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Corneal Tomography and Topography for Refractive and Cataract Surgeons

Session # IC-101
 Saturday July 24, 2021: 8:00 AM - 9:30 AM,
 Mandalay Bay Convention Center, Level 2, Lagoon KL

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Program

1. Renato Ambrósio Jr., MD, PhD
Scheimpflug Imaging
2. David Huang, MD, PhD
Corneal OCT
3. Li Wang, MD, PhD
Measuring the cornea for IOL calculations

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Scheimpflug Imaging for Enhanced Refractive Diagnosis

Renato Ambrósio Jr., MD, PhD



Renato Ambrósio Jr.
PROFESSOR TITULAR - UNIVERSIDADE

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Financial Disclosures

- Oculus
- Alcon/Wavelight
- Allergan
- Essilor
- Genom / União Química
- Ofta Vision Health
- Mediphacos
- ZEISS

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PROF. DR. AYSIANO MACHADO

PROF. DR. JOÃO MARCELO LYRA

PROF. DR. RENATO AMBRÓSIO



Edileuza V. Leão
PhD



Bernardo T. Lopes,
MD MPhil PhD

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MasterClass
Renato Ambrósio Jr.
 || *Multimodal Refractive Imaging*

Renato Ambrósio Jr.
 M.D., F.R.C.
 OPHTHALMOLOGIST • SURGEON SCIENTIST

MY/LEARNING
 MILTON YOGI STUDY GROUP

@dr.renatoambrosiojr @milton.yogi

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As for me, all I know is I know nothing.

~ Socrates

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Scheimpflug Imaging for Refractive Surgery

| | | |
|---|--|--|
| <i>Scheimpflug Imaging: Basic Concepts</i> | <i>Multimodal Imaging for Refractive Surgery</i> | <i>Enhanced Ectasia Belin / Ambrósio</i> |
| <i>Interpretation for Clinical Decision</i> | <i>Ancient Intelligence & Artificial Intelligence (AI)²</i> | <i>Clinical Examples</i> |

Renato Ambrósio Jr.
 FERRAZ DE ALVES, SÃO PAULO, BRAZIL

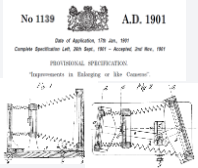
9

A little of History...

● Geometric principle was first described by Jules Carpentier (1901)

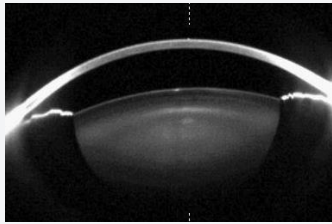


Jules Carpentier (1851 – 1921)



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Scheimpflug Imaging



Ronald Johnson
PROFESSOR OF OPTICS, UNIVERSITY OF GLASGOW

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Slit Lamp Scheimpflug



Ronald Johnson
PROFESSOR OF OPTICS, UNIVERSITY OF GLASGOW

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Evolution in Corneal Imaging

CORNEAL TOPOGRAPHY
(Corneal Surface Shape; Regularity)

CORNEAL TOMOGRAPHY
(3D Corneal Architecture;
front/back surfaces; elevation)

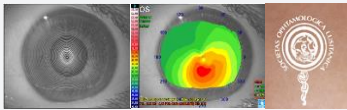


Ronald Anderson Jr
FACR, FRCO, FRCOphth, FRCOphthEd

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Corneal Topography

- from Greek τόπος (topos), meaning "place", and γράφω graphia, meaning "writing"
- Study of the front corneal surface
- Placido's Reflection

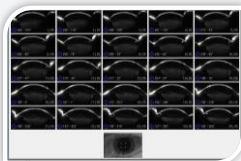
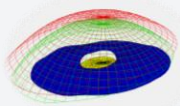


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Corneal Tomography


- from Greek τόμος (tomos), meaning "section, slice", and γράφω (graphia) "writing"
- 3D Study of the corneal architecture (front & back surfaces)
 - Scheimpflug



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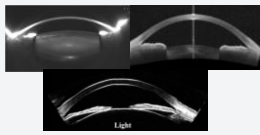
Scheimpflug ToMography




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Corneal ToMography


- from Greek τόμος (tomos), meaning “section, slice”, and γραφή (graphia) “writing”
- 3D Study of the corneal architecture (front & back surfaces)
 - Scheimpflug
 - OCT
 - VHF-US





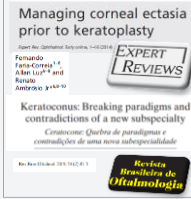
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Segmental or Layered ToMography



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Enhanced Diagnostics for Ectasia: The What's & The Why's



- Screening for ectasia Susceptibility
- Diagnosing Ectasia
- Staging the disease
- Clinical Management
- Follow-up

Renato Ambrósio Jr
PROFESSOR, TEREZINA, BRAZIL

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International English Edition

Highlights of Ophthalmology

ISSUE 08 VOLUME 12 • NUMBER 4

Multimodal Imaging in Refractive Surgery
 Louisa Pellegrino Gomes Esperante, MD
 Marcela G. Salomão, MD
 Nelson Batista Sene Jr, MD
 Jorge Haddad, MD
 (Brazil)
 David C. Dawson, MD
 (USA)
 Fernando Faria Corrales, MD, PhD, FEBOC-CE
 (Portugal)
 Renato Ambrósio Jr, MD, PhD
 (Brazil)

INDIAN JOURNAL OF OPHTHALMOLOGY Volume 08 Issue 12

Guest Editorial

Multimodal imaging for refractive surgery: Quo vadis?
 Prof. Renato Ambrósio Jr, MD, PhD



Renato Ambrósio Jr
PROFESSOR, TEREZINA, BRAZIL

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The Pentacam® Family

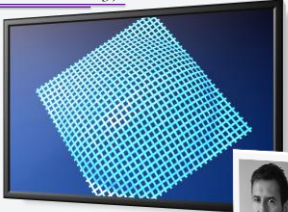


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*Elevation map is a Subtractor
from the Best Fit Reference (Float, Geometry)*

- Anterior e Posterior
- Reference
 - BFS, BFTE
 - Float
 - Zone for Calculation
- Pattern
- Central Elevation
- Elevation @ Thinnest Point
- Elevation @ Highest value within 4mm

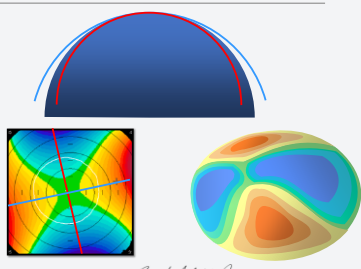


Courtesy of
Dariusz Gajda, MD, PhD
(France)

Randy Johnson Jr
PROGRESSIVE CONTACT MASTERS

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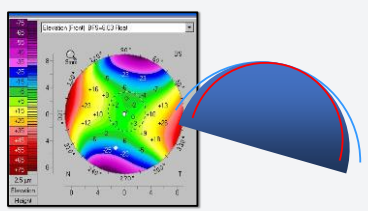
Registration of Elevation



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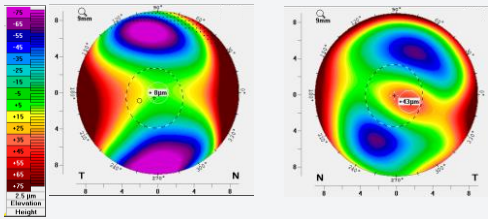
Astigmatism



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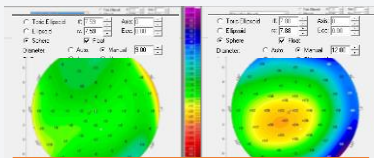
Astigmatism vs. Ectasia



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Corneal Zone Diameter For Calculating the Reference



Recommend BFS 8mm *Float*

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Best-Fit-Sphere (BFS) x Best-Fit Toric-Ellipsoid (BFTE)

The impact of reference body selection for calculating Posterior Corneal Elevation for Diagnosing Keratoconus: BFS x BFTE

Ana Laura C. Casado, MD
 Renato Aquilino, Jr, MD PhD, Kenneth Cornejo, MD, Leonardo Salcedo, MD, Brian Yabon, MD, Federico P. Guerra, MD, Michael W. Dolan, MD, FACS

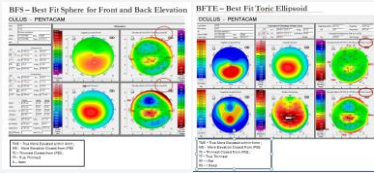
WORLD VI CORNEAL CONGRESS

Purpose: To compare the sensitivity and specificity for diagnosing keratoconus using elevation maps calculated from the Routed Best Fit Sphere (BFS) and Best Fit Toric-Ellipsoid (BFTE) for the 8mm area.

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*Best-Fit-Sphere (BFS) x
Best-Fit Toric-Ellipsoid (BFTE)*



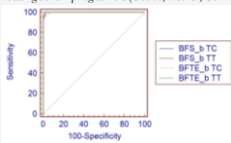
Ronald Johnson Jr
PROGRESSIVE LASER REFRACTIVES

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Corneal Elevation Back : Thinnest

- 498 normal corneas from 49 patients and 56 keratoconic corneas from 28 patients
- Pentacam HR rotating Scheimpflug camera (Oculus, Wetzlar, Germany).



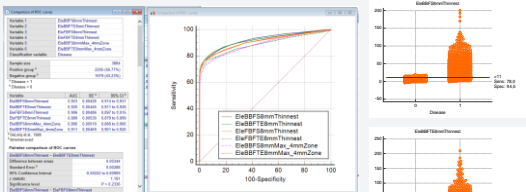
| P-value | BFS_b TC | BFS_b TT | BFE_b TC | BFE_b TT | ROC | ROC | ROC | ROC | ROC | ROC | ROC | ROC |
|----------|----------|----------|----------|----------|----------|------|------|-----|-------|----------|-----|-----|
| BFS_b TC | 1 | 0.436 | 0.094 | 0.145 | BFS_b TC | 100 | 96.9 | >17 | 0.999 | 0.0007 | | |
| BFS_b TT | | 1 | 0.067 | 0.131 | BFS_b TT | 100 | 100 | >19 | 1 | 0 | | |
| BFE_b TC | | | 1 | 0.093 | BFE_b TC | 94.6 | 100 | >16 | 0.998 | 0.00166 | | |
| BFE_b TT | | | | 1 | BFE_b TT | 100 | 98 | 12 | 0.999 | 0.000911 | | |

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PROGRESSIVE LASER REFRACTIVES

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**“Big Study” Data: No difference for All
(N x [KC+VA-E+VAE-NT])**

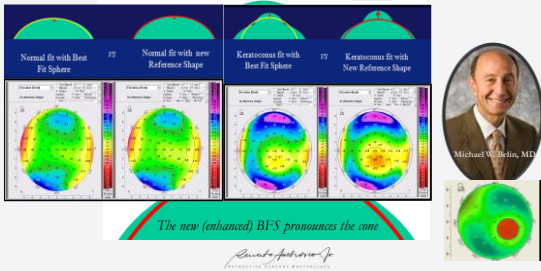


Ronald Johnson Jr
PROGRESSIVE LASER REFRACTIVES

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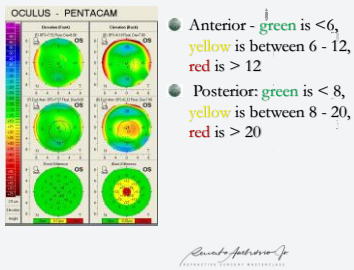


Enhanced BFS Elevation Concept (Belin)



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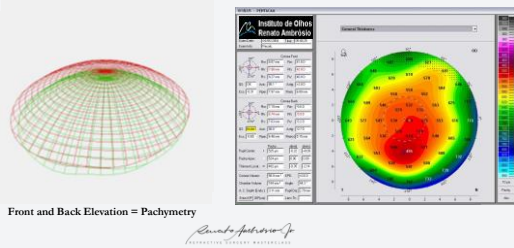
Enhanced BFS (Belin)



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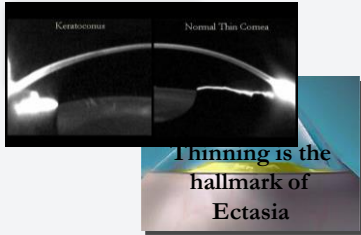
CORNEAL TOMOGRAPHY

3D Architecture



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Corneal Thickness Spatial Profile



Renato Ambrosio Jr.
PROFESSOR, VISUAL OPTICS

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Keratoconus: Spatial Variation of Corneal Thickness as a Diagnostic Test

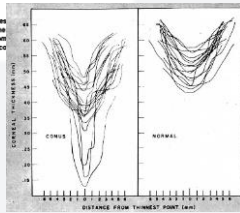
Robert B. Mandell, PhD, and Kenneth A. Polse, OD, Berkeley, Calif

The spatial variation in thickness for the horizontal meridian of eighteen keratoconic corneas was measured with the aid of a pachometer and a new automatic recording system. The significance of the results is discussed.

Arch Ophthalmol—Vol 82, Aug 1969



Robert Mandell, PhD Kenneth A. Polse, OD



Renato Ambrosio Jr.
PROFESSOR, VISUAL OPTICS

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Corneal Thickness Profile: CTSP & PTI

average of the thickness in the points within imaginary circles centered on the Thinnest Point with increasing diameters



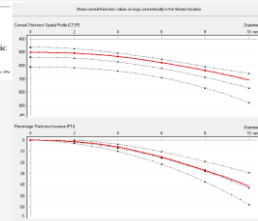
Application: Keratoconus

Corneal thickness spatial profile and corneal volume distribution: Tomographic indices to detect keratoconus

Renato Ambrosio Jr., MD, PhD



Renato Ambrosio Jr., MD, PhD



Renato Ambrosio Jr.
PROFESSOR, VISUAL OPTICS

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Post-LASIK Ectasia: Twenty Years of a Conundrum

Renato Ambrósio Jr. ^{1,2,3,4}

¹Department of Ophthalmology, Instituto de Olhos Renato Ambrósio, Rio de Janeiro, Brazil, ²Rio de Janeiro Corneal Topography and Biomechanics Study Group, Rio de Janeiro, Brazil, ³Department of Ophthalmology, Federal University of São Paulo, São Paulo, Brazil, and ⁴Department of Ophthalmology, Federal University of the State of Rio de Janeiro (UNIRIO), Rio de Janeiro, Brazil

ABSTRACT

Corneal ectasia has emerged as a serious complication of laser vision correction (LVC) procedures since the first report by Saito in 1988. Therefore, its prevention has become a major concern for refractive surgeons. Ectasia occurs due to biomechanical decompensation of the cornea, which may be related to a severe impact on corneal structure (i.e., attempted treatment for high myopia or the altered biomechanical properties preoperatively). The current understanding is that a combination from those factors determines stability or ectasia progression after LVC. Abnormal corneal topography has been the most important surrogate for lower biomechanical properties, but novel imaging technologies such as tomography and biomechanical assessment have proven to enhance the ability to detect mild ectasia, such as in the eyes with normal topography from patients with clinical ectasia in the fellow eye. Both are associated in a retrospective case series analysis data from 30,000 eyes from 16,720 documented eye eyes (80.9%) of seven patients that developed post-LASIK ectasia. This data supports the concept that the actual incidence of ectasia has decreased from 0.66% reported by Pallares in 2003. This has been the result of major development related to the advanced screening strategies. Nevertheless, mysterious cases of ectasia still challenge the field and stimulated research in this field. Ocular allergy and eye rubbing may be a factor that triggered ectasia in such series. Artificial intelligence (AI) and machine-learning algorithms may play a definitive role for further enhancing ectasia risk assessment. Reporting ectasia after LVC is needed.

Keywords: LASIK, ectasia, screening, risk, tomography

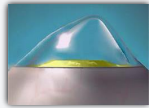
Renato Ambrósio Jr.
FEDERAL UNIVERSITY OF SÃO PAULO

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Pathophysiology of Ectasia

Biomechanical Decompensation of the Cornea

- Corneal Resistance (innate biomechanical properties)
- Impact from the environment
 - LVC procedures
 - Eye Rubbing



Any cornea may develop ectasia!

Renato Ambrósio Jr.
FEDERAL UNIVERSITY OF SÃO PAULO

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Violet June
THE KERATOCONUS

"Rubbing or scratching the eyes aggravates Keratoconus & may also cause Corneal Ectasia!"

The misinformation makes the patient suffer more than the disease."

Renato Ambrósio Jr., MD, PhD
Rio de Janeiro, Brazil

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What is Forme Fruste Keratoconus?

- Incomplete or abortive form of keratoconus (Amsler)
- "Mild" keratoconus (Krachmer)
- Fellow eye with no clinical signs from patients with very asymmetric (unilateral); Klyce, 2009)
- Fact: There is no consensus!
- Forme Fruste Keratoconus is defined as with **very high susceptibility** for ectasia progression (Ambrósio, 2009)

Ramato Ambrósio Jr
FUNDACION VITREO, MAQUILAS

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AMERICAN ACADEMY OF OPHTHALMOLOGY
 Proof of Participation: 2008 Annual Meeting and Subspecialty Day
 The following individual participated in the events listed below at the Academy's annual meeting and Subspecialty Day. The meeting was held November 7-12 at Georgia World Congress Center in Atlanta.

Ramato Ambrósio Jr MD
 REE12: Section VII: Business Strategies
 Participation: Moderator
 Presenter Time: 11:08:2008: 8:20 AM - 9:15 AM

COBB8: Elevation in the Ocular Way
 Participation: Presenter
 Presenter Time: 11:08:2008: 9:40 AM - 10:00 AM

REE18: Contact: Topographic and Biomechanical: New Concepts in Screening for Ectasia and its Susceptibility
 Participation: Paper Presenter
 Presenter Time: 11:08:2008: 4:14 PM - 4:19 PM

Very cordially,
David W. Pakis II
 David W. Pakis II, MD
 CEO
 American Academy of Ophthalmology

Ramato Ambrósio Jr
FUNDACION VITREO, MAQUILAS

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Ectasia Case Presentation

- Custom SBK-LASIK in 2008
- RSB=304µm; PTA=0.36
- Good vision in early postop, but developed progressive ectasia, confirmed at 14 months after LASIK




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Corneal Characterization


ARCHITECTURE or GEOMETRICAL CHARACTERIZATION (3D - TOMOGRAPHY)

↓


BIOLOGICAL PROPERTIES OF CORNEAL TISSUE



Leonardo Chelvarin, MD



Paulo Roberto Viegas



Rui Dreyfus, MD, PhD

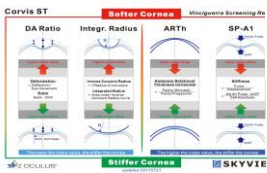
Corneal Wound Healing
Corneal Physical Properties (Biomechanics)

Paulo Roberto Viegas
FUNDADOR DO INSTITUTO DE OFTALMOLOGIA

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Dynamic Scheimpflug Imaging

- Oculus Corvis ST: Ultra High-Speed (UHS ST) Scheimpflug Technology taking 4,330 frames/sec with horizontal 8mm



Corvis ST

DA Ratio

Integr. Radius

ARTh


SP-A1

Stiffer Cornea | **SKYVIEW**

BIOMECHANICAL

Detection of Keratoconus With a New Biomechanical Index

Roberto Viegas, MD, Renato Ambrósio, Jr., MD, PhD, Aronoff Glazer, PhD, Carlos J. Simons, PhD, Bernardo Lopes, MD, Erasmo Henrique, PhD, Claudio Assini, MD, Paulo Viegas, MD



Paulo Roberto Viegas
ISRS Trondheim, Price: 2017


Paulo Roberto Viegas
FUNDADOR DO INSTITUTO DE OFTALMOLOGIA

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Evaluation of Corneal Shape and Biomechanics Before LASIK

Armedo R. et al. Int J Ophthalmol. 2014; 7(12): 2132-6.

- Renato Ambrósio, Jr. MD, PhD
- Leonardo P. Nguyen, MD
- Drogo L. Cordeiro, MD
- Bruno M. Fontes, MD
- Allen Lee, MD
- Jorge O. Costa, MD
- Victor Hsu-Avies, MD, PhD
- Michael W. Bell, MD, FACS



Rio de Janeiro
Corneal Tomography and
Biomechanics Study Group

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ARV
Ambrósio, Roberts & Vinciguerra
Tomography and Biomechanics Report

Random Forest
 "Leave one out" cross validation

TBI

Ambrósio, Roberts & Vinciguerra
 ASSOCIATION OF OPTOMETRIC RESEARCHERS

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(JRS, 2017) Integrated Scheimpflug Tomography and Biomechanics

Setting/Venues
 Humanitas Clinical and Research Center
 Italy
 Instituto de Olhos Renato Ambrósio
 Brasil

*94 VAE follow eyes criteria for normal topography

RISAN < 60
 I-S < 1.40 No PATHOLOG DETECTED
 Central K < 47D

100% sensitivity/100% specificity for frank ectasia (0.79)
 90.4% sensitivity/96% specificity for subclinical (0.29)

Ambrósio, Roberts & Vinciguerra
 ASSOCIATION OF OPTOMETRIC RESEARCHERS

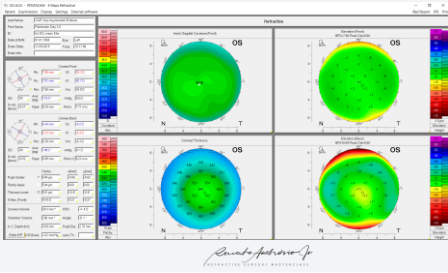
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Clinical Example
 masc, 53yo with interest in Refractive Surgery, reports gradual visual loss OD
 UDVA 20/60, J3 OD and 20/30, J3 OS
 MRx: -0.75 = -2.25 x 25° giving 20/20-1 OD // -0.50 = -0.75 x 120° giving 20/15 OS
 Add. +2.75, J1

Ambrósio, Roberts & Vinciguerra
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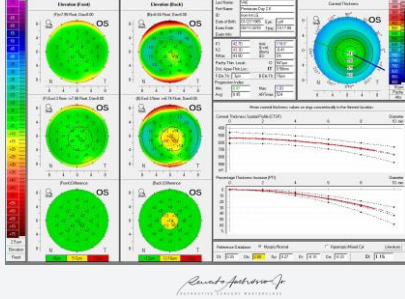
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Quad. Refractive Map



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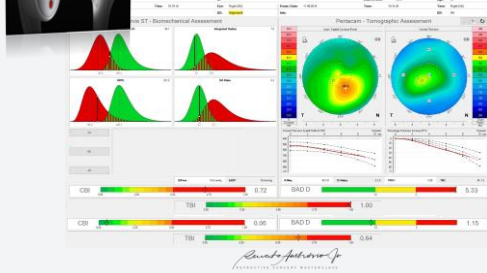
OCULUS - PENTACAM Belin / Ambrósio Enhanced Ectasia



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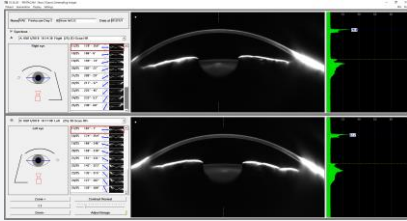
Integrated Tomography & Biomechanics

ST - Biomechanical/Tomographic Assessment (Ambrósio, Roberts & Vindiguerro)



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Lens Dysfunction Grade 2-3

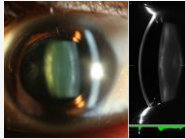


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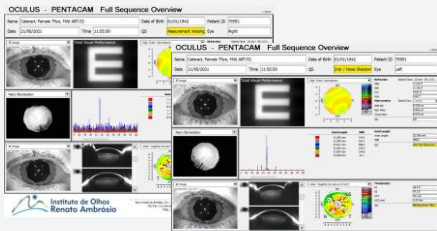
Lens Dysfunction Staging

- Stage I: early presbyopic (add up to +2.00D)
- Stage II: severe loss of accommodation (add >2.25D)
- Stage III: "mild cataract": loss of quality of vision *need for objective documentation
- Stage IV: loss of high contrast DCVA (Snellen <20/30)
- Stage V: moderate cataract with more significant loss of DCVA (Snellen <20/50)



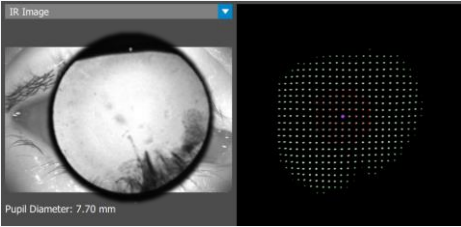
71

Female. 72 yo DCVA 20/30- OU



72

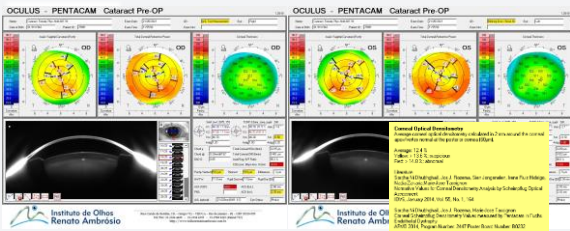
Wavefront Sensor & Retroillumination



Renzo Ambrosio
PENTACAM

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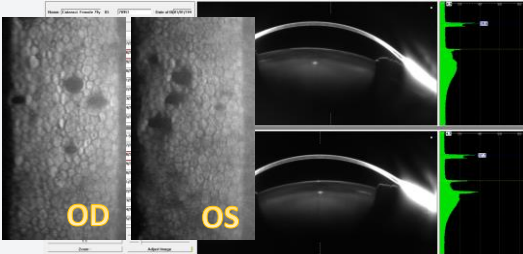
Female. 72 yo DCVA 20/30- OU



Renzo Ambrosio
PENTACAM

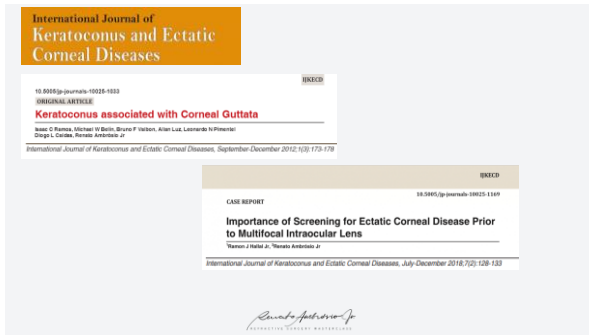
74

Female. 72 yo DCVA 20/30- OU

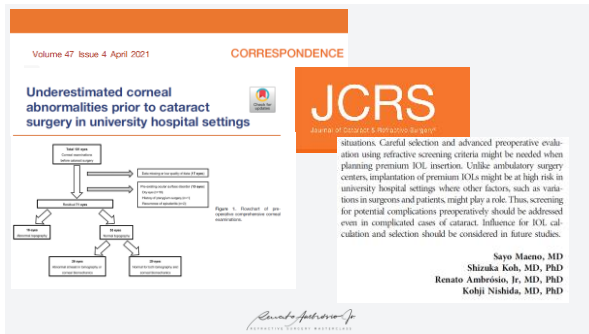


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PENTACAM

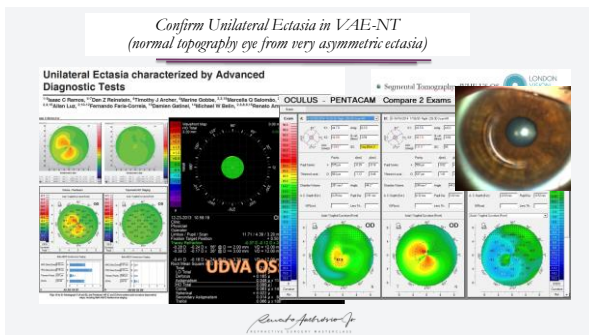
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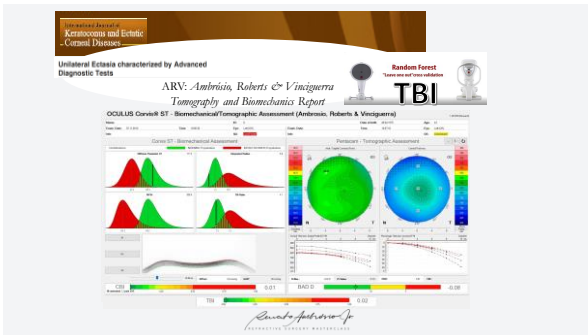


80

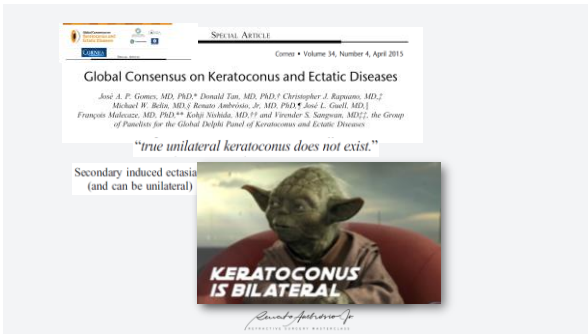


81

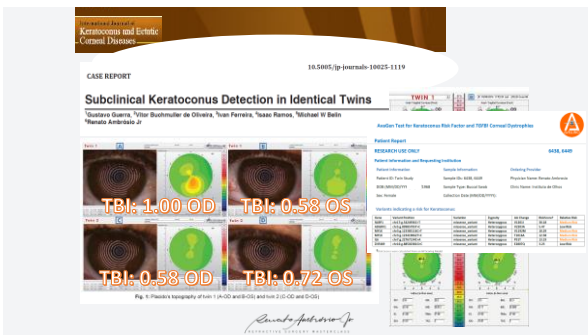




82

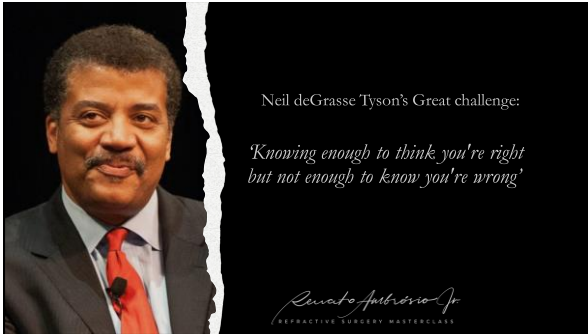


83



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Discussion: Recent TBI Studies

Eye and Vision

Biomechanical diagnostics of the cornea

| Author / Reference | NE | Chn | Est | Cut-off | Sensitivity | Specificity | AUC | VAE NT | Cut-off | Sensitivity | Specificity | AUC |
|--|-----|-----|-------|---------|-------------|-------------|-------|--------|---------|-------------|-------------|-------|
| Stromberg J / Refract Surg [1] | 105 | 95 | - | - | 98.00% | 100% | 0.998 | 327 | 0.11 | 72.00% | 71.00% | 0.825 |
| Karim R F / Contact Refract Surg [2] | 100 | 100 | >0.63 | - | 99.00% | 100% | 0.995 | 100 | >0.09 | 82.00% | 78.00% | 0.793 |
| Ferreira Mendonca J / Ann Ophthalmol [3] | 312 | 118 | 0.335 | - | 94.40% | 94.90% | 0.988 | 57 | 0.295 | 89.50% | 91.00% | 0.96 |
| Chan YC Y / Refract Surg [4] | 17 | 23 | - | - | - | - | - | - | 0.16 | 84.4% | 82.4% | 0.925 |
| Selgahat MB / Contact [5] | 137 | 145 | >0.49 | - | 100% | 100% | 1.000 | - | - | - | - | - |
| Koc M / Refract Surg [6] | 35 | - | - | - | - | - | - | 21 | 0.29 | 67.00% | 86.00% | 0.790 |
| Koh S / Refract Surg [7] | 70 | - | - | - | - | - | - | 23 | >0.259 | 52.17% | 88.57% | 0.751 |

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REFRACTIVE SURGERY MASTERSCLAS

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EURO TIMES
September 2020 | Vol 25 Issue 9

Artificial intelligence for KERATOCONUS DIAGNOSIS

Deep learning improves detection of mild ectasia
Flawed (Larkin reports)

Depth the accuracy of Keratoconus (KC) based on the detection of keratoconus using deep learning. The study was conducted by a team of researchers from the University of Illinois at Chicago (UIC) and the University of Michigan. The researchers used a deep learning algorithm to analyze a large dataset of corneal topography maps. The algorithm was able to detect KC with a sensitivity of 92% and a specificity of 98%. The researchers also found that the algorithm was able to detect KC in patients who had mild ectasia, which is a common finding in patients with KC. The researchers concluded that deep learning is a promising tool for the diagnosis of KC, particularly in patients with mild ectasia.


A revolution, in evolution...

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Methods: optimized TBI


- Multicentric retrospective study including twenty-five international centers
- Tomographic and biomechanical data from the Pentacam & Corvis ST (Oculus; Wetzlar, Germany).



| | n |
|-----------------|-------|
| * (1) CLIN NORM | 1,680 |
| * (2) KC | 1,181 |
| (3) VAE-E | 474 |
| (4) VAE-NT | 551 |

*one eye randomly selected


● **BAD-Dx3**
 cut off 1.98; Sensitivity=96.8%/Specificity=99.3% for KC + VAE-E
 cut off 1.27; Sensitivity=70.8%/Specificity=80.4% for VAE-NT



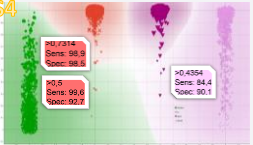
88

Results: RFS*4 X TBI (all cases)

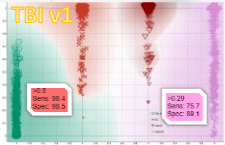
- Clin Ect. AUC: 0.999 x 0.999 (DeLong, $p=0.8180$)
- VAE-NT AUC: 0.945 x 0.899 (DeLong, $p<0.0001$)
- All cases AUC: 0.984 x 0.972 (DeLong, $p<0.0001$)




RFS4




TBI v1

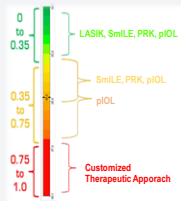




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Refractive Toolbox: Enhanced TBI






- 0 to 0.35: LASIK, SmILE, PRK, pIOL
- 0.35 to 0.75: SmILE, PRK, pIOL
- 0.75 to 1.0: Customized Therapeutic Approach

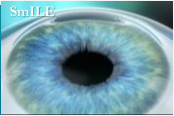
Surface Ablation



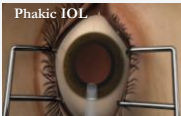
FemtoLASIK




SmILE



Phakic IOL





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Scheimpflug Imaging for Refractive Surgery

- Ancient Intelligence & Artificial Intelligence APPLIED: (A³I)²
- Treatment Planning (Customized) & Evaluation of Results
- Characterization of Ectasia Susceptibility
 - Screening, diagnosing, staging, prognosing, classifying, clinical follow-up for progression
- Characterization of Lens Dysfunction
- Refractive Imaging: *A true revolution, ... in evolution...*




Renato Ambrosio Jr.
 CONSULTOR SENIOR - OPORTUNIDADE

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Thank you, Obrigado!

dr.renatoambrosio@gmail.com

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Corneal OCT for Refractive & Cataract Surgeons

David Huang, MD, PhD
 Peterson Professor of Ophthalmology
 Professor of Biomedical Engineering
 Casey Eye Institute, Oregon Health & Science University
 Portland, Oregon

CASEY EYE Institute
OHSU

Financial Interests:
 OHSU and Dr. D. Huang have a significant financial interest in Optovue, a company that may have a commercial interest in the results of this research and technology. These potential conflicts of interest have been reviewed and managed by OHSU. Optovue, Inc.: patent royalty, equipment loan, stock ownership.

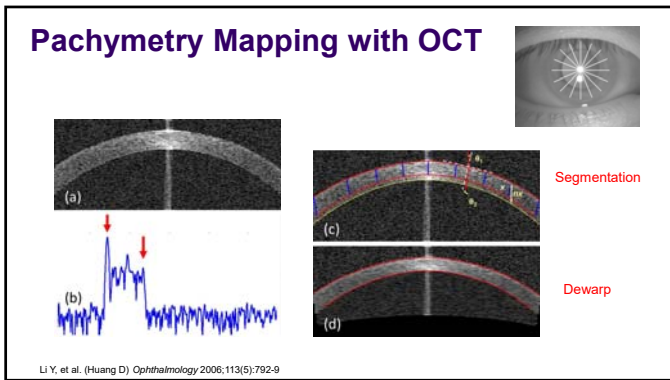
1

Corneal OCT Applications

CORNEAL PACHYMETRY AND EPITHELIUM MAPPING

2

Pachymetry Mapping with OCT



Li Y, et al. (Huang D) *Ophthalmology* 2006;113(5):792-9

3

OCT Pachymetry Agreement with Other Modalities

| N | OCT | OCT CCT (μm) | OCT- Other modalities (μm) | | |
|----|--------------------|--------------|--|----------------------------|---|
| | | | Scheimpflug (Pentacam [®] or Galle [®]) | Slit-scanning (Orbscan II) | Ultrasound (Sonogage [®] or Sonomed [®]) |
| 50 | RTVue ¹ | 536.9 | -- | -0.3±12.1 | -19.7±10.5 ^C |
| 66 | RTVue ² | 532.8 | -6.0±4.8 ^A | -- | -- |
| 50 | Casia ³ | 547.2 | -11.7±6.0 ^B | -7.2 | -9.2 ^D |

CCT = central corneal thickness

1. Li Y, et al. (Huang D) *J Cataract Refract Surg* 2010;36(5):826-831.
 2. Huang J, et al. (Wang Q) *PLoS One* 2014;9(5):e98316.
 3. Lee YW, et al. (Choi CY) *J Cataract Refract Surg* 2015;41(5):1018-1029.

4

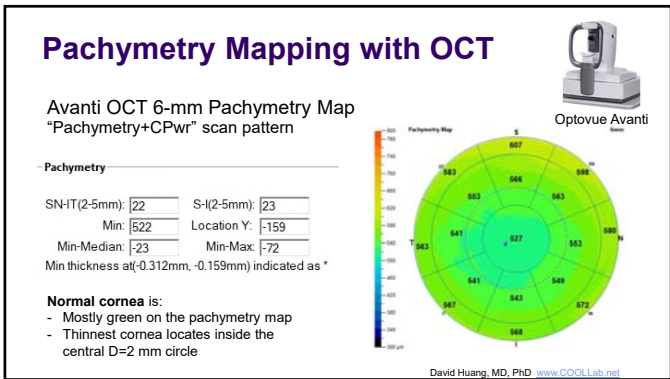
OCT Pachymetry Repeatability and Reproducibility

| | OCT | Central D<2mm | D=2~5mm |
|-----------------|---------------------|---------------|------------|
| Repeatability | RTVue ¹ | 1.3 μm | 1.8~3.8 μm |
| | RTVue ² | 2.1 μm | 2.9~5.5 μm |
| | Avanti ³ | 1.3 μm | 3.1~6.7 μm |
| Reproducibility | Casia ⁴ | 2.5 μm | 3.8~6.1 μm |
| | RTVue ⁵ | 2.1 μm | 3.6 μm |

1. Li Y, et al. (Huang D) *J Cataract Refract Surg* 2010;36(5):826-831.
 2. Huang J, et al. (Wang Q) *Ophthalmology* 2013;120(10):1951-1958.
 3. Unpublished data.
 4. Neri A, et al. (Neri A) *Acta Ophthalmol* 2012; 90:e452-e457.
 4. Prakash G, et al. (Agarwal A). *Am J Ophthalmol* 2009;148(2):282-290 e282.

5

Pachymetry Mapping with OCT



Optovue Avanti

Avanti OCT 6-mm Pachymetry Map
 "Pachymetry+CPwr" scan pattern

Pachymetry

SN-I(2-5mm): [22] S-I(2-5mm): [23]
 Min: [522] Location Y: [-159]
 Min-Median: [-23] Min-Max: [-72]
 Min thickness at(-0.312mm, -0.159mm) indicated as *

Normal cornea is:

- Mostly green on the pachymetry map
- Thinnest cornea locates inside the central D=2 mm circle

David Huang, MD, PhD www.COOLab.net

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OCT Keratoconus Pachymetric Parameters

| Parameter | Explanation |
|-----------|--|
| IT-SN | Average thickness of the IT octant minus that of the SN octant |
| I-S | Average thickness of the inferior (I) octant minus that of the superior (S) octant |
| Min | Minimum corneal thickness |
| Min - Med | Minimum corneal thickness - median corneal thickness |
| Y Min | Y coordinate of minimum corneal thickness |

1. Li Y, et al. (Huang D) *Ophthalmology* 2008;115(12):2159-2166.
 2. Qin B, et al. (Huang D) *J Cataract Refract Surg* 2013;39(12):1864-1871.

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OCT Pachymetry Based Keratoconus Risk Score

Keratoconus Risk Score Table
 Available for downloading @ <http://www.coolab.net/resources>

Patient Name: _____

| Variables (µm) | 0 | 1 | 2 | 3 | OD | OS |
|-------------------------------|-------|------------|-------------|--------|----|----|
| SN-IT | <33 | 33-42 | 43-51 | >51 | | |
| Minimum | ≥499 | 499-476 | 475-455 | <455 | | |
| Minimum-Median | ≥-21 | -21--25 | -26--29 | <-29 | | |
| S-I | <30 | 30-40 | 41-49 | ≥49 | | |
| Ymin | ≥-734 | -734--1069 | -1070--1353 | <-1353 | | |
| Keratoconus Risk Score | | | | | | |

Keratoconus risk:
 Keratoconus risk score 0-3: low risk, ≥4: high risk.

- Each variable will be assigned a score of 1, 2, 3 if it exceeds 20, 5, 1 percentile thresholds.
- The keratoconus risk score of the eye is the summation of all scores.

Qin B, et al. (Huang D) *J Cataract Refract Surg* 2013;39(12):1864-71.

8

Corneal Epithelial Imaging with FD-OCT

Li Y, et al. (Huang D) *Ophthalmology* 2012; 119(12):2425-33.

9

OCT Epithelial Thickness Map

iVue OCT 6-mm Epithelial Thickness Map
 "Cornea Pachymetry" scan pattern

| Epithelium | |
|---|--------------------|
| Epithelium statistics within central 5 mm | |
| S (2-5mm): | 51 I (2-5mm): 53 |
| Min: | 50 Max: 55 |
| Std Dev: | 1.4 Min-Max: -5 |
| Min/Max thickness indicated as +/- | |

Normal cornea is:

- Mostly green on the epithelial map
- Superior slightly thinner than the inferior
- Epithelial Std Dev < 3.6

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OCT Epithelial Map Repeatability

| OCT Device | Central D<2mm | D=2~5mm |
|------------------------|---------------|------------|
| RTVue ¹ | 0.7 µm | 0.7~1.1 µm |
| RTVue ² | 0.7 µm | 0.6~0.9 µm |
| Avanti ³ | 1.6 µm | 1.2~1.7 µm |
| Cirrus HD ⁴ | 1.5 µm | 1.3~1.5 µm |

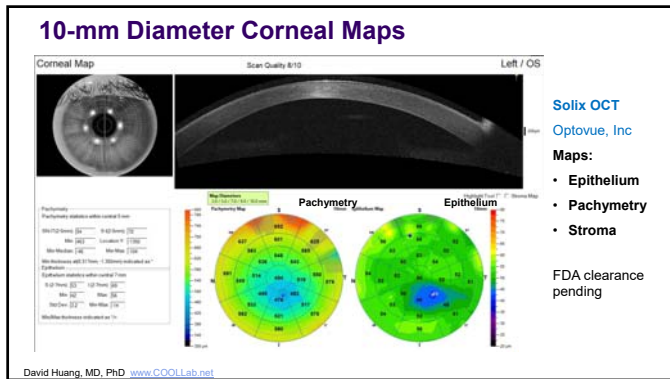
1. Li Y, et al. (Huang D) *Ophthalmology* 2012;119(12):2425-2433.
 2. Ma XJ, et al. (Koch DD) *Cornea* 2013;32(12):1544-1548.
 3. Hashmani N, et al. (Hashmani S) *Invest Ophthalmol Vis Sci* 2018; 59(3):1652-1658.
 4. Sha P, et al. (Durbin M) *Invest Ophthalmol Vis Sci* 2017; 58:3510

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Commercially Available Anterior Eye OCT

David Huang, MD, PhD www.COOLab.net * Pending FDA clearance for anterior eye imaging

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Yan Li, PhD Elias Pavlatos, PhD

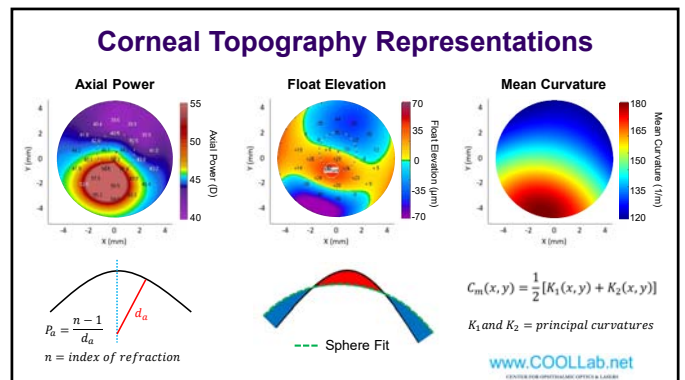
Corneal OCT Applications

CORNEAL TOPOGRAPHY WITH OCT

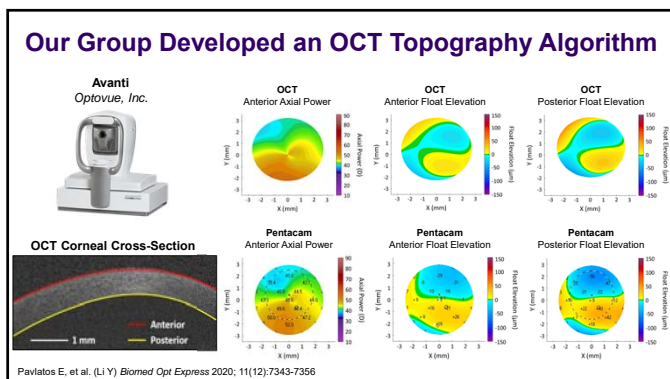
14



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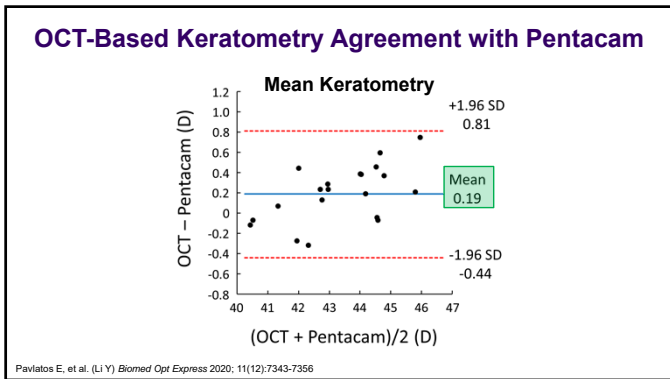
Motion Detection Improves Repeatability

| Diopters | No Motion Detection | Motion Detection |
|----------------------------------|---------------------|------------------|
| Anterior Mean Power | 0.28 | 0.14 |
| Anterior Astigmatism - Cardinal | 0.92 | 0.28 |
| Anterior Astigmatism - Oblique | 1.16 | 0.24 |
| Posterior Mean Power | 0.04 | 0.03 |
| Posterior Astigmatism - Cardinal | 0.13 | 0.05 |
| Posterior Astigmatism - Oblique | 0.15 | 0.05 |

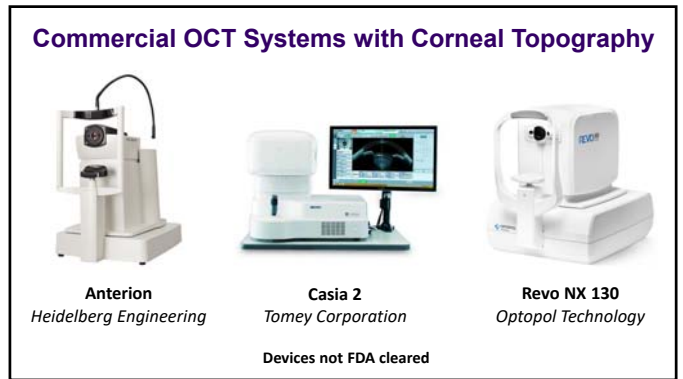
Pooled standard deviations for 20 eyes from 10 participants, 5 Repeated OCT Scans

Pavlatos E et al. *Biomed Opt Express*. 2020; 11(12):7343-7356

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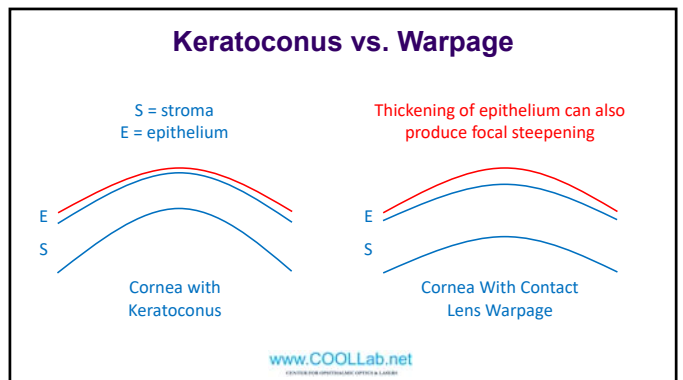
20

Yan Li, PhD Elias Pavlatos, PhD

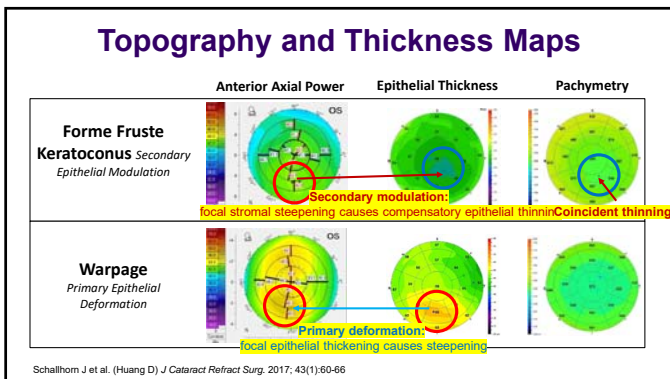
Corneal OCT Applications

DIFFERENTIATING KERATOCONUS FROM WARPAGE, DRY EYE, AND OTHER IRREGULARITIES

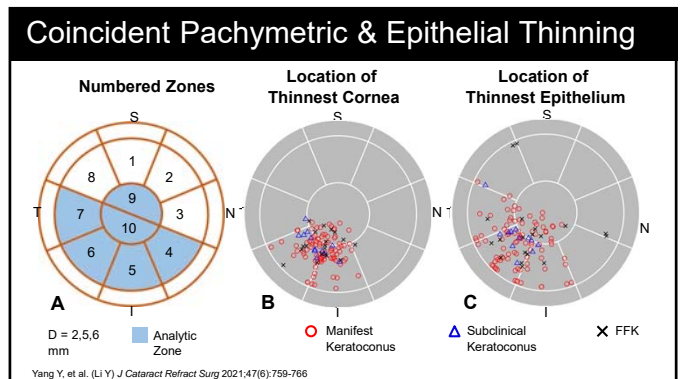
21



22



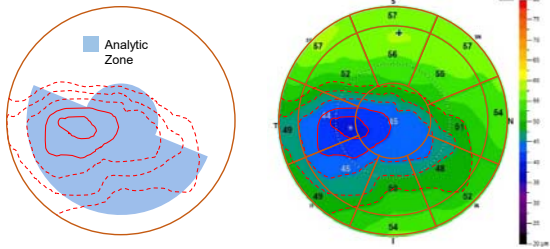
23



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Epithelial Concentric Thinning

- Epithelial thickness map has more than two color-scale step changes (>5 μm) inside the analytic zone
- At least one complete ring around the thinnest point

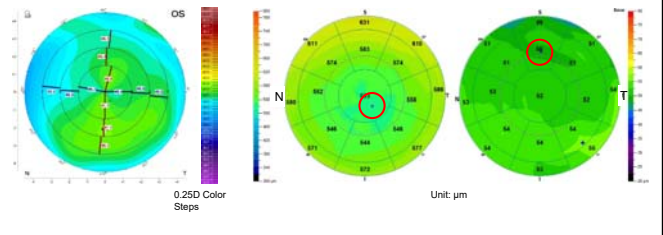


Yang Y, et al. (Li Y) J Cataract Refract Surg 2021;47(6):759-766

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Case 1 – Normal

Axial Power Pachymetry Epithelial Thickness

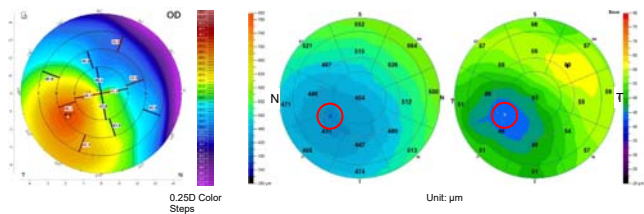


Yang Y, et al. (Li Y) J Cataract Refract Surg 2021;47(6):759-766

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Case 2 – Subclinical Keratoconus

Axial Power Pachymetry Epithelial Thickness

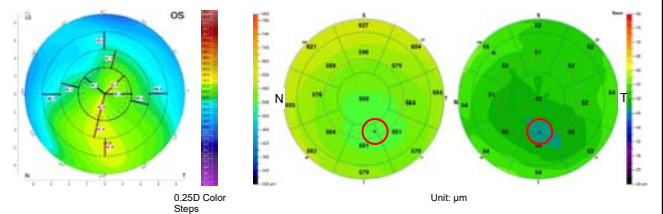


Yang Y, et al. (Li Y) J Cataract Refract Surg 2021;47(6):759-766

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Case 3 – Forme Fruste Keratoconus

Axial Power Pachymetry Epithelial Thickness



Yang Y, et al. (Li Y) J Cataract Refract Surg 2021;47(6):759-766

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Coincident Concentric Thinning Decision

Step 1
Map parameters

Pachymetric Min < 515 μm **OR**,
Pachymetric Min-Max < -71 μm **OR**,
Pachymetric SN-IT > 28 μm **OR**,
Epithelial Std Dev > 1.9 μm

YES
NO

Step 2
Map patterns

Coincident **AND**
concentric epithelial
thinning?

YES
NO

Not keratoconus

YES **Keratoconus**

NO **Not keratoconus**

Yang Y, et al. (Li Y) J Cataract Refract Surg 2021;47(6):759-766

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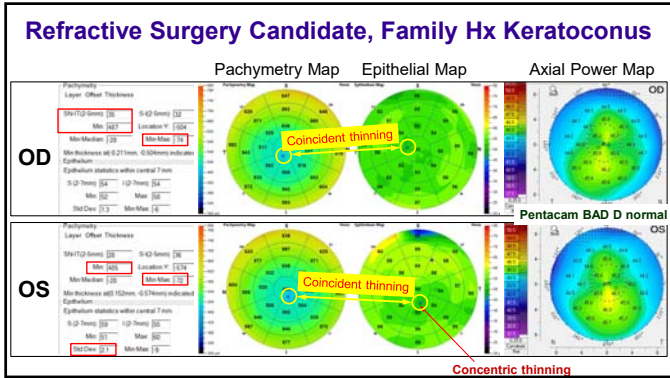
Classification Accuracy of Decision Tree

- **Manifest keratoconus** – abnormal topography and CDVA < 20/20
- **Subclinical keratoconus** – abnormal topography and CDVA ≥ 20/20
- **Forme Fruste keratoconus (FFK)** – fellow eyes of asymmetric keratoconus with normal or borderline topography and CDVA ≥ 20/20

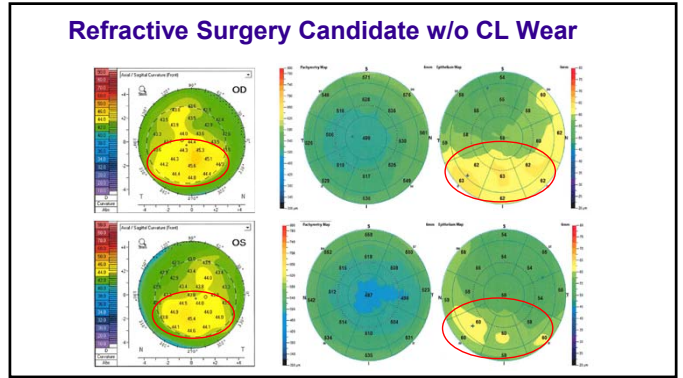
| Normal (54 eyes) | Manifest Keratoconus (91 eyes) | Subclinical Keratoconus (12 eyes) | Forme Fruste Keratoconus (19 eyes) |
|------------------|--------------------------------|-----------------------------------|------------------------------------|
| 100% | 97.8% | 100% | 73.7% |

Yang Y, et al. (Li Y) J Cataract Refract Surg 2021;47(6):759-766


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Yan Li, PhD

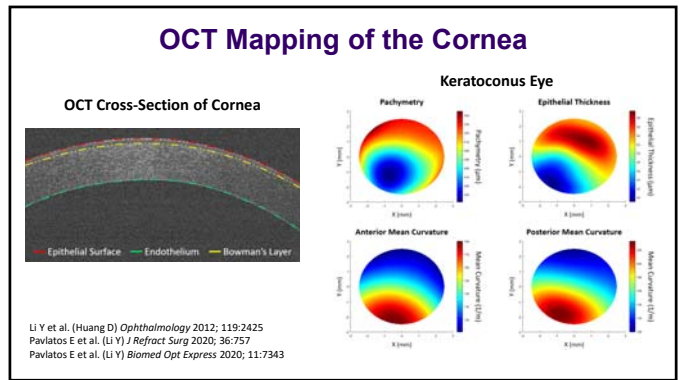


Elias Pavlatos, PhD

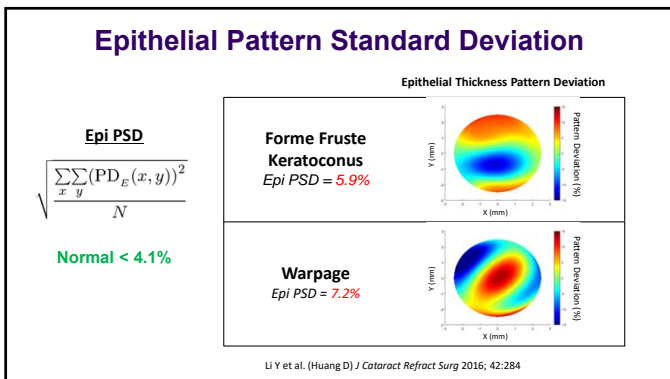
Corneal OCT Applications

QUANTITATIVE INDICES TO SEPARATE PRIMARY AND SECONDARY EPITHELIAL CHANGES

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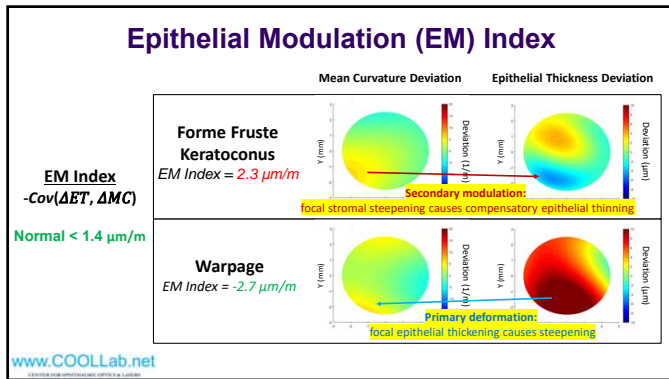
35

Epithelial PSD is a very sensitive detector of keratoconus

| | Keratoconus N= 35 eyes 150 control | 20/20 Keratoconus N=50 eyes 150 control | Forme fruste keratoconus N=8 150 control |
|-------------|--|---|--|
| Sensitivity | 100% | 96.0% | 87.5% |
| Specificity | 100% | 100% | 100% |

1. Li Y, et al. (Huang D) *Ophthalmology* 2012; 119:2425-33.
 2. Li Y, et al. (Huang D) *J Cataract Refract Surg* 2016; 42(2):284-95.

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Patient Groups

Keratoconus Groups

- Manifest Keratoconus (n = 89)**
 - Topographic signs of keratoconus
 - Corrected visual acuity < 20/20
- Subclinical Keratoconus (n = 16)**
 - Topographic signs of keratoconus
 - Corrected visual acuity $\geq 20/20$
- Forme Fruste Keratoconus (n = 26)**
 - Normal topography
 - Corrected visual acuity $\geq 20/20$
 - Keratoconus in fellow eye

Non-Keratoconus Groups

- Contact Lens Warpage (n = 18)**
 - Suspicious topographic steepening
 - Reduction in corrected visual acuity
- Normal (n = 32)**
 - Normal topography
 - Corrected visual acuity $\geq 20/20$

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Classification Accuracy of EM Index

Binary classification

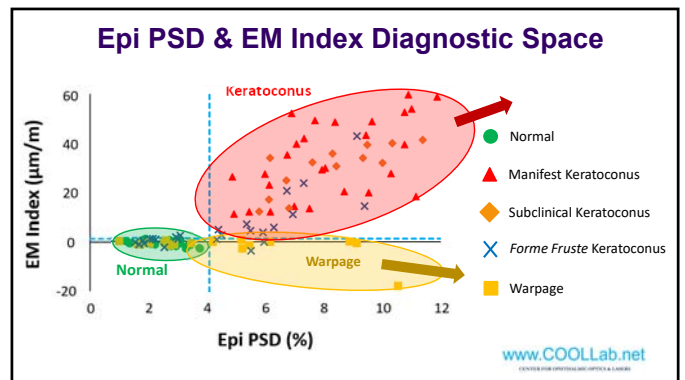
- Class 1 = non-keratoconus (normal and warpage)
- Class 2 = keratoconus (manifest, subclinical, forme fruste)

| EM Index | Classification Accuracy (%) | | | | |
|-----------------|-----------------------------|----------------|----------------------|-------------------------|----------------|
| Cutoff | Normal | Warpage | Manifest Keratoconus | Subclinical Keratoconus | FF Keratoconus |
| 1.39 ± 0.01 | 100 ± 0 | 98.9 ± 2.2 | 100 ± 0 | 100 ± 0 | 51.5 ± 1.9 |

• Cutoff determined at 50% probability by logistic regression
• 5-fold cross-validation repeated 5 times

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Concept Applies to Other Conditions

Primary Deformation


- Warpage
- Dry Eye
- EBMD

Secondary Modulation


- Keratoconus/ectasia
- Post-LASIK/PRK
- Stromal scars
- Stromal/Bowman dystrophies

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Yan Li, PhD

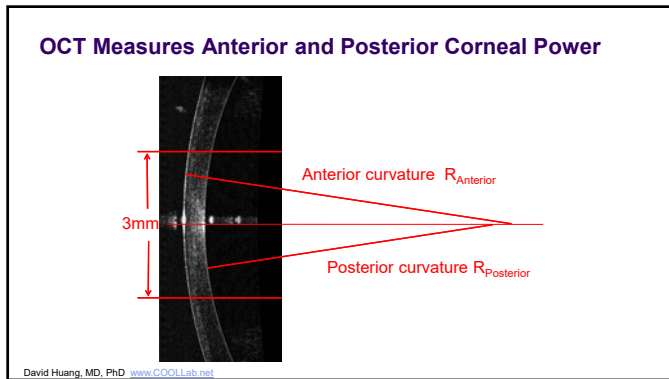


Clara Llorens Quintana, PhD

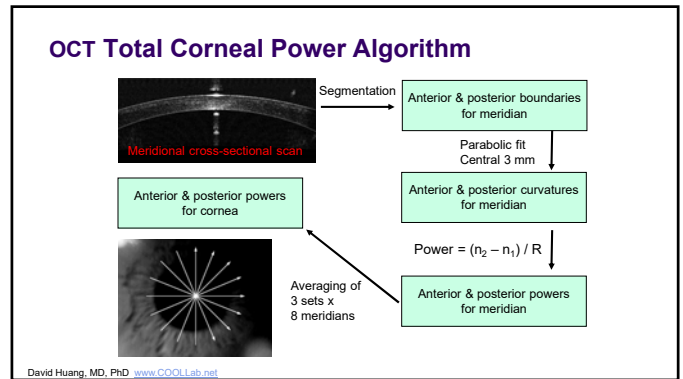
Corneal OCT Applications

TOTAL (NET) CORNEAL POWER AND ASTIGMATISM

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OCT Corneal Power Repeatability: Published Results

| | | RTVue ¹ | RTVue ² |
|-------------------|-------------|--------------------|--------------------|
| Corneal power (D) | Total (Net) | 0.19 | 0.10 |
| | Anterior | 0.19 | 0.11 |
| | Posterior | 0.02 | 0.02 |

1. Tang M, et al. (Huang D) *J Cataract Refract Surg*. 2010;36(12):2115-2122.
 2. Wang Q, et al. (Huang J) *Cornea*. 2015;34(10):1266-1271.

45

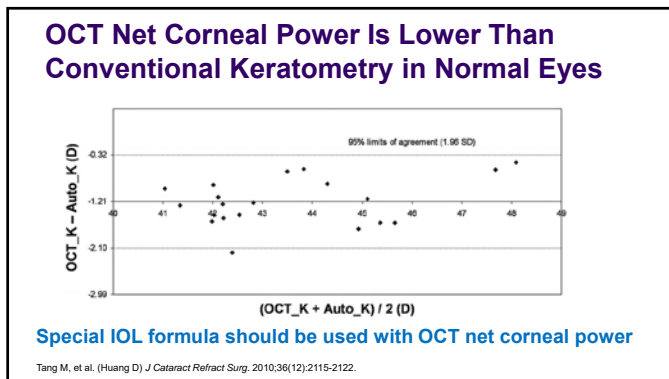
Repeatability: RTVue vs Avanti

| | | RTVue | Avanti |
|-------------------|-------------|-------|--------|
| Corneal power (D) | Total (Net) | 0.18 | 0.14 |
| | Anterior | 0.20 | 0.15 |
| | Posterior | 0.04 | 0.04 |

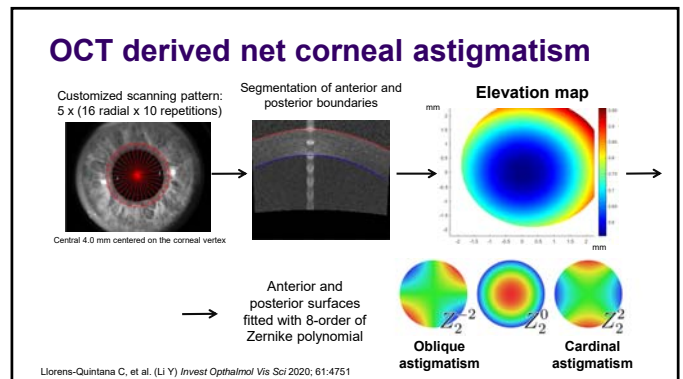
n = 24 eyes of 12 normal subjects

David Huang, MD, PhD www.COOLab.net

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Subjects

60 pseudophakic eyes (39 patients)

- With non-toric monofocal IOL
- Post Laser visual correction

↓

Manifest refraction astigmatism = ground truth

Llorens-Quintana C, et al. (Li Y) *Invest Ophthalmol Vis Sci* 2021;62:2026

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Net Corneal Astigmatism Repeatability

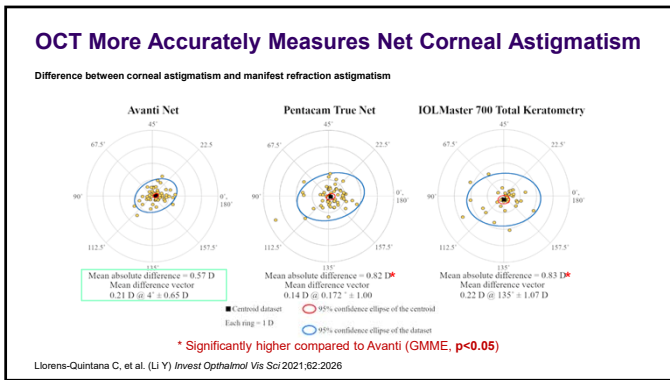
Coefficient of repeatability
Pooled standard deviation* 1.96 * sqrt(2)

| | Avanti* | Pentacam | Significance** |
|--------------|---------|----------|----------------|
| Cardinal (D) | 0.22 | 0.50** | $p < 0.05$ |
| Oblique (D) | 0.19 | 0.44** | $p < 0.05$ |
| Vector (D) | 0.29 | 0.67** | $p < 0.05$ |

* Experimental software, not FDA-cleared
** F test comparison with OCT Net astigmatism

Llorens-Quintana C, et al. (Li Y) *Invest Ophthalmol Vis Sci* 2021;62:2026

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OCT Corneal Mapping

- Four maps
 - Epithelial thickness
 - Pachymetry
 - Anterior topography
 - Posterior topography
- Distinguish ectasia from warpage
- May provide more accurate net corneal astigmatism measurement in aberrated corneas
- Advanced features still FDA pending

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Grants & Material Supports

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Unrestricted grant from Research to Prevent Blindness

Material support from Optovue, Inc.

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Jessica Mathum, Lab Manager

Loki Mathum, Events Manager

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Measuring the cornea for IOL power calculations

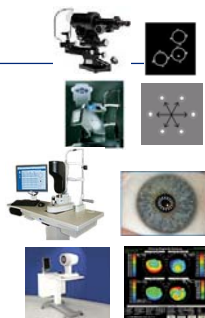
Li Wang, MD, PhD,
Cullen Eye Institute
Baylor College of Medicine,
Houston, Texas, USA

Consultant:
Alcon
Carl Zeiss Meditec

1

Corneal power measurements

- ✦ Range of devices:
 - ✦ 4 points: keratometer
 - ✦ 6-32 points: ocular biometers
 - ✦ >500 points: topographic / tomographic values averaged over the central 3-4 mm zone



2

Corneal power measurements

- ✦ Most of the commonly used devices measure only the anterior corneal curvature
- ✦ Total corneal power calculated
 - ✦ Assume constant ratio of posterior to anterior corneal curvature
 - ✦ Standard index of refraction 1.3375 (USA) or 1.332 (Europe)

3

Ratio of posterior/anterior radii of curvature from Galilei*

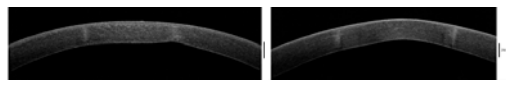
| | Ratio | "Standardized" index of refraction | Ratio in RK eyes (n=114) with Avanti OCT Mean: 0.94 Range: 0.66 – 1.30 |
|----------------------------|-------|------------------------------------|--|
| Gullstrand's schematic eye | 0.883 | 1.3325 | |
| Normal (n=94) | 0.816 | 1.3278 | |
| Myopic-LASIK/PRK (n=61) | 0.765 | 1.3246 | |
| Hyperopic-LASIK/PRK (n=9) | 0.857 | 1.3302 | |

* Wang L, Mahmoud AM, Anderson BL, Koch DD and Roberts CJ. IOVS 2011; 52:1716-22

4

Ratio of posterior/anterior radii of curvature

- ✦ Effect on calculation of total corneal power
 - ✦ Small in normal eyes
 - ✦ Key issue in post-refractive eyes



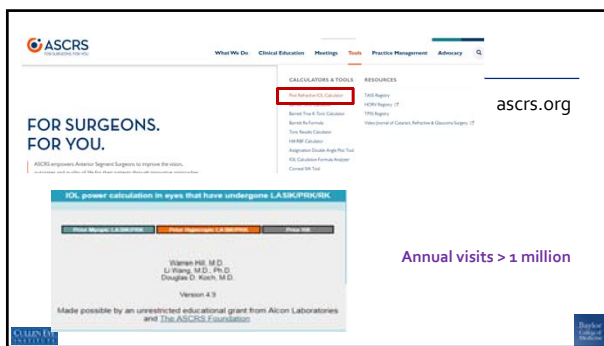
Posterior corneal power: -3.5 D

5

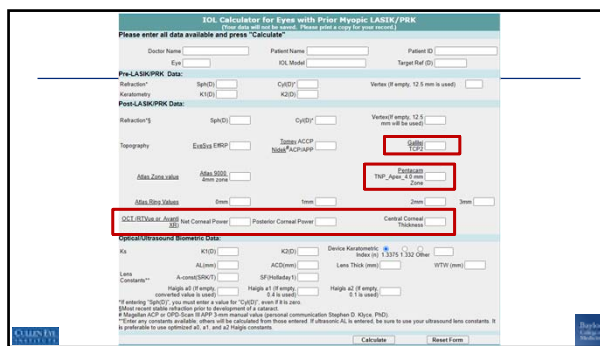
IOL power calculation in post-refractive eyes

- ✦ Traditionally using corneal power based on anterior corneal surface
- ✦ Formulas/methods using data from both anterior and posterior corneal surfaces emerging
 - ✦ OCT-based IOL calculation formula
 - ✦ Galilei TCP formula
 - ✦ Potvin-Hill Pentacam
 - ✦ Ray-tracing

6



7



8

What is ideally the best IOL formula?

- Ray tracing:
 - Incorporates all surfaces and aberrations
 - Cornea
 - IOL
 - Major limitation
 - Does not provide a solution to ELP
 - Outcomes with ray tracing formulas to date are not better than our other best formulas

9

Outcomes reported in literature: recent large studies including newer formulas

| | % ± 0.5 D |
|--------------------------|-----------|
| Myopic LASIK/PRK eyes | 40% - 85% |
| Hyperopic LASIK/PRK eyes | 38% - 73% |
| RK eyes | 29% - 62% |

- Reported more accurate formulas:
 - Average IOL power, OCT-based, Barrett True K, Haigis-L, Maskit

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Current best practices

- Use as many formulas as possible
- Rely more on
 - Average IOL power
 - OCT-based, Barrett True K No History, and Haigis-L
- Warn patients of inaccuracy and possible additional surgeries and costs

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"Holy Grail": postop adjustable IOL?

- RxSight: light adjustable lens
 - Curvature change
- Perfect Lens and Clerio
 - Localized refractive index change

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Posterior corneal astigmatism

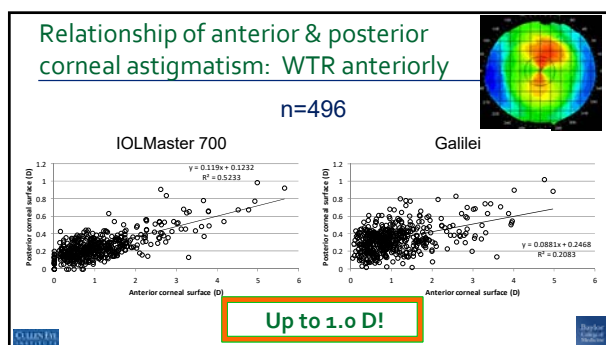
13

Magnitude of posterior corneal astigmatism

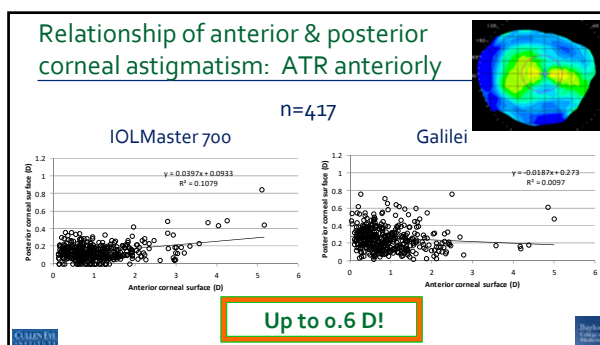
- ◆ Our paper: 0.30 ± 0.15 (0 – 1.10) (n = 715)
- ◆ Tonn...Kohlen: 0.33 ± 0.18 (0 – 1.35) (n = 3818)

Koch et al. JCRS 2012; 38:2080-2087
Tonn B, Klaproth OK, Kohlen T. IOVS 2014; 56:291-8

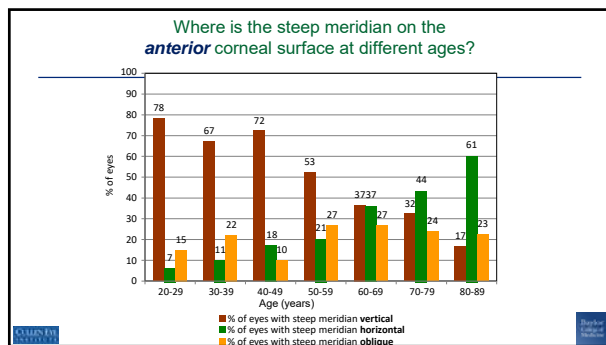
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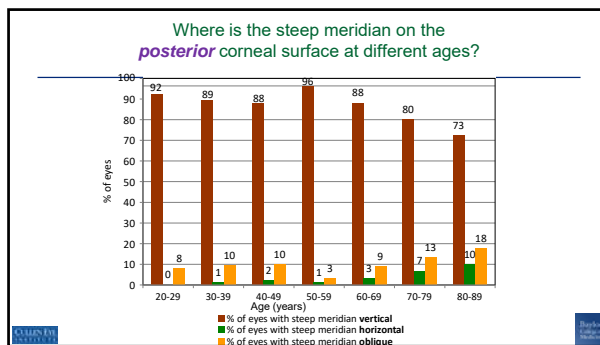
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Toric IOL calculation: how do we incorporate posterior corneal astigmatism (PCA)?

- Use regression/theoretical models
 - Baylor nomogram
 - Abulafia-Koch: Vector version of Baylor nomogram + clinical data
 - J&J (AMO): J&J clinical trial data + Baylor nomogram
 - Barrett toric calculator (standard, predicted PCA)
- Measure the posterior cornea
 - Barrett toric calculator (new, measured PCA)

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Baylor Toric IOL Nomogram, Version 2

Temporal clear corneal incision and target for postop astigmatism of 0.4 D WTR to account for ATR shift with age

| Effective IOL cylinder power at corneal plane (D) | WTR (D) | ATR (D) |
|---|--------------------|--------------|
| 0 | ≤ 1.69 (>1.0: PCR) | ≤ 0.39 |
| 1.00 | 1.70 - 2.19 | 0.40* - 0.79 |
| 1.50 | 2.20 - 2.69 | 0.80 - 1.29 |
| 2.00 | 2.70 - 3.19 | 1.30 - 1.79 |
| 2.50 | 3.20 - 3.79 | 1.80 - 2.29 |
| 3.00 | 3.80 - 4.39 | 2.30 - 2.79 |
| 3.50 | 4.40 - 4.99 | 2.80 - 3.29 |
| 4.00 | 5.00 - | 3.30 - 3.79 |

*Especially if specs have more ATR

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Barrett toric calculator (apacrs.org)

21

Problem with regression approaches for selecting toric IOLs: Scatter

- Wide individual variation
- Even "normal" corneas

22

Does measuring the posterior cornea improve toric outcomes?

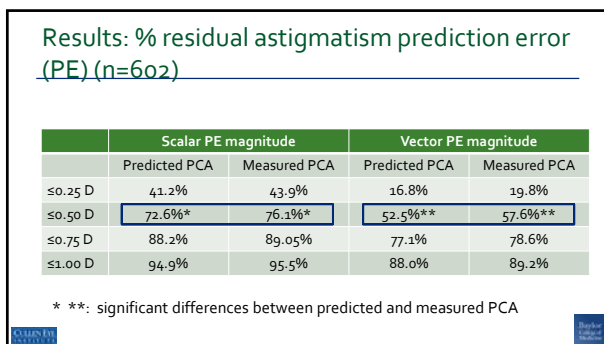
- Prior Scheimpflug studies:
 - No benefit to direct measurement
- Studies:
 - Regression models vs. direct measurements
 - More accurate with regression models

23

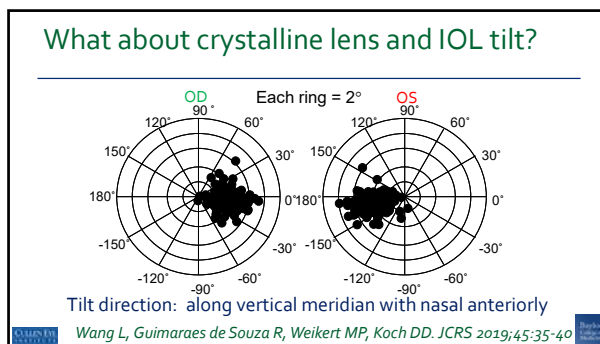
Our study

- To compare the accuracy of predicting residual astigmatism after cataract surgery using Barrett toric calculator with:
 - Predicted PCA
 - Measured PCA from IOLMaster 700
- Dataset from VERACITY surgical database
 - Included eyes with monofocal non-toric IOLs

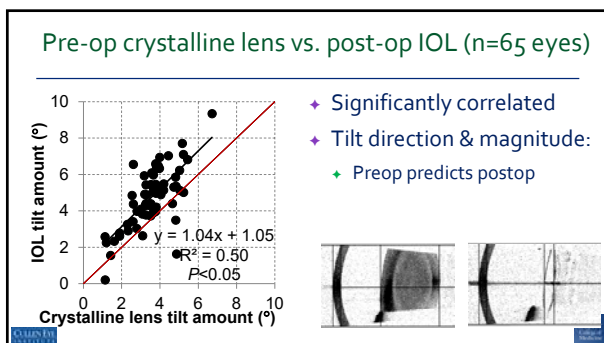
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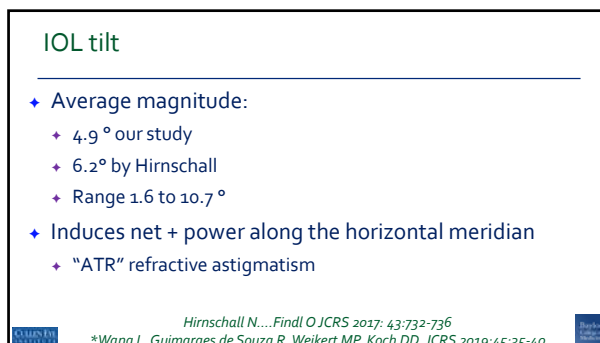
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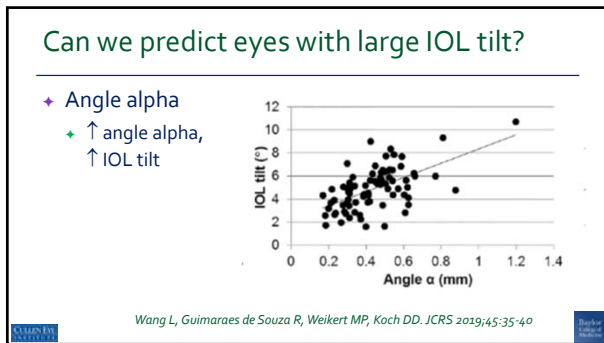
26



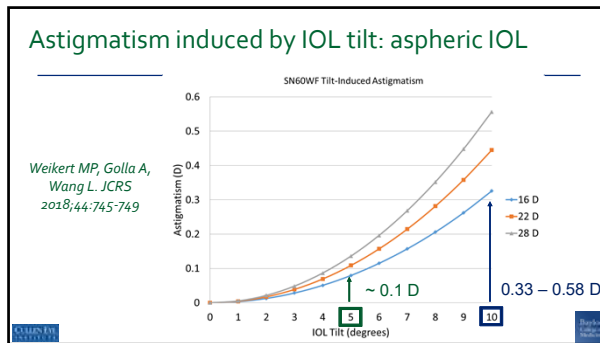
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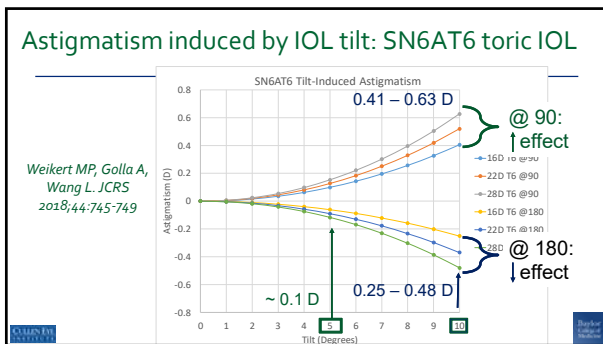
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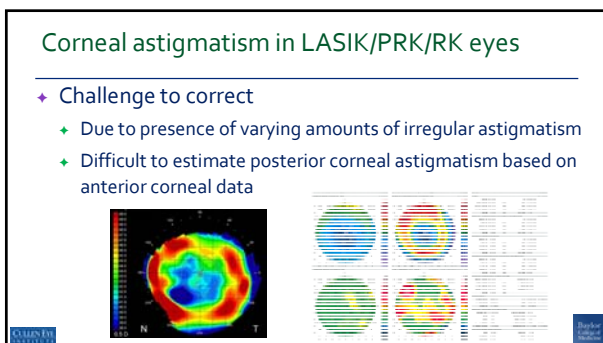
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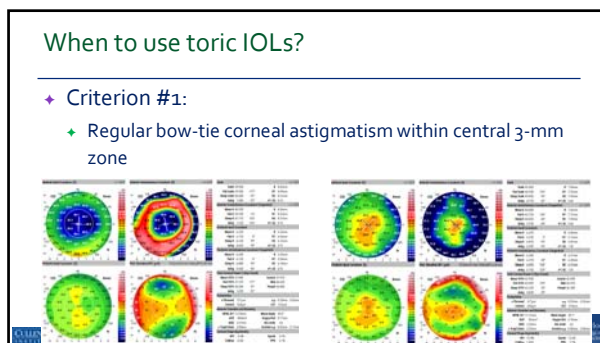
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- ### Toric IOL in LASIK/PRK/RK eyes
- ✦ High expectations following cataract surgery
 - ✦ Good uncorrected visual acuity
 - ✦ Spectacle independence
 - ✦ Corneal astigmatism common
 - ✦ LASIK/PRK performed to eliminate ocular refractive error including astigmatism
 - ✦ Residual or induced to compensate for lenticular astigmatism

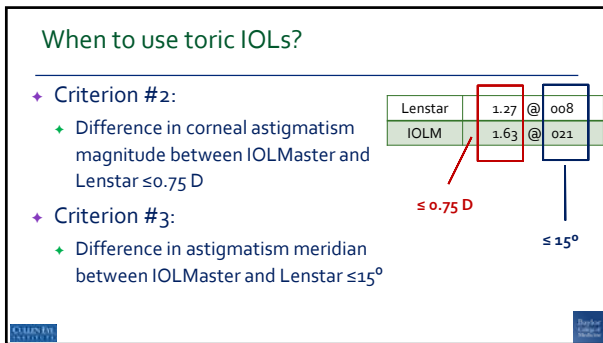
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- ### IOL toricity selection
- ✦ Posterior corneal astigmatism
 - ✦ Difficult to estimate
 - ✦ Estimate 0.3 D of against-the-rule refractive effect
 - ✦ Target for corneal astigmatism correction
 - ✦ 0.3 D WTR
 - ✦ Be conservative and not over-correct!

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IOL spherical power selection: ascrs.org

Post-refractive IOL calculator

Version 4.3
Made possible by an unrestricted educational grant from Alcon Laboratories and The ASCRS Foundation

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When corneas met all 3 criteria:

- % of eyes with refractive astigmatism ≤ 0.5 D
 - Post-myopic LASIK (n = 56): 80%
 - Post-hyperopic LASIK (n = 19): 84%
 - Post-RK (n = 41): 76%

Cao D, Wang L, Koch DD. JCRS . 46(4):534-539, 2020.

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Barrett True-K toric calculator

Myopic LASIK
2D @ 90
Net astigmatism: 1.39D@92

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Case #1

- 75 yo male
- s/p LASIK
- Cataract OS

40

Preop corneal astigmatism

| | |
|---------------|-------------------|
| MR | -0.75 x 1.0 x 175 |
| Lenstar | 1.29 @ 169 |
| IOLMaster 700 | 1.30 @ 169 |
| Galilei | |
| SimK | 0.99 @ 159 |
| TCP | 1.37 @ 165 |

- 19.0 D ZCT225@172
- POM#1
- UCVA = 20/20
- MR: plano

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Case #2: previous RK

- 60 yrs male cataract surgery OS
- MR: + 1.75 + 0.75 x 166
- 8 radial cuts and one superior T cut
- Target for D

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Corneal astigmatism

- + MR: $+1.75 + 0.75 \times 166$
- + IOLMaster: 0.93 D @150°
- + Lenstar: 1.04 D @149°
- + Galilei SimK: 1.01 D @157°
- + Galilei TCP: 1.19 D @173°
- + Atlas: 0.76 D @177°

Toric or nontoric IOL? ZCT150 @160°

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IOL spherical power: which would you choose??

Calculator formulas for target -0.5 D:
25.71 to 27.45 D

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26.5D ZCT150 @160 implanted targeting -0.50

POM #1:
UCVA = 20/100
-1.75 Sph = 20/20

POM #2:
UCVA = 20/50
-1.25 Sph = 20/20

POM #3:
UCVA = 20/70
-1.75 Sph = 20/25

What would you do now?

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Fit with -1.25 D CL

- + Realized how nice a little near vision is
- + Wants perfect distance vision OD!
- + It has 12 radial and 8 T cuts....

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Summary

- + Accurate total corneal power estimation is crucial
 - + Posterior corneal power measurements improve accuracy, especially in un-usual eyes
- + IOL power calculation in post-refractive eyes still a ways to go
 - + Especially in RK eyes
 - + More accurate corneal power measurements and IOL power formulas are needed

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Summary

- + Considering posterior corneal astigmatism in toric IOL selection improves accuracy
- + Toric IOLs can work well in post-refractive eyes
 - + Corneas met all 3 criteria
- + Postop IOL power adjustment is very promising

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