### Corneal biomechanics Mohamed Elhaw, MD, AL-Azhar university

### Aiming to $\rightarrow$

Understanding biomechanical properties and definitions

**Biomechanics of human cornea** 

Methods used for corneal biomechanics assessment

**Clinical application of biomechanics** 

#### **BIOMECHANICAL** DEFINITIONS

• The response of a biological tissue to a force.

• Depends on the **biochemical and physical nature** of the components present and their relative amounts.

# Elasticity





### Deforming Force

# Viscosity

## Stiffness





### Stiffness



# The cornea is considered to be viscoelastic Stiff but, still flexible

#### FACTORS DETERMINING CORNEAL BIOMECHANICAL PROPERTIES

#### Extracellular matrix components

glycoproteins, collagen and enzymes, providing biochemical and structural support to the surrounding cells

Collagen and elastin are responsible for the strength and elasticity of a tissue, while the ground substance is responsible for the viscoelastic properties

#### Hydration

- Environmental factors
- Hormonal fluctuations
- others



The mechanical structure of the cornea determines corneal integrity and shape...



#### ... and therefore optical and visual quality

# AVAILABLE INVESTIGATIONS FOR CORNEA

#### Placido-based TMS 5 (Tomey)



Placido-based Only anterior corneal surface Corneal tomography Both anterior & posterior corneal surfaces

#### Slit-scanning Orbsan (Bausch + Lomb)



#### Scheimpflug-based Pentacam HR (Oculus)

OCULUS - PENTACAM 4 Maps Selectable

#### OCT-based CASIA SS-2000 (Tomey)



#### METHODS USED FOR CORNEAL BIOMECHANICS ASSESSMENT

Technique		In-vivo/ex- vivo	Measurement variables	5	Detection of biomechanical change
St	ODA				Stress-strain behavior of cornea
Int	ORA				Corneal stiffness after CXL
Sp int					Corneal deformation, IOP detection
OF				nd	Parameters for detecting sub clinical keratoconus
СС	CORVI	SSI		ation	Detects ectatic diseases
Gc to				g force	IOP detection
Brillouin spectroscopy		In-vivo	3D Brillouin modulus		Brillouin modulus changes after CXL and in keratoconus



#### and dynamic corneal deformation imaging by air puff in vivo



# ORA

#### **Corneal hysteresis**

is the difference between the P1 and P2 values

#### corneal resistance factor

#### calculated according to the formula: **a** [P1-0.7P2] + **d**

where *a* and *d* are calibration and regression constants to maximize correlation with central corneal thickness



#### Ocular Response Analyzer

#### Classifying Corneal Pathologies



Thin Cornea with no ectasia



**Thin Cornea with Keratoconus** 



CH=11.2 CRF=10.8



CH=8.1 CRF=7.9

#### Ocular Response Analyzer

#### Pre / Post Lasik



#### Ocular Response Analyzer

#### Predicting Ectasia Risk

Pre & Post LASIK CRF with KC subjects



#### EVEN THOUGH STUDIES HAVE REPORTED CH AND CRF TO BE LOWER IN KC COMPARED TO HEALTHY CORNEAS

A CONSIDERABLE OVERLAP IN THE DISTRIBUTIONS OF BOTH PARAMETERS WAS OBSERVED

THE SENSITIVITY AND SPECIFICITY FOR KC DIAGNOSIS ARE RELATIVELY WEAK

# SO, WENEED MORE PARAMETERS

### CorVis ST

#### What is it?

- Air Tonometer
- Ultra high speed Scheimpflug camera

#### How does it work?

- Air puff indents cornea
- Video 4333 frames/sec
- Calculates a range of output measurements
- Parameters describe *in vivo* biomechanics



#### **Corvis ST – Parameters**

1st Appla	air puff (in milliseconds). The length of the applanation at this moment appears in parenthesis (in millimeters).					
Highest IST Applanation	mum concavity during the air puff (in milliseconds). The length of the distance between the two peaks of the i (in millimeters).					
2nd App 2nd Applanation	r puff (in milliseconds). The length of the applanation at this moment appears in parenthesis (in					
Maximur 2110 Applallation	a deformation during the air puff.					
Wing Dis						
Maximur HIGNEST CONCAVITY 50	(in meters per seconds [m/s]).					
Maximur	phase (in meters per seconds [m/s]).					
Curv Integrated Inverse Dedius						
Curv IIIEgidleu IIIVEISE Raulus ng the air puff (in millimeters).						
Cornea Thickness Measurement of the corneal thickness (in millimeters).						
Deformation Amplitude Patio 1						
Defo or 2						
IOP OF Z MM						
bIOP Biomechanically-corrected IOP						

### OCULUS Corvis® ST

Viewable biomechanical analysis with more than 4,000 images per second.





 Figure 1. Schematic diagram of air puff loading of the cornea with red arrows indicating direction of movement. The initial phase is Baseline Convex prior to deformation. The Loading Phase includes deformation through Inward Applanation (flattening) and into Concave phase in the Inward direction. The Unloading Phase includes initiation of recovery from Concave phase in the Outward direction through Outward Applanation to the Convex shape Recovered.

### CorVis ST - Parameters







- Applanation1
  - Time
  - Length
  - Velocity
- Highest Concavity
  - Deformation Amplitude
  - Radius of Curvature
  - Peak Distance
- Applanation2
  - Time
  - Length
  - Velocity

### Dynamic corneal Response Parameters

- Deformation Amplitude
- DA ratio 2mm
- Integrated Radius

• **CBI\_LVC**: Corneal stability after laser vision correction







#### Response to an air-pulse is not purely corneal



Deformation response to an air-pulse includes: corneal deformation and the globe being pushed into the socket

### **CORVIS ST - DCR - DA ratio 2 mm**



### **CORVIS ST - DCR - IR**



The central Radius of curvature is calculated. The inverse Radius (1 / R) is calculated and the area under this inverse Radius vs. time curve is determined. This area is called integrated Radius and is a very good parameter to quantify the effect of corneal cross-linking. If this parameter gets smaller it indicates a stiffening of the cornea.

### **CORVIS ST** — stiffness parameter (SP)

# Stiffness = force / displacement d T
#### CORVIS ST — stress strain curve SST



Value:	0.8		0.9
SD:	2.2		1.7
Diff. SD:	-0.5		
Change:	softer	not sign.	stiffer



CORVIS ST – CB

#### **OCULUS Corvis® ST - Vinciguerra Screening Report** 1.6(2031 Age: 35 Name: 00\_Demo\_Corvis, FFKC ID: monolateral kc OS Date of birth: 16.02.1980 Exam. Date: 10.11.2015 Time: 16:49:32 Left (OS) Eye: OK Info: QS: Deformation Amplitude Ratio (2mm) **Deformation Amplitude** Deformation Amplitude vs. bIOP DA Ratio vs. time DA Ratio vs. bIOP **Deformation Amplitude vs. time** [mm] [mm] [1] [1] A1 AT HC A2 HC A2 1.50 1.50 6.00 1.20 mm 5.67 6.00 1.00 1.00 4.00 4.00 0.50 0.50 2.00 2.00 0.00 0.00 20 30 ma 10 15 20 10 20 30 ms 10 15 25 mmHg 10 20 25 mmHg Inverse Concave Radius **Deflection Area** ~ Deflection Area vs. time Deflection Area vs. bIOP Inverse Concave Radius vs. time Inverse Concave Radius vs. bIOP [mm<sup>2</sup>] (mm ') (mm ') A1 A1 HC A2 HC A2 0.20 0.20 4.15 mm<sup>2</sup> 0.15 mm<sup>-1</sup> 4.00 4.00 3.00 3.00 0.10 - 0.10 2.00 2.00 1.00 1.00 0.00 0.00 30 ms 20 10 20 10 15 20 25 mmHg 10 30 ms 10 15 20 ŝ 25 mmHg Integr. Radius ARTh SP-A1 CBI SSI **DA Ratio** A1 N +3 SD -+3 SD +4 SD +3 SD 1.00 SP-A1: 73.2 0.84 - 5.7 -+2 SD +3 SD +2 SD +2 SD -- 73.2 -0.16 ARTh 356.2 - 356.2 -0.75 Stress [MPa] +2 SD +1 SD + +1 SD +1 SD 9.1 -HC **IOP** +1 SD 0 SD 0 SD 0 SD 0.50 IOPnct: 12.5 mmHg -1 SD 0 SD -1 SD --1 SD 0.25 DIOP: 13.8 mmHg -1 SD -2 SD -2 SD -2 50 0.04 A2 -2 SD -3 SD -3 SD -3 SD 0.00 Pachy Strain 0.00 OCT: . 0.00 ms 484 µm SD: SD: 1.65 SD: 2.05 CBI: 0.96 0% 2% 4% 6% 3.29 SD: 0.93

untreated O post LVC



CORVIS ST — TE



**CORVIS ST - CBH** 



#### **Clinical application**







# The current concept is that the pathophysiology of ectatic diseases is associated with

**Primary biomechanical abnormality** 



UNDERSTANDING THE **CORNEA'S BIOMECHANICAL BEHAVIOR** IS RELEVANT FOR THE DETECTION OF **SUBCLINICAL KC AS WELL AS FOR DETECTION OF ECTASIA PROGRESSION**, WHILE CHANGES IN TOPOGRAPHY ARE STILL INSUFFICIENT TO PROVIDE CONCLUSIVE EVIDENCE

#### Normal vs. Keratoconus See the difference!

Normal eye

Keratoconic eye



Provided by Renato Ambrosio Jr

#### No criterion = 100% sensitive and specific "Normal" can be "abnormal", but just below the limit of detection "Abormal" can be "normal", but just above the limit of detection



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## **DIFFERENT** SCENARIOS CASE 1->

















## **DIFFERENT** SCENARIOS $\mathsf{CASE} 2 \rightarrow$



Composite with the TBI display (Corvis ST + Pentacam) and the Placido disk-based topography Advanced corneal tomographic and biomechanical analysis also revealed abnormal findings

## **DIFFERENT** SCENARIOS $\mathsf{CASE} 3 \rightarrow$



## **IFFERENT** SCENARIOS CASE $4 \rightarrow$



## **DIFFERENT** SCENARIOS CASE $5 \rightarrow$



## FFERENT SCENARIOS $\mathsf{CASE} 6 \rightarrow$



## **DIFFERENT** SCENARIOS $\mathsf{CASE} 7 \rightarrow$







#### OCULUS Corvis® ST - Biomechanical/Tomographic Assessment (ARV)

ID: 64379 Date of birth: 01.2006 13 Name: Age: Exam. Date: 07.05.2019 Time: 17:25:14 Left (OS) Exam. Date: Time: 17:22:35 Left (OS) Eye: Eye: Info: QS: ost Points! Info: QS: OK **Biomechanical Assessment** Pentacam - Tomographic Assessment Corvis ST -0 0 444 Distributions: NORMAL Population KERATOCONUS Population 98.0 300 Axial / Sagital Curvature (Front) **Comeel Thickness** 88.0 340 Stiffness Parameter A1 45.4 Integrated Radius 11.8 90\* 901 Smin 9min OS OS 70.0 380 1460 633 500 46.0 44.0 Are: 620 660 -00.0 65.9 1063 8.1 10.5 586 475 577 750 ARTh 213.2 DA Ratio 5.7 740 604 10.0 900 D 10 jan N т N т 270\* 270" Curveture Pachy. 154.2 440.1 43 53 Abs Abs Percentage Thickness Increase (PT) Diameter Comeal Thickness Spetial Profile (CTSP) Diemeter :61 N 10 mm 10 mm 400 ------10 500 20 30 880 HC: -40 700 50 60 70 800 900 A2 10 1000 7.79 ms K Max. **IOP**nct DIOP. 9.4 mmHg 14D PRFL 0.99 TKC KC 2 75mmHg 55.5 D **IS Value** BAD D CB 7.36 1.00 0.25 0.50 0.75 1.00 1.6 Interest C post Ly TBI 1.00 0.00 0.25 0.50 0.75 1.00

#### The Ambrósio, Roberts & Vinciguerra (ARV) Display from the VAE-E

1.6b1970 (Research)
#### OCULUS Corvis® ST - Biomechanical/Tomographic Assessment (ARV)

1.6b1970 (Research)



The Biomechanical and Tomographic Display showing the Corvis Biomechanical Index (CBI), tomographic biomechanical index (TBI) from the VAE-NT case with uncorrected distance visual acuity of 20/20.



(A)

1.5/1902

OCULUS Corvis® ST - Biomechanical/Tomographic Assessment (Ambrosio, Roberts & Vinciguerra)



Author / Reference	NE (n)	Clin Ectasia (n)	Cut- off	Sensitivity (%)	Specificity (%)	AUC	VAE- NT	Cut- off	Sensitivity (%)	Specificity (%)	AUC	Observation
Steinberg J et al. [71]	105	96	-	98.00	100	0.998	32	0.11	72.00	71.00	0.825	VAE NTT: 18 eyes Sensitivity: 67% / Specificity: 65% / AUC: 0.732
Kataria P et al. [ <u>62</u> ]	100	100	> 0.63	99.00	100	0.995	100	> 0.09	82.00	78.00	0.793	-
Ferreira-Mendes J et al. [ <u>16]</u>	312	118	0.335	94.40	94.90	0.988	57	0.295	89.50	91.00	0.96	-
Chan TCY et al. [72]	37	23	-	-	-	-	-	0.16	84.40	82.40	0.925	-
Sedaghat MR et al. [ <u>61</u> ]	137	145	> 0.49	100	100	1.000	-	-	-	-	-	-
Koc M et al. [ <u>73</u> ]	35	-	-	-	-	-	21	0.29	67.00	86.00	0.790	-
Koh S et al. [ <u>74</u> ]	70	-	-	-	-	-	23	> 0.259	52.17	88.57	0.751	-

*NE*= normal eyes, *VAE-NT*= very asymmetric eyes with normal topography, *NTT*= eyes with normal topography and tomography, *AUC*= area under the receiver operating characteristics curve



Biomechanical comparison displayed from the right eye in 2017 (Exam A) and 2019 (Exam B). Note that the cornea becomes softer and thinner, considering the first and second consultations, as noted comparing the variables DA ratio, integrated radius, SP A1, and ART h.



#### **OCULUS Corvis® ST - Vinciguerra Screening Report**



145/370 (Passer)/









Comparative Corvis ST display before (A in red) and after CXL (B in blue), including the overlap image at higher deformation, the SSI (Stress-Stain Index), and the stress-strain curves, along with comparative DA ratio, integrated radius, and the Stiffness Parameter at first Applanation (SPA1) indicating stiffer behavior after the procedure.

## Pre- and Post CLX in keratoconus Greater stability post op

Pre op

10 month after crosslinking



president in the second devices of

### Effect of corneal rings Less oscillation

Pre op

Post op

🖈 z oculus®

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# HOME MESSAGES

- Biomechanics would be useful in several clinical applications, including management of, ectasia risk-profiling, and the degree, depth of CXL and precise intraocular pressure measurements based on biomechanically corrected intraocular pressure (bIOP) readings.
- The integration of tomographic and biomechanical data has demonstrated potential to improve the accuracy of detection of ectatic disease and identify susceptibility to develop this complication after laser vision correction .
- Further integration with other data, such as ocular wavefront, axial length, segmental layered (epithelium) and microlayer (Bowman) tomography is also promising.

