#### UPDATE ON DERMATOLOGIC DRUG THERAPY

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### UPDATE ON DERMATOLOGIC DRUG THERAPY

Professor Jasna Lipozenčić and co-authors

### **DEDICATION**

We would like to dedicate this book to our patients who helped us to understand the meaning of dermatologic drug therapy, side effects on it and who have given us insights into how to design better research studies to deal with enormous knowledge for the dermatologic drug therapy.

To the memory of glorious 51-year history of Croatian Academy of Medical Sciences and those dermatologic forebears who were fascinated by drug allergy in the skin, who advanced knowledge of skin therapy and who cared for and about patients.

### **PREFACE**

We are honored and pleased with the idea that Croatian Academy of Medical Sciences is organizer of Congress and a book "Update in Dermatological Drug Therapy". The Contents presented in this book represent a scientific and professional challenge, dedicated to drug therapy.

Skin is the largest organ of our body, which with its appendages hair, nails, and glands has complex structure. The skin is permanently exposed to different environmental influence and is a mirror of internal diseases influence. According definition of the European Union of Medical Specialists (UEMS), Section of Dermatology and Venereology: "Dermatology is organ specialized that is responsible for the diagnosis and treatment (both medical and surgical) and prevention of diseases of the skin and subcutaneous tissue." Medicine is a science, which integrates the current scientific knowledge on health and diseases, in this book, caused by drugs. We believe that today, more than ever before in the history of medicine, it is essential to achieve a good balance between science and humanity not letting technology engulfs the doctor-patient relationship, which is really at the core of medical practice.

Besides dermatovenerologists with a main role, there are important internal medicine specialists, immunologists, infectologists, gynecologists included with their knowledge on recognizing of drug side-effects. About 30% of patients have dermatologic problems in every day routine of family physicians.

Everyone has during lifetime an problem with a skin. Each patient or client is an individual and has different response to therapy, that is why we wanted to give them a practical knowledge on update on dermatological drug therapy and individual approach is needed. The progress in treatment of skin diseases in past decade is impressive as well as more and more new diagnostic tests, new classification, and modern therapies are available. Besides different topical agents, there are available more and more physiological remedies.

Treatment of skin disease has changed remarkably during the last 10 years. Thus, novel targets have been identified and specific drugs developed which directly interfere with or alter the processes. Many of these new agents are administered systematically, either orally or by subcutaneous injection. And yet, as with all forms of drug therapy, these highly efficacious agents can be associated with severe side effects and drug-induced toxicity. To understand

the etiopathogenetic mechanisms of drug induced as well as adverse events why a certain disease occurs and how to find the best cure for it. With the rapid development of novel therapeutics, there has been a major evolution in the clinical practice of dermatovenerology. In European countries, patients with skin disease are often treated as in-patients by dermatologists whereas in United States, this occurs only rarely and the patients are treated under Internal Medicine and dermatologists are consulted on their management. Throughout medical history and diagnostic approach, authors are on viewing the disease and how to help to patients. However, in the last few decades we have witnessed to the development of ever more and more the adverse drug reactions. The purpose of human medicine is always and before all to help the patients. Today we can do much more in terms of prevention of drug-induced diseases. Not many doctors have large experience with a certain drug and a type of drug hypersensitivity reactions. As drug hypersensitivity characteristically appears unexpectedly, patient - oriented research is difficult. Thus, person oriented, individual approach is welcomed. The aim is to provide a general view on this topic so that can find relevant information practicing physician, dermatologists, the allergologists, immunologists, pharmacologists, pharmacists, immunotoxicologists and pharmaceutical company officer. It combines pharmacology with antigen specific immunology and a concept of T cells stimulated by drugs. There is involved interdisciplinary group of physicians, and scientists who work together to better understanding these disease and drug hypersensitivity reactions. In this book, 48 of colleagues from various professions from Croatia, Europe, America, and Australia have created together with members of Croatian Academy of Medical Sciences uniform book conceived to address dermatologic management as well as most drug-induced changes. Personally, we are proud to be organizers of this prestigious team which promote this book, contributing to a update of medicine in third millennium. There is assembled on outstanding set of papers on conceptual basis data based research, novel therapeutic procedures that are based on understanding the pathophysiology of skin diseases and drug induced diseases from various medical specialties and other relevant fields authored by experts from Croatia and worldwide. Experts have reviewed all manuscripts carefully. We are thankful to publisher ing. Cindro, LaserPlus d.o.o. of Zagreb, Croatia to organize this book. We are grateful to all of authors for their experience and wisdom the refocusing of dermatologic drug therapy in content.

The issue is the final product of the efforts and cooperation of an extraordinary team. We offer our sincere thanks to the contributors who have giving their valuable time to ensure the success of this book.

Jasna Lipozenčić and Željko Reiner

### **ACKNOWLEDGEMENT**

The idea for this book came from me, Jasna Lipozenčić as principle author. I am grateful to all contributors who made possible this book as prestige work of the Croatian Academy of Medicial Sciences.

Professors Dunja Beritić and Tatjana Jeren were engaged and read through draft after draft and page proof after galley.

Professor Josip Čulig has overseeing every detail of the work at it being printed. Academician Reiner made valuable additions to the Preface.

Professor Branka Marinović her Department of dermatology in Zagreb supplied valuable with technical help and photographs that appear in the book.

Mrs. Gordana Dučkić organizes the entire project in regard to all manuscripts and the more than hundred photograph that were screened and selected with finality. The all coworkers also participated in several of the intensive working processes during which the text was reviewed and finalizing.

Without this pleasant refuge for long productive hours, this book might never have never been written.

# **CONTENTS**

PREFACE  Jasna Lipozenčić, Željko Reiner
I. INTRODUCTION OF DERMATOTHERAPY
Basic Principles of Pharmacology –  Zdravko Lacković, Lidija Bach-Rojecky
2. Personalized medicine as it affects Dermatology –  Jasna Lipozenčić, Branka Marinović
3. Safety of Dermatologic Drug Therapy – Petra Turčić, Jasna Lipozenčić
4. Special Pharmacology – Pharmacoeconomics – Petra Turčić
5. Principles of Corneotherapy –  Jasna Lipozenčić, Zrinka Bukvić Mokos, Sanja Gregurić
6. Drugs and chemical compounds as initiators and promotors of skin tumors – Neira Puizina Ivić, Antoanela Čarija, Lina Mirić-Kovačević, Dubravka Vuković
7. The new Hedgehog Inhibitors for Advanced BCC – The ERIVANCE trial – Aleksandar Sekulić
8. Dermatologic Drug Therapy in Children – Nives Pustišek
9. Management of STIs and other Genitourethral Disorders – Michael Waugh
10. Dermatologic Drugs during Pregnancy and Lactation – Ksenija Makar Aušperger

11. SYSTEMIC AND TOPICAL DRUGS IN DERMATOTHERAPY
11. Recent Advances in Anti-infective Treatment in Dermatovenereology – Mihael Skerlev
12. Antihistamines in Dermatology –  Jasna Lipozenčić, Marcia Ramos-e-Silva.
13. Drugs used with UV or Visible Light –  Krešimir Kostović, Zrinjka Paštar
14. Laser Therapy and Drugs –  Zrinka Bukvić Mokos
15. Biological Therapeutics – Romana Čeović
16. Systemic Drugs in Bullous dermatoses – Branka Marinović
17. Antiandrogens and Androgen Inhibitors – Dinka Pavičić Baldani, Lana Škrgatić, Marina Šprem Goldštajn
18. Topical Corticosteroids – Different Strengths, Different Indications – Andrija Stanimirović, Maja Kovačević
19. The Current Role of Calcineurin Inhibitors – Slobodna Murat Sušić
20. Dermatologic Surgery –  Daška Štulhofer Buzina
21. Management of Epidermolysis Bullosa Acquisita – Bonnie Fergie, Lizbeth Ra Intong, Dedee F Murrell
22. An Update of the Use of Corticosteroids in Autoimmune Blistering Diseases – John W Frew, Dedee F Murrell
23. Injectable and Mucosal Routes of Drug Administration –  Marinka Mravak Stipetić
III. ADVERSE DRUG REACTIONS
24. Serious Adverse Effects from Systemic Drugs –  Ksenija Makar Aušperger
25. Complication of Dermatologic Therapies –  Ronni Wolf, Jasna Lipozencic, Bataya B. Davidovici

V.	PATHOMECHANISMS OF ADVERSE DRUG REACTIONS
	26. Drugs as Haptens, Antigens and Immunogens – Drago Batinić
	27. HIV Drug Therapy and Hypersensitivity Reactions – Šime Zekan, Josip Begovac
	28. Toxic Epidermal Necrolysis (TEN), Stevens-Johnson Syndrome and DRESS Syndrome –
	Ružica Jurakić Tončić, Jaka Radoš, Jasna Lipozenčić
V.	CLINICAL MANIFESTATIONS AS ADVERSE DRUG REACTIONS OF DERMATOLOGICAL THERAPIES
	29. Urticaria and Anaphylaxis due to Penicillins and Cephalosporins – Liborija Lugović Mihić
	30. Allergy to Pyrazolones, Quinolones and other Non-β-Lactam Antibiotics – Mateja Dolenc-Voljč
	31. Hypersensitivity Reactions to Iodinated Contrast Media – Suzana Ljubojević
	32. Maculopapular Drug Eruptions and Vasculitis allergica – Marinka Mravak Stipetić
	33. Drug Induced Pemphigus –  Branka Marinović
	34. Drug Induced Nephritis – Petar Kes, Nikolina Bašić-Jukić
	35. Drug Induced Aspirin and other NSAIDs reactions – Zrinjka Paštar, Gordana Štajminger, Jasna Lipozenčić
VI.	DIAGNOSIS OF DRUG HYPERSENSITIVITY
	36. Approach to the Patient with a Drug Hypersensitivity – Jasna Lipozenčić, Branka Marinović, Ružica Jurakić Tončić
	37. Drug Skin Tests –  Branka Marinović, Jasna Lipozenčić, Ružica Jurakić Tončić, Sandra Marinović Kulišić
	38. In vitro Tests of T-Cell-Mediated Drug Hypersensitivity and Basophil Activation Tests –
	Sabina Rabatić, Valerija Vojvoda

### VII. DESENSITIZATION

39.	Desensitization on Antibiotics, Chemotherapeutics and Biologicals –
	Franjo Gruber
40.	Specific Immunotherapy in Atopic Dermatitis – Current Possibilities and Update Knowledge –
	Magdalena Czarnecka Operacz

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### **AUTHOR INDEX**

Bach-Rojecki, L. • 1

Bašić-Jukić, N. • 34

Batinić, D. • 26

Begovac, J. • 27

Bukvić Mokos, Z. • 14

Czarnecka-Operacz, M. • 40

Čarija, R. • 6

Čeović, R. • 15

Davidovici, BB, • 25

Dolenc Voljč, M. • 30

Fergie, B. • 21

Frew, JW • 22

Gregurić, S. • 5

Gruber, F. • 39

Jurakić Tončić, R. • 28, 36, 37

Kes, P. • 34

Kostović, K. • 13

Kovačević, M. • 18

Lacković, Z. • 1

Lipozenčić, J. • 2, 3, 5, 12, 25, 28, 35, 36, 37

Lugović Mihić, L. • 29

Ljubojević, S. • 31

Makar Ausperger, K. • 10, 24

Marinović, B. • 16, 33, 36, 37

Marinović Kulišić, S. • 37

Mirić-Kovačević, L. • 6

Mravak Stipetić, M. • 23, 32

Murat Sušić, S. • 19

Murrel, D.F. • 21, 22

Paštar, Z. • 13, 35

Pavičić Baldani, D. • 17

Pustišek, N. • 8

Puizina Ivić, N. • 6

Rabatić, S. • 38

Radoš, J. • 38

Ra Intong, L. • 21

Ramos-e-Silva, M. • 12

Sekulić, A. • 7

Skerley, M. • 11

Stanimirović, A. • 18

Škrgatić, L. • 17

Šprem Goldštajn, M. • 17

Štajminger, G. • 35

Štulhofer Buzina, D. • 20

Turčić, P. • 3, 4

Vojvoda, V. • 38

Vuković, D. • 6

Waugh, M. • 9

Zekan, Š. • 27

### **ABBREVIATIONS**

AD

Atopic Dermatitis

**ADEs** 

Adverse Medication Effects

ADRs

Adverse Drug Reactions

ΑE

Adverse events

**AED** 

AntiEpileptic Drugs

**AGEP** 

Acute Generalized Exanthematic Pustulosis

ΔGW

AnoGenital Warts

AΚ

Aktinic Keratosis

**ALA** 

AminoLevulinic Acid

A DC

Antigen Presenting Cells

**ASA** 

AcetylSalicylic Acid

BAT

**Basophil Activation Test** 

**BCC** 

Basal Cell Carcinoma

BD

Bowen Disease

**BLs** 

Beta Lactams

BPG

British Photodermatology Group

CLA

Lymphocyte-Associated Antigen

CM

Contrast Media

COX

Cyclo-OXygenase

CS

Cephalosporins

CTCL

Cutaneous T-cell Lymphoma

DFS

DiethylStilbestrol

рінс

Drug-Induced Hypersensitivity Syndrome

DI OI

Dermatology Life Quality Index

**DRESS** 

Drug Rush with Eosinophilia and Systemic

Symptoms

**EAACI** 

European Academy of Allergy and Clinical

Immunology

**EBA** 

Epidermolysis Bullosa

**ELISA** 

Enzyme-Linked ImmunoSorbent Assay

**ELISPOT** 

Enzyme-Linked ImmunoSorbent Spot assay

**EMLA** 

**Eutectic Mixture of Local Anesthetics** 

**FDA** 

Food and Drug Administration

**FEIA** 

FluoroEnzyme ImmunoAssay

**GM-CSF** 

Granulocyte-Macrophage Colony-Stimulating Factor

**HAART** 

Highly Active AntiRetroviral Treatment

**HOCM** 

High-Osmolality Contrast Media

**HPV** 

Human PapillomaVirus

**ICM** 

Iodinated Contrast Media

ICS

Intracellular Cytokine Stain ing assay

**IDEC** 

Inflammatory Dendritic Epidermal Cells

**IDSA** 

Infectious Diseases Society of America

**IDT** 

IntraDermal Test

INF

**INterFeron** 

INL

International Normalized Ratio

**IRF** 

Independent review facility

**laBCC** 

Locally advanced Basal Cell Carcinoma

LC

Langerhans Cells

LE

Lupus Erythematosus

**LED** 

Light Emitting Diodes

LG\

LymphoGranuloma Venereum

LOCM

Los-Osmolality Contrast Media

LTT

Lymphocyte Transformation Test

MAL

Methyl AminoLevulinic acid

ΜΔΩ

MonoAmine Oxidase

m<sub>BCC</sub>

Metastatic basal cell carcinoma

**MBT** 

Molecular Biologic Techniques

MC

Mollusca Contagiosa

MCV

MollusCipox Virus

**MDM** 

Minor Determinant Mixture

MEL

Monochromatic Excimer Light

Mes

Medication Errors

MHC PID

Major Histocompatibility Complex Pelvic Inflammatory Disease

MDE PPL

Masulopapular Drug Eruption Penicilloil Polylysine

MPE F

MaculoPapular Exanthema Pharmacologic interaction

MRSA PsA

Methyiciline Resistant Staphylococcus aureus Psoriatic Arthritis

MSM PSF

Men who have Sex with Men Progression Free Survival

MTZs RCM

Microscopic Treatment Zones RadioContrast Media

NAAT SJS

Nucleic Acid Amplification Test Stevens-Johnson Syndrome

NSAIDs SPT

NonSteroidal AntiInflammatory drugs Skin Prick Test

OLL STI

Oral Lichenoid Lesion Sexually Transmitted Infections

OLP TCI

Oral Lichen Planus Topical Calcineurin Inhibitors

ORR TCR:

Objective response rate T Cell Receptors

OS TEN

Overall survival Toxic Epidermal Necrolysis

OTC TMP

Over-the-Counter TriMethylPsoralen

PASI TMP-SMX

Psoriasis A>rea and Severity Index TriMetoPrim-SulfaMethoXazole

PDT TNI

PhotoDynamic Therapy Tumor Necrosis Factor

PGA WAO

Physician's Global Assessment World Allergy Organization



INTRODUCTION OF DERMATOTHERAPY