#### Uveitis: Management with Vitreoretinal Surgery?

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No Financial Interests to Declare



# Professor Ingrid Kreissig The first woman in Germany who got a chair for Surgery in Ophthalmology (1979-2000)



## Always Energetic and Enthusiastic





# Magic Course in 2015



Deeply touched by Professor Kreissig's

Passion in Teaching



# Dr. Kreissig is Enjoying the Contact with the Juniors



## My Journey

- Trained as vitreoretinal surgeon at Taipei Veterans General Hospital, Taipei, Taiwan
- Research fellow at Doheny Eye Institute with Dr Steve Ryan
- President of Taiwan Retinal Society (2013-2015)
- President of Taiwan Ocular Inflammation Society (2020-)
- International society member: EURETINA, Jules Gonin Club, the Retinal Society, International Ocular Inflammation Society (IOIS)

# Happy Learning Time at Doheny Eye Institute



Research meeting with Ryan's team

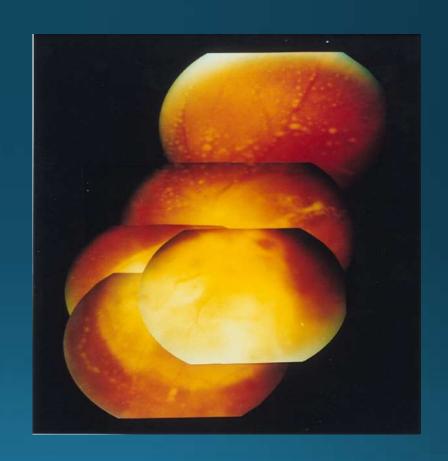
Professor Rao implant the concept of uveitis

# Exciting Experience for a Surgeon



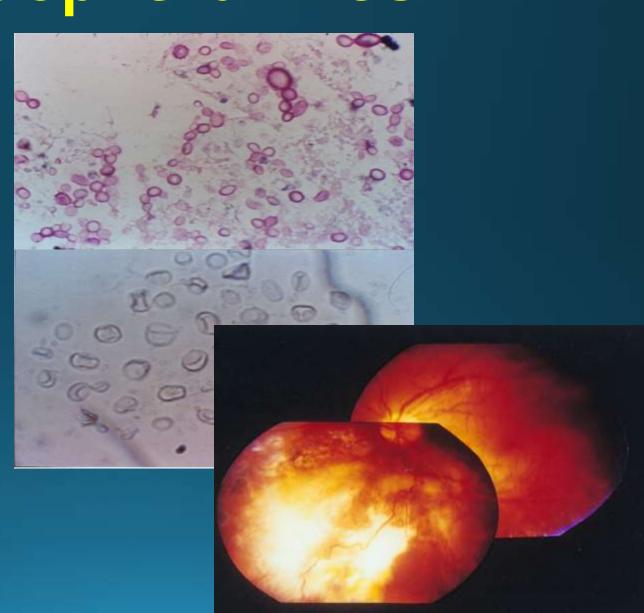
# My First Retinal Biopsy for the Diagnosis of Uveitis

- 45 y/o female, SLE under steroid and cytotoxic agents
- Panuveitis with a huge retinal abscess
- No response to IV/IVI ceftazdime/vancomycin
- Pars plana vitrectomy with retinal biopsy



# Cryptococcal Endophthalmitis

- 45 y/o female
- SLE under steroid and cytotoxic agents
- No response to IV/IVI ceftazdime/vancomycin
- Pars plana vitrectomy with retinal biopsy - Cryptococcus neoformans
- IV/IVI with amphotericin B
- Inflammation resolved



## Vitreoretinal Surgery in Uveitis

#### **♦**Diagnostic:

- > Atypical clinical presentations
- Rapid progressive course with no conclusive results
- Non- or sub-optimal response with standard treatment and masquerade syndrome.

#### **♦**Therapeutics:

- Management of uveitis: complications, cataract, glaucoma, hyphema, hypotony, epiretinal membrane, retinal detachment, vitreous hemorrhage.
- Clear visual opacities and decrease inflammation in cases of persistent inflammation.

## Microincision Vitrectomy (MIVS)

- Compared to traditional 20G, MIVS offers
- smaller wound
- less intra-operative bleeding
- higher patient comfort postoperatively
- ◆ Compared to vitreous tap for vitreous sampling
- PPV can obtain larger sample volumes for diagnosis
- avoids vitreous traction created by the needle
- allows directly proceeding to therapeutic vitrectomy

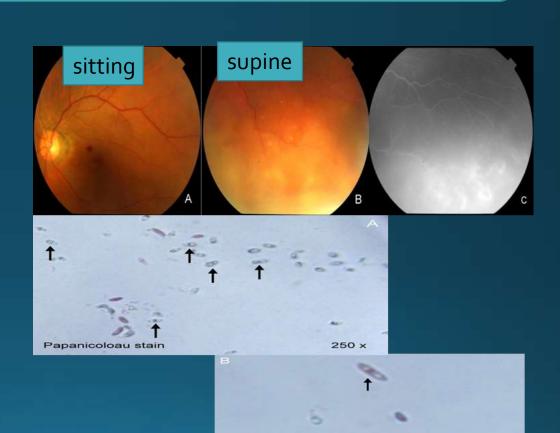
#### Most Common Etiologies for Diagnostic PPV

- 405 consecutive diagnostic PPV, 1973-1994
- Infection in 14.8% of all PPV
- Neoplasm in 14.3% (lymphoma) of all PPV
- Some examples: acute retinal necrosis, vitreo-retinal lymphoma, amyloidosis, sarcoidosis, endophthalmitis
- Overall positive rate: 14-64% (Infection: 67%, Lymphoma: 33%)

#### Diagnostic PPV for Infection

- Vitreous samples
  - Smear-positive rate (Gram stain): 66%
  - Culture-positive rate: 44%–67%
- Other examples for molecular diagnosis
  - PCR: viral retinitis, toxoplasmic chorioretinitis, tubercular uveitis, Cutibacterium acnes, and fungal/bacterial endophthalmitis
  - Multiplex PCR, DNA arrays or next generation sequencing

- 44 y/o male
- Intermittent blurred, OS for 2 yr
- Recurred 3 days after cataract surgery
- Presented as panuveitis with exudative RD initially, but progressed to RRD during systemic survey
- PPV with FGE and laser



- 50 y/o female
- CC: blurred painful eye, OD for 1 month treated with oral and topical steroid
- Received dental procedure about 2 months before onset of disease
- Systemic survey (-)



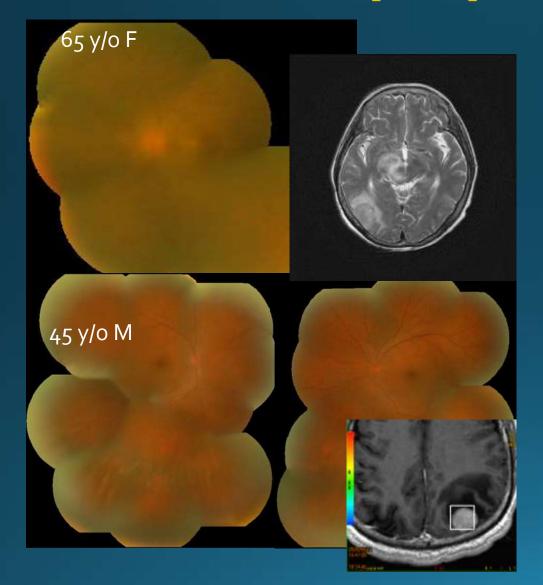
## Masquerade Syndromes

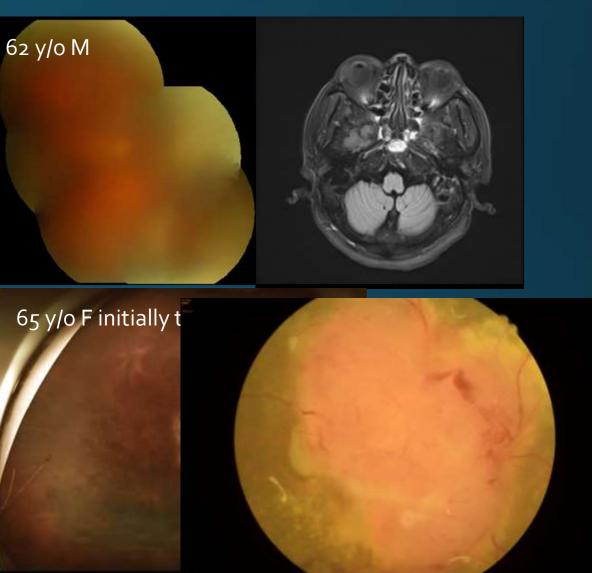
- One of the major indications for diagnostic vitrectomy
- Simulating a chronic idiopathic uveitis
- Having an underlying primary cause that is not immune mediated, but with an apparent clinical picture of intraocular inflammation
- They are usually poorly, if not at all, responsive to corticosteroid treatment.
- Classification: malignant/non-malignant

#### Diagnostic PPV for Intraocular Lymphoma

- Patients who have been under systemic or local corticosteroids are advised to discontinue the corticosteroids for a few weeks to allow more vitreous infiltration.
- Vitreous biopsy specimens need to be transported quickly for laboratory analysis
  - Lymphoma cells immediately undergo morphological degradation
- Cytokine analysis: IL10:IL6 ratio>1 suggestive of intraocular lymphoma

# Intraocular Lymphoma

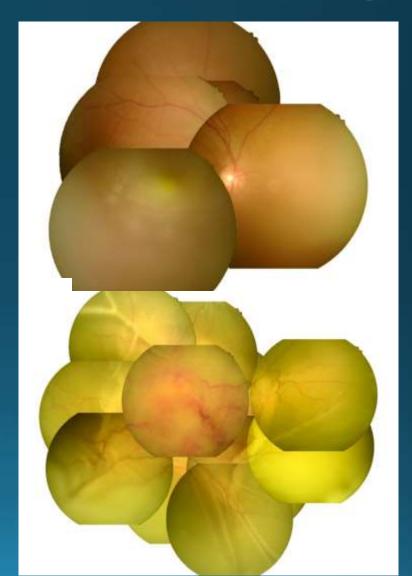




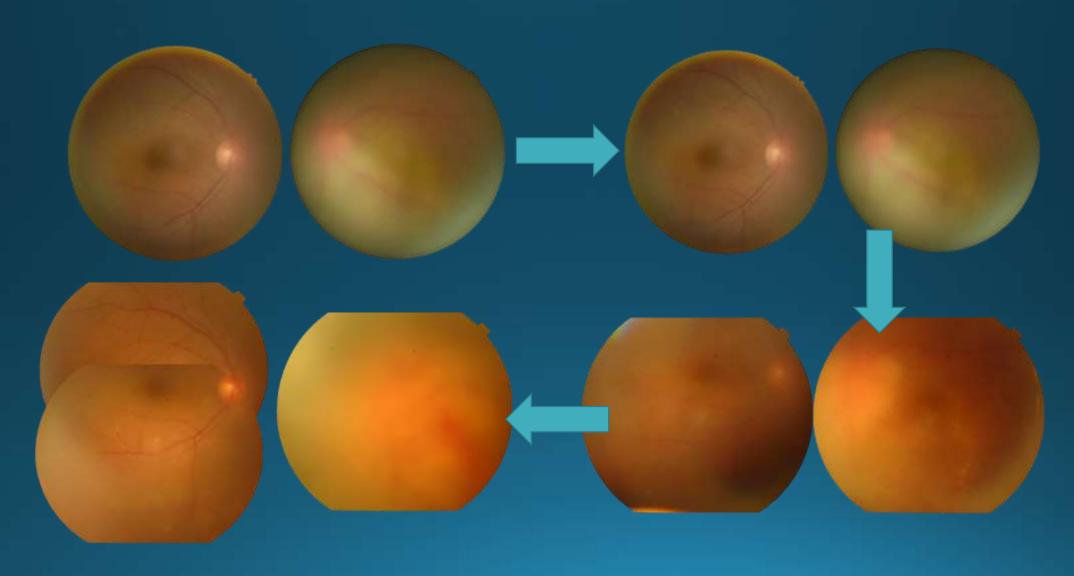
## 62 y/o female, blurred OS->OD ½ yr

- DM(+), HT(-), cataract op, OU
- Ever treated as panuveitis by other center but in vain, even progressed
- Diagnostic PPV OS 20210324: culture (-), malignant cell (-)
- 20210412 (first visit)

Va o.6/SL-, AC cell+-, vitreous cell++ OU

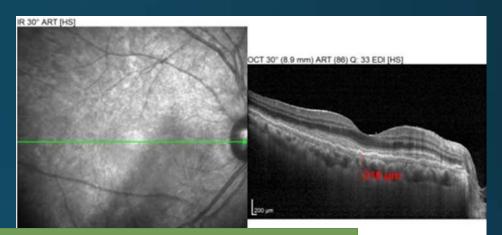


HLA B27+, HSV/VZV/CMV IgG+, vitreous sampleprednisolone, ciclosporin, mycophenolate, valacyclovir

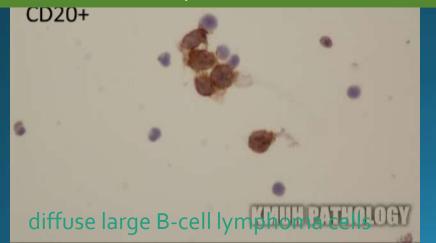


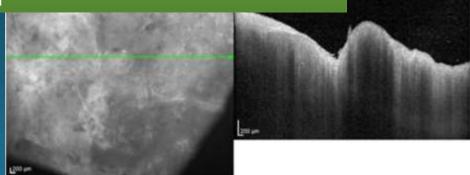
## Vitreous aspiration for cytology OS





Repeated aspiration cytology should be tried in highly suspected cases even when the vitreous sample from PPV did not get positive results.



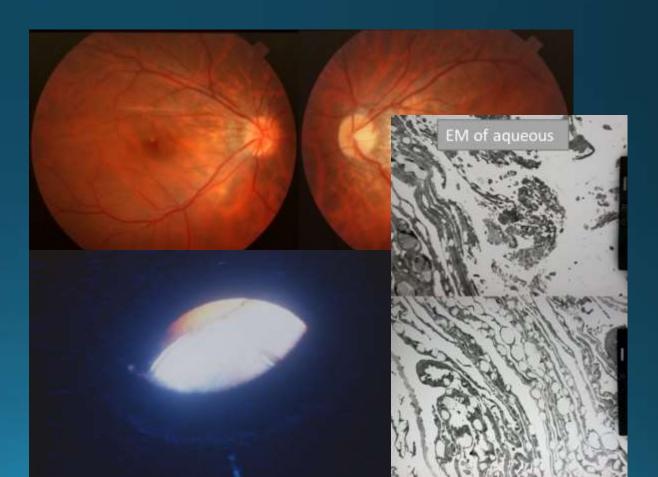


# Schwartz Matsuo Syndrome

13 y/o boy atopia, panuveitis with RD

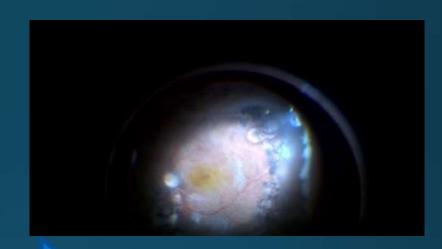
17 y/o M atopia, panuveitis with RD



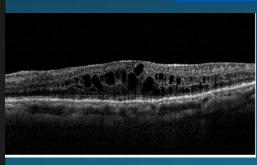


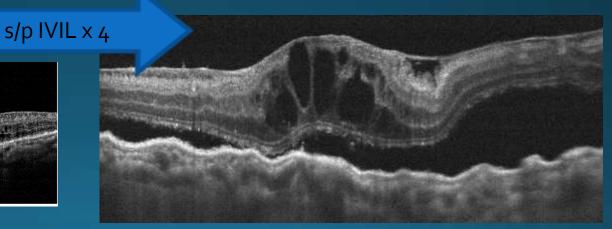
# Chronic RD Masquerading as Uveitis

- 63 y/o F
- ME developed 8 yrs after phacovitrectomy for PDR & TRD
- Panuveitis with shallow RD









### **Preoperative Inflammation Control**

- Previous experience suggests aggressive control of perioperative inflammation using topical, regional, and systemic steroids with or without systemic steroid-sparing immunomodulatory therapy for at least 3 months
- Recent studies showed that PPV is effective even if complete control of intraocular inflammation is not achieved prior to the surgery
- It depends on how urgent of the PPV is.

#### Reduce Inflammation

- PPV may decrease inflammation in eyes with noninfectious uveitis by presumably removing the inflammatory cytokines
- PPV and lensectomy are used for control of inflammation in lensassociated uveitis
- PPV may be an alternative to immunomodulatory therapy, where
   44% can be stopped after PPV
- PPV can reduce recurrence of intermediate, posterior and panuveitis

## Post-operative Endophthalmitis

- The most common indication for PPV to be done in an inflammed eye.
- Modern MIVS in exogenous endophthalmitis with much better initial vision than light perception is regarded safe and with low rate of complications
- The timing and indication of MIVS PPV should be earlier than those suggested by the EVS study.
- Vitrectomy can be considered once the clinical presentation progresses after standard intravitreal antimicrobial injections or for those with dense vitritis or fulminant appearance regardless of initial vision

#### Acute Post-operative Endophthalmitis

- 90 y/o male
- DM+
- Acute visual loss and pain, OD
   6 days after cataract surgery
- Has been on topical and subconjunctival antibiotics for one week
- Va LP

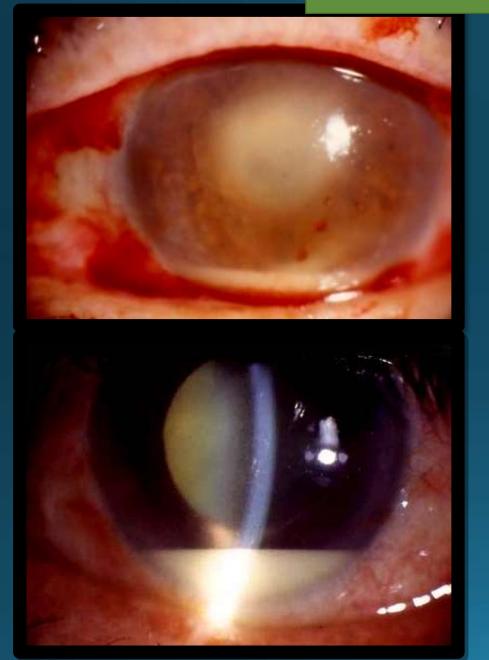




## Endogenous Endophthalmitis

- Systemic symptoms are usually non-specific and easily missed causing a delayed diagnosis leading to possibly devastating outcomes.
- Klebsiella pneumoniae (KP) EBE most common cause of endogenous endophthalmitis in Asia, including Taiwan.
- Diabetes was significantly associated with the endophthalmitis (Sheu, 2011)
- 8/42 (19%) had ocular symptoms before diagnosis of liver abscess (Sheu 2011).

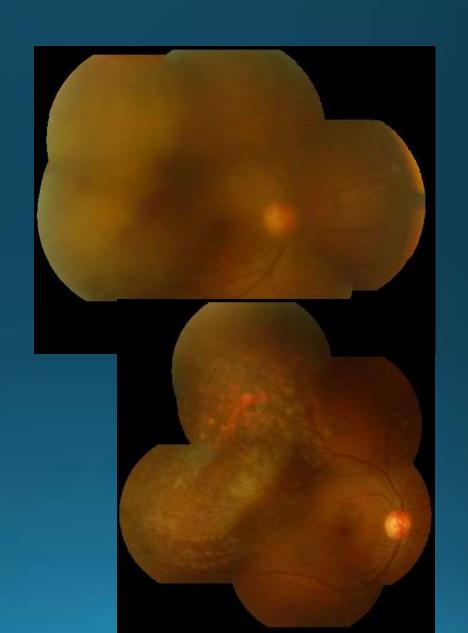
#### Different Presentations of KP EBE





## 35 y/o M

- Blurred OD for one month
- Liver chirossis, HBV, alchoholism, multiple site KP infection
- 6/20, AC cell++
- Fd retinitis patch with retinal hemorrhage and vitreous opacity
- PPV+ pc+ IVI with ceftazidime
   2.25mg/0.1cc
- Va at last follow up (1/2 yr) –
   6/7.5



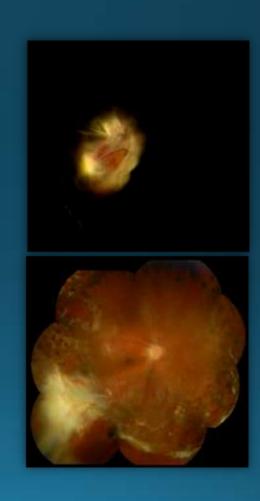
## 63 y/o M

- Hypertension+, CAD+, DM-Endocarditis, KP liver abscess
- Diagnosed and treated at 90 days after onset of KP PLA
- IVI with ceftazidime 2.25mg and vancomycin 1mg
- Phacovitrectomy then 2nd IOL, OS
- VA: HM → 4/60



## 48 y/o M plastic surgeon

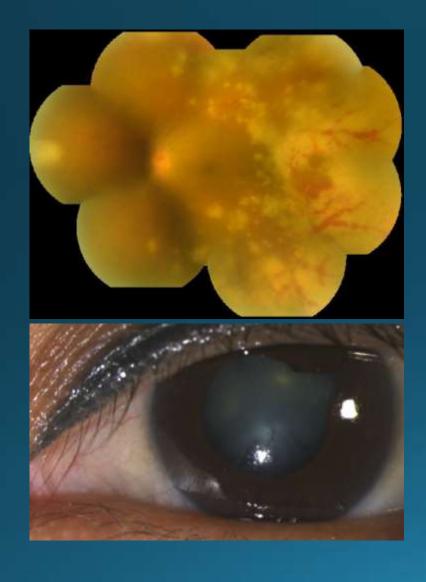
- Initially treated as uveitis with steroid then, IVI & VT/cat op for complicated RD and cataract due to KP endophthalmitis at other hospital
- Presenting as panuveitis with small fixed pupil and PVR
- 3rd op: vitrectomy + relaxing retinotomy + silicon oil tamponade at 67 days after onset

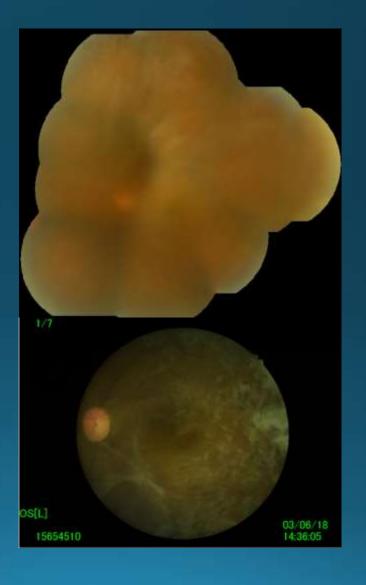


#### **Retinal Detachment**

- Uveitis is an independent risk factor for RRD
- Panuveitis and infectious uveitis are the most common entities
- Proposed maneuver to prevent RD in ARN (with conflicting evidence)
  - Prophylactic laser photocoagulation
  - Early vitrectomy
- RD after ARN: High risk of PVR, multiple holes in an atrophic thinned retina and the presence of ischemic optic neuropathy.
- MIVS PPV with silicon oil tamponade is the treatment of choice.

#### 37 y/o F, ARN followed by RRD with PVR

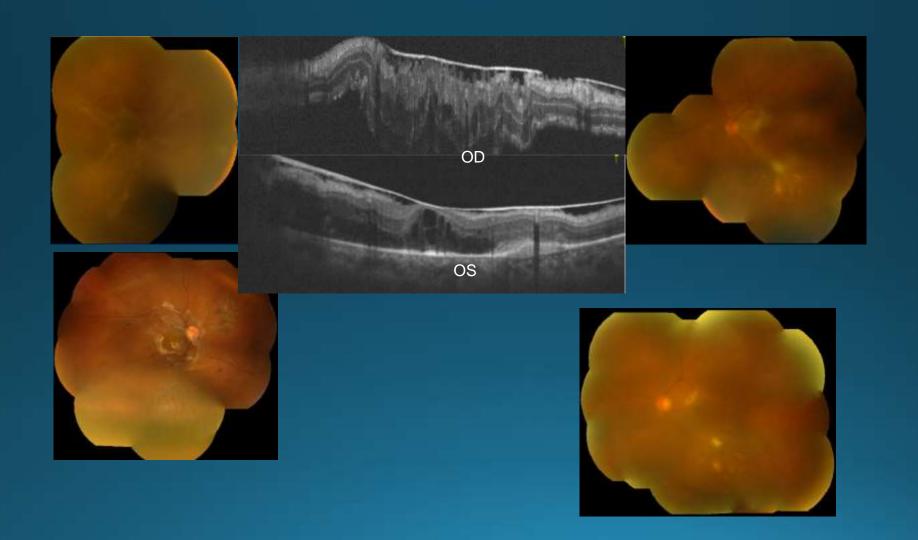


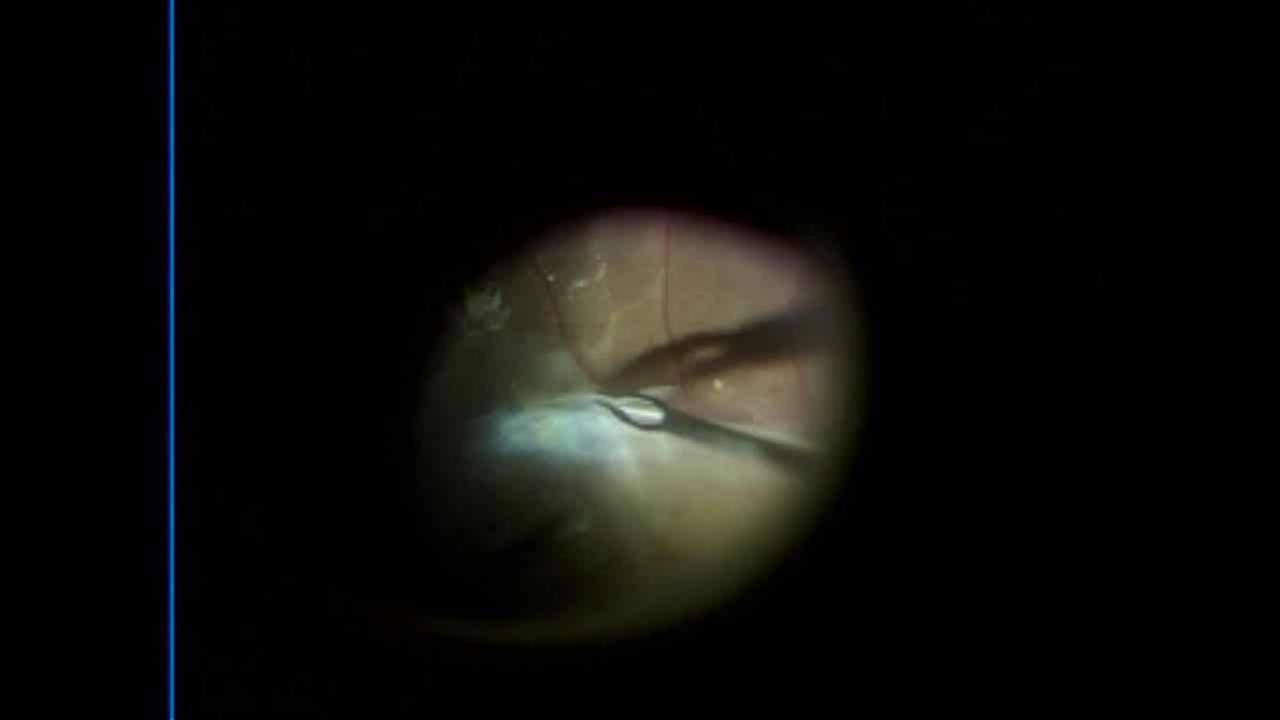


#### Epiretinal Membrane (ERM) in Uveitis

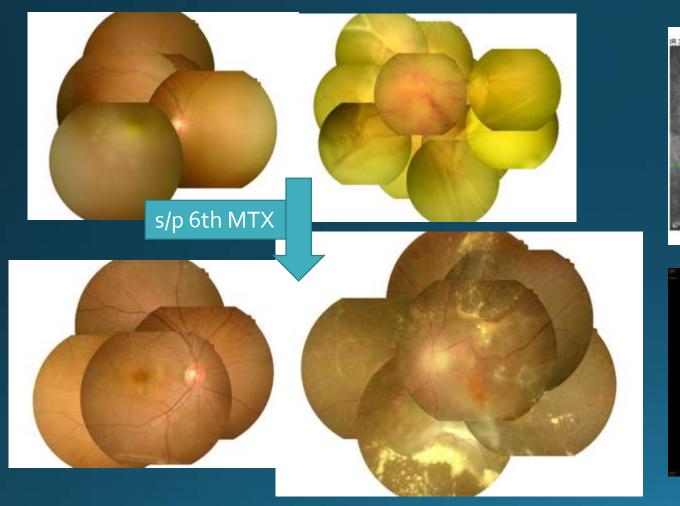
- ERM represents a common complication of uveitis
- ERM 2nd to uveitis is pathogenically distinct from those that are idiopathic.
- PPV is indicated in ERM with sight-threatening macular edema or if progression occurs despite maximal medical treatment.
- Tangential traction is usually stronger in uveitis-related ERM.

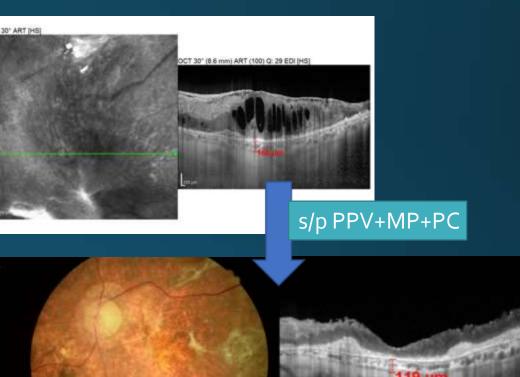
#### Macular Pucker following CMV Retinitis





# PVR in Intraocular Lymphoma







#### Complications of PPV in Uveitic Eyes

- Overall complication rate: 42-54% hypotony, VH, RD, ERM, cataract or worsen inflammation (Bansal 2017; Oahalou 2014)
- Generally more risk in those associated with systemic risk factors.
- We must weigh the benefit and risk of the surgery for the patients.

#### Conclusions



- VR surgeries provide more choices in the management of uveitis.
- MIVS makes the procedure easier, harbors less complications and more patients' comfort.
- Achieving optimum control of inflammation before PPV is recommended for non-emergent therapeutic indications.
- Recent indications are extended to include some active inflammation for diagnostic vitreous sampling or medically unresponsive inflammation.