The Neuroscience of Clinical Psychiatry

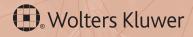
The Pathophysiology of Behavior and Mental Illness

Edmund S. Higgins, MD

Clinical Associate Professor of Psychiatry and Family Medicine Medical University of South Carolina Charleston, South Carolina

Mark S. George, MD

Distinguished Professor of Psychiatry, Radiology, and Neurosciences Layton McCurdy Endowed Chair Director, Brain Stimulation Laboratory (BSL) Editor-in-Chief, *Brain Stimulation* Medical University of South Carolina Charleston, South Carolina



Philadelphia • Baltimore • New York • London Buenos Aires • Hong Kong • Sydney • Tokyo Acquisitions Editor: Chris Teja Editorial Coordinator: Emily Buccieri Strategic Marketing Manager: Rachel Mante Leung Production Project Manager: Linda Van Pelt Design Coordinator: Elaine Kasmer Manufacturing Coordinator: Beth Welsh

Prepress Vendor: TNQ Books and Journals Third edition Copyright © 2019 Wolters Kluwer.

© 2013, 2007 by LIPPINCOTT WILLIAMS & WILKINS, a WOLTERS KLUWER business

All rights reserved. This book is protected by copyright. No part of this book may be reproduced or transmitted in any form or by any means, including as photocopies or scanned-in or other electronic copies, or utilized by any information storage and retrieval system without written permission from the copyright owner, except for brief quotations embodied in critical articles and reviews. Materials appearing in this book prepared by individuals as part of their official duties as U.S. government employees are not covered by the above-mentioned copyright. To request permission, please contact Wolters Kluwer at Two Commerce Square, 2001 Market Street, Philadelphia, PA 19103, via email at permissions@lww.com, or via our website at lww.com (products and services).

9 8 7 6 5 4 3 2 1

Printed in China

Library of Congress Cataloging-in-Publication Data

Names: Higgins, Edmund S., author. | George, Mark S. (Mark Stork), 1958-author.

Title: The neuroscience of clinical psychiatry : the pathophysiology of behavior and mental illness / Edmund S. Higgins, Mark S. George.

Description: Third edition. | Philadelphia : Wolters Kluwer, [2019] | Includes bibliographical references and index. Identifiers: LCCN 2017058724 | ISBN 9781496372000 (alk. paper)

Subjects: | MESH: Mental Disorders--physiopathology | Nervous System--physiopathology | Neuropsychology--methods Classification: LCC RC483 | NLM WM 140 | DDC 616.89--dc23 LC record available at https://lccn.loc.gov/2017058724

This work is provided "as is," and the publisher disclaims any and all warranties, express or implied, including any warranties as to accuracy, comprehensiveness, or currency of the content of this work.

This work is no substitute for individual patient assessment based upon healthcare professionals' examination of each patient and consideration of, among other things, age, weight, gender, current or prior medical conditions, medication history, laboratory data and other factors unique to the patient. The publisher does not provide medical advice or guidance and this work is merely a reference tool. Healthcare professionals, and not the publisher, are solely responsible for the use of this work including all medical judgments and for any resulting diagnosis and treatments.

Given continuous, rapid advances in medical science and health information, independent professional verification of medical diagnoses, indications, appropriate pharmaceutical selections and dosages, and treatment options should be made and healthcare professionals should consult a variety of sources. When prescribing medication, healthcare professionals are advised to consult the product information sheet (the manufacturer's package insert) accompanying each drug to verify, among other things, conditions of use, warnings and side effects and identify any changes in dosage schedule or contraindications, particularly if the medication to be administered is new, infrequently used or has a narrow therapeutic range. To the maximum extent permitted under applicable law, no responsibility is assumed by the publisher for any injury and/or damage to persons or property, as a matter of products liability, negligence law or otherwise, or from any reference to or use by any person of this work.

LWW.com.

To my two sons, who assisted with the artwork on Tuesday mornings at local coffee shops while waiting for school to start.

-ESH

To my many mentors—four formally designated in research fellowships, numerous others who've just helped and taught along the way—and the students and patients who have taught me so much as well. May we all be lifelong students!!

-MSG

Preface

Neuroscience is the basic science of psychiatry. Neuroscience describes the brain mechanisms that

- gather information from the external and internal world,
- analyze the information, and
- execute the best response.

Psychiatric disorders are the result of problems with these mechanisms.

The increased accessibility to the workings of the brain in the past 30 years has resulted in an explosion of information about neuroscience. Different lines of research such as brain imaging and animal studies along with more traditional postmortem analysis, study of medication effects, and genetic studies have transformed the way we conceptualize normal and abnormal behavior.

Bits and pieces of the neuroscience literature have filtered up to the practicing clinician, but a comprehensive understanding of the field is almost inaccessible to all but the most dedicated self-educators. The jargon is foreign and difficult to navigate. The standard textbooks are thick with contributions from multiple authors and almost impossible to read cover to cover. The relevance to the practice of psychiatry can sometimes be hard to appreciate.

We hope this book will provide a way for residents and practicing clinicians to gain a thorough appreciation for the mechanisms within the brain that are stimulating (or failing to stimulate) their patients. We also hope that the reader will have more accurate answers for the patient who asks, "What's causing my problem?" Likewise, we hope the reader will be better prepared for the increasingly difficult neuroscience questions that appear on board certification tests.

If we've learned anything from our studies on the brain, it is that LEARNING IS WORK! The brain increases its metabolism when conducting academic assignments. The process of focusing one's attention, understanding the concepts, and storing the new information requires energy. There is no passive learning.

Consequently, when learning is interesting and relevant it requires less energy. We have made every effort to make this material appealing and easy to consume. Pictures, drawings, and graphs have been liberally incorporated to allow the reader to learn the concepts quickly and efficiently. Every effort has been made to keep the material short and concise, but not too simple. Finally, we think information that is relevant to the reader is easier to retain, so we have tried to keep bringing the focus back to the practice of psychiatry.

We intend our book to be for three populations. First, it is for those in training: psychiatrists, psychologists, counselors, and allied physicians. Second, it is for psychiatric residents seeking to review the topics in preparation for their board examinations. And last, it is for the practicing clinician who was trained before the revolution in neuroscience and who would like to become more up-to-date and familiar with the field.

We hope that the reader will have a thorough—soup to nuts—understanding of the important topics in neuroscience and will henceforth be able to read and comprehend the future research in this field.

Edmund S. Higgins, MD Mark S. George, MD

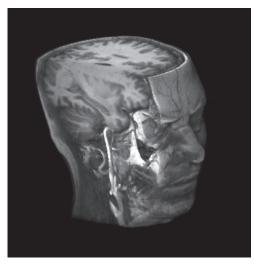
Acknowledgments

The authors wish to thank the following people for their assistance with this manuscript: Sherri A. Brown for her assistance with the artwork; Pamela J. Wright-Etter, MD, and Robert J. Malcolm Jr, MD, who reviewed individual chapters; and Laura G. Hancock, DO, and L. William Mulbry, MD, residents who reviewed the entire book. We also wish to thank all those readers who pointed out the typos in the first edition.

Figures 3.5–3.7 and 22.2 and the dolphin in Chapter 15, Sleep, were drawn by Fess Higgins.

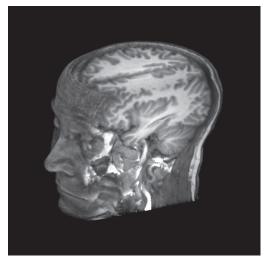
For the third edition we would like to thank Eric Brueckner, DO, and Edward M. Kantor, MD, for facilitating a review of the book by the MUSC fourth year residents—and all the helpful feedback they offered.

About the Authors



Edmund S. Higgins, MD

Edmund S. Higgins, MD, is a Clinical Associate Professor of Psychiatry and Family Medicine at the Medical University of South Carolina (MUSC). He received his medical degree from Case Western Reserve University School of Medicine. He completed residencies in family practice and psychiatry at MUSC. He is currently providing psychiatric care for South Carolina Department of Corrections and has a tiny private psychiatric practice. He lives on Sullivan's Island, SC.



Mark S. George, MD

Mark S. George, MD, is a Distinguished Professor of Psychiatry, Radiology, and Neuroscience as well as Director of the Brain Stimulation Division, Psychiatry at the Medical University of South Carolina (MUSC), Charleston. He also holds the Layton McCurdy Endowed Chair in Psychiatry and was the original director of the Center for Advanced Imaging Research at MUSC. He received his medical degree and completed dual residencies at MUSC in both neurology and psychiatry and is board certified in both areas. After a fellowship in London and 4 years at the National Institute of Mental Health, he returned to Charleston where he has conducted pioneering work with functional imaging of the brain, transcranial magnetic stimulation and vagus nerve stimulation, and other forms of brain stimulation. He is on several editorial review boards, has published over 500 scientific articles or book chapters, and has written or edited 6 books. He has been the editor-in-chief of Brain Stimulation for the past decade since it began, a journal published by Elsevier. He also resides on Sullivan's Island, SC.

Contents

SECTION I The Neuroscience Model

1	Introduction	2
2	Neuroanatomy	15
3	Cells and Circuits	29
4	Neurotransmitters	41
5	Receptors and Signaling the Nucleus	53
6	Genetics and Epigenetics	64

SECTION II Modulators

7	Hormones and the Brain	79
8	Plasticity and Adult Development	92
9	Immunity and Inflammation	108
10	The Electrical Brain	122

SECTION III Behaviors

11	Pain	133
12	Pleasure	148
13	Appetite	164
14	Anger and Aggression	177
15	Sleep	191
16	Sex and the Brain	207
17	Social Attachment	222
18	Memory	237
19	Intelligence	249
20	Attention	261

x Contents

SECTION IV Disorders

21	Depression	274
22	Anxiety	288
23	Schizophrenia	302
24	Alzheimer's Disease	316
Bibliography		328
Answers to End-of-Chapter Questions		346
Index		348