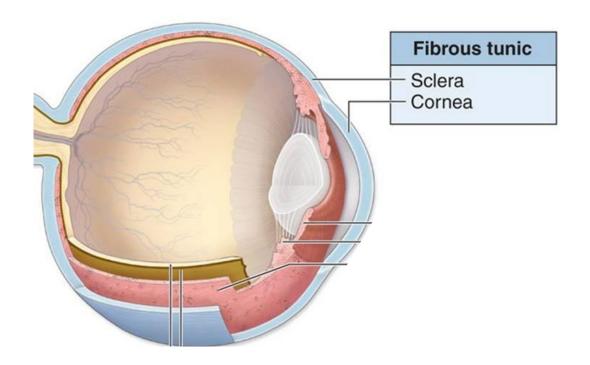
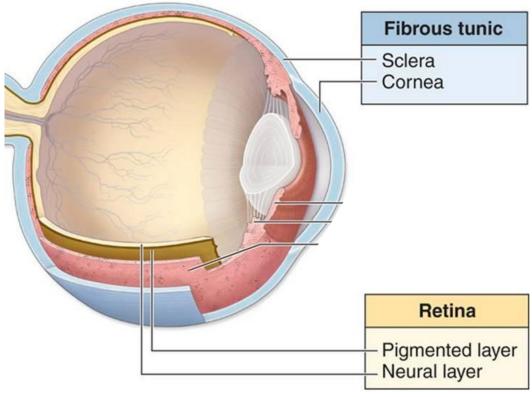
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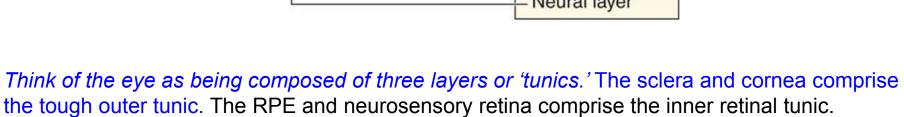




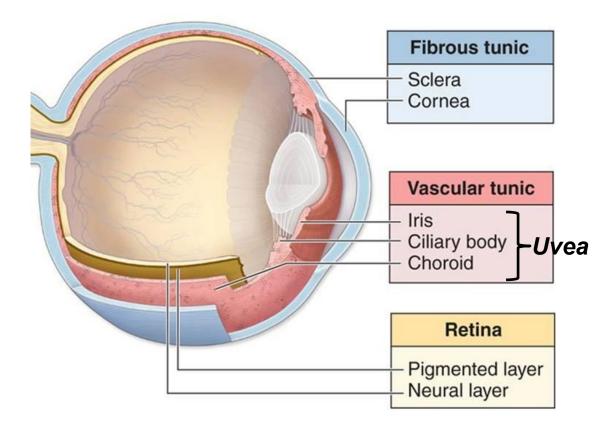


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3



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Uveal tissue. Note its deep purple hue

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The location of the uveitis

- --Anterior
- --Intermediate
- -Posterior
- --Panuveitis

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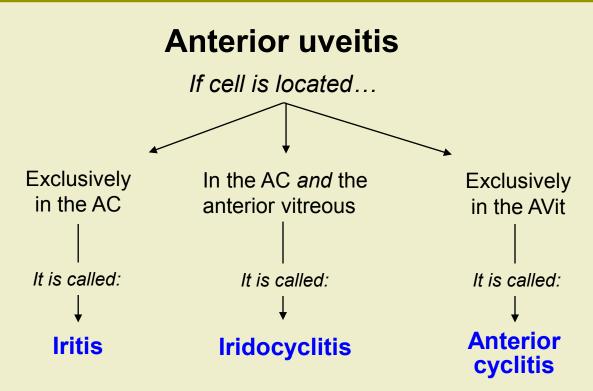
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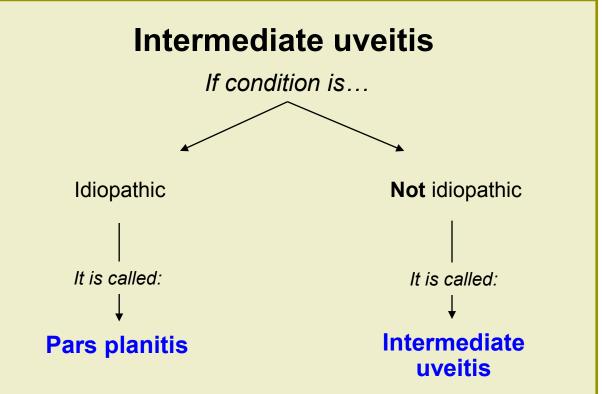
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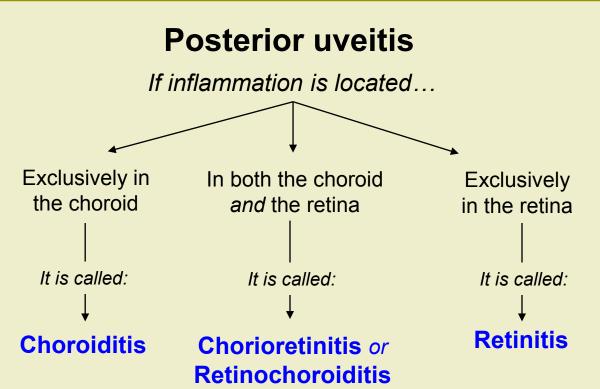
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In posterior uveitis, the site of inflammation is the retina and/or choroid (the optic nerve head can be involved too)

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#### In panuveitis, all three locations are equally involved

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- 3) A differential diagnosis list is generated
- 4) Studies are obtained to identify the etiology



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- 1) The uveitis is profiled
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- 3) A differential diagnosis list is generated
- 4) Studies are obtained to identify the etiology
- 5) Treatment appropriate for the etiology is initiated



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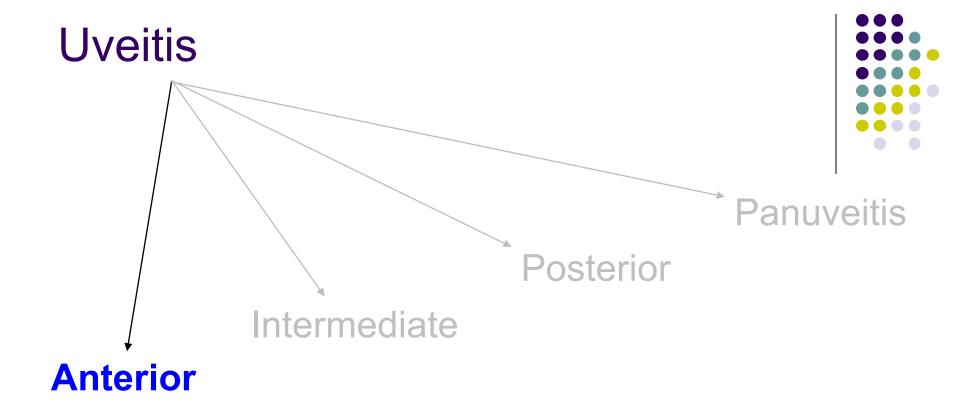
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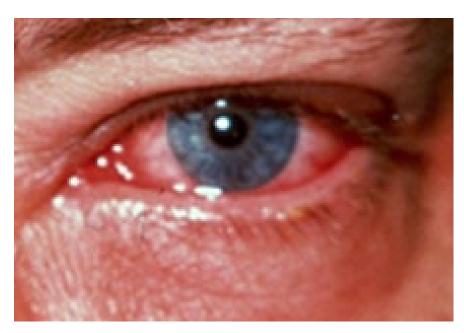


Let's drill down on anterior uveitis. Specifically, let's look at how the BCSC organizes it by presentation



Anterior uveitis is by far the most common form encountered clinically. The classic symptoms are **pain** and **photophobia**, along with some degree of **reduced vision**. Patients will also complain of **surface injection** (which presents often in a so-called 'ciliary flush' pattern).





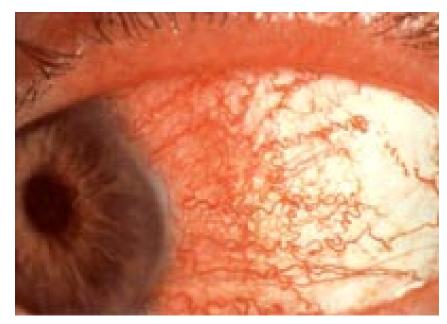
Injection in conjunctivitis

In surface disorders (eg, conjunctivitis), redness is either distributed uniformly across the eye, or it tapers off near the limbus.





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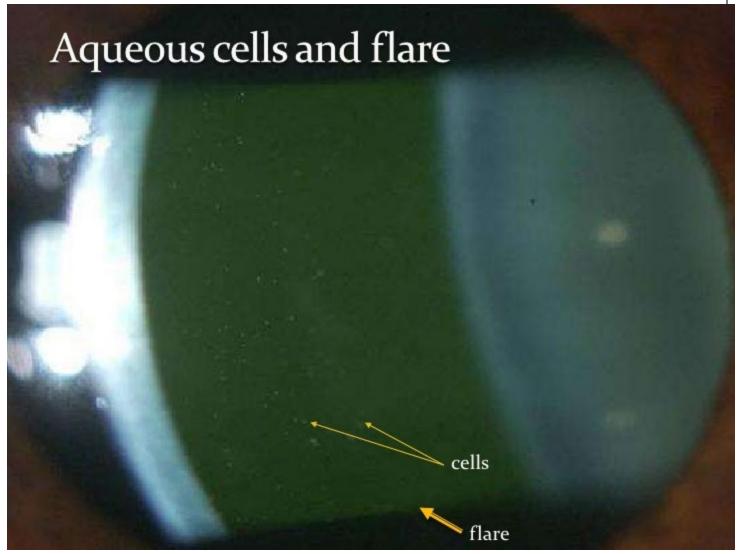


Injection in anterior uveitis, aka ciliary flush

In surface disorders (eg, conjunctivitis), redness is either distributed uniformly across the eye, or it tapers off near the limbus. In contrast, redness associated with anterior uveitis is usually most intense at and just behind the limbus. This is because this area overlies the inflamed ciliary body—hence the term *ciliary flush* for this presentation.



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# Uveitis Anterior Uveitis Granulomatous Nongranulomatous



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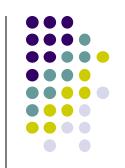
The *Uveitis* book employs an organizational tree on which it hangs the common causes of anterior uveitis. The first branch point in this tree is whether the inflammation is **granulomatous** or **nongranulomatous**. In clinical practice the term *granulomatous* refers to a particular slit-lamp appearance of KP—large, grayish, and 'greasy.'





Granulomatous KP

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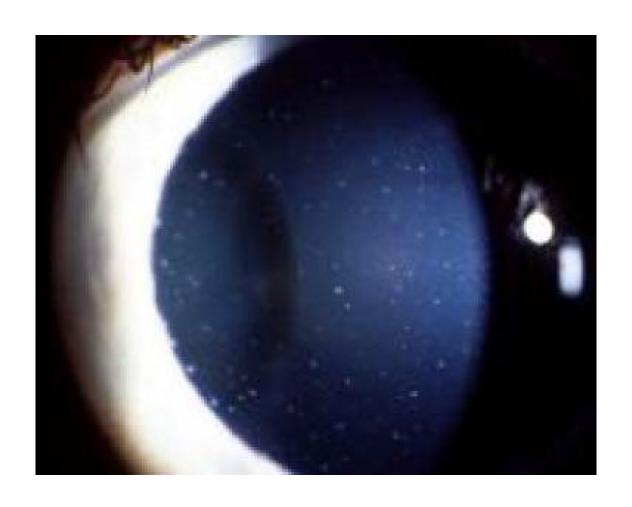


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In contrast, *nongranulomatous KP* are smaller, lighter in color, and do not look greasy. (Note: If no KP are present, the inflammation is considered nongranulomatous.)





Nongranulomatous KP

#### **Anterior Uveitis**

#### Granulomatous

TB

Sarcoid

**Syphilis** 

HSV

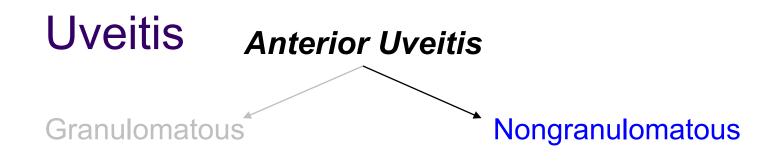
VKH

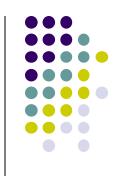
Toxoplasmosis

Lyme

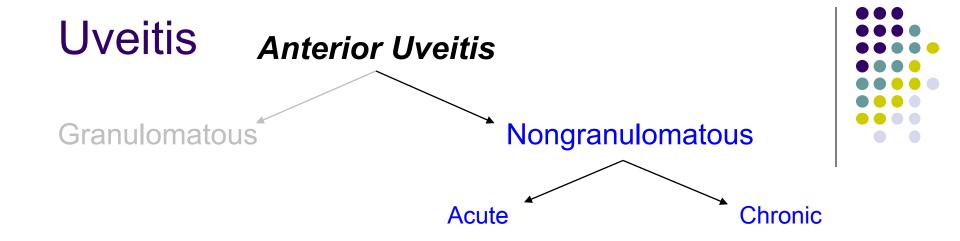
#### Nongranulomatous

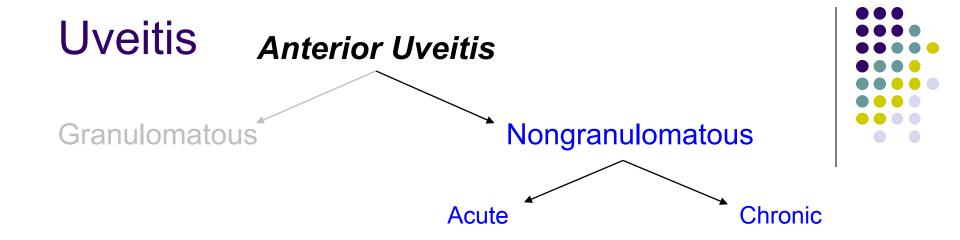
These are the common entities that can produce a granulomatous anterior uveitis. (Note: For some of these, the granulomatous anterior findings are part of an overall panuveitic presentation, ie, they typically do not present as an *isolated* anterior uveitis.)



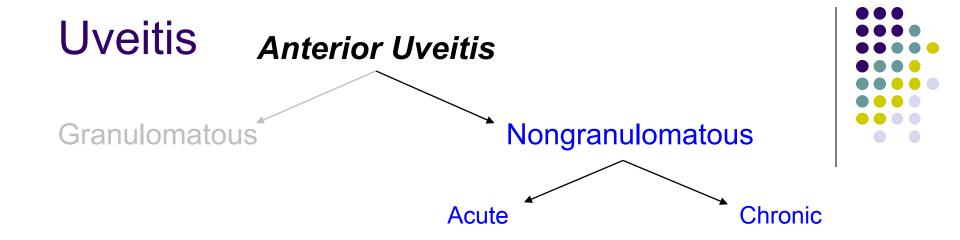


The rest of the anterior-uveitis classification tree concerns **nongranulomatous** dz.

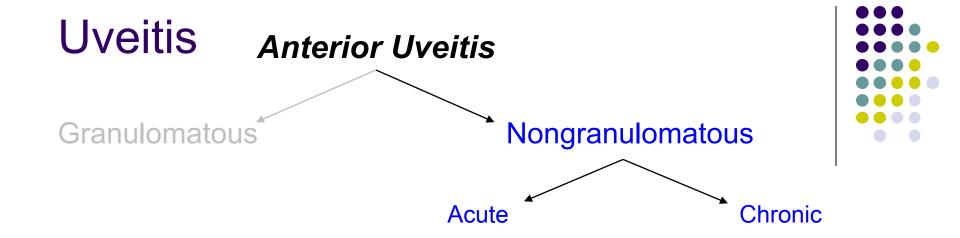




Acute uveitis comes on suddenly and resolves fairly quickly.

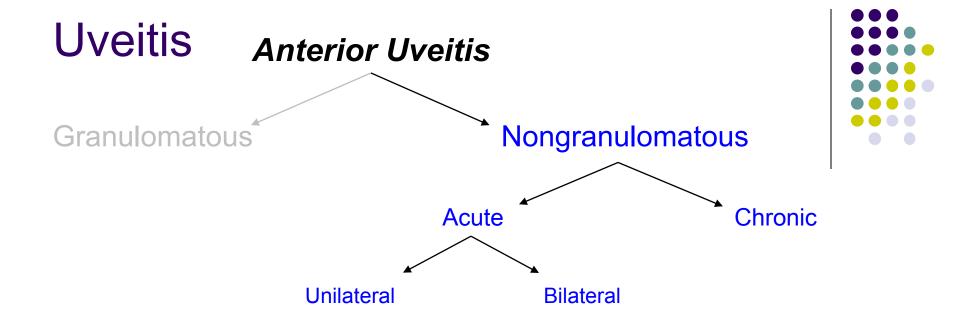


Acute uveitis comes on suddenly and resolves fairly quickly. Chronic uveitis also resolves, but once treatment is withdrawn, it relapses within three months.

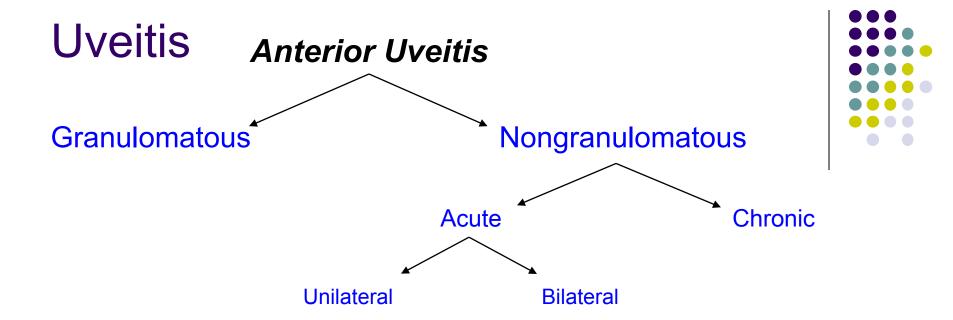


Acute uveitis comes on suddenly and resolves fairly quickly. Chronic uveitis also resolves, but once treatment is withdrawn, it relapses within three months.

(FYI: If a uveitis eventually relapses but is quiescent off-treatment for **longer** than three months, it is termed a *recurrent* uveitis.)

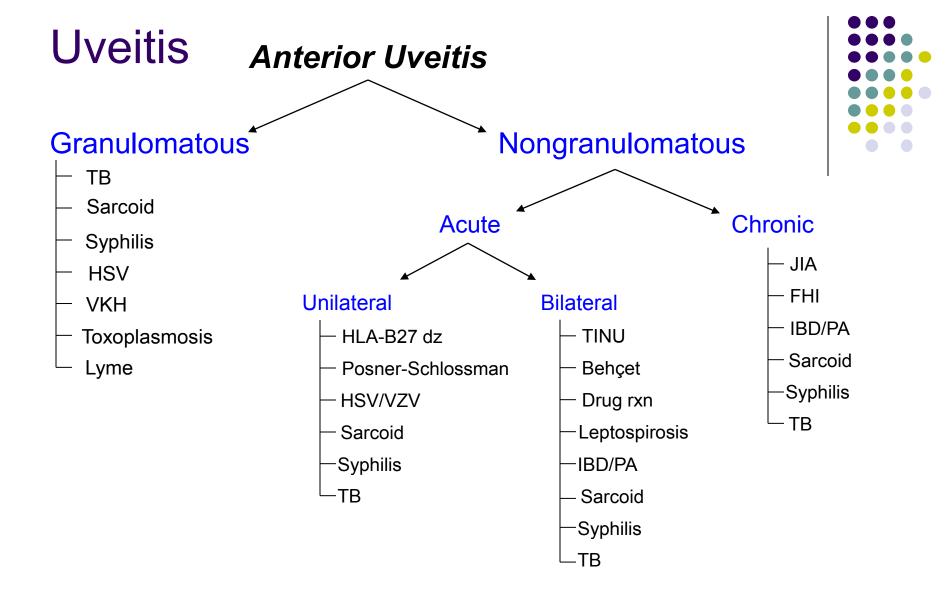


Finally, the acute uveitides are divided into those that present **unilaterally** vs those that tend to present **bilaterally**.

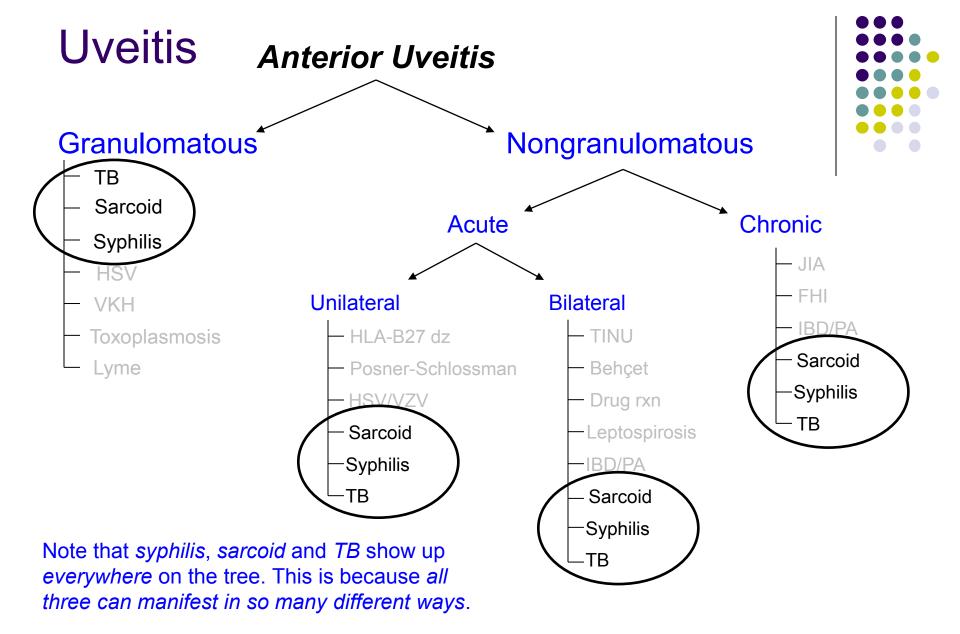


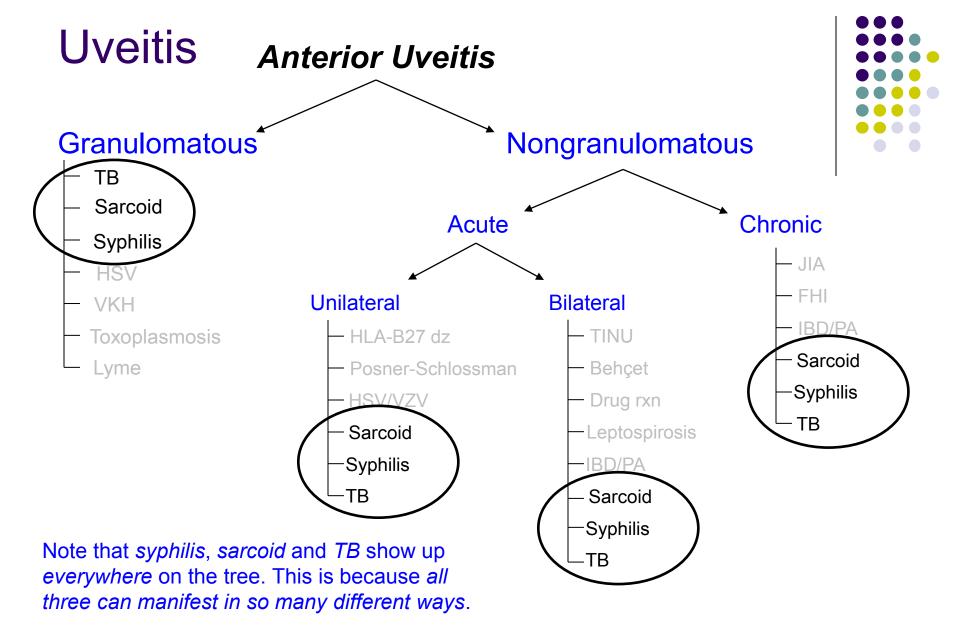
The rest of the anterior-uveitis classification tree concerns nongranulomatous dz. The first branch-point divides the etiologies into those that produce acute dz vs that a good look at this slide—it represents how you should think about anterior uveitides encountered in the clinic or on the OKAP.

[It wouldn't be a bad idea to commit this to memory at this juncture.]



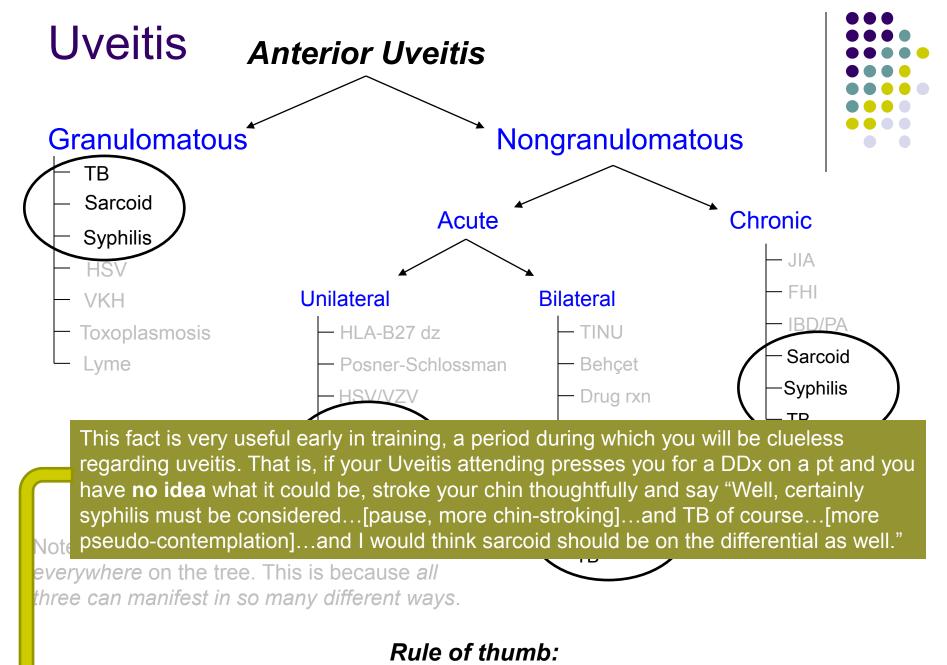
Just as an FYI, these are the anterior uveitides that are covered in detail in the *Uveitis* book. **Don't** *try to memorize all this at this juncture!* (They will stick better if you learn them in their naturally-occurring groupings.)



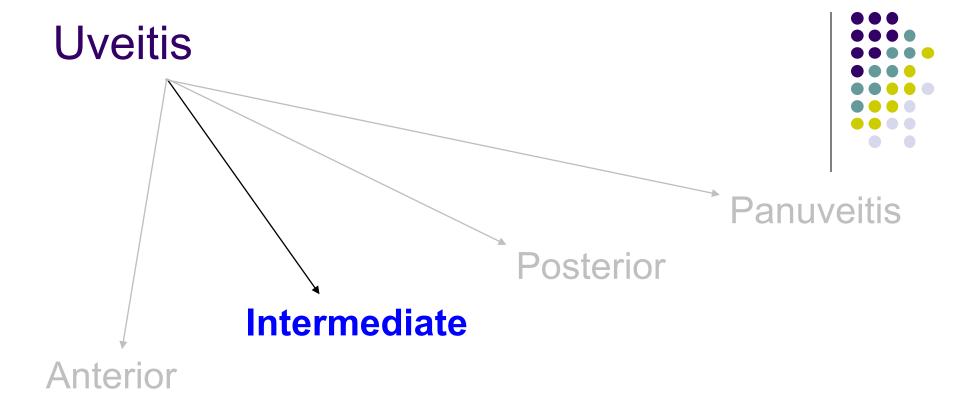


#### Rule of thumb:

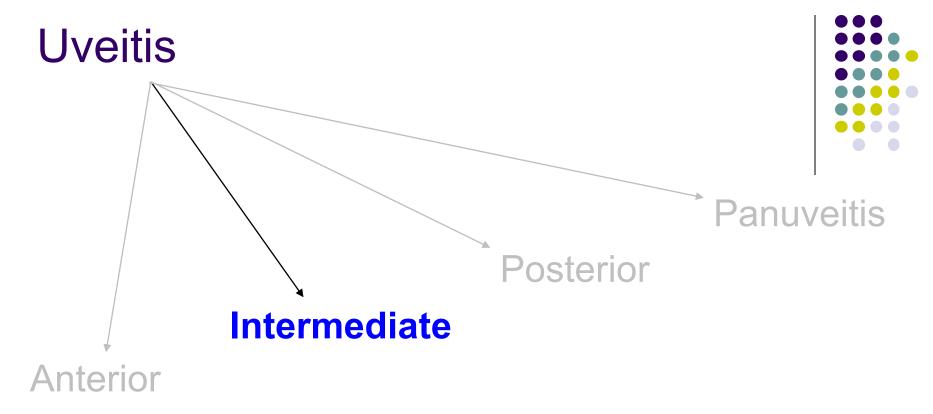
Syphilis, sarcoid and TB are on the DDx for every pt with any form of uveitis!



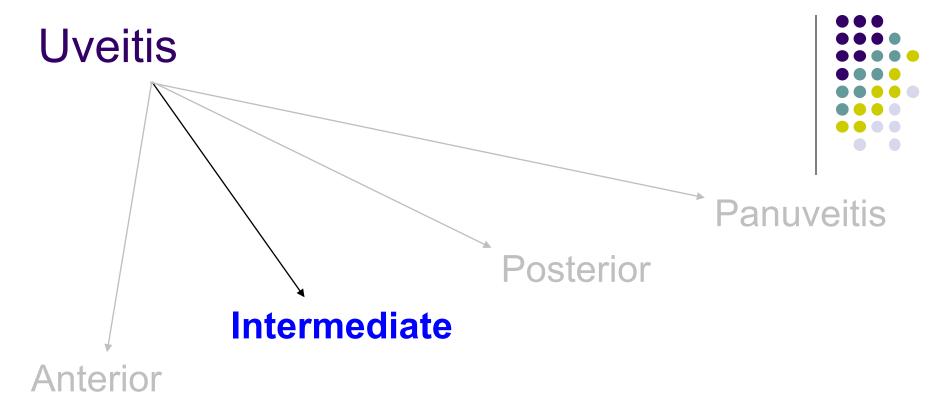
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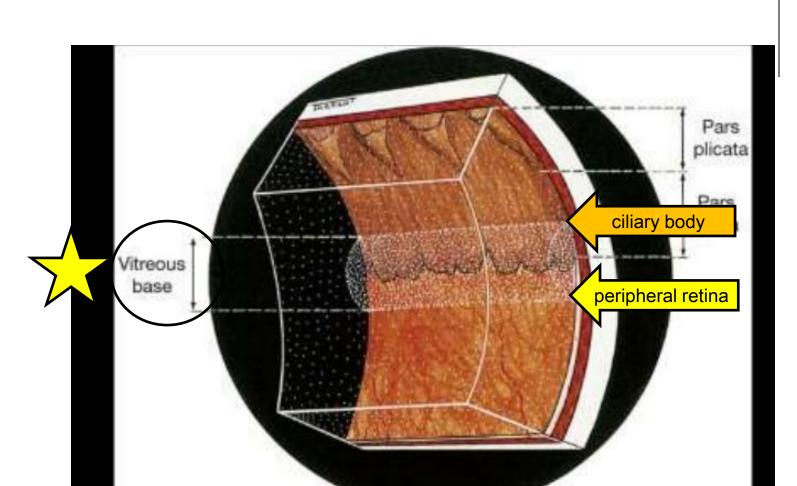
Next let's look at intermediate uveitis



The hallmark of **intermediate uveitis** (IU) is inflammation in the anterior vitreous that involves the vitreous base.

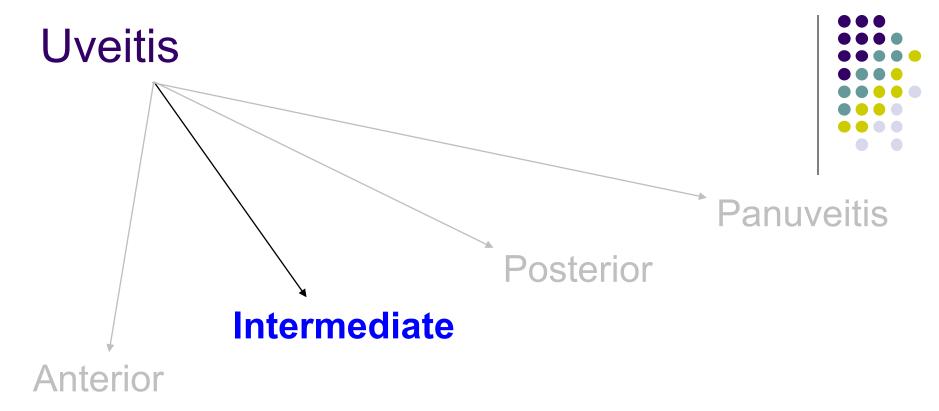


The hallmark of **intermediate uveitis** (IU) is inflammation in the anterior vitreous that involves the vitreous base. The vitreous base is the primary attachment point of the vitreous; it forms a ~5 mm-wide band that straddles the ora serrata (the location where the anteriormost retina meets the posteriormost portion of the ciliary body).

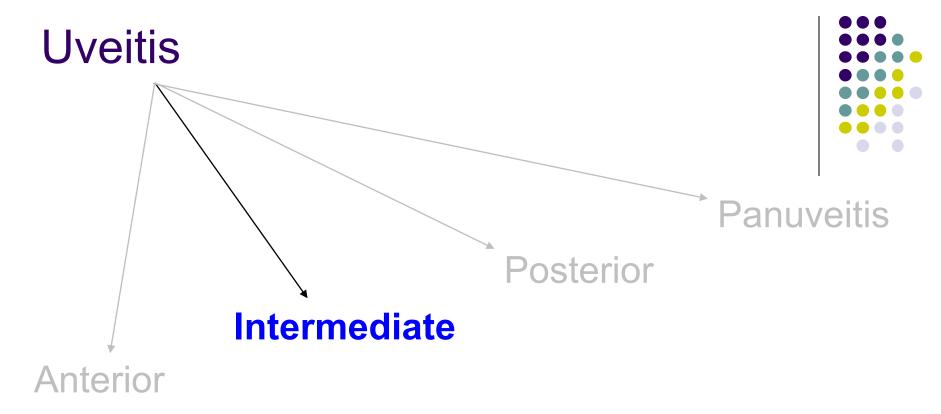


The vitreous base





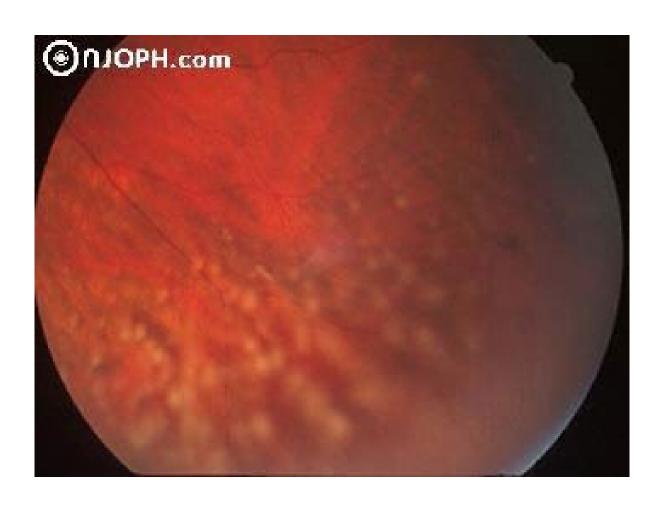
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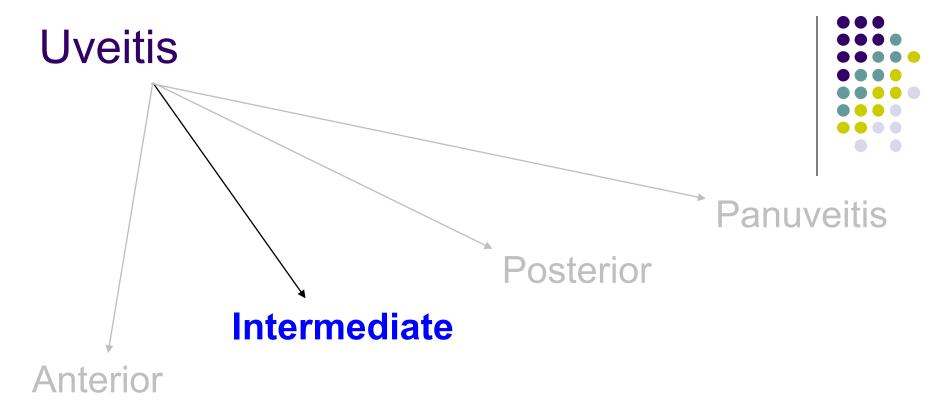
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Snowballs are a classic finding in IU. Snowballs—so named because of their appearance—are clumps of inflammatory cells and detritus in the vitreous.





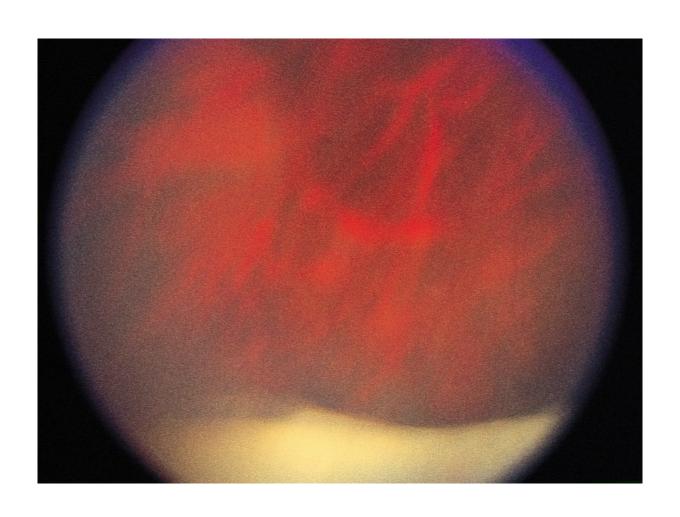
Snowballs in intermediate uveitis



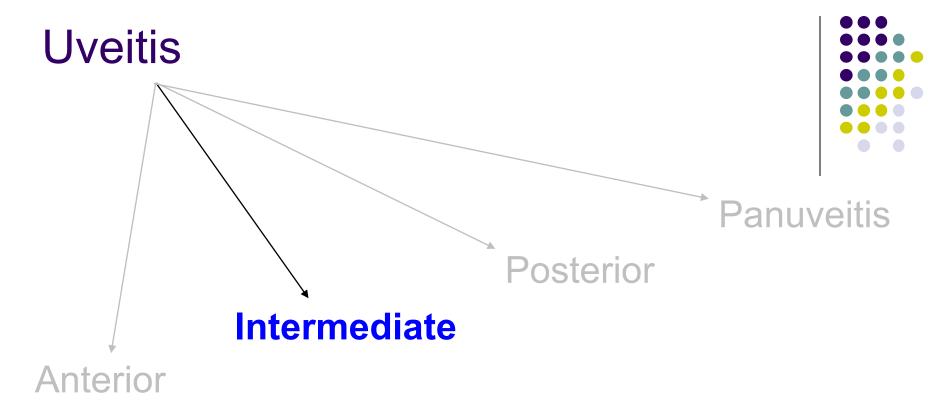
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Snowballs are a classic finding in IU. Snowballs—so named because of their appearance—are clumps of inflammatory cells and detritus in the vitreous. When this material accumulates in broad swaths along the inferior pars plana, it is referred to as *snowbanking*.



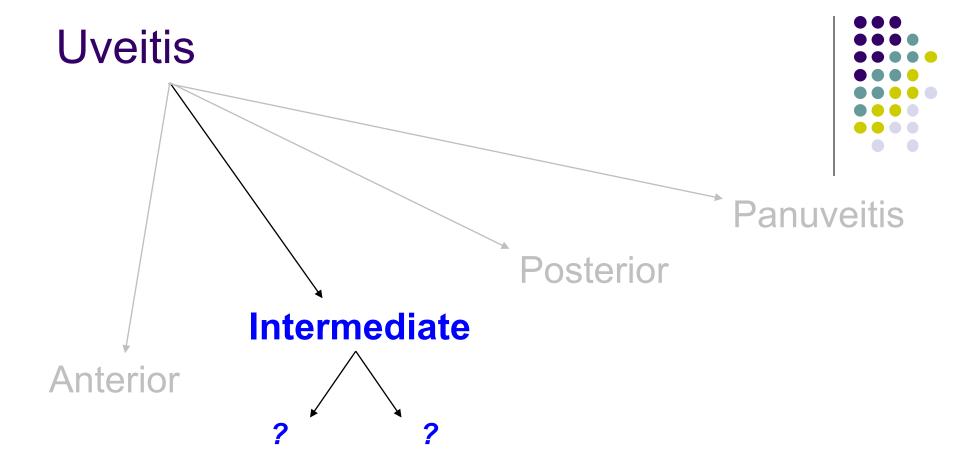


Snowbanking in intermediate uveitis

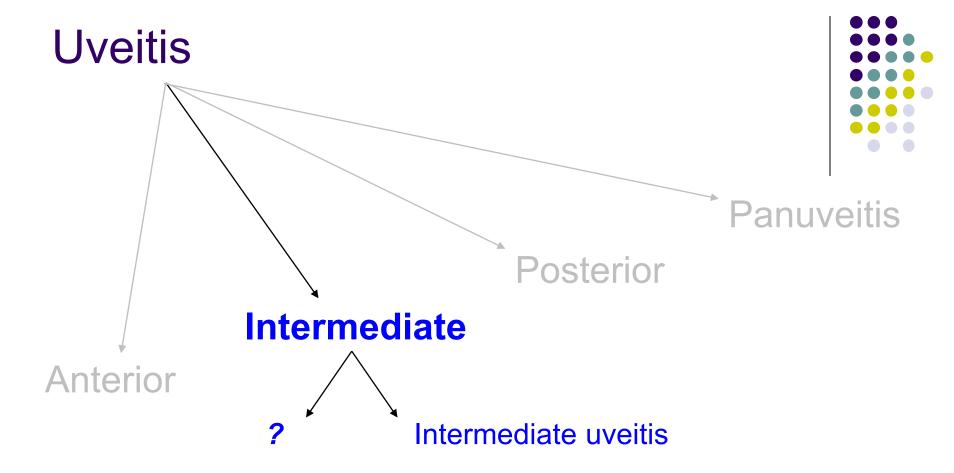


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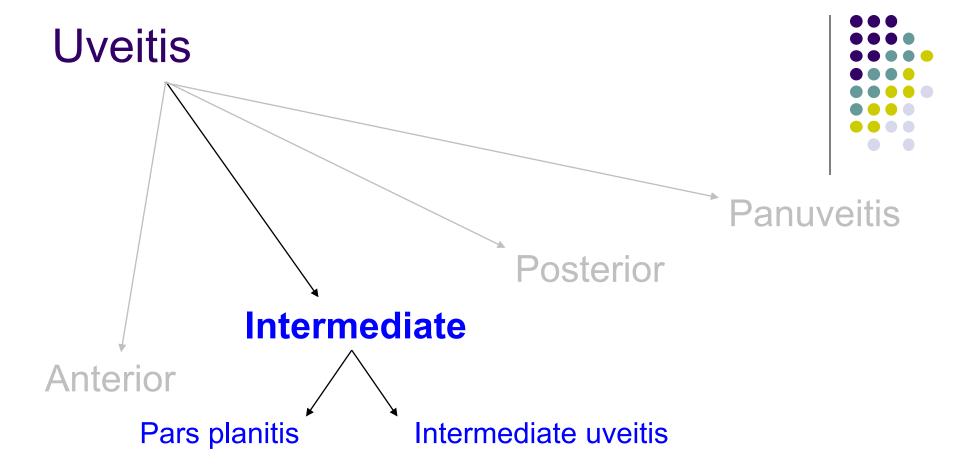
IU tends to be a dz of young people—teens through 40 or so. It is bilateral in most (80%) cases, although it can be quite asymmetric.



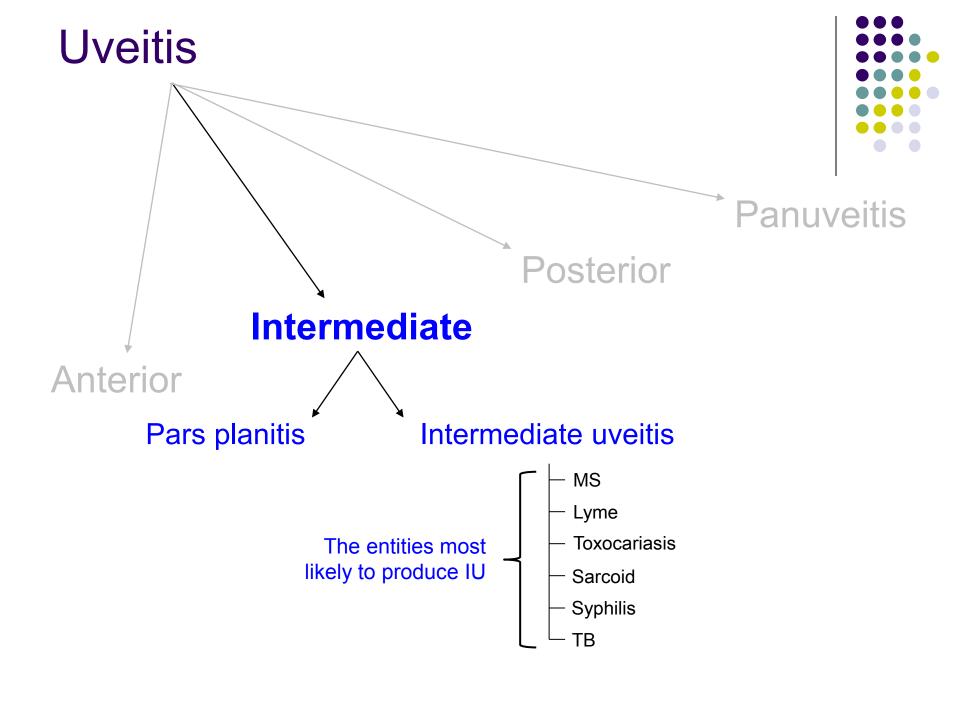
IU is divvied up into two categories.

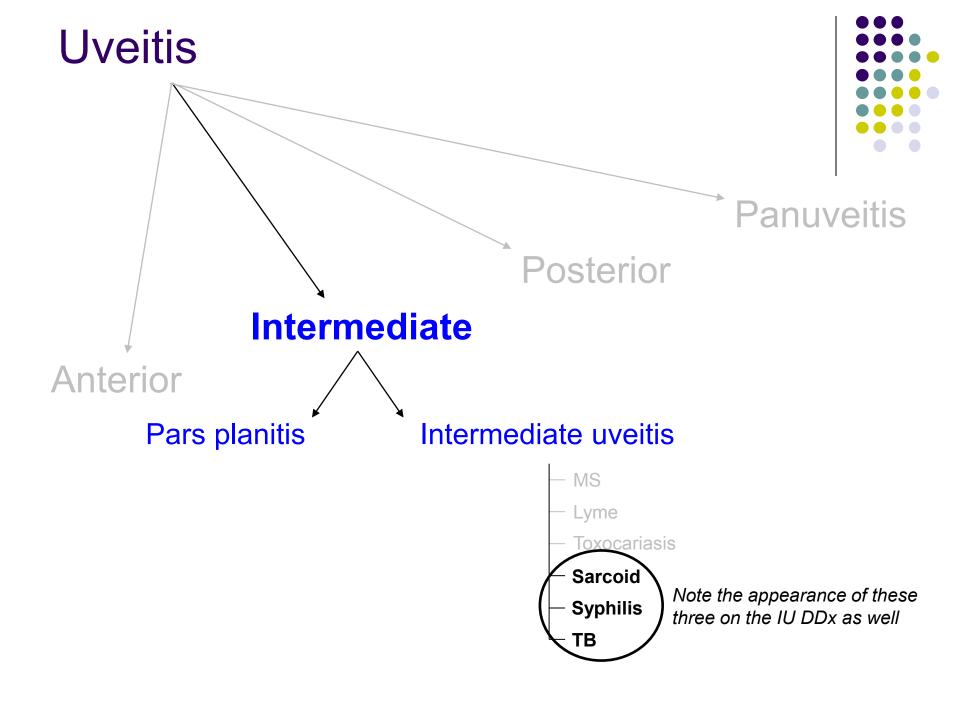


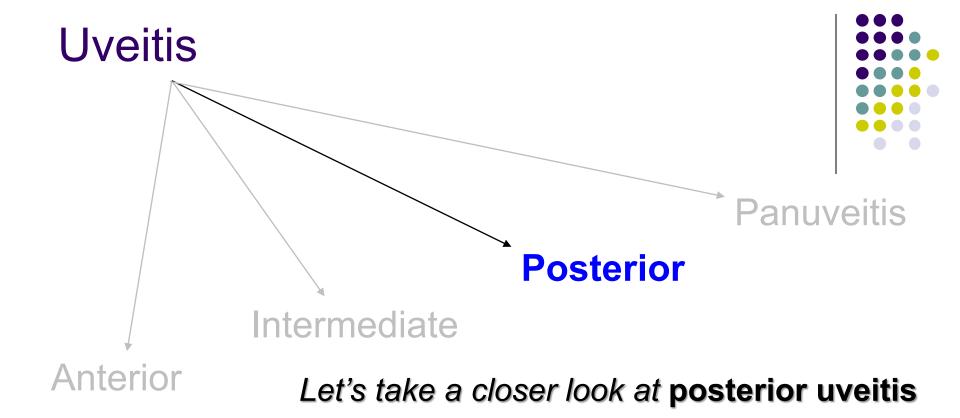
IU is divvied up into two categories. If the inflammation is associated with an identifiable condition, the uveitis is called **IU**.



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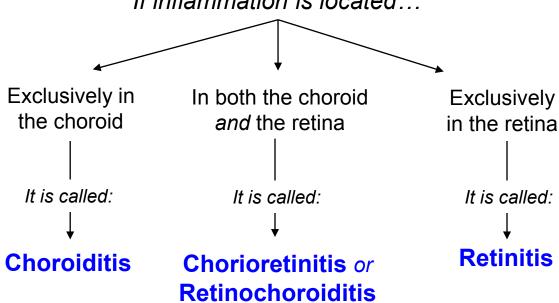






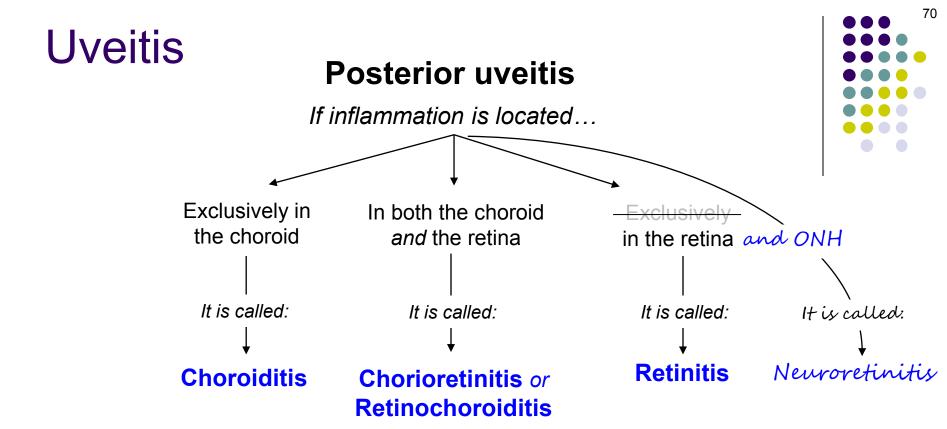
#### **Posterior uveitis**

If inflammation is located...

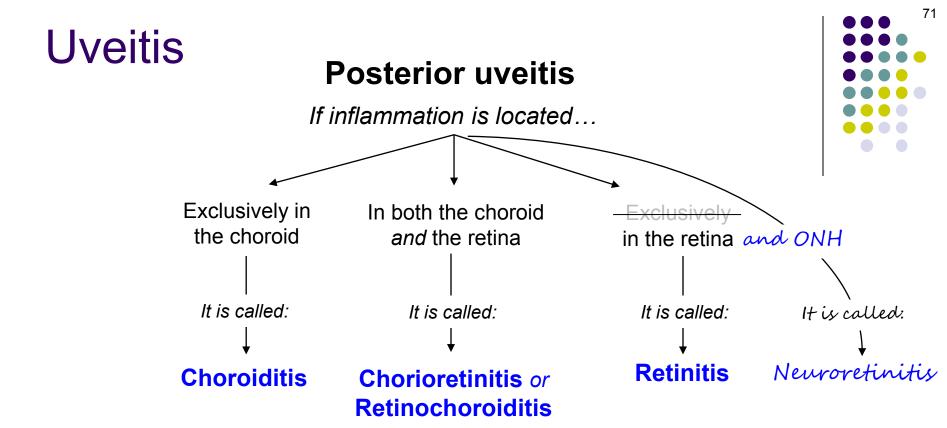


As presented previously, here are the ways **posterior uveitis** can manifest.



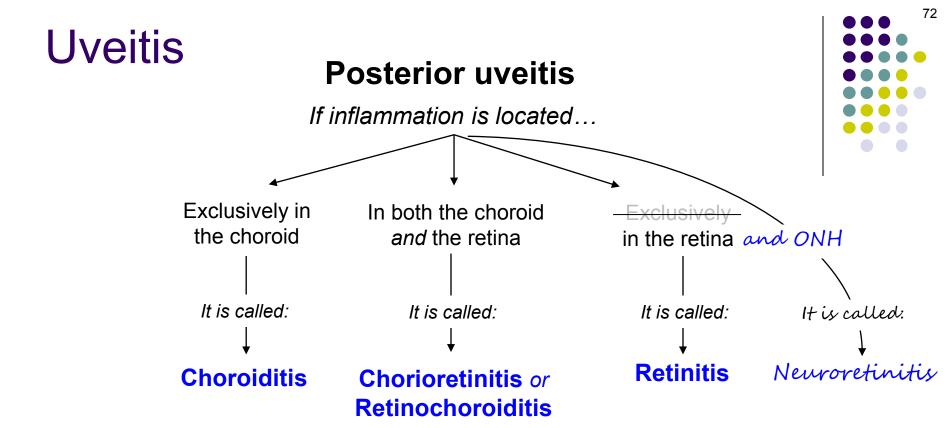


As presented previously, here are the ways **posterior uveitis** can manifest. One more—*neuroretinitis*, inflammation involving both the retina and optic nerve—should be added for completeness sake.

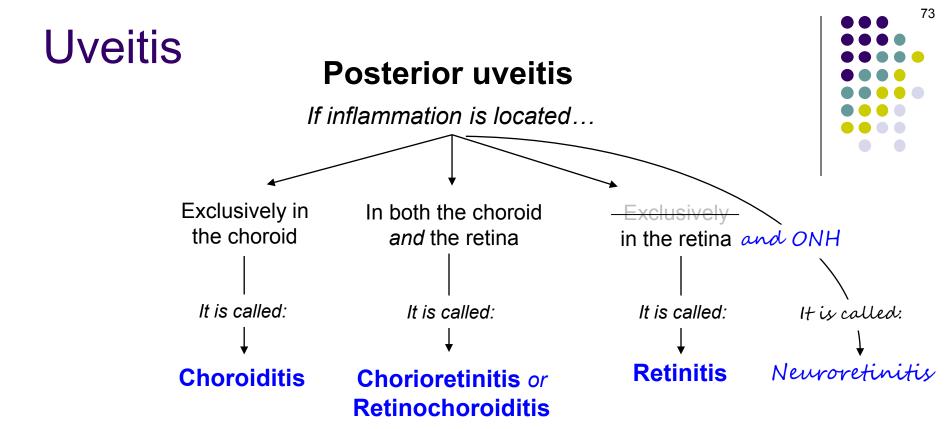


As presented previously, here are the ways **posterior uveitis** can manifest. One more—*neuroretinitis*, inflammation involving both the retina and optic nerve—should be added for completeness sake.

About 80% of **anterior** uveitis cases are **non**infectious in origin. *The opposite is true for posterior uveitis*: most cases are infectious—weirdly, also about 80%.



**Toxoplasmosis** is a common, classic cause of posterior uveitis. It is infectious, the bug being *Toxoplasma gondii*, an obligate intracellular parasite. The cat is its definitive host. *T gondii* has a worldwide distribution; an estimated one billion people are infected. Humans usually acquire the parasite via consumption of unwashed produce or undercooked meat.



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Oocyst form

Tachyzoite form

Tissue cyst

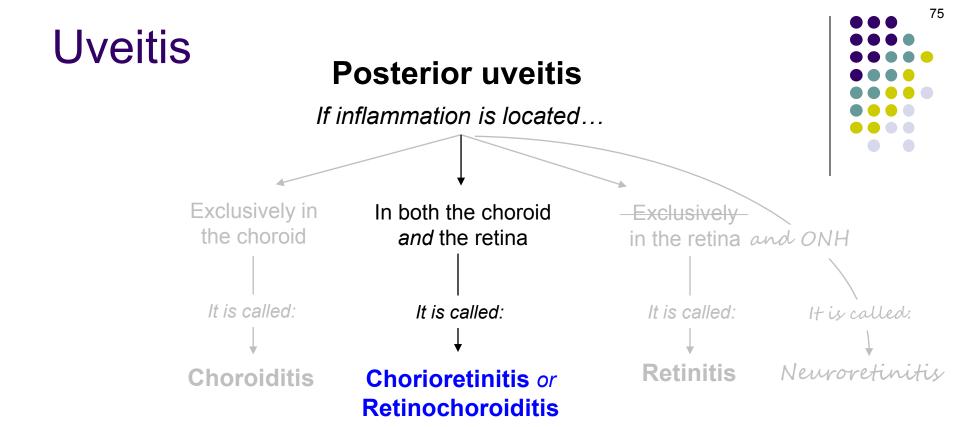
--Found in GI tract of cat (shed in feces)

--Acquired via ingestion of unwashed produce

--Found in circulatory system
of infected mother
--Responsible for
transplacental infection

--Found in tissue of infected livestock
--Acquired via consumption of
undercooked meat

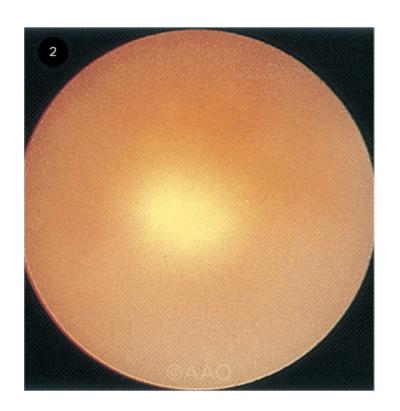
Toxoplasma gondii: Three infectious forms



**Toxoplasmosis** is a common, classic cause of posterior uveitis. It is infectious, the bug being *Toxoplasma gondii*, an obligate intracellular parasite. The cat is its definitive host. *T gondii* has a worldwide distribution; an estimated one billion people are infected. Humans usually acquire the parasite via consumption of unwashed produce or undercooked meat. Another crucial mechanism of transmission is transplacentally, which leads to devastating congenital manifestations in affected infants (it is one of the TORCH syndrome etiologies).

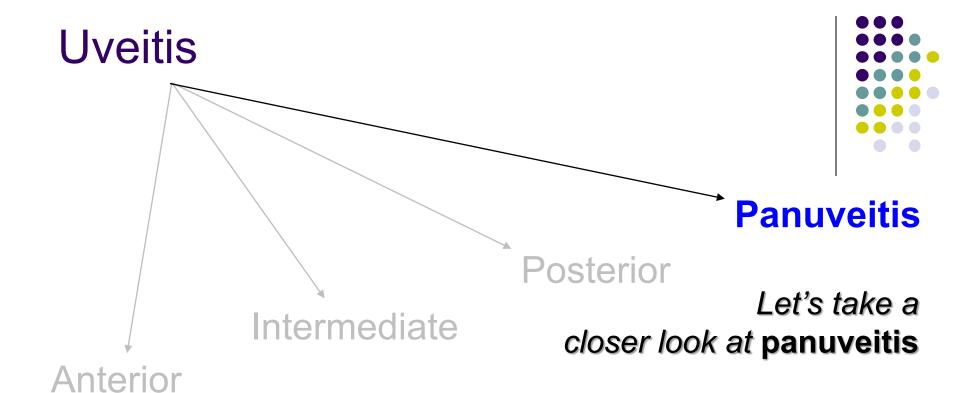
The typical posterior-uveitis manifestation of toxoplasmosis is a retinochoroiditis accompanied by a dense overlying vitritis. Taken together, the appearance has been likened to a 'headlight in the fog.'

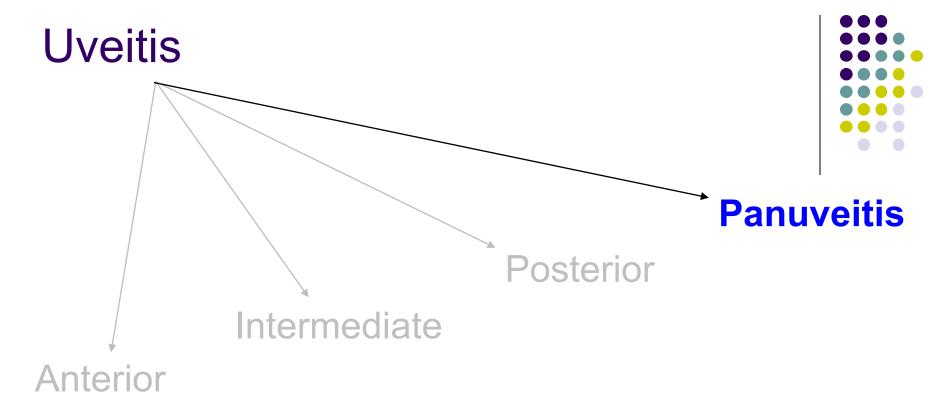


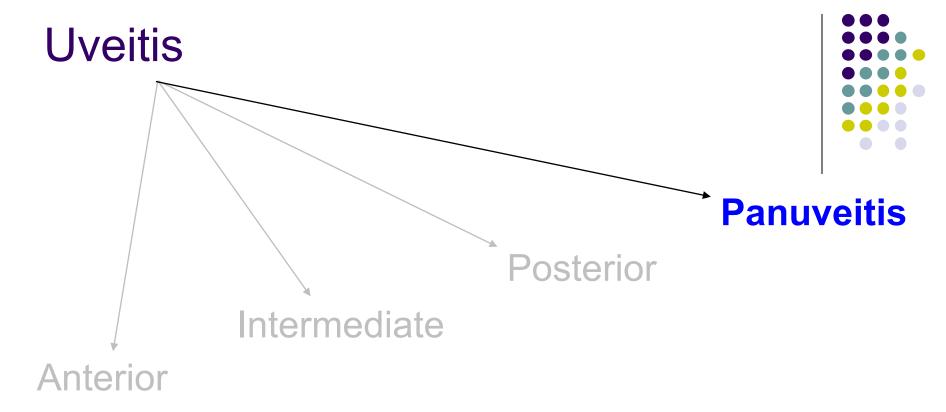


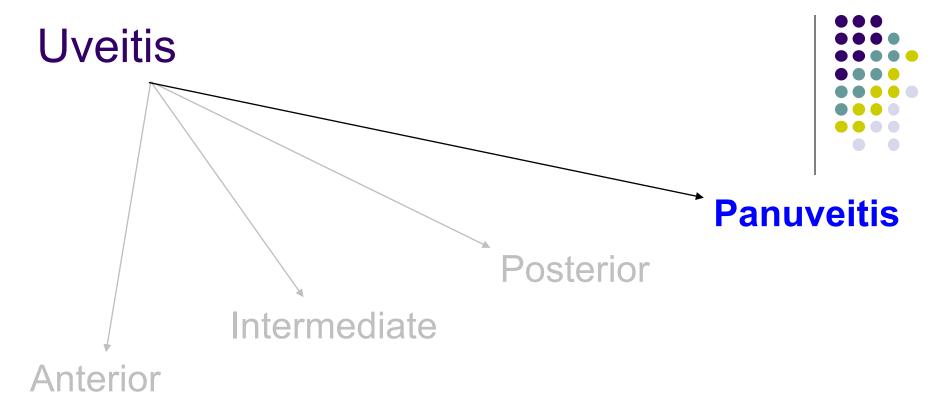


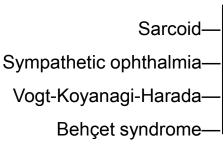
Ocular toxoplasmosis: Headlight in the fog

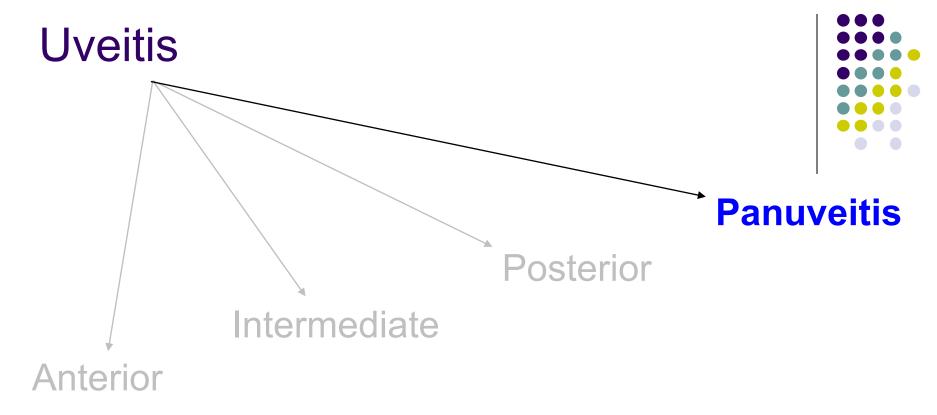


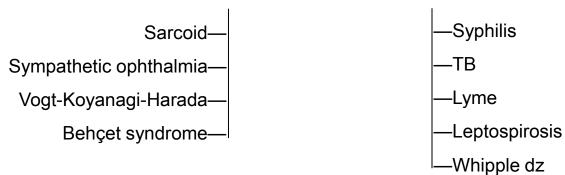


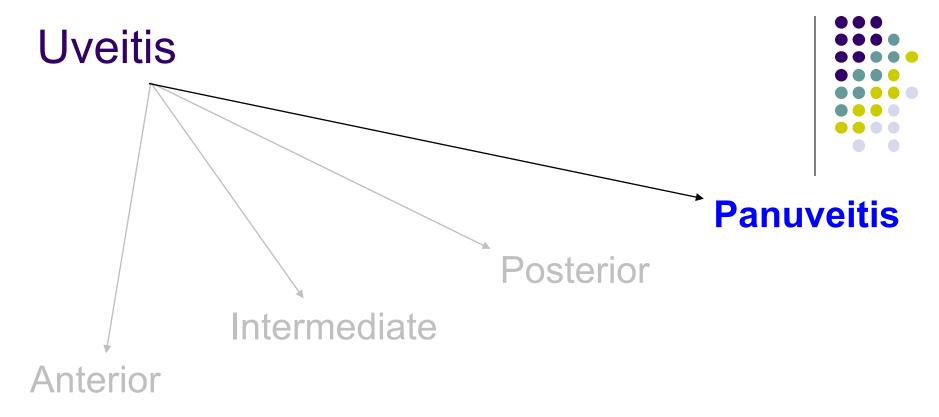












Sarcoid— Sympathetic ophthalmia—	Note that these three appear yet again	—Syphilis —TB
Vogt-Koyanagi-Harada—		—Lyme
Behçet syndrome—		—Leptospirosis
		—Whipple dz

### Endophthalmitis

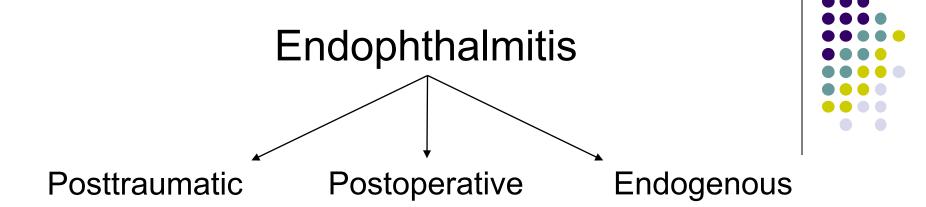


Now we'll change gears and look at endophthalmitis

### Endophthalmitis



The *Uveitis* book defines **endophthalmitis** as an inflammatory process involving both the AC and vitreous cavities that is 2ndry to bacterial or fungal infection.



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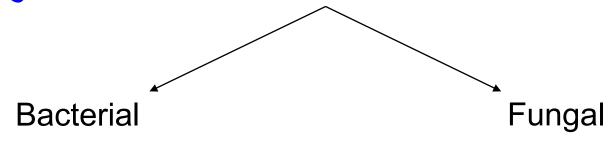
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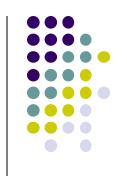
Endogenous endophthalmitis involves hematogenous spread of infection from a remote location to the eye. It is uncommon, accounting for less than 10% of all cases of endophthalmitis. Individuals at increased risk of endogenous endophthalmitis include those with impaired immune status, those who recently underwent an invasive medical procedure, and those subjected to chronic and/or repeated breaching of the body's outer barrier.



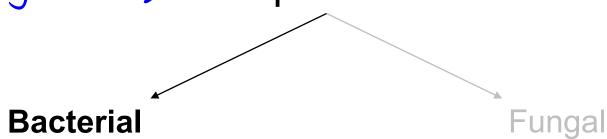
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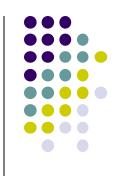
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As mentioned, endogenous endophthalmitis can be bacterial or fungal.





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Bacterial endophthalmitis

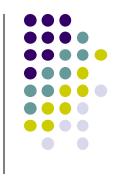


Bacterial Fungal



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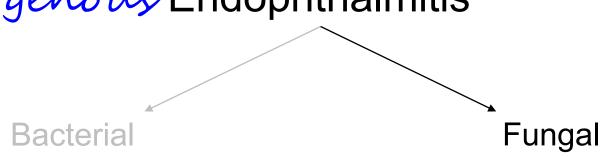
Some classic bacterial pathogen associations in endogenous endophthalmitis:

--Endocarditis: Strep --Skin infections: Staph

--IVDU: Bacillus

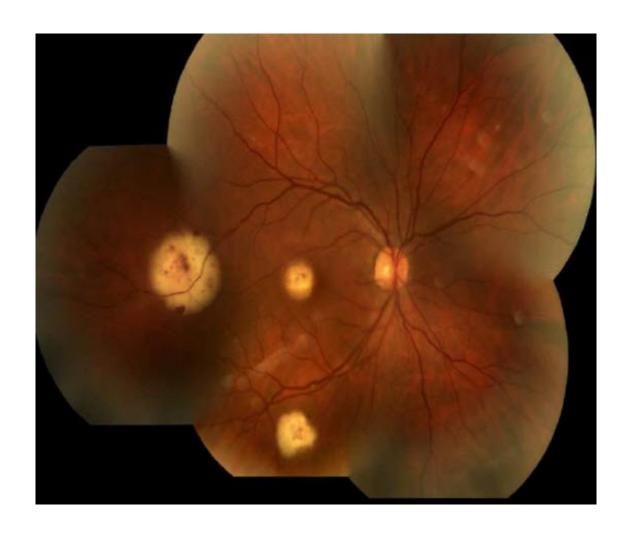
--Liver abscess: Klebsiella

the anterior segment.

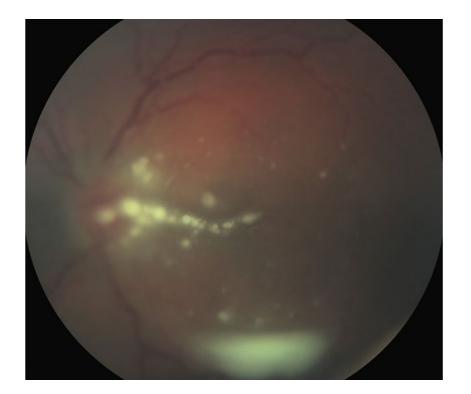


In contrast to the bacterial version, endogenous **fungal** endophthalmitis tends to be more insidious in onset. It generally progresses in a particular fashion. First, isolated choroidal metastatic lesions appear. With time, these break through Bruch's membrane to involve the retina. Eventually, the bug reaches the vitreous, and (if still unchecked)





Candida endophthalmitis: Choroidal lesions







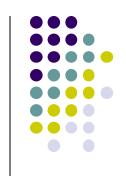
Candida endophthalmitis: Classic 'string of pearls' vitreous involvement





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## Some classic fungal pathogen associations in endogenous endophthalmitis:

- --Chronic indwelling lines/catheters: Candida
- --HIV/AIDS: Cryptococcus
- --Hx liver transplantation: Aspergillus
- --San Joaquin valley: Coccidioides



### **Scleritis**

Next let's take a look at scleritis



#### **Scleritis**

Scleritis is an inflammatory condition characterized by painful infiltrative scleral edema and congestion of the deep episcleral plexus. It can be extremely painful, and can lead to blindness and loss of the eye. Women are more likely to be affected than are men. It is rare in children.



#### **Scleritis**

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To make matters worse, it can herald the presence or worsening of a systemic conditions that may be potentially lethal. About 40% of scleritis pts have an identifiable systemic inflammatory condition, the most common of which is rheumatoid arthritis.



Scleritis is divvied up with respect to whether the...



Scleritis is divvied up with respect to whether the...

Anterior sclera is affected, vs the Posterior sclera.



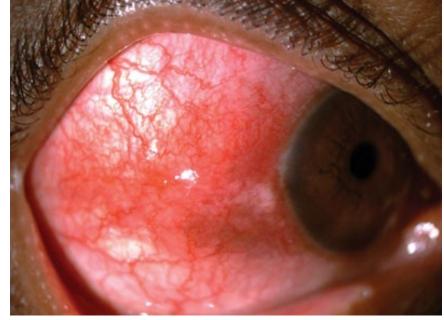
### **Scleritis**



There are three classic signs of anterior scleritis:

--Scleral edema

-





Anterior scleritis: Scleral edema



Anterior scleritis: Scleral edema. Note the thickening of the limbal sclera (b) in comparison to the unaffected fellow eye (a)



#### **Scleritis**



There are three classic signs of anterior scleritis:

- --Scleral edema
- --Sclera has a violaceous hue

\_\_\_





Anterior scleritis: Violaceous hue

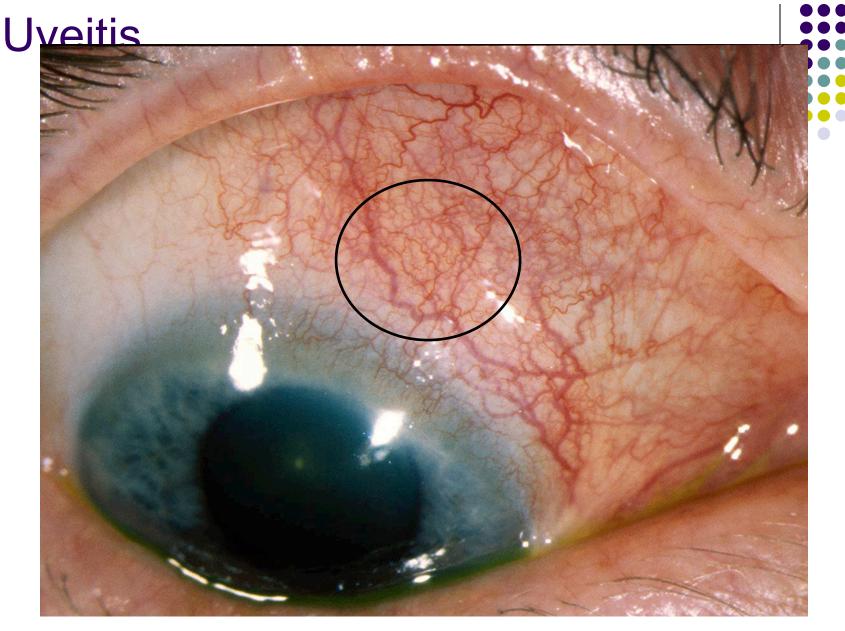


#### **Scleritis**



There are three classic signs of anterior scleritis:

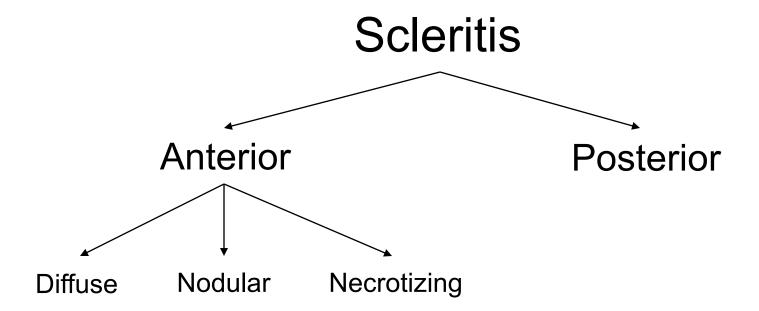
- --Scleral edema
- --Sclera has a violaceous hue
- --Inflamed vasculature has a criss-cross pattern



108

'Criss-cross' injection of the deep vasculature in anterior scleritis. To see it, you have to look past the brighter injection of the inflamed overlying conj vessels



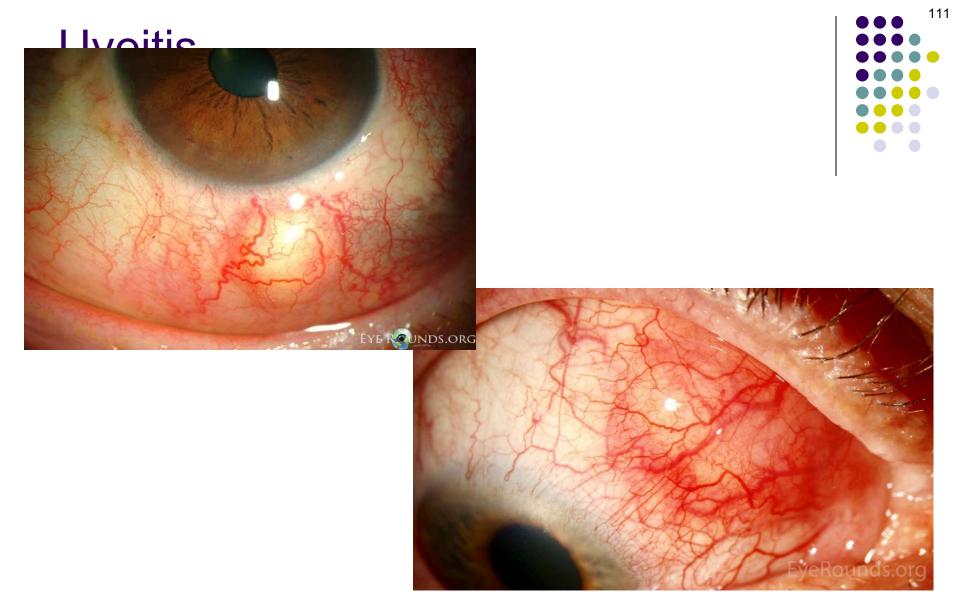


Anterior scleritis comes in three forms: Diffuse, nodular and necrotizing



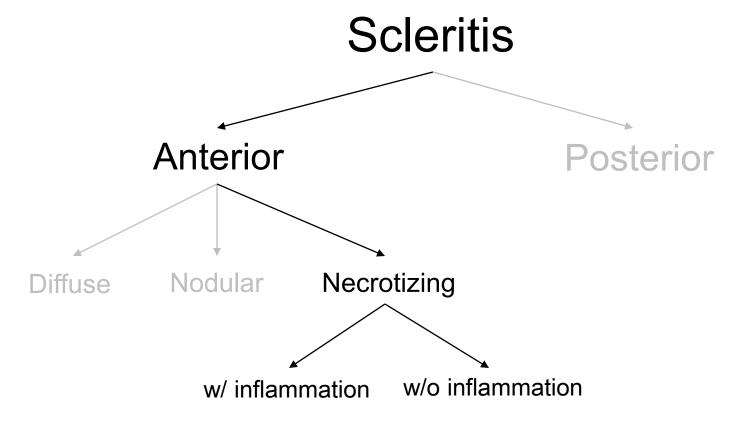


Diffuse anterior scleritis



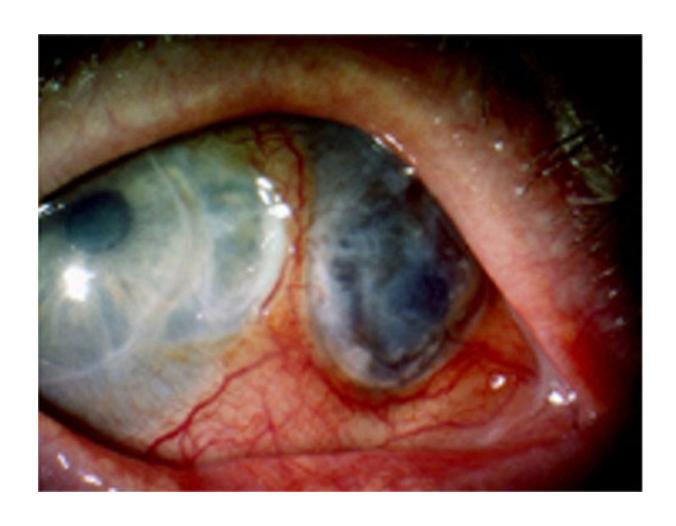
Nodular anterior scleritis





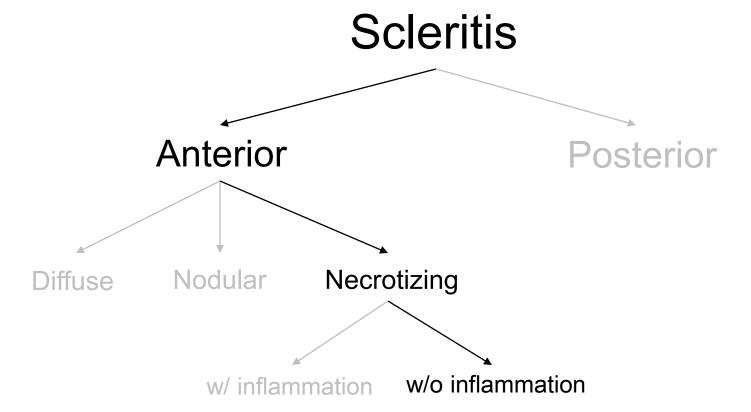
Necrotizing anterior scleritis comes in two forms: with and without inflammation





Necrotizing anterior scleritis with inflammation



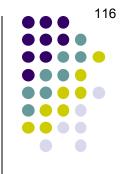


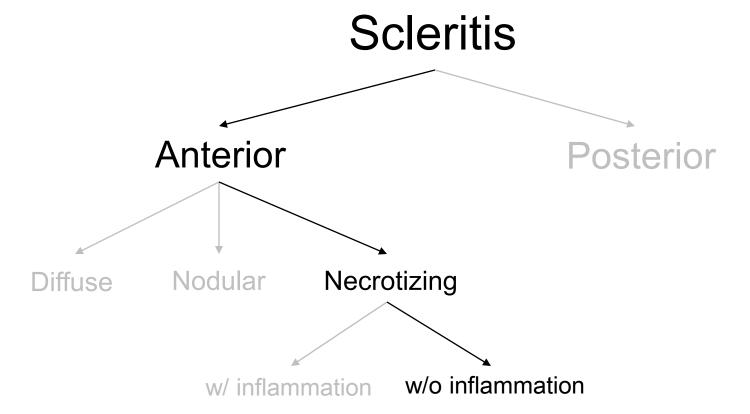
Contrary to the implications of the name, inflammation is present in *necrotizing scleritis w/o inflammation*. It is so named because, unlike its 'with inflammation' cousin, it is typically painless, and the eye does not *appear* inflamed.





Necrotizing anterior scleritis without inflammation

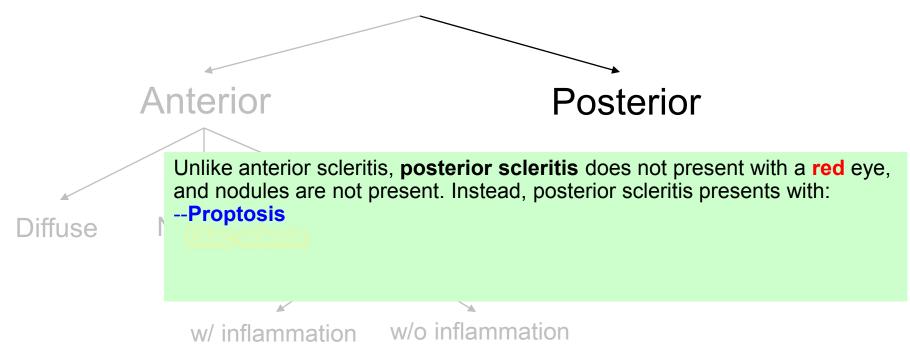




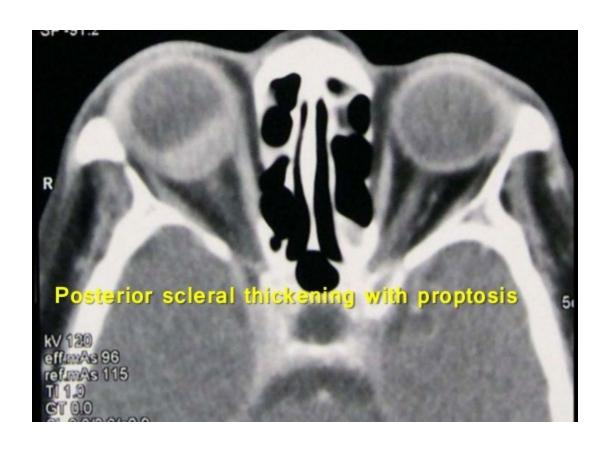
Contrary to the implications of the name, inflammation is present in *necrotizing scleritis w/o inflammation*. It is so named because, unlike its 'with inflammation' cousin, it is typically painless, and the eye does not *appear* inflamed. Necrotizing scleritis w/o inflammation is also known as **scleromalacia perforans**. It is strongly associated with RA.



## **Scleritis**



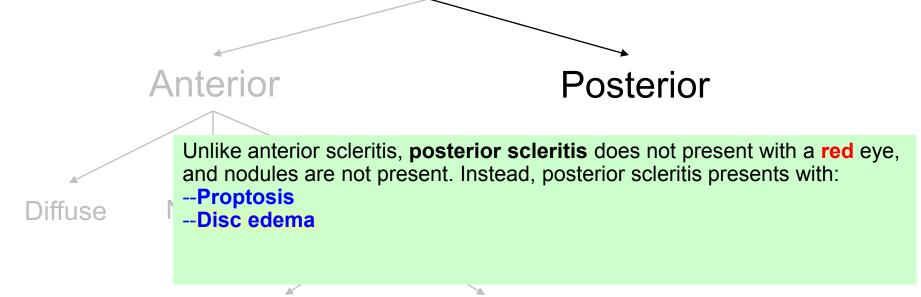




Posterior scleritis: Proptosis



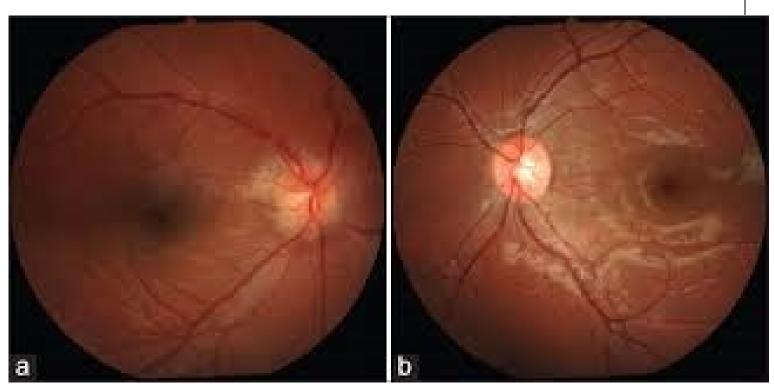
# **Scleritis**



w/ inflammation

w/o inflammation





Posterior scleritis OD: Optic nerve edema

Diffuse



# Scleritis



Unlike anterior scleritis, **posterior scleritis** does not present with a **red** eye, and nodules are not present. Instead, posterior scleritis presents with:

- --Proptosis
- -- Disc edema
- --Retinal/choroidal findings

w/ inflammation w/o inflammation





Posterior scleritis producing retinal folds

Diffuse



## Scleritis

Anterior Posterior

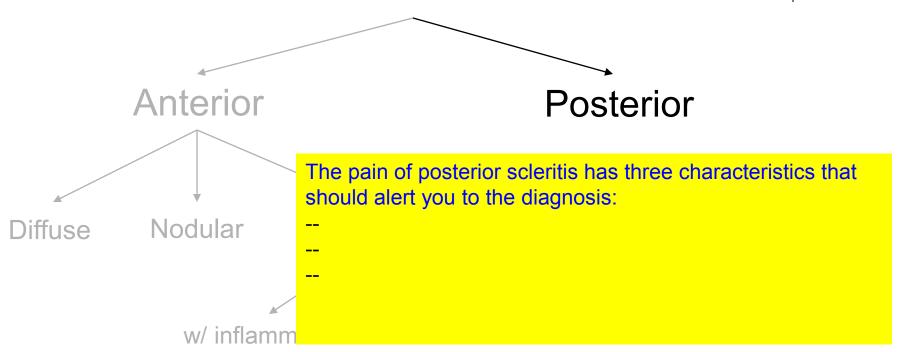
Unlike anterior scleritis, **posterior scleritis** does not present with a **red** eye, and nodules are not present. Instead, posterior scleritis presents with:

- --Proptosis
- -- Disc edema
- --Retinal/choroidal findings
- -- Motility disorders

w/ inflammation w/o inflammation

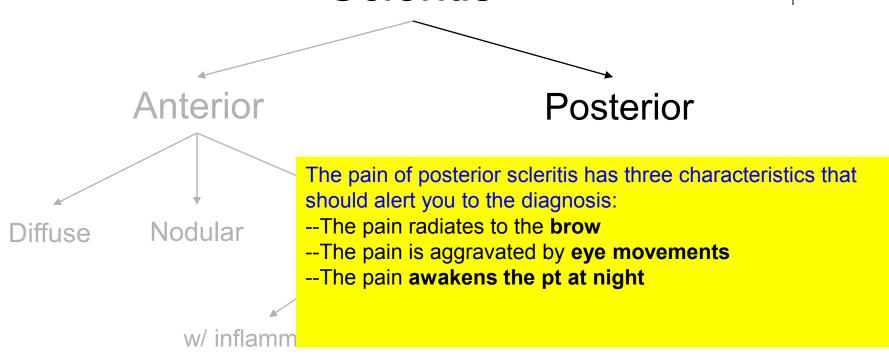






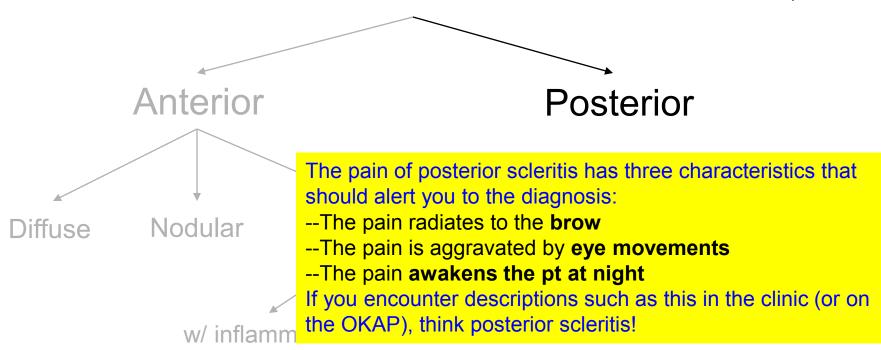


# **Scleritis**



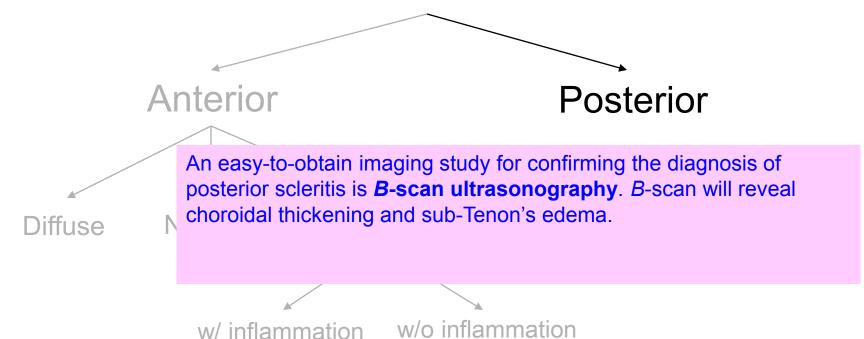








# **Scleritis**







Posterior scleritis: Sub-Tenon's edema



#### **Scleritis**

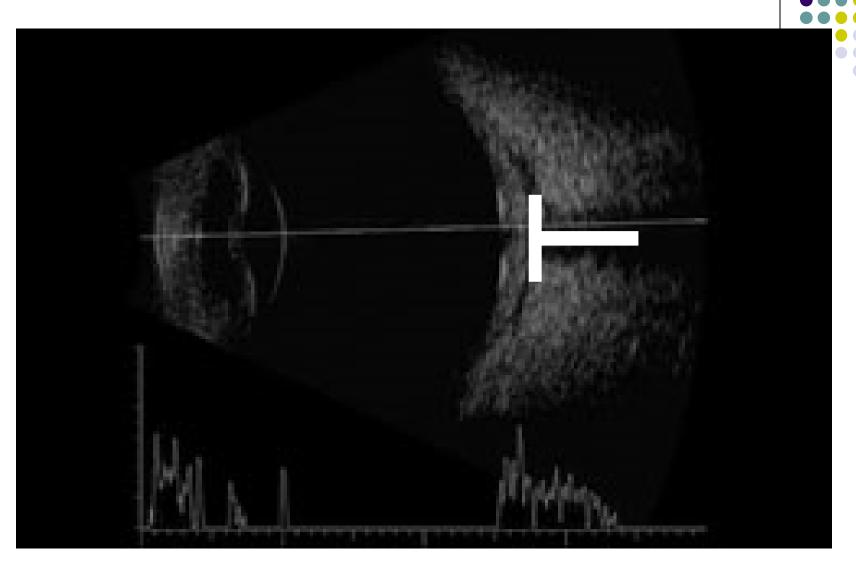


Diffuse

An easy-to-obtain imaging study for confirming the diagnosis of posterior scleritis is **B-scan ultrasonography**. **B-scan will reveal choroidal thickening and sub-Tenon's edema**. When sub-Tenon's edema involves the space around the optic nerve, the classic **T sign** finding will result.

w/ inflammation

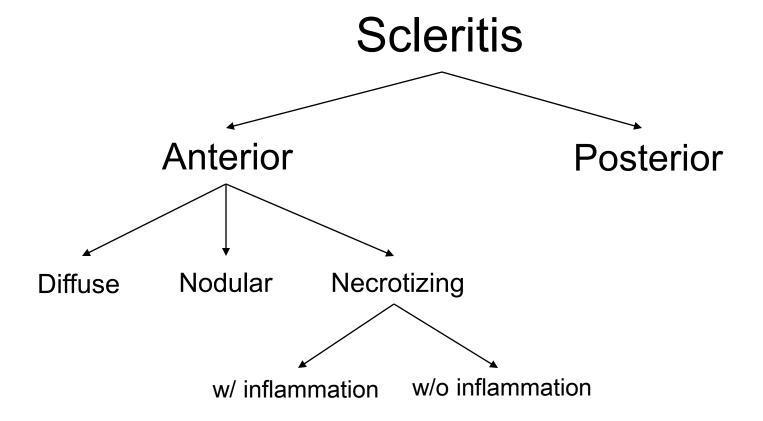
w/o inflammation



130

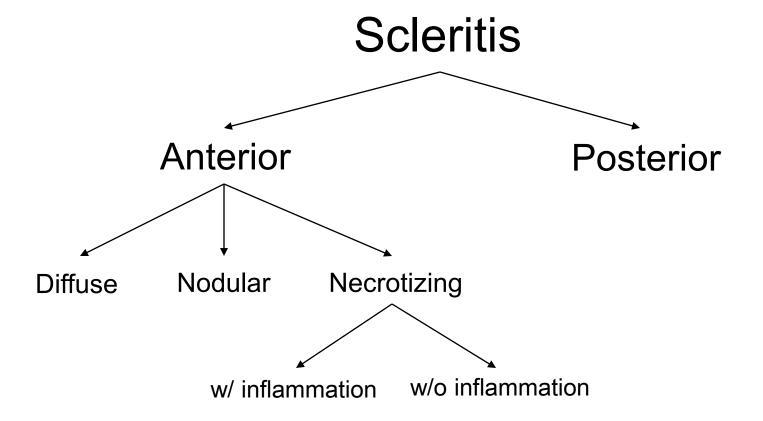
Posterior scleritis: T-sign





**Scleritis requires systemic treatment**. Diffuse scleritis might respond to PO NSAIDs—try these first. For the others, PO steroids are usually the first-line med, although NSAIDs may be tried. More powerful immunosuppression is frequently required.





Scleritis requires systemic treatment. Diffuse scleritis might respond to PO NSAIDs—try these first. For the others, PO steroids are usually the first-line med, although NSAIDs may be tried. More powerful immunosuppression is frequently required. Subconj depot steroids, long considered contraindicated, have recently gained wide acceptance as a treatment option.

# Masquerade Syndrome



Finally, we will look at masquerade syndromes

# Masquerade Syndrome refers to entities



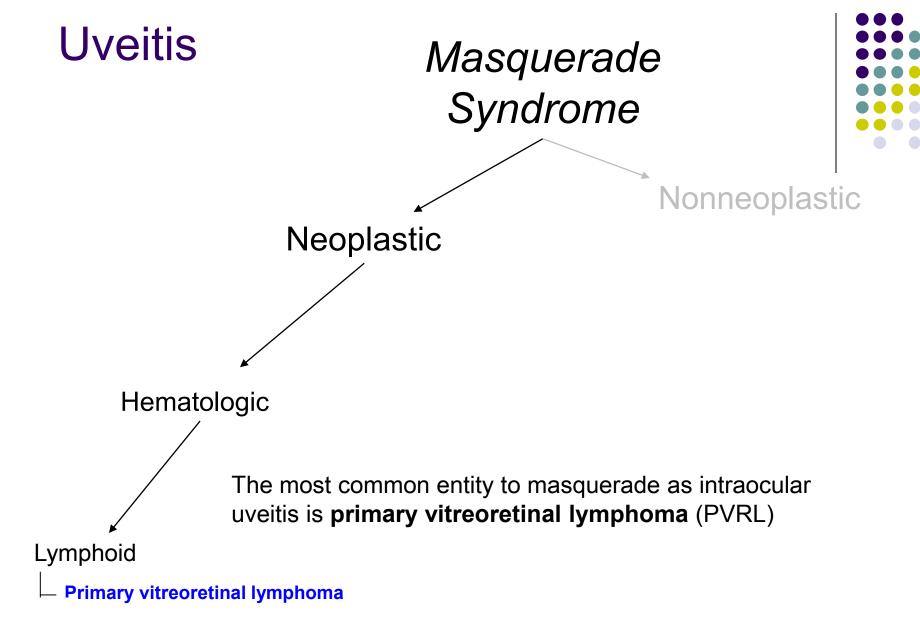


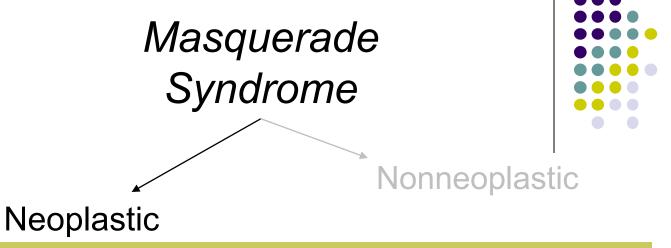




that mimic immune-mediated dz.

The entities can be broadly divided into Nonneoplastic and Neoplastic causes.





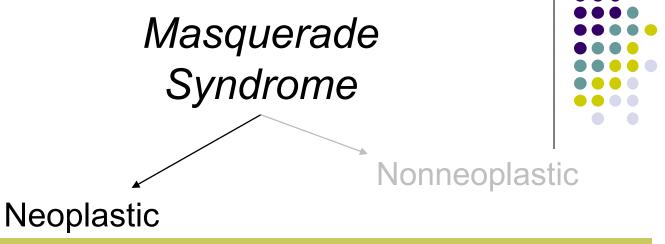
Virtually all PVRLs are non-Hodgkin B-cell lymphomas. The typical PVRL pt is an adult in their 50s-60s. They usually present with complaints of decreased vision and/or floaters.



The most common entity to masquerade as intraocular uveitis is **primary vitreoretinal lymphoma** (PVRL)

Lymphoid

**Primary vitreoretinal lymphoma** 

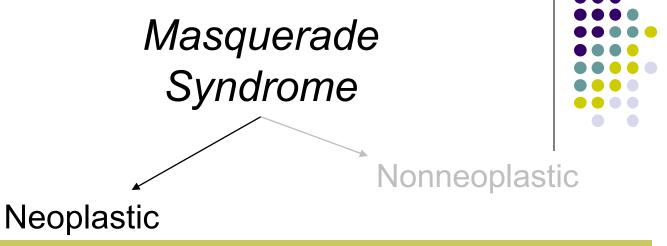


Virtually all PVRLs are non-Hodgkin B-cell lymphomas. The typical PVRL pt is an adult in their 50s-60s. They usually present with complaints of decreased vision and/or floaters. Importantly, many will also manifest evidence of CNS involvement, the most common being changes in behavior or personality. Other, more obvious S/S include seizures, cerebellar signs, hemiparesis and cranial nerve palsies. Confusion, weakness, and memory loss may also occur.

Lymphoid

The most common entity to masquerade as intraocular uveitis is **primary vitreoretinal lymphoma** (PVRL)

└ Primary vitreoretinal lymphoma



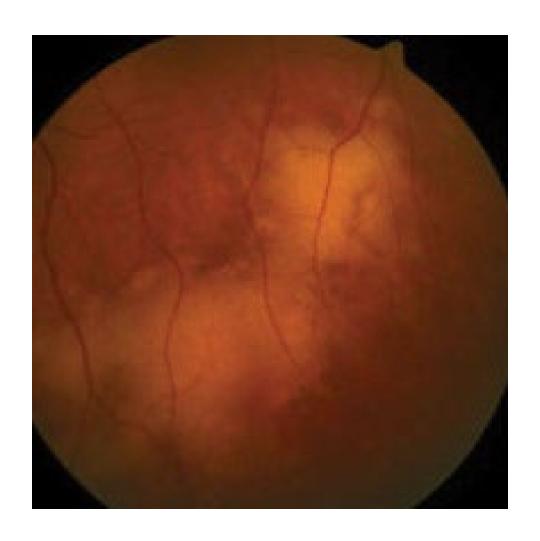
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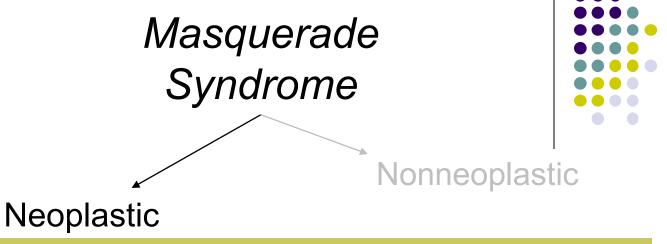
Lymphoid

**Primary vitreoretinal lymphoma** 





PVRL: Typical white-yellow subretinal infiltrates



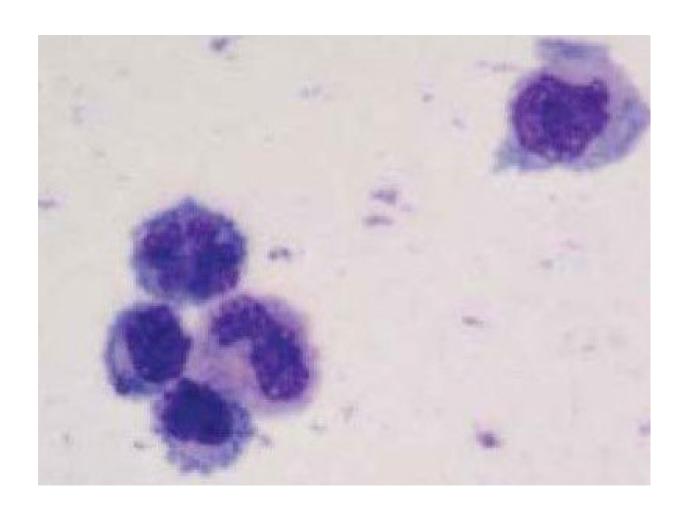
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The most common entity to masquerade as intraocular uveitis is **primary vitreoretinal lymphoma** (PVRL)

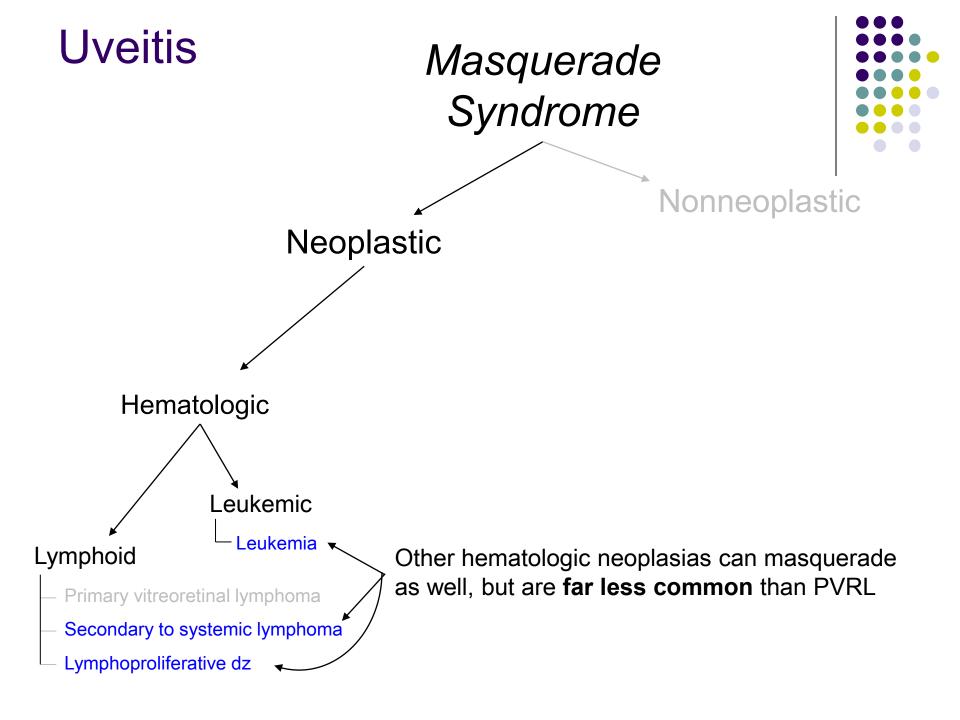
Lymphoid

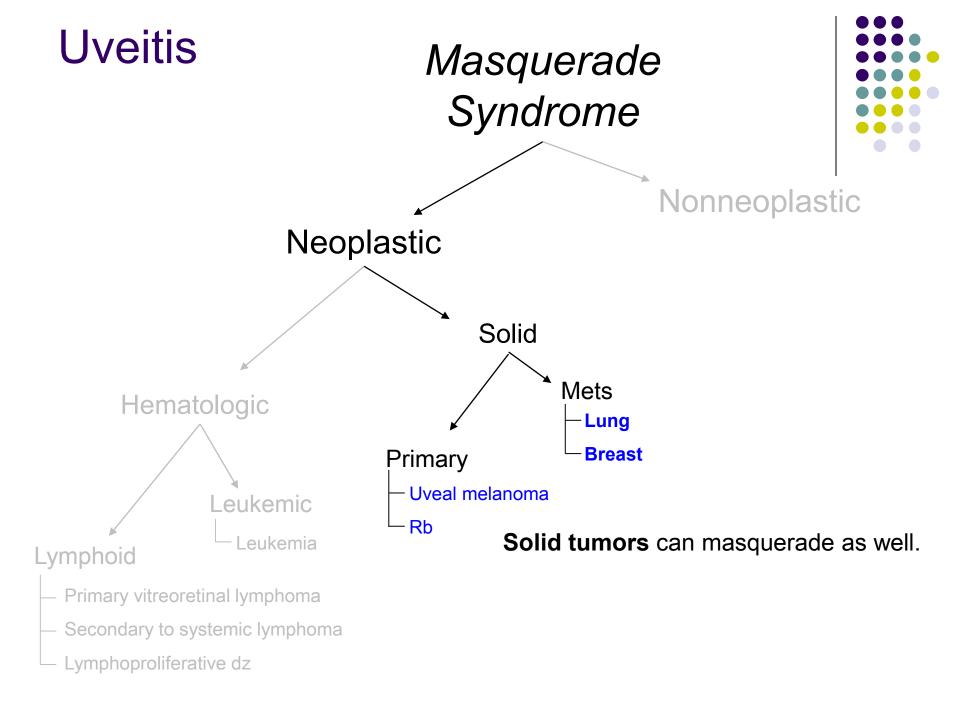
**Primary vitreoretinal lymphoma** 

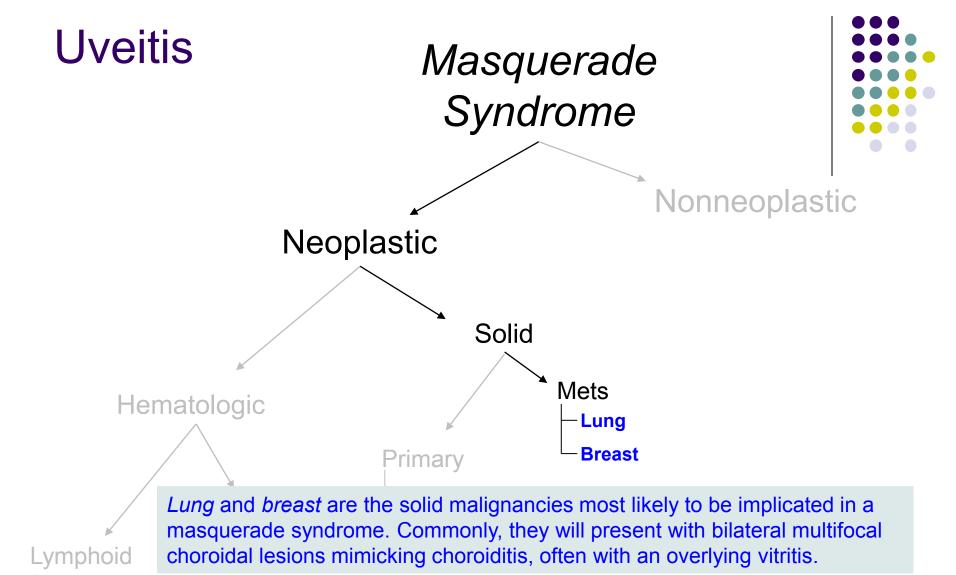




Typical cytology of PVRL cells from the vitreous showing several atypical lymphoid cells with basophilic cytoplasm and large prominent irregular nuclei







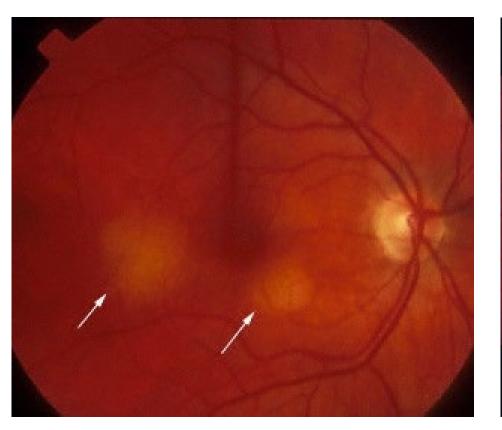
Primary vitreoretinal lymphoma

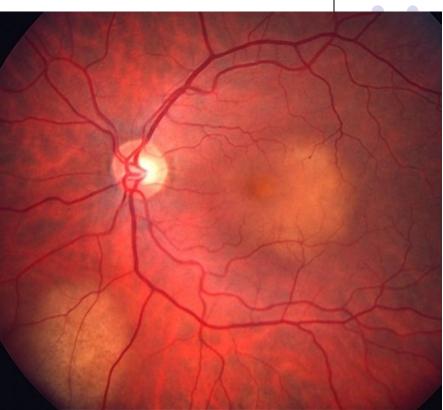
Secondary to systemic lymphoma

Lymphoproliferative dz

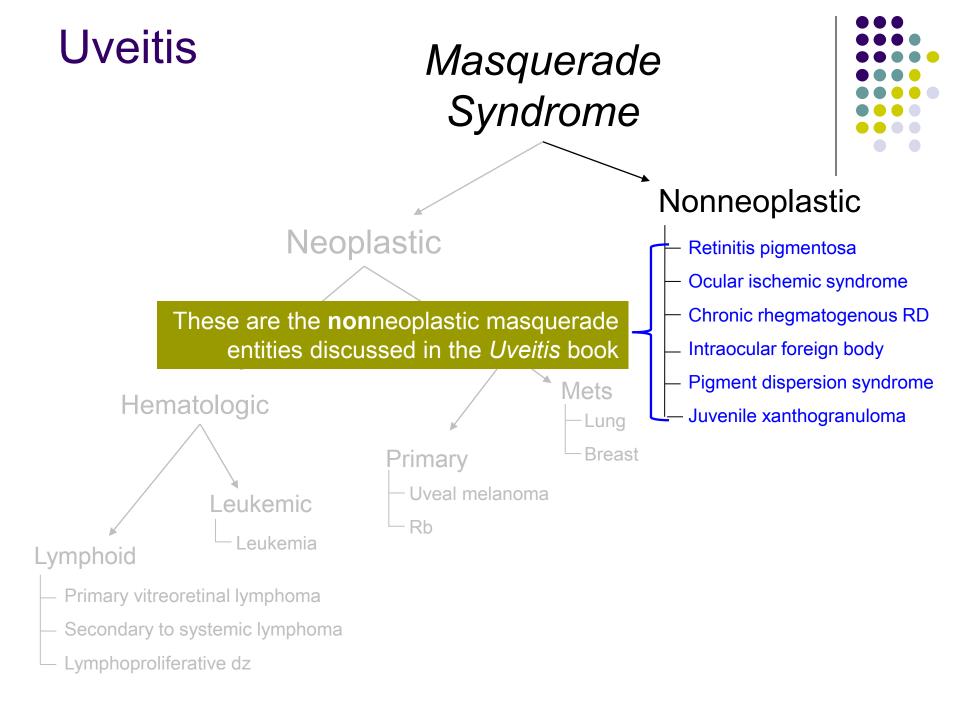








Bilateral metastatic breast cancer



# Masquerade Syndrome



Ocular ischemic syndrome (OIS) is a constellation of ocular abnormalities stemming from chronic hypoperfusion of the globe. The classic cause is carotid stenosis ipsilateral to the eye in question. The typical pt is an elderly vasculopathic male.

Nonneoplastic



S

ıng

east

Chronic rhegmatogenous RD

Intraocular foreign body

Pigment dispersion syndrome

- Juvenile xanthogranuloma

Leukemic

Leukemia

- Rb

Lymphoid

Primary vitreoretinal lymphoma

Secondary to systemic lymphoma

Lymphoproliferative dz

# Masquerade Syndrome

S

ıng

east



Ocular ischemic syndrome (OIS) is a constellation of ocular abnormalities stemming from chronic hypoperfusion of the globe. The classic cause is carotid stenosis ipsilateral to the eye in question. The typical pt is an elderly vasculopathic male.

Four findings, common in OIS, can (mis)lead one to conclude the pt has uveitis:

- --AC cell and flare
- --Low IOP
- --Neovascularization of the iris and/or angle
- -- Cataract more advanced on that side

Leukemia Steal Melanema

— Rb

— Leukemia

#### Lymphoid

Primary vitreoretinal lymphoma

Secondary to systemic lymphoma

Lymphoproliferative dz

#### Nonneoplastic



Chronic rhegmatogenous RD

Intraocular foreign body

Pigment dispersion syndrome

- Juvenile xanthogranuloma

# Masquerade Syndrome



RP

OIS

Chronic rhegmatogenous RD

\_ IOFB

**Pigment dispersion syndrome** 

Juvenile xanthogranuloma

Neoplastic

The hallmark of *PDS* is the liberation of pigment from the posterior aspect of the iris. This pigment subsequently migrates into the anterior chamber, where the pigment granules can be mistaken for inflammatory cells.

eukemic

Leukemia

Primary vitreoretinal lymphoma

Secondary to systemic lymphoma

Lymphoproliferative dz

Lymphoid

Uveal melanoma

Rb

# Masquerade Syndrome



Neoplastic

The hallmark of *PDS* is the liberation of pigment from the posterior aspect of the iris. This pigment subsequently migrates into the anterior chamber, where the pigment granules can be mistaken for inflammatory cells. Typically, retroillumination of the iris will reveal transillumination defects with a radial orientation.

- Uveal melanoma

- Rb

Lymphoid

Primary vitreoretinal lymphoma

Secondary to systemic lymphoma

eukemic

eukemia

Lymphoproliferative dz

- RP

- OIS

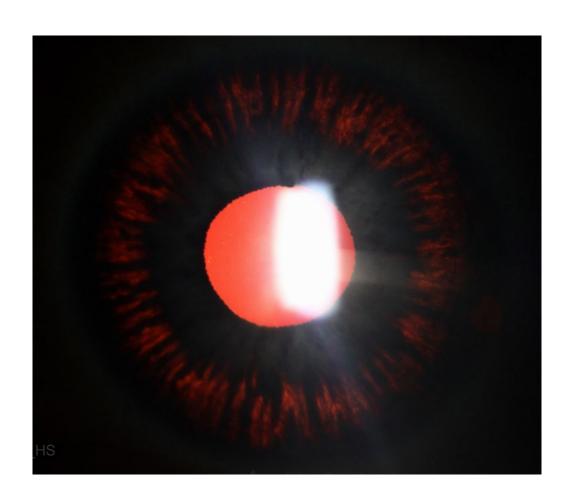
Chronic rhegmatogenous RD

– IOFB

**Pigment dispersion syndrome** 

Juvenile xanthogranuloma





PDS: Radial transillumination defects



That's it! Go through this slide-set a couple of times (at least) until you feel like you have a handle on it. When you're ready, do slide-set *U10*, which covers this material in a Q&A format (and more detail).