#### 10th ANNIVERSARY





## **Prevent** Blindness

Focus on Eye Health National Summit

#### **OUR CHANGING VISION**

July 14-15, 2021



# **Aging Eye Summit**

## Financial Disclosures 2021 – 2022

- None relevant to presentation
- Other financial relations:
- Wolters Kluwer Health
- NSF AWD010114 (MPI Argento & Moroi)
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  - R21 EY030363 (MPI Musch & Moroi)
  - R01 EY025752 (PI Komaromy)



• US Patent #10575723 Larry Kagemann, Joel Schuman, Sayoko Moroi, U Pitt & UM (application #15/550,021initiated 5/29/2000; notified 2/12/2020)





#### Strategic Vision: Where are we? What are the key challenges & opportunities? Sayoko Moroí, MD, PhD

**Chair & William H. Havener, MD Endowed Professor** 

#### VISION RESEARCHERS AT THE OHIO STATE UNIVERSITY

STEERING COMMITTEE: Sayoko Moroi, MD, PhD (P30 PD, Dept. Chair); Colleen Cebulla, MD, PhD; Andrew Fischer, PhD; Raymond Gao; PhD; Lisa Jordan, MS, PhD; Srinivasan Parthasarathy, PhD
 Advisory Group: Anish Arora, PhD; Samir Ghadiali, PhD; Peter Mohler, PhD; Karla Zadnik, OD, PhD

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NEI R01; SAYOKO MOROI (R01EY022124) CYNTHIA ROBERTS+ (R01EY027399) SHIGEO TAMIYA (R01EY030060) NEI other; Raymond Gao (R01EY027315) Nagaraj Kerur (R21EY030651) Sayoko Moroi (R01EY025752, R21EY030363) NIH other: Mohamed Abdel-Rahman (R01CA) Raymond Gao (RF1AG060472) Nagaraj Kerur (R01AI148741) NSF, DOD, VA: Mohamed Abdel-Rahman (DOD) Colleen Cebulla (DOD) Sayoko Moroi (NSF) Other Research Support: Mohamed Abdel-Rahman Mona Adeli Colleen Cebulla Fred Davidorf Gloria Fleming Andy Hendershot Nagaraj Kerur (Alzheimer's Assn) Tom Mendel Sayoko Moroi Matthew Ohr Mark Slabaugh Ana Suelves	TIMOTHY PLAGEMAN (R01EY026910) NEI other: Heather Anderson (R01EY024590) Heather Chandler (R01EY030621) Nathan Doble (R01) Stacey Choi+ (R01) Lisa Jordan (UG1EY023206, R34EY030582) Donald Mutti (UG1EY023210) Jeffery Walline (UG1EU023208) Deyue Yu (R01EY025658) NSF, DOD, VA: Andrew Hartwick (DOD) Other Research Support: Melissa Bailey Angela Brown Stacey Choi+ Nathan Doble+ Jennifer Fogt Nicklaus Fogt Bradley Dougherty Marjean Kulp Gladys Mitchell Donald Mutti Teng Leng Ooi Timothy Plageman Thomas Raasch Heidi Wagner Phillip Yuhas Karla Zadnik (U10FY08893)	Science & Engineering NEI R01: NEI other: Aleix Martinez (R01EY020834) NIH other: Srinivasan Parthasarathy NSF, DOD, VA: Srinivasan Parthasarathy (NSF) Federal other: Srinivasan Parthasarathy Dept of Blomedical Engineering+ NEI R01; JUN LIU (R01EY025358) NSF, DOD, VA; Matthew Reilly (DOD)+ Other Research Support: Katelyn Swindle-Reilly+ Cynthia Roberts+ Ronald Xu	NEI R01:      NEI other:      Julie Golomb (R01EY025648)      Other Federal Funding:      Lindsey Delwin      COLLEGE OF      YE TERINARY      MEDICINE      Anne Metzler      Eric Miller      Georgina Newbold      NATIONWIDE      CHILDREN'S      HOSPITAL
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Dept of Neuroscience

#### NEI R01:

ANDY FISCHER (R01EY022030, R01EY032141) JAMES JONTES (R01EY027003) STUART MANGEL (R01EY029777) BENJAMIN SEGAL (R01EY028350, R01EY029159) NEI other: Andrew Sas (K08EY029362) NIH other: Benjamin Segal (R01NS105385) NSF, DOD, VA: Warren Campbell

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CATALYST		
Ann McAlearny Timothy Huerta (UL1TR002733) Federal other: Ann McAlearny Timothy Huerta		

## Biomechanics & Imaging

**Biomechanics** 

- Cornea biomechanics: develop biomechanical markers of disease (glaucoma, ocular hypertension, diabetes, keratoconus vs normal)
- Model optic nerve damage by combing intraocular pulsations with cerebrospinal fluid pressure across the lamina cribrosa

## Cynthia J. Roberts, PhD

Imaging with

- ANTERION (anterior segment OCT with axial length measurement) provide reference markers for imaging that can be used to calculate biomechanics
- Flex Arm OCT with acquisition in sitting, supine, and Trendelenburg provide measurement of translaminar cribrosa gradient
- Corneal and Ocular Wavefront Analysis with high spatial resolution to assess index of refraction
- Corneal Tomography and Topography provide precision to assess keratoconus treatment outcomes









#### Biomechanical Imaging -- PI: Jun Liu, PhD



#### **Out-of-Plane Shear**





### Genetics & biology Mohamed Abdel-Rahman, MD, PhD

#### **Ocular cancer genetics**

1- Identify genes associated with hereditary predisposition to uveal melanoma.

2- Clinical spectrum & management of BAP1-tumor predisposition syndrome (coleader of the BAP1 international consortium).

- 3- Role of BAP1 in tumorigenesis of different cancers.
- 4- Molecular mechanisms of sexual dimorphism in cancer.



Molecular mechanisms of ocular pigmentation in various eye diseases (uveal melanoma, AMD, toxic retinal injury and retinal detachment).



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#### Ocular Oncology & retina cell biology, Colleen M. Cebulla, MD, PhD

#### Uveal Melanoma

- What causes uveal melanoma?
  - BAP1 Tumor Predisposition Syndrome
- The Cancer Genome Atlas Project (TCGA) for Uveal Melanoma
- How can we better treat metastatic disease?
  - Crizotinib adjuvant clinical trial
- Long vs short survivors why?
  - Genetic and inflammatory mechanisms
- OSU Ocular Oncology Study Group
- BIG Consortium



Animal models of retinal detachment (RD):

- Mouse & chick
- Inflammatory mechanisms of retinal cell death / scarring
- Targets for therapy (proteins & genes)

Macrophage migration inhibitory factor (MIF):

- Neuroprotection of photoreceptors & reduces scarring
- MIF gene variations of epiretinal membrane vs proliferative vitreoretinopathy



#### Dr. Stacey Choi and Dr. Nathan Doble

#### High Resolution Optical Imaging of the Human Retina

- No dyes or contrast agents
- Pupil dilation / paralyze accommodation only
- Visible imaging wavelengths
- Use Adaptive Optics (AO) to correct for all of the optical distortions in the eye





AO-OCT-SLO in Rm. 5011 Havener



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#### Precision Medicine PI: Xiaoyi Raymond Gao, PhD

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Our research goals are to advance our understanding of the genetic architecture of complex human diseases / traits, e.g. glaucoma and intraocular pressure, and to improve our ability to predict disease, target prevention to high-risk individuals and tailor treatment based on individual genomic differences.



#### Genetic risk prediction

#### Machine learning / Al

3090E

0.16896



LCD10\_POAG



Department of Biomedical Engineering William G. Lowrie Department of Chemical and Biomolecular Engineering Department of Ophthalmology and Visual Science

#### Swindle-Reilly Lab for Biomimetic Polymeric Biomaterials

#### **Ocular Biomaterials Research:**

- Age-related vitreous liquefaction
- Injectable hydrogel biomimetic vitreous substitutes
- Controlled release of therapeutic agents to prevent oxidative damage
- Lens epithelial cell (LEC) response to biomaterials
- IOL design to prevent posterior capsule opacification



Katelyn Swindle-Reilly, Ph.D. reilly.198@osu.edu biomaterials.engineering.osu.edu

#### Ocular Drug Delivery Research:

- Biodegradable injectable implants
- Drug delivery systems for macular degeneration
- Corneal drug delivery
- Treatments for traumatic optic neuropathy



## **Ocular Aging and Trauma Laboratory**

PI: Reilly; Applying engineering principles to prevention, diagnosis, and treatment of visual problems arising from age or trauma

Ciliary Muscle

Vitreous

Zonule

**Biomechanics, Mechanobiology, and Biochemistry of the Aging Lens** 







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### **Precision Medicine**

#### PI: Sayoko Moroi, MD, PhD

Goals:

- 1) variations in eye pressure and glaucoma drug response (EY022124)
- 2) gene therapy approaches for the trabecular meshwork (EY025752)
- 3) the biomechanics of peri-limbal scleral and aqueous veins (NSF 1760291)
- 4) interplay between vision, fear of falling, and falls in 'SWAN' (EY030363)



#### Molecular pathways of retina scarring – Shigeo Tamiya, PhD

- Cell studies on fibroblast cells and myocardin-related transcription factor (MRTF)
  [cartoon from Velasquez LS et al, PNAS 110:16850, 2013]
- 'Druggable targets' for proliferative vitreoretinopathy

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#### Age-related Macular Degeneration – Nagaraj Kerur, PhD



#### Innate Immune Pathways:

- Inflammasome
- DNA sensor cGAS
- 'druggable' targets NLRP3 & cGAS
- telomerase



#### Laser model of neovascularization:

- VEGF & telomerase cross talk
- Angiogenesis transcriptome profiling
- 'druggable' anti-oligos or small molecules

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- Harness data
- Analyze data (artificial intelligence/machine



- Break barriers of health disparities
- Prevent blindness



# Thank you for your attention





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