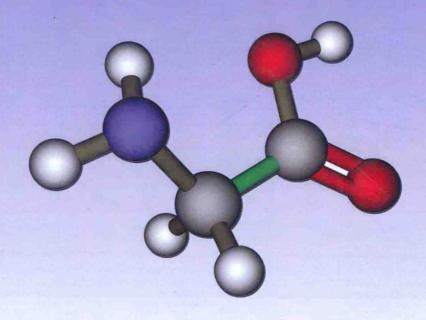
# AMINO ACIDS

**Biochemistry and Nutrition** 



**GUOYAO WU** 



# AMINO ACIDS

Biochemistry and Nutrition 版 一 章 Guoyao Wu



CRC Press Taylor & Francis Group 6000 Broken Sound Parkway NW, Suite 300 Boca Raton, FL 33487-2742

© 2013 by Taylor & Francis Group, LLC CRC Press is an imprint of Taylor & Francis Group, an Informa business

No claim to original U.S. Government works

Printed on acid-free paper Version Date: 20130220

International Standard Book Number-13: 978-1-4398-6189-9 (Hardback)

This book contains information obtained from authentic and highly regarded sources. Reasonable efforts have been made to publish reliable data and information, but the author and publisher cannot assume responsibility for the validity of all materials or the consequences of their use. The authors and publishers have attempted to trace the copyright holders of all material reproduced in this publication and apologize to copyright holders if permission to publish in this form has not been obtained. If any copyright material has not been acknowledged please write and let us know so we may rectify in any future reprint.

Except as permitted under U.S. Copyright Law, no part of this book may be reprinted, reproduced, transmitted, or utilized in any form by any electronic, mechanical, or other means, now known or hereafter invented, including photocopying, microfilming, and recording, or in any information storage or retrieval system, without written permission from the publishers.

For permission to photocopy or use material electronically from this work, please access www.copyright.com (http://www.copyright.com/) or contact the Copyright Clearance Center, Inc. (CCC), 222 Rosewood Drive, Danvers, MA 01923, 978-750-8400. CCC is a not-for-profit organization that provides licenses and registration for a variety of users. For organizations that have been granted a photocopy license by the CCC, a separate system of payment has been arranged.

**Trademark Notice:** Product or corporate names may be trademarks or registered trademarks, and are used only for identification and explanation without intent to infringe.

#### Library of Congress Cataloging-in-Publication Data

Wu, Guoyao, 1962-

Amino acids: biochemistry and nutrition / Guoyao Wu. pages cm

"A CRC title."

Includes bibliographical references and index. ISBN 978-1-4398-6189-9 (hardcover: alk. paper)

1. Amino acids. 2. Amino acids in human nutrition. I. Title.

QP561.W8 2013 612'.015756--dc23

2013003657

Visit the Taylor & Francis Web site at http://www.taylorandfrancis.com

and the CRC Press Web site at http://www.crcpress.com

### **Preface**

Amino acid biochemistry and nutrition are interesting, dynamic, and challenging subjects in biological sciences. They span a wide range from chemistry, metabolism, and physiology to reproduction, immunology, pathology, and cell biology. In the nearly half-century since the last two volumes of the monograph Biochemistry of the Amino Acids were published by Alton Meister in 1965, this field has developed rapidly. Important technical and conceptual advances include (1) the analysis of amino acids by high-performance liquid chromatography and mass spectrometry; (2) isotopic measurements of synthesis and degradation of proteins and amino acids in cells, tissues, and the whole body; (3) interorgan metabolism of amino acids involving key tissues and organs such as liver, skeletal muscles, small intestine, and kidneys; (4) amino acids in cell signaling and regulation of gene expression; (5) nonlysosomal protein degradation by ATP- and ubiquitin-dependent multicatalytic proteasomes; (6) molecular and cellular regulation of intracellular protein turnover and amino acid metabolism; (7) determination of true ileal digestibilities of amino acids; (8) molecular cloning of transporters for amino acids and small peptides; (9) development of the ideal protein concept; and (10) elucidation of dietary requirements of functional amino acids by humans and other animals. Comprehensive and systematic coverage of these new scientific developments in a well-organized book will benefit researchers and students in both biomedical and agricultural disciplines to improve health and the nutritional value of foods.

Amino Acids: Biochemistry and Nutrition consists of 13 chapters. The text starts with the discoveries and basic concepts of amino acids, peptides, and proteins. It then advances to protein digestion in the gastrointestinal tract and the absorption of small peptides and free amino acids in the small intestine. This chapter is followed by detailed coverage of cell- and tissue-specific synthesis and catabolism of amino acids and related nitrogenous substances (including nitric oxide, polyamines, glutathione, creatine, urea and uric acid) in animals. After the use of isotopes in studying nitrogen metabolism is introduced in Chapter 7, the book continues with intracellular protein turnover, short- and long-term regulation of amino acid metabolism, physiological functions of amino acids, and inborn errors of amino acid metabolism. Finally, the text ends with dietary requirements of amino acids by humans and other animals. While the classical concepts and principles of amino acid biochemistry and nutrition are emphasized throughout the book, every effort has been made to include the most recent progress in this ever-expanding field so that readers in various biological disciplines can integrate amino acid biochemistry with nutrition, health, and disease in mammals, birds, and other animal species. At the end of each chapter, selected references are listed to provide readers with both comprehensive reviews of the chosen topics and original experimental data on which modern concepts in amino acid biochemistry and nutrition are based. Reading the scientific literature is essential for a thorough understanding of the history of the field and also provides "food" for creative thinking and for rigorous development as a productive scientist.

xviii Preface

In the Index, a list of key words, phrases, and abbreviations is provided to help readers quickly find information presented in all the chapters.

This book owes its origin to the lecture notes of a graduate course ANSC/NUTR 613 "Protein Metabolism" the author has taught at Texas A&M University for the past 21 years. Its conception was motivated largely by the lack of a suitable text-book for teaching such an advanced course for students majoring in animal science, biochemistry, biomedical engineering, biology, human medicine, kinesiology, veterinary medicine, nutrition, physiology, toxicology, and other related disciplines. Besides its use as a textbook, all of the chapters also provide useful references to general and specific knowledge on amino acid biochemistry and nutrition for researchers in biomedicine and agriculture (including animal science and plant breeding).

The sciences of amino acid biochemistry and nutrition have been built on the shoulders of many giants and pioneers worldwide. Their seminal contributions to the field have made this book possible. The author must apologize to those whose published works are not cited in the text due to limited space. Sincere thanks are also extended to the author's past and current students for their constructive comments on the "Protein Metabolism" course and for their stimulating discussions to further improve its content.

### Acknowledgments

This book was initiated at the invitation of Dr. Ira Wolinsky, a consulting editor for CRC Press, and completed with the patience and guidance of its editorial staff, Randy L. Brehm and Kari A. Budyk. The author would like to thank Sudath Dahanayaka for drawing structures and metabolic pathways of amino acids, Dr. Gregory A. Johnson for illustrating enterocytes of the small intestine, and Syed Mohamad Shajahan and his team at Techset Composition for their professional typesetting of the entire manuscript. Sincere thanks also go to a group of accomplished scientists who critically reviewed various chapters and provided constructive suggestions for improvement: Drs. Vickie E. Baracos, Fuller W. Bazer, Brian J. Bequette, John T. Brosnan, Teresa A. Davis, Catherine J. Field, Nick E. Flynn, C. Andy Hu, Darrell A. Knabe, Errol B. Marliss, James C. Matthews, Cynthia J. Meininger, Sidney M. Morris Jr., J. Marc Rhoads, James R. Thompson, and Malcolm Watford.

The author is appreciative of important contributions of his former and current students, postdoctoral fellows, visiting scholars, and technicians to the conduct of research and valuable discussions, particularly Drs. Houssein Assaad, Douglas G. Barstow Jr., Arturo G. Borbolla, Christopher M. Dekaney, Zhaolai Dai, Sujay Datta, J. Todd Davidson, E. Lichar Dillon, Michael E.R. Dugan, Nick E. Flynn, Wenjiang Fu, Haijun Gao, Fei Ji, Wenjuan Jobgen, Xiangfen Kong, Hyuk-Jung Kwon, Arantza Lassala, Jian Lei, Hui Li, Peng Li, Xilong Li, Maria E. Mahata, Ronald D. Mateo, Jason R. McKnight, Yose Rizal, M. Carey Satterfield, Jon T. Self, Bie Tan, Carmen D. Tekwe, Junjun Wang, Xiuqi Wang, Jiawei Wei, Zhenlong Wu, Pengbin Xi, Haipeng Yang, Ying Yang, Kang Yao, and Yulong Yin as well as Professor Lixiang Chen, Sudath Dahanayaka, Mou Dasgupta, Paula K. Davis, Sean P. Flynn, Chenxing Fu, Merrick Gearing, Tony E. Haynes, Sichao Jia, Scott C. Jobgen, Ning Jiao, Ripla Kohli, Katherine A. Kelly, Gang Lin, Chuang Liu, Sabina A. Meier, Reza Rezaei, Robin R. Simon, Yuli Sun, Bin Wang, Weiwei Wang, Xiaoqiu Wang, Natalie A. Morrow, Wene Yan, Wei Ying, and Chen Zhang.

The author is grateful to his graduate advisors Professor Sheng Yang and Dr. James R. Thompson, as well as postdoctoral mentors Dr. Errol B. Marliss and Dr. John T. Brosnan for training in amino acid biochemistry and nutrition, as well as their enthusiastic support and inspiration for a life-long pursuit of this discipline. Productive and enjoyable collaborations with colleagues at Texas A&M University (particularly Drs. Fuller W. Bazer, Robert C. Burghardt, Douglas G. Burrin, Raymond J. Carroll, Timothy A. Cudd, Teresa A. Davis, Kathrin A. Dunlap, William C. Ellis, L. Wayne Greene, Cristine L. Heaps, Laurie A. Jaeger, Gregory A. Johnson, Sung Woo Kim, Darrell A. Knabe, Joanne R. Lupton, Catherine J. McNeal, Cynthia J. Meininger, Judy M. Muller-Delp, Wilson G. Pond, M. Carey Satterfield, Joseph R. Sharkey, Stephen B. Smith, Thomas E. Spencer, Yanan Tian, Nancy D. Turner, Shannon E. Washburn, Beiyan Zhou, and Huaijun Zhou), at other U.S. institutions (particularly Drs. Matthew A. Boegehold, Gautam Chaudhuri, Susan K. Fried, David A. Geller, Bernhard Hennig, Louis J. Ignarro, Duane H. Keisler, Sidney M. Morris, Jr., Juan

B. Ochoa, J. Marc Rhoads, Penelope Perkins-Veazie, Nathalie L. Trottier, Kimberly A. Vonnahme, Malcolm Watford, and Feng-Qi Zhao), and in China (particularly Drs. Yongqing Hou, Zongyong Jiang, Defa Li, Ning Li, Shiyan Qiao, Haizhou Sun, Fenglai Wang, Junjun Wang, Zhenlong Wu, Weiyun Zhu, Yulong Yin, and Yongliang Zhang) are gratefully acknowledged. The author is indebted to Drs. David H. Baker, Fuller W. Bazer, H. Russell Cross, Susan K. Fried, Harris J. Granger, Bryan H. Johnson, Peter J. Reeds, and Malcolm Watford for their mentoring of career development and for their professional support.

Work in the author's laboratory was supported, in part, by grants from Ajinomoto Inc. (Tokyo, Japan), American Heart Association, The Chinese Academy of Sciences, Gentech Inc. (Shanghai, China), Houston Livestock Show and Rodeo, International Council of Amino Acid Science (Brussels, Belgium), International Glutamate Technical Committee (Brussels, Belgium), JBS United (Sheridan, Indiana), Juvenile Diabetes Research Foundation (USA), National Institutes of Health (USA), National Natural Science Foundation of China, Pfizer Inc., Scott & White Hospital (Temple, Texas), Texas A&M AgriLife Research, Texas A&M University, the Thousand-People Talent Program at China Agricultural University, United States Department of Agriculture, U.S. National Corn Growers Association, U.S. National Watermelon Promotion Board, and U.S. Poultry & Egg Harold E. Ford Foundation.

Special thanks are extended to the author's parents for being role models of patience, kindness, strong work ethics, integrity, and respect for others. Finally, the great assistance and support of the author's wife (Yan Chen), son (Neil D. Wu), and daughter (Jennifer M. Wu) in preparing the entire manuscript are gratefully acknowledged.

#### **Author**

**Dr. Guoyao Wu** is a university distinguished professor, university faculty fellow, and Texas A&M AgriLife Research senior faculty fellow at Texas A&M University. He received a BS in animal science from the South China Agricultural University at Guangzhou, China (1978–1982), an MS in animal nutrition from China Agricultural University at Beijing, China (1982–1984), and an MSc (1984–1986) and PhD (1986–1989) in animal biochemistry from The University of Alberta at Edmonton, Canada. Dr. Wu completed his postdoctoral training in diabetes, nutrition, and biochemistry at McGill University Faculty of Medicine in Montreal, Canada (1989–1991) and Memorial University of Newfoundland Faculty of Medicine in St. John's, Canada (1991). He joined the Texas A&M University faculty in October 1991. His sabbatical leave was to study human obesity at the University of Maryland School of Medicine in Baltimore, USA (2005).

Dr. Wu has taught graduate (experimental nutrition, general animal nutrition, protein metabolism, and nutritional biochemistry) and undergraduate (problems in animal science, nutrition, and biochemistry) courses at Texas A&M University for the past 21 years. He has given numerous lectures at other institutions in the United States, Canada, Mexico, Brazil, Europe, and Asia. His research focuses on the biochemistry, nutrition, and physiology of amino acids (AA) in animals at genetic, molecular, cellular, and whole-body levels. Research interests include (1) functions of AA in gene expression (including epigenetics) and cell signaling; (2) biochemical mechanisms that regulate intracellular synthesis and catabolism of proteins and AA; (3) hormonal and nutritional regulation of homeostasis of metabolic fuels; (4) biology and pathobiology of nitric oxide and polyamines; (5) key roles of AA in preventing metabolic diseases (including diabetes, obesity, and intrauterine growth restriction) and associated cardiovascular complications; (6) essential roles of AA in survival, growth and development of embryos, fetuses, and neonates; (7) dietary requirements of AA and proteins in the life cycle; and (8) animal models (e.g., pigs, rats, and sheep) for studying human metabolic diseases. He has published 365 papers in peer-reviewed journals, including Amino Acids, American Journal of Physiology, Annual Review of Nutrition, Biochemical Journal, Biology of Reproduction, British Journal of Nutrition, Cancer Research, Clinical and Experimental Immunology, Comparative Biochemistry and Physiology, Diabetes, Diabetologia, Endocrinology, FASEB Journal, Frontiers in Bioscience, Gut, Journal of Animal Science, Journal of Biological Chemistry, Journal of Nutrition, Journal of Nutritional Biochemistry, Journal of Pediatrics, Journal of Physiology (London), Livestock Science, and Proceedings of National Academy of Science (USA), and 48 book chapters.

Dr. Wu has received numerous prestigious awards from China, Canada, and the United States, which include the China National Scholarship for Graduate Studies Abroad (1984), the University of Alberta Andrew Stewart Graduate Prize (1989), the Medical Research Council of Canada Postdoctoral Fellowship (1989), the American Heart Association Established Investigator Award (1998), the Texas

xxii Author

A&M AgriLife Faculty Fellow (2001), the Texas A&M University Faculty Fellow (2002), the Nonruminant Nutrition Research Award from American Society of Animal Science (2004), the Outstanding Young Investigator Award from National Science Foundation of China (2005), the Texas A&M Agriculture Program Vice Chancellor's Award for Excellence in Team (2006) and Individual (2008) Research, and in Diversity (2011), the Changjiang Scholar Award from China (2008), the Texas A&M University Distinguished Research Achievement Award (2008), the Texas A&M Agrilife Research Senior Faculty Fellow Award (2008), the Chutian Scholar Award from Hubei Province of China (2008), the FASS-AFIA New Frontiers in Animal Nutrition Research Award from the Federation of Animal Science Societies and American Feed Industry Association (2009), the Dingying Scholar Award from South China Agricultural University (2009), the Thousand-People-Talent Award from China (2010), and the Samburu Collaboration Award from the International Association of Giraffe Care Professionals (2010).

Dr. Wu is a member and an elected fellow of the American Association for the Advancement of Science, as well as a member of the American Heart Association, the American Society of Animal Science, the American Society of Nutrition, and the Society for the Study of Reproduction. He has served on editorial advisory boards for the *Biochemical Journal* (1993–2005), the *Journal of Animal Science and Biotechnology* (2010–present), the *Journal of Nutrition* (1997–2003), and the *Journal of Nutritional Biochemistry* (2006–present), as well as an editor of *Amino Acids* (2008–present), an editor of the *Journal of Amino Acids* (2009–present), and a managing editor of *Frontiers in Bioscience* (2009–present).

## Contents

Acknowledg	gments	.xix
Author		.xxi
Chapter 1	Discovery and Chemistry of Amino Acids	1
	Definition and Nomenclature of AA	
	Definition of AA	1
	Definition of Imino Acids	3
	Isomers of AA	3
	Protein and Nonprotein AA	5
	Free and Peptide (Protein)-Bound AA	5
	Discovery of AA	
	L-AA and Glycine	
	β- and γ-AA with Physiological Significance	
	D-AA	
	Presence of D-AA in Foods	
	Presence of D-AA in the Animal Kingdom	
	Chemical Properties of AA	
	Physical Appearance and Melting Points of Crystalline AA	18
	Tastes of Crystalline AA	
	Solubility of AA in Water and Organic Solvents	
	Zwitterionic Form of AA	
	Stability of AA in Water and Buffered Solutions	22
	Stability of AA in Water under High Pressure and High	
	Temperature	23
	Stability of AA in Strong Acid and Alkaline Solutions	23
	Chemical Reactions of AA	24
	Chemical Reactions of the Amino Group in α-AA	24
	Chemical Reactions of the Carboxyl Group in α-AA	26
	Chemical Reactions of the Side Chain in α-AA	27
	Chemical Reactions Involving Both the Amino and	
	Carboxyl Groups of α-AA	27
	Proteins	30
	Peptides versus Proteins	30
	Summary	31
	References	
Chapter 2	Protein Digestion and Absorption of Peptides and Amino Acids	33
	Classification and Content of Protein in Diets	34
	Classification of Protein	

	Content of Foodstuff Proteins of Animal and Plant Origin	
	Definitions of Digestion and Absorption	
	Digestion	35
	Absorption	
	Developmental Changes in Intestinal Digestion and Absorption	. 37
	Protein Digestion and Absorption of Peptides	
	and AA in Monogastric Animals	38
	Historical Perspective of Protein Digestion and Absorption	38
	Digestion of Dietary Protein in the Gastrointestinal Tract	
	Determination of Protein Digestibility	
	Absorption of Free AA and Small Peptides by the Small	
	Intestine	42
	Transport of Free AA via Transmembrane Transporters	
	Transport of AA via the γ-Glutamyl Cycle	
	Transport of Small Peptides	
	Net Balance of AA across the Small Intestine in Fed Animals	
	Extensive Recycling of Nitrogen in the Intestine of Animals	
	Protein Digestion and Absorption of Peptides and AA	70
	in Ruminants	51
	Nutritional Significance of Protein Digestion in Ruminants	
	Digestion of Dietary Protein in the Gastrointestinal Tract	
		32
	Absorption of Free AA and Small Peptides by the Small	55
	Intestine	)
	Nitrogen Recycling in Ruminants and Its Nutritional	
	Implications	
	Protecting High-Quality Protein from Rumen Degradation	
	Heating	
	Chemical Treatments	
	Polyphenolic Phytochemicals	
	Physical Encapsulation of Proteins or AA	
	Inhibition of AA Degradation	
	Summary	
	References	60
Chapter 3	Synthesis of Amino Acids	63
ompro.		
	Synthesis of AA in Tissues and Cells of Animals	
	General Considerations	
	Cell-, Tissue-, and Species-Specific Synthesis of AA	
	General Pathways for Synthesis of AA in Animal Cells	
	Overall Pathways for Synthesis of AA in Animal Cells	
	Historical Aspects of AA Transamination	
	Specific Pathways for Synthesis of AA in Animal Cells	
	Synthesis of Alanine, Glutamine, and Glutamate in Animal Cells.	72
	Synthesis of Arginine, Citrulline, and Ornithine in Animal Cells.	74
	Synthesis of Cysteine and Taurine in Animal Cells	76

Contents

	Synthesis of Glycine and Serine in Animal Cells	76
	Synthesis of Methylarginines in Animal Cells	
	Synthesis of Proline and Hydroxyproline in Animal Cells	81
	Synthesis of Tyrosine in Animal Cells	82
	Formation of β-Alanine in Animal Cells	
	Pathways for Synthesis of AA in Microorganisms	83
	Overall Pathways for Synthesis of AA in Microorganisms	83
	Pathways for Synthesis of NEAA in Microorganisms	
	Pathways for Synthesis of EAA in Microorganisms	87
	Synthesis of D-AA in Animal Cells and Bacteria	91
	Conversion of D-AA to L-AA in Animal Cells and Bacteria	92
	Summary	93
	References	94
Chapter 4	Degradation of Amino Acids	97
	General Characteristics of AA Degradation in Animal Cells	
	Overall View of AA Catabolism	
	Cell- and Tissue-Specific Degradation of AA	
	Pathways for Degradation of AA in Animal Cells	
	Historical Perspective	105
	Catabolism of Alanine, Aspartate, Asparagine, Glutamate,	
	and Glutamine	
	Catabolism of Arginine, Citrulline, and Ornithine	
	Catabolism of BCAA	
	Catabolism of Glycine and Serine	
	Catabolism of Histidine	
	Catabolism of Lysine	
	Catabolism of Phenylalanine and Tyrosine	
	Catabolism of Proline	
	Catabolism of Sulfur-Containing AA	
	Catabolism of Threonine	
	Catabolism of Tryptophan	
	Catabolism of D-AA in Animal Cells	138
	Catabolism of L-AA and D-AA in Microorganisms	
	Summary	
	References	143
Chapter 5	Synthesis and Catabolism of Special Nitrogenous Substances	
	from Amino Acids	145
	Production of Dipeptides Consisting of Histidine	
	or Its Methylated Derivatives	145
	History of Research on Carnosine and Related Dipeptides in	
	Animal Tissues	145

Synthesis of Carnosine and Related Dipeptides	
Catabolism of Carnosine and Related Dipeptides	149
Synthesis and Degradation of GSH	149
History of GSH Research	149
Concentrations of GSH in Physiological Fluids and Tissues	150
GSH Synthesis	150
Regulation of GSH Synthesis	152
Transport and Degradation of GSH	153
Production of Gly-Pro-Hydroxyproline	153
Abundance of Tripeptide Gly-Pro-Hydroxyproline in Milk	
and Plasma	153
Utilization of Gly-Pro-Hydroxyproline by Animals	154
Synthesis and Catabolism of Polyamines	
History of Polyamine Research	
Polyamine Synthesis	
Pathways of Polyamine Synthesis	
Regulation of Polyamine Synthesis	
Polyamine Degradation	
Synthesis and Utilization of Creatine	
History of Creatine Research	
Creatine Synthesis through Interorgan Cooperation	
Regulation of Creatine Synthesis	
Degradation of Creatine	
Synthesis and Catabolism of L-Carnitine	161
History of Carnitine Research	
Carnitine Synthesis through Interorgan Cooperation	
Regulation of Carnitine Synthesis	
Catabolism of Carnitine	
Synthesis and Catabolism of Purine and Pyrimidine	
Nucleotides	165
History of Purine and Pyrimidine Research	165
Purine and Pyrimidine Bases in Nucleotides	
Synthesis of Purine Nucleotides	
Synthesis of Pyrimidine Nucleotides	168
Regulation of Pyrimidine Nucleotide Synthesis	
Catabolism of Purines and Pyrimidines	170
Heme Synthesis and Catabolism	
History of Heme Research	171
Pathways of Heme Biosynthesis	
Regulation of Heme Biosynthesis	174
Regulation of Heme Synthesis in Hepatocytes	
Regulation of Heme Synthesis in Erythroid Cells	
Catabolism of Heme	
Synthesis and Catabolism of Histamine	
Synthesis of Histamine	
Catabolism of Histamine	178

Energy Requirement for Uric Acid Synthesis	212 214 215 215 216 216 217
Use of Isotopes for Studying Amino Acid Metabolism	221
Basic Concepts about Isotopes	221
Significance of SR and IE of a Tracer	229
Calculation of Product Formation or Metabolic Flux	
Using Radioisotopes	229
Calculation of Product Formation or Metabolic Flux	
Using Stable Isotopes	230
	232
References	243
Protein Synthesis	24
Historical Perspectives of Protein Synthesis Pathway	24
I alliways of I fotom Synthesis in the Cytopiashi and	
	Synthesis of Uric Acid from Ammonia via Adenosine

	Pathway of Protein Synthesis in the Cytoplasm	
	Gene Transcription	
	Initiation of Translation	248
	Peptide Elongation	253
	Termination of Peptide Chain Elongation	255
	Posttranslational Modifications of Newly Synthesized	
	Proteins	256
	Pathway of Protein Synthesis in Mitochondria	258
	Biochemical Characteristics and Significance of Protein Synthesis	
	Energy Requirement	
	Physiological Significance of Protein Synthesis	
	Physiological Functions of Proteins	261
	Regulation of Protein Concentrations	
	Replacement of Cells	
	Wound Healing	
	Immune Responses	
	Measurements of Protein Synthesis	
	Measurement of Protein Synthesis In Vitro	
	General Considerations	
	In Vitro Preparations	
	Choosing a Labeled AA Tracer	265
	Measuring Rate of Protein Synthesis	
	Measurement of Protein Synthesis In Vivo	
	General Considerations	
	General Terminologies Used in Measuring Protein	
	Turnover In Vivo	267
	Pulse Labeling of Proteins by Single Administration of a	
	Labeled AA	269
	Flooding Dose Technique	
	Continuous Infusion of a Tracer AA	
	Leucine Oxidation Method	
	Summary	
	References	
	References	211
Chapter 9	Intracellular Protein Degradation	279
	Historical Perspectives of Intracellular Protein Degradation	
	Proteases (Peptidases) for Intracellular Protein Degradation	
	Classification by Reaction Type	
	Exopeptidases	
	Endopeptidases	
	Classification by Catalytic Site	
	Classification by Evolutionary Relationship	
	Intracellular Proteolytic Pathways	
	Lysosomal Proteolytic Pathway	
	Nonlysosomal Proteolytic Pathway	286

	Ca <sup>2+</sup> -Dependent Proteolytic System (Calpain System)	
	Caspases	.286
	ATP-Dependent and Ubiquitin-Independent Proteolytic	
	System	.286
	ATP-Dependent and Ubiquitin-Dependent Proteolytic	
	System	.287
	Characteristics and Physiological Significance of Intracellular	
	Protein Degradation	. 291
	Biological Half-Lives of Proteins	
	ATP Requirement for Intracellular Protein Degradation	
	Physiological Significance of Intracellular Protein Degradation	
	Measurements of Intracellular Protein Degradation	
	Measurement of Intracellular Protein Degradation In Vitro	
	General Considerations.	
	Tracer and Nontracer Methods for Measuring Protein	
	Degradation In Vitro	.295
	Measurement of Intracellular Protein Degradation In Vivo	
	General Considerations.	
	Pulse Labeling of Proteins by Single Administration of a	
	Labeled AA	298
	Leucine Oxidation Method	
	Urinary Excretion of 3-Methylhistidine	
	Summary	
	References	
	references	. 501
CI	D 12 64 1 4 11 4 11 11	200
Chapter 10	Regulation of Amino Acid Metabolism	. 303
	Basic Concepts in Metabolism	.305
	Chemical Reactions	
	Laws of Thermodynamics as Applied to AA Metabolism	
	First Law of Thermodynamics	
	Second Law of Thermodynamics	
	Concept of Free Energy Unifies the First and Second	
	Laws of Thermodynamics	.307
	Concept of Equilibrium	
	Near-Equilibrium (Reversible) and Nonequilibrium	
	(Irreversible) Reactions	.309
	Enzymes as Biochemical Reactions	
	Enzymes as Biological Catalysts	
	Reversible Inhibition of Enzymes	
	Irreversible Inhibition of Enzymes	
	Intracellular Compartmentation of Metabolic Pathways	
	Metabolic Design Principles	
	Regulation of AA Metabolism	
	Allosteric Regulation	
	Reversible Phosphorylation and Dephosphorylation of Protein	
	relation in the priority and in the priority and the first in the state of the priority and the priority	