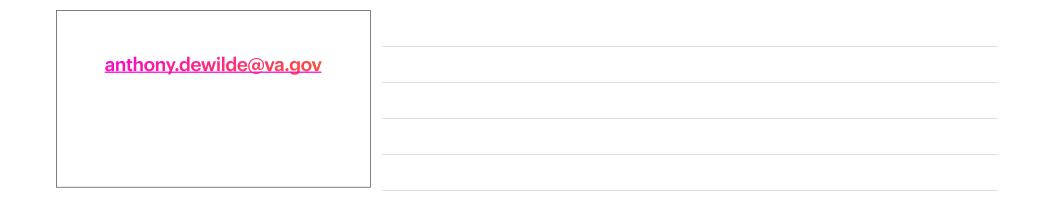
Advanced Technology in Eyecare Biologics and OCT Grand Rounds	
Anthony DeWilde, OD FAAO	



No Financial Disclosures	
NO FINANCIAI DISCIOSURES	









# **Biologics Utility**

Most common uses Anti-TNF Anti-Interleukin Anti-CGRP

## **Biologics Utility**

#### Autoimmune Disease

Inflammatory Bowel Disease Psoriasis Ankylosing Spondylitis Graves' Disease Myasthenia Gravis

Biologics Utility	
<b>Headache</b> Migraine	
Cluster Headache	

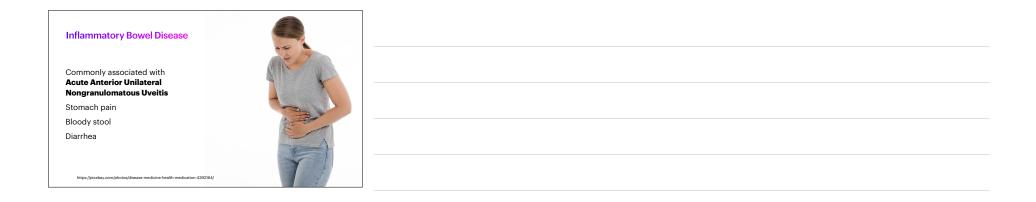




Side Effects Injection site rash Infection Headache	Cimzia - My First Injection(s)	

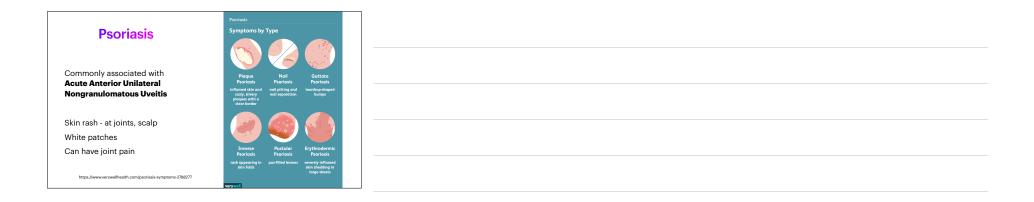






Sees Gastroenterologist	
Sees Gastroenterologist Endoscopy Colonoscopy Stool samples CT/MRI	
Steel complete	
Stool samples	
CT/MRI	

Treatment includes	
Corticosteroids	
Aminosalicylates	
Immune suppression	
(Azathioprine, Methotrexate)	
Biologics	
(Remicade, Humira, Stelara, etc)	
(	







#### **Ankylosing Spondylitis**

Commonly associated with Acute Anterior Unilateral Nongranulomatous Uveitis

Lower back pain Worse with rest Better with movement Better with NSAID



Possible combination of clinical, laboratory, or imaging SpA features	Post-test probability
IBP plus family history	51%
IBP plus heel pain	35%
IBP plus uveitis	54%
IBP plus synovitis	39%
IBP plus dactylitis	42%
IBP plus family history plus heel pain	78%
IBP plus uveitis plus NSAID*	85%
IBP plus heel pain plus synovitis plus alternating buttock pain	89%
IBP plus family history plus heel pain plus NSAID*	95%
IBP plus heel pain plus HLA-B27	83%
IBP plus NSAIDs* plus HLA-B27	88%
IBP plus heel pain without HLA-B27	6%
IBP plus NSAIDs* without HLA-B27	8%
IBP plus dactylitis plus ESR/CRP	62%
IBP plus HLA-B27 plus ESR/CRP	78%
IBP plus HLA-B27 without ESR/CRP	47%
IBP plus HLA-B27 plus MRI	93%
IBP plus HLA-B27 without MRI	14%
IBP plus heel pain plus HLA-B27 without MRI	35%
The pretest probability of low back pain is assumed to be 5%. IBP-inflammatory bac CRP=C-reactive protein. *A good response to NSAIDs is needed. Adapted from Radu	
BMJ Publishing Group.	valence al with permission of

On an Dharman ta la mu	
Sees Rheumatology	
X-ray CT MPI	
X ruy, or, mitt	
Sees Rheumatology X-ray, CT, MRI Possible HLA B-27 testing	
Possible HLA B-27 testing	

** IL-17	
Treatment includes	
NSAID	
Physical Therapy	
Anti-TNF biologics	
(Humira, Enbrel, Remicade, Cimzia)	
Anti-IL biologics**	
(Cosentyx, Taltz)	



Classic vs Common Unilateral, painful headache Aura Nausea, vomiting Light/sound sensitivity

Mostly women





Nearly 1 in 4 US households includes someone with Migraine	
https://migraineresearch/oundation.org/about-migraine/migraine-facty/	
https://migraineresearchfoundation.org/about-migraine/migraine-facts/	

Sees Neurology CT, MRI - rule out other causes	
May need other testing (EEG, Lumbar Puncture)	

Treatment includes	
Preventative	
Preventative	
Abortive	

Preventative
Topamax
Beta Blockers
Verapamil
Amitriptilyine
Botox injections
Anti-CGRP (Ajovy, Emgality, Aimovig)





CGRP medication also approved for Cluster Headaches
--



		7
Ocular Disease	<b>Ocular Disease</b>	

#### **Giant Cell Arteritis**

Ocular concern is Arteritic ION Severe vision loss

Systemic concern is CVA/MI



ystemic Symptoms	
Fever	
Malaise	
Headache	
Scalp Tenderness	
Neck pain	
Jaw claudication	

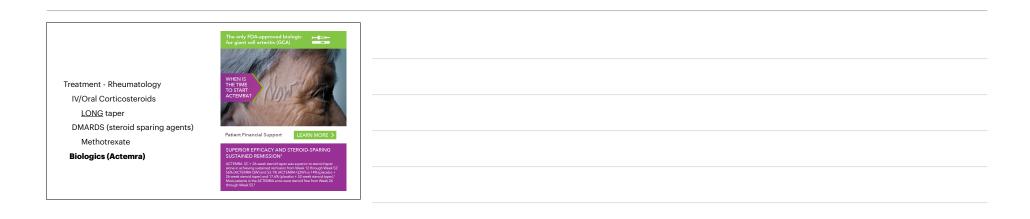
1/3 of AION patients become bilateral within 14 days	

Ocular Signs & Symptoms
"Pallid" Edema
Amaurosis Fugax (1/3)
Rarely diplopia
CRAO (5-15%)



Management of ischemic optic neuropathies Indian Journal Ophthal 2011. Vol 59, 2, 123-136





itudy	
Groups	
Actemra weekly vs bi-weekly (with 26 week pred taper)	
lacebo (with 26 week pred taper)	
lacebo (with 52 week pred taper)	
Stone, J H, et al. Trial of Tocilizumab in Giant Cell Arteritis. N Engl J Med 2017; 377:317-328	



Cumulative Dose of Prednisone	
Actemra weekly 1862 mg Actemra bi-weekly 1862 mg	
Placebo 26 week 3296 mg Placebo 52 week 3818 mg	

Mostly non-serious (like injection site reaction) More serious reaction in prednisone alone 25% vs 15%



Pain is much less than appearance would suggest	
Fair is much less than appearance would suggest	
Irritation	
Foreign Body Sensation Epiphora	
Epiphera	
Ерірпога	
Blur	
Prone to infection	

Lubrication	
Tears, gels, ointments	
Bandage CL	
Tarsorrhaphy	
Amniotic membranes	
Plasma Rich Protein (PRP) Biologics (Oxervate)	
Biologics (Oxervate)	



	7
If paperwork is filled out, patient can get much cheaper	
8-week course (patient is shipped 1 week at a time!!)	
Refrigerate - take 6x/day	

Studies	
1. Placebo healed 43% vs Oxervate 70%	
2. Placebo healed 29% vs Oxervate 70%	
1 - Bonini S, et al. Phase II randomized, double-masked, vehicle-controlled trial of recombinant human nerve growth factor for neurotrophic keratitis. Ophthalmology. 2018;125:1332–1343	
<ol> <li>Pflugfelder SC, et al. Topical recombinant human nerve growth factor (cenergermin) for neurotrophic keratopathy: A multicenter randomized vehicle-controlled pivotal trial. Ophthalmology. 2020;127:14–26.</li> </ol>	



O a con Dhanna a chun an Earda an ia chun an	
Sees Rheumatology, Endocrinology	
Thyroid scan	
myrold scan	
Lab tests (thyroid panel)	
Not just T3/T4 and TSH	
Needs Thyroid antibody testing	

Treatments Help Ocular problems Lubrication	
Diplopia	



Ocular Surgeries	
Ocular Surgeries	
Orbital Decompression	
Orbital Decompression	
Strabismus surgery Eyelid retraction	
Eyelid retraction	

What about Tepezza?	
---------------------	--



CAS	For initial assessment, only score items 1–7	
1	Spontaneous orbital pain	
2	Gaze evoked orbital pain	
3	Eyelid swelling; considered due to active TED	
4	Eyelid erythema	
5	Conjunctival redness; considered due to active TED	
6	Chemosis	
7	Inflammation of caruncle or plica	
	Follow-up assessment at I-3 months can be scored	
	out of 10	
8	Increase of >2mm in proptosis	
9	Decrease in uniocular excursion in any one direction of >8	
	degrees	
10	Decreased acuity equivalent to 1 Snellen line	

Tepezza is for active Graves' Orbitopathy	
Now approved for INACTIVE Thyroid Eye Disease!!	
Total of 8 infusions	
One infusion every 3 weeks Cost is <b>\$46,000</b> for 3 vials!!!	

FDA approved	
Inactive TED	
Muscle spasm Hyperglycemia (10%) Hearing loss (38%**)	
Live events (10%)	
Hypergiycernia (10%)	
Hearing loss (38%**)	

#### **Exudative AMD**

Choroidal Neovascular Membranes Well know benefit of Anti-VEGF These are all biologics



Ranibizu <b>mab</b> (Lucentis)	
Bevacizu <b>mab</b> (Avastin)	
Brolucizu <b>mab</b> (Beovu)	







### **Uveitis**

Humira is now FDA approved for treatment of noninfectious uveitis Most beneficial for patients with multiple bouts Reduce intensity Reduce recurrence



#### **Other conditions**

Juvenile Idiopathic Arthritis Myasthenia Gravis (still under research) Diabetes Rheumatoid Arthritis (Enbrel, Humira) Lupus Multiple Sclerosis

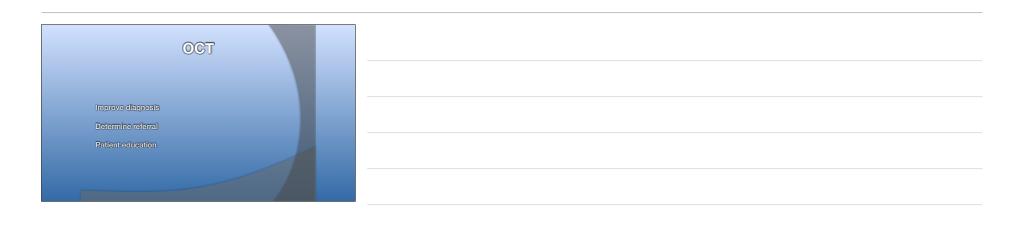
Other conditions	
Cancer	
Sickle Cell Anemia Osteoporosis	

GLP-1 Agonists
Trulicity Victoza
Ozempic**

Future Research
How do Biosimilars compare to Biologics?
Less frequent dosing?
More conditions approved?
Oral dosage?

Summary
Biologics hold a lot of promise
Improved outcomes Reduced side effects
But presently can be cost prohibitive

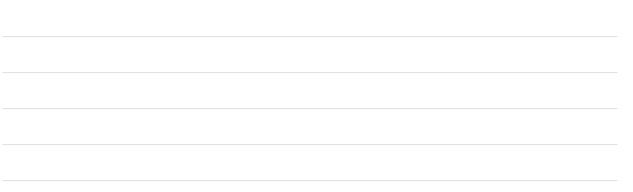


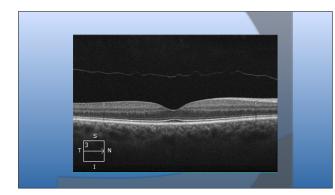


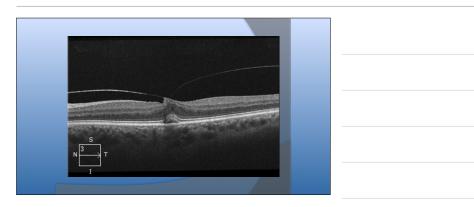


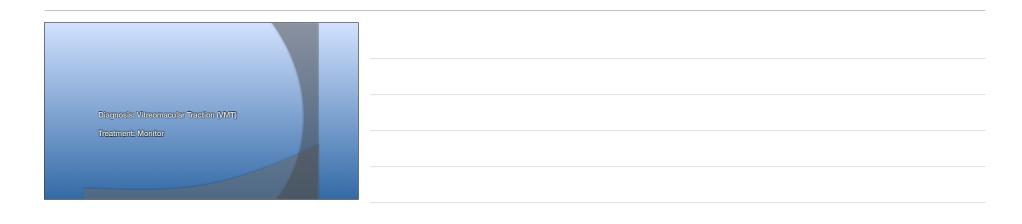


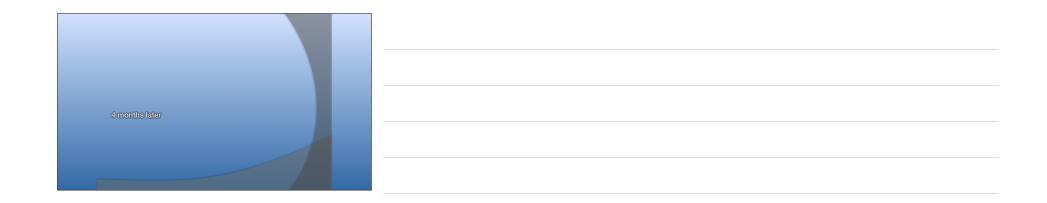


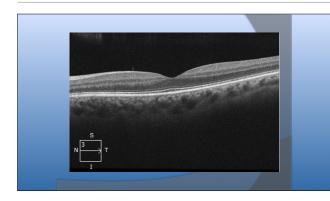






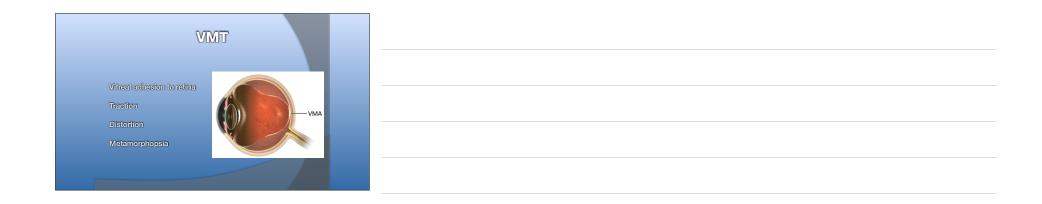








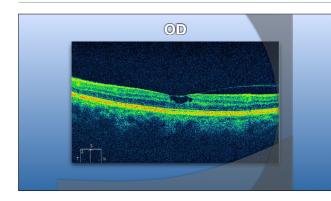




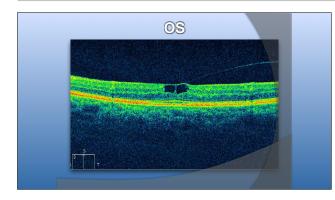




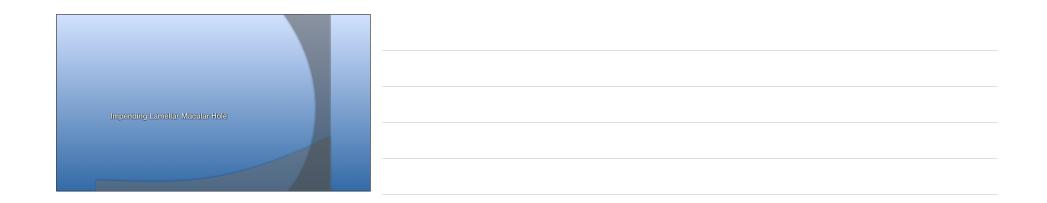


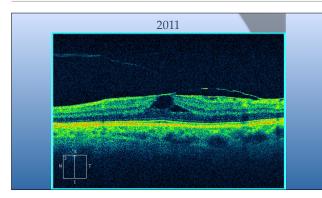




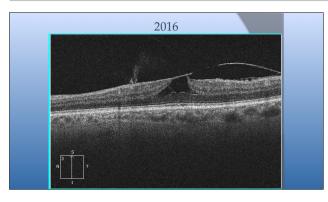


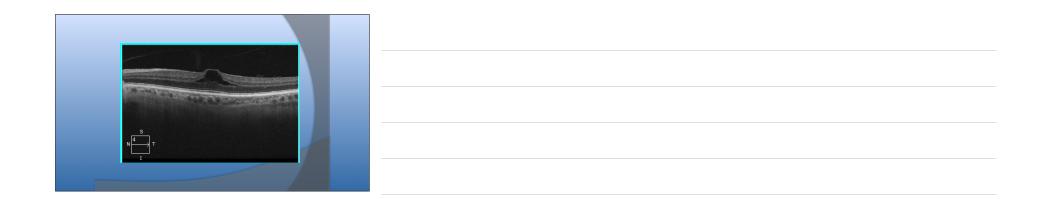








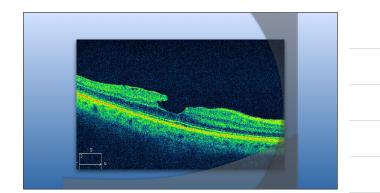
















Acuity typically better	
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Difficult to treat

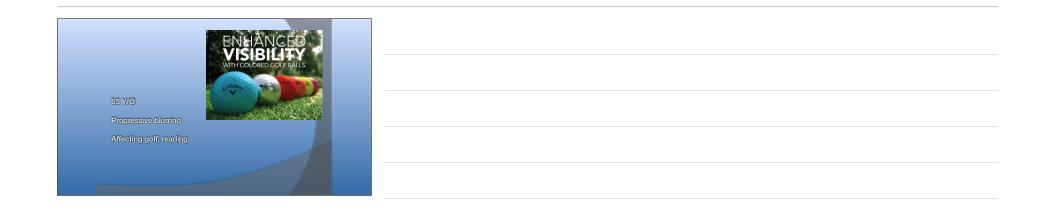
Vitrectomy and Peel

May be more selective on when to treat

Before OCT

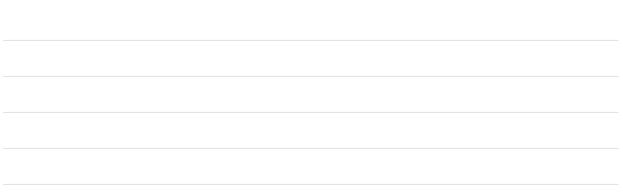
Diagnosis less certain

Less known about Tx/prognosis

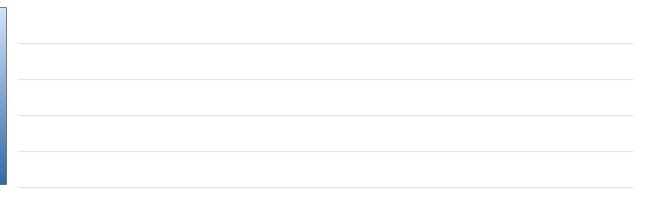


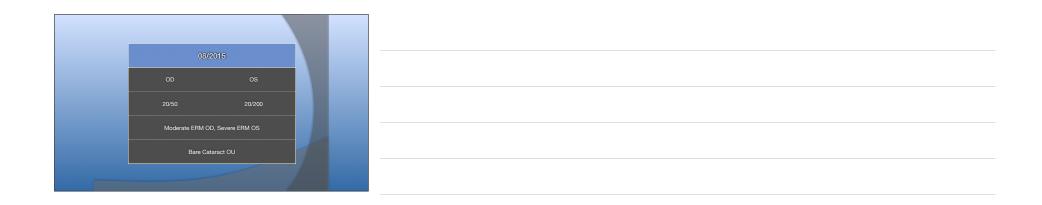
7/2014		
OD	os	
20/20	20/25	
Normal Systemic Health		
Bare Cataract OU		



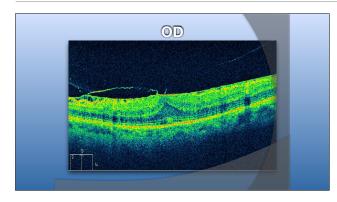


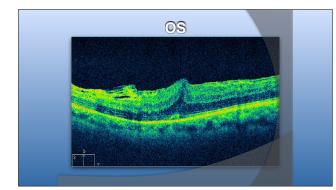
02/2		
OD	os	
20/30	20/50	
ERM		
Bare Cat		







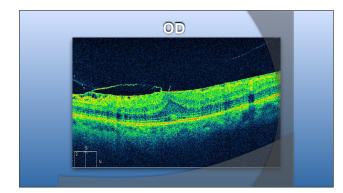




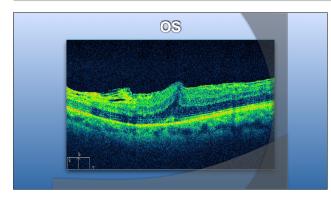


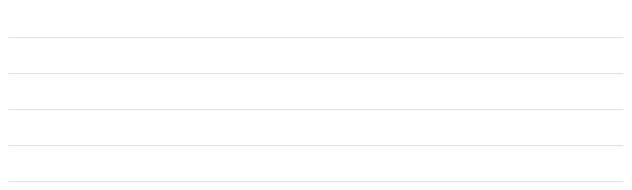


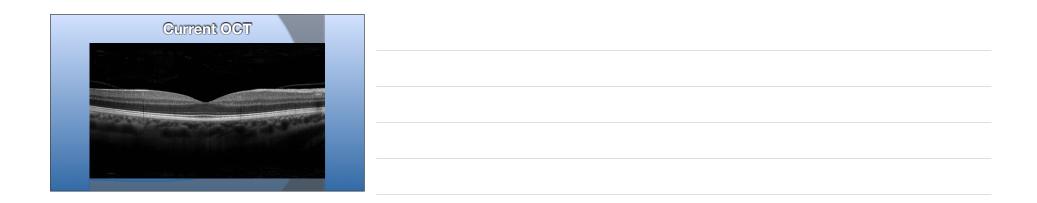














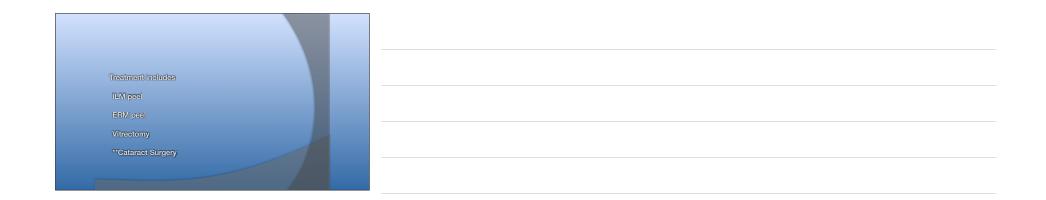




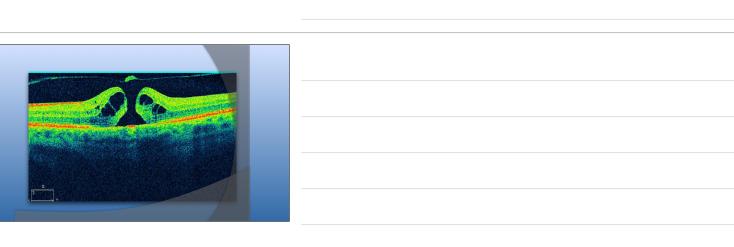
From vitreal detachment	
PRP or other laser	
Cellular growth/traction	
20% of patients over 75	

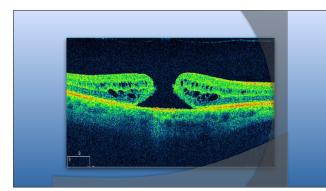
Blur	
Distortion	
Metamorphosis	
and an include the second s	
Represent country (20/20 - 20/400)	
Range of acuity (20/20 - 20/400)	
How many progress?	

Typically asymptomatic	
If acuity reduced refer to retinal specialist	
What acuity should be treated??	



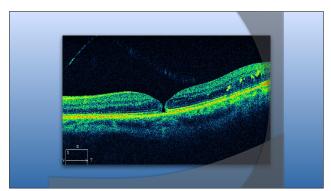


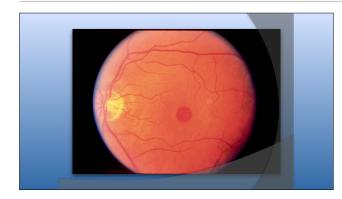


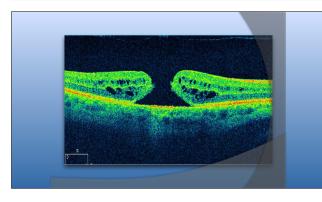




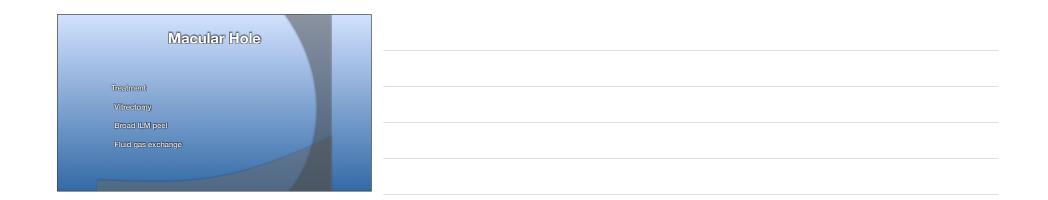




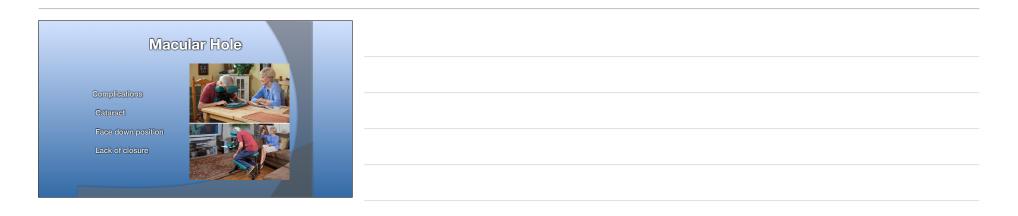










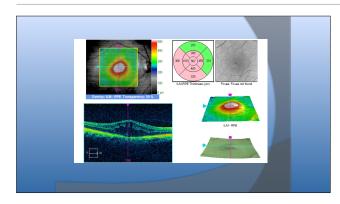


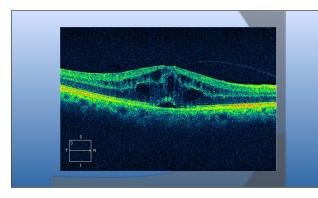
Macular Hole
Modern Treatment
3 days mostly face down Then sleep on side
Then sleep on side

Macular Hole
Marlow Frankran
Modern Treatment If concurrent cataract - treat first
Risk of CME disrupting outcome



Anterior Segment Trace Cell OD Mild K Edema (Fuch's Dystrophy)

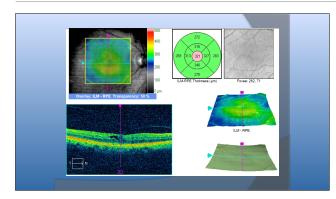


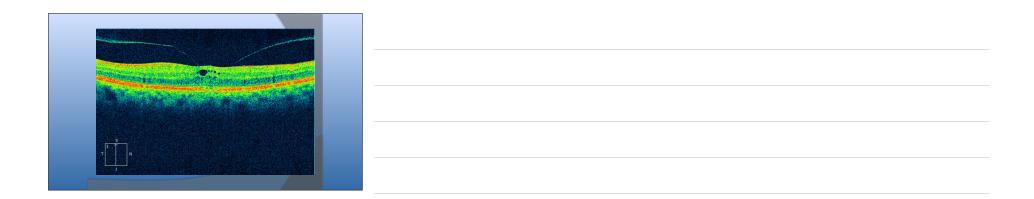












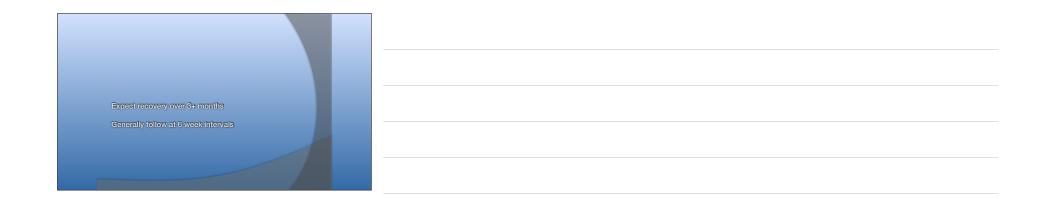
Irvine Gass CME
Need to dafine GME
Vein occlusion
ERM
RP
Diabetes
Post-op

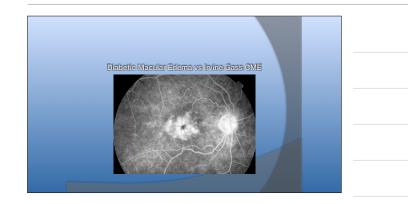


Risk Factors	
Posterior capsule rupture	
IOL dislocation	
Iris fixed IOL	
ACIOL	



Treatment Topical NSAID/Steroid QID*
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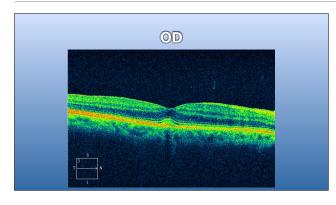


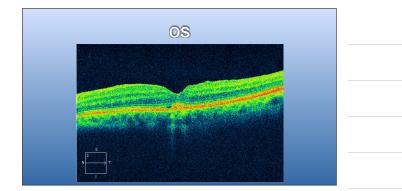




# IOP 8/10 on Latanoprost

Posterior segment Glaucoma Normal vasculature One small drusen OU











https://images.app.goo.gl/57bripcyYLyDTdsT8

Adult Vitelliform
Similar to Best's
Later in life
Smaller lesions
Less complications

#### Adult Vitelliform

Lipofuscin accumulation

Between RPE and Photoreceptor

5-15% develop CNVM

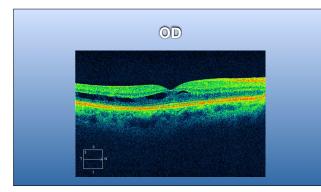
Typically mild and asymptomatic

Optometry (2006) 77, 156-166

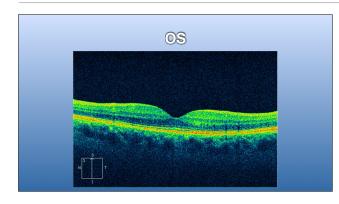




Posterior segment			
Normal ONH			
Normal vasculature			
Macula abnormal			
	· · · · · · · · · · · · · · · · · · ·		

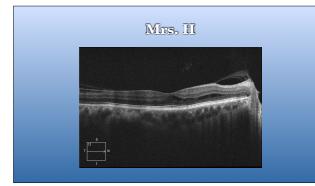






# Juvenile Retinoschisis

Incidental finding Patient asymptomatic Treated with CAI (Trusopt) - no benef





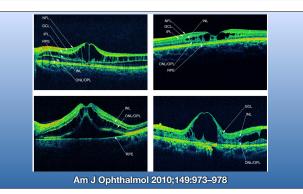
## Juvenile Retinoschisis

Genetic malformation (x-linked) - typically male Difficult to diagnose funduscopically Amblyopia masquerader

#### SNIFR

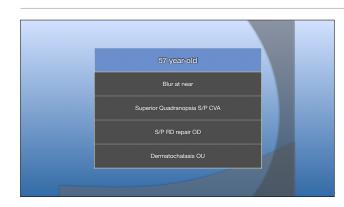
Stellate Nonhereditary Idiopathic Foveomacular Retinoschisis

Similar to Juvenile Retinoschisis - no gender predilection

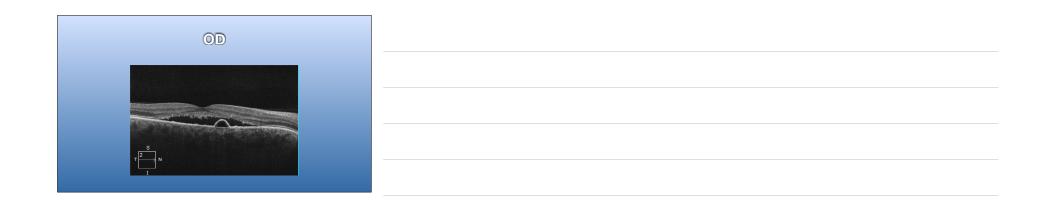


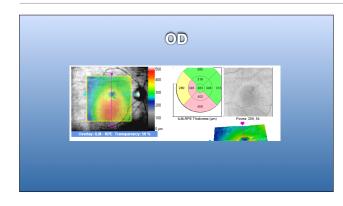


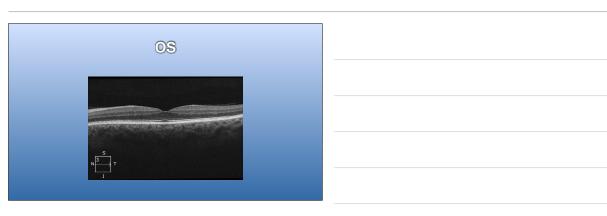
Juvenile Retinoschisis	
Variable layers affected Variable appearance	
Variable appearance	



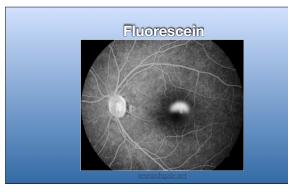












### **Central Serous**

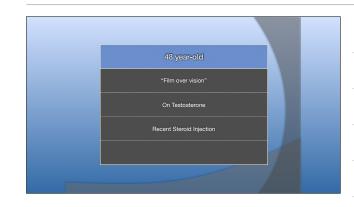
Serous elevation of retina

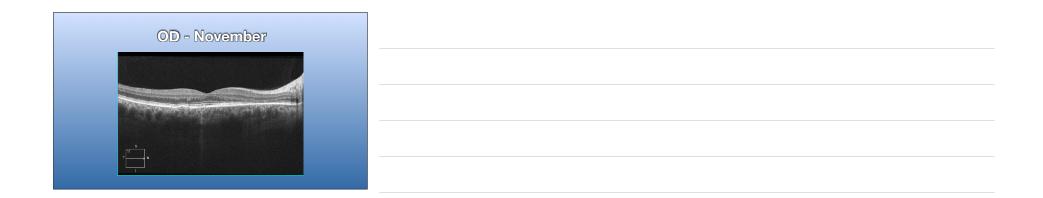
Males (20-50 years old) Stress/Cortisol Steroid

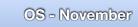


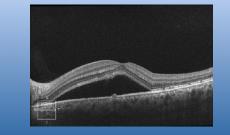
Self-limiting - 3-4 months	
Laser	
PDT	

30% can become chronic	
Mineralocorticoid Receptor Antagonists	
Spironolactone, Eplerenone	

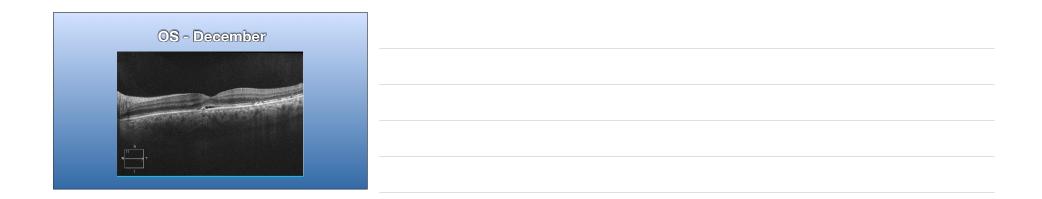








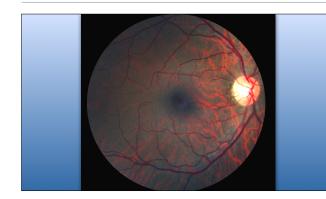


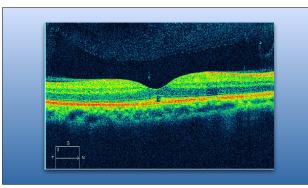


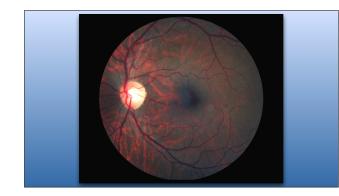


Alerted POP to side effects of Testosterone, Steroids Self-resolved	







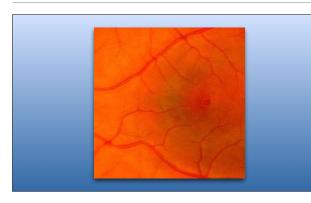


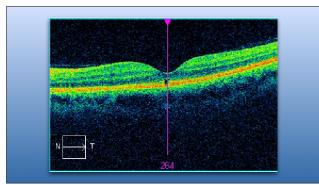


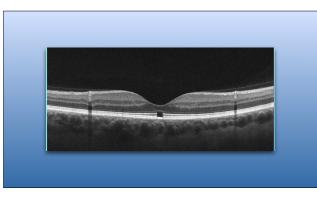
Diagnosis?	

Solar Maculopathy
Thermal Burn
Psych Diagnosis Eclipse
Drugs

_		
	Adjustment Disorder	
	Alcohol Abuse	· · · · · · · · · · · · · · · · · · ·
	Inadequate Housing	
	Inadequate Housing Depressive Disorder Tobacco Dependence	
	The Development	
	Tobacco Dependence	







# Solar Maculopathy

No ocular treatment - non progressive Mental health referral?

Plaquenil
2016 Recommendations
Continues testing recommendations from 2011

# Plaquenil

Low risk drug if correct dosage

If proper dosage, should never see "bull's eye maculopathy"

# Plaquenil

Consider modifying risk by modifying dosage

Only come in 200 mg tablets

Patient may want to take fewer than 14 tablets/week



Plaquenil
Study of 500 patients started on Plaquenil 50% were on too high dose for ocular safety
Ophthalmology 2017;124:604- 608

Plaquenil
Height to take med safely based on IDEAL weight
Men - 5'5" Women - 5'7"
Women - 5 /

Plaquenil
For every 2" below ideal, subtract 1 tablet per week

## Plaquenil

Safe weight for 400 mg/day - based on REAL weight 180 lbs For every 13 lbs less than this subtract 1 tablet/week

#### Recommendations

Baseline Examination

If low risk, then test after 5 years

If high risk, test yearly

#### Recommendations

High Risk

Kidney Disease

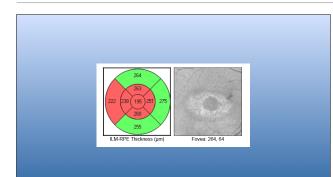
Liver Disease

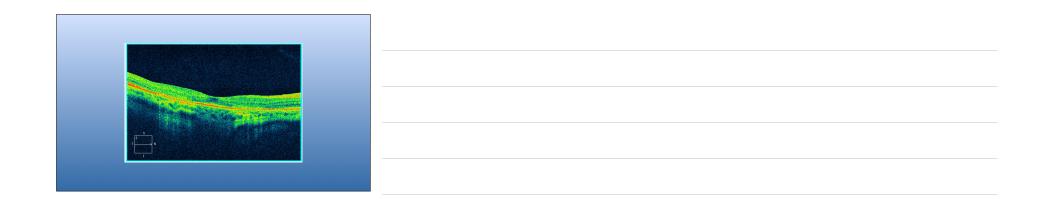
Cumulative Dose

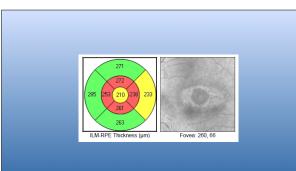
Tamoxifen Use

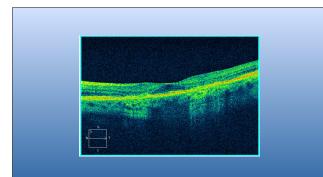
Case 1
60 Year Old White Female
Blur at distance
Photophobia Dry eye
Dry eye

Case 1
Plaquenīl x 20 years 400 mg/day
Lupus
5'4"
105 lbs - safe dosage is 200 mg









Plaquenil toxicity

Should have been on lower do

Need to discontinue medicine

86 Year Old White Male

Blur at distance

Ocular History: Early AMD

Medical History Psoriatic Arthritis HTN CAD Hyperlipidemia

Mediceitions	
Plaquenil	
Fluocinonide	
Coreg	
Coreg ASA 325 mg	

57° fal 130 lbs - would need 10 tablets/week

Patient at VA since 1999 Every exam: RPE changes OS>OD

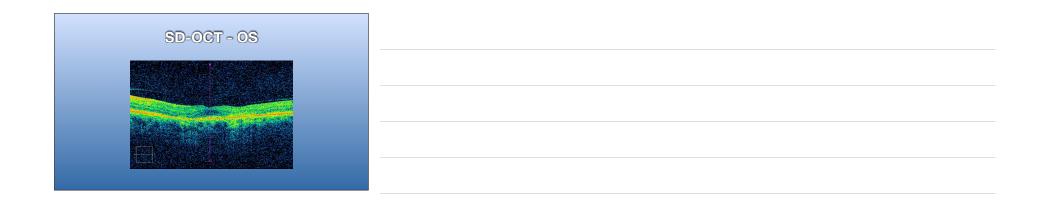
Called AMD

20/20- OD and OS

September 2010 Ring shaped atrophy OS Fundus Photos/OCT









Management
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Discontinue Plaquenil
See Rheumatologist Patient refused initially
See Aneumatologist Patient refused mitany

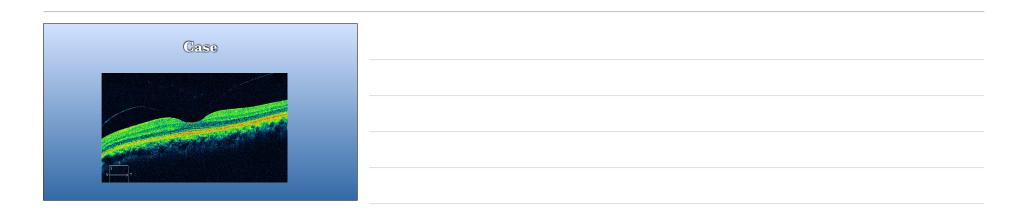
Management
Patient eventually discontinued meds
Asymptomatic Good outcome?
Good outcome?

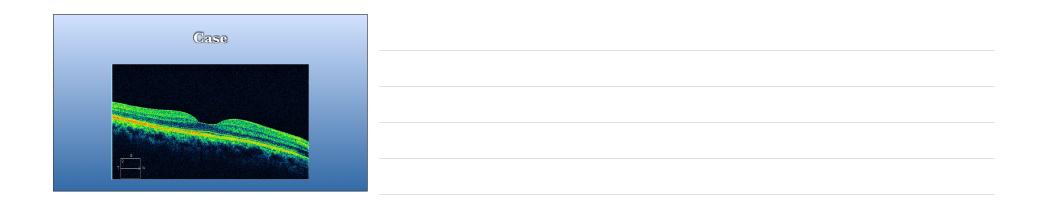
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Case 3
71 Year Old White Male
Evaluation for Plaquenil Maculopathy
No vision complaints



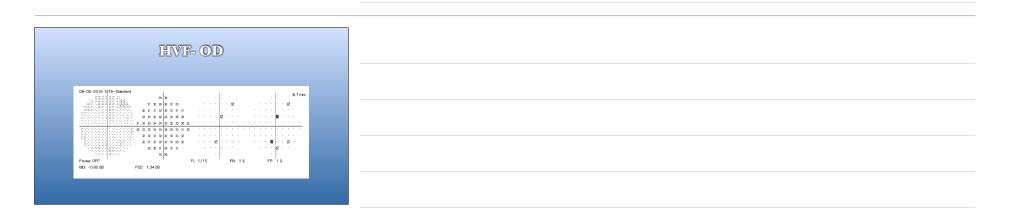
Case 3	
5'11" tall 155 lbs - should be on 12 tablets/week	







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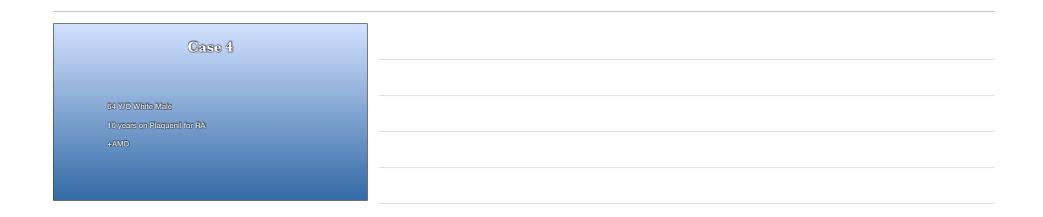
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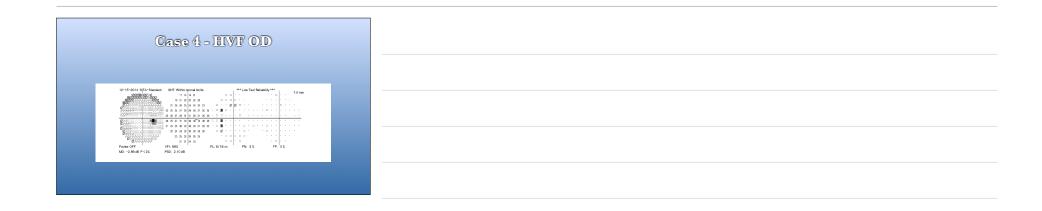
Case 3
Normal macula Normal OCT
Normal OCT
?HVF

Case 3				
Do we continue medication?				
Do we discontinue medication?				

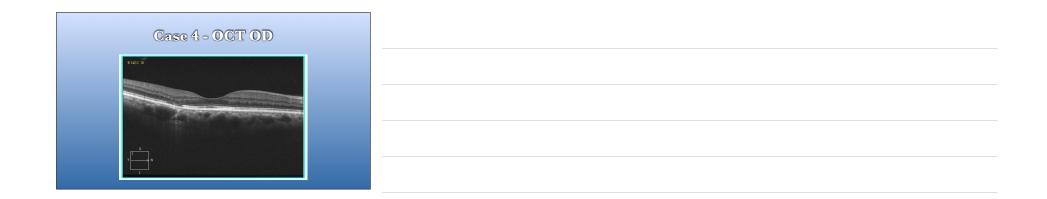
Case 3
Plan
Continue medication
RTC 6 months - retest



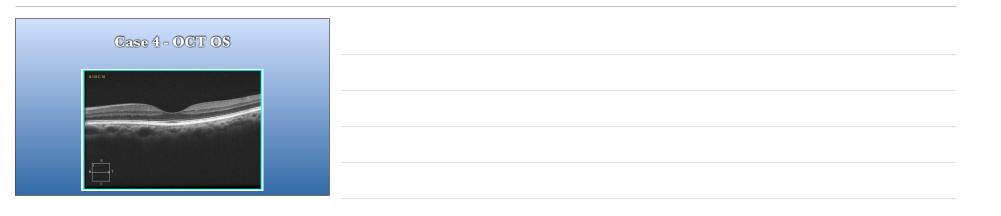
Case 4	
5111" 182 lbs	
102.105	





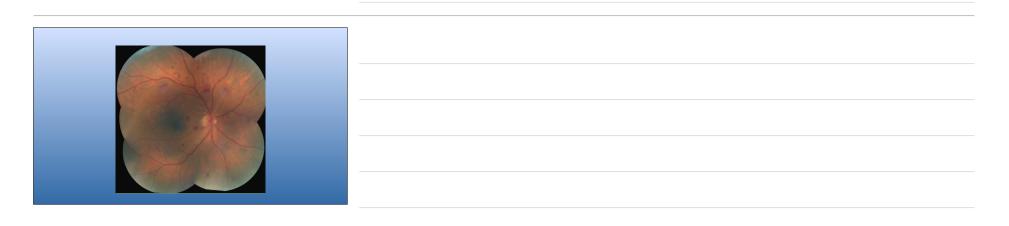


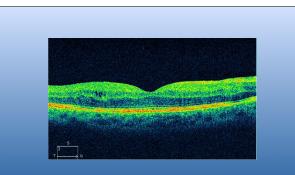




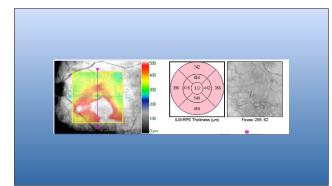
Case 4
Informed Rheumatologist
They discontinued Plaquenil Was this the right call?
Could we have done better?



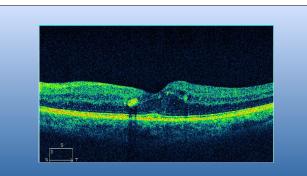












Diagnosis NPDR OU No GSME OD CSME OS

## CSME

Retinal thickening 1 disc area - any part within 1 disc diameter of center of fovea

Hard exudates within 1/3 disc diameter of center of fovea with adjacent thickening

Retinal thickening within 1/3 disc diameter of center of fovea

Treatment Griteria - Anti-VEGF	
Central retinal thickening ≤20/30	

Treatment	
Prompt referral to retinal specialist	
Under treatment with IVI Avastin	

Prognosis?	
Floghosis	
	4
	4
	4

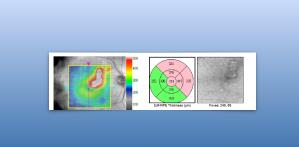
How many injections are necessary?	

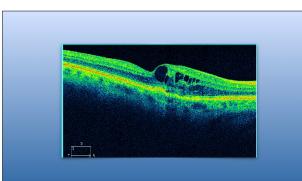










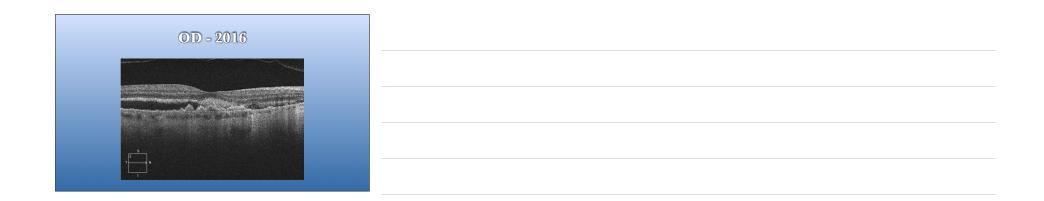


BAYO OD	
BRVO OD 20/25 acuity Refer?	
Defer)	
neren i	

With the first first of with large @ 2 manufact		
Historically treated with laser @ 3 months Now treated with Anti-VEGF injections		
Now treated with Anti-VECF injections		

Prognosis? How many injections needed?







AMD OU
Serous elevation
Refer to retina
Intravitreal Anti-VEGF



08 - 2018

Vision	
20/40	
20/25 Can we expect better?	
Can we expect better?	

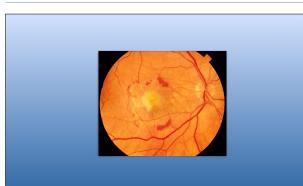
## Macular Degeneration

Four potential findings Sensory detachment

Pigment epithelium detachment (PED)

Sub-Retinal hemorrhage

Sub-RPE hemorrhage



## Macular Degeneration

Historically treated with

Nothing

Laser

Visudyne

Studies
90% maintain acuity with treatment Only 50% untreated maintain
Only 50% untreated maintain





Treat and Extend
Treat every month for 3 months If stable, extend out

Treat and Extend
90% had stability at 2 years 45% had 20/40 acuity
Ophthalmology 2015;122:1212-1219

	Treat and Extend	
	Fewer injections (13 versus 17) ~ over 2 years Fewer Visits	
	Less SS	
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