

Retinal Hemodynamics: Parafoveal Capillary Leukocyte Velocity

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Why is studying retinal hemodynamics important?

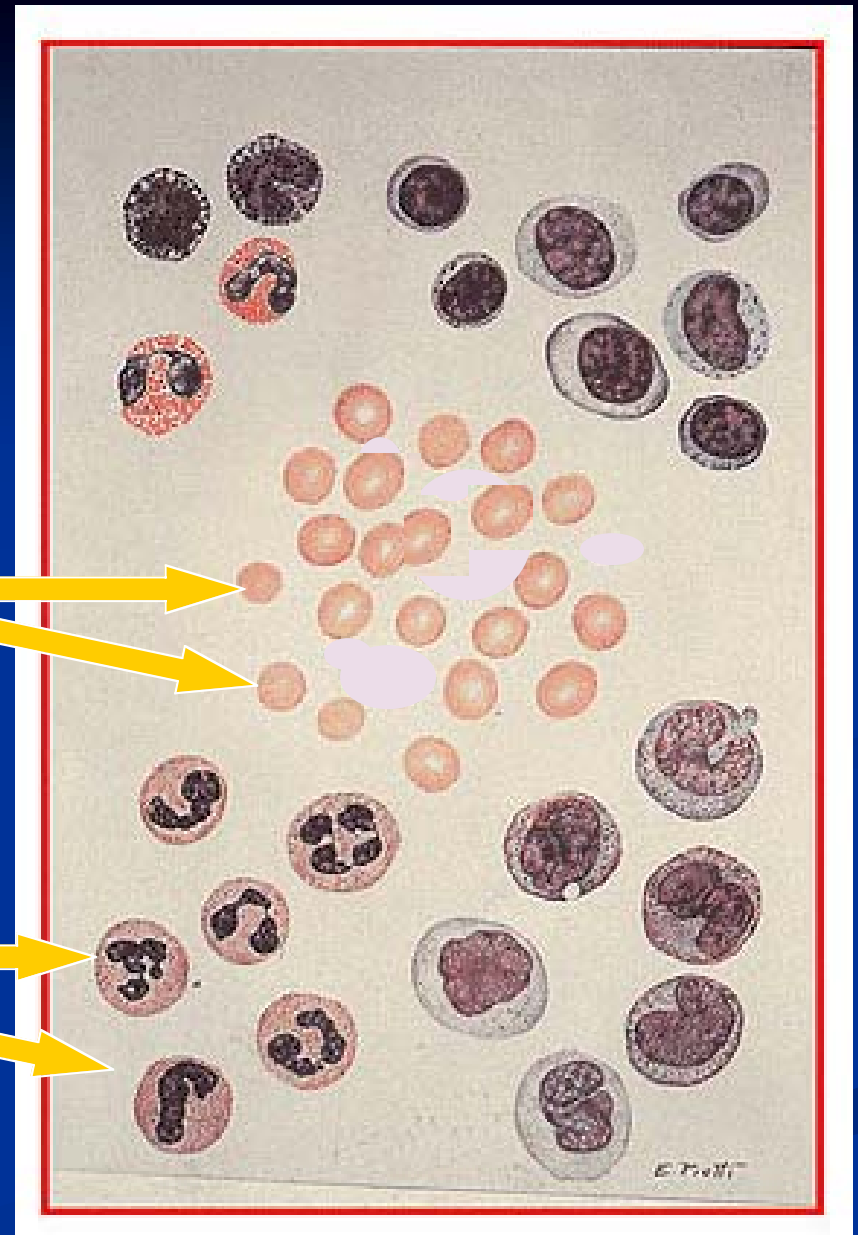
- Blood flow has been implicated in several retinal diseases including: ARMD, Diabetic Retinopathy, and Glaucoma
- Mechanisms of the Disease
 - Developing better understanding of normal and abnormal hemodynamics and the role of vascular changes in disease processes
- Early Diagnosis
 - Retinal circulatory alterations are thought to be an early indicator of pathology
- Treatment of the Disease
 - Research and Development of pharmaceuticals to aid in management or treatment of retinal disease

**RBC=4.2-6.2
million/mm₃ : WBC=5-10
thousand/mm₃**

**Erythrocytes
(RBC)**

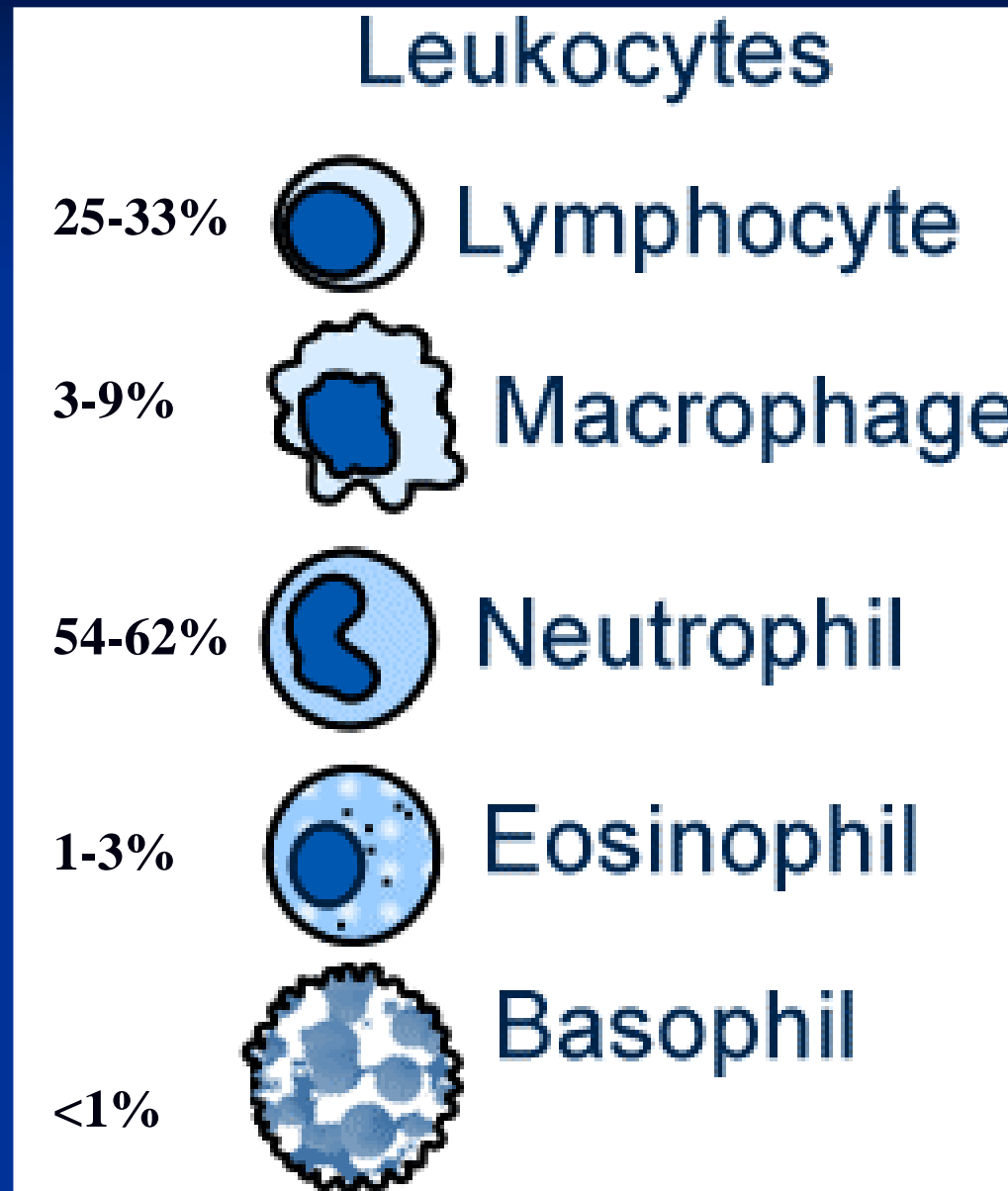
**Leukocytes
(WBC)**

**2-3x Larger than
Erythrocytes**



*Hole's Human Anatomy and
Physiology, Shier et al., 1996*

Proportion of Leukocytes



*Hole's Human
Anatomy and
Physiology
Shier et al., 1996*

Dimensions

- Retinal Capillary
 - Diameter ~ 5 microns
 - Smaller diameters located closest to the fovea
- Erythrocyte
 - Diameter ~7.5 microns
- Leukocyte
 - Diameter ~ 9-12 microns

Snodderly et.al., J. of Neuroscience, 1992

Color Textbook of Histology, 2001

The Retinal Circulation, 1971

Macula Organization

Macula

Parafovea

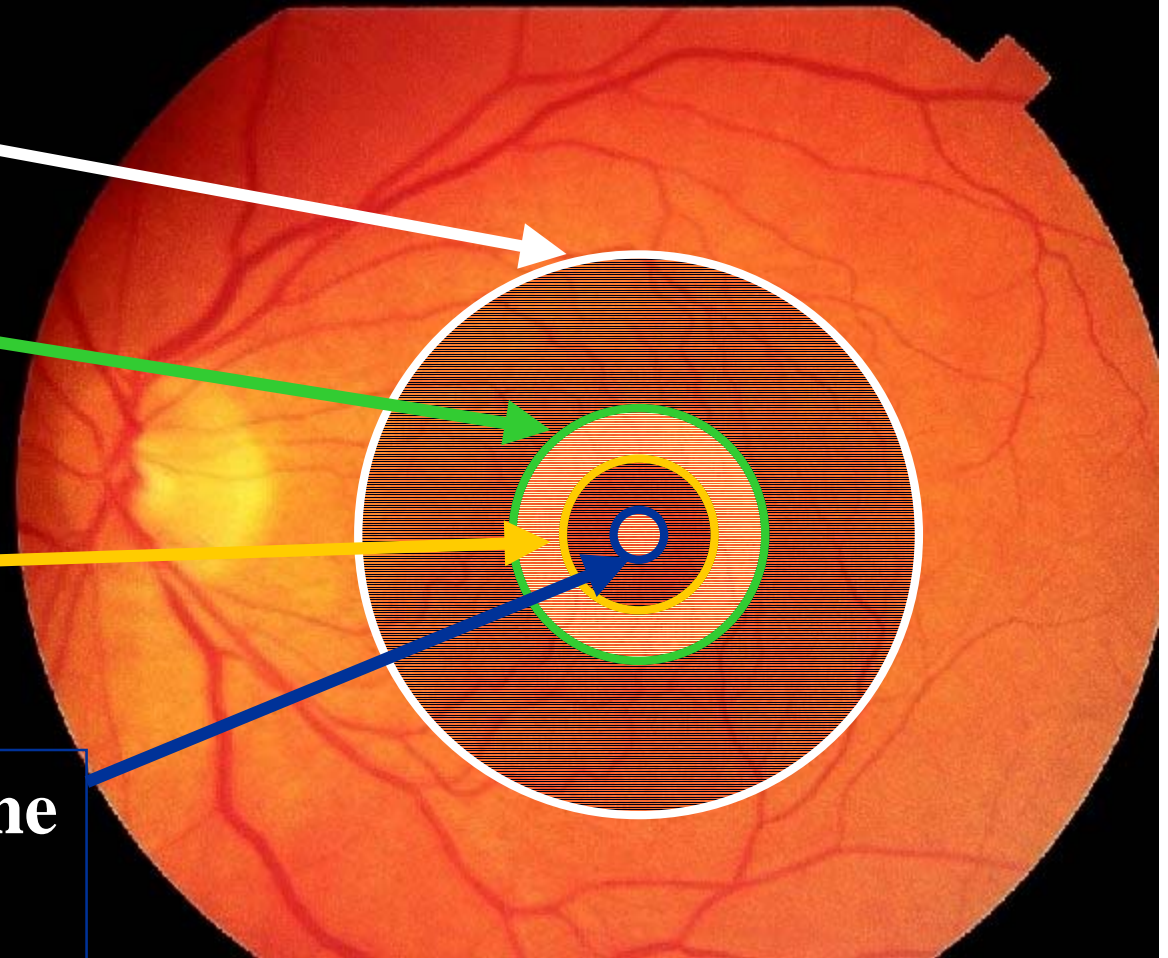
~8.4° diameter

Fovea

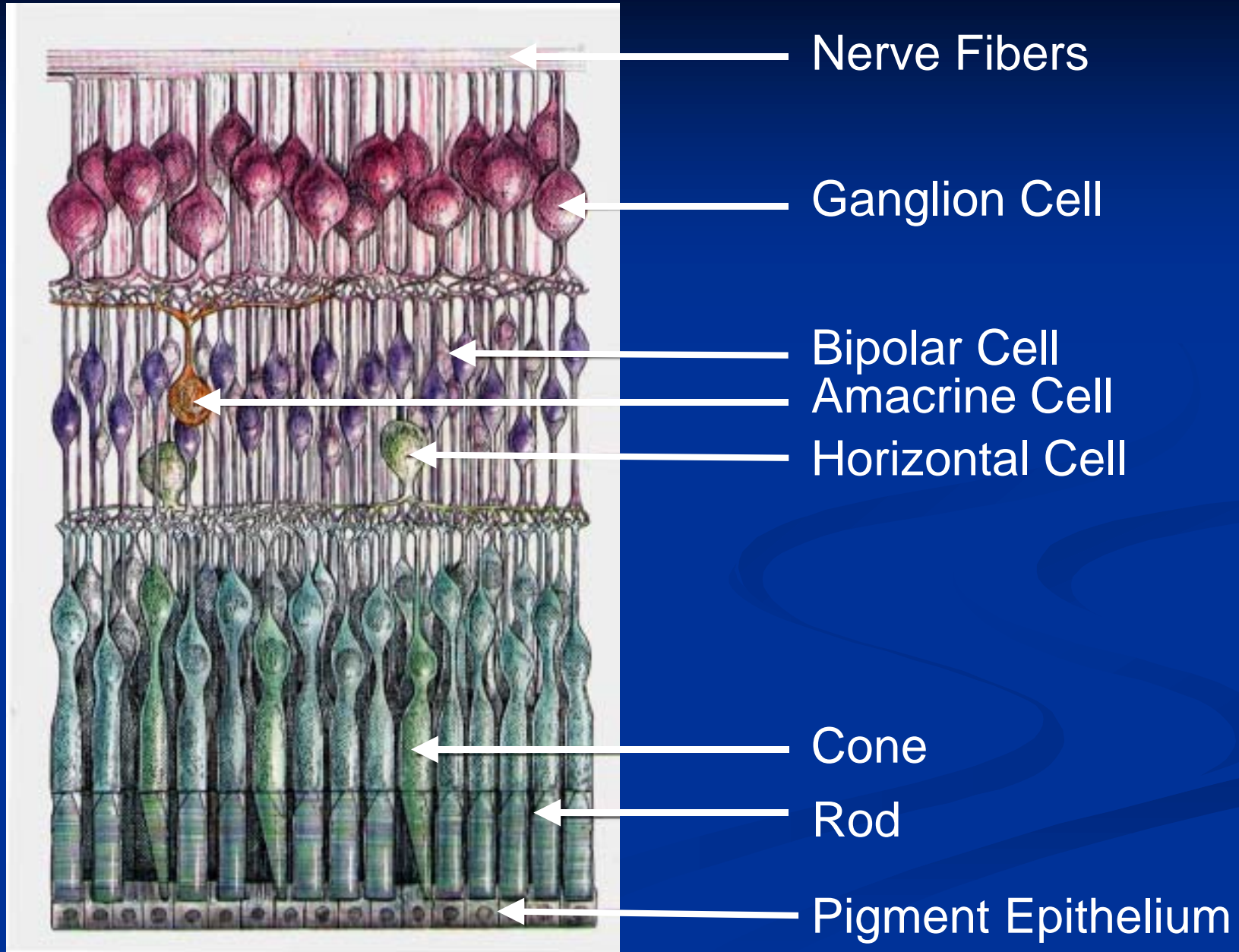
~5°

Foveal Avascular Zone

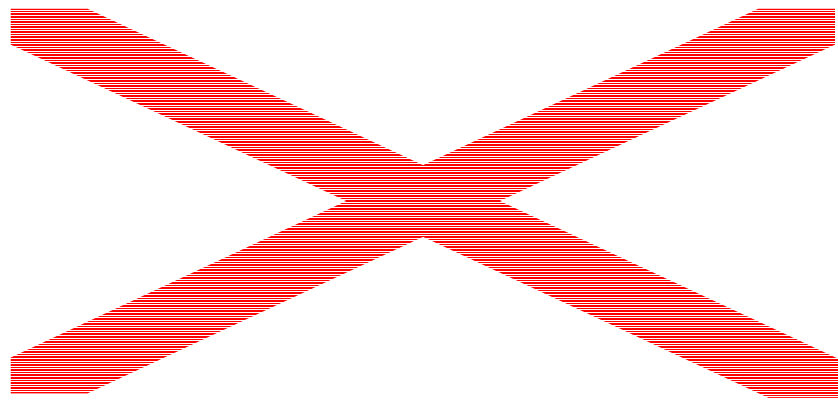
~1.3 - 1.6° diameter



The Retina is a Thick, Multi-layered Structure



Diabetic Retinopathy Lesions



Current techniques for studying retinal hemodynamics:

- Leukocyte Velocity
 - Blue Field Entoptic Phenomenon (Riva and Petrig, 1980)
 - SLO with Fluorescein Angiography (O. Arend)
- Erythrocyte Velocity
 - Retinal Function Imager (Optical Imaging Ltd.)
 - Laser Doppler Velocimetry (C. Riva, 1972)
- Blood Flow
 - Heidelberg Retina Flowmeter
 - Laser Speckle tissue blood flow analysis

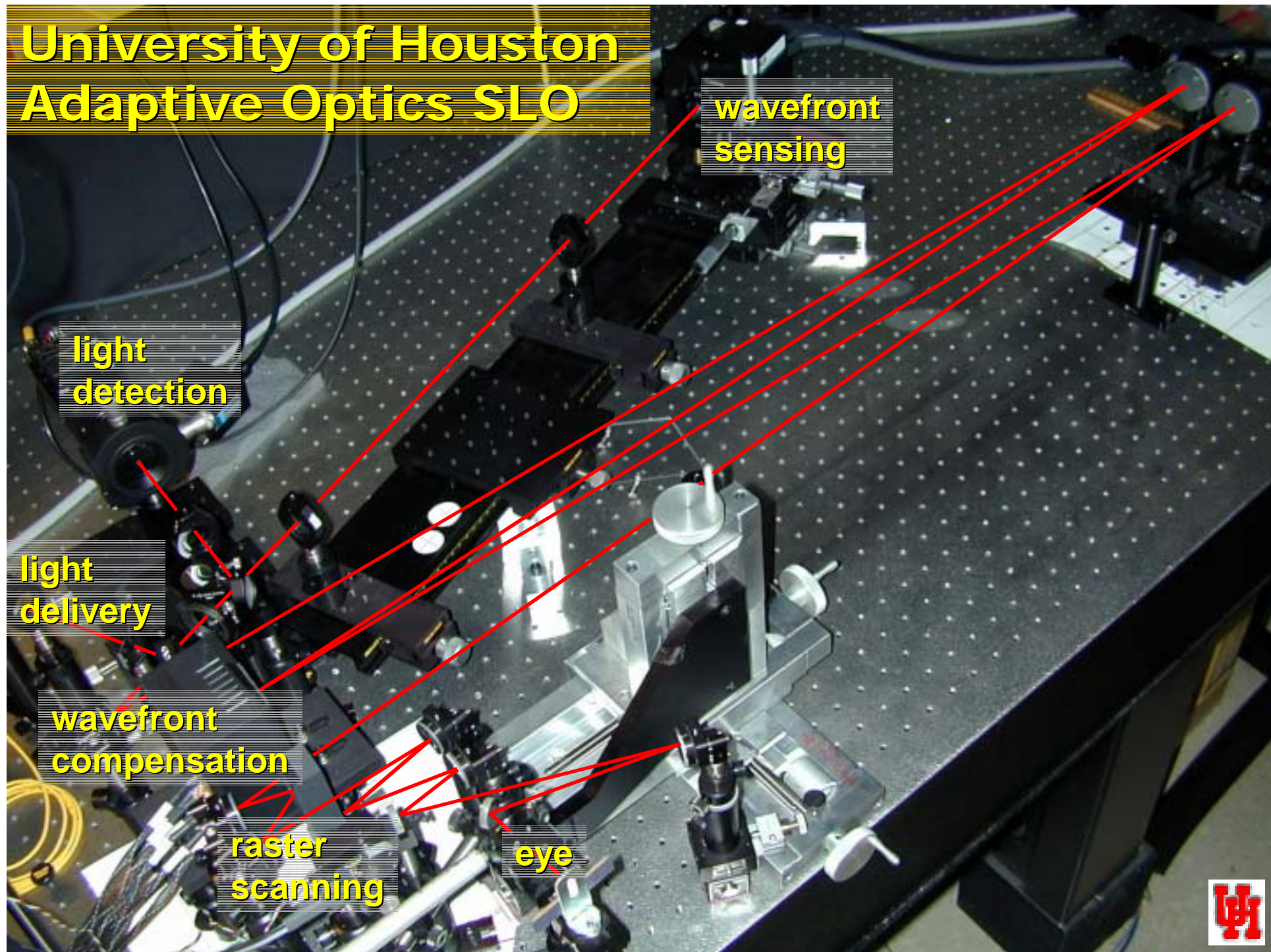
Current Technique's Limitations

- Comparison between techniques is difficult due to variability in location of measurement
- No single technique offers long term, direct, objective, and non-invasive blood flow measurement

Research Purpose

- Develop a technique that allows for long term, direct, and non-invasive retinal parafoveal capillary blood flow measurement

University of Houston Adaptive Optics SLO



wavefront sensing

light detection

light delivery

wavefront compensation

raster scanning

eye



AO ON / OFF

AR left eye

Houston AOSLO

02/05/02



1.5 deg = 0.45 mm

**Photoreceptors
and
Blood Flow**

AR OS 02/05/02



1.5 deg = 0.45 mm

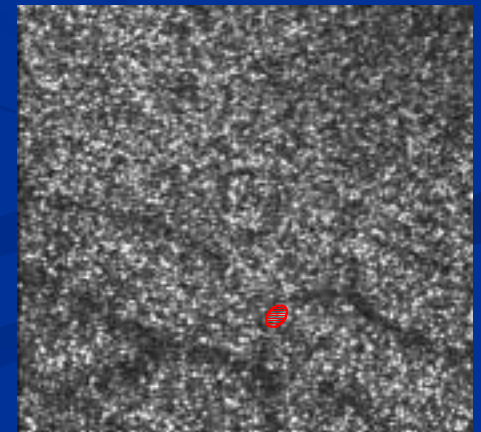
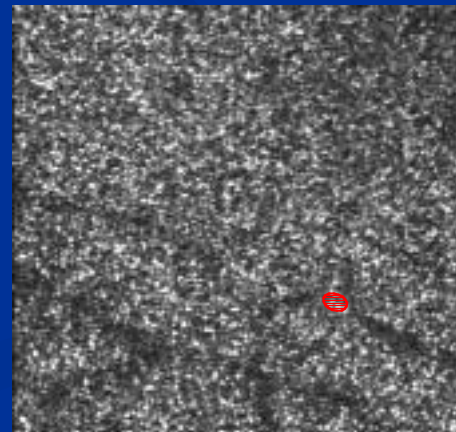
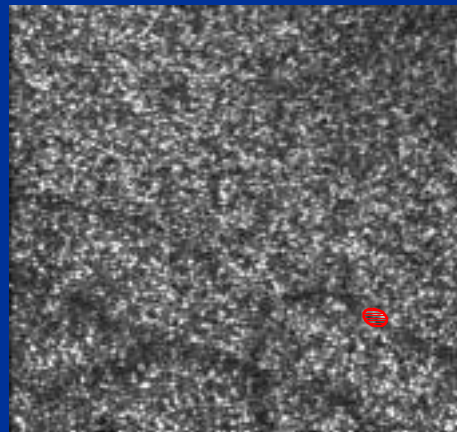
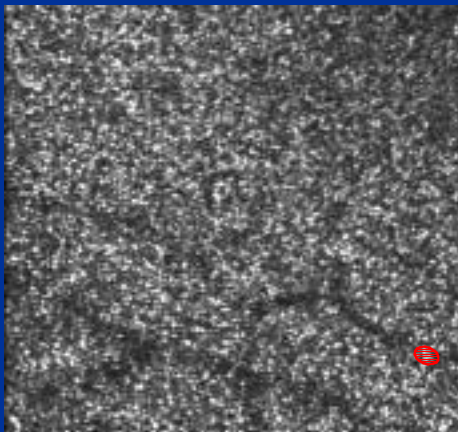
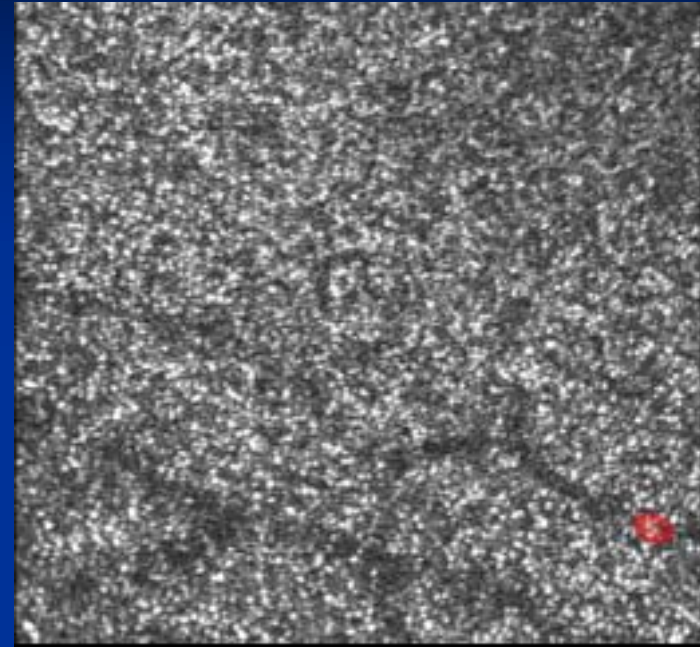
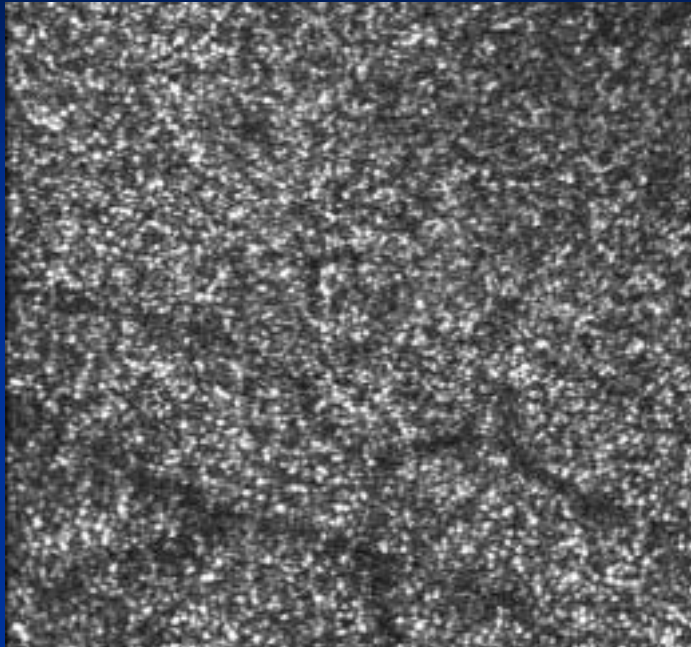
Materials and Methods

- 7 normal healthy subjects were imaged using an Adaptive Optics Scanning Laser Ophthalmoscope (AOSLO)
- 17-35 years of age
- Velocity was determined near the foveal avascular zone in the parafoveal region

Materials and Methods

- Imaging wavelength: 660nm
- Frame rate: 30 Hz
- Velocity was measured directly from movie segments with clearly visible leukocytes
- Five repeated manual measurements were performed to determine individual leukocyte velocity

Leukocyte Velocity Measurement

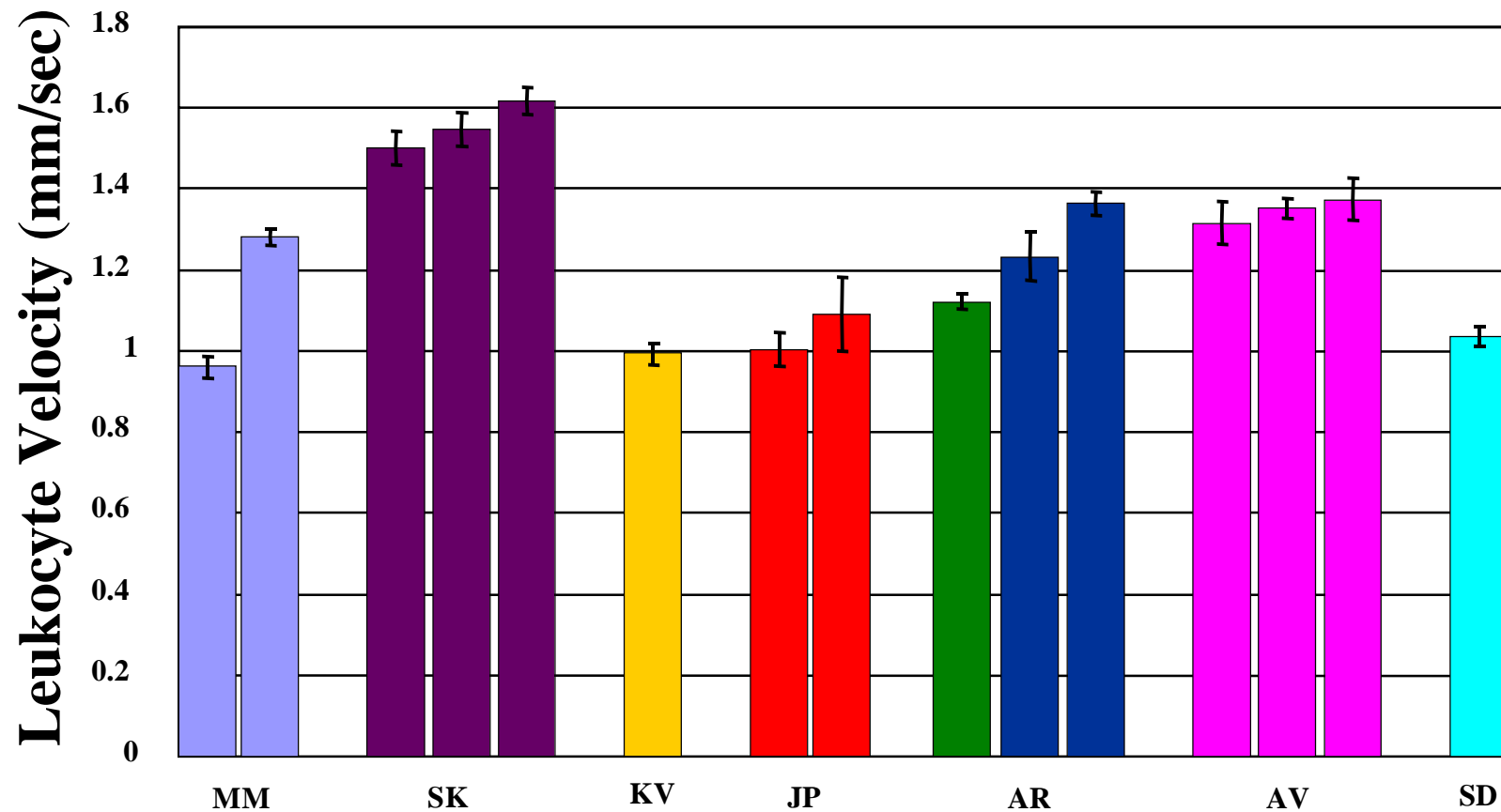


MM Superior Temporal

Results

- Leukocyte velocity was determined in all seven subjects
- Leukocytes were not visible in all parafoveal capillaries
- Mean Velocity = 1.19 mm/sec
- Range: 0.96 – 1.62 mm/sec

Parafoveal Capillary Leukocyte Velocity

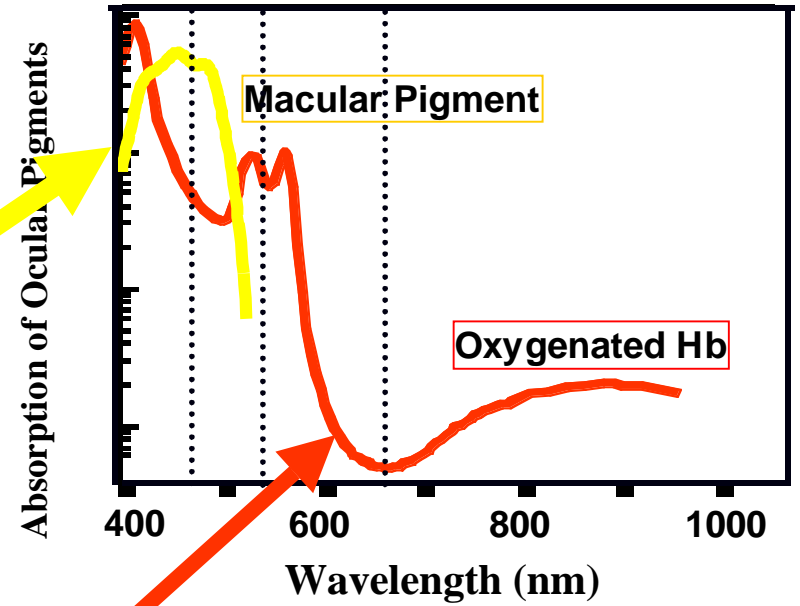
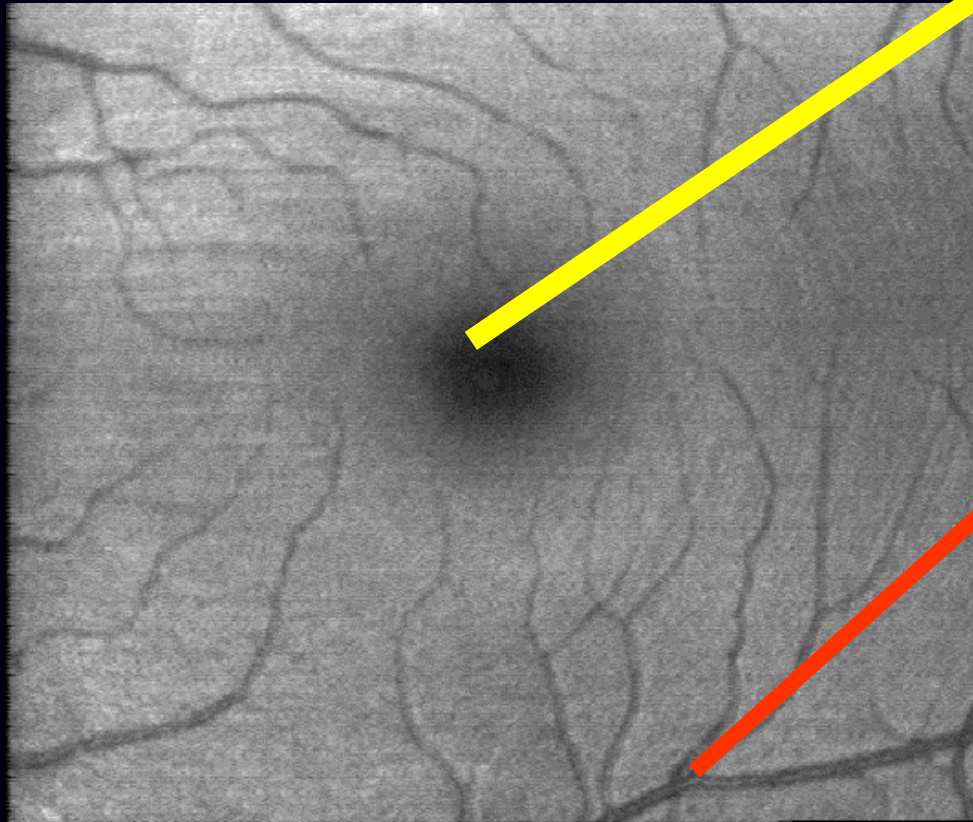


Discussion

Improving Leukocyte Contrast

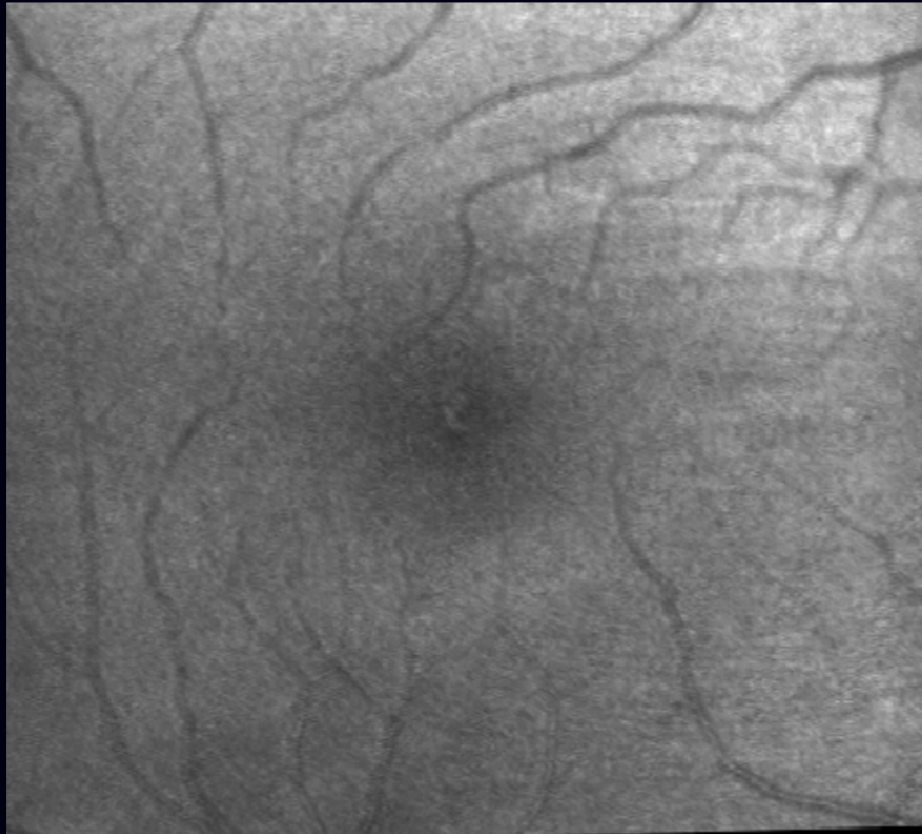
- Currently AOSLO system is imaging with red wavelength
- Use of shorter green wavelength:
 - Maximize contrast of capillary and leukocytes

488 nm



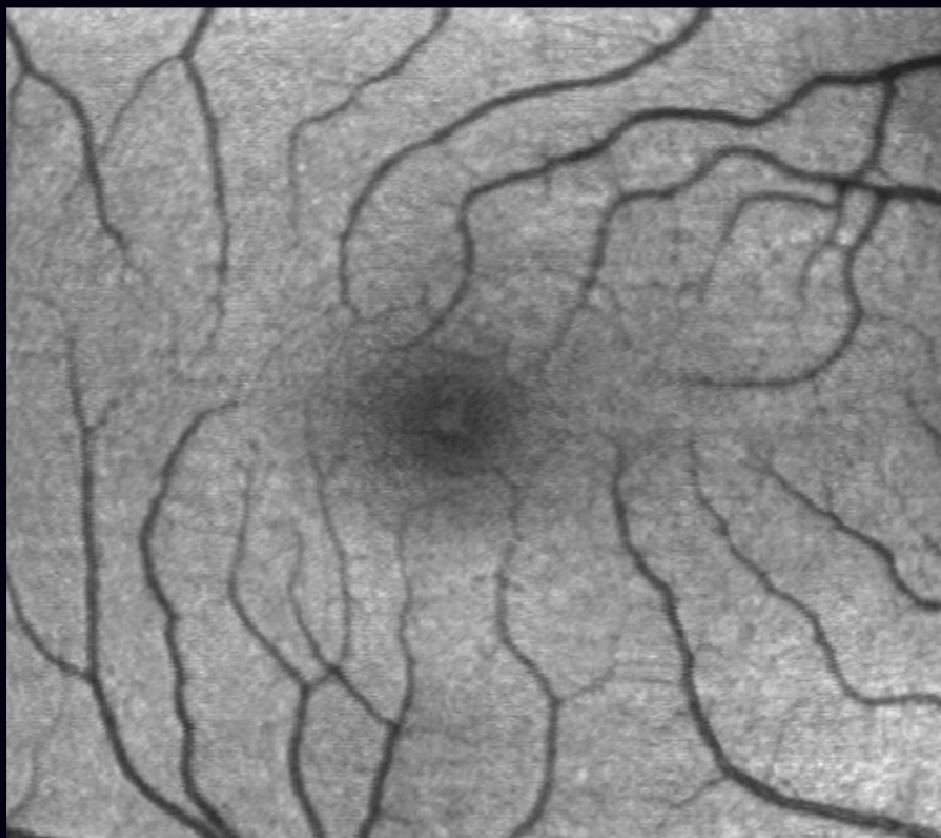
*Courtesy of Steve Burns and Ann Elsner,
Schepens Eye Research Institute, Boston, MA*

633 nm



*Courtesy of Steve Burns and Ann Elsner,
Schepens Eye Research Institute, Boston, MA*

543 nm



*Courtesy of Steve Burns and Ann Elsner,
Schepens Eye Research Institute, Boston, MA*

Limitations

- Need for clear optical media which can be decreased in some subjects:
 - Cataracts and dry eye
- Bite bar used to aid in patient stabilization

Advantages

- Non-invasive
- Direct velocity measurement
- Visualization of capillaries during measurement to determine status as perfused or ghost vessels
- Confocal AOSLO allows for vessel depth localization
- Longitudinal retinal hemodynamics studies are made possible

Conclusions

- Parafoveal capillary leukocyte velocity can be directly and non-invasively measured using an AOSLO

Long Term Scientific Plans

- Measure leukocyte velocity and blood flow in normal healthy subjects
- Measure leukocyte velocity and blood flow in diseased populations: Type I diabetes, glaucoma, and ARMD

Technical Plans

- Confocal imaging of capillaries allowing calculation of vessel diameter and blood flow
- Computer automation of blood velocity and flow measurements
- Use of shorter imaging wavelength
- Replace bite bar with pupil tracking

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