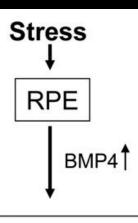
Wet Age-related Macular Degeneration Neovascular AMD

Mallika Goyal, MD Apollo Hospitals, Hyderabad

Pathogenesis Multifactorial



Sustained BMP4

Early AMD RPE Senescence

Senescence Activated Secretory Pathway (SASP)

Inflammatory Mediators (eg. TNF-α)

↓

BMP4↓ VEGF↑

Geographic Atrophy

RPE Senescence RPE Apoptosis Neovascular AMD

IL-8 1

RPE Activation
Choroidal Angiogenesis

Investigations

- Fundus Fluorescein Angiography FFA
- Optical Coherence Tomography OCT
- ICG

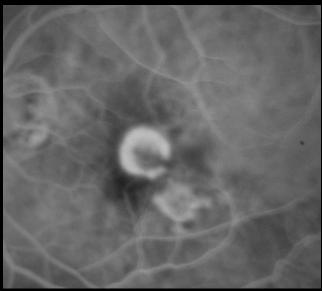
FFA in AMD Indications

- At presentation only if doubt about diagnosis
- When response inadequate
- Pree-PDT for lesion diameter

Classic CNVM

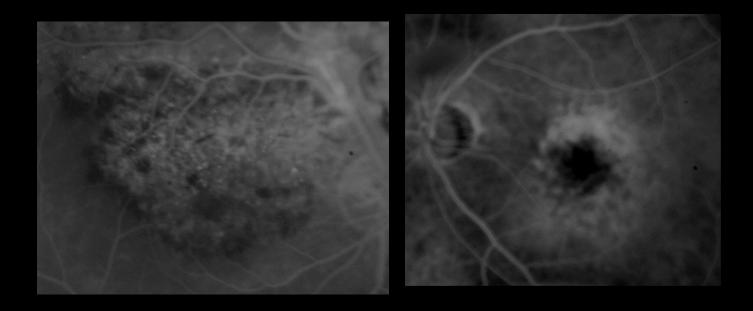
- Sub-neuroretinal / -sensory retinal
- Bright leak
- Borders distinct

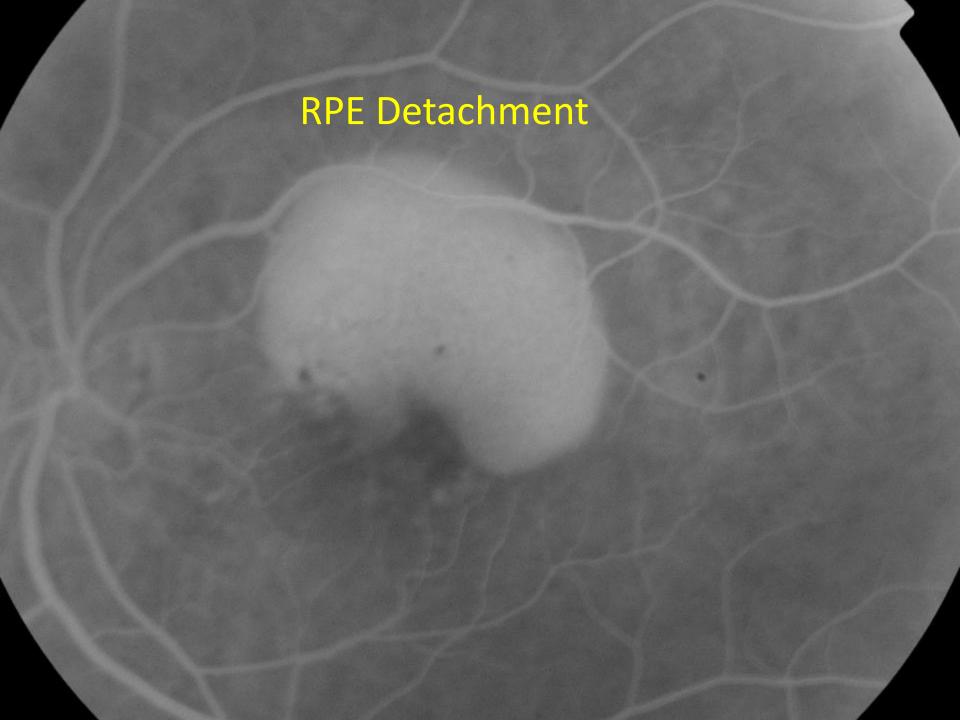


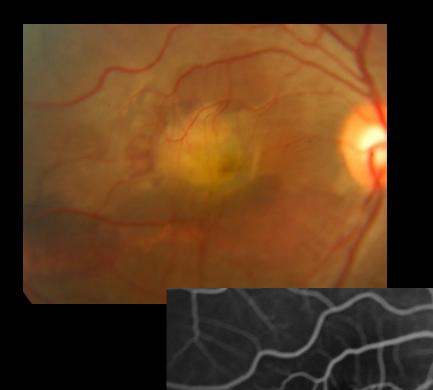


Occult CNVM

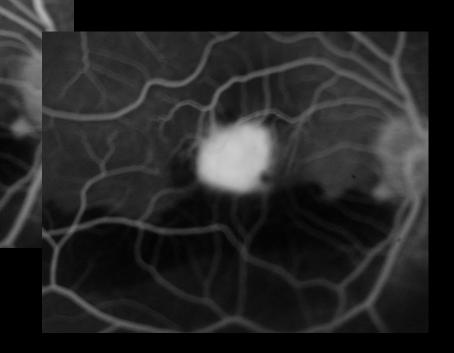
- Sub-RPE lesion
- Borders ill-defined
- Punctate hyperfluorescence
- Diffuse ooze membrane





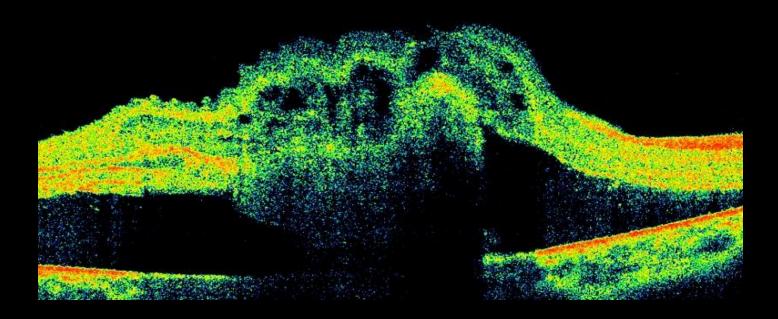


Retinal Angiomatous Proliferans (RAP)



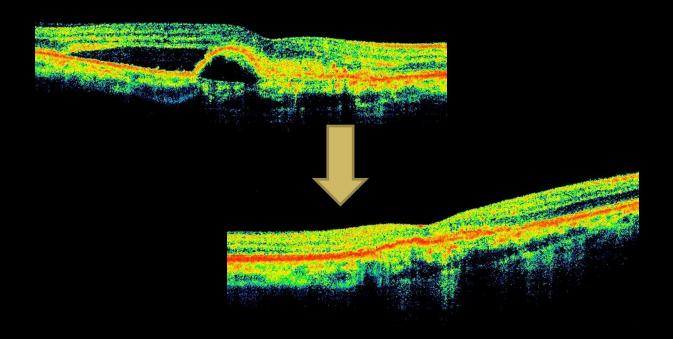
OCT RAP lesion

- Massive fluid in nuclear layers
- Hyper-reflective fluid
- Shadowing of outer layers, RPE



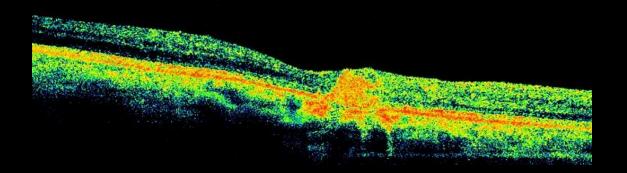
OCT

- High resolution tomogram 5 um
- Picks up minimal amounts of fluid
- Differentiates scar from active CNVM
- Allows quick comparison on serial follow-up



When to stop treatment

No fluid: no activity, can observe

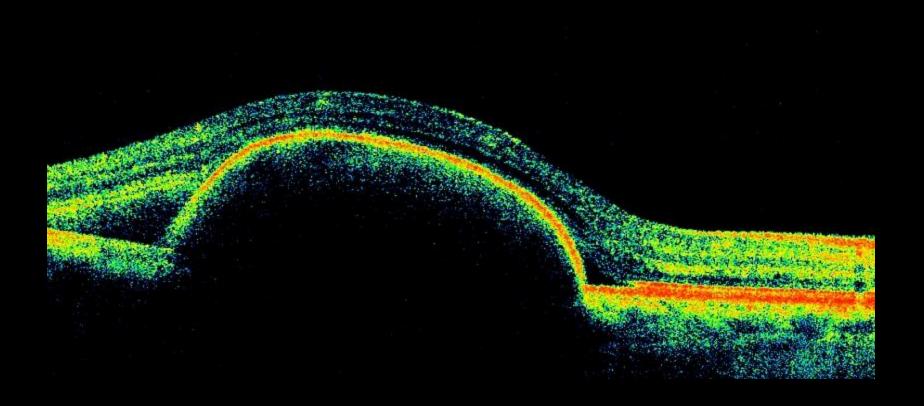


Haemorrhagic lesion wet AMD

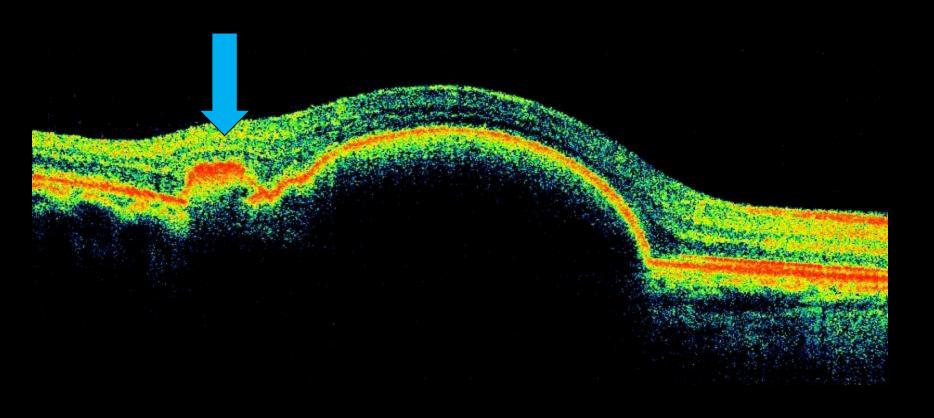
Blocked choroidal fluorescence

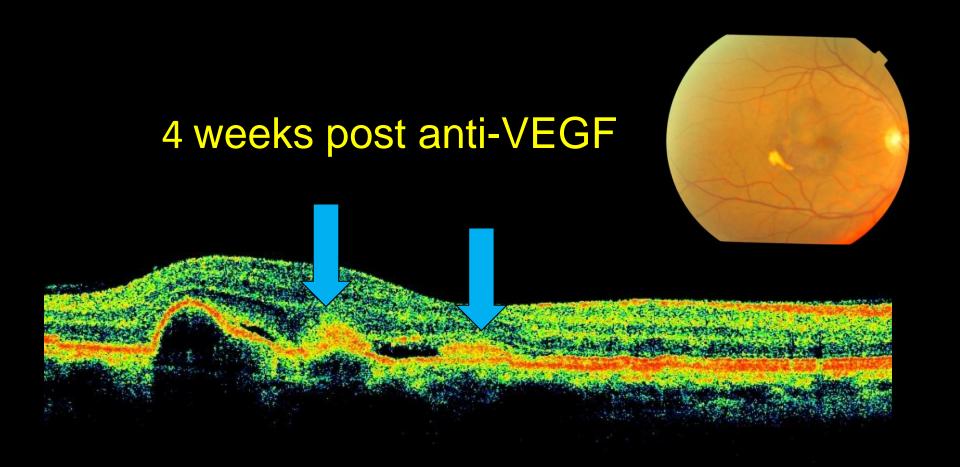
Blocked choroidal fluorescence

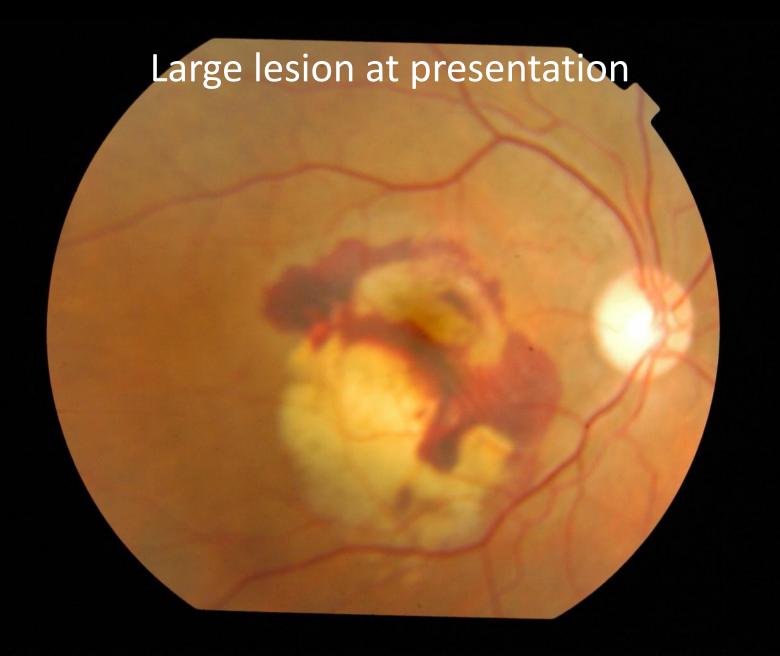
- Sensory retinal detachment
- All outer retinal layers compressed over lesion



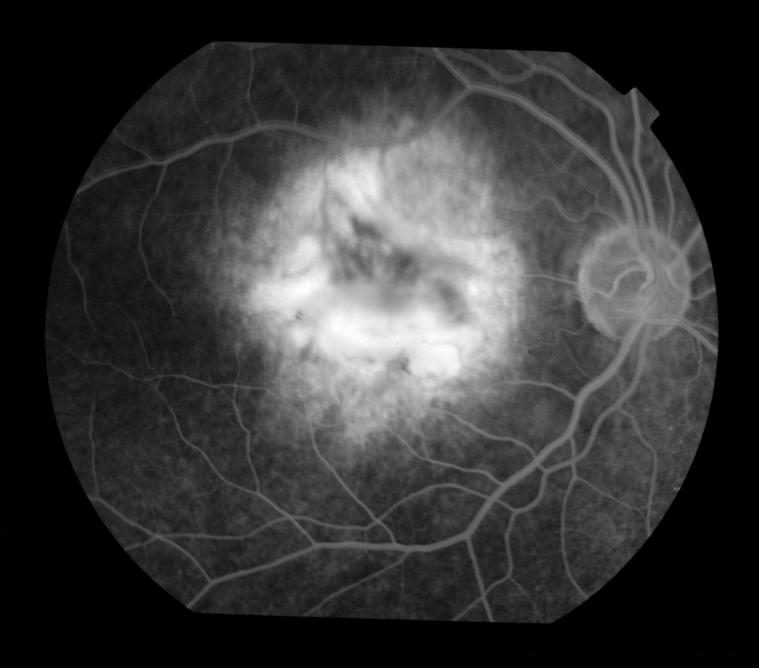
2 weeks post anti-VEGF





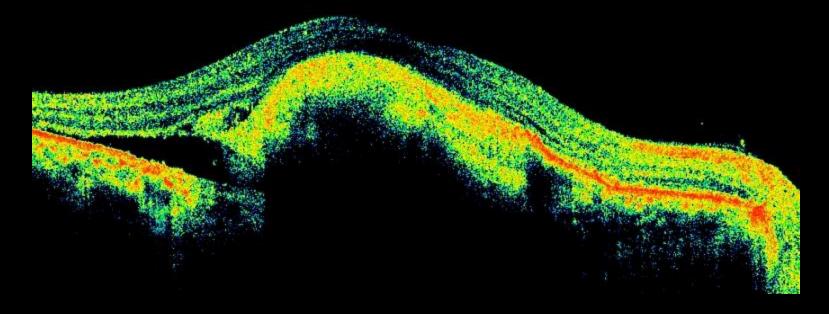






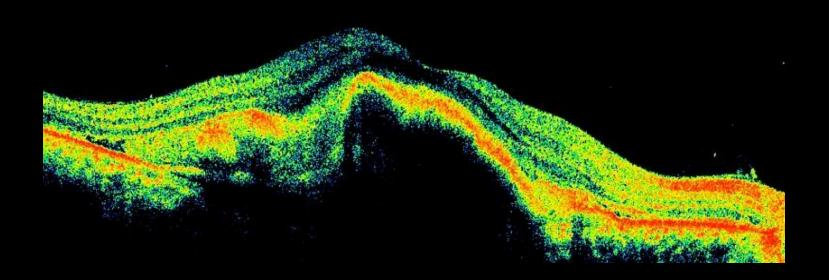
OCT more informative than FFA would be

- Large RPED
- Subretinal fluid
- Large scar



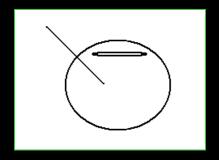
OCT more informative than FFA would be

Resolution of fluid on treatment



Anti-VEGF therapy Therapy of first choice

- Ranibizumab (Lucentis): 0.3 /0.5 mg
- Bevacizumab (Avastin): 1.25 /2.5 mg
- Aflibercept (Eylea/ VEGF Trap)2.0 mg



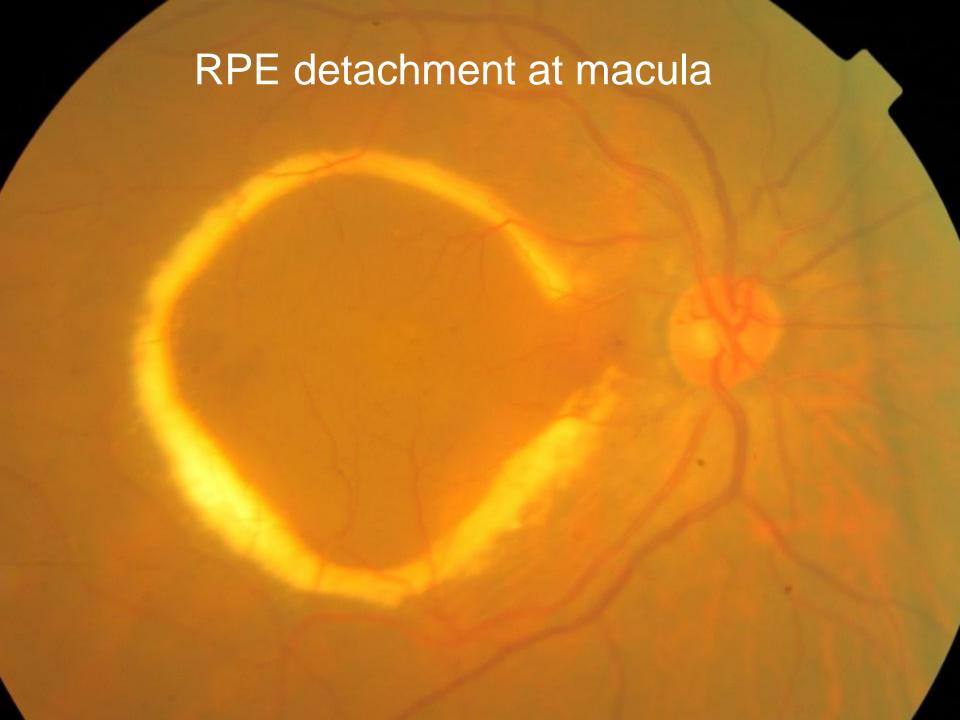
Complications of AMD Treatment Trial CATT

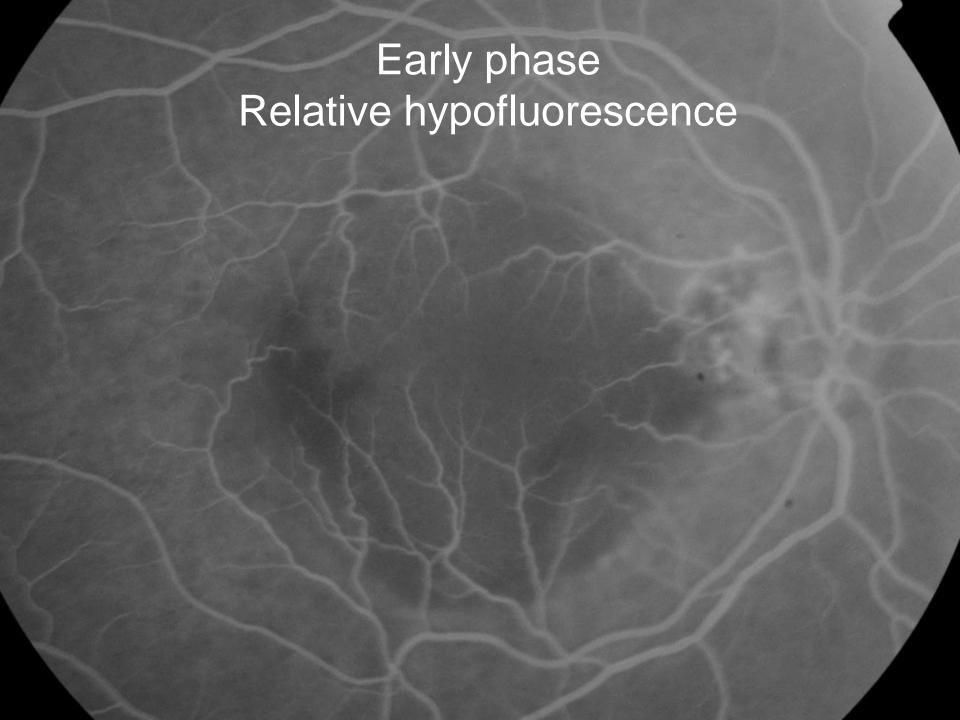
- Results support the use of either bevacizumab or ranibizumab for AMD
- As-needed regimen is an acceptable alternative to a monthly regimen
- Monthly evaluation necessary

High dose Avastin

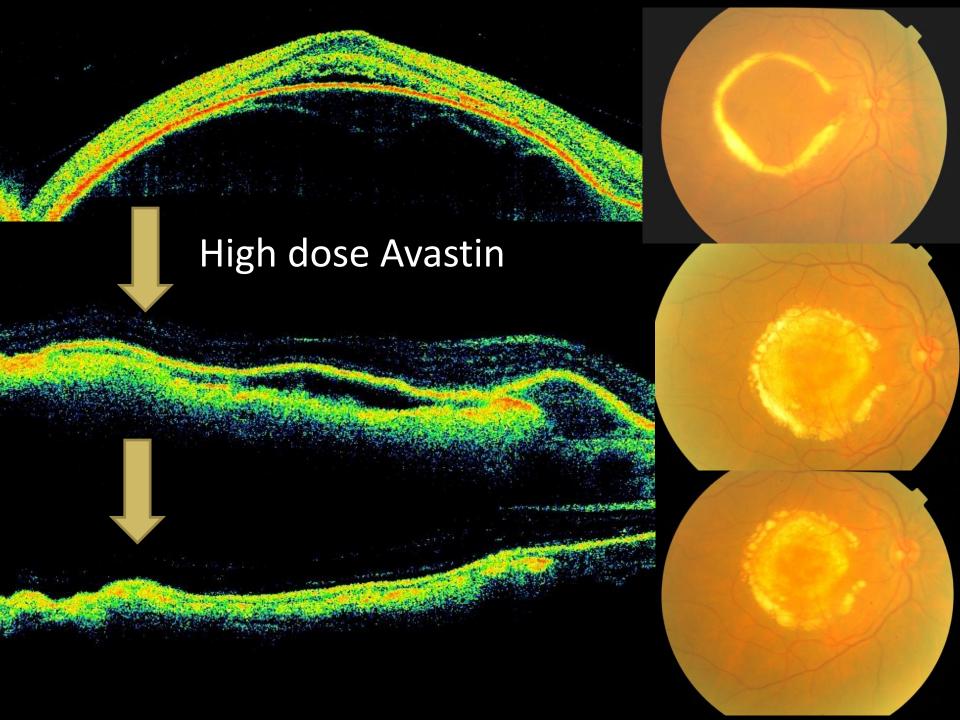
2.5 mg/ 0.1 ml every 2-4 weeks

Regular dosing: 1.25 mg/ 0.05 ml every
 6 weeks



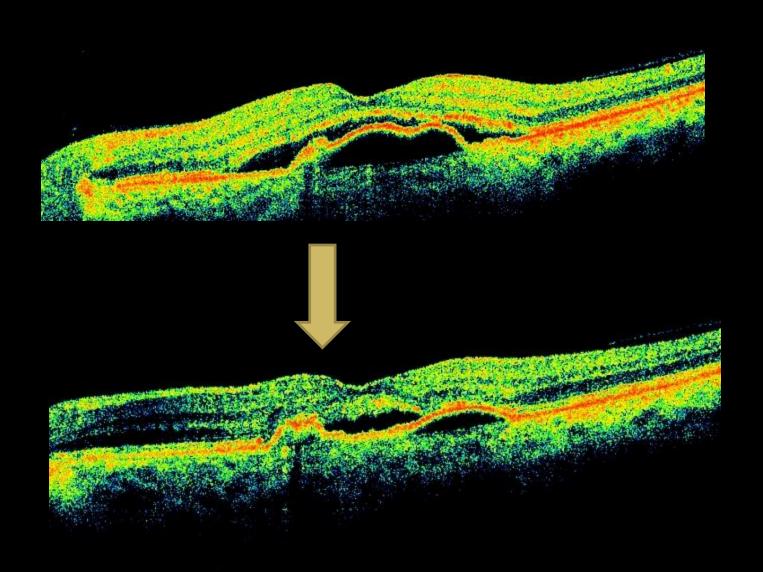


Late Phase Pooling of dye

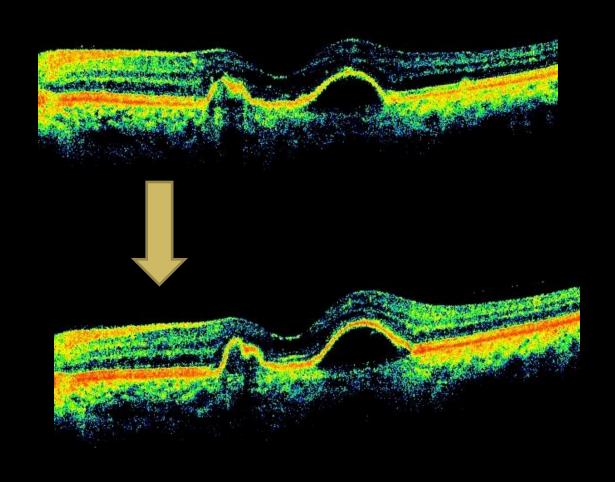


Anti-VEGF Switch

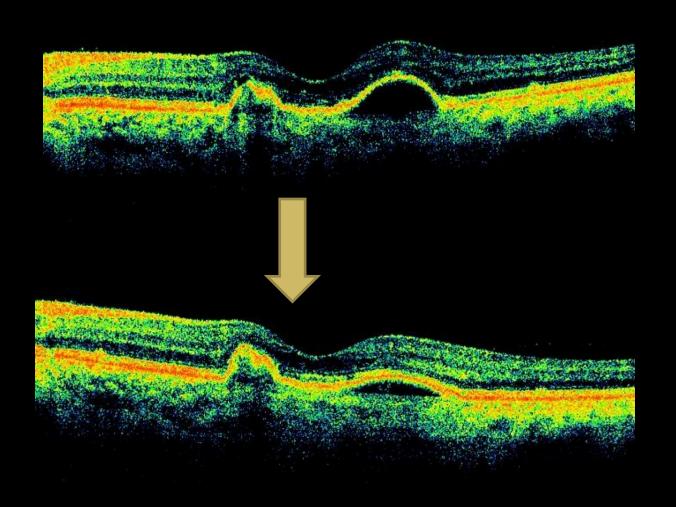
Mr Chatterjee Within 2 months of Avastin



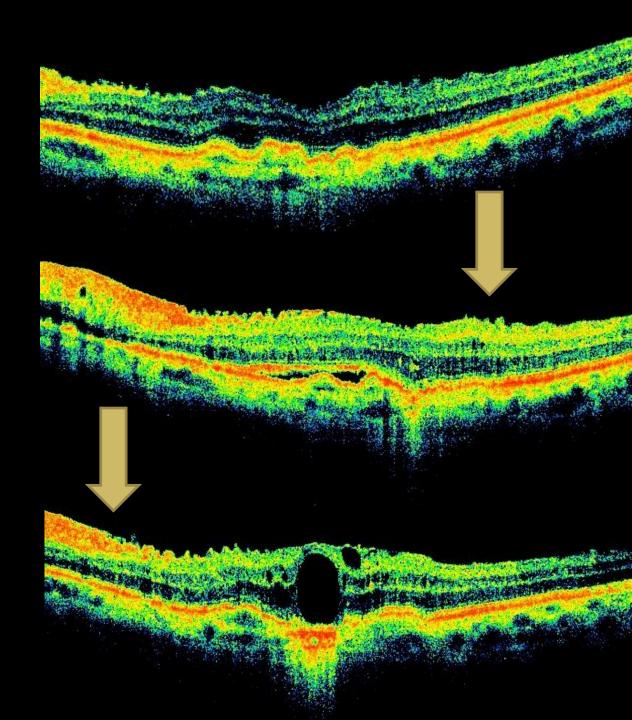
No change on High dose Avastin 1 year Sep 2013 to Aug 2014



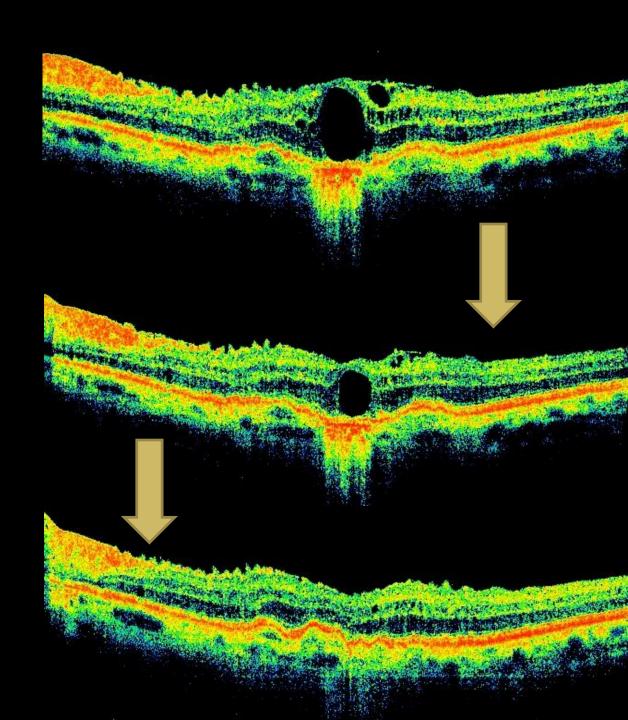
Following Lucentis Sep 4, 2014



Mr Chhatwal
Worsening on
Avastin



After 2 Lucentis



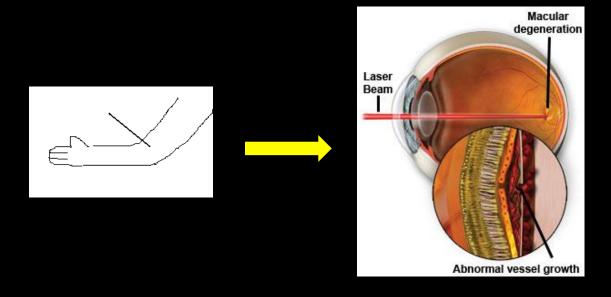
Photodynamic Therapy (PDT) with Verteporfin in AMD

Combination Therapy PDT + anti-VEGF therapy

- PDT addresses pre-existing CNVM
- Anti-VEGF prevents growth of new vessels

A 2-step Procedure

- Verteporfin infusion over 10 minutes
 (6 mg/m2 BSA; 3 ml / minute)
- 2. Red Light 689 nm, 50 J/cm² light, 600 mW/cm² for 83 seconds at 15 minutes

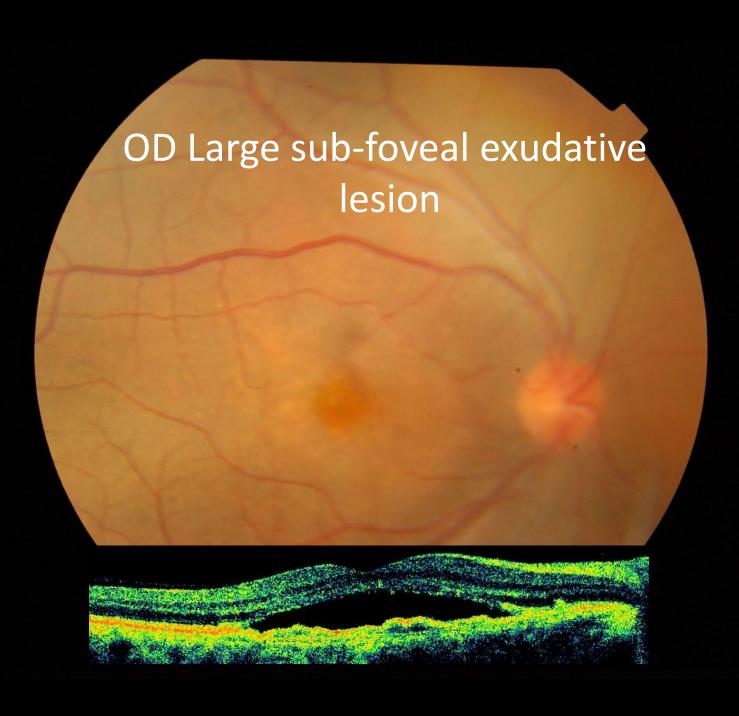


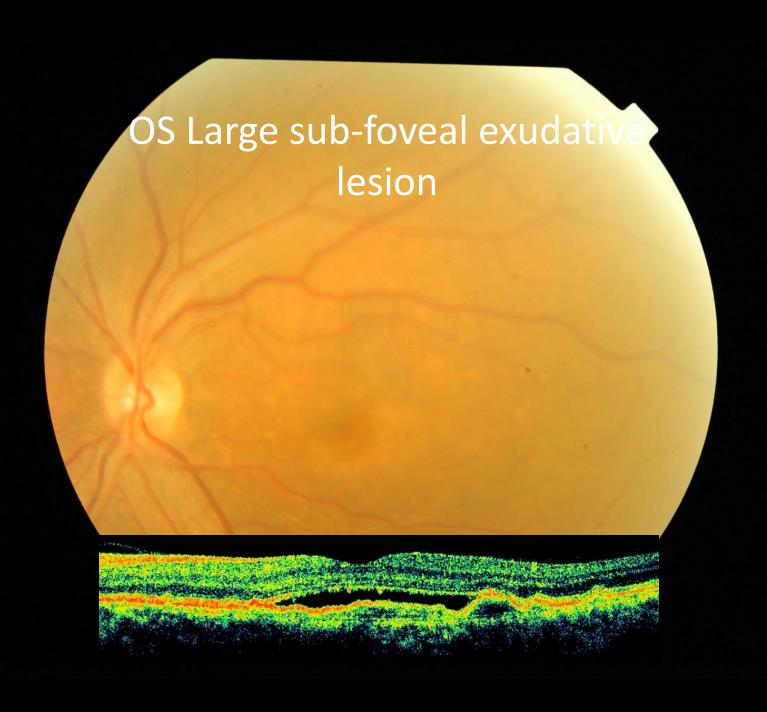
Indications for PDT in Macular Lesions

- CNVM non responding to anti-VEGF monotherapy
- RAP lesions
- PCV
- CSCR

Case 1. Presentation July 4, 2012

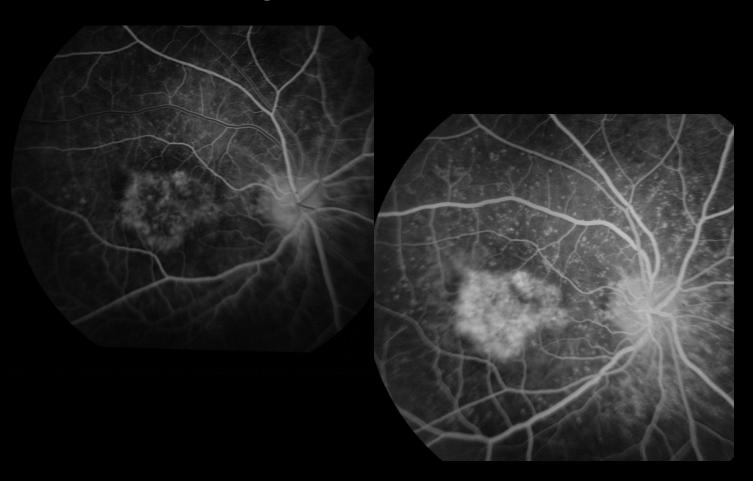
- 61 year old male
- c/o bilateral vision drop for several months
- Visual acuity was OD 20/100 OS 20/40



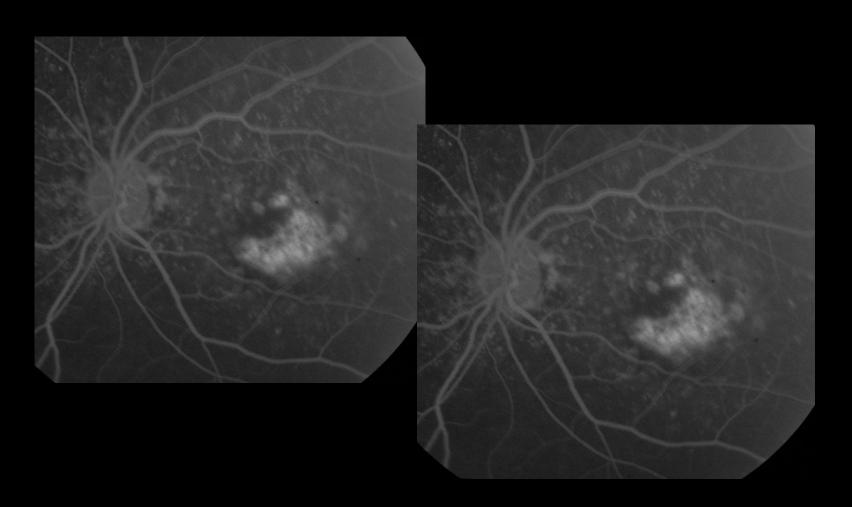


Fluorescein angiogram

OD Large classic CNVM

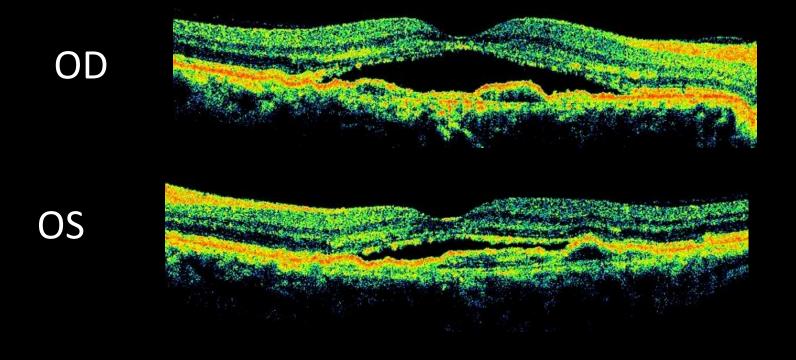


OS Large classic CNVM



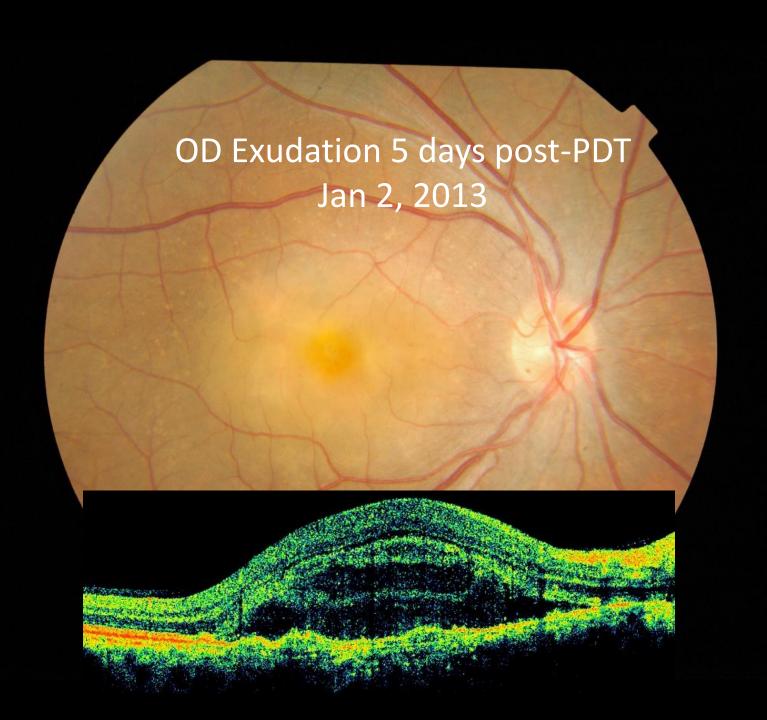
Monthly Bevacizumab for 6 months

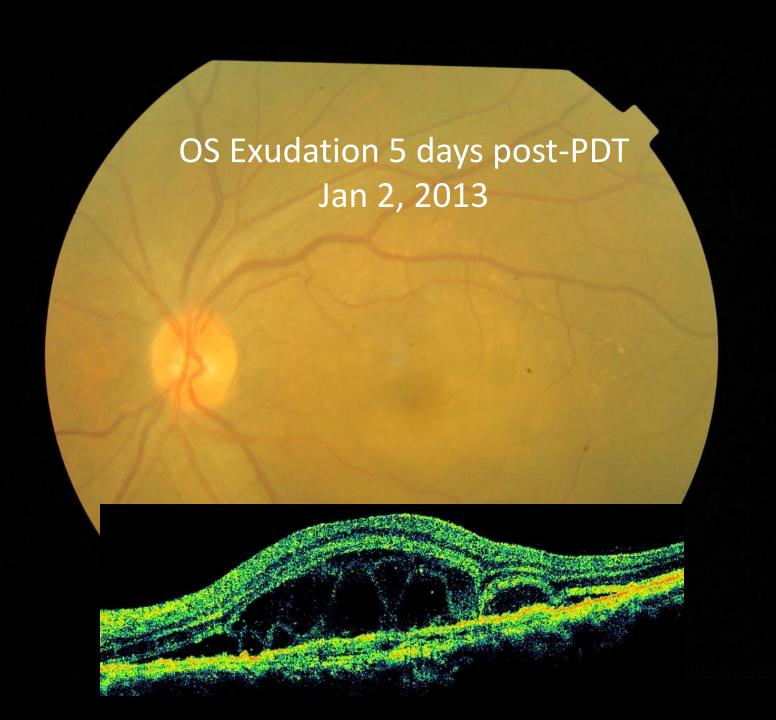
No Improvement



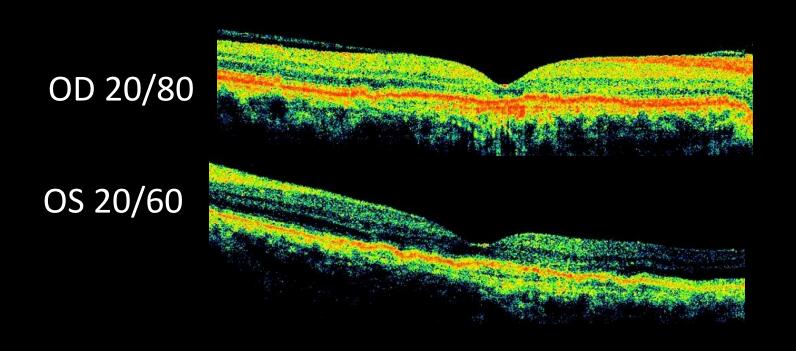
Dec 28, 2012

Bilateral Photodynamic Therapy Standard Fluence, Spot Size 3500 um

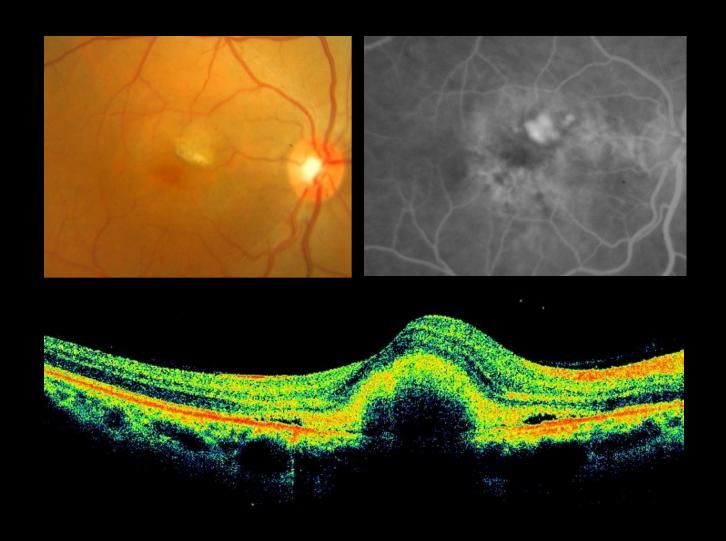




4 weeks post PDT, Feb 2013 to date Maintained on anti-VEGF monotherapy



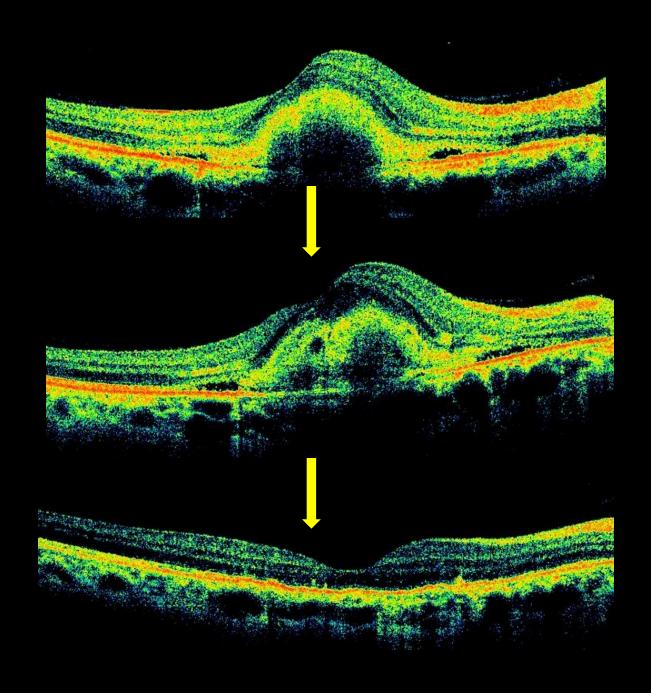
Case 2. Wet AMD, classic CNVM Worsening on anti-VEGF therapy over 4 months



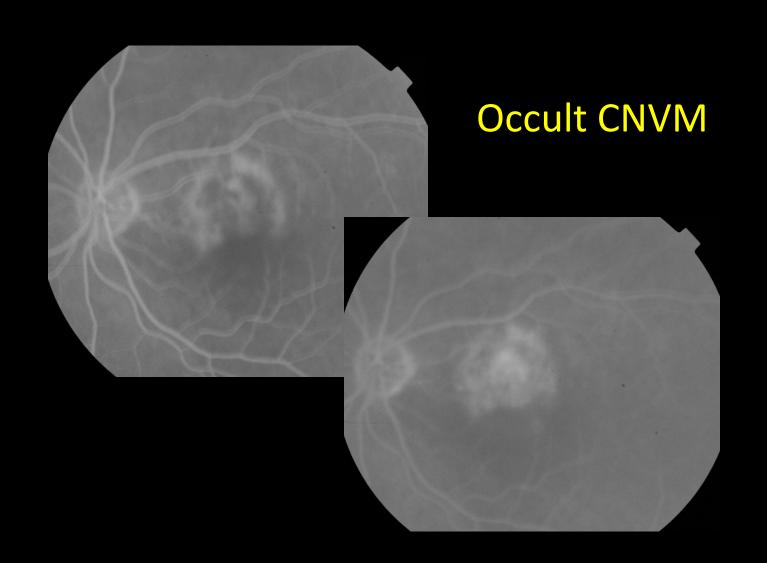
Case 2.

Increasing fluid on bevacizumab over 4 months

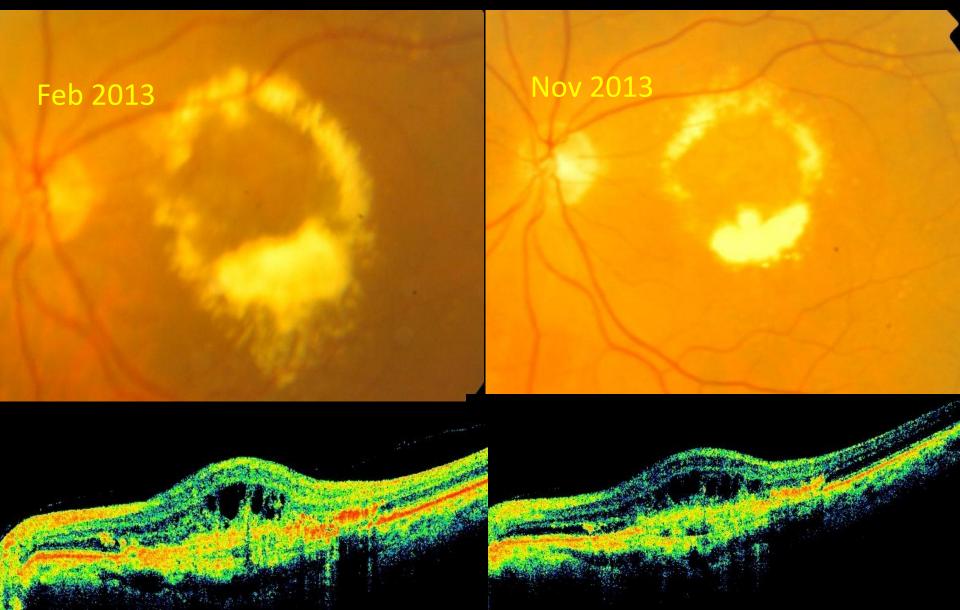
Complete resolution 4 weeks post-PDT



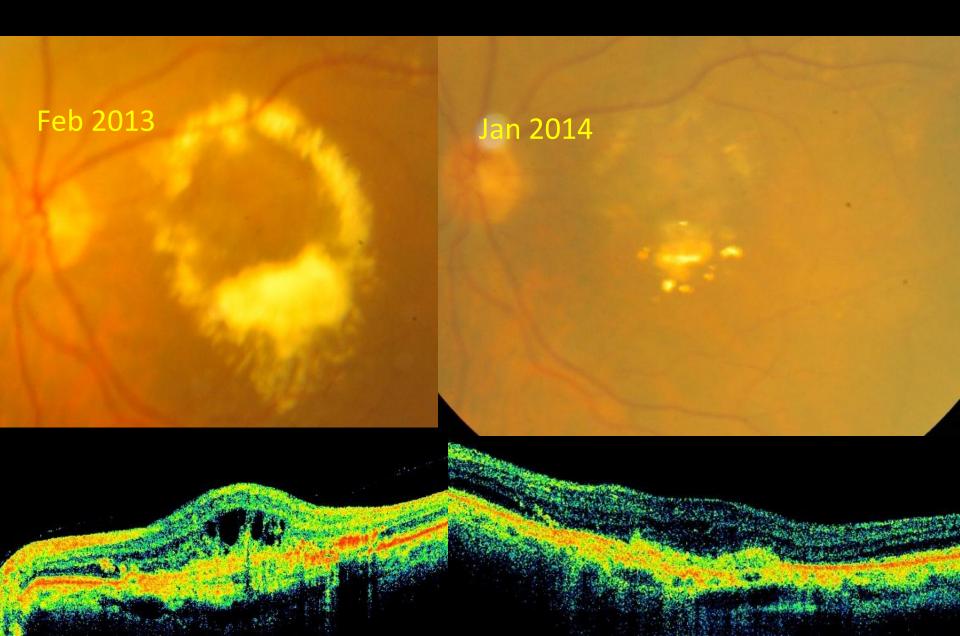
Case 3. Occult CNVM with exudates Ms Leela Feb 2013



Comparative over 9 months Anti-VEGF monotherapy



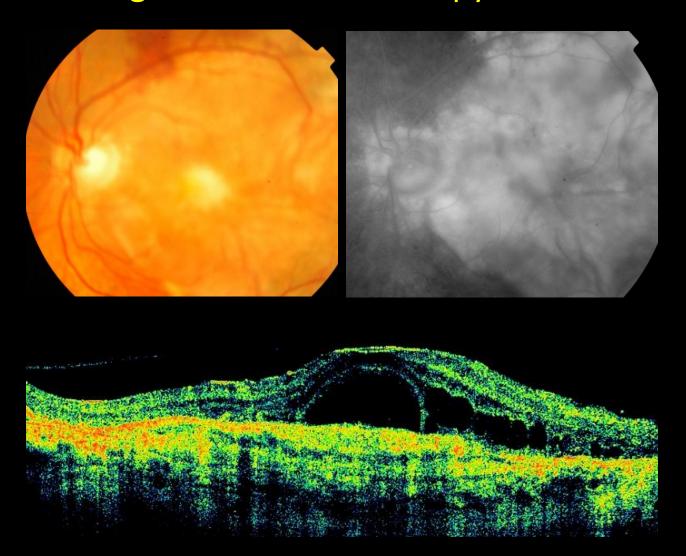
Post PDT 3 months

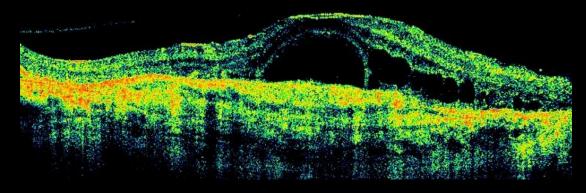


Steroids in wet AMD

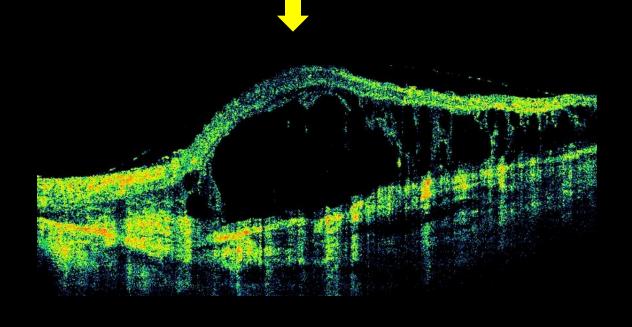
 Intravitreal steroid can be added to anti-VEGF therapy in some unresponsive cases

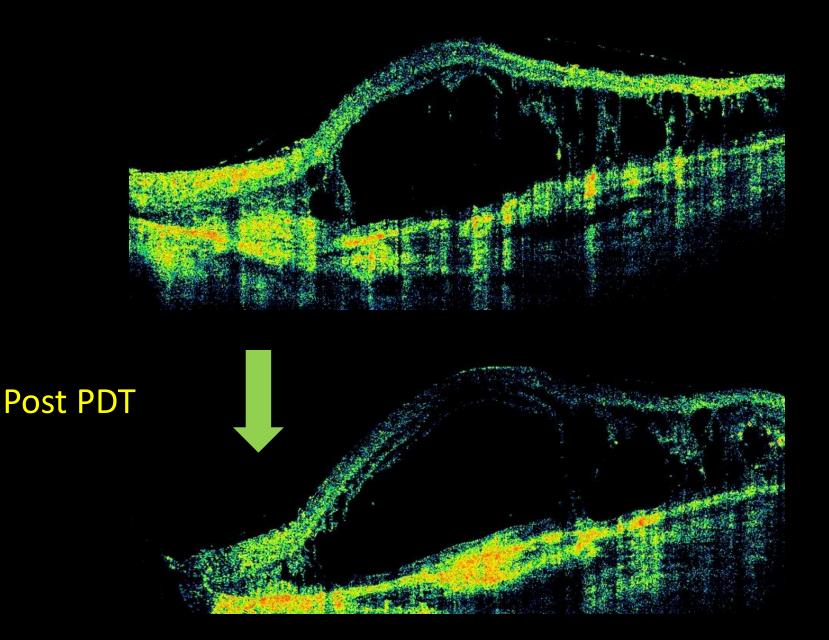
Case 1. Large exudative AMD lesion Worsening on anti-VEGF therapy over 4 months

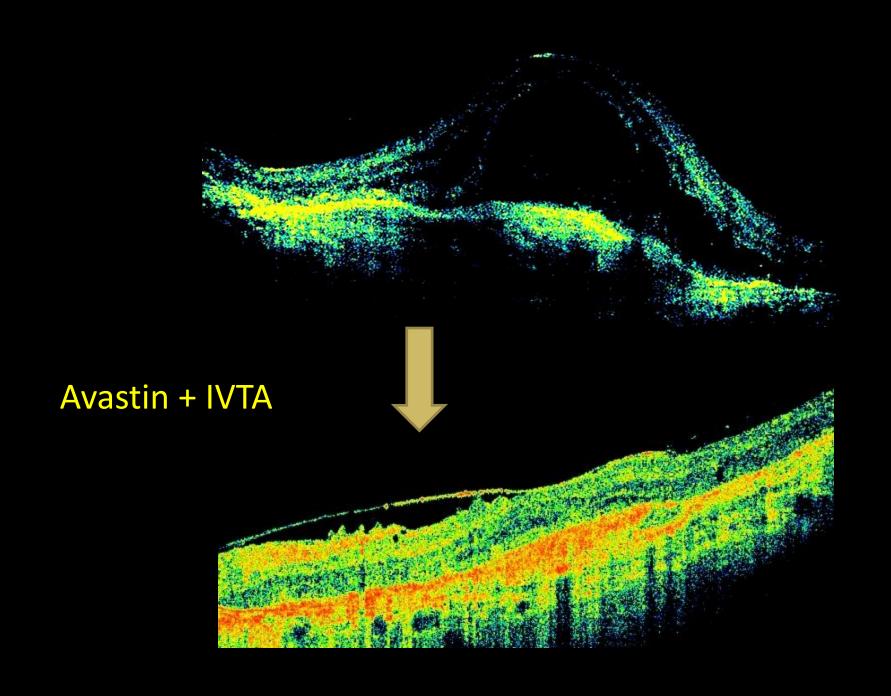




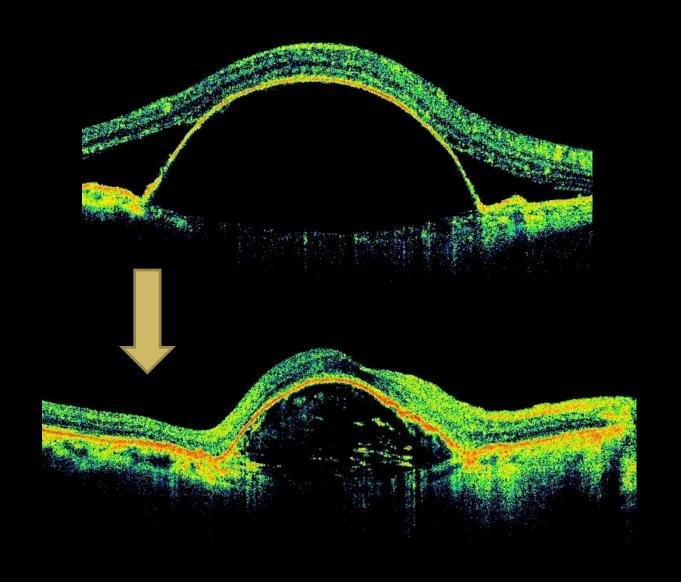
On Avastin for 4 months



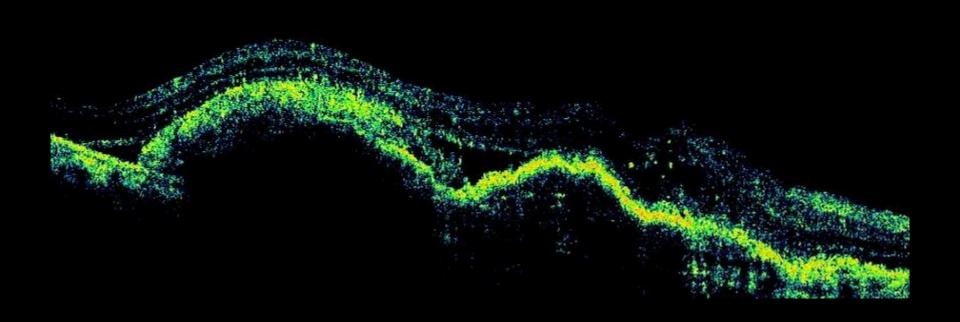




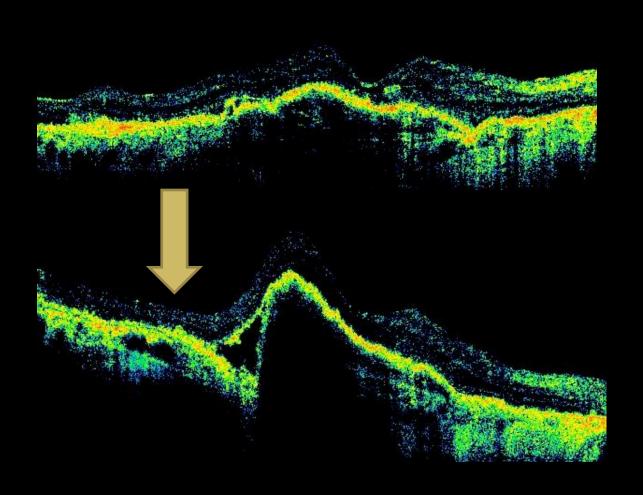
Case 2. Improvement with bevacizumab



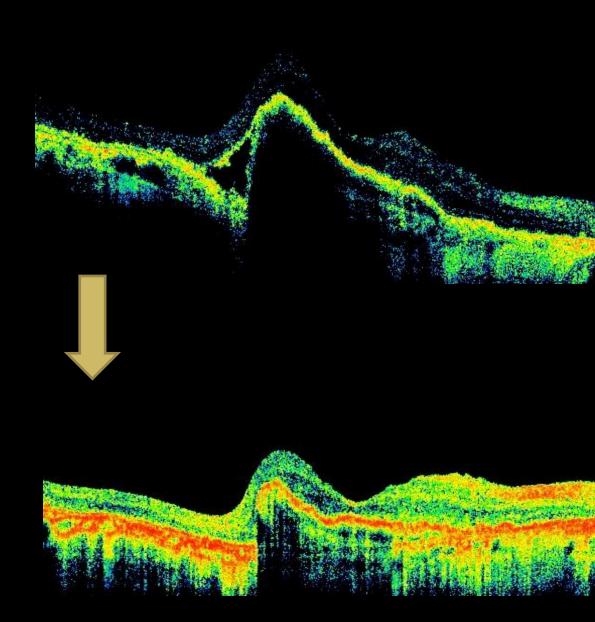
After a 9 month treatment free interval



Persistent fluid with avastin



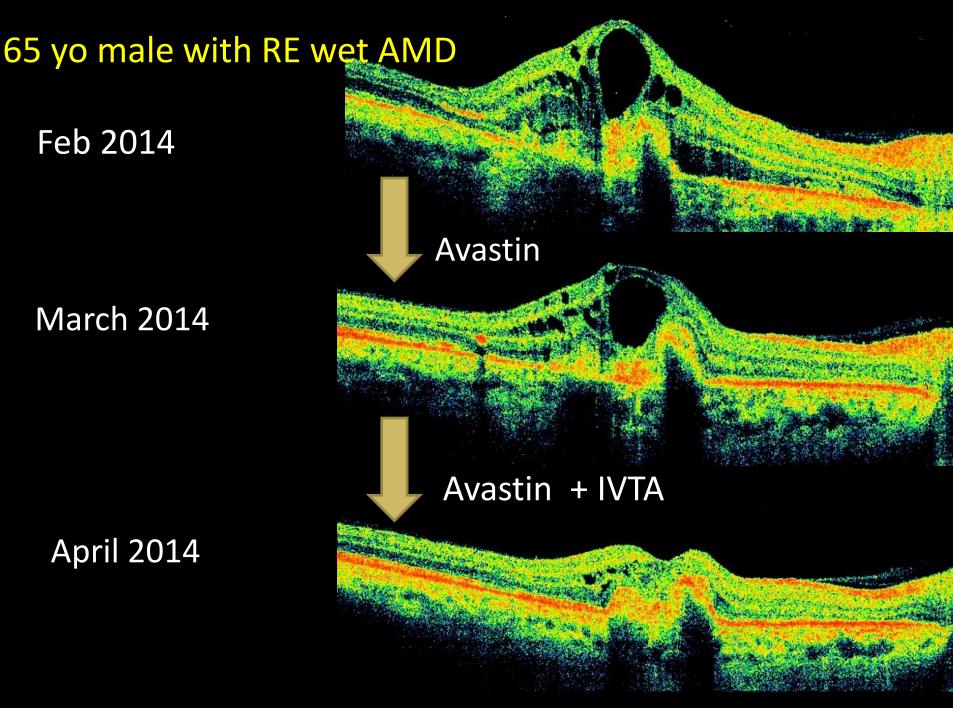
RE with steroid



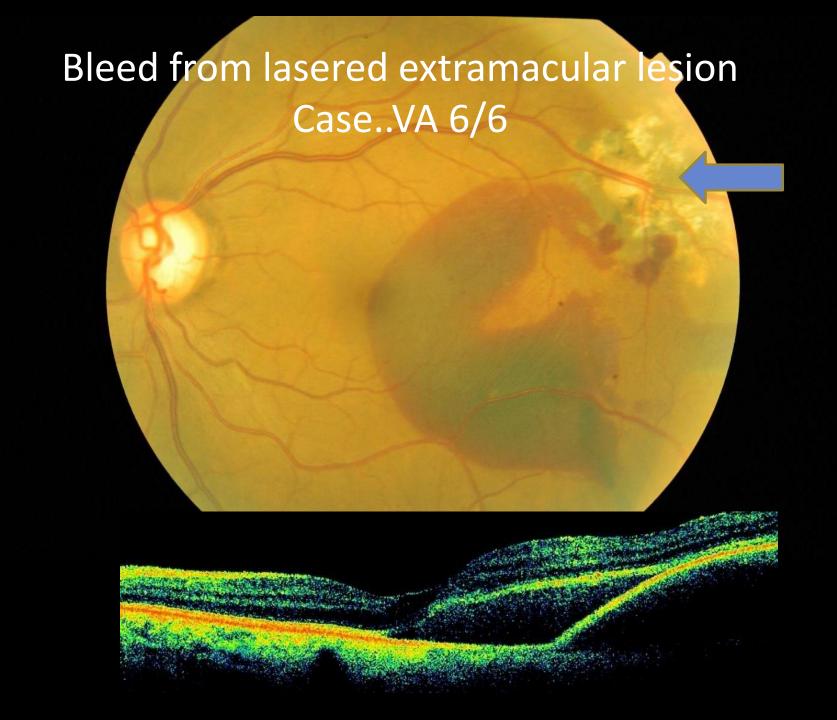
Feb 2014

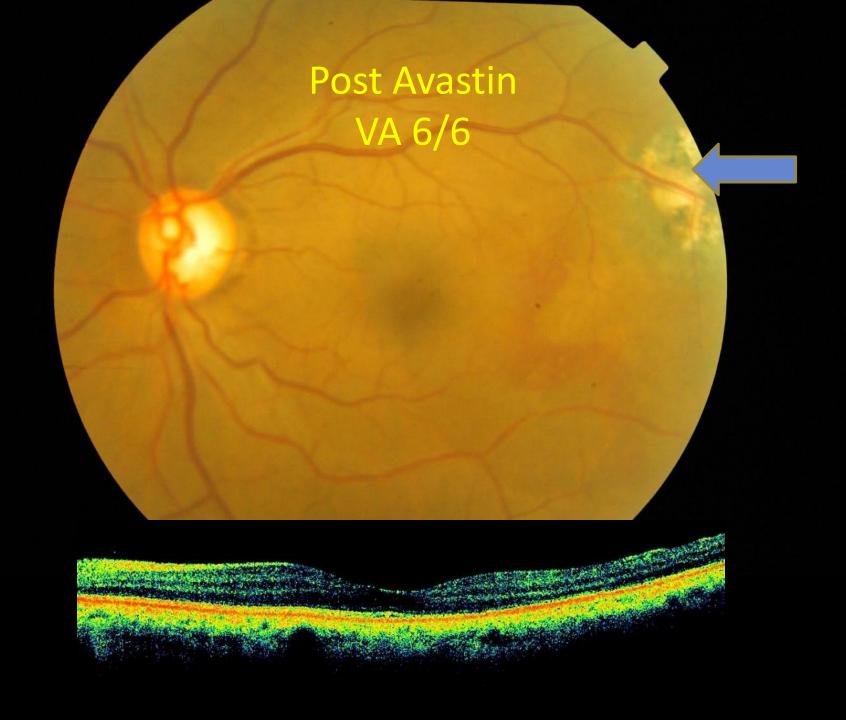
March 2014

April 2014



Extramacular lesions not always safe



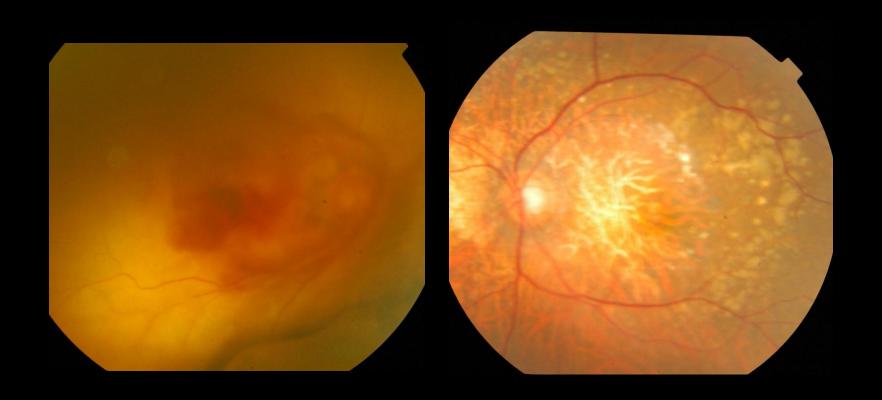


Concurrent Dry AMD Progression

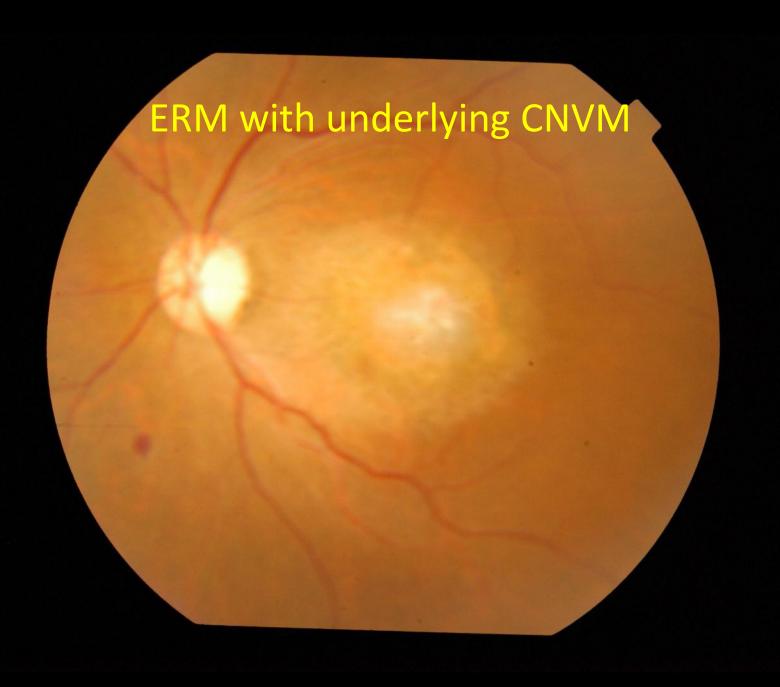
Antioxidants should continue

Vision loss from concurrent dry AMD progression

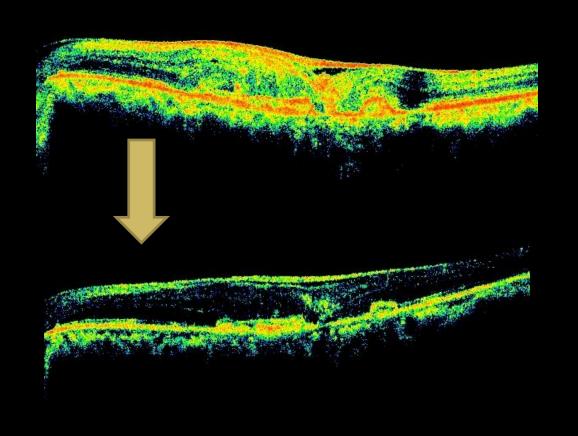
One eye wet AMD, other dry



Co-existing Problems



Avastin Sep – Jul 2014



Thank you!