

Focusing on Quality 1st...

- Patient-dependent factors

 Pupil size, TF, central media opacity
- 2. Operator-dependent factors
- 3. Device-dependent factors

Review Article: Factors Affecting Clarus-IID OCT Optic Disc Scan Quality: A Review with Case Examples Jobs 5. Neths, timost 12065, Vol. C. Steins, State Chin, and Emmers You

J Ophth 2013





• OCT confused about location of RPE

• Numeric values of macular thickness are erroneously low





- 34 AA Male
- Smoker
- Schizophrenic • Fluphenazine
 - Risperidone
- 20/30 OD, OS, down from 20/20
- Metamorphopsia
- Color vision-normal
- 10-2: clear



Hmmm...??

- Macular cyst?
- Vitreomacular traction?
- Inherited disorder?
- Toxicity?
- What else...???



Ganglion Cell Analysis



What Else Can Cause Central Inner Retinal Thinning?



- Glaucoma
 - Demyelinating disease
- ONH drusen
- Toxicity
- Retrograde degeneration of GCL from optic tract lesion
- Lebers Hereditary Optic Atrophy
- ION
- Overlying choroidal ischemia

Photoreceptor Integrity Line (PIL) Junction PR I/O segment



Clinical Significance of PIL

- Central blip=elongation on cone outer segment
- Rod monochromat missing central PIL
- RP PIL present centrally and then fades away
- Lamen-Moon Bardet-Biedle (LMBB) Syndrome – genetic disease with polydactyly and missing PIL (retinal normal looking), no VEP activity



- Toxicities (e.g. Tamoxifen)
- VMT
- Juxtafoveal Macular Telangiectasia
- Welder's retinopathy
- Stargardt's
- Acute Retinal Pigment Epitheliitis
- Or....

Patients admits to "staring at the sun"....

- Solar maculopathy?
- Damage to outer segment of PR layer, RPE, \uparrow choroidal thickness
- Often subtle but permanent vision loss OCT findings:
 - Abrupt interruption to OS/IS junction
 (PIL)
 - Slight separation between RPE and PR



Solar Retinopathy -FAF

 Pinpoint hypo at fovea with subtle hyper surrounding it





Differentials?

Adult Vitelliform Pattern Dystrophy VMT CSR PED





Adult Vitelliform Dystrophy

- Typically bilateral pattern dystrophy

 1/3-1/2 DD size yellowish elevation
 Above RPE, below retina
- 4th-6th decade
- Acuity 20/50 or better
- Complications:
 - RPE pigmentation
 - PR disruption
 - CNVM 11%





Pattern Dystrophy as a Differential for AMD..

Group 1: adult-onset foveomacular vitelliform dystrophy. Group 2: butterfly-shaped pigment dystrophy. Group 3: reticular dystrophy of the RPE. Group 4: multifocal pattern dystrophy simulating fundus flavimaculatus.

Group 5: fundus pulverulentus.

25://www.reviewofoptometry.com/article/pattern-recognition-how-to



Fundus Autofluorescence – clinical significance Lipofuscin – hyperfluoresces with 500-800nm Pro • Byproduct of phagocytosis of PR OS Noninvasive, easy Expensive addition to retinal camera, and does not • Accumulates in the RPE (and eventually in Attached to camera (some) allow extra billing Early detection and the potential for predicting progression of many retinal diseases outer retinal layers) Quantification of intensities is subjective Varying reports/data of clinical use naturally with age • excess accumulation is a sign of pathology: • View of many different retinal disease processes from a activity in RPE cell death process metabolic perspective Creates • Lipofuscin exposed to light 500 to 800 nm ightarrow• Outer segment PRs shed \rightarrow phagocytosis via RPE autofluorescence lipofuscin molecule exited \rightarrow when exposed to →Lipofuscin (predominant ocular *fluorophore*) spontaneously releases a photon of light wavelengths of in Vitr (2016) 2:12 is formed as a result with a longer wavelength \rightarrow light 500-800nm visible in photograph

FAF Image Interpretation

- Hyper AF= RPE is metabolically sick LF accumulation
- Hypo AF=
 - RPE is dead or not present (LF gone)
 - Signal blockage
- FAF intensity variables
 - agemedia opacity

 - pupil size
 - FAF device used



41 AA Male; Never had an eye exam



Stargardt Disease

- Defective outer segment degradation, lipofuscion accumulation, and center degeneration of RPE and PRs
- OCT: disruption/loss IS/OS of photoreceptor layer
- FAF: Useful in Dx
 - Early atrophy/flecks not visible otherwise
 - correlates well with vision function







LUXTERNA[™] (voretigene neparvovec-rzyl)

- FDA approved December 2017 for children and adults with mutations of both copies of RPE65 genes
 - Lebers Congenital Amaurosis, RP
- One time gene therapy inserts a normal copy of the gene into retinal cells
 - Subretinal injection
 - One eye at a time, ≥6 days apart
- Improved in visual function based on navigation of an obstacle course at low light levels

Be proactive in...

- Genetic testing / counseling
- Low vision referrals
- Defining legal blindness / advocating disability



Dry AMD with drusenoid PED

73 Caucasian female 6-year history of dry AMD OD>OS















On the other hand...

82 WF



Anglography Analysis : Anglography 3x3 mm

OD O O OS







Pigment Epithelial Detachment

- MC cause is AMD (drusenoid, hemorrhagic) • Other etiologies/associations:
 - CSR
 Pattern dystrophies
 - Idiopathic
- 75%---self-limiting

 Better prognosis if <55
 Worse if turbid fluid or associated hemorrhage
- Visual loss can occur
 RPE atrophy
 Up to 30% lead to CNVM
 10% of patients over 55 develop RPE tear
- Overlying serous sensory RD and CSR



Idiopathic PED: management

Monitor carefully

- IVFA
- SD-OCT

• Avastin?

- persistent PED
- PED with CNVM
- Height of PED predictor for risk of tear after Avastin injection Chan et al. Retina, Feb 2010
- However....Majority of larger PEDs from CNVM in AMD do not respond to iniections Freeman, et al. *Retina* Oct 2011. Parodi, et al. *Am J Opth*, Jan 2013.

Non-exudative Neovascular AMD

- Quiescent CNV
- Not visible on FA no leakage
- Treat or not to treat ?



Paleiwala et al. Retina 2015.

Geographic AMD

- 11% have quiescent CNV
 - 25% will become exudative
 - Irregular PEDs
 - (-) leak
- Typical anterior displacement of choroidal vessels - don't confuse with CNV
 - Medium sized vessels
 - Visible in CC slab 100% of GA

Capuano V. Am J Ophth 2017. Nesper P. Retina 2019.



FAF - Geographic AMD

- More dramatic visualization
- Hypo-AF: no PRs and no lipofuscin in RPE cells
- Hyper-AF at borders RPE is overworked
 - · Important risk factor for progression
 - · Indicates an active expansion



New Treatments for Geographic Atrophy (GA)in 2023

 Targets the complement system at different places Goal: prevention of





Syfovre (pegcetacoplan)

- FDA approval in February 2023 15 mg q25 to 60 days
- Targets C3
- OAKS and DERBY trials rate of GA progression reduced by up to 22% with monthly injection
- adverse reactions (>5%)
- ocular discomfort
 Wet AMD
 Floaters
 conjunctival hemorrhage





Change in appearance after treatment

- Type 1
 - Mature, longstanding
 - · Resistant to tx
- Type 2
 - Decreased size and VD of CNV after anti-VEGF
 - Trunk/feeder vessels unchanged
 - No reduction in lesion area but reduction in blood flow within CNVM
- Suggestion is to repeat OCT-A x 2 wks to determine need for re-tx



RETINA, THE JOURNAL OF RETINAL AND VITREOUS DISEASES 2017

Neovascular AMD

- Earlier, non-invasive detection
- Can detect actual vasculature, w/o relying on leakage, exudation
- Sub-type, location, size
- Monitor response to Tx/need for re-tx
- Inactive membranes
- Choroidal ischemia



Vs. Polypoidal choroidal vasculopathy

- Subset of AMD, and/or Pachychoroidal diseases
- · Often missed or confused with classic AMD
- Mostly pigmented races
- Hemorrhagic PED and serosanguinous maculopathy vs. macular drusen
- ICG gold standard for dx • EDI useful : choroidal thickening • OCT-A
- PDT often used instead of/along with anti-
- VFGF Eleftherios I, et al. Eye, July 2018.



Choroidal Thickness

- SS-OCT or SD-OCT with EDI
- vertical distance from RPE and inner surface of the sclera
- Subfoveally + parafoveally





OCT with Enhanced Depth Imaging (EDI)

- Pre-selected setting in the acquisition of OCT
- Higher resolution in deep / outer retinal findings
- Suggested useful for
 - Choroidal thickness
 - CSR
 - AMD
 - Choroidal nevi/melanoma
 - Optic disc drusen • Scleral contour

 - Normal tension glaucoma •

44 Hispanic FM

- Blurry vision R eye x 1 monťh
- 20/200, PHNI
- PMH:
 - HTN GERD
 - Knee pain: Recent history of oral prednisone and fluticasone injection for knee pain











Plan:

D/C steroids

Topical NSAID QID

Oral spironolactone 50mg QD



Central Serous Retinopathy – Revisited

- Now considered one of the pacychoroid diseases
- Risk Factors, Liu et al, Retina 2016
 - Hypertension Helicobacter pylori infection
 - steroid usage sleeping disturbance .
 - autoimmune disease
 - psychopharmacologic medications Type A behavior
- Generally a self-limiting disease, however...
 RPE damage → diffuse retinal pigment epitheliopathy (DRPE)
 CNVM



CSR (CSC) and Choroidal Thickness

- pachychoroid diseases include
- CSC
- pachychoroid pigment epitheliopathy
- · pachychoroid neovasculopathy
- polypoidal choroidal vasculopathy
- · Looking at subfoveal choroidal thickness from EDI
- increased in CSC eyes and fellow eyes
 - · eyes with hypermeability also had increased choroidal thickness

Chen G[,] PLoS One. 2017 Zhonghua, 2012

Central Serous Retinopathy

- Initially hypo FAF due to blockage / absorption by subretinal fluid
- As progresses/current activity:
- Fluid present affects RPE function
- Granular hyper FAF
- Longstanding: hypo FAF due to atrophy
 - · May identify previous events



Phosphodiesterase-5 Inhibitors retinopathy?

- Indications: Erectile dysfunction, benign prostate hyperplasia, pulmonary hypertension
- Explanation:
 - Choroid is vascularly similar to corpus cavernosum
 - 10% activity against PD6 (retinal phototransduction)
- Suggested Associations:
 - CSR
 - Choroidal thickening

Arora S, Surakiatchanukul T, Arora T, Cagini C, Lupidi M, Chhablani J. Sildenafil in ophthalmology: An update. Surv Ophthalmol. 2022 Mar-Apr;67(2):463-487. doi: 10.1016/j.survophthal.2021.06.004. Epub 2021 Jun 25. PMID: 34175342.

Elizalde J, Laiseca A, García de Oteyza G.Retin Cases Brief Rep. 2022 Nov 1;16(6):779-782. doi: 10.1097/ICB.000000000001082.PMID: 33165301



Diabetic Retinopathy – Update on Treatment and Protocols





Procotol V: Other considerations

- Central thickness 400 : marker for more severe DME
- Applicable to all eyes with CI-DME with good VA?
 - patients had well-controlled diabetes with a mean A1c of 7.6, and
 - enrolled eyes had earlier stages of DR, mild DME







Isolating Blood Flow Data at a Specific Depth













FAF: Plaquenil Toxicity

- May contribute to earlier detection

 Before RPE degeneration develops
- berere in 2 degener
- Early change:
 - hyper AF
 - 2-6 degree ring around fovea
 Early photoreceptor damage subtle
- Later change:
- hypo AF
 - Coalesces into dark absence of FAFcells are dead



OCT – what to look for early

- Scans
 - High resolution
 - Raster or radial
 - Enhanced depth imaging (EDI)
 - Macular cube quantifiable

Outer retinal thinning

Disruption to PIL



Toxicity is Dose-Dependent Other Risk Factors • Proper dose <1% with toxicity</pre> • Kidney disease • 6.5mg/kg based on IDEAL weight • (consider that it is commonly used in conditions that themselves affect the • Typically 200mg BID – ok for normal size patient kidneys such as SLE) • Not ok if short/small • Existing retinal/macular disease • Not ok if obese • Concurrent use of tamoxifen • Not ok if kidney disease • Mostly clinically significant cases of toxicity are iatrogenic • Overdosing (>10%) • Lack of screening Inadequate screening/missed findings Melles RB, Marmor MF.JAMA Ophthalmol. 2014 Dec;132(12):1453-60. doi: • Damage can progress up to 3 years after discontinuation.... Browning DJ Hydroychloroquine and Chloroquine Retinopathy 2015 10.1001/jamaophthalmol.2014.3459.PMID: 25275721

But what if Plaquenil is life-saving?

• A patient with systemic lupus erythematosus on hydroxychloroquine (HCQ) was taken off the drug due to toxic maculopathy. Four years later, HCQ was resumed because of a compelling need to do so to control her disease activity. At seven years since the resumption of HCQ, there has been only mild, slow progression of the retinopathy.

> Sunness JS, Sunness RH, Hellmann DB. Hydroxychloroquine can be resumed with close monitoring after retinopathy has developed, without major visual loss: Case Report. Retin Cases Brief Rep. 2023 Dec 12. doi: 10.1097/CB.0000000000000000000536. Epub ahead of print. PMID: 38150578.

Differentiating Suspicious Choroidal Nevi

52 year old Caucasian male



How Many Choroidal Nevi Did You See This Week?

- prevalence in the posterior pole
 - 4.1% in whites
 0.7% in blacks
 - 1.2% in Hispanics
 - 0.04% in Chinese
- 3 features changed with age
 - 1. thickness
 - 2. multiplicity
 - 3. drusen
- Annual transformation rate to melanoma: estimated at 1 in 8,845 (0.0113%)

Shields C et al. Retina 2014.



\geq 3 = 50% chance of transformation in 5 years *These 3 = 69% risk!

High Risk Features for Small Nevus Conversion to Melanoma			
To*	Thickness	Thickness of >2mm (as measured by ultrasound) - bad	
Find	Fluid	Overlying subretinal fluid - bad	
Small*	Symptoms	Symptoms of flashes/floaters/blurry vision - bad	
Ocular	Orange pigment	Presence of lipofuscion – bad	
Melanomas*	(Disc) Margin	Nevus ≤3mm from disc margin - bad	
Use	Ultrasound	Hollowness noted on ultrasound - bad	
Helpful	Hollowness		- Le
Hints	Halo	Presence of halo - good	
Daily	Drusen	Presence of overlying drusen – good	Shields C et al. Retina 2014. Shields C, et al. JAMA 2012.





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FAF

- Orange pigment = lipofusion
 Naturally fluorescent with bright light
- Indistinct borders
- Fluid









