



# **SUNCOAST SEMINAR**

Presented by the  
**Pinellas Optometric  
Association**

**Course Syllabus**

# Suncoast Seminar 2024

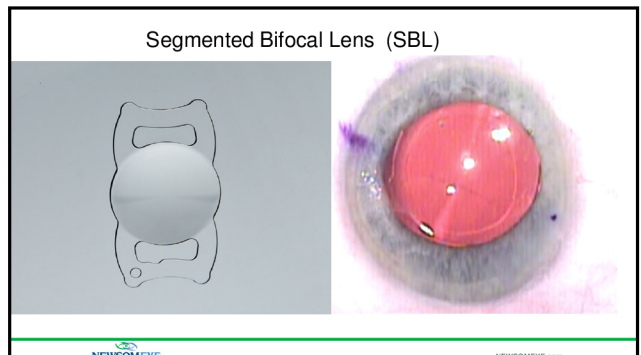
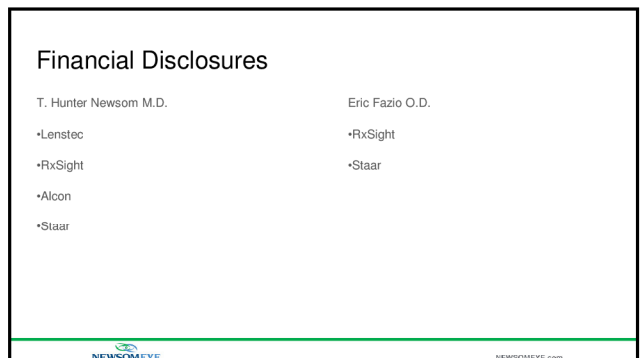
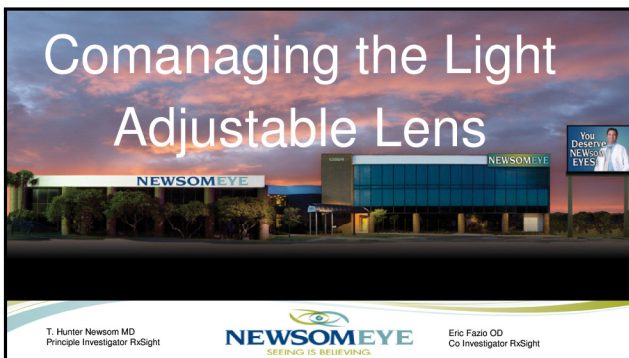
## Schedule of Events

### Saturday, April 27, 2024

- 7:45 am – 8:15 am **Registration**  
**Continental Breakfast** - sponsored by Eye Institute of West Florida  
Exhibit Hall open
- 8:15 am – 9:55 am **Co-Managing the Light Adjustable Lens (90938-PO)**  
T. Hunter Newsom, M.D., Brian Szabo, D.O., and Eric Fazio, O.D.
- 9:55 am – 10:40 am **Break** - sponsored by Updegraff Vision  
Exhibit Hall open
- 10:40 am – 12:20 pm **Emerging Trends in Macular Disease (TQ) (90790-TD)**  
Sherrol A. Reynolds, O.D.
- 12:20 pm – 1:10 pm **Lunch** - sponsored by St. Luke's Cataract & Laser Institute  
Exhibit Hall open
- 1:10 pm – 1:20 pm **Lighthouse of Pinellas Update**
- 1:20 pm – 1:30 pm **F.O.A. Update**
- 1:30 pm – 3:10 pm **Eye on Systemic Disease (TQ) (90791-SD)**  
Sherrol A. Reynolds, O.D.
- 3:10 pm – 3:30 pm **Break** - sponsored by Sight360
- 3:30 pm – 5:10 pm **The ODs Role in Diabetes (TQ) (86739-TD)**  
Sherrol A. Reynolds, O.D.

### Sunday, April 28, 2024

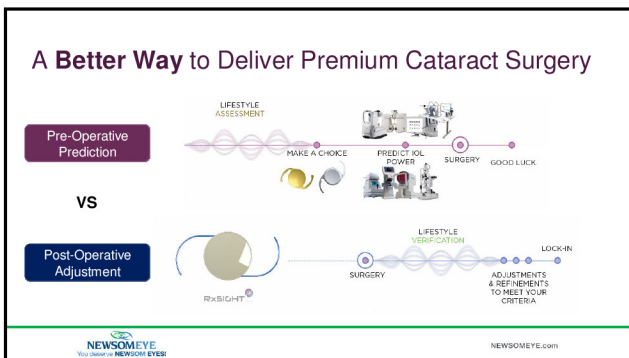
- 7:30 am – 8:00 am **Registration**  
**Continental Breakfast** - sponsored by Next Vision Instruments
- 8:00 am – 9:40 am **Neural Pearls (TQ) (89379-NO)**  
Joe Sowka, O.D.
- 9:40 am – 10:00 am **Break** – sponsored by Suncoast Seminar
- 10:00am – 11:40 am **Prevention of Medical Errors (89825-EJ)**  
Joe Sowka, O.D.
- 11:40 am – 12:00 pm **Break** – sponsored by Suncoast Seminar
- 12:00 pm – 1:40 pm **Florida Jurisprudence (89275-EJ)**  
Joe Sowka, O.D.





### Light Adjustable Lens (LAL)

- 3-piece IOL design
- Blue PMMA modified-C haptics
- Photosensitive silicone
- 6.0mm biconvex optic
- Overall length 13.0mm
- Diopter Power: +10.0 to +30.0 D
- Posterior layer UV absorber
- Optic edge: Square and Round



### 20/Happy in 2020: CORPORATE EDUCATION SESSION

#### Mastering Your Patients' Quality of Vision

Faculty

20/HAPPY IN 2020

Shifting the Perspective: Presbyopia and Patient Needs in the Modern Age

20/happy = 20/imperfect results

LAL = 20/Perfect



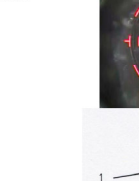
### LAL Surgery

- Standard cataract implant procedure
- Surgery scheduling:
  - Schedule the second eye one week after the first
  - Let both eyes heal for ~two weeks
- \*\*\*At this point the co-managing OD needs to see the patient and:
  - Get a killer refraction
  - Measure IOP and check health just like any other postop cataract
  - If any issue please communicate quickly with Dr. Fazio (cell)
  - 1 week after Comanaged OD refraction adjustments start
  - All treatment visits for LDD are already scheduled

### Description of Technology: LAL Composition

- Photoinitiator**
  - Activated using a specific wavelength of light
- Polymer Matrix**
  - High mol. wt. poly(siloxane)
  - Low glass transition temperature (~-125 °C)
  - Consistently bonded UV absorber
  - Relatively rapid diffusion
- Macromer**
  - Low mol. wt. poly(siloxane)
  - Non-volatile
  - Insoluble in water
- Photo Polymerizable End groups**

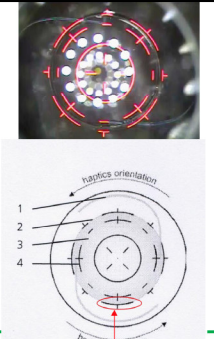
### Light Adjustable Lens

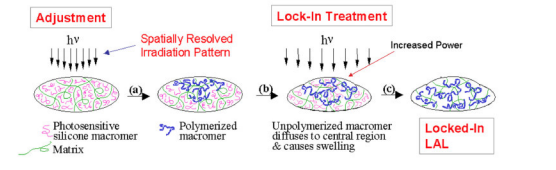
Myopic

Hyperopic

Astigmatic



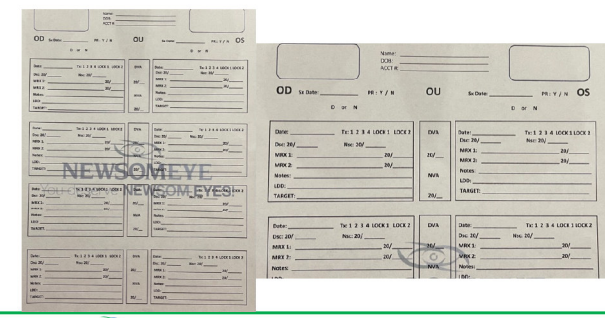
### How does it work?

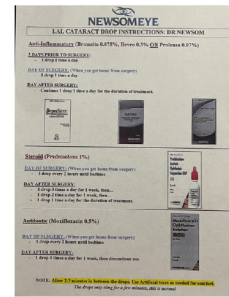


**Adjustment:**  $h\nu$  + Spatially Resolved Irradiation Pattern → Photosensitive silicone macromer Matrix → Polymerized macromer

**Lock-in Treatment:**  $h\nu$  + Increased Power → Unpolymerized macromer diffuses to central region & causes swelling → Locked-In LAL

NOTE: 365 nm ultraviolet light

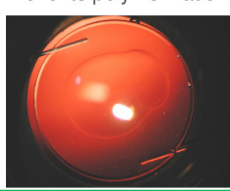



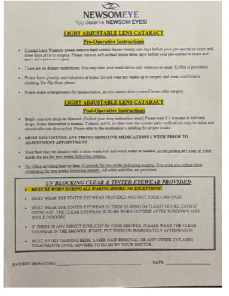


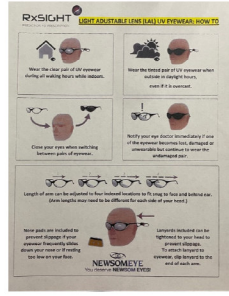
	ADD	MINUS	SPH	CYL	AX	PRISM	BASE	AX
Right	0	0	0	0	0	0	0	0
Left	0	0	0	0	0	0	0	0

### UV Glasses Compliance

UV blocking glasses post-op  
Prevents polymerization





**Introducing a New Level of UV Protection in the Light Adjustable Lens**



ActivShield™ is a revolutionary UV protection layer built into the newly updated Light Adjustable Lens®








**ActivShield™ UV Protector**



**How does it work?**

During light treatments, the ActivShield automatically opens to allow delivery of the precise light from the Light Delivery Device to adjust the lens. After the treatment is complete, ActivShield is automatically engaged to once again protect the lens from outside UV rays.

**Patient Selection**

- Any patient that desires the best quality of vision
- Post Refractive Surgery / Type A personality
- Understands and willing to commit to the process
- Compliance with UV Glasses
- Pupils dilate to 6.5 mm
- No h/o herpes keratitis



**LAL Designed for Predictable Personal FDA Results (n=29)**

**Monocular UVCA one week post lock-in #2:**

- 93% (27/29) of eyes achieved 20/20 or better (Alcon Toric 38%)
- 100% (29/29) of eyes achieved 20/25 or better

**MRSE one week post lock-in #2:**

- 70% (20/29) of eyes within +/- 0.25 D of plano
- 100% (29/29) of eyes within ±0.50 D of plano






**Refractive Accuracy of first 172 eyes**

The refractive accuracy of these 86 subjects was excellent.

The table to the right shows the mean residual refractive error at Lock-in for these subjects.

	Near Eye (N = 66)	Distance Eye (N = 86)	All Eyes (N = 172)
Mean Residual Cylinder (D)	0.08	0.07	0.08
Mean Residual Absolute MRSE from Target(D)	0.11	0.18	0.14

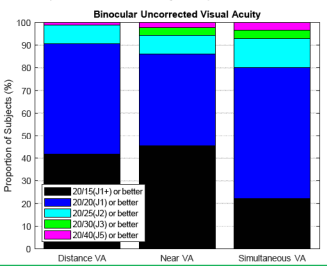





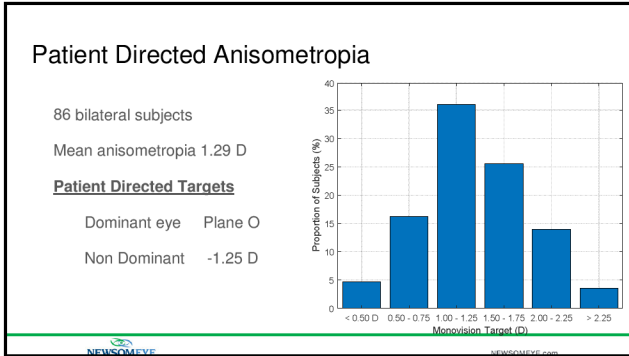
**Post FDA Visual Outcomes (First 172 eyes)**

Uncorrected Binocular Visual Acuity at Distance and Near was measured at visits after all refractive adjustments were complete.

The outcomes of these 86 subjects most recent visit is shown on the right.

We see over 80% (69/86) subjects saw at least 20/20 at distance and J1 at near.



### Customized Monovision

During the course of light treatments, 73% of these 86 subjects chose to change the amount of anisometropia.

On average, subjects chose to increase the anisometropia by 0.37 D.

The table to the right shows initial and final target anisometropia by percentage of subjects with that starting amount.

Final Anisometropia Target (D)	Initial Anisometropia Target (D)			
	< 0.50 D (n = 7/8%)	0.50 - 0.75 D (n = 38/25%)	1.00 - 1.25 D (n = 34/42%)	1.50 - 1.75 D (n = 13/15%)
< 0.50 D	57%	0%	0%	0%
0.50 - 0.75	43%	77%	8%	0%
1.00 - 1.25	0%	43%	47%	8%
1.50 - 1.75	0%	23%	22%	54%
2.00 - 2.25	0%	3%	19%	31%
> 2.25	0%	3%	3%	8%

### It's all about the Method

Conventional Cataract Surgery 1<sup>st</sup> Eye, 2nd Eye 1 week Later

3 weeks post op 1<sup>st</sup> Eye back to Comanaging OD for Quality MRX—**MUST HAVE**

Do not discuss Monovision

First we get the distance than we get the near

As we give you more near you will lose some distance

Any routine post op concerns patients directed to Comanaging OD

4 Weeks post op 1<sup>st</sup> Eye back on Weekly basis for 3-5 Adjustments (Avg 4)

First 3-4 visits adjustments, Last visit is Lock In

### Adjustment Method for Custom Distance

Initial surgical outcomes target hyperopic outcome of +0.75

1<sup>st</sup> adjustment takes patient from +0.75 to +0.25

Initial treatment of +0.50 of hyperopia gives EDOF into the lens.

"First we give you distance vision than the next adjustments we give you Near"

2<sup>nd</sup> Adjustment patient directs if you keep distance or add more near.

"How is your distance vision, computer vision, and distance vision?"

"We always take from the distance to add to the near!!"

### Adjustment Method for Presbyopia

Initial surgical outcomes target hyperopic outcome

Dist eye target +0.75 ----- 1<sup>st</sup> adjustment target for +0.25

Near eye target +0.00----- 1<sup>st</sup> adjustment targets for -0.75

Initial treatment of +0.50 of hyperopia gives EDOF into the lens.

"First we give you distance vision than the next adjustments we give you Near"

2<sup>nd</sup> Adjustment patient directs to add more near, stay the same, or push distance

"How is your near vision, computer vision, and distance vision?"

"We always take from the distance to add to the near!!!"

### Medication Schedule

Antibiotic- tid x 1 week

Steroid- tid x 3 weeks and qd until adjustments completed

NSAID- qd until all adjustments completed

### Late Adjustments

If patients desire to adjust distance or near eye  
A "Late Adjustment" can be performed after 3-6 months post op  
Corneal surgery (Lasik) or piggyback IOL  
Very rare  
Relieves patient's anxiety about IOL being "LOCKED IN"



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### Summary

Light Adjustable Lens is the most accurate IOL in the world  
Moving away from "20/Happy" to "20/Perfect"  
Corrects distance and near vision  
Best option for Post Refractive IOL



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### Thank You!!



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### About me...

- Dr. Fazio is a board certified optometrist and the Medical Director of Primary Eye Care at Newsom Eye.
- Dr. Fazio manages Newsom Eye's accredited one year Residency Program with Salus University.
  - Specializes in ocular disease diagnosis and treatment, dry eye, surgical co-management (over 50,000 cases of cataracts and LASIK)
- EDUCATION
  - Residency Training in Primary Eye Care & Ocular Disease, PCO, PA
  - Doctor of Optometry, University of Waterloo, Canada
  - Honors Bachelor of Science, University of Waterloo, Canada
- NOTEWORTHY ACCOMPLISHMENTS
  - Sub-Investigator of Restor 3.0 Toric post-FDA study (ongoing)
  - Sub-Investigator of CXL eye-on
  - Sub-Investigator for Segmented Bifocal Intraocular Lens, Lensteq, Inc (just approved)
  - Sub-Investigator for the Light Adjustable Lens, Lamou Vision, Inc.
  - Assisted in the Development of the Newsom Bladeless Laser Cataract Technique
  - Published in multiple journals
- PROFESSIONAL ASSOCIATIONS
  - American Board of Optometry, Diplomate
  - American Optometric Association
  - Florida Optometric Association
  - Hillsborough Society of Optometrists Member and Past President



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### LAL Co-Management – What it Takes

- Dr. Newsom has laid out the data and the details of just how good the results can be for this technology
- But how do we get there as Ods co-managing?
- The work for this lens is definitely AFTER the surgery to get the results – NOT before.
- What do we need to know and do to be successful?
- We are going to go over everything you should do and know in pre and post-op care.



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### LAL Patient Selection

Identifying appropriate patients for LAL is key to ensuring high patient satisfaction


Patients Wanting Best Quality Vision

Toric Patients


Precision Mono Vision

Post-Refractive Corneas


Patients who may be candidates include those who are:




Looking to optimize their vision and outcomes



Able to make and keep the additional 2 to 4 appointments needed for optimal vision with the RxLight LAL



Found to have preexisting corneal astigmatism of  $\geq 0.75$  diopters



Able to comply with wearing UV protective glasses until final light treatment

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### Preparing for LAL Surgery


- **Preoperative examination** is unchanged from conventional cataract surgery
- **Reduced dependence** on IOL calculation, although the more accurate the determination the greater range of postop adjustability
- **Lens selection/formula**
  - Use A-constant of 118.4
  - Must dilate well → >6.5 mm pupil
  - Target plano

+10.0 D to +30.0 D

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### Light treatments

Light treatments are painless, non-invasive, and take approximately 90 seconds



LIGHT TREATMENT SCHEDULE

Initial Light Treatment

At least 17 days after surgery

Secondary Light Treatment

At least 3 days after initial light treatment


Additional Light Treatments (if required)

At least 3 days after each prior light treatment

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### Uv protective glasses

At the end of surgery, LAL patients are provided with UV protective glasses to help protect the LAL from indoor and outdoor sources of UV light




Exposure to UV light, such as sunlight, can cause uncontrolled changes to the LAL

The patient must wear UV protective glasses during all waking hours, from the time of lens implantation until 24 hours after the final light treatment has been completed

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### Interactive post-op process


- Refraction optimized after healing is complete and ocular media clear
- Patient selects preferred prescription
- First ever "patient trial" of final outcome



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### Customizing Vision with the LDD


- **Make sure tear film is healthy!**
- **Perform excellent refractions**
- **Involve patient in visual outcomes**
- **LDD treatment scheduling**
  - Handled by a personal RC
  - All treatments are done by THN
  - ? Future for FL ODs to do these???



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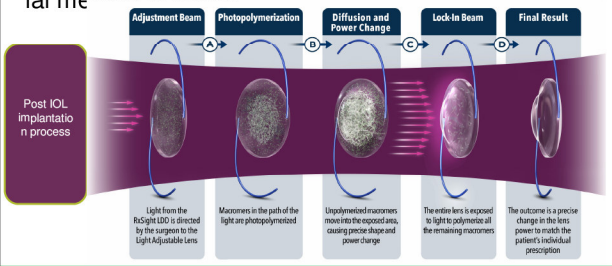
### LDD Targeting

- LAL has negative spherical aberration
- LDD planning
  - Approach from slightly hyperopic (patients get better each time)
  - if goal is plano, target +2SD or +.5D
- Monovision example:
  - Goal OD: Plano OS: -1.00D
  - Target OD: +.5D OD: -.5D



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### Final method of action



Post IOL implantation process

- Adjustment Beam: Light from the R/Sight LDD is directed by the surgeon to the Light Adjustable Lens.
- Photopolymerization: Macromers in the path of the light are photopolymerized.
- Diffusion and Power Change: Unpolymerized macromers move into the exposed area, causing precise shape and power change.
- Lock-In Beam: The entire lens is exposed to light to polymerize all the remaining macromers.
- Final Result: The outcome is a precise change in the lens power to match the patient's individual prescription.

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### Precision monovision with the light adjustable lens

- The LAL addresses the limitations of traditional monovision by:
  - Reducing residual refractive error
    - Precision LAL adjustability
  - Reducing monovision intolerance
    - Adjustable and reversible via patient input
  - Negative SA (LAL and LDD) extends depth of focus to blend near and intermediate UCVA

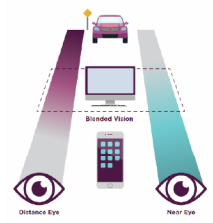
*All while preserving the visual quality of a monofocal lens!*

	Monovision	Trifocal	LAL
<b>Pros</b>	<ul style="list-style-type: none"> <li>Good distance and intermediate (or near) UCVA</li> <li>Low rate of dysphotopsias</li> </ul>	<ul style="list-style-type: none"> <li>Good distance, intermediate, and near UCVA</li> </ul>	<ul style="list-style-type: none"> <li>Good distance, intermediate, and near UCVA</li> <li>Low rate of dysphotopsias</li> </ul>
<b>Cons</b>	<ul style="list-style-type: none"> <li>Residual refractive error</li> <li>Intolerance to monovision</li> <li>Limited UCVA at near or intermediate (depending on target)</li> </ul>	<ul style="list-style-type: none"> <li>Dysphotopsias</li> <li>Residual refractive error</li> </ul>	<ul style="list-style-type: none"> <li>Post-operative UV spectacles</li> <li>Additional post-operative visits</li> </ul>

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### LAL Blended VISION Process

- LAL Surgery
  - Use 118.4 A constant to deliver +0.50 D OU
  - LAL's baseline negative SA leaves most patients with slightly broadened defocus curve
- Light Treatments with LDD
  - Target emmetropia in distance eye
  - Myopic target in near eye doubles the baseline negative SA value (via LDD)
  - Final target refractions are guided by patient interaction



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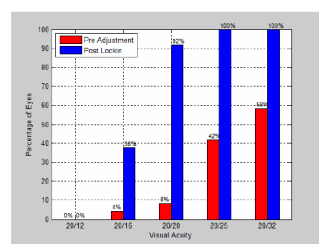
### Refractive Accuracy (24 LAL eyes)

- ❖ 67% (16/24) of eyes are within 0.25D of the target MRSE of plano at 1 week post-lock-in.
- ❖ 100% (24/24) of eyes are within 0.50D of the target MRSE of plano at 1 week post-lock-in.

Caution: Investigational device. Limited by Federal Law to Investigational use.

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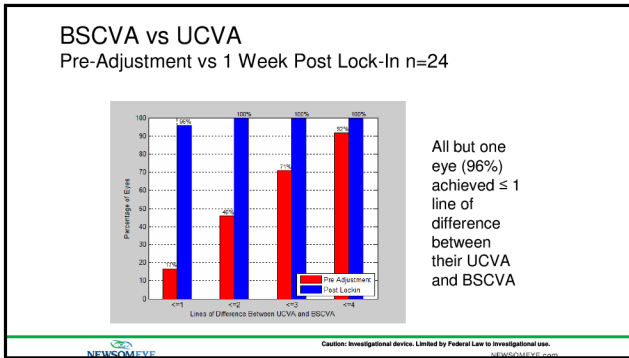
### UCVA: Pre-Adjustment vs 1 Week Post Lock-In n=24



Visual Acuity	Pre Adjustment (%)	Post Lock-In (%)
0/18	0%	0%
20/15	0%	33%
20/20	0%	92%
20/25	42%	100%
20/32	58%	100%

After lock-in, 93% (22/24) of eyes achieved an UCVA of 20/20 or better.

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	PRE-OP	ADJ#1	ADJ#2	LOCKIN#1	LOCKIN#2
PT #1	UCVA	20/100	20/25+2	20/16-2	20/12.5-2
	MIX	(+4.25+0.75X090)	(-0.50+1.00X080)	(-0.25SPH)	(+0.25SPH)
PT #2	UCVA	20/20-1	20/20-2	20/16-1	20/12.5-2
	MIX	(-2.50+0.75X135)	(+0.25+0.50X130)	(-0.75+0.75X130)	(-0.25SPH)
PT #3	UCVA	20/40	20/40-2	20/25+1	20/20-1
	MIX	(-1.00+1.00X155)	(+0.25+1.00X095)	(-0.25SPH)	(PLSPH)
PT #4	UCVA	20/40+2	20/25+1	20/20-2	20/20-1
	MIX	(-2.00+0.50X140)	(PL+1.00X100)	(PL SPH)	(PLSPH)
PT #5	UCVA	20/300	20/30-2	20/12.5-2	20/12.5-1
	MIX	(-8.50+0.50X010)	(+0.75SPH)	(PLSPH)	(+0.25SPH)

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### Toric vs LAL

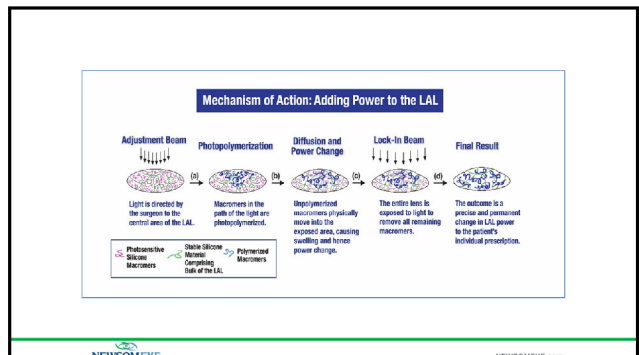
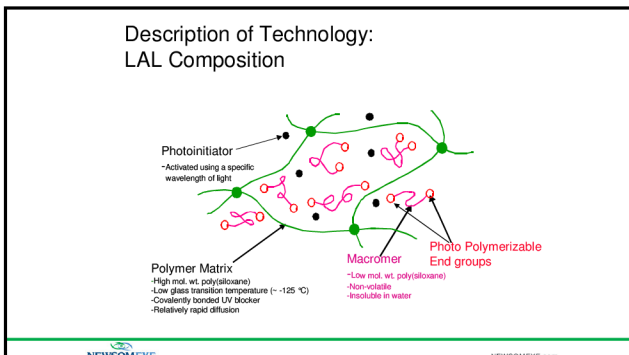
	Pre OP	Adjustment 1	Adjustment 2	1 wk PO Lock in 2	6 Month PO
MIX	1.25 + 1.75 X 005	0.75 + 1.00 X 160	-0.25 + 0.50 X 120	-0.25 + 0.25 X 120	-0.25
Keratometry	44.25@150 / 43.00@060	44.50@165 / 43.50@075	N/A	44.50 @ 165 / 43.50 @ 075	44.75 @ 165 / 43.75 @ 075
UCVA	20/200	20/40+2	20/16+1	20/20-2	20/16
BCVA	20/20-1	20/16	20/12.5-2	20/16-2	20/16+1

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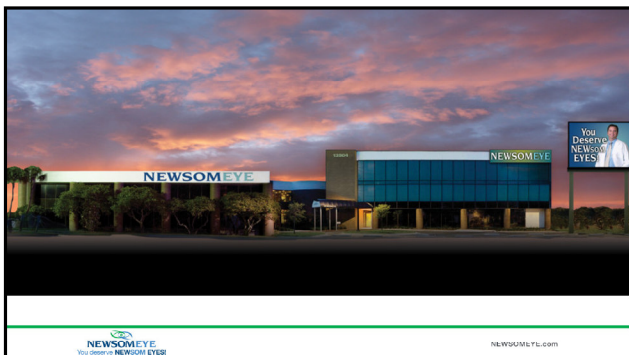
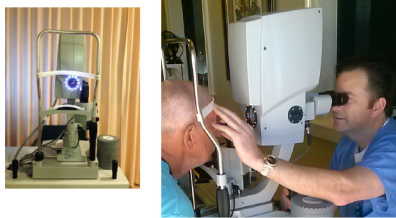
### In Conclusion

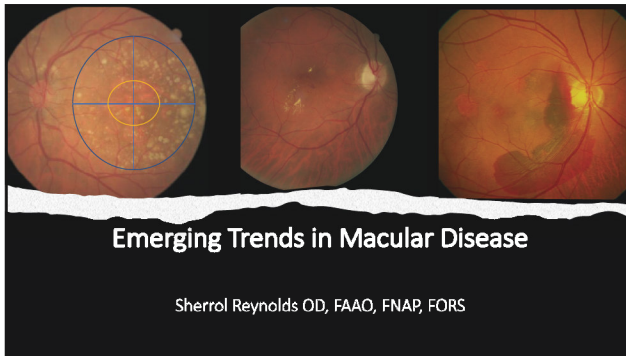
- Reduces post op refractive error / astigmatism
- Reduces glare and halos
- Successfully works in monovision
- Reduces dependence on preop testing
- True Custom IOL for each patient
- Learn that small refractive errors matter
- Results based - Truly the next revolutionary change in cataract surgery!

Caution: Investigational device. Limited by Federal Law to Investigational use.



Light Delivery Device





1

**COPE Disclosure**

Sherrol A. Reynolds OD, FAAO, FNAP, FORS  
Professor of Optometry  
Nova Southeastern University (NSU)  
College of Optometry

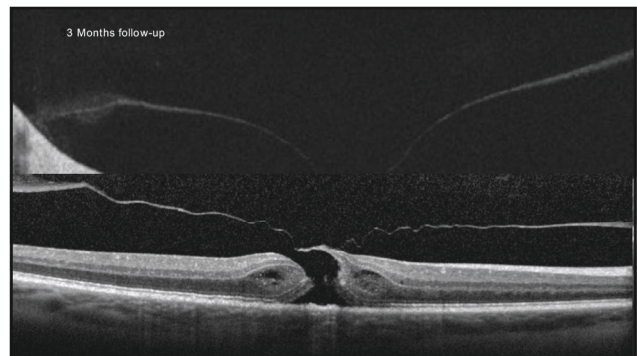
Serve on the speaker bureau, advisory, or contributed board for:  
Vision Service Plan (VSP), American Diabetes Association (ADA), Allergan

2

**Course Agenda**

- Describe the essential tools in examination of macular for assessment of abnormalities leading to a proper diagnosis.
- Appreciate specific clinical and diagnostic findings of most common macular disease.
- Understand the management strategies and preferred practice guidelines for commonly encountered macular disease.
- Know the clinical course, expected outcomes, and latest research for commonly encountered macular disease.

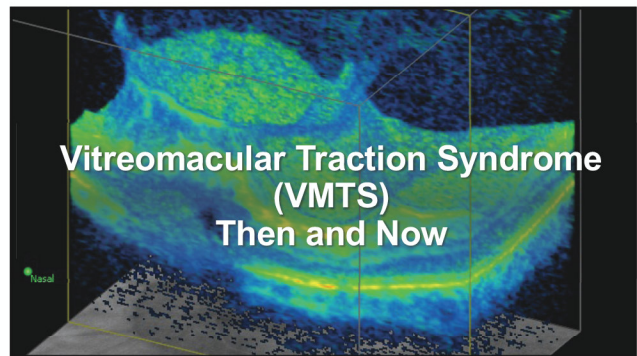
3



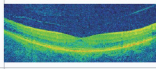
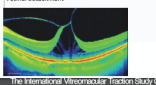
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Entity	OCT-based definition	Additional features	Symptom	Corresponds to full thickness macular hole (FTMH) stage:
<b>Vitreomacular adhesion (VMA)</b>	The following must be present on at least one OCT B-scan image: (i) Partial vitreous detachment as indicated by elevation of cortical vitreous above the retinal surface in the perifoveal area (ii) Persistent vitreous attachment to the macula within a 3-mm radius from the center of the fovea (iii) Acute angle between posterior hyaloid and inner retinal surface (iv) Absence of changes in foveal contour or retinal morphology		None	Stage 0 (Other eye should have full thickness macular hole)
<b>Vitreomacular traction (VMT)</b>	The following must be present on at least one OCT B-scan image: (i) Partial vitreous detachment as indicated by elevation of cortical vitreous above the retinal surface in the perifoveal area (ii) Persistent vitreous attachment to the macula within a 3-mm radius from the center of the fovea (iii) Acute angle between posterior hyaloid and inner retinal surface (iv) Presence of changes in foveal contour or retinal morphology (distortion of foveal surface, intraretinal structural changes such as pseudocyst formation, elevation of fovea from the retinal pigment epithelium (SPE), or a combination of any of these three features) (v) Absence of full thickness interruption of all retinal layers	Foveal pseudocyst, macular thickening, retinal capillary leakage (typically isolated VMT alone does not cause leak on fluorescein angiography), macular schisis, cystoid macular edema, macular schisis, cystoid macular edema, retinal detachment 	Reduced or distorted vision	Stage 1 (VMT only, i.e. impending macular hole) OR: Stage 2 (VMT with small/medium FTMH) OR: Stage 3* (VMT with medium/large FTMH)

The International Vitreomacular Traction Study Group Classification of Vitreomacular Adhesion, Traction, and Macular Hole. Ophthalmology. 2013;120(12):2611-2619

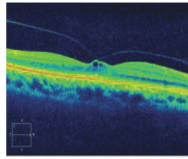
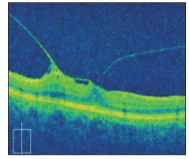
7

### Vitreomacular Traction Study (IVTS) Group (Duker, 2013)

<b>Vitreomacular adhesion (VMA)</b>	(i) Focal: Width of attachment $\leq 1500 \mu\text{m}$ (ii) Broad: Width of attachment $> 1500 \mu\text{m}$
<b>Vitreomacular traction (VMT)</b>	(i) Focal: Width of attachment $\leq 1500 \mu\text{m}$ (ii) Broad: Width of attachment $> 1500 \mu\text{m}$

(i) Concurrent: Associated with other macular abnormalities (e.g. age-related macular degeneration, retinal vein occlusion, diabetic macular edema)  
(ii) Isolated: Not associated with other macular abnormalities

(i) Concurrent: Associated with other macular abnormalities (e.g. age-related macular degeneration, retinal vein occlusion, diabetic macular edema)  
(ii) Isolated: Not associated with other macular abnormalities

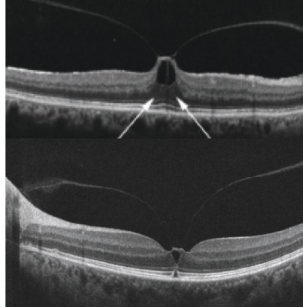



Ophthalmology 2013;120(12):2613-9

8

### Predictors of Vitreomacular Traction Release

- VMT- Classified by the degree of inner-only versus both inner and outer retinal involvement
- Spontaneous resolution
- 10–32% of VMT with only inner retinal distortion are more likely to have spontaneous resolution of traction compared with those who had both inner and outer retinal distortions.**

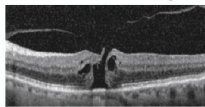
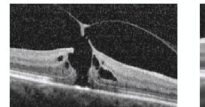
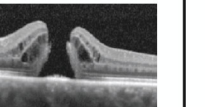


Clinical course of vitreomacular traction managed initially by vitrectomy. Ophthalmic Surg Lasers Imaging Retina 2012; 43(5):373-376

9

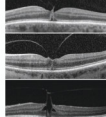
### VMT/ Macular Hole

#### Macular Hole: Updated Classification

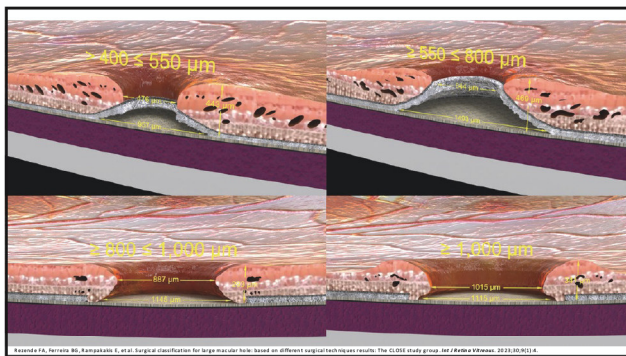
Small  $\leq 250 \mu\text{m}$       Medium  $>250-399 \mu\text{m}$       Large  $> 400 \mu\text{m}$

**Cause -- primary or secondary/ Presence of absence of VMT**



The International Vitreomacular Traction Study Group Classification of Vitreomacular Adhesion, Traction, and Macular Hole. Ophthalmology. 2013;120(12):

10

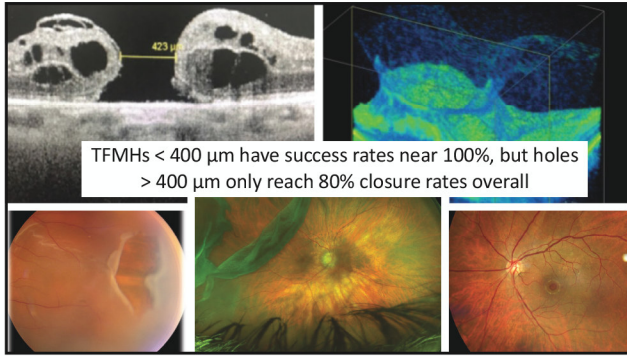


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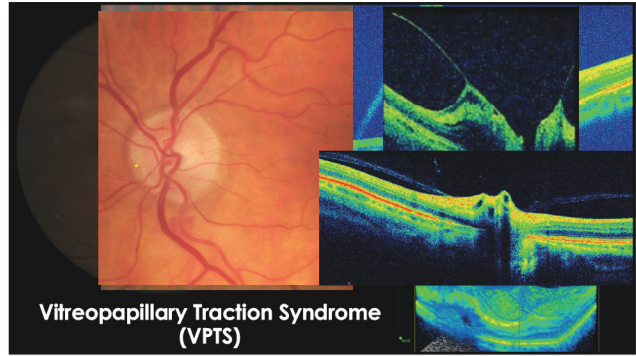
TABLE 3. MEAN BCVA GAINS BASED ON SURGICAL TECHNIQUE (LOGMAR)				
Surgical Technique	Large	X-Large	XX-Large	Giant
ILM Peeling	-0.5293	-0.4248	-0.3858	NA
ILM Flap	-0.3602	-0.3778	-0.2338	-0.2694
Macular Hydrodissection	NA	-0.4748	-0.3441	-0.5664
Human Amniotic Membrane	-0.4902	-0.5177	-0.5342	-0.3497
Autologous Retinal Transplantation	0.2202	-0.3561	-0.4633	-0.4178

NA = not enough numbers available

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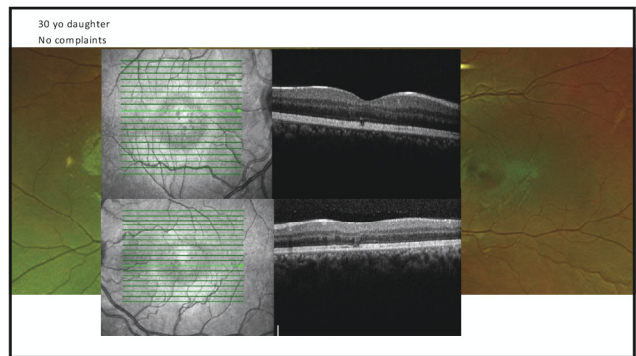
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### Stargardt (STGD) disease

Stargardt macular dystrophy, juvenile macular degeneration, or fundus flavimaculatus	Common cause of central vision loss in adults under the age of 50 • Most common childhood recessively inherited macular dystrophy	Autosomal Recessive (AR) • Genetic basis due to mutations in the <i>ABCA4</i> gene
Lipofuscin storage disease - affects the RPE/photoreceptors through a sequence variant in <i>ABCA4</i> gene • Mutations in this gene result in accumulation of N-retinylidene-N-retinyl-ethanolamine (A2E)	VA- between 20/50 and 20/200	Risk of CVM

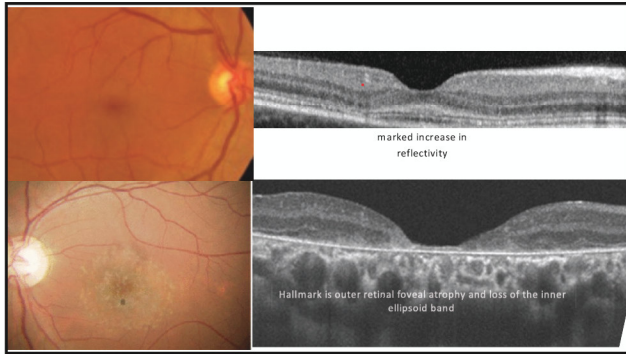
17

### Spectrum of disease

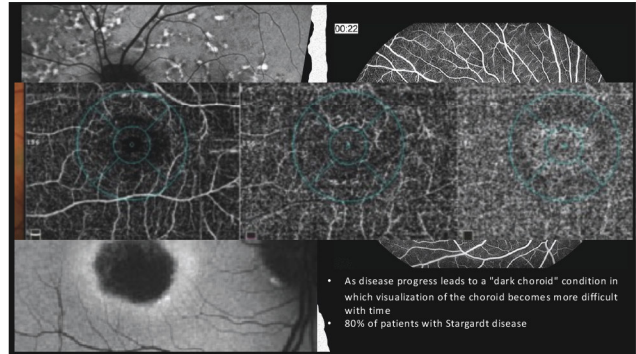
No maculopathy	Maculopathy with/without flecks	
Retinal pigment epithelium (RPE) and choriocapillaris atrophy	Yellow or white fish-shaped flecks with no associated maculopathy	
Extensive atrophy (looks like RP)		

*Fundus Flavimaculatus*  
"fish-tail" or pisciform lesions in 2/3 of cases

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### Monogenic Disease in the Eye: Inherited Retinal Diseases (IRDs)

- Family of genetic disorders that cause retinal degeneration<sup>1</sup>
  - Leber Congenital Amaurosis (LCA)
  - Retinitis Pigmentosa (RP)
  - Cone-Rod/Rod-Cone Dystrophy
  - Choroideremia
  - Stargardt Macular Degeneration
- Rare** – Accurate diagnosis can be difficult and require multiple physician visits<sup>2</sup>
- Few/no treatments available<sup>1</sup>** – Gene therapy could potentially provide new treatments

References: 1. 2018; 2. 2018. © 2018. All rights reserved. 10.1016/j.ophtha.2018.08.026

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### IRD Gene Therapy

#### LUXTURNATM (voretigene neparvovec-rzyl)

subretinal injection in a total volume of 0.3 mL

- A prescription gene therapy product used for treatment of patients with inherited retinal disease due to mutations in both copies of the **RPE65** gene.
- LUXTURNA uses the adeno-associated viral Vector serotype 2 (AAV2) to carry a functional copy of the RPE65 gene into the retinal pigment epithelial (RPE) cells to compensate for the RPE65 mutation.

**Luxturna is \$850,000 per a one-time treatment**

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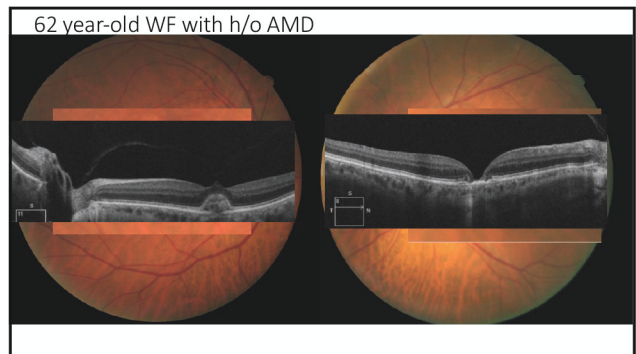
### Effective gene therapy of Stargardt disease with PEG-ECO/pGRK1-ABCA4-S/MAR nanoparticles

Da Sun<sup>1</sup> • Wenyu Sun<sup>1</sup> • Song-Qi Gao • ... Elias I. Traboulsi • Krzysztof Palczewski • Zheng-Rong Lu

Open Access • Published: August 23, 2022 • DOI: <https://doi.org/10.1016/j.omtn.2022.08.026>

ABCA4 Expression after 3 Treatments

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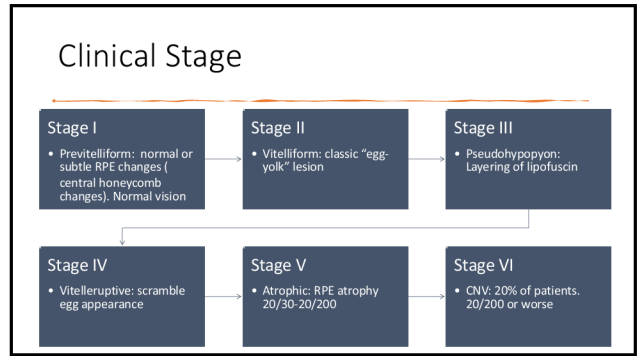
24



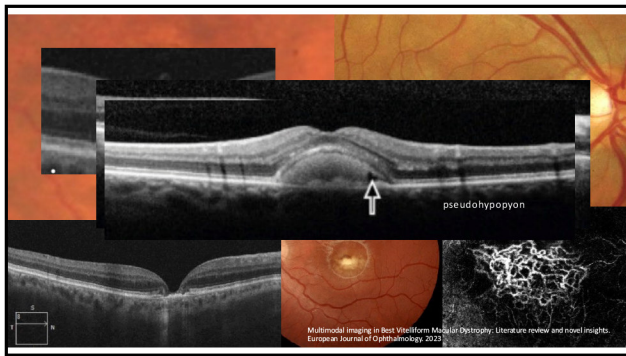
### Adult-onset Macular Vitelliform Dystrophy (AVMD)

- Adult-onset foveovitelliform macular dystrophy (AFVD)
- Autosomal dominant (AD) disease
- Bilateral vitelliform (egg-like) lesions in the macula
  - closely resembles Best's disease
  - **Bestrophinopathies**
- *BEST1* mutation/ *PRPH2*
  - Lipofuscin deposits in the RPE layer
- Abnormal electrooculogram (EOG)
- Risk of CVM

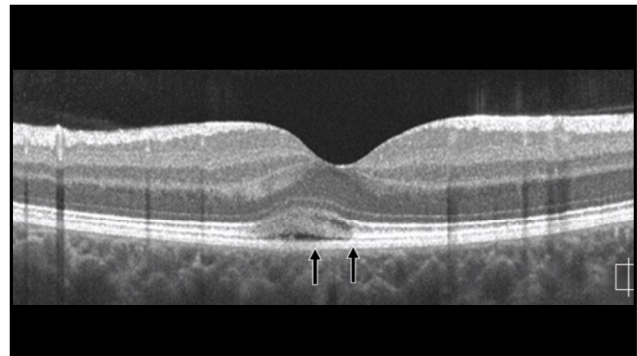
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### Pattern Dystrophy of the RPE

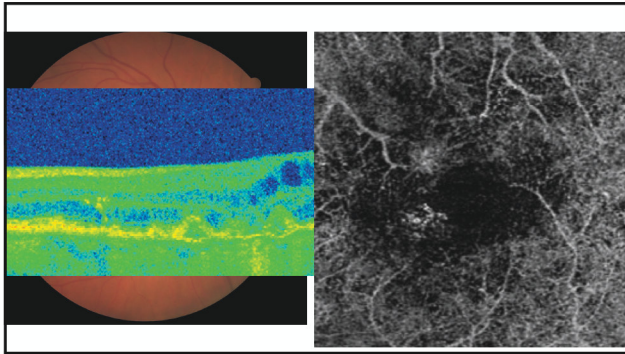
- Clinical picture
  - Symptoms vary – not correlated to maculopathy
  - Mild visual disturbances
    - Blurred vision
    - Metamorphopsia/Relative scotomas
- Heterogenous group of A/D inherited maculopathies
- Human retinal degeneration slow (RDS)/peripherin gene on chromosome
- Adult-onset manifestation: **30-50 yo**

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### Pattern Macular Dystrophy

- Butterfly-shaped pigment dystrophy
- Reticular dystrophy of the RPE
- Multifocal pattern dystrophy
  - simulating fundus flavimaculatus

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### Prevalence of Age-Related Macular Degeneration (AMD) 2024

- AMD affects ~20 million American
- 2 million Americans have Advanced AMD
  - Increase to >5 million by 2050
- 8 million Americans have Intermediate AMD
- 8 million have Geographic Atrophy (GA)

Rein DB, Witterborn JS, Burke-Cotta Z, et al. Prevalence of age-related macular degeneration in the US in 2019. JAMA Ophthalmol. November 3, 2022

The Pathophysiology of Geographic Atrophy Secondary to Age-Related Macular Degeneration and the Complement Pathway as a Therapeutic Target. Retina. 2017;37(8):819-835

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### AMD – Risk Factors

#### Non-Modifiable

- Older-age
- Gender
  - Women are 1.3 times at greater risk for AMD
- Caucasian
- Light iris > darker
  - 2X greater risk
- AMD in the fellow-eye
- Genetic predisposition (eg, family history of AMD)
  - Genetic differences account for ~ 55% of total variability in disease risk.

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### AMD Pathogenesis

- Oxidative stress to the retinal pigment epithelium (RPE)
- Reduced mitochondrial function
- Chronic low-grade inflammation of the retina
- Angiogenesis cascade
- Stress-induced cell death

#### Multiple Cells/Structures Involved

- Photoreceptors
- RPE
- Buch's membrane
- Choriocapillaris
- Choroid

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### AREDS Classification of AMD

Classification	AREDS Category	Clinical signs
No AMD	1	None or a few small drusen (<=3 microns)
Early AMD	2	Any or all of the following: multiple small drusen, few intermediate drusen (63 to 124 microns in diameter), or retinal pigment epithelium abnormalities
Intermediate AMD	3	Any or all of the following: extensive intermediate drusen, at least one large drusen (>=125 microns in diameter), or geographic atrophy not involving the centre of the fovea
Advanced AMD	4	Geographic atrophy involving the fovea and/or any of the features of neovascular AMD

Age-Related Macular Degeneration Study Research Group. Arch Ophthalmol. 2005;123(11):1570-1574

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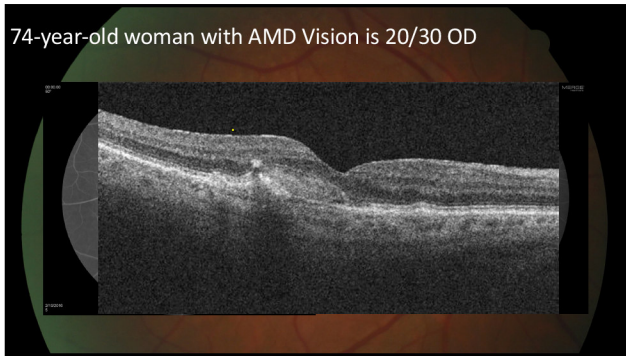
### Monitoring AMD Patients

- Amsler grid
  - Low compliance
- HOME MONITORING DEVICE
  - FORESEE HOME
  - OCT HOME

Hyperscopic Visual Field Test About 3 minutes / Eye at Home

Less than 6.5 minutes indicates impaired function

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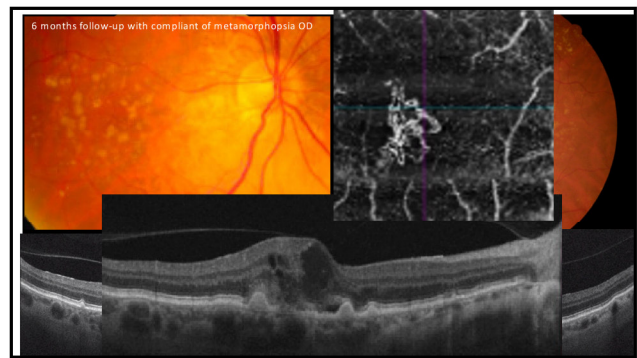
### SD-OCT: AMD High Risk Features

- Loss of RPE integrity
- Disruption of Photoreceptor
- Intraretinal Hyper-reflective foci overlying druse (pigment migration)
- Hypo-reflective foci within druse ('softening of drusen')

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### Current ARMD Treatment Options

- Photodynamic therapy – used Rarely – PCV, chronic leaking growing lesions with scars
- Macugen – used “maybe never” very ineffective but still available
- Avastin – used with step therapy and for cost reasons
- Lucentis
- Eylea
- Beovu – used rarely – unresponsive CNV
- Vabysmo – increasing usage due to improved duration and efficacy
- Susvimo – rarely used – new technology – few trained surgeons
- Biosimilar Lucentis – usage will start soon and be dictated by insurance

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### Latest Treatment

#### Anti-Vegf Biosimilars

- “A biosimilar is a biological product that is approved based on data showing that it is highly similar to a biological product already approved by the FDA (reference product) and has no clinically meaningful differences in terms of safety, purity and potency (i.e., safety and effectiveness) from the reference product, in addition to meeting other criteria specified by law.”
- Currently, 2 FDA approved Ranibizumab Biosimilars
  - Byoviz (Samsung) approved Sept 2021
  - Cimerli (Coherus) approved Oct 2022
- Many in development
  - Ranibizumab  $\cong$  5
  - Aflibercept  $\cong$  8
  - Bevacizumab  $\cong$  1 Outlook Pharmaceuticals (Lytenava)

#### High Dose Aflibercept (Eylea)


- PULSAR (AMD) and PHOTON (DME) Studies
  - Looked at 8 mg vs 2 mg of Eylea
  - Demonstrated non-inferior and clinically equivalent vision gains at 48 weeks with 8 mg at 12 and 16 week dosing after 3 initial doses compared to Eylea every 8 weeks after initial dosing
- Eylea HD FDA approved 8/18/2023 for AMD, DME and DR
  - Recommended dose 1 injection every 4 weeks for first 3 mos for all indications, then every 8-16 weeks (2-4 mos) for AMD and DME and every 8-12 weeks (2-3 mos) for DR

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2024 SUNCOAST SEMINAR

### Vabismo (faricimab)

- Roche/Genentech
- FDA approved January 3, 2022 for AMD and DME
- First bi-phasic antibody for intraocular use
- One arm: Vegf-A inhibitor
- Other arm: Angiopoietin-2 (Ang-2)inhibitor
  - growth factor that promotes vascular destabilization and inflammation
- Dual inhibition of VEGF and Ang-2 have proven more effective than inhibiting either target alone
- Multiple studies show similar results to monthly Lucentis/Eylea but able to object less frequently, many pts q 16 weeks
- FDA approved for RVO



### Susvimo

Previously called Genentech Port Delivery System (PDS)

Refillable port placed under conjunctiva to allow steady supply of Lucentis

Studies (LADDER, ARCHWAY) demonstrated equivalent results to monthly Lucentis at 40 weeks

Large % of pts did not need refill prior to 6 or 12 mos

FDA approved 10/1

Recalled 10/22

Issue with implants breaking when refilled

May be available again late 2023/early 2024

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## AREDS UPDATE

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AREDS Annual Meeting Abstract | June 2021

### The Results of the 10 Year Follow-on Study of the Age-Related Eye Disease Study 2 (AREDS2)

Emily Y. Chew, Traci E. Clemons, Tianran D. Li, Keenan J. Elvira Agrios, Claire E. Maltby, Anitha Domalpally

**Purpose:** To assess the long-term effects of adding lutein/zeaxanthin and omega-3 fatty acids to the Age-Related Eye Disease Study (AREDS) supplements on age-related macular degeneration (AMD) progression and adverse side-effects.

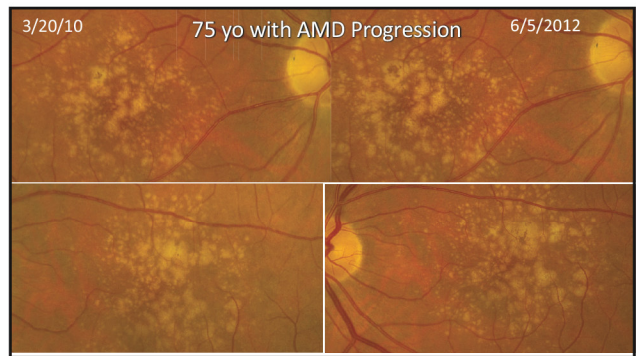
**Conclusion:** The 10-year Follow-on study replicated the findings of the randomized clinical trial at 5 years. Lutein/zeaxanthin, when compared with beta-carotene, had an incremental beneficial effect on progression to late AMD. Beta-carotene doubled the risk of lung cancer, providing support for lutein/zeaxanthin as a replacement of beta-carotene in the AREDS2 supplements

**Methods:** The AREDS2 clinical trial randomly assigned participants with bilateral intermediate AMD or late AMD in one eye to lutein/zeaxanthin and/or omega-3 fatty acids or placebo. Secondary randomization also evaluated varying doses of beta-carotene (0 vs. 15 mg/d and 25 vs. 50 mg/d). At the end of the clinical trial, a follow-up study was conducted with 6-monthly telephone calls to the surviving AREDS2 participants from the central coordinating center to collect outcome data and adverse events for safety monitoring for an additional 5 years. Medical records were obtained from treating physicians to validate any self-reported diagnosis or treatment of late AMD and cataract and side-effects. AREDS2 supplements with lutein/zeaxanthin, vitamin C and E, and zinc plus copper were provided to all participants during this additional follow-up. Repeated measures logistic regression was used in the primary analyses.


**Results:** 6360 study eyes (3887 participants) were analyzed and 3047 (48%) progressed to late AMD. The main effects of lutein/zeaxanthin vs. no lutein/zeaxanthin and of omega-3 fatty acids vs. no omega-3 fatty acids resulted in hazard ratios of 0.91 (95% CI: 0.89-0.93) (p<0.05) and 1.00 (0.92-1.09) (p<0.91), respectively. When the lutein/zeaxanthin main effect analysis was restricted to those randomized secondarily to beta-carotene, the HR was 0.50 (0.49-0.52) (p<0.002). On direct analysis of lutein/zeaxanthin vs. beta-carotene, the HR was 0.85 (0.74-0.98) (p<0.026). For the comparison of low vs. high zinc and no beta-carotene vs. beta-carotene, the HRs were 1.04 (p=0.48) and 1.04 (p<0.05), respectively. For those randomized to beta-carotene, the odds ratio (OR) of developing lung cancer was 1.92 (1.11-3.33) (p<0.02) while the OR for those randomized to lutein/zeaxanthin was 1.19 (0.82-1.73) (p<0.35).

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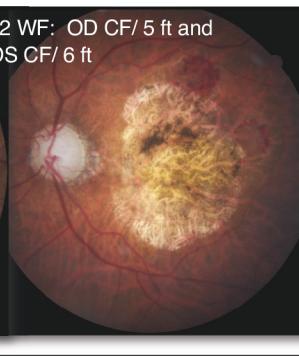
3/20/10
75 yo with AMD Progression
6/5/2012



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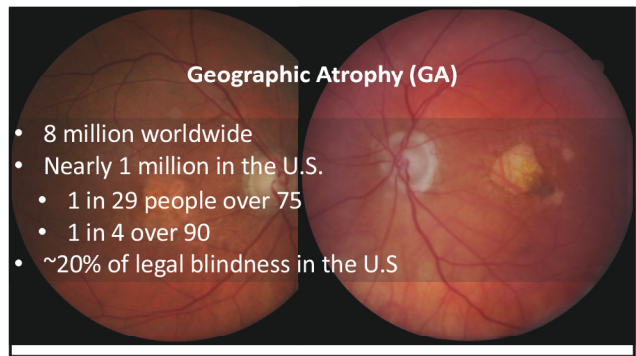
92 WF: OD CF/ 5 ft and OS CF/ 6 ft



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### Geographic Atrophy (GA)

- 8 million worldwide
- Nearly 1 million in the U.S.
- 1 in 29 people over 75
- 1 in 4 over 90
- ~20% of legal blindness in the U.S



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### DRY AMD Geographic ATROPHY (GA) Progression

- Measure the atrophic area to check for progression
- Identification of peri-lesional abnormalities
- Hyper-FAF borders- Sick RPE
- Predict future troubles

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### GA Progression- Classification of Atrophy Meeting (CAM)

- Color fundus photography (CFP)
- Fundus autofluorescence (FAF)
- Near-infrared reflectance (NIR)
- OCT
  - Cross sectional B-scan ( line raster)
  - En-face

Consensus Definition for Atrophy Associated with Age-Related Macular Degeneration on OCT Classification of Atrophy Report 3. Ophthalmology. 2018; 125(4): 537-548

50

### Incomplete RPE and Outer Retinal Atrophy (iRORA)

- “Impending GA”
- Subsidence of the OPL & INL and a hypo-reflective wedge
- Signal hypertransmission into the choroid with corresponding attenuation/disruption of the RPE

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### Complete RPE and Outer Retinal Atrophy (cRORA)

Absence of the RPE and photoreceptors  
 ≥ 250µm in diameter  
 Homogenous choroidal hyper-transmission

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**RPE Analysis**

Advanced RPE Analysis : Macular Cube 512x128

Prior Visit      Current Visit

RPE Elevation Map

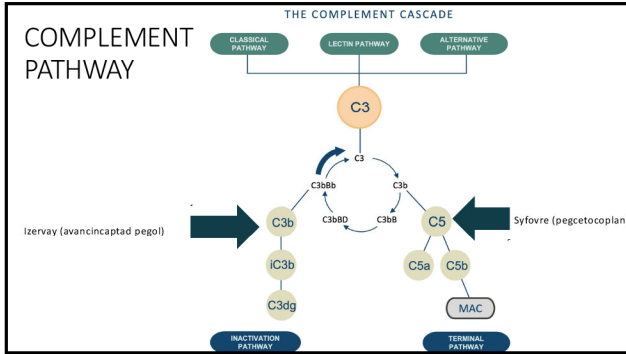
Sub-RPE Map

RPE Profile™

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### OCT Biomarker for GA Progression

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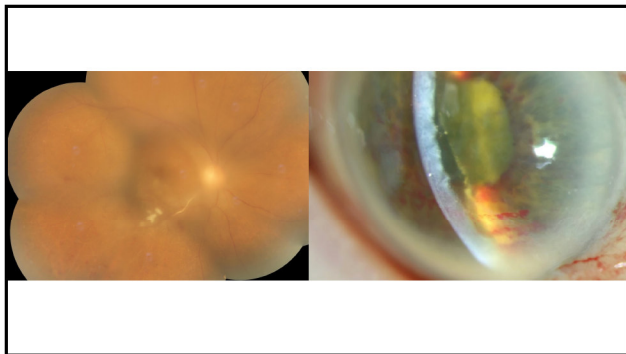


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### Latest Treatment for Geographic Atrophy (GA)

- Syfovre (pegcetocoplan)**
  - Apellis
  - FDA approved Feb 17, 2023
  - One injection every 25 to 60 days
  - Slows progression of GA lesions by blocking C3
    - OAKS Study: 22% qm, 18% qom
    - DERBY: 18% qm, 17% qom
    - Increasing effect with time
  - Recent concern of vasculitis
- Izervay (avancincaptad pegol)**
  - Iveric Bio
  - FDA approved Aug 5, 2023
  - One injection per mos for up to 12 mos
  - Slows progression of GA by blocking C5
    - GATHER1/GATHER 2 studies:
      - At 12 mos, 27% (2 mg) and 28% (4 mg) less GA growth
      - At 18 mos, 28% and 30%
      - Reduced rate of vision loss noted
      - Good safety profile

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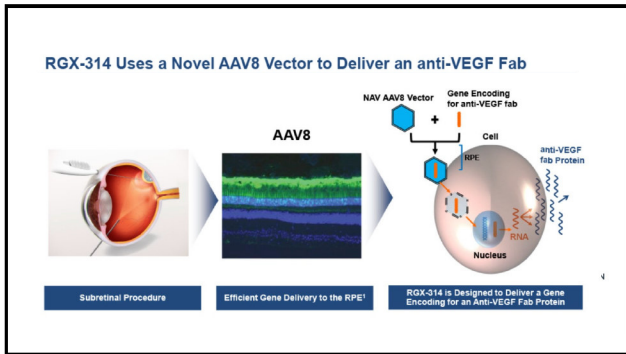
### Gene therapy in AMD: Promises and challenges

- Gene therapy are now being investigated for AMD
- Significant challenges remain, particularly with delivering the vector to the back of the eye
- Long-term benefit of gene therapies in AMD remains to be seen

Therapy name (sponsor)	Vector	Indication	Study Phase	Mechanism of action	Delivery method
RGX-314 (Regeneron)	AAV8	Neovascular AMD	I/IIa	Encodes an anti-VEGF Fab protein similar to ranibizumab	Surgical subretinal injection
ADVM-22 (Aduro Biotechnologies)	AAV2	nAMD	I	Promotes production of anti-inhibitor protein	Intravitreal injection
AAV2-sFLT1 (Sightline)	AAV2	nAMD	I	Encodes sFLT-1 to neutralize vascular endothelial growth factor	Intravitreal injection
AAV2-CDS9 (Altera Biosciences)	AAV2	nAMD and non-nAMD	I	Stable form of CDS9 to inhibit intermediate attack complex formation	Intravitreal injection
GTO05 (Oryscop Therapeutics)	AAV2	Non-nAMD	III	Targets complement cascade	Surgical subretinal injection

Wang J, Khoury S, Zhou L, et al. Intravitreal injection of AAV2-sFLT1 in patients with advanced neovascular age-related macular degeneration. *Appl Clin Biotechnol*. 2023;9(6):61. doi:10.1002/apcb.10000. PMID: 37500000

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### AMD and Stem Cell-Based Therapies

Joseph C Giacalone, David H Parkinson, Daniel A Balkov, Rajesh C Rao

PMID: 38148879 | PMID: 381678350 (available on 2025-01-01) | DOI: 10.1093/ibd/0000000000000000

**Abstract**

Age-related macular degeneration (AMD) is a prevalent and complex disease leading to severe vision loss. Stem cells offer promising prospects for AMD treatment as they can be differentiated into critical retinal cell types that could replace lost cells or provide trophic support to promote host retinal cell survival. However, challenges such as immune rejection, concerns regarding homogeneity and genomic integrity must be addressed. Clinical trials with stem cell-derived retinal pigment epithelial cells have shown preliminary safety in treating dry AMD, but improvements in manufacturing and surgical techniques for delivery are needed. Late-stage AMD poses additional hurdles, possibly requiring multi-layered grafts. Advancements in automation technologies and gene correction strategies show potential to enhance iPSC-based therapies. Stem cell-based treatments offer hope for AMD management, but further research and optimization are essential for successful clinical implementation.

Printed October 2023

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> Int Ophthalmol Clin. 2024 Jan 1;64(1):21-33. doi: 10.1097/IIO.0000000000000510. Epub 2023 Dec 26.

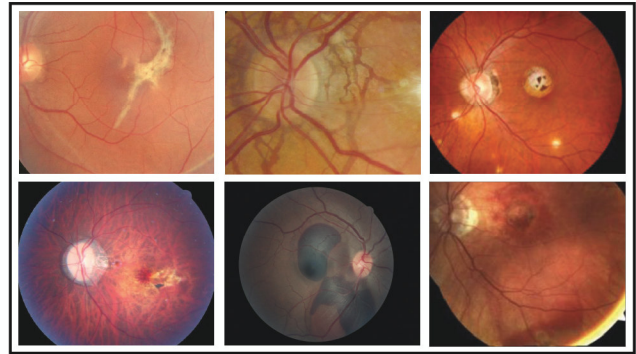
### AMD and Stem Cell-Based Therapies

Joseph C Giacalone, David H Parkinson, Daniel A Balkov, Rajesh C Rao

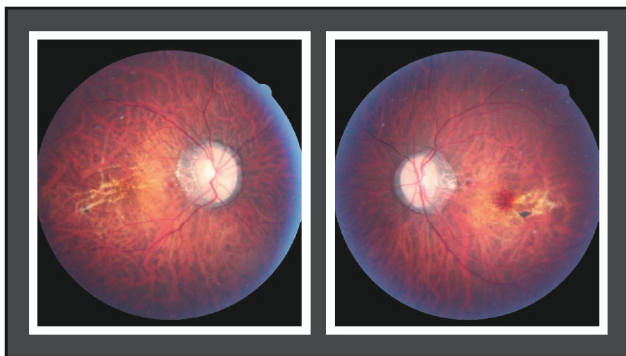
PMID: 38146879 PMCID: PMC10783850 (available on 2025-01-01)  
DOI: 10.1097/IIO.0000000000000510

**Abstract**  
Age-related macular degeneration (AMD) is a prevalent and complex disease leading to severe vision loss. Stem cells offer promising prospects for AMD treatment as they can be differentiated into critical retinal cell types that could replace lost host retinal cells or provide trophic support to promote host retinal cell survival. However, challenges such as immune rejection, concerns regarding tumorigenicity, and genomic integrity must be addressed. Clinical trials with stem cell-derived retinal pigment epithelial cells have shown preliminary safety in treating dry AMD, but improvements in manufacturing and surgical techniques cell delivery are needed. Late-stage AMD poses additional hurdles, possibly requiring multi-layered grafts. Advancements in automation technologies and gene correction strategies show potential to enhance iPSC-based therapies. Stem cell-based treatments offer hope for AMD management, but further research and optimization are essential for successful clinical implementation.

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### Myopia 2024

**Non-pathologic myopia**

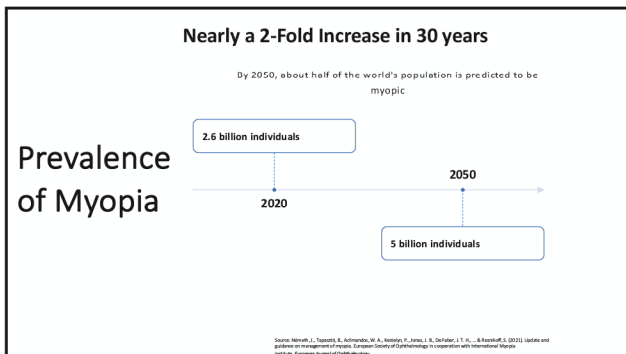
- Usually minimal to moderate (< 6.00 diopters)
- "high myopia"

**Pathologic myopia**

- (> -6.00D)
- Axial length ≥26.5 mm
- Major cause of visual impairment worldwide
- 3% of the world's population
- Highest prevalence in Asian populations
- A common cause of legal blindness in young individuals

Task Force on Myopia. Reducing the global burden of vision by slowing the onset of myopia and reducing myopia progression in children. The Lancet's Task Force on Myopia. Ophthalmology. 2023;130(8):1610-1618.

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### Risk Factors

#### PATHOLOGIC M

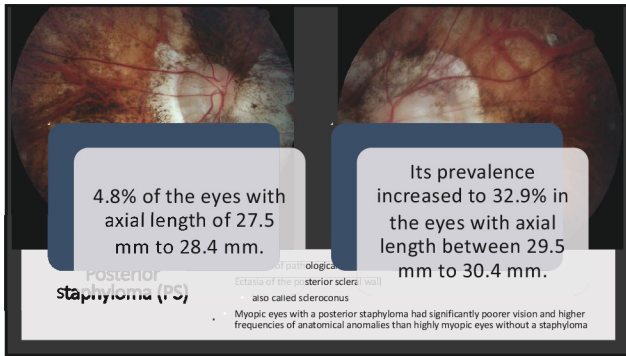
- Posterior staphyloma
- Atrophy (peripapillary)
- Thin sclera and choroid / Tessellations
- Holes in the retina
- Optic disc tilting
- Lacquer cracks / Lattice degeneration
- Ovoid patchy atrophy
- Glaucoma (↑ risk)
- Intrachoroidal cavitations (peripapillary)
- Choroidal neovascularization

**Macular schisis**

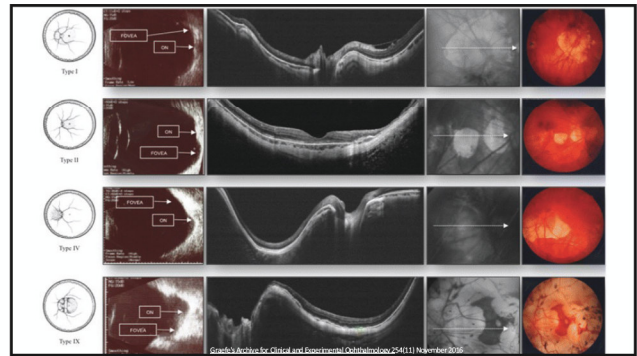
- tilted optic disc
- ovoid patchy atrophy
- tessellation
- thinned sclera
- iris transillumination opening
- post-staphyloma
- breaks in Bruch's membrane
- macular schisis
- pigmented CNV (Fuchs spot)

"The most significant predictor of visual acuity in highly myopic eyes with no macular pathology is 'suprachoroidal thickness'" (BCCO Retina 2022-2021: 213)

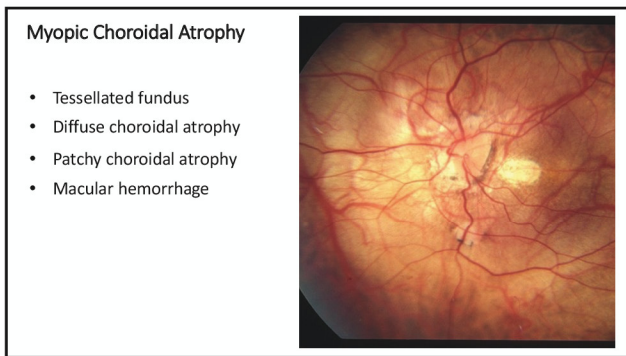
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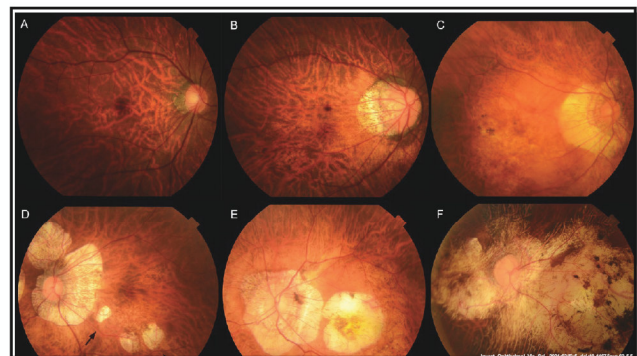
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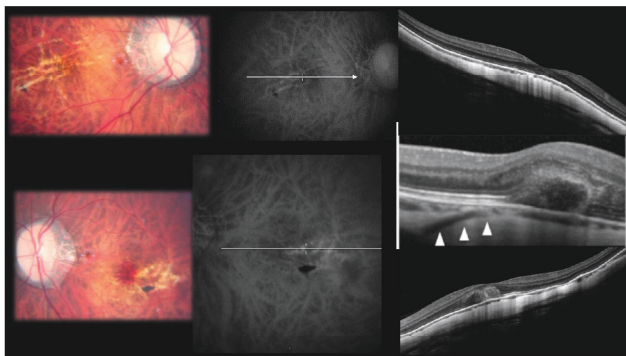
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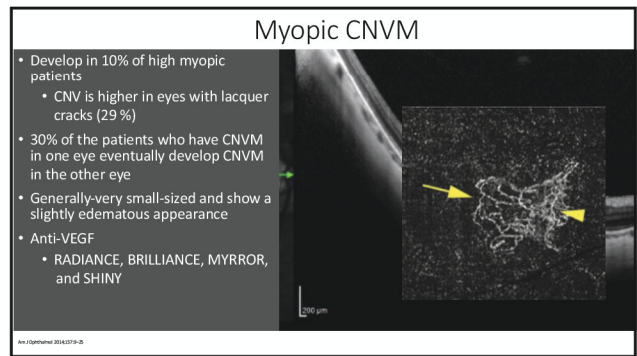
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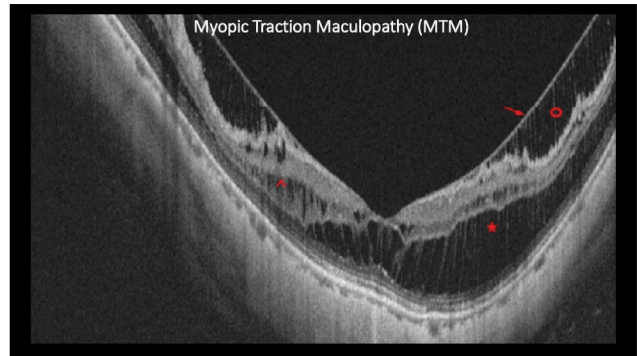
**Myopic CNV**

Three phases

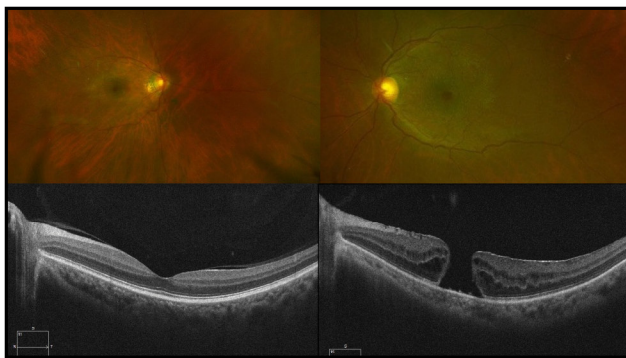
- Active phase with proliferation of a fibrovascular membrane including CNV, exudation, and hemorrhage
- Scar phase exemplified by a Fuchs spot
- Atrophic phase



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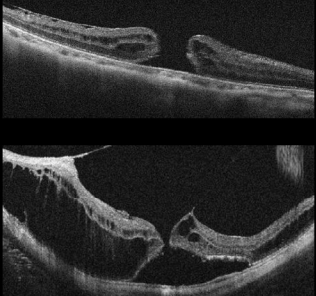


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**Myopic Macular hole**

Two types of macular holes in highly myopic eye

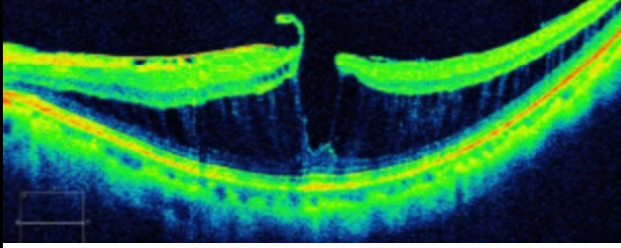
1. One is the type with the edge of the hole thickened with retinal cysts
2. Myopic foveoschisis



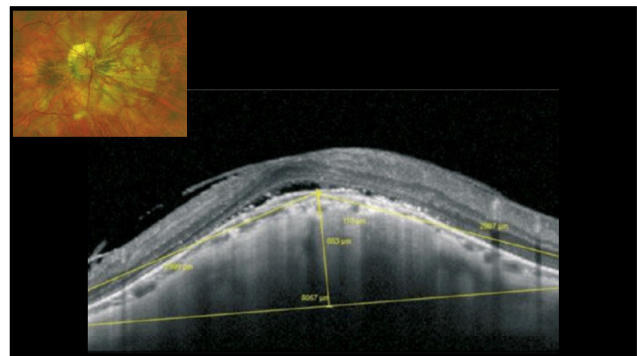
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**Myopic foveoschisis**

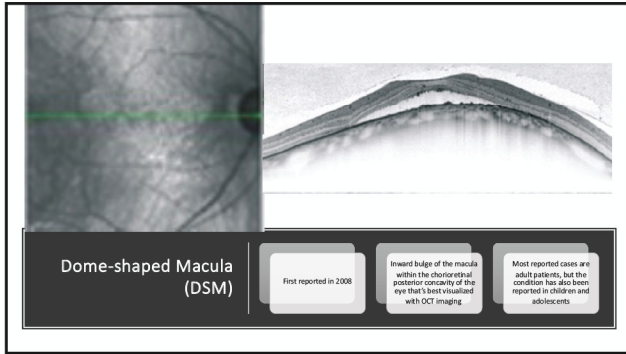
- Myopic macularschisis
- Vitreoretinal interface (VRI) traction
- First described by Takano and Kishi in 1999
- Splitting in the inner nuclear layer and outer plexiform layer



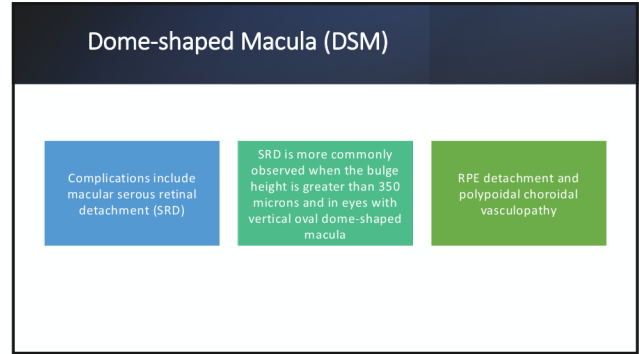
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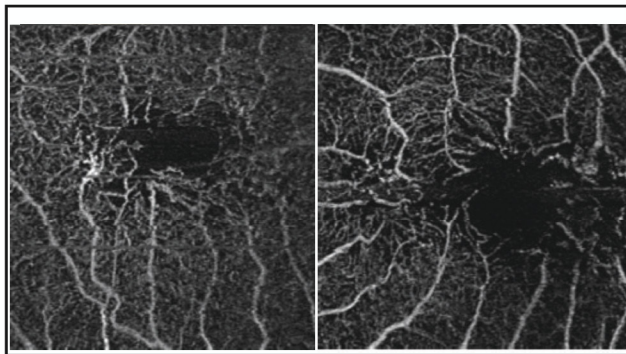
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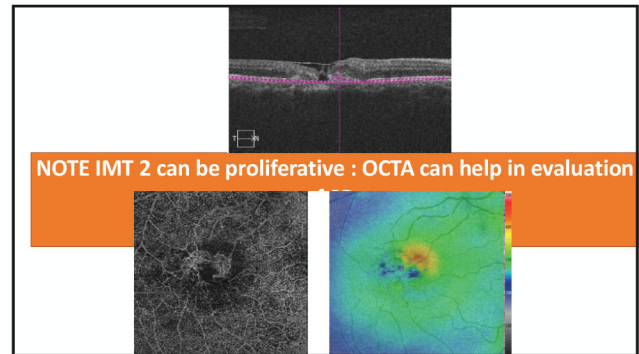
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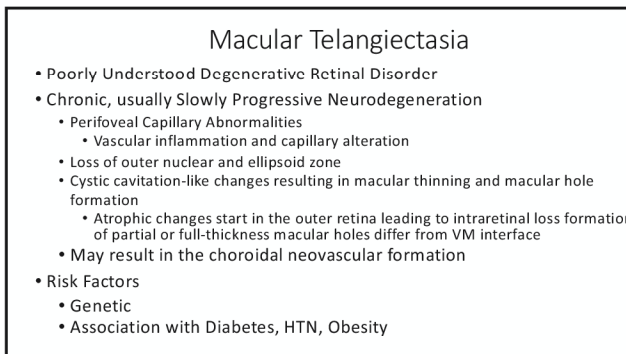
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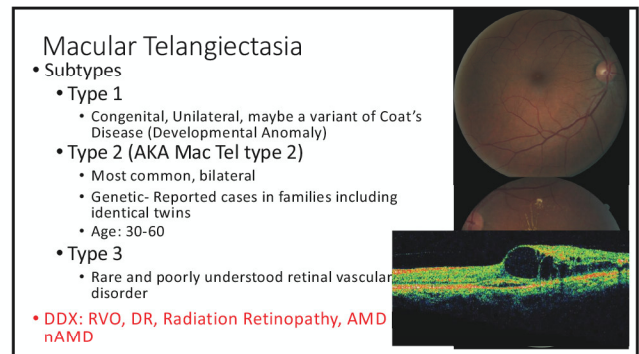
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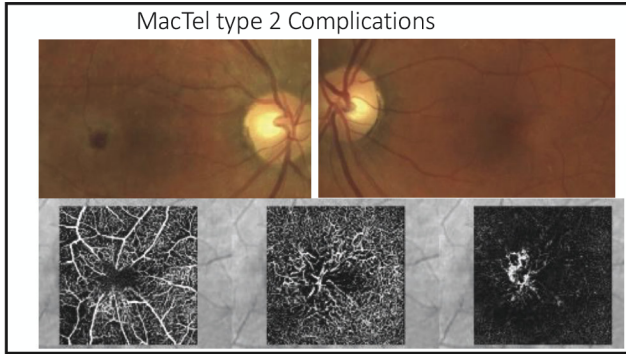
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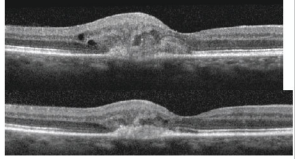
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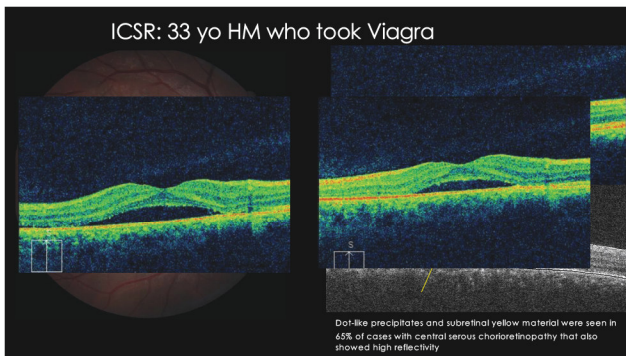
85

### Management and Prognosis

- Focal laser and PDT (Type 1)
- Steroid Injections
  - Reduce Inflammation and Fluid but does not reverse outer retinal loss
- Anti-VEGF
  - For CNVM
  - Improves fluid but does not reverse outer-retinal loss
- Ciliary neurotrophic factor (CNTF)
  - Under Investigation



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### Central Serous Chorioretinopathy (CSCR)

Fourth most common retinopathy

**Classification**

- Acute
- Isolated or multifocal RPE or serous detachment(s)
- Bullous serous retinal detachment
- Chronic CSCR
- Retinal pigment epithelium depigmentation

**Etiology**

- Type A personality-catecholamine release, which increases choroidal permeability
- Psychological stress and depression, sleep apnea, medications (steroids, MEK inhibitors, pseudoephedrine, Methylenedioxymethamphetamine (MDMA) or ecstasy, sildenafil), *H. pylori* infection, HTN, endocrine disorders

**Recurrences**

- Recurrence reported to be as high as 50%

Central serous chorioretinopathy. Ophthalmologica 2014; 232: 65-76

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### Choroidal Morphology

The thickness of the choroid varies throughout the posterior pole  
 Typically, thickest beneath the fovea, where its average thickness is 320µm  
 The temporal choroid is thicker than the nasal choroid

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### Pachychoroid Disease Spectrum (PDS)

Central Serous Chorioretinopathy (CSCR)	Pachychoroid Pigment Epitheliopathy (PPE)	Pachychoroid Neovascularopathy (PNV)	Polypoidal Choroidal Vasculopathy (PCV)
Peripapillary Pachychoroid Syndrome (PPS)	Focal Choroidal Excavation (FCE)	Peripapillary Pachychoroid Neovascularopathy	Peripheral Exudative Hemorrhagic Chorioretinopathy

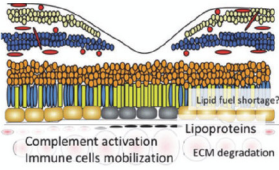
© Brown R, Mohan S, Chhablani J. Pachychoroid Spectrum Disorders: An Updated Review. J Ophthalmic Vis Res. 2023 Apr 29;38(2):212-229.  
 © Brown R, Mohan S, Chhablani J. Pachychoroid Spectrum Disorders: An Updated Review. Ophthalmology. 2023 Apr; 30(4):1212-1220. Available on PubMed. Copyright 2023 for National Eye Institute. All rights reserved. When text changes are indicated, please refer to the original source. 2023 31(1):118-124. doi:10.1016/j.ophtha.2023.04.018

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### Pachychoroid Disease Spectrum (PDS)

Pachychoroid = thicken choroid

- Abnormal and permanent increase in choroidal thickness
  - (choroidal thickness  $>320 \mu\text{m}$ )
- Larger Haller layer vessels (**Pachyvessels**) and medium vessels of Sattler layer & choriocapillaris present or effaced (atrophy)
- Reduced fundus tessellation- thinning of the overlying inner choroid
- Retinal pigment epithelium (RPE) abnormalities
- Choroidal vascular hyperpermeability (CVH)
- A lack of soft-drusen (an exception is made for **pachydrusen**, which are irregular, scattered yellow-white deposits across the posterior pole)

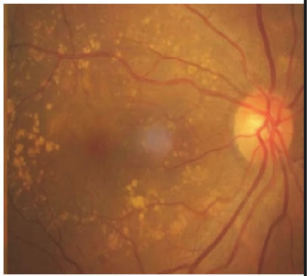


**Keynote Article**  
 Pachychoroid Spectrum Disorders: An Updated Review  
 Brown N, Mahan S, Chhabri S. Pachychoroid Spectrum Disorders: An Updated Review. J Ophthalmic Vis Res. 2023 Apr;18(2):212-229.  
 Gallego-Ponce A, et al. Med Hypotheses Disor Inve. 2014;10(4):111-114.

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### Pachydrusen

- Pachydrusen are large
  - Typically,  $>125 \mu\text{m}$
  - sub-RPE deposits that are yellow-white in color
- Deposits are distributed across the posterior pole and are isolated or clustered in small groups.
- The deposits appear with irregular, complex shapes but have distinct borders.
- Another important distinguishing feature is that these drusenoid lesions are associated with the presence of thickened choroid.



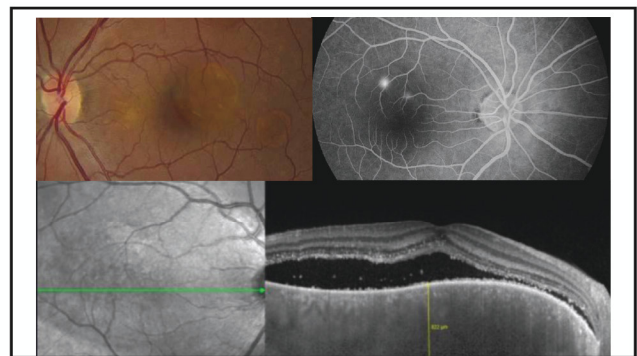
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### Pachychoroid Disease Spectrum (PDS)

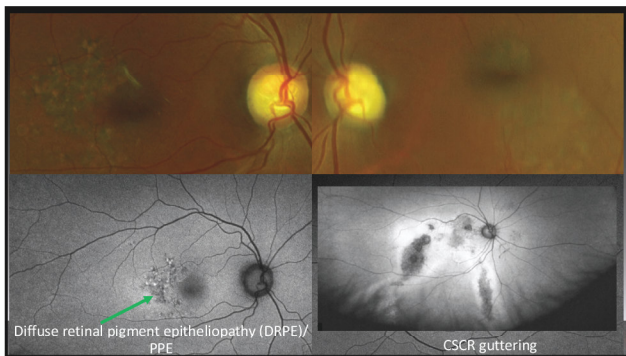
- Choroidal vascular congestion/ attenuation
  - Thickened sclera
  - Lengthened intrascleral course of vortex veins
- Physiologic Factors
  - Excess choroidal interstitial fluid/choroidal vascular hyperpermeability (CVH)
  - Precapillary arteriolar hypertension
  - Altered intravascular osmolality ( serum proteins (albumin))
  - Pharmacologic agents
    - Corticosteroids
    - Phosphodiesterase (PDE) inhibitors
  - Alterations in interstitial tissues in the choroid

**Keynote Article**  
 Pachychoroid Spectrum Disorders: An Updated Review  
 Brown N, Mahan S, Chhabri S. Pachychoroid Spectrum Disorders: An Updated Review. J Ophthalmic Vis Res. 2023 Apr;18(2):212-229.  
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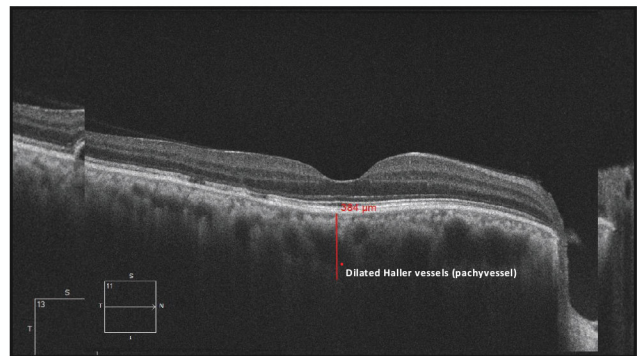
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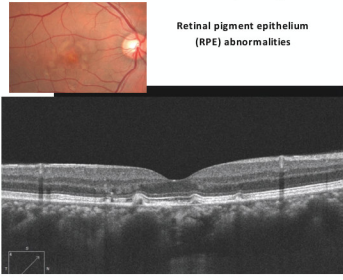
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### PACHYCHOROID PIGMENT EPITHELIOPATHY (PPE)

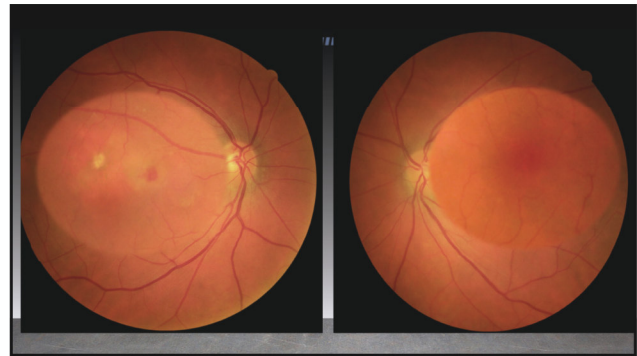
- Forme Fruste of CSC to Chronic CSC
- Orange red fundus appearance
- Absence of normal fundus tessellation
- RPE changes mistaken for ARMD or pattern dystrophy
- OCT scattered RPE elevations, small serous PEDs, thick choroid
- ICG shows mid-phase hyperfluorescence suggestive of hyperpermeability
- Fundus Autofluorescence shows granular hypoautofluorescence and stippled mixed areas of hyper and hypoFAF
- Management: Observation for progression to CSC, PVN, or even PCV



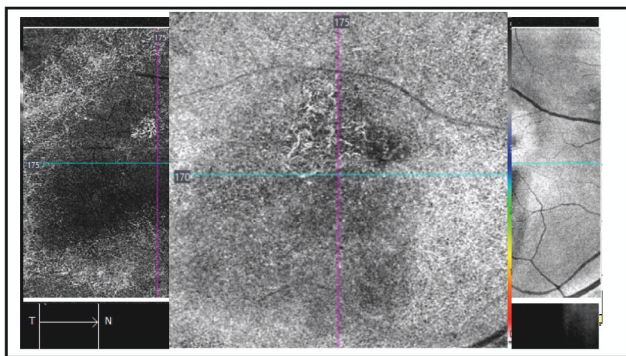
Retinal pigment epithelium (RPE) abnormalities

Galligo-Pinazo R, et al. Med Hypothesis Discov Innov. 2014;X(6):111-114

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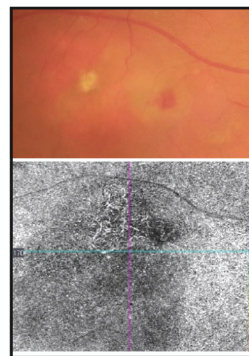
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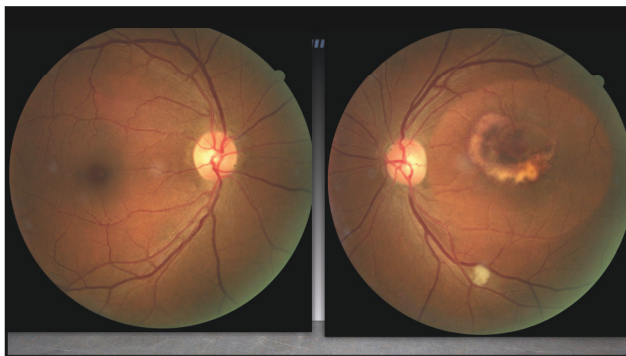
### Pachychoroid Neovascularopathy (PNV)

- Characteristics:
  - OCT Findings of CSC and/or PPE
  - OCT (A)
    - Type 1 sub RPE CNV
    - With or without subretinal fluid
    - Double-layer Sign

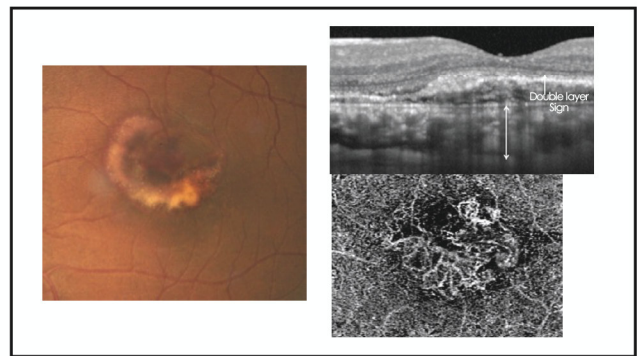


Galligo-Pinazo R, et al. Med Hypothesis Discov Innov. 2014;X(6):111-114.

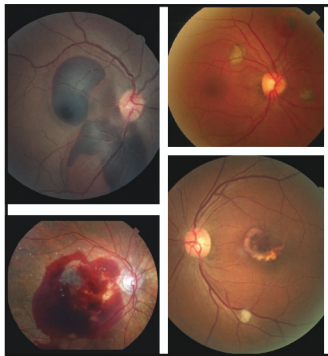
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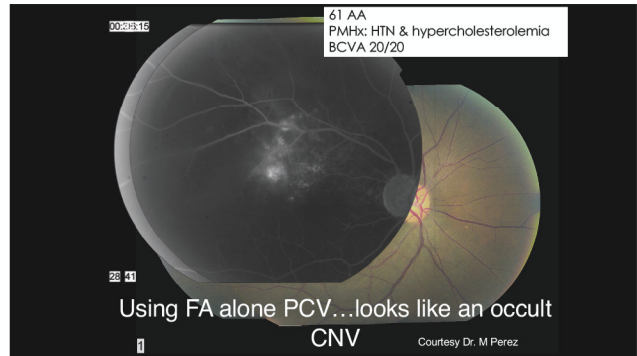
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**POLYPOIDAL CHOROIDAL VASCULOPATHY (PCV)**

- INTRACHOROIDAL vascular abnormalities
  - Abnormal branching vascular network (BVN) with terminal aneurysmal red spheroidal dilations (polypoidal lesions) Dilated, thin-walled vessel of the choriocapillaris
- Recurrent hemorrhage and leakage
- Two Types:
  - Idiopathic type
  - nPCV
    - Type I CNVM variant
    - #1 Misdiagnosis is AMD

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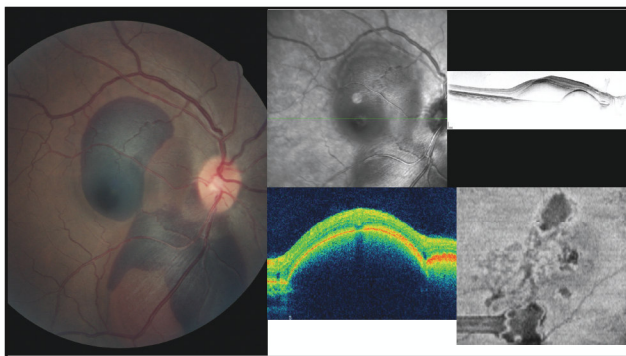


61 AA  
PMHx: HTN & hypercholesterolemia  
BCVA 20/20

Using FA alone PCV...looks like an occult CNV

Courtesy Dr. M Perez

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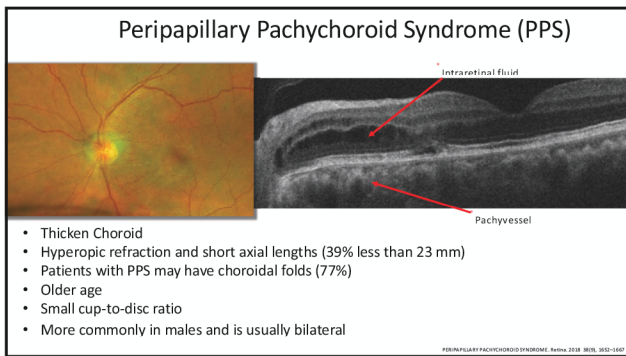
**PCV Treatment**

- Asymptomatic lesions may be observed
  - Lesions may spontaneously resolve
- Anti-VEGF, PDT, or both
  - EVEREST-II trial showed combination therapy of ranibizumab plus verteporfin PDT are superior to ranibizumab monotherapy
- Thermal laser photocoagulation of feeder vessels or polyps
- Prognosis is better than neovascular AMD

EVEREST study efficacy and safety of verteporfin photodynamic therapy in combination with ranibizumab or alone versus ranibizumab or verteporfin in subjects with symptomatic neovascular polypoidal choroidal vasculopathy. Retina. 2012;32(12):1435-46.

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**Peripapillary Pachychoroid Syndrome (PPS)**

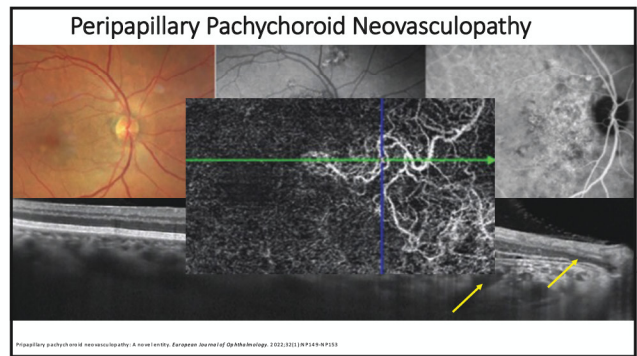


- Thicken Choroid
- Hyperopic refraction and short axial lengths (39% less than 23 mm)
- Patients with PPS may have choroidal folds (77%)
- Older age
- Small cup-to-disc ratio
- More commonly in males and is usually bilateral

PERIPAPILLARY PACHYCHOROID SYNDROME. Retina. 2018; 38(5): 952-957

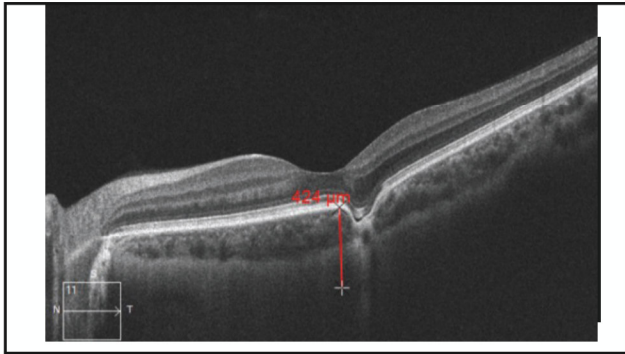
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**Peripapillary Pachychoroid Neovascularopathy**

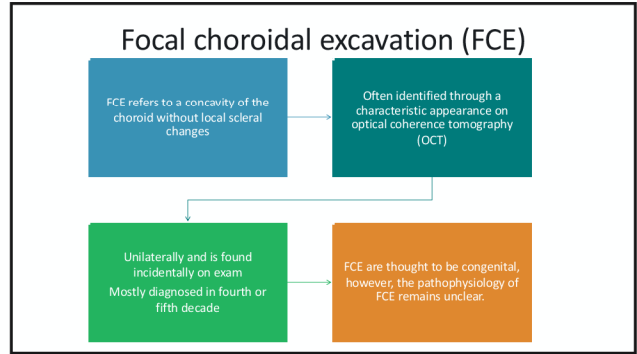


Peripapillary pachychoroid neovascularopathy: A novel entity. European Journal of Ophthalmology. 2022;32(12):1814-1823

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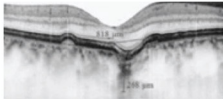
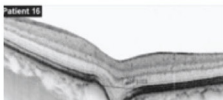
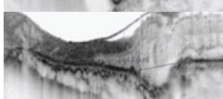


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### Focal Choroidal Excavation (FCE)

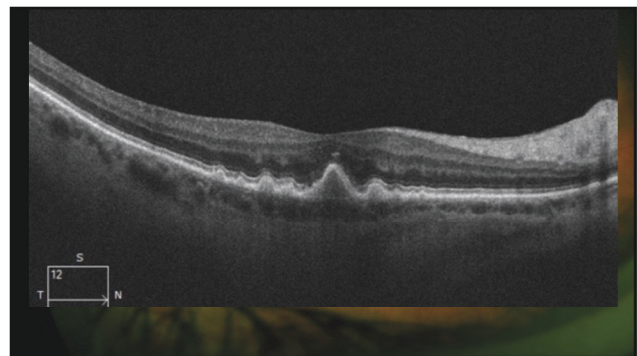
**Three patterns**

1. cone shaped—Most common pattern; all these cases had regular RPE on OCT and less degenerative changes on angiography, hence bearing a better prognosis.
2. Bowl shaped—Higher incidence of RPE disruptions on OCT and degenerative changes on angiography.
3. Mixed morphology—This pattern has features of both cone- and bowl-shaped FCEs.

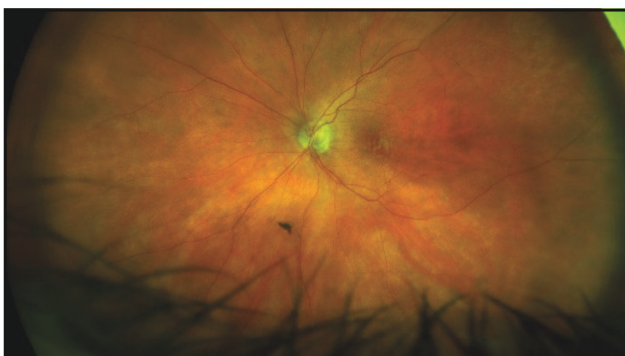




Shinojima A, Kawamura A, Mori R, et al. Morphologic features of focal choroidal excavation on spectral domain optical coherence tomography with simultaneous angiography. Retina. 2014;34(1):467-74

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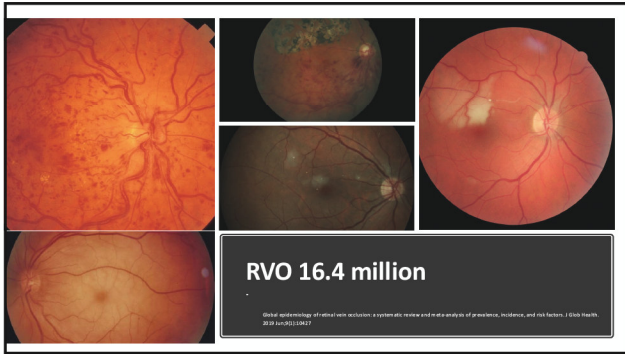
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### Peripheral Exudative Hemorrhagic Chorioretinopathy (PEHCR)

First described by Annesley in 1980 as Peripheral Exudative hemorrhagic chorioretinopathy (PEHCR)	Bilateral	Predominantly affects older women	Signs include Peripheral subretinal hemorrhage, subretinal fluid (SRF), exudation, and PED
Patients may be asymptomatic or have decreased visual acuity and flashes and floaters	Has been postulated to be part of the Pachychoroid disease spectrum	One study found that the choroid in eyes with PEHCR is thickest in the temporal periphery	Club-shaped choroidal contour

Peripheral exudative hemorrhagic chorioretinopathy: a new addition to the spectrum of pachychoroid disease? Retina 2021;41:1318-1326.

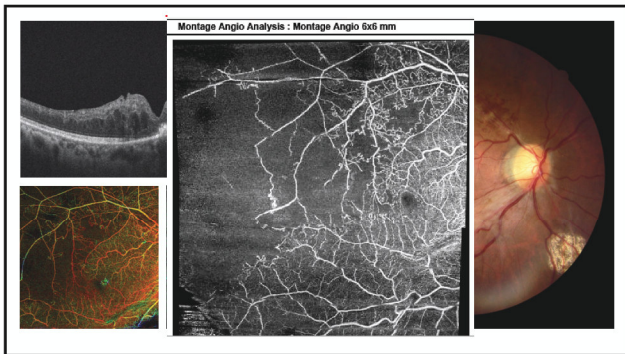
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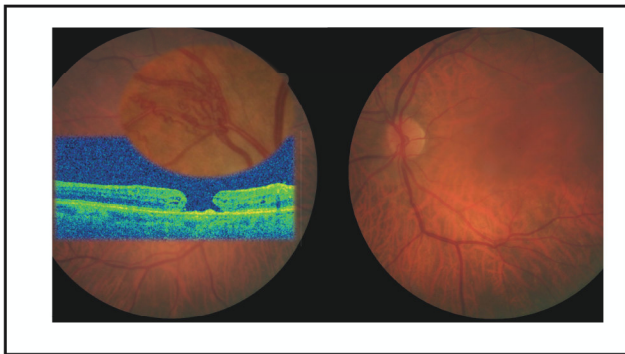
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### Blood Pressure Categories

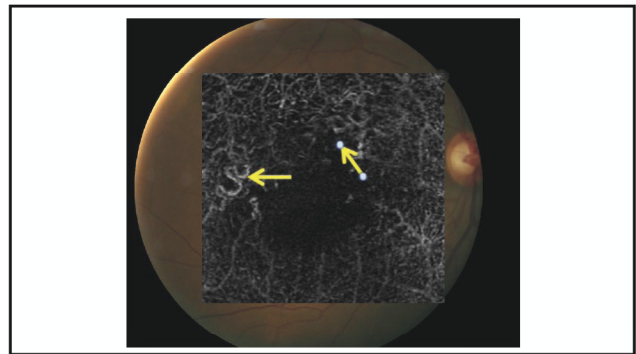
BLOOD PRESSURE CATEGORY	SYSTOLIC mm Hg (upper number)		DIASTOLIC mm Hg (lower number)
NORMAL	LESS THAN 120	and	LESS THAN 80
ELEVATED	120 - 129	and	LESS THAN 80
HIGH BLOOD PRESSURE (HYPERTENSION) STAGE 1	130 - 139	or	80 - 89
HIGH BLOOD PRESSURE (HYPERTENSION) STAGE 2	140 OR HIGHER	or	90 OR HIGHER
HYPERTENSIVE CRISIS (consult your doctor immediately)	HIGHER THAN 180	and/or	HIGHER THAN 120

BP indicates blood pressure (based on an average of  $\geq 2$  careful readings obtained on  $\geq 2$  occasions, as detailed in DBP, diastolic blood pressure, and SBP systolic blood pressure.)

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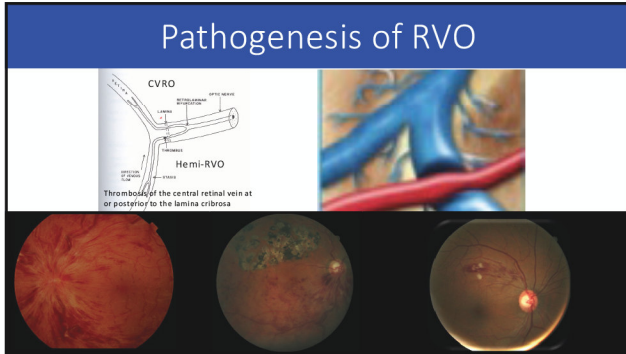


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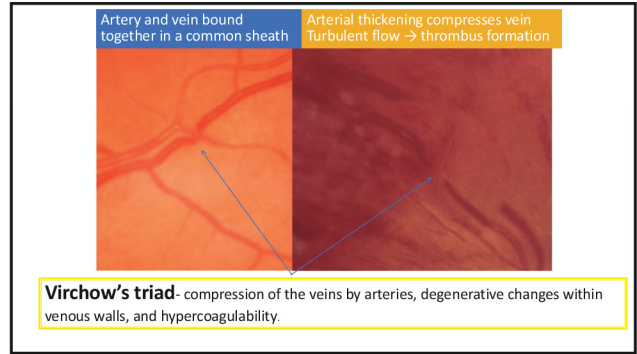


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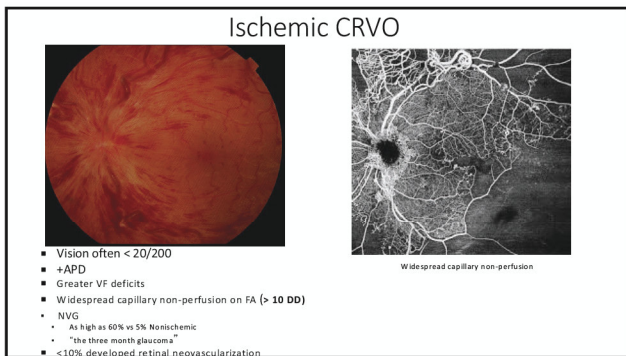




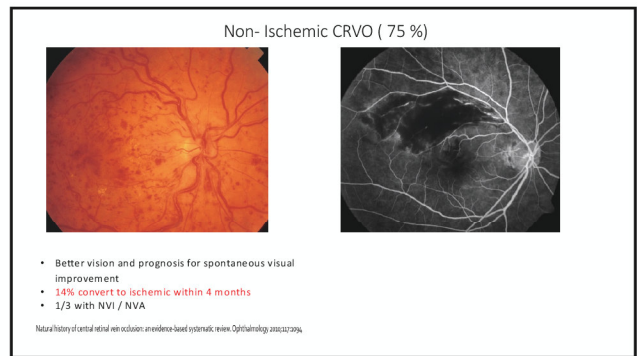
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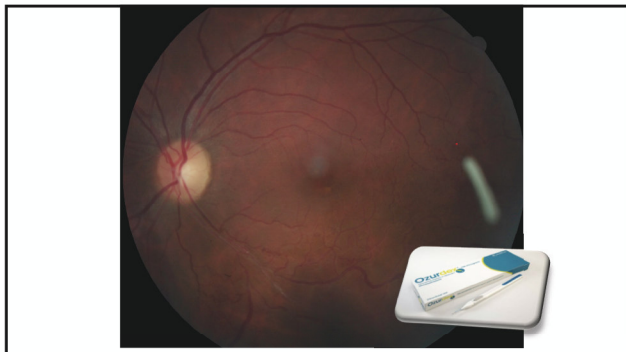
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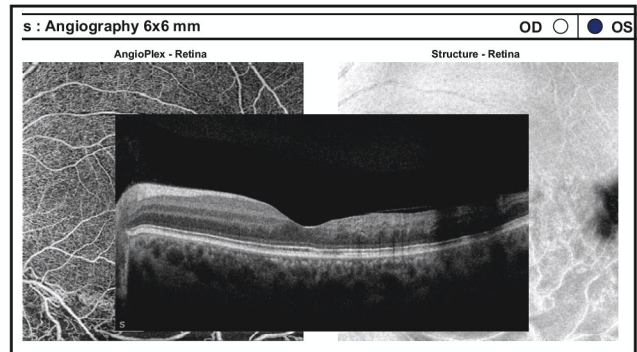
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### VABYSMO (Faricimab) in AMD

FDA approved January 2022

10.27.2023

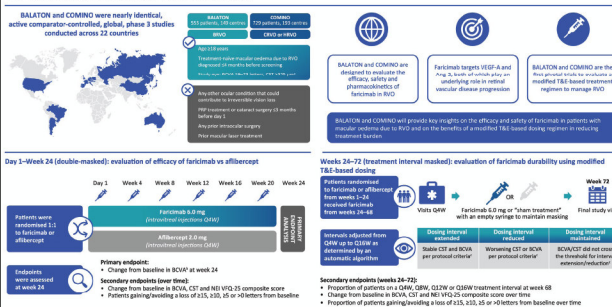
#### FDA Approves Genentech's Vabysmo for the Treatment of Retinal Vein Occlusion (RVO)



A humanized, bispecific monoclonal antibody, specifically designed for intravitreal use, that simultaneously binds and neutralizes both VEGF-A and Angiopoietin-2

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### Study Design of the BALATON and COMINO Phase 3 Randomised Controlled Trials of Faricimab in Patients With Retinal Vein Occlusion



BALATON and COMINO were nearly identical, active comparator-controlled, global, phase 3 studies conducted across 22 countries.

- MAXIMIZE:** 500 patients, 250 centers
- MINIMIZE:** 272 patients, 167 centers

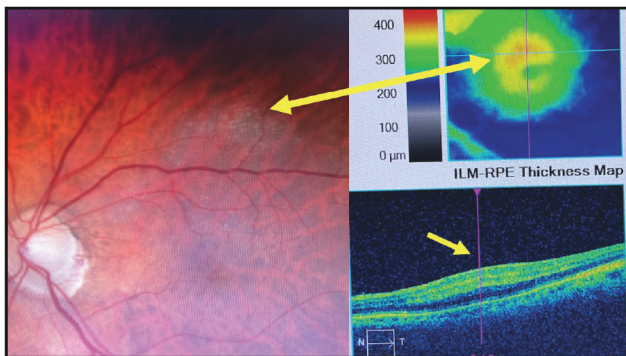
**Key Features:**

- Treatment naïve macular edema due to RVO (presence of RVO) before screening
- Any other ocular condition that could interfere with the study
- RPV treatment or vitreal surgery (3 months before Day 1)
- Any prior intravitreal surgery
- RPV resolved before treatment

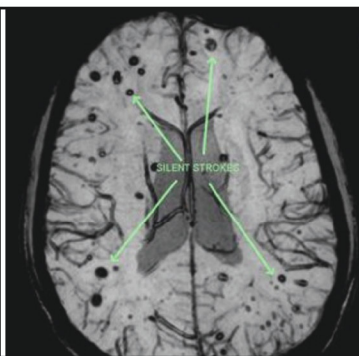
**Study Objectives:**

- Primary endpoint:** Change from baseline in BCVA at week 24
- Secondary endpoints (over time):**
  - Change from baseline in BCVA, CST and NEI VFQ-25 composite score
  - Patients gaining/avoiding a loss of ≥15, ≥30, ≥45 or ≥60 letters from baseline
- Secondary endpoints (weeks 24-72):**
  - Proportion of patients in CSW, CSW, CS2W or CS2W treatment interval at week 68
  - Change from baseline in BCVA, CST and NEI VFQ-25 composite score over time
  - Proportion of patients gaining/avoiding a loss of ≥15, ≥30, ≥45 or ≥60 letters from baseline over time

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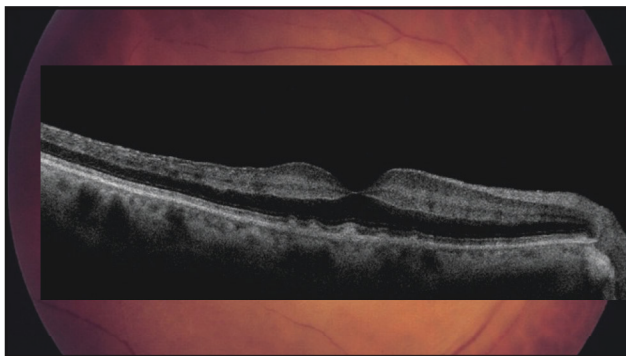
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### Diffusion-weighted imaging (DWI) MRI

- DWI- MRI revealed: signs of 2 small silent strokes
- Pt placed on blood thinner

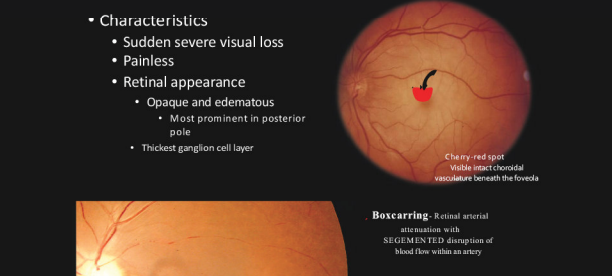
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### Central Artery Occlusion (CRAO)

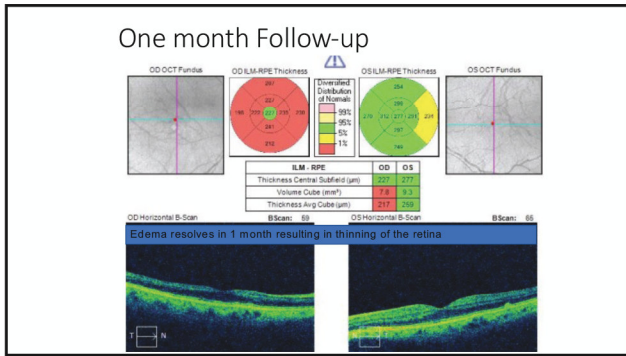
- Characteristics**
  - Sudden severe visual loss
  - Painless
  - Retinal appearance
    - Opaque and edematous
      - Most prominent in posterior pole
    - Thickest ganglion cell layer



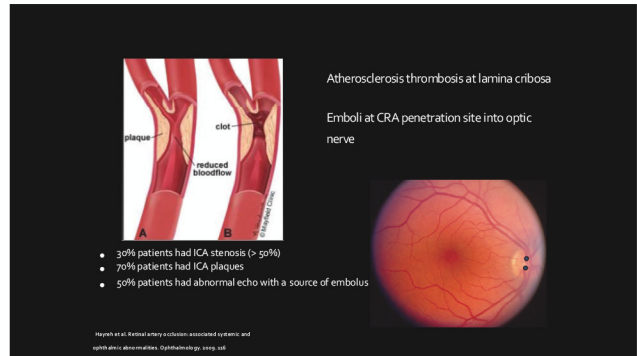
**Cherry-red spot:** Visible intact choroidal vasculature beneath the fovea

**Boxcarring:** Retinal arterial attenuation with SEGMENTED dysregulation of blood flow within an artery

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- Non-Arteritic CRAO ~66.9%
  - 93.2% CF or worse, none better than 20/40
- Non-Arteritic CRAO w/ clioretinal sparing ~14.3%
  - 60% CF or worse, 20% better than 20/40
  - Patient clioretinal artery improve visual prognosis
- Transient Non-Arteritic CRAO ~4.5%
  - CRA temporarily occluded
  - Fall in perfusion pressure, drop arterial BP or rise IOP, vasospasm
- Arteritic CRAO ~16%
  - Secondary to Giant Cell Arteritis (GCA)
  - Posterior ciliary artery and central retinal artery occlusion
  - Occult GCA – no systemic symptoms, always order labs to r/o

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### Update on the Management of Central Retinal Artery Occlusion

Michael Dattilo, MD, PhD<sup>1,2</sup>, Valérie Biousse, MD<sup>1,2,3,4,5</sup>, Nancy J. Newman, MD<sup>1,2,3,4,5</sup>

**KEYWORDS**  
 • Central retinal artery occlusion • Rate for stroke peaks (~60%) within 1 week s/p RAO onset • Thrombolysis • Management • Treatment • Thrombolysis • Ischemia

**CRAO, BRAO, & TVOs = a stroke & needs to be recognized as an EMERGENCY!**  
**Urgent referral to ER (with stroke center) is CRUCIAL**  
**Management requires identifying and treating risk factors + neuro consult + cardio eval**

**FURTHERMORE**  
**HTN/Hypercholesterolemia or other risk factor need to be identified and treated**

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### Retinal Artery Occlusion Emergent Stroke Evaluation

CRAO, BRAO, & TIAs = a stroke & needs to be recognized as an EMERGENCY!

- Silent POSITIVE DWI-MRI strokes are seen in:
  - >55% of CRAO
  - ~31% of BRAO
  - ~18% OF TVO
- Pts with (+) DWR-MRI silent strokes have a High risk for MAJOR stroke
  - Especially during the next week to month
  - Rate for stroke peaks (~60%) within 1 week s/p RAO onset
- Urgent referral to ER (with stroke center) is CRUCIAL
  - Management requires identifying and treating risk factors + neuro consult + cardiology evaluation

**Update on the Management of Central Retinal Artery Occlusion**

Michael Dattilo, MD, PhD<sup>1,2</sup>, Valérie Biousse, MD<sup>1,2,3,4,5</sup>, Nancy J. Newman, MD<sup>1,2,3,4,5</sup>

**KEYWORDS**  
 • Central retinal artery occlusion • Branch retinal artery occlusion • Stroke • Ischemia • Management • Treatment • Thrombolysis

**KEY POINTS**

• Acute central retinal artery occlusion (CRAO) and branch retinal artery occlusion (BRAO) are the retinal equivalent of a cerebral infarction in the anterior circulation.

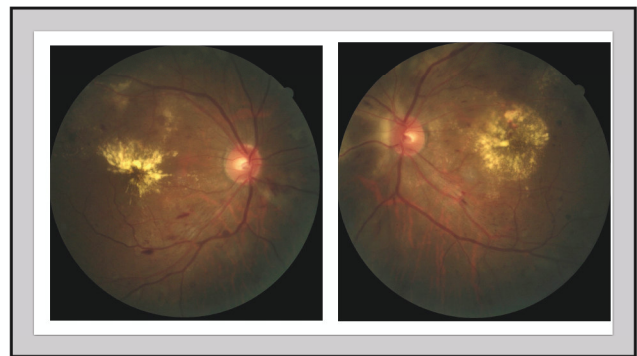
• The risk factors for a CRAO or BRAO and acute central ischemia are very similar.

• Patients with acute CRAO and BRAO need to be evaluated emergently in a stroke center similar to patients with cerebral ischemia.

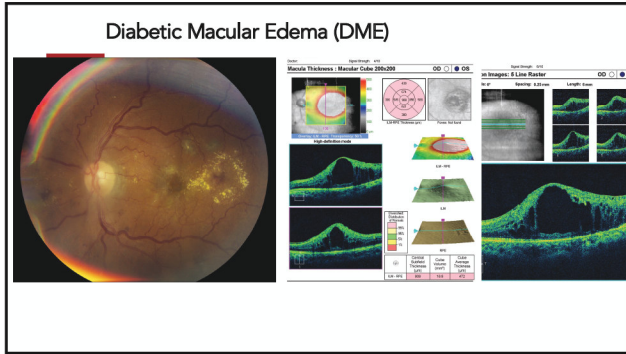
• Up to 20% of patients with acute retinal ischemia have concomitant cerebral infarctions on brain CT/MRI performed 1-10 days after onset.

• Because the current benefit of intravenous thrombolysis has been shown to improve visual outcome compared with the initial history of CRAO, management of CRAO should be focused on secondary prevention of cerebral events, such as cerebral ischemia, myocardial infarction, and cardiovascular death.

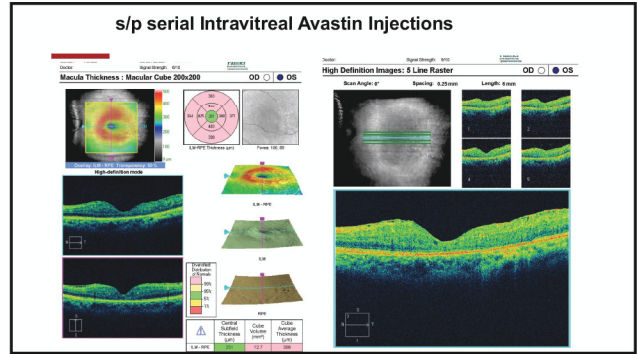
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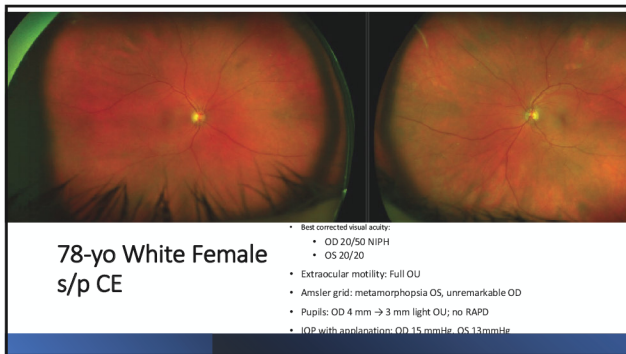
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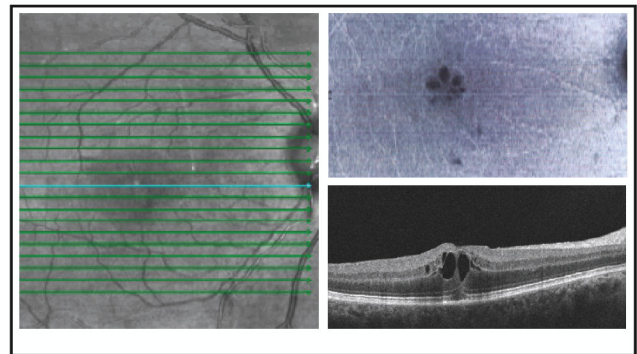
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**Irvine-Gass Syndrome**

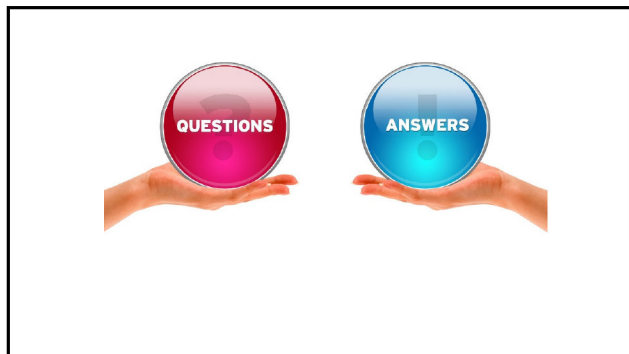
- Most common causes of visual loss after uneventful cataract surgery
- Benign, self-limiting, and resolves spontaneously without visual impairment
  - 26.8% of eyes with pseudophakic CME did not recover
- Represents breakdown of blood-retinal barrier due to inflammation
  - VEGF

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**Pseudophakic Cystoid Macular Edema**

- RTCX 3-4 weeks to determine improvement
- Persistent CME
  - steroid injection
  - Anti-VEGF drugs
  - Surgical therapy
    - Pars plana vitrectomy (PPV)

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