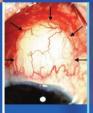
## **ESSENTIALS IN OPHTHALMOLOGY**

G.K.KRIEGLSTEIN · R.N.WEINREB Series Editors



Glaucoma



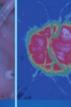
Cataract and Refractive Surgery

Uveitis and Immunological





Vitreo-retinal Surgery



Medical Retina



Oculoplastics and Orbit

500



Pediatric Ophthalmology, Neuroophthalmology, Genetics

Cornea and External Eye Disease

# Glaucoma

## **PROGRESS III**

Edited by F.GREHN R.STAMPER



#### **Essentials in Ophthalmology**

#### Glaucoma

F. Grehn R. Stamper Editors

#### Essentials in Ophthalmology

G. K. Krieglstein R. N. Weinreb Series Editors Glaucoma

Cataract and Refractive Surgery

**Uveitis and Immunological Disorders** 

Vitreo-retinal Surgery

**Medical Retina** 

**Oculoplastics and Orbit** 

Pediatric Ophthalmology, Neuro-Ophthalmology, Genetics

Cornea and External Eye Disease

Vitreo-retinal Surgery

Editors Franz Grehn Robert Stamper

## Glaucoma

With 67 Figures, Mostly in Colour and 8 Tables



#### Series Editors

#### Günter K. Krieglstein, MD

Professor and Chairman Department of Ophthalmology University of Cologne Kerpener Straße 62 50924 Cologne Germany

#### Robert N. Weinreb, MD

Professor and Director Hamilton Glaucoma Center Department of Ophthalmology University of California at San Diego 9500 Gilman Drive La Jolla, CA 92093-0946 USA

#### **Volume Editors**

#### Franz Grehn, MD

Professor and Chairman Department of Ophthalmology University of Wuerzburg Josef-Schneider-Straße 11 97080 Wuerzburg Germany

#### **Robert Stamper, MD**

Director of Glaucoma Service Department of Ophthalmology University of California 10 Kirkham Street, Rm K301 San Francisco CA 94143 USA

ISBN 978-3-540-69472-4

e-ISBN 978-3-540-69475-5

ISSN 1612-3212

Library of Congress Control Number: 2008932232

© 2009 Springer-Verlag Berlin Heidelberg

This work is subject to copyright. All rights are reserved, whether the whole or part of the material is concerned, specifically the rights of translation, reprinting, reuse of illustrations, recitation, broadcasting, reproduction on microfilms or in any other way, and storage in data banks. Duplication of this publication or parts thereof is permitted only under the provisions of the German Copyright Law of September 9, 1965, in its current version, and permission for use must always be obtained from Springer-Verlag. Violations are liable for prosecution under the German Copyright Law.

The use of general descriptive names, registered names, trademarks, etc. in this publication does not imply, even in the absence of a specific statement, that such names are exempt from the relevant protective laws and regulations and therefore free for general use.

Product liability: The publishers cannot guarantee the accuracy of any information about dosage and application contained in this book. In every individual case the user must check such information by consulting the relevant literature.

Cover Design: WMXDesign GmbH, Heidelberg, Germany

Printed on acid-free paper

987654321

springer.com

### Foreword

The Essentials in Ophthalmology series represents an unique updating publication on the progress in all subspecialties of ophthalmology.

In a quarterly rhythm, eight issues are published covering clinically relevant achievements in the whole field of ophthalmology. This timely transfer of advancements for the best possible care of our eye patients has proven to be effective. The initial working hypothesis of providing new knowledge immediately following publication in the peer-reviewed journal and not waiting for the textbook appears to be highly workable.

We are now entering the third cycle of the Essentials in Ophthalmology series, having been encouraged by readership acceptance of the first two series, each of eight volumes. This is a success that was made possible predominantly by the numerous opinion-leading authors and the outstanding section editors, as well as with the constructive support of the publisher. There are many good reasons to continue and still improve the dissemination of this didactic and clinically relevant information.

**G.K. Krieglstein R.N. Weinreb** Series Editors September 2008

## Preface

This third volume in the series, Essentials of Ophthalmology, just like the first, seeks to bring the ophthalmic practitioner up to date in the important new advances or changes in glaucoma diagnosis or management that have occurred over the last ten years. The last decade has seen significant changes in our understanding of the pathophysiology of some glaucomas, in our diagnostic approaches and in our management of them. Toward the goal of providing the most up-to-date information in a readable fashion, we have asked some of the world's experts to discuss areas to which they have contributed in a way that will be useful for the practicing doctor. For example, one of the pioneers in the imaging of live ganglion cells is Dr. Francesca Cordeiro. Her studies could lead to a potentially significant breakthrough as, in the future, clinicians may be able to determine the health and number of ganglion cells in the retina as both a diagnostic and monitoring test. As the prevalence of glaucoma increases in our aging population, epidemiology has become more important as a methodology to identify risk factors; Drs. Giangiacomo and Coleman discuss what we have recently learned that is relevant to our clinical understanding of glaucoma. Drs. Doshi, Weinreb and colleagues describe the diurnal fluctuation of intraocular pressure, how those fluctuations impact on glaucoma, the relationship of postural change to that fluctuation, and what it means for managing glaucoma. Detecting progression of glaucoma can be tricky. Imaging techniques may be helpful. Strouthidis and Garway-Heath tell us how. Our concepts of and terminology for angle-closure glaucoma have undergone major changes over the last few years. Sharma, Low and Foster describe these changes and introduce the new-now internationally agreed upon-terminology. The association of uveitis and glaucoma has been known and has frustrated those caring for patients with these two concurrent conditions for many years; Drs. Nagpal and Acharya discuss the interrelationship between uveitis and glaucoma, what the doctor should look for, and how to manage these difficult patients. New approaches to glaucoma surgery have been described recently. Drs. Mendrinos and Shaarawy describe the techniques and results of nonpenetrating glaucoma surgery. Drs. Tam and Ahmed describe and discuss several new approaches to glaucoma surgery using special shunts that have appeared in the past few years. As electronic medical record systems gain popularity around the world, Drs. Schargus and Grehn describe the European Glaucoma Society's electronic glaucoma record and their agreement on what is important to include in such a system. We hope that all the topics and authors that we have selected are helpful in improving the understanding of the many faces of glaucoma and, ultimately, will contribute to reduced visual loss and better care for our patients.

Franz Grehn Robert L. Stamper

## Contents

#### Chapter 1 Imaging Individual Ganglion Cells in the Human Retina

Nicholas E.H. Wood, Li Guo, M. Francesca Cordeiro

1.1	Introduction	1
1.2	Description of the Imaging	
	Techniques	1
1.3	The Imaging Techniques	2
1.3.1	Scanning Laser Polarimetry	2
1.3.2	High-Resolution Reflectance	
	Imaging	2
1.3.3	Optical Coherence Tomography	3
1.3.4	Confocal Scanning Laser	
	Ophthalmoscopy	5
1.3.5	Adaptive Optics	6
1.4	Applications to RGC Imaging	7
1.4.1	Retrograde Labelling	7
1.4.2	RGC-Specific Fluorescent Protein	
	Expression	7
1.4.3	The Detection of Apoptosing Retinal	
	Cells (DARC)	8
1.5	The Future	9
	References	10

#### Chapter 2 The Epidemiology of Glaucoma

Annette Giangiacomo, Anne Louise Coleman

21	Introduction	13
2.2	Primary Open-Angle Glaucoma	13
2.2.1	Increased IOP	14
2.2.2	Age	14
2.2.3	Family History	15
2.2.4	Sex	15
2.2.5	Ethnicity	15
2.2.6	Муоріа	16
2.2.7	Other Risk Factors	17
2.3	Primary Angle-Closure Glaucoma	17
2.3.1	Risk Factors	18
2.3.2	Prevalence	18
	References	19

#### Chapter 3 Circadian Changes in Intraocular Pressure

Amish B. Doshi, John H.K. Liu, Robert N. Weinreb

3.1	Introduction	23
3.2	Normal IOP Curve	23
3.3	Sources of Circadian Control	24
3.4	Glaucoma and 24-Hour IOP	25
3.5	Medical Management of	
	24-Hour IOP	26
	References	27

#### Chapter 4

#### **Detecting Glaucoma Progression by Imaging**

Nicholas G. Strouthidis, David F. Garway-Heath

4.1	Introduction	29
4.1.1	The Principles of Progression	29
4.1.2	Historical Perspective: Optic	
	Nerve Head Photography	30
4.1.3	The Potential of Optic Nerve	
	Head Imaging Devices	30
4.2	HRT	31
4.2.1	HRT Progression: Available	
	Techniques	31
4.2.2	HRT Progression: Stereometric	
	Parameter vs. Pixel-Based	
	Techniques	31
4.2.3	HRT Progression: Stereometric	
	Parameter Event Analyses	34
4.2.4	HRT Progression: Stereometric	
	Parameter Trend Analyses	34
4.2.5	HRT Progression: Pixel-Based	
	Technique	35
4.3	Detecting Progression by	
	GDx-VCC	35
4.4	Detecting Progression by OCT	37
4.5	Frequency of Testing	37
4.6	Lack of Concordance	38
	References	39

## Chapter 5

#### The Classification of Primary Angle-Closure Glaucoma

Tarun Sharma, Sancy Low, Paul J. Foster

5.1	Background
5.2	The Purposes of Disease
	Classification
5.3	The Evolution of Classification
	Schemes for Angle-Closure
	Glaucoma
5.4	Definition of an "Occludable" or
	Narrow Angle
5.5	Primary Open-Angle Glaucoma is a
	Diagnosis of Exclusion
5.6	Classification of Angle Closure in
	Epidemiological Research
	(ISGEO Scheme)
5.7	Trabecular Meshwork Damage in
	Angle Closure
5.8	An Anatomical Basis for the Primary
	Angle Closure Mechanism
5.9	Classification System for
	Angle-Closure Glaucomas
5.9.1	Level I: Iris and Pupil
5.9.2	Level II: Ciliary Body
5.9.3	Level III: Lens-Induced Angle
	Closure
5.9.4	Level IV: Ciliolenticular Block/Aqueous
	Misdirection/"Malignant Glaucoma"
5.10	Gonioscopy
	References

#### Chapter 6 **Uveitic Glaucoma**

Agnieszka G. Nagpal, Nisha R. Acharya

6.1	Introduction
6.2	The Epidemiology of Uveitis-Related Ocular Hypertension (OHT) and Secondary Glaucoma
6.3	Pathogenesis of Uveitic Glaucoma
6.3.1	Aqueous Dynamics in Uveitic
	Glaucoma
6.3.2	Mechanical Causes of Uveitic
	Glaucoma
6.3.3	Steroid-Induced Glaucoma
6.4	Common Uveitic Entities Associated
	with OHT and Secondary
	Glaucoma
6.4.1	Glaucomatocyclitic Crisis: Posner-
	Schlossman Syndrome
6.4.2	Fuchs' Heterochromic
	Iridocyclitis

	6.4.3	Herpetic Disease	53
	6.4.4	Juvenile Inflammatory Arthritis (JIA)	53
	6.4.5	Pars Planitis	53
	6.4.6	Toxoplasmosis	54
	6.4.7	Sarcoidosis	54
42	6.4.8	Syphilis	54
	6.5	Treatment of Uveitic Glaucoma	55
42	6.5.1	Medical Treatment	55
	6.5.2	Surgical Treatment	55
	6.6	Conclusion	56
43		References	56

#### Chapter 7

43

#### 44 Nonpenetrating Glaucoma Surgery

Efstratios Mendrinos, Tarek Shaarawy

44	7.1	Introduction	59
	7.2	Deep Sclerectomy	59
45	7.2.1	Superficial Scleral Flaps	59
15	7.2.2	Deep Sclerectomy and Exposure	57
45	1.2.2	of Trabeculo-Descemet's	
15		Membrane	60
45	7.3	Deep Sclerectomy Technique	60
45	7.4	The Use of Implants	61
46	7.5	Viscocanalostomy	61
	7.6	Mechanisms of Filtration	62
46	7.6.1	Flow Through the TDM	62
	7.6.2	Aqueous Humor Resorption	62
46	7.7	Nd:Yag Goniopuncture	63
47	7.8	Technique	64
47	7.9	Indications for Nonpenetrating	•••
		Glaucoma Surgery	64
	7.9.1	Primary Open Angle Glaucoma	64
	7.9.2	Glaucoma in High Myopia	64
	7.9.3	Pseudoexfoliation and Pigmentary	
		Glaucoma	64
	7.9.4	Uveitic Glaucoma	64
49	7.9.5	Congenital and Juvenile Glaucoma	64
	7.9.6	Glaucoma Associated with Sturge-	
		Weber Syndrome	65
49	7.9.7	Glaucoma in Aphakia	65
50	7.10	Contraindications for Nonpenetrating	
		Glaucoma Surgery	65
50	7.10.1	Relative Contraindications	65
	7.10.2	Absolute Contraindications	66
50	7.11	Complications of Nonpenetrating	
51		Glaucoma Surgery	66
	7.11.1	Intraoperative Complications	66
	7.11.2	Early Postoperative Complications	67
51	7.11.3	Late Postoperative Complications	69
	7.12	Clinical Experience with	
51		Nonpenetrating Glaucoma	
		Surgery	70
52	7.12.1	Viscocanalostomy	70

7.12.2	Deep Sclerectomy
7.12.3	Studies Comparing Trabeculectomy
	and Nonpenetrating Glaucoma
	Surgery
	References

#### Chapter 8 New Glaucoma Surgical Devices

Diamond Y. Tam, Iqbal Ike K. Ahmed

Introduction	75
Basic Review of the Anatomy and	
Physiology of Aqueous Outflow and	
Drainage Devices	76
Subconjunctival Filtration	77
Schlemm's Canal Outflow	77
Subconjunctival Filtration Device:	
the Ex-PRESS Shunt	79
Schlemm's Canal Devices: Canaloplasty/	
iScience, Glaukos Trabecular	
Micro-Bypass Stent,	
and the Trabectome	81
Ab Externo Schlemm's Canal	
Approaches: Nonpenetrating	
Schlemm's Canaloplasty	81
Trabecular Micro-Bypass Stent	86
Trabectome	89
Suprachoroidal Filtration Device:	
The SOLX Gold Microshunt	91
	Basic Review of the Anatomy and Physiology of Aqueous Outflow and Drainage Devices

8.6	Conclusion	94
	References	96

## 71 Chapter 9

70

#### Digital Glaucoma Patient Record and Teleconsultation Systems for Glaucoma Specialists: The European Glaucoma Society Glaucocard Project

Marc Schargus, Franz Grehn, The Glaucocard Workgroup

9.1	Introduction	99
9.2	History of Telemedicine in	
	Ophthalmology	100
9.3	The Concept of an Electronic	
	Glaucoma Patient Health	
	Record System	101
9.3.1	General Issues for	
	Implementation	101
9.3.2	Important Classifications for	
	Electronic Glaucoma Medical	
	Record Systems	104
9.4	The EGS Glaucocard Project	106
9.5	Future Prospects	109
9.6	Conclusion	109
	Acknowledgments	110
	References	110
Index		113
acA	•••••••••••••••••••••••••••••••••••••••	115

Contents

## Contributors

#### Nisha R. Acharya

Francis I. Proctor Foundation University of California San Francisco 95 Kirkham St., San Francisco CA 94143, USA

#### Iqbal Ike K. Ahmed

University of Toronto Toronto Ontario, Canada

#### Anne Louise Coleman

Jules Stein Eye Institute/UCLA 100 Stein Plaza Los Angeles, CA 90095 USA

#### M. Francesca Cordeiro

Glaucoma & Retinal Degeneration Research Group UCL Institute of Ophthalmology Bath Street London EC1V 9EL, UK

#### Amish B. Doshi

Hamilton Glaucoma Center Department of Ophthalmology University of California San Diego, CA USA

#### **Paul Foster**

Department of Epidemiology & International Eye Health UCL Institute of Ophthalmology 11-43 Bath Street London EC1V 9EL, UK

#### David F. Garway-Heath

Moorfields Eye Hospital and UCL Institute of Ophthalmology NIHR Biomedical Research Centre 162 City Road London, UK

#### Annette Giangiacomo

CB 7040, 5109 Bioinformatics Building Department of Ophthalmology University of North Carolina-Chapel Hill Chapel Hill, NC 27599-7040 USA

#### Franz Grehn

University Eye Hospital Wuerzburg Josef Schneider Str. 11 97080 Wuerzburg Germany

#### Li Guo

Glaucoma & Retinal Degeneration Research Group, UCL Institute of Ophthalmology Bath Street London EC1V 9EL, UK

#### John H.K. Liu

Hamilton Glaucoma Center Department of Ophthalmology University of California San Diego CA, USA

#### Sancy Low

Glaucoma Service Moorfields Eye Hospital London, UK Department of Epidemiology and International Eye Health UCL Institute of Ophthalmology Bath Street, EC1V 9EL London, UK

#### **Efstratios Mendrinos**

Department of Ophthalmology Glaucoma Unit Geneva University Hospitals 1211 Geneva 14 Switzerland

#### Agnieszka G. Nagpal

Francis I. Proctor Foundation University of California, San Francisco 95 Kirkham St. San Francisco, CA 94143 USA

#### Marc Schargus

University Eye Hospital Wuerzburg Josef Schneider Str. 11 97080 Wuerzburg Germany

#### **Tarek Shaarawy**

Glaucoma Unit Department of Ophthalmology Geneva University Hospitals Alcide-Jentzer 22 1211 Geneva 14 Switzerland

#### Tarun Sharma

Glaucoma Service Moorfields Eye Hospital London, UK

#### Nicholas G. Strouthidis

Moorfields Eye Hospital and UCL Institute of Ophthalmology NIHR Biomedical Research Centre 162 City Road London, UK

#### **Diamond Y. Tam**

University of Toronto Toronto Ontario, Canada

#### **Robert N. Weinreb**

Department of Ophthalmology University of California 9500 Gilman Drive La Jolla, CA 92093 USA

#### Nick Wood

Glaucoma & Retinal Degeneration Research Group UCL Institute of Ophthalmology Bath Street EC1V 9EL London, UK

#### Chapter 1

## Imaging Individual Ganglion Cells in the Human Retina

Nicholas E.H. Nick Wood, Li Guo, M. Francesca Cordeiro

#### **Core Messages**

- Retinal ganglion cells (RGCs) are the key cells implicated in glaucoma, and their assessment could lead to effective treatment and monitoring regimens
- Scanning laser polarimety (SLP) gives a good measure of RNFL thickness and RGC axonal loss but cannot provide focussed information about RGCs
- High-resolution reflectance imaging uses highquality CCDs(charge-coupled device), which can use much more information from simple funduscopic observa-tions but again provide little information on RGCs
- Optical coherence tomography (OCT) is a rapidly developing technology which is now enabling

#### 1.1 Introduction

Glaucoma is a leading cause of blindness worldwide [1] and it is expected that the number of people with the disease will rise dramatically by 2020 [2]. Diagnosis is traditionally from changes in the optic nerve head (ONH) and visual field loss, but these can only detect the disease after significant (25–40%) loss of retinal ganglion cells (RGCs), the key cell implicated in this process [3, 4].

The inner retinal layers, being optical media that are therefore transparent to visible-frequency light, are inherently low contrast. This presents a significant challenge for traditional imaging such as fundus imaging. Modern technologies now use many different properties of light to differentiate between the retinal structures and these technologies are enabling us to observe fine detail, such as the photoreceptor layers, in vivo [5]. Combined with other techniques, they allow the examination of individual RGCs [3, 5–7]. In vivo imaging also enables longitudinal studies [3, 5], which brings great possibilities for elucidating disease pathways and developing new treatments [8, 9]. retinal cellular, functional and 3D imaging, but its role in RGC imaging is still uncertain

- Most promising technologies use the established confocal scanning laser ophthalmoscope (cSLO) combined with other methodologies to improve RGC visualization
- Imaging in experimental research has permitted the direct assessment and successful evaluation of RGCs in disease models
- Some safe techniques developed in animal models are beginning to make the crossover into clinical glaucoma detection
- Ideally, methodologies enabling the visualization of healthy and "sick" RGCs would provide a comprehensive assessment of glaucomatous changes and disease states

Recent advances have allowed unprecedented access to the retinal layers, creating the possibility of potentially visualizing ganglion cells in order to provide a new and early clinical parameter for glaucomatous injury. This chapter aims to cover the current research achievements in RGC imaging and the promising directions they are taking visual science.

#### 1.2 Description of the Imaging Techniques

- Scanning laser polarimetry (SLP): A confocal imaging system with a polarimeter to measure the birefringence caused by the retinal nerve fibre layer (RNFL)
- High-resolution reflectance imaging: Based around a fundus camera with a high-quality CCD camera, this system can take a sequence of rapid images which can measure wavelength-dependent reflectance changes with very high temporal resolution
- Optical coherence tomography (OCT): A low-coherence interferometry-based imaging system where changes in reflectivity are measured in a volume of the retina with very high axial resolution