PROGRESS IN BIOORGANIC CHEMISTRY AND MOLECULAR BIOLOGY

Editor Yu. A. Ovchinnikov

PROGRESS IN BIOORGANIC CHEMISTRY AND MOLECULAR BIOLOGY

Proceedings of the International Symposium on Frontiers in Bioorganic Chemistry and Molecular Biology held in Moscow and Alma-Ata, USSR, on 19-24 June, 1984. Sponsored by the Federation of European Biochemical Societies, the International Union of Biochemistry and the International Council of Scientific Unions

Editor

Yu. A. Ovchinnikov

1984

ELSEVIER SCIENCE PUBLISHERS Amsterdam · New York · Oxford



ICSU PRESS Paris All rights reserved. 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 written permission of the publisher, Elsevier Science Publishers B.V., Biomedical Division, P.O. Box 1527, 1000 BM Amsterdam, The Netherlands. Special regulations for readers in the USA - This publication has been registered with the Copyright Clearance Center Inc. (CCC), Salem, Massachusetts. Information can be obtained from the CCC about conditions under which photocopies of parts of this publication may be made in the USA. All other copyright questions, including photocopying outside the USA, should be referred to the copyright owner, Elsevier Science Publishers B.V., unless otherwise specified.

ISBN 0 444 80643 1

Published by: Elsevier Science Publishers B.V. P.O. Box 211 1000 AE Amsterdam The Netherlands

Sole distributors for the USA and Canada: Elsevier Science Publishing Company Inc. 52 Vanderbilt Avenue New York, NY 10017 USA

Library of Congress Cataloging in Publication Data

International Symposium on Frontiers in Bioorganic Chemistry and Molecular Biology (1984: Moscow, R.S.F.S.R., and Alma-Ata, Kazakh S.S.R.) Progress in bioorganic chemistry and molecular biology.

"Sponsored by the Federation of European Biochemical Societies, the International Union of Biochemistry, and the International Council of Scientific Unions."

Includes indexes.

1. Bioorganic chemistry--Congresses. 2. Molecular biology--Congresses. I. Ovchinnikov, \widehat{IU} . A. (\widehat{IU} rii Anatolevich), 1934- . II. Federation of European Biochemical Societies. III. International Union of Eiochemistry. IV. International Council of Scientific Unions. V. Title. QP550.I58 1984 574.19'2 84-25859 ISBN 0-444-80643-1 (U.S.)

PREFACE

There is nothing new in the statement that the scientist nowadays has to spend more and more of time and effort simply trying to follow what is being done by his colleagues all over the world. Daily library work is essential for this, but there is one more effective (and pleasant) way to be constantly in touch with the latest advances in your field - that is to meet your colleagues during scientific meetings - to exchange information, to establish new contacts. The number of such conferences, national and international, is growing every year. This book deals with the one of such meetings, which is memorable in many ways.

The International Symposium "Frontiers in Bioorganic Chemistry and Molecular Biology" was held in Moscow and Alma-Ata, June 19-24, 1984 as a satellite conference of the 16th FEBS Meeting. It was also co-sponsored by the International Council of Scientific Unions and the International Union of Biochemistry. The Symposium which in fact became a "summit" meeting of world top scientists in the rapidly developing field of physicochemical biology, was also meant to commemorate the 25th anniversary of the Shemyakin Institute of Bioorganic Chemistry of the USSR Academy of Sciences.

The Symposium programme covered a wide range of aspects of modern physicochemical biology and biotechnology: investigations into structure-functional relations in peptides and proteins, nucleic acids, carbohydrates, lipids, and other bioregulators, elucidation of the structure, synthesis, study of conformational peculiarities, mechanism of biological action, membrane and receptor systems responsible for transfer of genetic information, regulation, as well as the most challenging problems of immunology, neurobiology, oncology, endocrinology, in order to solve them novel approaches and methods of chemistry and physics are employed.

Naturally, fundamental researches in these areas have become vitally important for modern medicine, agriculture, and industry. The advances of genetic and cell engineering, chemical and enzymatic synthesis, hybridom analysis, that is of biotechnology, are widely applied to facilitate their development.

The plobal geography of the countries represented, the variety of the areas discussed, together with an exceedingly high level of presentation, make this book particularly valuable.

I believe, the atmosphere of the Symposium promoted mutual understanding, strengthening of friendly ties and collaboration among scientists throughout the world. The Organizing Committee of the Symposium "Frontiers in Bioorganic Chemistry and Molecular Biology" expresses deep gratitude to all participants for their activity and interesting presentations.

Professor Yu.A. Ovchinnikov

LIST OF PARTICIPANTS

N.G.	Abdulaev	USSR	Yu.A.	Chizmadzhev	USSR
G.I.	Abelev	USSR	A.	Caldas	Spain
٧.Ι.	Agol	USSR	M.	Cortijo	Spain
Yu.B	.Alakhov	USSR	T.B.	Darkanbaev	USSR
A.D.	Alshtein	USSR	S.P.	Datta	Great Britain
S.A.	Andronati	USSR	٧.G.	Debabov	USSR
L.N.	Andreev	USSR	V.I.	Deigin	USSR
R.	Arnon	Israel	٧.٧.	Demin	USSR
I.G.	Atabekov	USSR	Р.	Desnuelle	France
M.A.	Aytkhozhin	USSR	B.A.	Dmitriev	USSR
Α.	Azzi	Switzerland	V.N.	Dobrynin	USSR
Ph.C	.Babychev	USSR	N.P.	Dubinin	USSR
B.K.	Bachhawat	India	K.M.	Dyumaev	USSR
A.A.	Baev	USSR	J.P.	Ebel	France
S.B.	Balmukhanov	USSR	٧.٨.	Efimov	USSR
E.A.	Barnard	Great Britain	G.B.	Elyakov	USSR
L.I.	Barsukov	USSR	N.M.	Emanuel	USSR
S.V.	Belyaev	USSR	٧.٨.	Engelgardt	USSR
N.D.	Beklemyshev	USSR	L.	Ernster	Sweden
L.D.	Bergelson	USSR	Α.	Eschenmoser	Switzerland
S.G.	Van den-Bergh	The Netherlands	R.P.	Evstigneeva	USSR
A.J.	Birch	Australia	P.	Fasella	Belgium
Ε.	Blout	USA	N.	Fedoroff	USA
A.A.	Bogdanov	USSR	L.D.	Frye	Japan
P.D.	Boyer	USA	S.	Fukui	Japan
A.E.	Braunstein	USSR	I.M.	Gelfand	USSR
1.3	Budowsky	USSR	G.P.	Georgiev	USSR
E.B.	Burlakova	USSR	I.I.	Gitelzon	USSR
R.G.	Butenko	USSR	٧.١.	Goldansky	USSR
V.F.	Bystrov	USSR	S.I.	Gorodetsky	USSR
E.	Carafoli	Switzerland	Ε.	Gre11	FRG
R.	Castro de la Mata	Peru	E.Ya.	Gren	USSR
D.	****	Great Britain	N.	Grubhofer	FRG
	Chapman Chazov	USSR	M.	Grunberg-	France
	Chertov	USSR	M.V.	Manago Gusev	USSR
	Chizhov	USSR	M.S.	Gylyarov	USSR
J.J.	CHIZHOV	0337			

E.V.	Grishin	USSR	A.A.	Lev	USSR
B.P.	Gottikh	USSR	С.	Liebecq	Belgium
	Gvozdev	USSR	С.	Lin	China .
P.J.	Henderson	Great Britain	A.W.	Linnane	Australia
В.	Hess	FRG	V.M.	Lipkin	USSR
R.D.	Henriques	Cuba	٧.Κ.	Lishko	USSR
R.L.	Hill	USA	W.R.	Loewenstein	USA
5.	Ho dgkin	Great Britain	E.M.	Lukanidin	USSR
۶.	Hucho	W.Berlin	L.G.	Magazanick	USSR
A.N.	llyaletdinov	USSR ·	W.	Manz	Great Britain
G.R.	Ivanitsky	USSR	L.N.	Markovsky	USSR
S.G.	Inge-Vechtomov	USSR	I.V.	Martynov	USSR
0.	Jardetzky	USA	J.	Mathieu	France
J.	Jaz	France	G.Kh	.Matsuka	USSR
Р.	Jorgensen	Denmark	F.	Mayor	Spain
٧.٨.	Kabanov	USSR	N.N.	Mełnykov	USSR
Υ.	Kagawa	Japan	B.R.	Merrifield	USA
Yu.T	.Kalinin	USSR	Н.	Metzner	FRG
D.	Kanazir	Yugoslavia	Ρ,	Minakakis	Greece
K.P.	Kashkin	USSR	A.D.	Mirzabekov	USSR
E.	Katchalski-Katzir	Israel	Н,	Mislin	Switzerland
A.S.	Khokhlov	USSR	E.N.	Mishustin	USSR
A.Ya	.Khorlin	USSR	Yu.	Miyazaki	Japan
М	Kikutani	Japan	N.N.	Modyanov	USSR
L.L.	Kiselev	USSR		Monastyrskaya	USSR
L.K.	Klyshev	USSR		Muranov	USSR
D.G.	Knorre 、	USSR	T.A.	Muranova	USSR
N.K.	Kothetkov	USSR	T.	Nagatsu	Japan
M.N.	Kolasov	USSR	S.M.	Navashin	USSR
J.K.	Kostrzewski	Poland	N.K.	Nadyrov	USSR
P.G.	Kostyuk	USSR		Nephedov	USSR
A.	Kotyk	CSSR	V.A.	Nesmeyanov	USSR
A.A.	Krasnovsky	USSR		Ogarkov	USSR
E.M.	Kreps	USSR		Oesterhelt	FRG
٧.G.	Korobko	USSR		Okudzhava	USSR
V.P.	Kukhar	USSR		Ostrovsky	USSR
A.M.	Kunaev	USSR		Ovchinnikov	USSR
M.	Lazdunski	France		Ovchinnikov	USSR
Yu.S.	Lazurkin	USSR	_	Ozawa	Japan
Ε.	Lederer	France			Brazil
					U. U. C. I I

_		·. ·			
F.	Palmieri	Italy	M.Kh	.Shagaeva	USSR
	.Pankov	USSR	G.K.	Skryabin	USSR
S.	Papa	I taly	K.G.	Skryabin	USSR
G.	Pasternak	GDR	٧.Р.	Skulachev	USSR
L.	Pauling	USA	V.N.	Smirnov	USSR
	Petrov	USSR	Ε.	Smith	USA
	Phokin	USSR	G.	Snatzke	FRG
Th.	Photakis	Greece	٧.٤.	Sokolov	USSR
R.J.	Planta	The Netherlands	A.A.	Sozinov	USSR
٧.	Prelog	Switzerland	A.S.	Spirin	USSR
M.A.	Prokophyev	USSR	S.	Stendah!	Sweden
A.	Pullman	France	F.B.	Straub	Hungary
В.	Pullman	France	E.D.	Sverdlov	USSR
Ε.	Quagliariello	Italy	K.M.	Sytnik	USSR
K.	Rajewsky	FRG	J.	Szentagothai	Hungary
S.M.	Rapoport	GDR	R.	Thunberg	Sweden
0.A.	Reutov	USSR	J.	Tigyi	Hungary
Α.	Rich	USA	Α.	Toepfer	FRG
J.	Řiman	ČSSR	J.	Tooze	FRG
A.B.	Rubin	USSR	I.V.	Torgov	USSR
R.S.	Rychkov	USSR	R.	Tsanev	Bulgaria
A.S.	Sadykov	USSR	V.I.	Tsetlin	USSR
R.I.	Salganik	USSR		Turpaev	USSR
R.K.	Salyaev	USSR		Ugolev	USSR
L.S.	Sandakhchiev	USSR	D.	Vazquez	Spain
Α.	Sato Sato	Peru		Vainstein	USSR
J.C.	Schell	FRG		.Vasiliev	USSR
Κ.	Scherrer	France		Vasilov	USSR
Ε.	Schram	Belgium	M.	Villavicencio	Peru
Κ.	Šebes ta	ČSSR	•••	Nunez	
М.	Sela	Israel	Ε.	Vogel	FRG
G.	Semenza	Switzerland	٧.L.	Voeykov	USSR
	Serebryakov	USSR	M.V.	Volkenstein	USSR
	Severin	USSR	Υ.	Wang	China
	Severin	USSR	J.E.	Walker	Great Britain
	Sharma	India	J.H.	Weil	France
	Shabarova	USSR	Th.	Wieland	FRG
	Shemyakin	USSR	М.	Wiewiorowski	Poland
	Shibaev	USSR	R.J.	Williams	Great Britain
-	Shirokov	USSR		Wittmann	W.Berlin
J. U.	31110804	033K			

В.	Wittmann-	W.Berlin	M. Yomtov	Bulgaria
	Liebold		H.G. Zachau	FRG
_ (,	Woodward	USA	H. Zahn	FRG
K.	Wuthrich	Switzerland	S.Z. Zairov	USSR
ν	Vaci	12.02.0	S.Z. Zairov	033K

CONTENTS

Preface	, ,
List of participants	vii
PEPTIDES AND PROTEINS. SPATIAL ORGANIZATION AND FUNCTION	
Peptide and protein research at the Shemyakin Institute Yu.A. Ovchinnikov	3
Understanding the surfaces of proteins R.J.P. Williams, G.R. Moore and G. Williams	31
NMR analysis of large peptide and small protein conformation in solution V.F. Bystrov	41
A method for the definition of the solution structure of proteins from NMR and other physical measurements: The <i>lac</i> -repressor headpiece 0. Jardetzky	55
Solution conformation of E. coli lac-repressor DNA binding domain by 2D NMR: Sequence location and spatial arrangement of three $\alpha\text{-helices}$ E.R.P. Zuiderweg, M. Billeter, R. Kaptein, R. Boelens, R.M. Scheek and K. Wüthrich	65
Compartmentation of proteins of the translation machinery on eukaryotic polyribosomes A.S. Spirin and L.P. Ovchinnikov	71
Crystallization of membrane proteins T. Ozawa	83
Labeling of hydrophobic segments of intrinsic membrane proteins by means of a photogenerated carbene J. Brunner and G. Semenza	91
Solid phase peptide synthesis: The role of sequence and conformation in the action of the cecropins R.B. Merrifield, D. Andreu, H.G. Boman and H. Steiner	101
Thermodynamic characterization of the conformational transition induced by the allosteric activator in glycogen phosphorylase P.L. Mateo, C. Baron, J.S. Jimenez and M. Cortijo	109
Three-dimensional organization of catalases B.K. Vainshtein, W.R. Melik-Adamyan, V.V. Barynin and A.A. Vagin	117
Importance of substrate conformation for renin activity M.C.F. Oliveira, C.R. Nakaie, J.L. Pesquero and A.C.M. Paiva	127
Receptor-mediated mutual requlation of tyrosine-hydroxylase activity and the biopterin cofactor level T. Nagatsu, Y. Hirata and T. Yamaguchi	133
Carbohydrate binding proteins (lectins) in rat macrophages M.A. Lehrman, R.S. Haltiwanger, A.E. Eckhardt and R.L.Hill	141

The specificity of neurons and neuron connexions i exact localization of mediators and brain pept J. Szentágothai	n the light of more iides 151
Prospects for therapeutic application of peptides E.I. Chazov	157
BIOENERGETICS, BIOMEMBRANES AND TRANSPORT	
Dynamics as basic attributes of living states B. Hess, M. Markus and D. Kuschmitz	165
Active site in proton translocating ATPase Y. Kagawa, M. Yoshida, H. Kihara and V.R. Rama	ukrishnan 175
Respiration and energy conversion in biomembranes S. Papa	183
Energy-linked changes of enzyme conformation in AT bioenergetic processes P.D. Boyer	P formation and other
Cytochrome c oxidase: Molecular aspects of a proto A. Azzi, M. Muller, G. Petrone, P. O'Shea and	
On sodium energetics V.P. Skulachev	207
Structure, mechanism and differentiation of voltage channels which generate action potentials M. Lazdunski, J. Barhanin, M. Borsotto, M. Fos G. Romey and H. Schmid	
Theoretical studies on ion-channel interactions: G A. Pullman ,	Gramicidin A 219
Ionic channels - New problems P.G. Kostyuk	225
Membrane transport in the messenger function of ca E. Carafoli	lcium 233
Specificity and dynamic aspects of cation transpor E. Grell, G. Krause, E. Lewitzki, G. Mager and	t systems in membranes I H. Ruf 239
Intracellular pH distribution and driving forces f transport in yeast cells A. Kotyk and J. Slavík	for secondary active 247
Sugar-proton transport proteins of <i>Escherichia col</i> P.J.F. Henderson, E.O. Davis, H. Hirata, Y. Ka Jones-Mortimer and A.J.S. Macpherson	<i>i</i> gawa, M.C. 255
The active phosphate carrier from heart mitochondr Single band of the hydroxylapatite eluate F. Palmieri, F. Bisaccia, G. Prezioso, A. Rizz	
The GABA receptor complex: Purification, analysis of the proteins and their production from mess E.A. Barnard, G. Bilbe, C. Mamalaki, F. Sigel	and reconstitution senger RNA and F.A. Stephenson 275

	XIII
Transport mechanisms of neurotransmitter amino acids and of neuro- transmitter precursors in the CNS F. Mayor, F. Valdivieso, C. Gimenez, M.C. Aragon, F. Mayor Jr, J.G. Marvizon, J. Diez-Guerra, E. Herrero and F. Zafra	287
Transport of nuclear coded proteins into mitochondria E. Marra, M. Greco, M.R. Montemurro, E. Perlino, S. Passarella, S. Doonan, C. Saccone and E. Quagliariello	295
Biochemistry of oxygen toxicity L. Ernster	303
Teleonomic regulated oxidative damage - The role of lipoxygenase in the maturation of reticulocytes S. Rapoport	311
Activity control of the hepatic endoplasmic cytochrome P-450 dependent monooxygenase system W. Scheler, H. Rein, J. Blanck and K. Ruckpaul	319
Pyruvate degradation in archaebacteria: Radicals, ferredoxins and evolutionary aspects D. Oesterhelt, L. Kerscher, S. Mathews and F. Lottspeich	327
Characterization of photopolymerized, diacetylenic phospholipids in liposomes and reconstituted systems J.A. Hayward, F. Castelli, M.A. Whittam, D.S. Johnston and D. Chapman	335
STRUCTURE AND FUNCTION OF NUCLEIC ACIDS	
Affinity chemical modification of nucleic acids D. Knorre	345
Total synthesis of yeast alanine transfer RNA Y.L. Wang	353
Nucleotide sequence of four chloroplast transfer RNA genes from <i>Vicia</i> faba JH. Weil, G. Bonnard, F. Michel and A. Steinmetz	363
Structure and evolution of small nuclear RNAs JP. Ebel, A. Krol, E. Myslinski, E. Lazar, H. Gallinaro, B. Haendler, M. Jacob and C. Branlant	371
Right-handed and left-handed double helical DNA A. Rich	381
Recognitory interaction of non-intercalating ligands with DNA B. Pullman	395
Comparative structural analysis of selected salts of putrescine, spermidine and spermine in the aspect of their stereochemical interactions with nucleic acids M.D. Bratek-Wiewiórowska, M. Jaskólski, M. Alejska, A. Perkowska and M. Wiewiórowski	403
Regulation of gene expression of <i>E. coli</i> phenylalanyl and threonyl tRNA synthetases M. Grunberg-Manago, M. Springer, J.A. Plumbridge, C. Sacerdot, M. Graffe, G. Fayat, P. Lestienne, JF. Mayaux and S. Blanquet	411

Immunoglobulin genes: Rearrangements, clustered point mutations, amplification, transpositions, and gene conversion H.G. Zachau, M. Pech, HG. Klobeck, HR. Jaenichen, P.S. Neuma HD. Pohlenz and B. Straubinger	ier, 423
MOLECULAR IMMUNOLOGY	
The immune system and its molecular engineering M. Sela	433
Lymphocyte reactivity in normal and malignant cell proliferation aga a phylogenetically conserved fetal antigen G. Pasternak, B. Schlott, G. Gryschek, S. Albrecht, J. Reinhöfer E. Matthes and B. von Broen	
Muramyl-peptides as immunomodulators, sleep factors and vitamins E. Lederer	447
Highly immunogenic artificial complexes based on synthetic poly-ions (artificial antigens and vaccines) R.V. Petrov	455
New trends in non-isotopic immunoassays S. Stendahl and E. Soini	463
LOW MOLECULAR WEIGHT BIOREGULATORS	
Inorganic 'enzymes': A new approach to organic synthesis A.J. Birch	471
Metabolism of flavins in mammals K. Yagi, N. Ohishi and H. Ohkawa	479
Structure-function relationship of vitamin B_{12} coenzyme (adenosylcobin the diol dehydrase system. Function of the adenine moiety of Coß (upper) ligand in the coenzyme activity S. Fukui and T. Toraya	alamin) the 487
A bio-organic walk through the world of mushrooms T. Wieland	495
Study of nerve conduction with spin-labelled anesthetics J. Tigyi, K. Hideg and T. Lakatos	501
Psychosocial (emotional) stress, steroid hormones and carcinogenesis Molecular aspects. Facts and speculations	
D.T. Kanazir, R. Djordjević-Marković and R. Grossarth-Maticek	509
Author index	521
Subject index	523

PEPTIDES AND PROTEINS SPATIAL ORGANIZATION AND FUNCTION



PEPTIDE AND PROTEIN RESEARCH AT THE SHEMYAKIN INSTITUTE

YURI A OVCHINNIKOV

Shemyakin Institute of Bioorganic Chemistry, USSR Academy of Sciences, ul. Vavilova 32, Moscow V-334 (USSR)

The 50s are marked by great advances of the world science in protein-peptide chemistry. Since 1949 the α -helix proposed by L.Pauling triumphed all over the world that opened wide the doors for penetrating into the structure of protein molecules.

V. du Vigneaud completed the chemical synthesis of the most important peptide hormone, oxytocin, later F.Sanger decipherd the amino acid sequence of the first protein, insulin. Undoubtedly studies of peptide-protein substances occupied a prominent place and captured scientists' attention. Just that time the USSR Academy of Sciences faced the necessity of organizing an Institute engaged in studies of life science, especially its chemical aspects and in 1959 the Institute for Chemistry of Natural Products was founded.

The impetuous progress in the protein and peptide chemistry made us change the traditional trends in the study of natural low molecular weight compounds of alicyclic or heterocyclic series: steroids, alkaloids, antibiotics of chloramphenical or tetracycline groups, etc., and side with those for whom the main notions were amino acid, peptide bond, α -helix, β -structure.

M.M.Shemyakin initiated the study of peptides of a new type and since the first years of the Institute's foundation systematic investigations of atypical naturally occurring peptides, depsipeptides, containing hydroxy and amino acid residues started. Antibiotics-depsipeptides, valinomycin and enniatins, were the first in this row. In 1955 H.Brockmann and G.Schmidt-Kastner isolated valinomycin from extracts of the strain *Streptomyces fulvissimus*. The structure (cyclo-octadepsipeptide) proposed by. H.Brockmann and H.Geeren was erroneous. As shown valinomycin is a cyclododecadepsipeptide, built of three identical tetradepsipeptide moieties. The structures for enniatin antibiotics isolated by P.Plattner in 1947^{4,5} from cultural liquids of *Fusarium* turned out to be also incorrect. The total synthesis showed that enniatins A and B were not cyclotetradepsipeptides, but cyclohexadepsipeptides^{6,7}. The Institute developed a novel synthesis of depsipeptides: natural cyclodepsipeptides, valinomycin (Fig. 1), enniatins A, B, C, beauvericin (Fig. 2), and their numerous analogues.

The structure-functional studies of cyclodepsipeptides permitted the use of a topochemical principle of the peptide molecule modification for the analysis of many biologically active peptides. Now active analogues are constructed by changing the molecule as a whole, but not local regions, to maintain stereoelectron

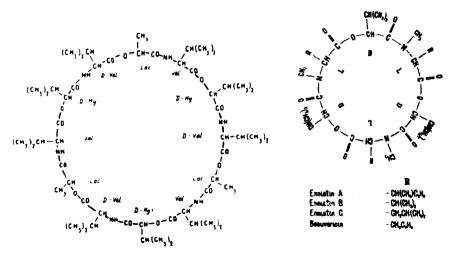


Fig. 1. Valinomycin.

Fig. 2. Enniatin ionophores.

characteristics of the peptide. Topochemical transformations include change in the direction of acylation of residues (retro-isomers), in configuration of asymmetric centers (enantio-isomers), make possible obtaining retroenantiomers, cyclization of linear peptides, and other transformations. The topochemical approach was successfully used for investigation of structure-functional dependence for membrane-active cyclopeptides, valinomycin, enniatins, gramicidin S, and antamanide $^{9-11}$.

The work in the laboratory of Prof. V.Prelog (Switzerland), a world-known stereochemist, stimulated this approach to the peptide system modification. First publications summarize data on simple cyclopeptides, built of glycine and alanine residues 12,13.

The proper studies of depsipeptides of valinomycin and enniatin groups resulted in their wide use when studying ion permeability of artificial and biological membranes. That became possible only when B.Pressman discovered the effect of valinomycin on mitochondria, American and Soviet investigators developed the concept of ionophore action of depsipeptides 15,16 . The elucidation of the mechanism of ${\mbox{K}}^{+}$ ion-binding to valinomycin, the bracelet structure for the valinomycin ${\mbox{K}}^{+}$ complex proposed in the USSR crowned the research 15 . Just this elegant molecule decorates the entrance to the Shemyakin Institute of Bioorganic Chemistry.

As established later 17 enniatins are also effective complexones of Na $^+$, K $^+$, Rb $^+$, and Cs $^+$ ions, correlation between their antimicrobial and complexing activities was discovered. Spectral methods were employed to determine enniatin B conformations in solutions in free state as well as in complexes with alkali metal ions. Enniatins A, B, C and beauvericin in solutions are characterized by equilibrium of two forms, polar (P) and nonpolar (N) 18 . Conformation of enniatin B complexes