### FUNDUS AUTOFLUORESCENCE

Syed M Shahid ST2 Ophthalmology Hillingdon Hospitals NHS Trust

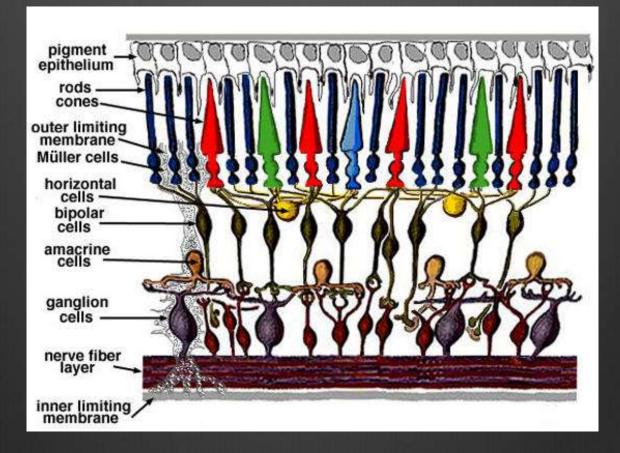
## INTRODUCTION

 Spontaneous emission of light by biological structures after absorption of light – used to distinguish light originating from artificially added fluorescent markers

 'Fluorophores' – chemical structures that possess fluorescent properties when exposed to light of an appropriate wavelength

 Absorb light moving to a higher energy state triggering light at a wavelength longer than the source

### Layers of the retina



## Retinal Pigment Epithelium

Wit A circulation

- Synthesise extra-cellular matrix
- Transport molecules
- Phagocytose outer disc of photoreceptors

- Affected by age and in disease
- Leads to accumulation of '*lipofuscin*'
- Plays a role in AMD and other hereditary macular disease

### Lipofuscin

Intrinsic fundus AF due to lipofuscin

Excitation with short to medium wavelength, leads to autofluorescence with peak emission of 630 nm

O Other fluorophores also identified ~10

One of the major fluorophores of lipofuscin is A2E

## Retinal conditions identifiable from AF patterns

- Hereditary
- Age-related
- Inflammatory
- Retinal pigment change due to laser therapy

### Natural fluorophores

- Lipofuscin
- Optic nerve drusen
- Astrocytic hamartomas
- Aging crystalline lens



Edward S. Harkness Eye Institute Columbia University

## Clinical applications

 Macula pigment density and pigment densities elsewhere in the retina to evaluate diseases

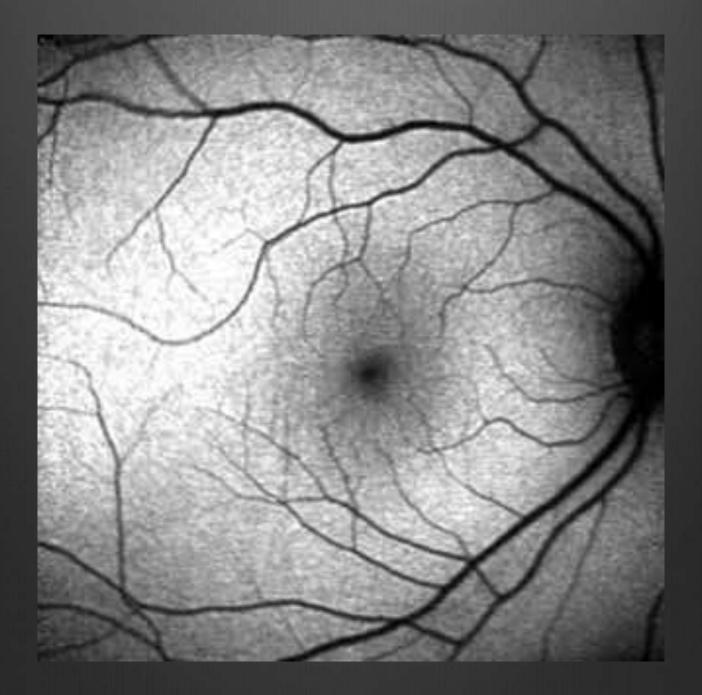
 Newer applications using macula pigment density to evaluate oxygen dependent cell metabolism

### Normal AF pattern

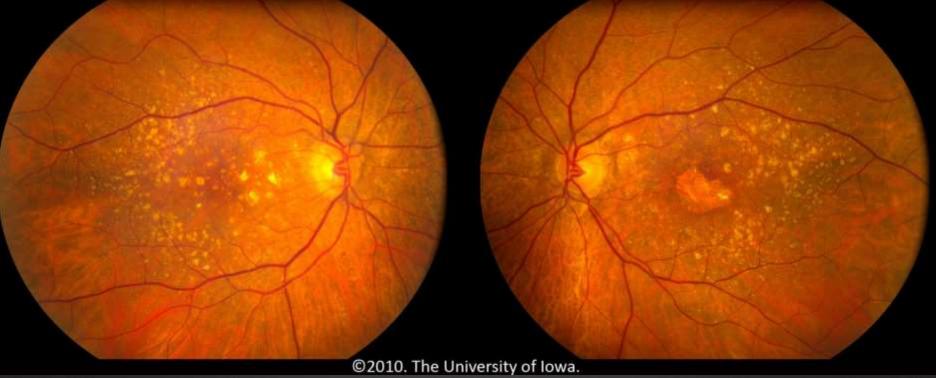
Decreased AF in perifoveal area – lutein and zeaxanthin

ON and vessels appear dark

Area around vessel arcades has the highest intensity of AF decreasing towards the periphery

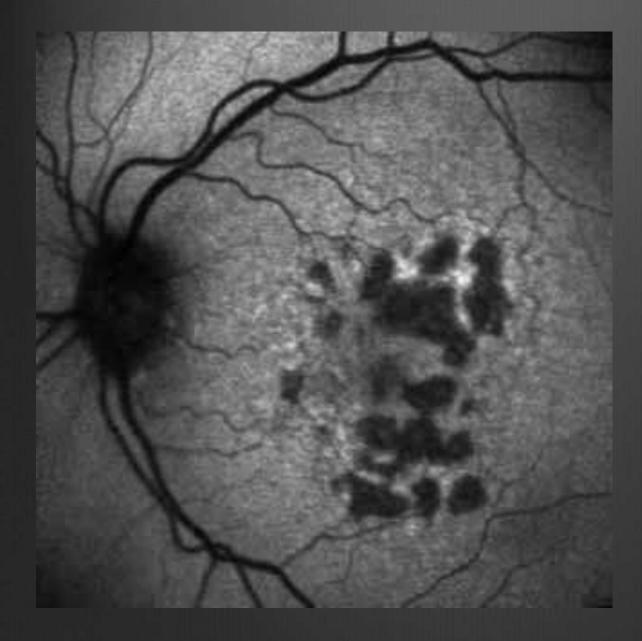


## AGE RELATED MACULAR DEGENERATION

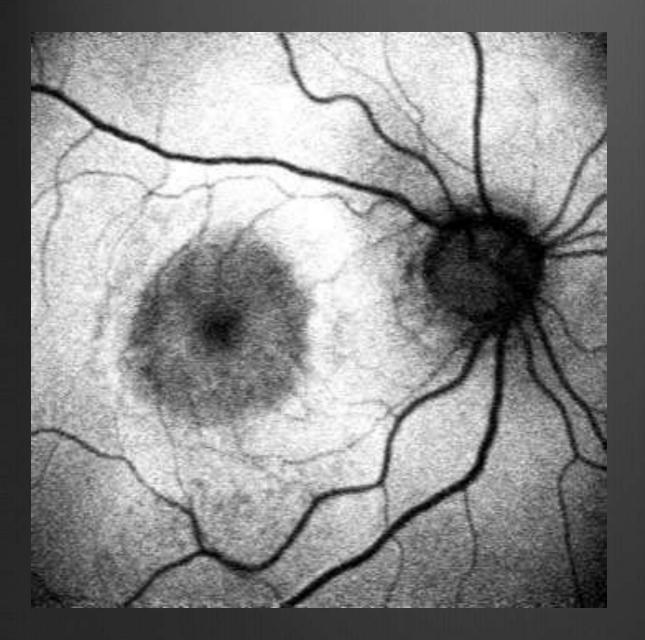




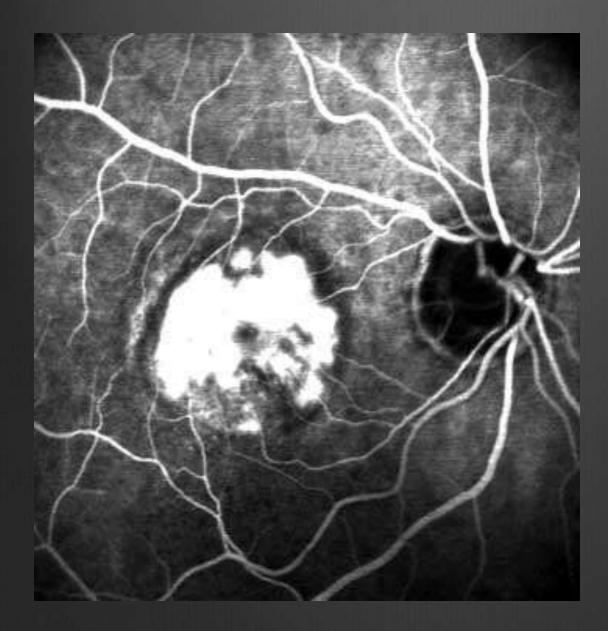




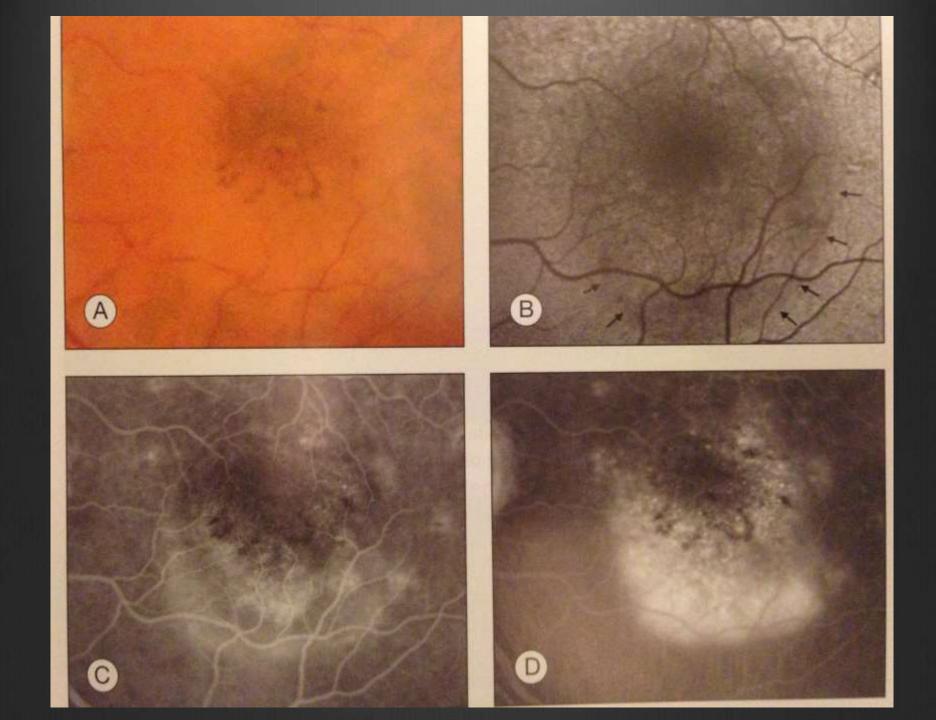
- Geographic atrophy – Advanced AMD. RPE cell loss causes dark AF
- Increased AF in the junctional zone – lipofuscin accumulation
- Atrophy will spread to areas of increased AF



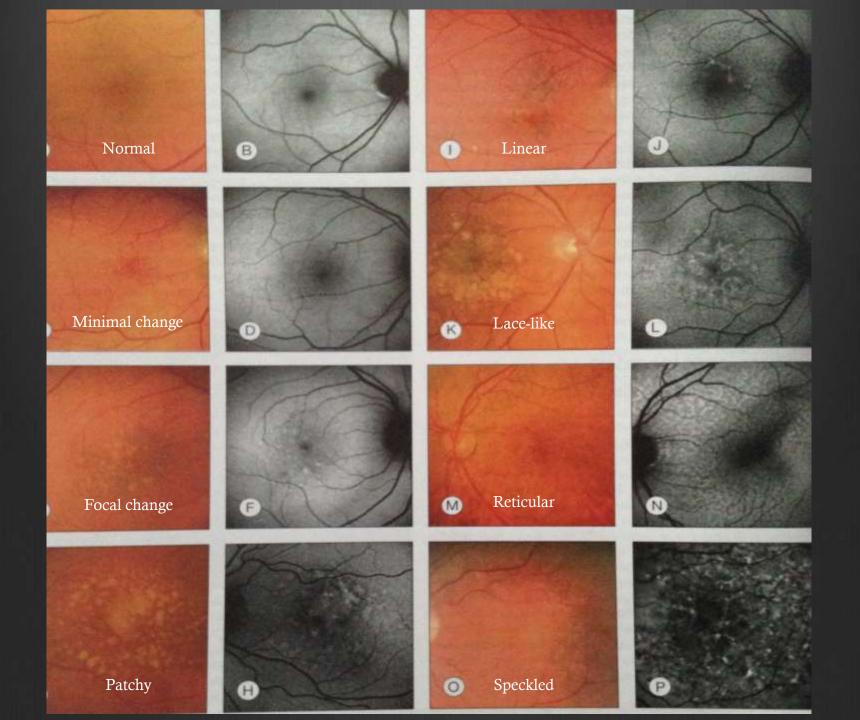
Well demarcated
blockade of AF
due to CNVM
lying above RPE



 Corresponding FA shows a classic
CNVM due to advanced AMD



- AF patterns vary in early forms of AMD define and characterize the clinical picture
- Tool for giving valuable prognosis in those with early AMD
  - Enhanced perifoveal AF might lead to earlier and faster progression to geographic atrophy (higher levels of intracellular lipofuscin in the junctional zone and thus higher risk of cell death)
- Distinguish between classic and occult CNVM
  - Decreased AF in classic over the whole area surrounded by slightly enhanced AF; slightly irregular and enhanced AF in occult
- Heterogenous AF patterns in CNVM indicates progression to sudden irreversible visual loss – macular oedema, SRH and scarring



# DRUG TOXICITY

### Chloroquine & Hydroxychloroquine

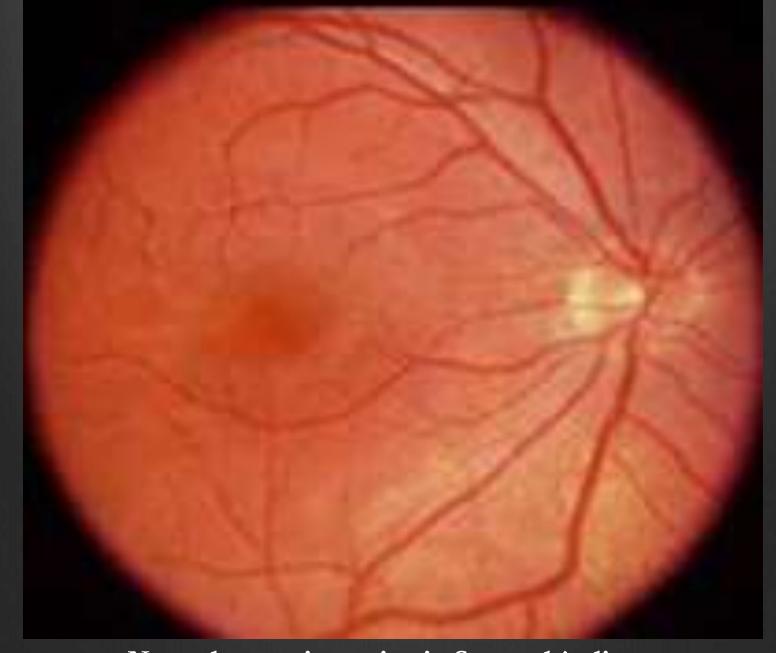
- Early on a pericentral ring of increased AF intensity
  - Associated with pericentral reduction in multifocal ERG amplitudes and pericentral disruption of PRC IS –OS junction on OCT imaging
- Advanced stages mottled appearance with increased and decreased AF intensity in the pericentral macula



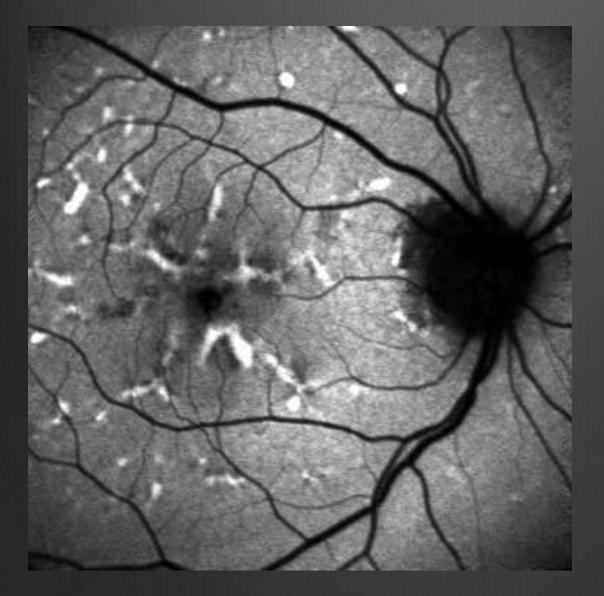
# HEREDITARY DISEASES



### Stargardt's diseae

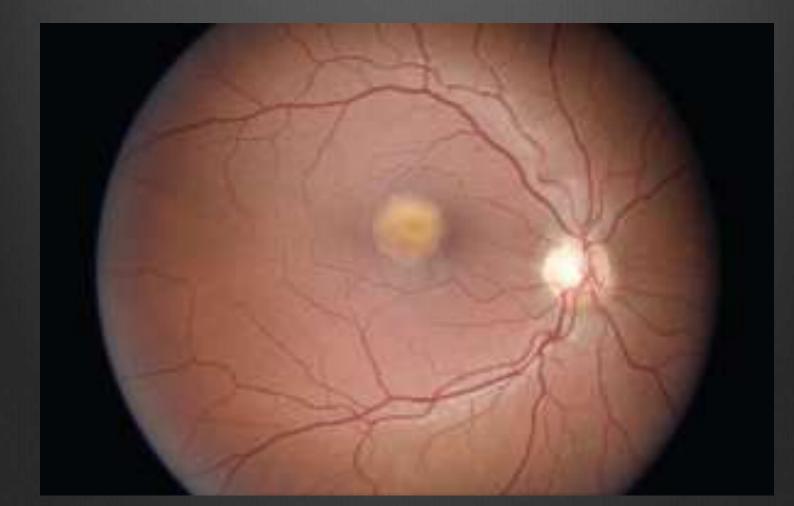


### Normal appearing retina in Stargardt's disease



- Diffusely enhanced and focally increased AF seen in normal appearing retina
- Pathologic accumulation of lipofuscin
- AF pattern might also be very heterogeneous (correlations of AF phenotype with genotypic phenotypes)

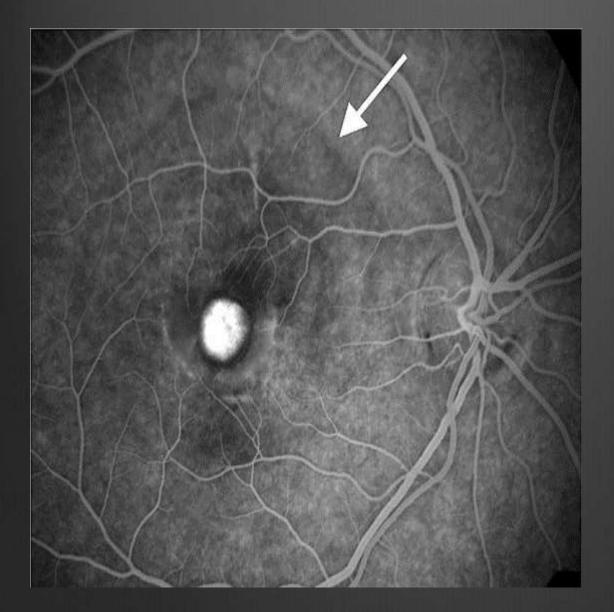
### Best Disease



'Egg-yolk' appearance

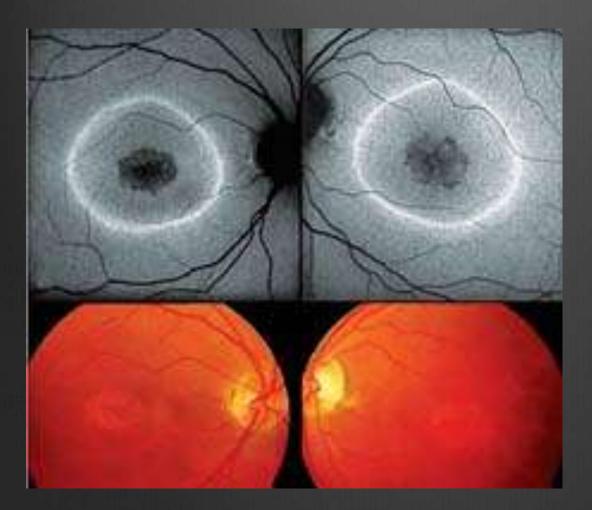


 Excessive accumulation of lipofuscin in the central area showing highly increased AF



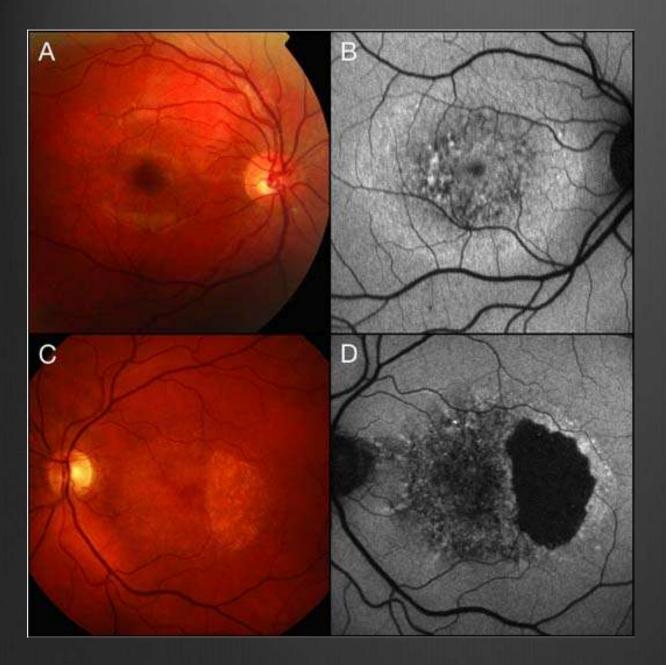
FFA imageBEST disease

#### MACULAR DYSTROPHY



Ring of increased autofluorescence in the parafovea which is not visible on fundus photography. Functional testing reveals that this ring correlates with functional abnormalities and represents a demarcation line between normal and abnormal

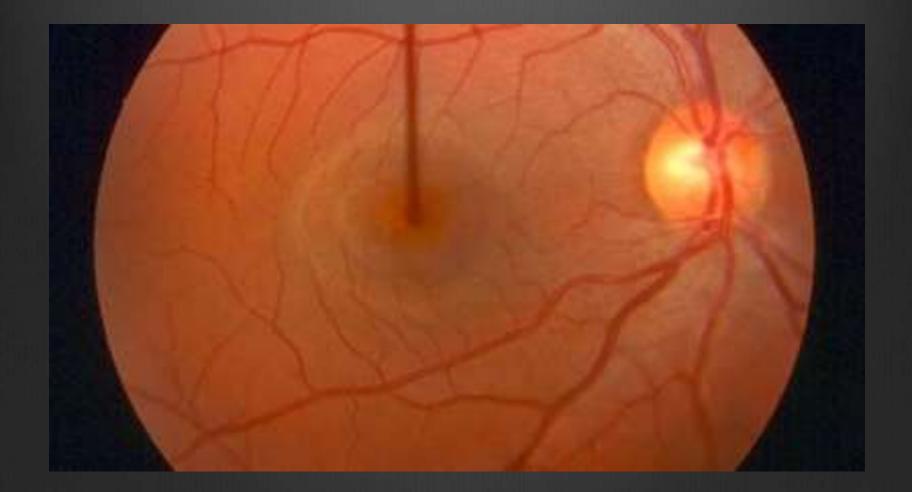
functional retina.

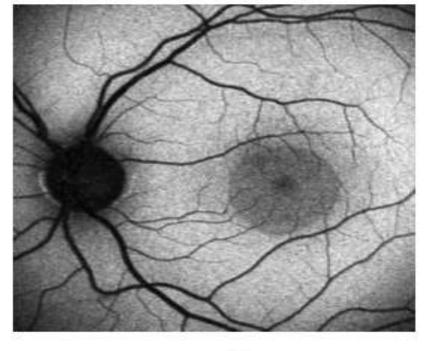


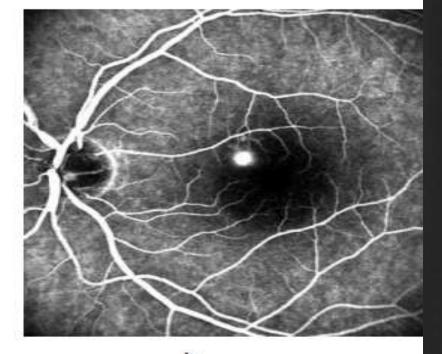
Pattern dystrophy with corresponding autofluorescence images

## INFLAMMATORY CONDITIONS AFFECTING RPE

### Central Serous Retinopathy

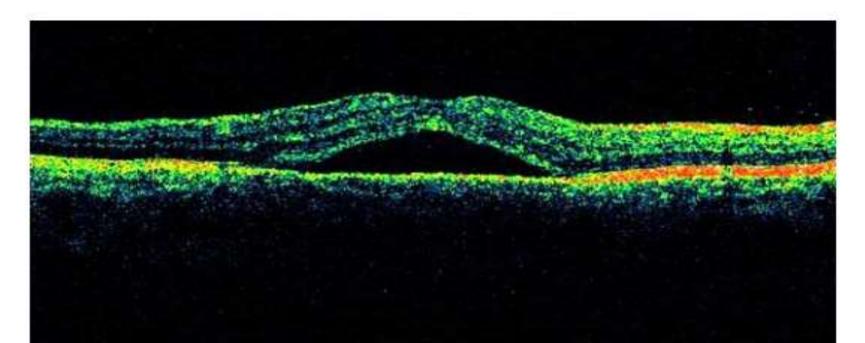




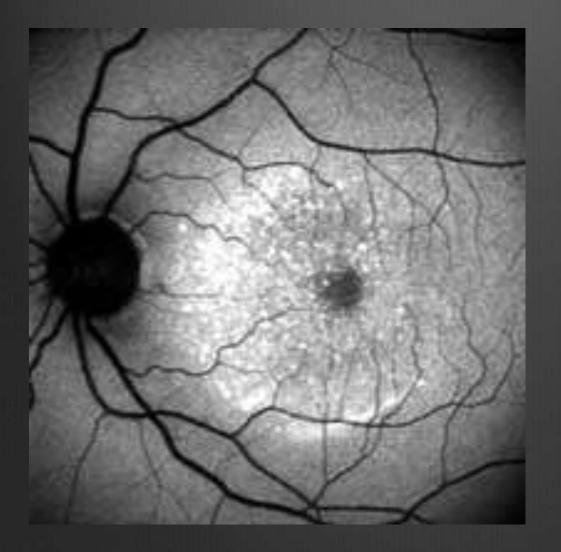


a

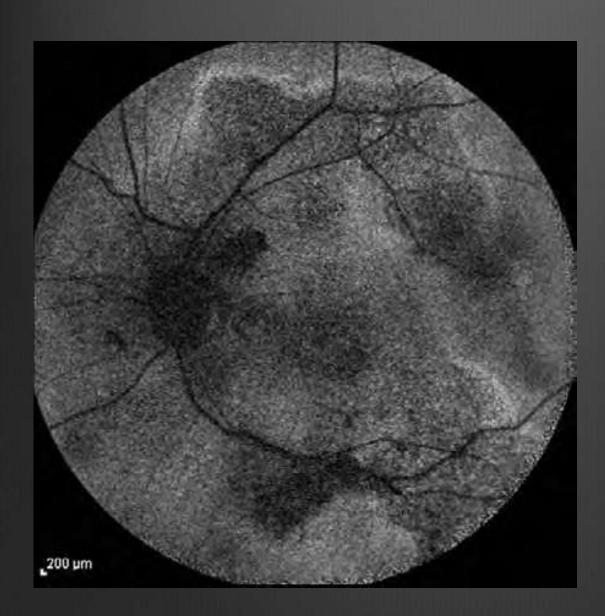




### Resolved CSR



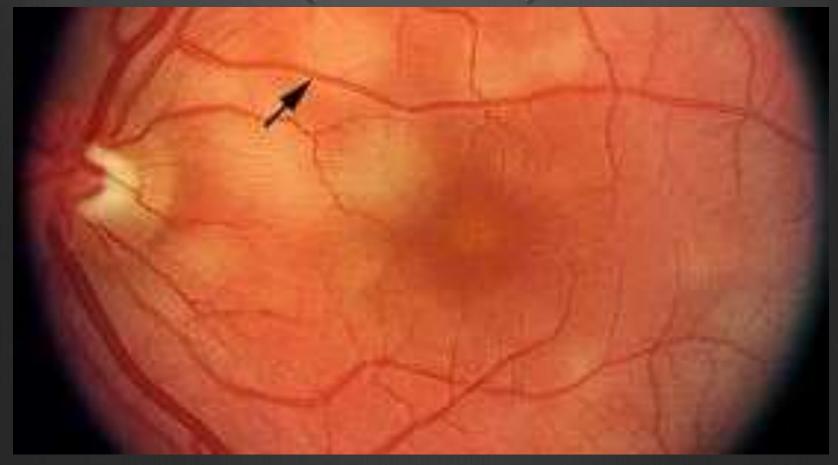
 Increased AF within the former neurosensory elevation most likely due to higher metabolism of RPE cells from the protein enriched subretinal fluid



- Extensive AF pattern in patient with chronic CSR
- AF can help differentiate the edge of RPE dysfunction
- There is hyperautofluorescence either due to shed PRC or abnormal accumulation of lipofuscin A2E



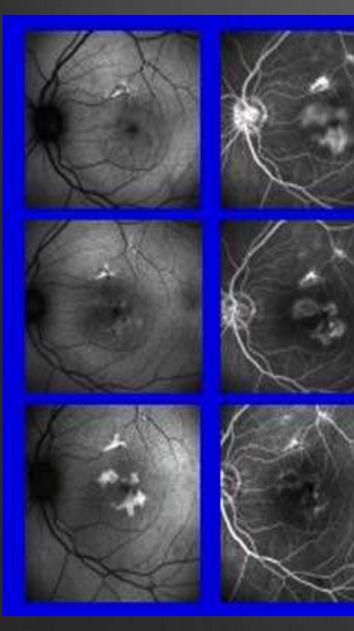
### Acute posterior multifocal placoid pigment epitheliopathy (AMPPE)



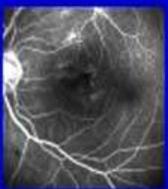
### AMPPE

AF sequence of an eye with APMPPE

- Affected RPE cells demonstrate increased AF in the early phase
- An increase in AF is seen up to 3 weeks
- Gradual fade away after this
- Atrophy occurs after 1 year leading to decrease AF in former lesions







6wo

3d

3wo



12mo

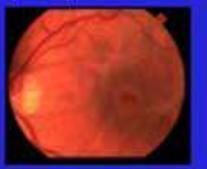
# MACULAR HOLE SURGERY



Bare RPE cell layer
causes increased AF
with no blockage from
macular pigments

#### Macular Hole Surgery

#### preop



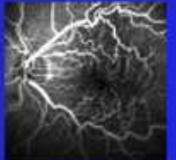


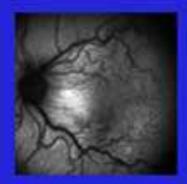


Visual acuity 0,1; macular hole and epiretinal gliosis

#### postop







Visual acuity 0,2 (2 months postop)

• Pre op increase in AF and disappearance after successful closure

## SUMMARY

- AF imaging is a novel tool of intrinsic fluorescence RPE layer
- It is primarily a result of intracellular accumulation of lipofuscin with age and alterations in different disease states
- Improvement in fundus cameras have enhanced imaging from AF
- Non-invasive, inexpensive and fast
- Can detect and add information for a variety of macular diseases
- Monitoring tool after surgery and laser treatment
- May contribute as an adjunctive tool for new treatment modalities
- Wastly underused diagnostic tool