAL AND NATURAL THERAPIES	855
67	Vitamins, minerals and amino acids	856
68	Enteral and parenteral nutrition	876
69	Herbal medicines	886
SECTION XIV	THE MODULATION OF CELLULAR GROWTH AND PROLIFERATION	907
70	Introduction to chemotherapy	909
71	Sulfonamides and trimethoprim	917
72	Antibacterial agents	923
73	Antituberculotics and antileprotic agents	947
74	Antiseptics and disinfectants	960
75	Antiparasitic agents	969
76	Antimalarial agents	981
77	Antiviral agents	991
78	Antifungal agents	1012
79	Immunomodulating agents	1025
80	Cytotoxic chemotherapeutic agents	1050
81	Gene therapies	1080
SECTION XV	MEDICINES USED TOPICALLY	1089
82	Medicines used in diseases of the skin	1090
83	Medicines and the eye	1111
APPENDIX	A Common prescription terminology	1135
	B Common American generic drug names	1136
	C SI units	1137
	D Medicine calculations	1140
	E Common symbols used in medication charts	1145
	F Common word mix-ups	1146
	G Drug-herbal interactions	1148
	H Orphan drugs	1150
Glossary		1153
Index		1165

FIGURES AND TABLES

FIGURE	1.1	A time line highlighting some major pharmaceutical events	4
	1.2	Medicinal plants commonly found in suburban gardens	6
	2.1	Prescription indicating that brand substitution is permitted	13
	7.1	Appropriate technique for administering eye drops	62
	7.2	Appropriate technique for administering ear drops	62
	7.3	Location of sublingual and buccal sites	63
	7.4	Design of patches used for transdermal administration of medicines	65
	7.5	Nebuliser unit with nebuliser, oxygen tubing and mask	72
	7.6	Use of a metered-dose inhaler	72
	7.7	Use of a spacer in a young child	72
	9.1	Intravenous fluid chart	84
	9.2	The national inpatient medication chart	86
	9.3	Example of a completed medication chart	87
	9.4	Request for cross-matching	88
	9.5	Once-only medicines and health professional-initiated medications chart	89
	9.6	Additive label	89
	9.7	Additive label	89
	9.8	Observation chart	90
	9.9	Diabetic chart	91
	10.1	Sequence of events and checking procedure for medicine administration	94
	13.1	An example of drug nomenclature	129
	13.2	Molecular structures of three tricyclic antidepressants	131
	14.1	Drug absorption between body compartments	134
	14.2	Factors influencing drug absorption	136
	14.3	Protein-bound drug reservoir	137
	14.4	Ion trapping	138
	14.5	Apparent volume of distribution	139
	15.1	Drug metabolism	143
	15.2	Differences in spatial molecular arrangements of glucose and galactose	144
	15.3	The enterohepatic cycle	145
	15.4	Hepatic first-pass effects	146
	15.5	Drug excretion	147
	15.6	The development of a steady-state concentration	148
	15.7	The effect of a loading dose on plasma concentration	149
	15.8	Kinetics of drug metabolism	149