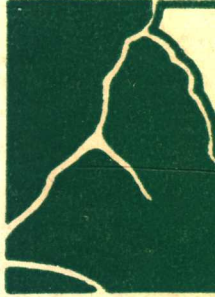
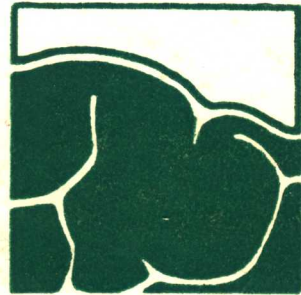


ANTIEPILEPTIC DRUGS

Edited by:

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Preface

This volume is the result of a collaborative effort exerted for over two years by a large number of investigators who are interested in epilepsy and the antiepileptic drugs. The aims of this endeavor originated from deliberations of the Public Health Service Advisory Committee on the Epilepsies and the NINDS Ad Hoc Committee on Anticonvulsant Drugs. The goal was simply to create an authoritative reference work that would present in a single source all of the recent advances in knowledge concerning the antiepileptic drugs as well as an in-depth review of basic pharmacologic data from both animals and man. In addition to presenting a definitive statement of the basic pharmacologic and clinical principles of the drugs used to treat epilepsy, it was hoped that this effort would serve to stimulate further research in the field. Although the book is not intended as a manual on how to treat epilepsy, it should provide all of the currently available pharmacologic knowledge, except mechanisms of action, required by clinicians for a thorough understanding and use of the antiepileptic drugs.

New studies on both old and new drugs were already underway in several laboratories and clinics when a workshop composed of clinical and basic research investigators from diverse disciplines related to epilepsy and the pharmacology of antiepileptic drugs was held in Warrenton, Virginia, in June 1970 to consummate the plans for a work that would meet the objectives of this undertaking. It was the consensus of the participants that (1) the volume should include data from animal experimentation for support and understanding of that obtained from man, though the contributors should clearly differentiate the findings because of species differences; (2) the practical limitations of space and publication time demanded that the mechanisms of action of the antiepileptic drugs be omitted, since considerable discussion of the neuropharmacologic and neurochemical mechanisms of epilepsy would be required to thoroughly cover this important subject; (3) continuation chapters on certain topics would be necessary to present the latest unpublished findings by other investigators and to provide adequately for different interpretations of the data; (4) the book should be structured according to the classes of drugs and arranged to present the basic pharmacologic and clinical principles of each drug within the classes; and (5) the volume should contain an introductory section concerning general principles.

Contributors were selected from the participants and other investigators suggested at the workshop. It was recommended that another meeting be organized to provide an opportunity for the contributors to profit from the critical discussion of their colleagues working in the field and to learn of pertinent questions from members of the scientific audience that might otherwise go unanswered in their final contributions.

At the second meeting in Scottsdale, Arizona, September 1971, the highlights of the proposed chapters were presented and discussed, and the preliminary manuscripts were reviewed by the Editorial Board. The contributors then submitted their revised manuscripts for final approval of the Editorial Board.

This volume, then, is the result of the enthusiastic cooperation and endeavors of such multidisciplinary investigators as pharmacologists, neurologists, toxi-

cologists, pediatricians, specialists in internal medicine, and biostatisticians. For the most part, therefore, any deficiencies in the scope of the material should be credited to the editors. The material included, however, represents the views and opinions of the individual authors and does not necessarily reflect those of the Editorial Board and the National Institutes of Health. Because of the many priorities required to maintain the rapid publication schedule, it was not possible to include an in-depth discussion of all the published information on a number of the subjects. Instead, an attempt has been made to provide the pertinent references. More than half the pages of this book represent information that has not been available previously. Setting the new information in context with previously known data provides a current review of knowledge on the subject.

It has become apparent from the contributors and their colleagues that new investigative efforts are now underway. This volume should serve as the initial milepost in the more rapid development of knowledge of the biotransformation and interaction of the antiepileptic drugs and their metabolites, which has been slow in evolving. Furthermore, the new information concerning the relationship of plasma concentration to clinical control and toxicity should not only stimulate further investigation on the efficacy and toxicity of these drugs, but also should provide immediately for improved utilization of the existing drugs. Although the development of new antiepileptic drugs is a slow, tedious, and expensive process, they will be used more effectively when they appear.

If the authors and editors of this work have adequately accomplished their task, the remaining responsibility for the fulfillment of the goals set forth for this endeavor rests with the publisher, at a time when the careful production, rapid publication, and adequate distribution of such a multiauthored volume are more difficult than ever. For this accomplishment, we are indebted to Dr. Alan M. Edelson of Raven Press and his capable managing editor, Virginia B. Martin. Finally, we acknowledge the invaluable assistance of Bettie Jean Hessie, Teresina M. Williams, and Lawrence D. Smith throughout this project and the support of the National Institute of Neurological Diseases and Stroke, which made this publication possible.

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