## Pearls of Glaucoma Management

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### **Foreword**

If you have ever uttered the commonly expressed lament, "Glaucoma is so confusing!" then this text is for you. You will no longer be bewildered.

Why practitioners may be confused about how to be of help to patients with glaucoma – in its many incarnations and reincarnations – is easily understood. The issue seems to be overwhelming when one considers that the already massive population of those with glaucoma is increasing rapidly as the world's population increases and ages.

During the past 50 years the fundamental definition of glaucoma has changed almost 180°, and the indications for treatment have become more variable and controversial, some advising early therapy and others strongly cautioning against such an approach: Various diagnostic tests have come and gone and are interpreted in such different ways that there seems to be no consensus; surgical techniques come in and out of fashion in perplexing ways. There seems to be a constantly shifting, sandy foundation on which are built unsteady schools of ever-varying advice. Why practitioners, patients, and the public are often bewildered is understandable.

The current text was designed to be relevant, scientific, and practical. The editors have accomplished their objective well. The authors chosen to share their wisdom are expert practitioners who recognize the dangers of basing treatment on theory. They, the leaders in their fields, create an understanding of glaucoma and conditions related to glaucoma that is sound, scientific, and effective. The editors clearly instructed their contributors to avoid speculation, to be practical, and to insist on evidence, not opinion (and where good evidence was lacking, to indicate such a lack). The result is a cohesive picture that should be of immense help to all those trying to make sense of what to many seems to be confusing.

It is perhaps not surprising that this text accomplishes its objective so admirably. The senior editor is a vastly experienced physician, equally at home in the clinic, the operating room, the classroom, and in a basic research laboratory. The contributing authors come from many different institutions and cultures; some are younger and others older. The current text, however, does not present information that must be sifted by a discerning reader in order to come up with appropriate advice. Rather, the authors simplify, clarify, organize, and explain practically and scientifically. Those wanting to know how to approach patients with glaucoma or those many, many patients in whom it is not clear whether glaucoma is present or not will find this a treasure trove of sound science blended with critical experience.

The need for this intellectually vigorous, practical approach to caring for patients with conditions related to intraocular pressure and optic nerve disease is great. There is probably truth in the belief that all persons will eventually develop glaucoma if they

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live long enough. As the world population ages and increases, as resources become ever more precious, as cost considerations become more confining, there is increasing urgency for guidelines that concentrate on the essentials and that will help achieve the goal of caring for the sick and for the well, specifically, the greatest good for the greatest number, while still addressing the needs and wants of each individual person.

Currently there is much interest in "translational research." This book is highly successful in translating vast amounts of disparate, sometimes disconcerting information into understandable sentences, paragraphs, and illustrations that will result in more effective and more relevant care.

Pennsylvania, USA

George Spaeth

#### **Preface**

This book was developed based on the questions that clinicians taking care of glaucoma patients, fellows, and residents have asked us as consultants. Most textbooks on glaucoma provide a broad overview of the basic science and clinical literature, which is very useful for students learning about glaucoma. However, these textbooks may leave many questions unanswered for the clinician specifically searching for advice on how to manage a specific problem. We have intentionally included topics that are not yet traditionally found in textbooks, such as new surgical technologies and measurement of optic nerve blood flow, because these and other areas are being discussed at meetings, and clinicians have specific questions about them.

In addition to asking the questions that frequently arise in managing patients with glaucoma, the goal of this textbook was to have the authors who are familiar with the world literature digest that information in the context of their own clinical experience. We asked authors to answer questions the way they might answer a physician's questions over the phone. We asked them to state their opinions on how they like to manage clinical situations, where appropriate, but to also point out that their preferred management is not the only way to manage the problem if other acceptable means are available. The questions are organized by topic and cover diagnostic testing and interpretation, risk factors, medical treatment, procedural treatments, various glaucoma subtypes, and complications.

We must thank all the consulting physicians, students, residents, and fellows who we have encountered, and who inspired this text. Special thanks to Ms. Trini Phan who helped jump-start this project by sending out the invitations to contribute to our world-expert authors, and to Mr. Doug Hoffman whose technical assistance was invaluable.

Los Angeles, California, USA

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### **Abbreviations**

ACC Acute angle closure
ACG Angle closure glaucoma

AGIS Advanced Glaucoma Intervention Study

AL Axial length

ALT Argon laser trabeculoplasty
APON Acquired pit of the optic nerve

AUC Area under the receiver operating curve

BAK Benzalkonium chloride
CAI Carbonic anhydrase inhibitor
CCT Central corneal thickness

C/D Cup-to-disc ratio
CDI Color Doppler imaging
CDR Cup-to-disc ratio
CH Corneal hysteresis

CIGTS Collaborative Initial Glaucoma Treatment Study

CLBF Canon laser blood flowmetry

CNTGS Collaborative Normal Tension Glaucoma Study

CSLO Confocal scanning laser ophthalmoscopy

CTD Congenital tilted disc

dB Decibels

DCT Dynamic contour tonometer ECD Endothelial cell density ECM Extracellular matrix

ECP Endoscopic cyclophotocoagulation EGPS European Glaucoma Prevention Study EMGT Early Manifest Glaucoma Trial

FBCF Fornix based conjunctival flap

FDT Frequency doubling technology perimetry

5FU 5-Flurouracil

GAT Goldmann applanation tonometry
GATE German adaptive threshold estimation

GCP Glaucoma change probability GDD Glaucoma drainage device

GDX Brand name of machine that performs scanning laser polarimetry

GDX-ECC GDX with enhanced corneal compensation

GDX-NFA GDX nerve fiber analyzer

GDX-VCC GDX with variable corneal compensation

xxx Abbreviations

Glaucoma hemifield test **GHT Guided Progression Analysis GPA GPS** Glaucoma Probability Score **HRP** High pass resolution perimetry Heidelberg retinal tomography HRT **HFA** Humphrey visual field analyzer **HRF** Heidelberg retinal flowmeter **ICE** Iridocorneal endothelial syndrome

ILM Inner limiting membrane IOP Intraocular pressure

ISNT Mneumonic for the thickest (inferior) to thinnest (temporal)

neuroretinal rim

LASIK Laser in situ keratomileusis
LBCF Limbal based conjunctival flap
LDF Laser Doppler flowmetry
LPI Laser peripheral iridotomy
LTP Laser trabeculoplasty
MD Mean deviation

mf ERG Multifocal electroretinogram mf VEP Multifocal visual evoked potential

MMC Mitomycin-C

MPHSD Mean pixel height standard deviation

MRA Moorfield regression analysis

MTD Myopic tilted disc

NFI Nerve fiber index (used on GDX)

NFL Nerve fiber layer

NFLT Nerve fiber layer thickness NTG Normal tension glaucoma NV Neovascularization

NVG Neovascular glaucoma
OAG Open angle glaucoma
OBF Ocular blood flow

OCT Optical coherence tomography

OHTS Ocular Hypertension Treatment Study

OND Optic nerve drusen
ONH Optic nerve head

ONTT Optic Neuritis Treatment Trial

OPA Ocular pulse amplitude
OPP Ocular perfusion pressure
ORA Ocular response analyzer
PAS Peripheral anterior synechiae
POAG Primary open angle glaucoma
POBF Pulsatile ocular blood flowmeter

PPA Peripapillary atrophy PSD Pattern standard deviation

PXF Pseudoexfoliation
RBP Rarebit perimetry
RGC Retinal ganglion cell
RNFL Retinal nerve fiber layer

Abbreviations xxxi

RPE Retinal pigment epithelium RVA Retinal vessel analyzer SAP Standard automated perimetry

SD Standard deviation

SD-OCT Spectral domain optical coherence tomography (also known as

Fourier domain OCT)

SITA Swedish Interactive Threshold Algorithm

SLO Scanning laser ophthalmoscope SLP Scanning laser polarimetry SLT Selective laser trabeculoplasty

SWAP Short wavelength automated perimetry

SWS Sturge Weber syndrome

TCA Topographical change analysis
TCP Transcleral cyclophotocoagulation
TOP Tendency oriented perimetry

TSNIT Temporal, superior, nasal, inferior, temporal

VA Visual acuity

VCDR Vertical cup to disc ratio

VF Visual field VFI Visual field index

ZEST Zippy estimation by sequential testing

3D 3-Dimensional

Claude F. Burgoyne

#### **Core Messages**

- > The principle insult in glaucoma occurs within the neural, cellular, and connective tissues of the optic nerve head (ONH).
- > Intraocular pressure at all levels has biomechanical effects on the optic nerve tissues.
- Clinical cupping is one manifestation of the pathophysiology of glaucomatous damage, but is not the pathophysiology itself.
- The variable appearance of the ONH in all optic neuropathies is the predictable result of ONH tissue biomechanics.
- As our clinical tools for characterizing ONH biomechanics improve, so too will our ability to understand normal ONH aging and its contributions to the clinical behavior and susceptibility of the ONH.

# 1.1 Why Is the Optic Nerve Important in the Diagnosis and Management of Glaucoma?

Glaucoma is an optic neuropathy. Although there are several pathophysiologies that must be managed in the clinical care of the glaucoma patient, what defines all forms of glaucoma is an optic neuropathy that

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demonstrates classic and recognizably variable [1–6] structural and functional behaviors.

#### 1.1.1 The Optic Nerve Head (ONH) Is the Principal Site of Glaucomatous Damage to the Visual System

Although glaucomatous damage likely encompasses important pathophysiology within the retinal ganglion cell (RGC) stroma [7–12], photoreceptors [13–17], lateral geniculate body [18–20], and visual cortex [20], strong evidence suggests that damage to the retinal ganglion cell axons within the lamina cribrosa of the ONH [21–26] is the central pathophysiology underlying glaucomatous vision loss. Recent studies in monkeys [25–30], rats [31–33], and mice [34] support the importance of the ONH in glaucoma by describing profound alterations at the earliest detectable stage of the disease within the prelaminar, laminar, and peripapillary scleral tissues of the ONH.

The ONH tissues make up a dynamic environment wherein 1.2–2.0 million RGC axons converge, turn, and exit the eye through the inner (Bruch's membrane opening) and outer (scleral) portions of the neural canal (Fig. 1.1). Within the scleral portion of the canal, the bundled axons pass through a 3-dimensional meshwork of astrocyte-covered, capillary-containing connective tissue beams known as the lamina cribrosa (Fig. 1.1). Within the lamina, axonal nutrition is dependant upon the movement of oxygen and nutrients from the laminar capillaries, through the laminar beam extracellular matrix (ECM), into the laminar astrocyte processes within the beam, finally reaching the peripheral and central axons of each bundle, via cell processes [35].

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