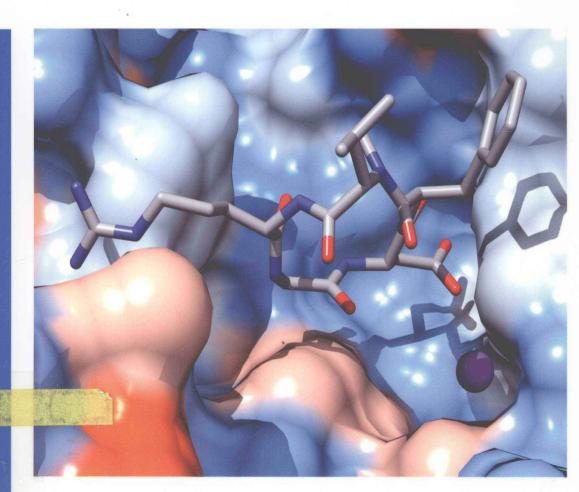
# Peptides: Chemistry and Biology

Second, Revised and Updated Edition



## Norbert Sewald and Hans-Dieter Jakubke

## Peptides: Chemistry and Biology

Second, Revised and Updated Edition





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#### Preface to the second edition

Peptides are continuously gaining increasing attention for application as lead compounds for the development of drugs, as drug molecules, as molecular tools for diagnostic purposes in biochemistry, medicinal sciences as well as proteome research. Since the first edition of this book has been published, a storming development in the peptide field took place. While formerly the development of synthetic methodology prevailed, currently peptide application in a biochemical, medicinal or analytical context predominates. Many different peptide conjugates have been tailored according to specific scientific questions.

The second thoroughly revised edition of this book especially takes into account these recent developments. Many sections have been updated carefully and some of them have been re-organized and extended significantly. This is especially true for Chapter 3, dealing with the biology of peptides, Chapter 5 that focuses on synthesis concepts for peptides and proteins, as well as Chapter 6 on special peptides and peptide conjugates. The recent developments of peptide drugs required an in-depth revision of Chapter 9 which is now an up to date reference on currently available peptide and protein therapeutics. The emerging role of peptides in an analytical context called for the addition of a new Chapter 10 providing details on the role of peptides on proteomics.

The second edition of our book also very much relies on advice, help and contributions by several colleagues. While it would lead too far to mention all of them, the most significant contributions have to be acknowledged in this preface. Chapter 2 was critically revised by Hans-Jörg Hofmann, University of Leipzig. The Section 6.2 was re-written by Luis Moroder, Max-Planck-Institute of Biochemistry, Martinsried. Dirk Ullmann and Volker Schellenberger usefully contributed to updating Chapter 9 on the application of peptides and proteins. Dr. Katherina Sewald prepared excellent graphical material and molecular formulae. Jens Conradi designed the new cover picture. Dr. Frank Weinreich, Lesley Belfit, Maike Peterson, Dr. Rosemary Whitelock, and Claudia Zschernitz, who were responsible for editing and production at Wiley-VCH, are acknowledged for carefully dealing with the manuscript and converting it into a book of high quality.

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We very much hope that this second edition will meet the same positive response and enthusiastic comments by our colleagues as the first edition did.

Finally we would like to dedicate this book on Chemistry and Biology of Peptides to the memory of two pioneers in the field who favorably commented on the first edition, but passed away in the meantime: Robert B. Merrifield and Murray Goodman.

Bielefeld and Dresden-Langebrück August 2008

Norbert Sewald

Hans-Dieter Jakubke

#### Preface to the first edition

The past decades have witnessed an enormous development in peptide chemistry with regard not only to the isolation, synthesis, structure identification, and elucidation of the mode of action of peptides, but also to their application as tools within the life sciences. Peptides have proved to be of interest not only in biochemistry, but also in chemistry, biology, pharmacology, medicinal chemistry, biotechnology, and gene technology.

These important natural products span a broad range with respect to their complexity. As the different amino acids are connected via peptide bonds to produce a peptide or a protein then many different sequences are possible - depending on the number of different building blocks and on the length of the peptide. As all peptides display a high degree of conformational diversity, it follows that many diverse and highly specific structures can be observed.

Whilst many previously published monographs have dealt exclusively with the synthetic aspects of peptide chemistry, this new book also covers its biological aspects, as well as related areas of peptidomimetics and combinatorial chemistry. The book is based on a monograph which was produced in the German language by Hans-Dieter Jakubke: Peptide, Chemie und Biologie (Spektrum Akademischer Verlag, Heidelberg, Berlin, Oxford), and first published in 1996. In this new publication, much of the material has been completely reorganized and many very recently investigated aspects and topics have been added. We have made every effort to produce a practically new book, in a modern format, in order to provide the reader with profound and detailed knowledge of this field of research. The glossary, which takes the form of a concise encyclopedia, contains data on more than 500 physiologically active peptides and proteins, and comprises about 20% of the book's content.

Our book covers many different issues of peptide chemistry and biology, and is devoted to those students and scientists from many different disciplines who might seek quick reference to an essential point. In this way it provides the reader with concise, up-to-date information, as well as including many new references for those who wish to obtain a deeper insight into any particular issue. In this book, the "virtual barrier" between peptides and proteins has been eliminated because, from

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the viewpoint of the synthesis or biological function of these compounds, such a barrier does not exist.

This monograph represents a personal view of the authors on peptide chemistry and biology. We are aware however that, despite all our efforts, it is impossible to include all aspects of peptide research in one volume. We are not under the illusion that the text, although carefully prepared, is completely free of errors. Indeed, some colleagues and readers might feel that the choice of priorities, the treatment of different aspects of peptide research, or the depth of presentation may not always be as expected. In any case, comments, criticisms and suggestions are appreciated and highly welcome for further editions.

Several people have contributed considerably to the manuscript. All the graphical material was prepared by Dr. Katherina Stembera, who also typed large sections of the manuscript, provided valuable comments, and carried out all the formatting. We appreciate the kindness of Professor Robert Bruce Merrifield, Dr. Bernhard Streb and Dr. Rainer Obermeier for providing photographic material for our book. Margot Müller and Helga Niermann typed parts of the text. Dr. Frank Schumann and Dr. Jörg Schröder contributed Figures 2.19 and 2.25, respectively. We also thank Dirk Bächle, Kai Jenssen, Micha Jost, Dr. Jörg Schröder, and Ulf Strijowski for comments and proofreading parts of the manuscript.

Dr. Gudrun Walter, Maike Petersen, Dr. Bill Down, and Hans-Jörg Maier took care that the manuscript was converted into this book in a rather short period of time, without complications.

Bielefeld and Dresden-Langebrück April 2002

Norbert Sewald

Hans-Dieter Jakubke

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