

# DIABETIC RETINOPATHY



# Diabetic eye disease

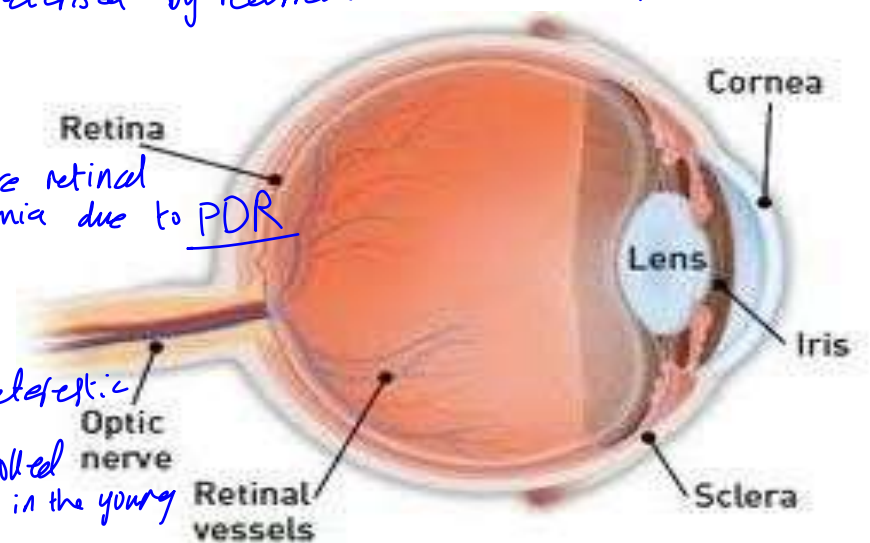
Refers to a group of eye problems that people with diabetes may face as a complication of diabetes.

All can cause variable degrees of vision loss or visual symptoms (pain and Diplopia )

# Diabetic eye disease

From anterior to posterior

- Corneal abnormalities → characterized by recurrent attacks of corneal abrasions
- Iris and angle  
Neovascularization. due to severe retinal ischemia due to PDR
- Neovascular Glaucoma
- Cataracts... snowflake cataracts in young pts and greater frequency and earlier onset of age related cataract. characteristic to uncontrolled DM in the young
- Ocular Neuropathies. → nerves 3<sup>rd</sup>, 4<sup>th</sup>, 6<sup>th</sup> CN → usually present with diplopia
- Diabetic Retinopathy. → the most common, serious, severe complication



Diabetic cataract, or “snowflake” cataract,

(rare type specific for DM in younger  
(uncontrolled DM))



+ most common cataract type  
that occurs with DM  
is senile cataract  
(that comes with aging)

# Diabetic retinopathy :

It is a progressive microangiopathy of the retinal blood vessels caused by chronic hyperglycemia.

Diabetic retinopathy - most common cause of moderate to severe vision loss between ages 25 and 74 years. → working age group

\* most common cause of visual impairment + vision loss in diabetic retinopathy is diabetic macular edema

other causes may include: macular ischemia, vitreous hemorrhage, preretinal hemorrhage, tractional retinal detachment, neovascular glaucoma.

# The Retina

A structure that lines the inside of the globe

- **Two major layers:**

- Inner neurosensory retina (NSR): transparent, has the photoreceptors (rods and cones), light sensitive
- Outer retinal pigment epithelium (RPE).

→ between those 2 layers there's a potential space called subretinal space → usually is dry, if fluid is there → retinal detachment occurs (separation of 2 layers by fluid build up)

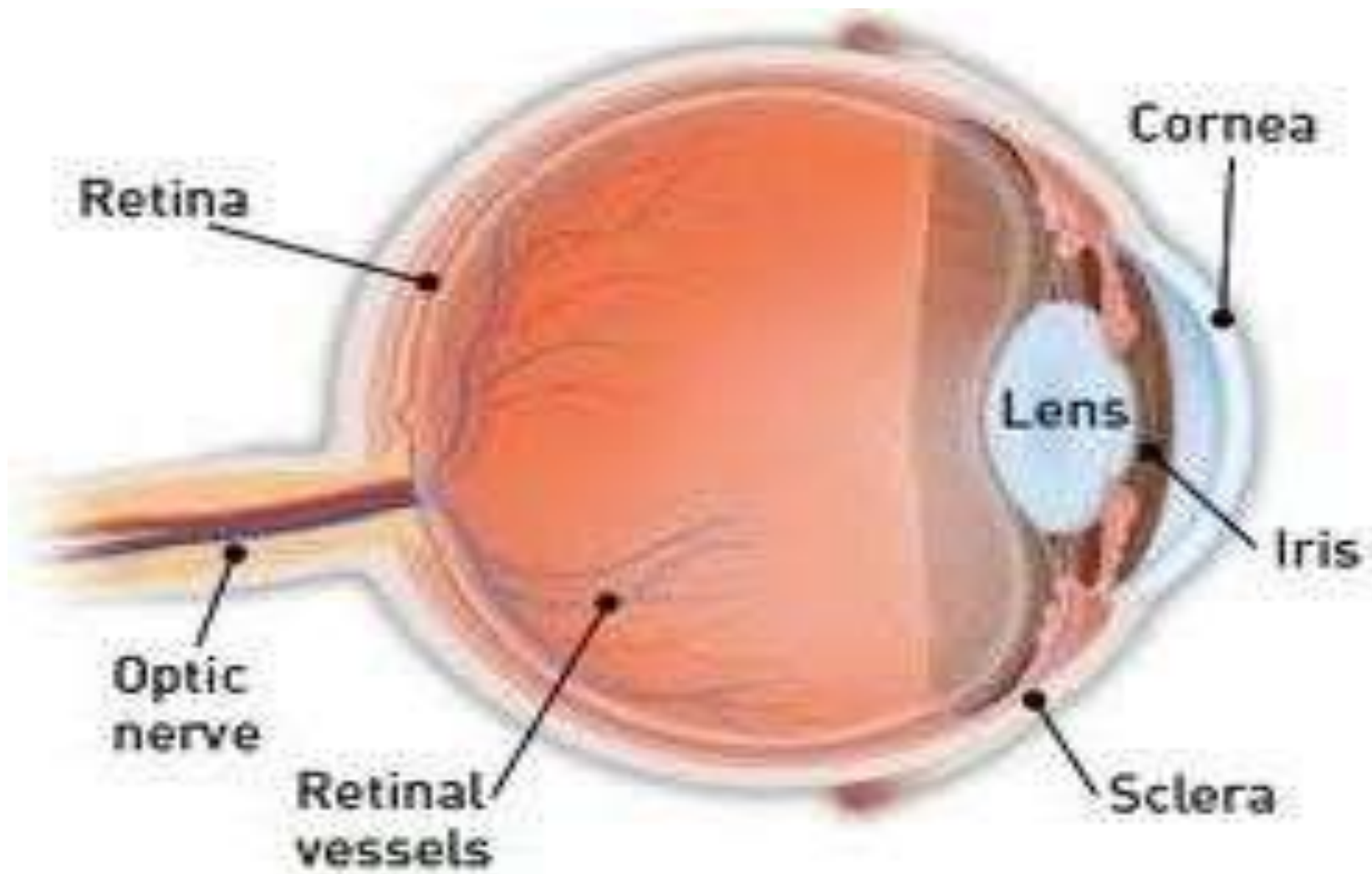
- **Retinal blood supply:**

From central retinal artery and choroidal circulation.

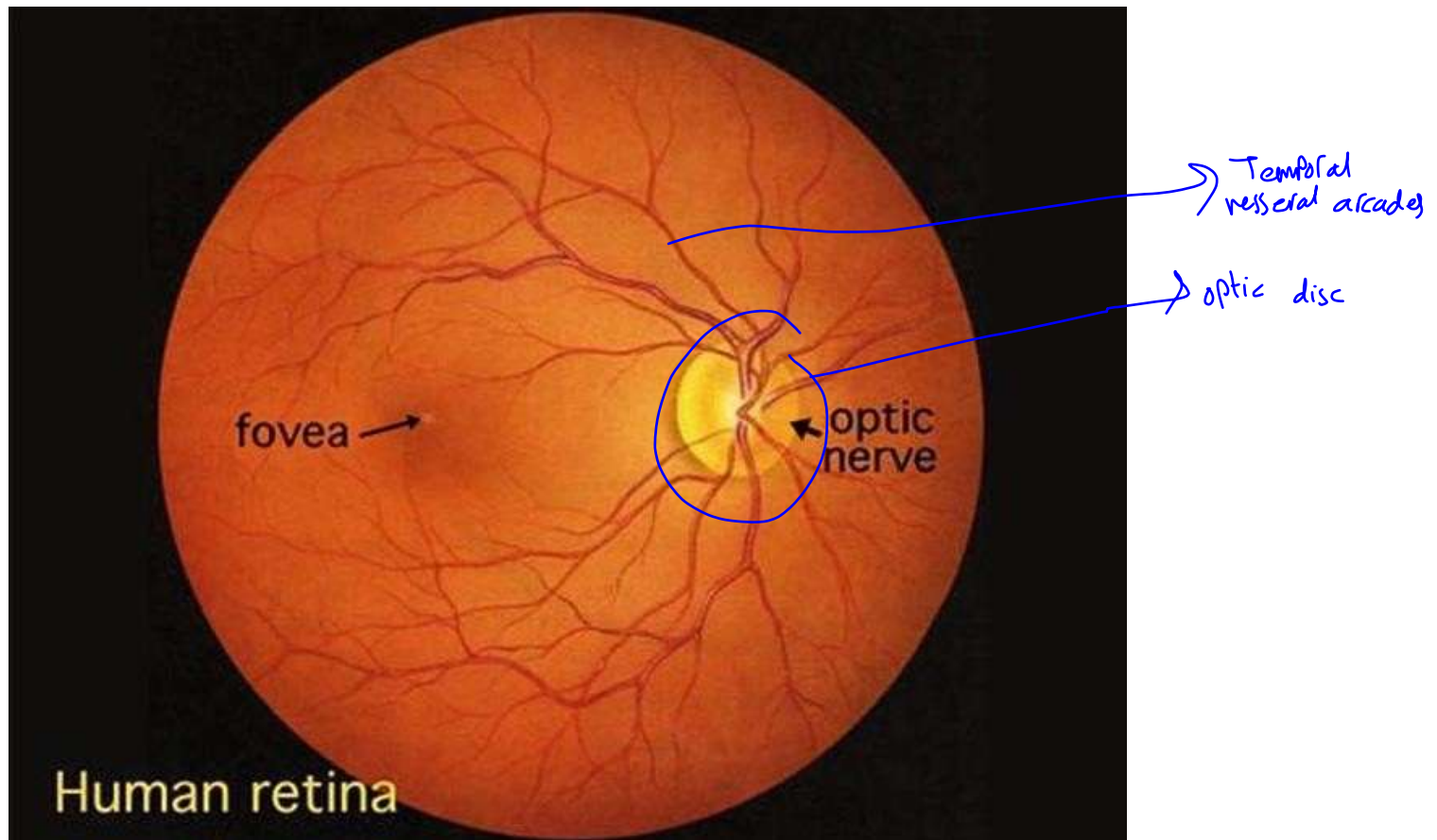
↳ supplies the inner part

(short ciliary artery)

↳ supplies the outer part of retina



# Retinal Anatomy *as viewed by indirect ophthalmoscope*





# RISK FACTORS:

Duration of diabetes ✕

Poor control of diabetes ✕

most important risk factors

⇒ most important modifiable risk factor

Hypertension

Nephropathy

hyperlipidemia

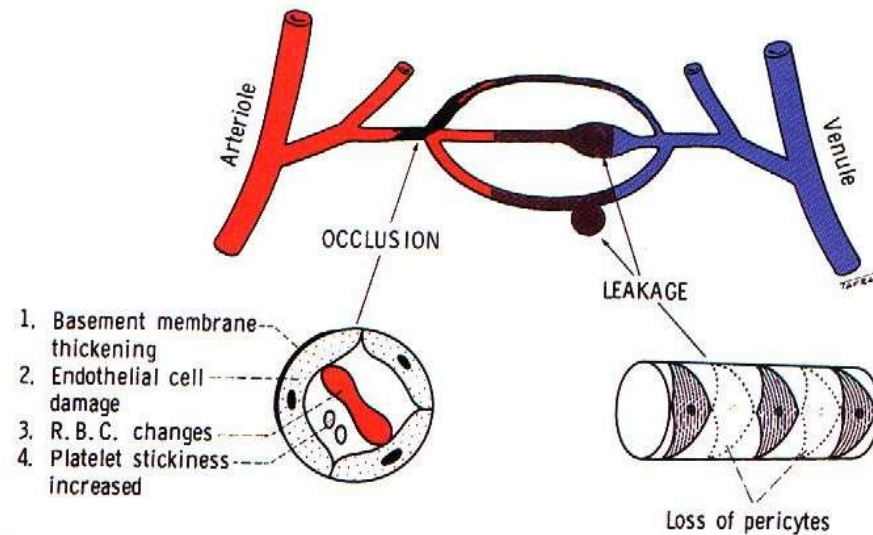
Smoking

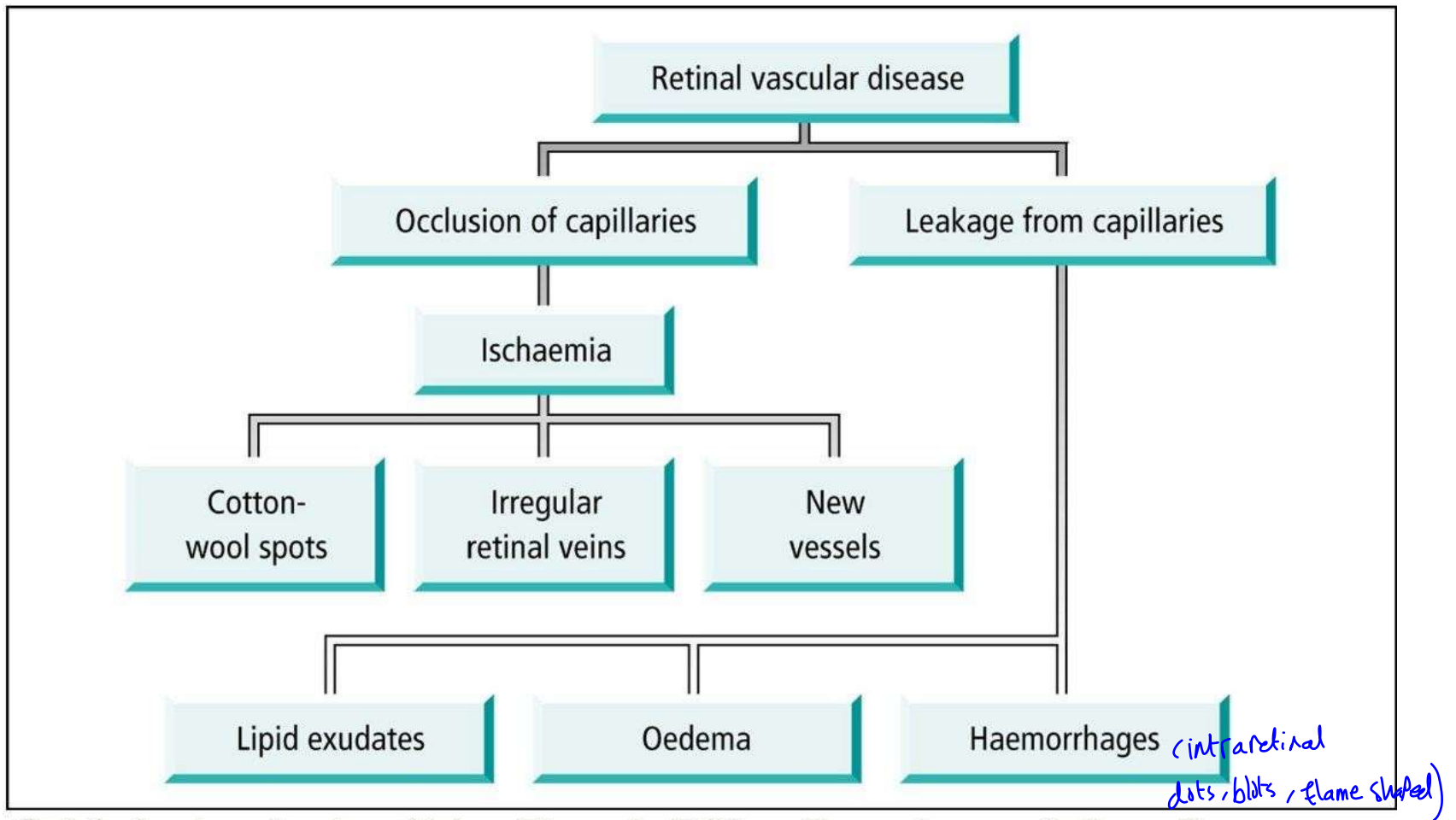
Obesity

Pregnancy

# Pathogenesis

Microangiopathy which has features of both microvascular leakage and occlusion.





*Ophthalmology Lecture Notes*, Eleventh Edition. Bruce James, Anthony Bron.  
 © 2011 Bruce James and Anthony Bron. Published 2011 by Blackwell Publishing Ltd.

**Figure 12.1** The building blocks of retinal vascular disease. Capillary leakage and occlusion often occur together.

# Microvascular leakage

Loss of pericytes results into :

Distention of capillary wall producing  
*microaneurysms*

Disruption of the inner Blood-retinal barrier p  
causing plasma constituents to leak into the retina  
*retinal edema, hard exudates*

# Microvascular occlusion

Basement membrane thickening, endothelial cell damage, deformed RBCs, platelet stickiness and aggregation

Vascular Endothelial Growth Factor (VEGF) is produced by **hypoxic retina**

VEGF stimulates the shunt and growth of new vessels

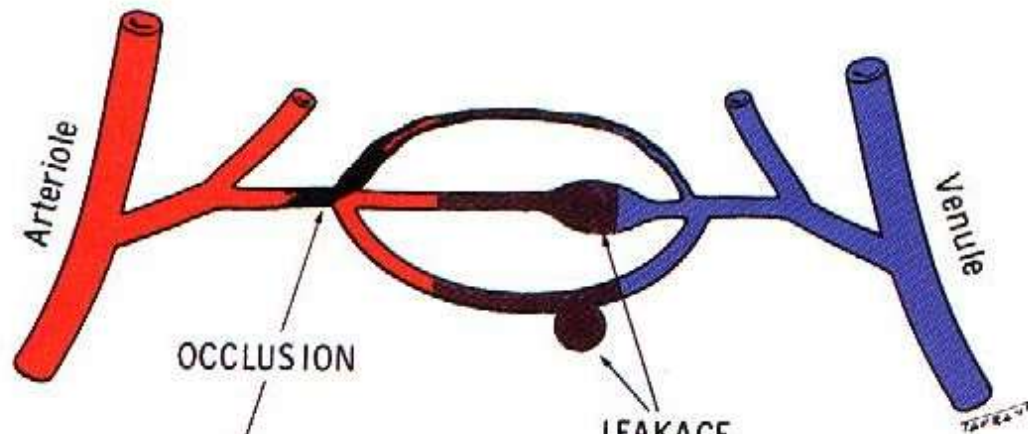
# New vessels is the hallmark of proliferative diabetic retinopathy

These new vessels forms at or near the optic disc (NVDs), anywhere in the retina (NVEs) or at iris (NVI)s

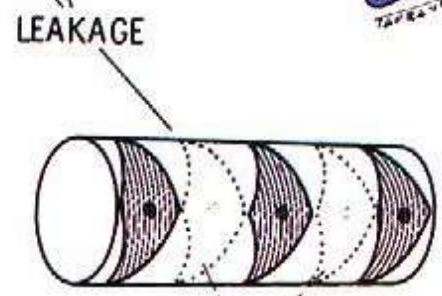
The new vessels break easily and leak into the vitreous gel producing vitreous hemorrhage.

\* leak between retina + vitreous gel → preretinal / subhyaloid hemorrhage

With time the fibrous component of new vessels contracts and results into traction retinal detachment



1. Basement membrane thickening
2. Endothelial cell damage
3. R. B. C. changes
4. Platelet stickiness increased



Loss of pericytes

## Signs of diabetic retinopathy :

Early signs (signs of non proliferative DR) :

- ❖ Microaneurysms
- ❖ Dot and blot hemorrhages
- ❖ Flame-shaped hemorrhages
- ❖ Cotton-wool spots
- ❖ Hard exudates
- ❖ Edema
- ❖ Venous changes ( beading and looping )
- ❖ Intraretinal microvascular abnormalities (IRMAs)

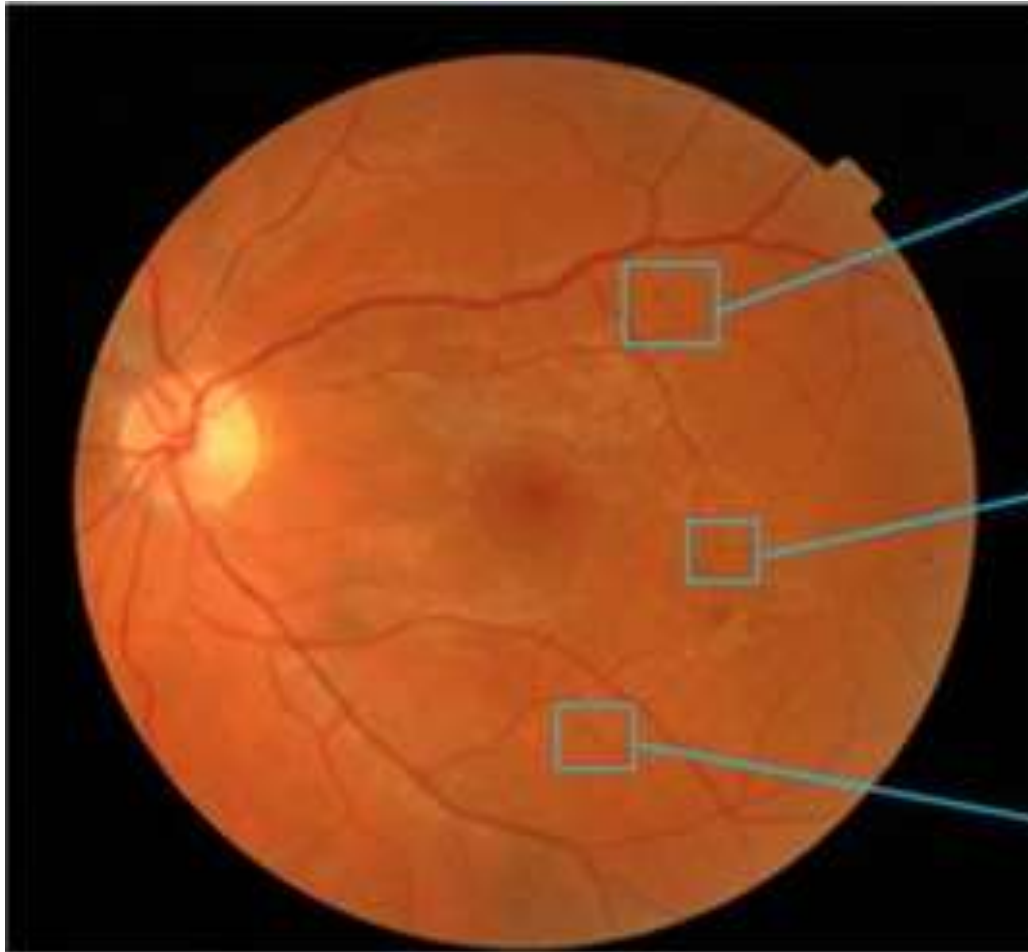


## ❖ Microaneurysms :

-Earliest clinical sign of diabetic retinopathy.

-Appear as small red dots in the superficial retinal layers

-Rupture produces dot, blot and flame shaped hemorrhages → rupture of superficial vessels



Microaneurysms  
First detectable  
sign of DR  
They are hard  
to distinguish  
between them  
and dots  
hemorrhages

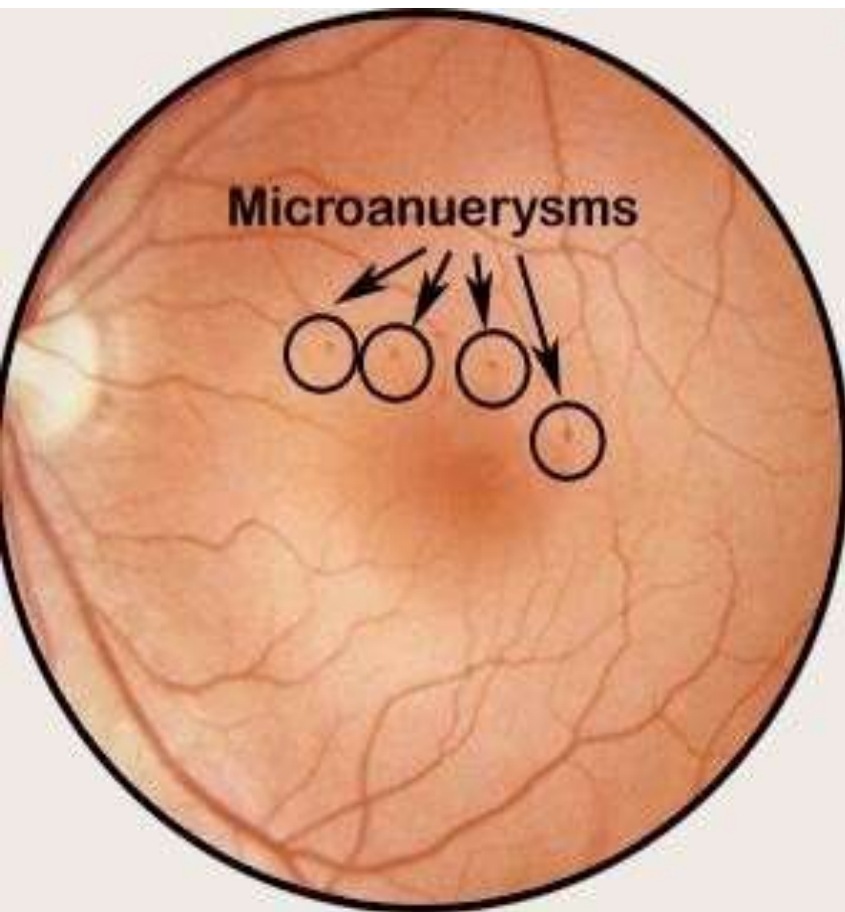


↓  
We can use  
fluorescent



aneurysms →  
hyperfluorescent  
hemorrhages  
→ hypo fluorescent

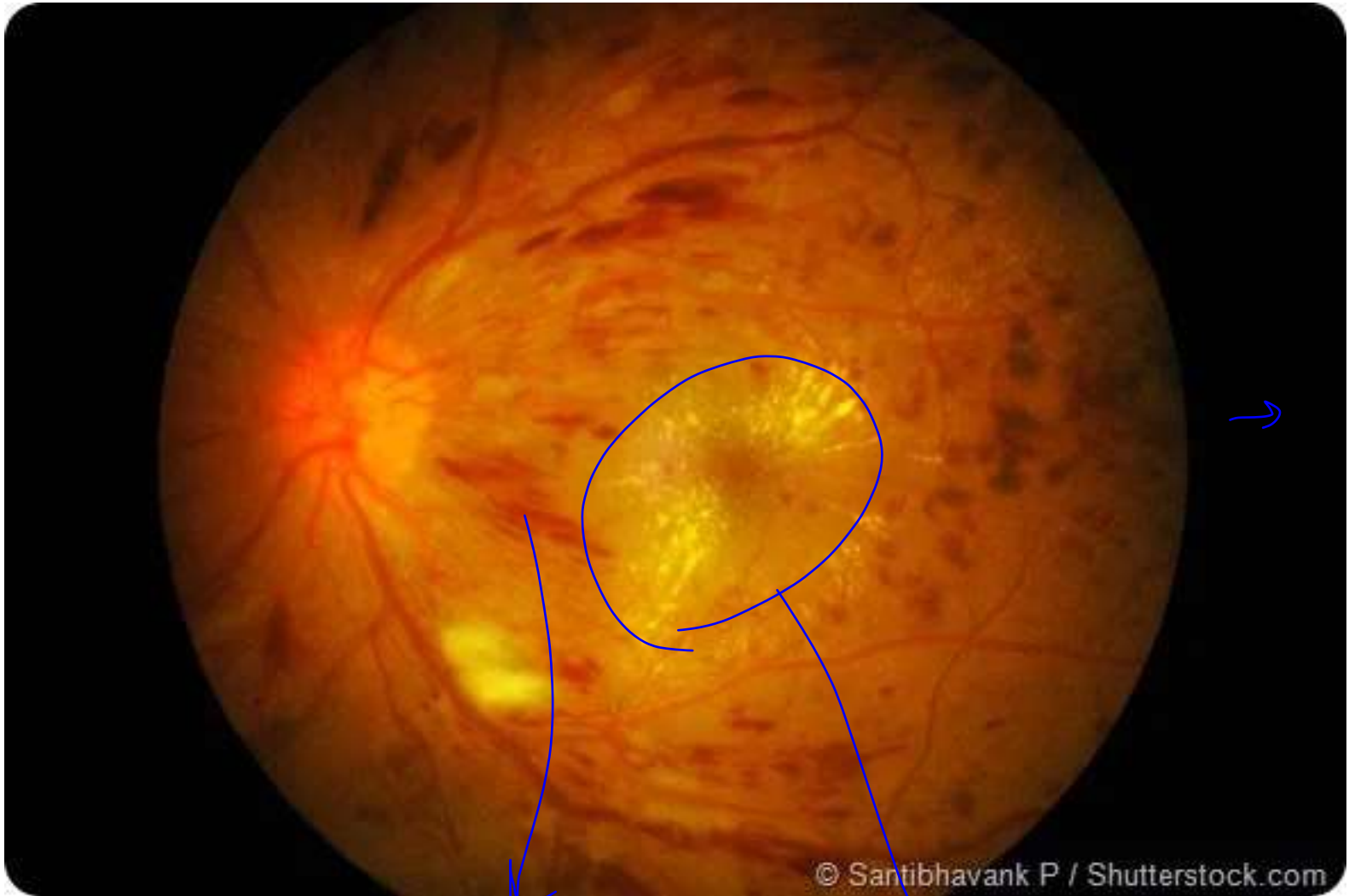
⇒ both of  
them appear  
as red dots  
clinically



## ❖ Dot and blot hemorrhages

Occur as microaneurysms rupture in the deeper layers of the retina (similar to microaneurysms if they are small, distinguish by fluorescein angio).

❖ - Splinter or flame shaped hemorrhages, superficial.



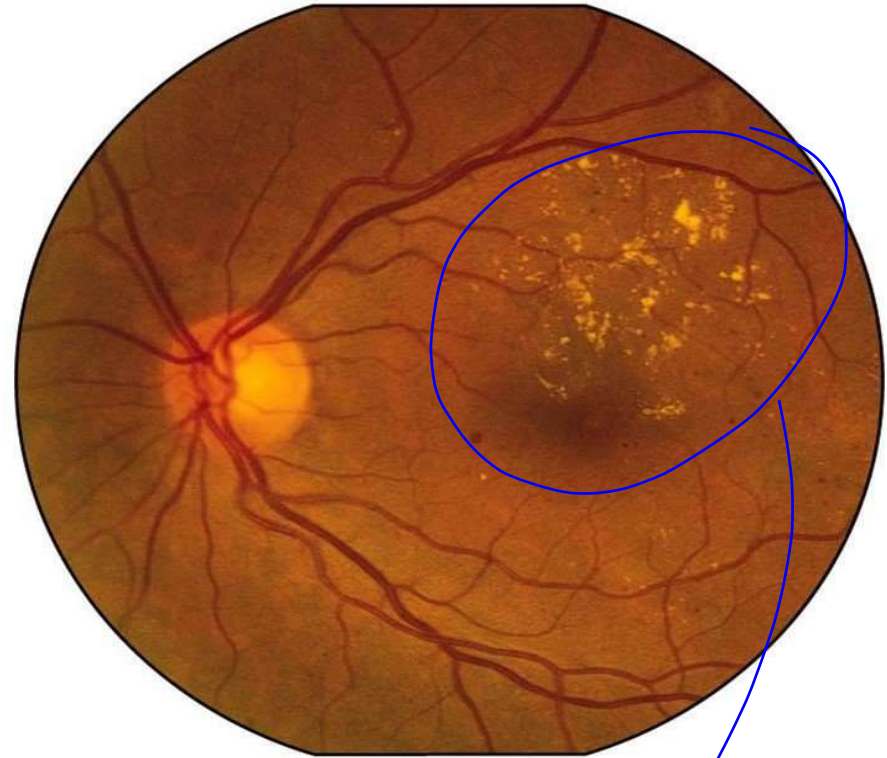
blots  
hemorrhages

red streaks are flame hemorrhages

hard exudates

## ❖ Hard exudates

-Caused by the breakdown of the blood-retina barrier, allowing leakage of serum proteins and lipids, from the vessels.



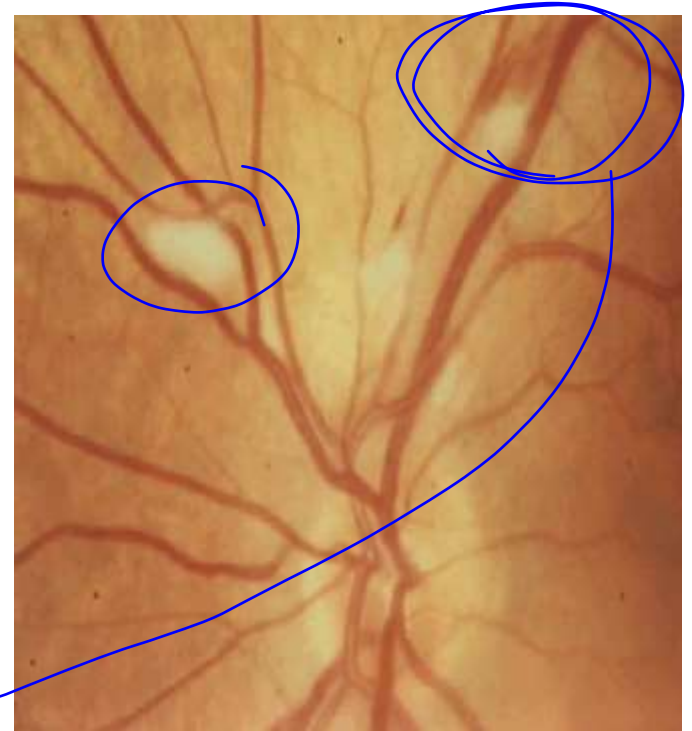
appear as yellow, well-defined  
lesions, usually within  
macular area  
circinate pattern

## ❖ Cotton-wool spots

Nerve fiber layer infarction from occlusion of precapillary arterioles

Fluorescein angiography - No capillary perfusion

→ white/pale, illdefined lesions, <sup>(fluffy)</sup> around optic nerve head.  
Can be associated with flame shaped hemorrhages

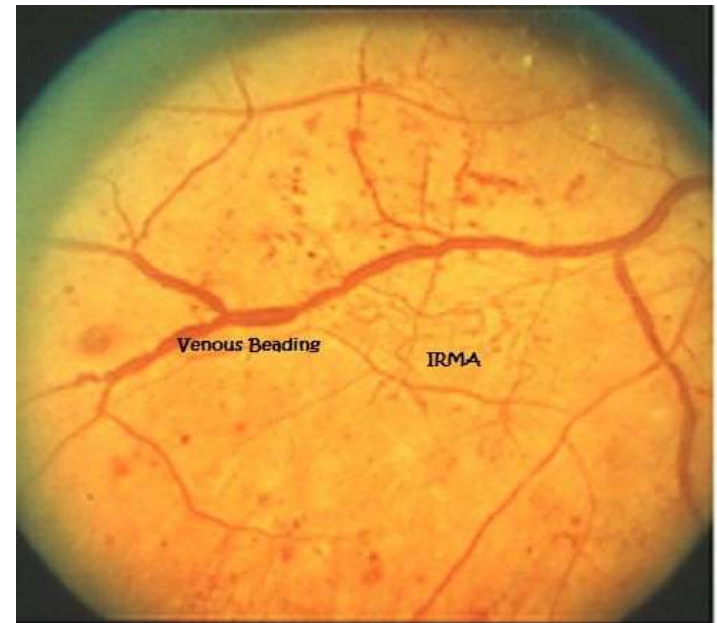


# ❖ Intraretinal microvascular abnormalities

*happens at areas of ischemia  
in addition to venous abnormalities*

abnormal branching,  
sinuous shunt vessels  
that typically develop  
adjacent to areas of capillary  
non perfusion

It is a sign of severe NPDR ~~\*\*\*~~  
*↳ non proliferative*





# Diabetic Macular Edema

presence of edema and/or exudates at macular area

⇒ Diabetic maculopathy



## International Clinical Diabetic Macular Edema (DME) Disease Severity Scale:

- **DME absent:**  
No retinal thickening or hard exudates (HE) present in the posterior pole.
- **DME present:**  
Some retinal thickening or hard exudates (HE) present in the posterior pole.

- If DME present, it can be categorized as follows:

- **Mild DME**:

Some retinal thickening or HE present in the posterior pole but distant from the center of macula.

- **Moderate DME**:

Retinal thickening or HE approaching the center of the macula but not involving its center.

- **Severe DME**:

Retinal thickening or HE involving the center of the macula.

# Clinically significant macular edema

↳ indication for treatment  
if not significant → just observation

the Early Treatment Diabetic Retinopathy Study classification protocol as the presence of:

PHO

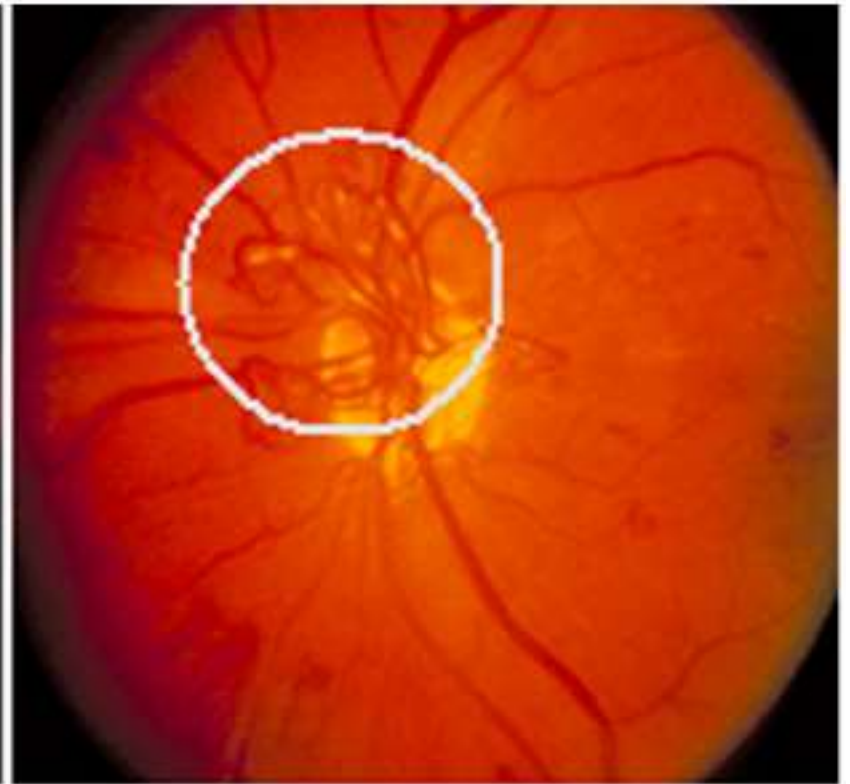
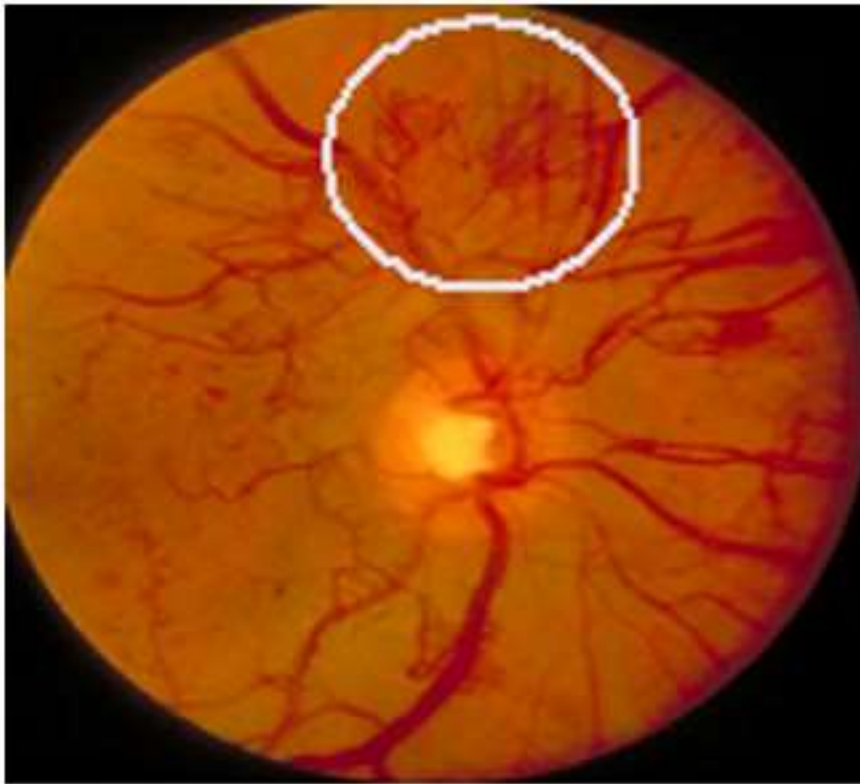
Retina thickening at or within 500  $\mu\text{m}$  from the center of the macula

- Retinal hard exudate at or within 500  $\mu\text{m}$  of the center of the macula if associated with edema.
- Zone of thickening one disc diameter, at least part of which is within one disc from the center of the macula.

## Signs of diabetic retinopathy :

Late signs (signs of proliferative DR) :

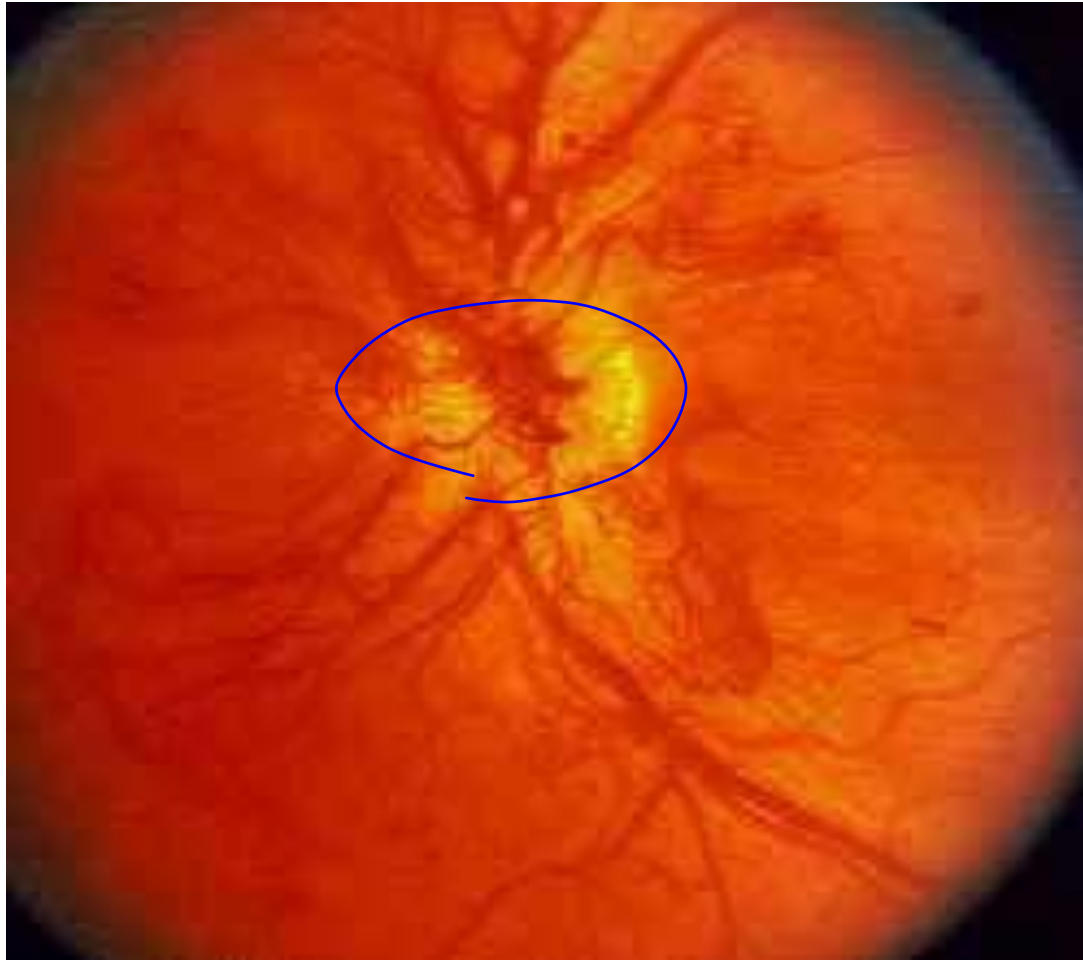
- ❖ Neovasclerization (NVDs, NVEs and NVIs)
- ❖ Vitreous hemorrhages
- ❖ Per retinal hemorrhages (subhyaloid)
- ❖ Traction retinal detachment
- ❖ Neovascular glaucoma



**(a) New vessels elsewhere (NVE)**      **(b) New vessels on disc (NVD)**

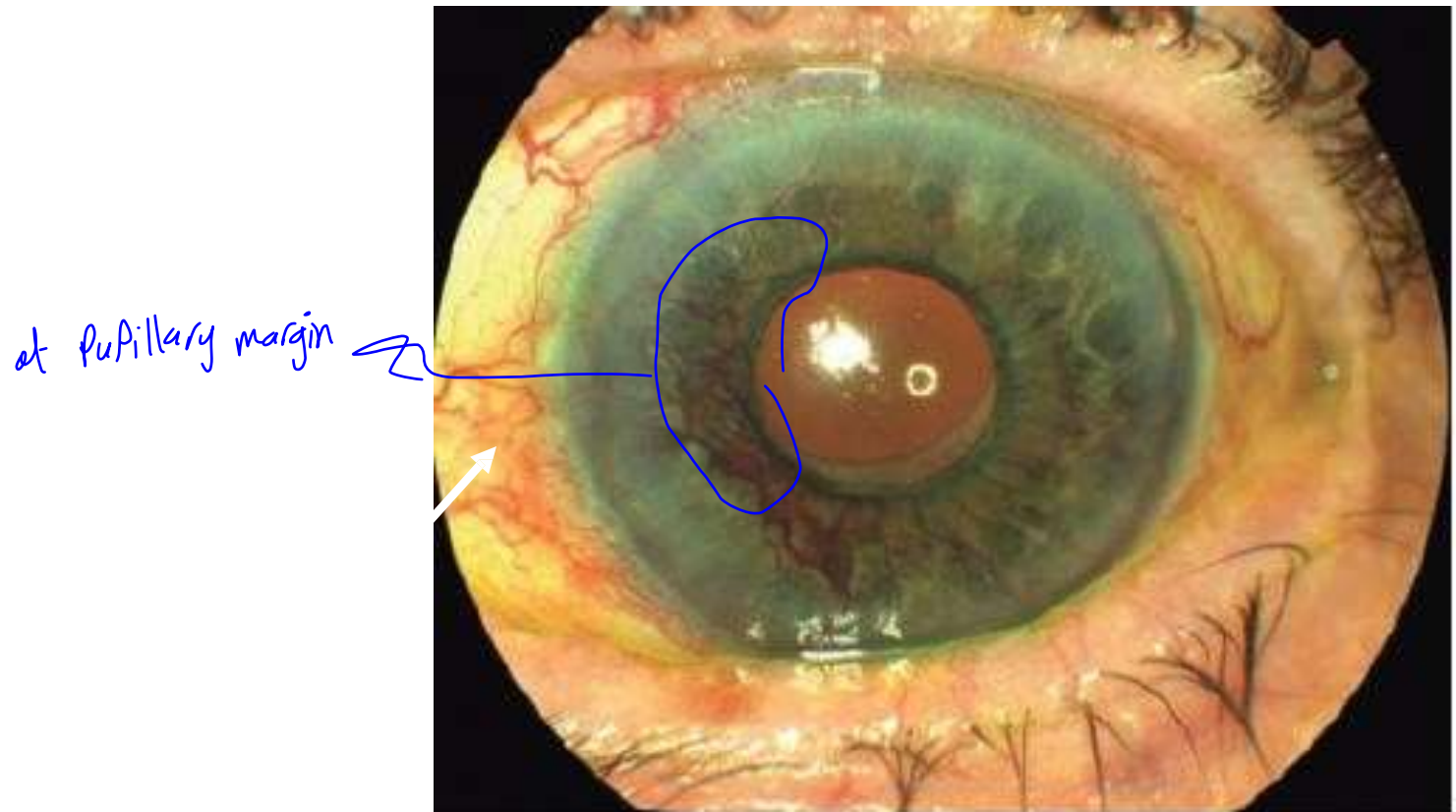
hallmark of PDR

# NVDs



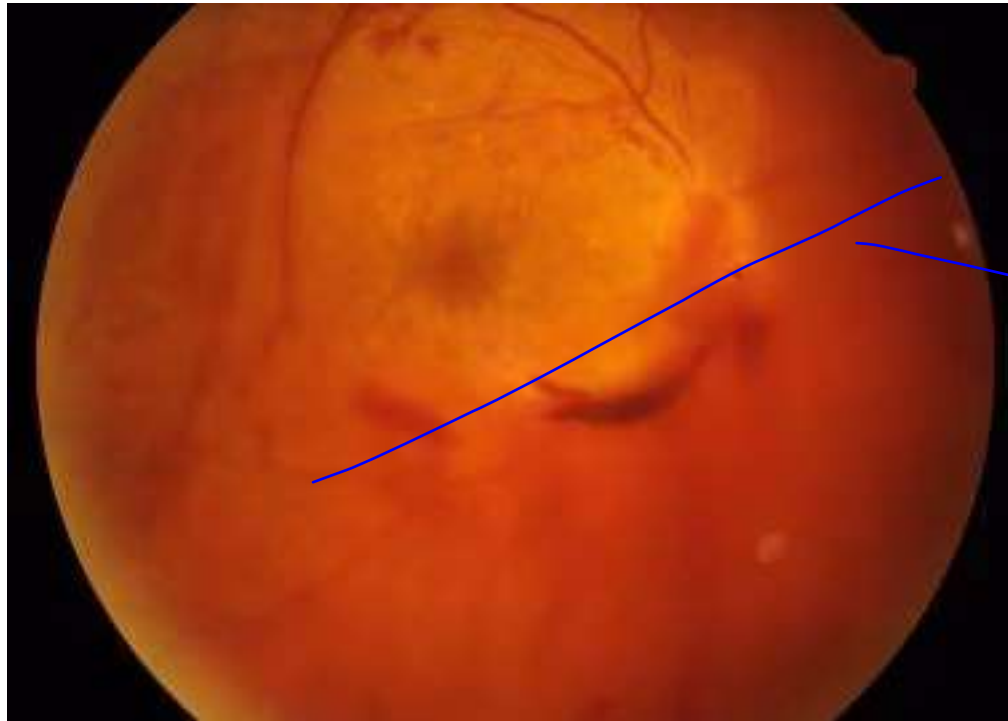
# Rubeosis Iridis

Neovascularization of the iris.





## ❖ Vitreous hemorrhages



above the line we can see the details

from here down details of the retina do not appear → v. hemorrhage inferiorly

comes from NVDs or NVEs when they rupture

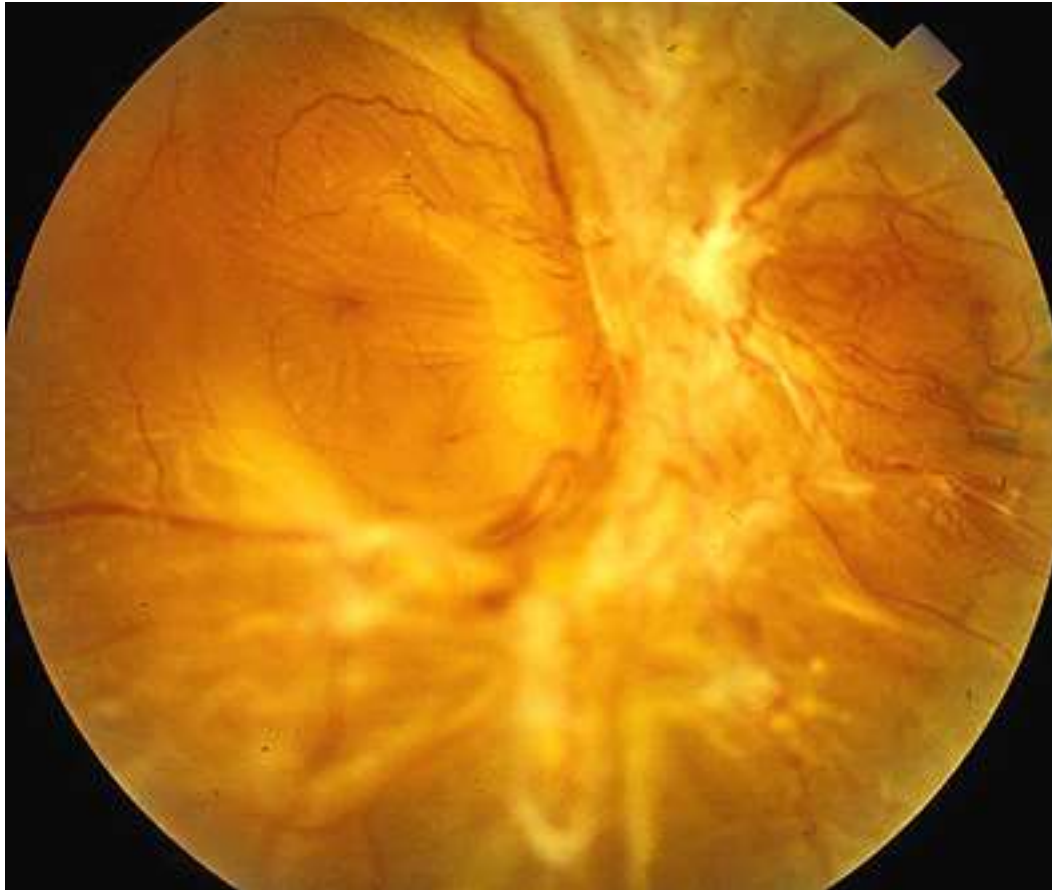
# Pre-retinal or sub-hyloid Hemorrhage

Vitreous is clear

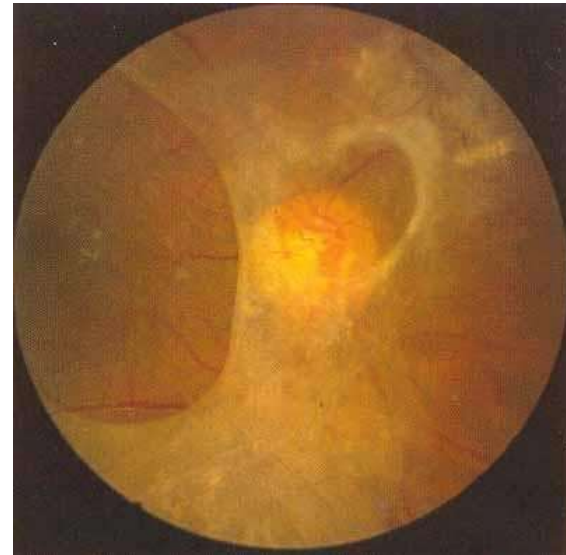
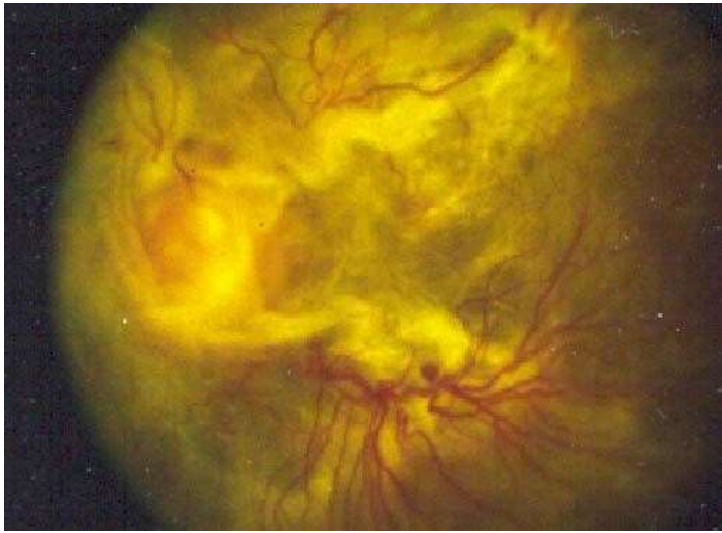
Sign of PDR



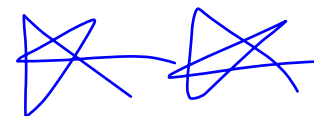
# Tractional retinal detachment



# Tractional retinal detachment



# Classification of the American Academy of Ophthalmologists

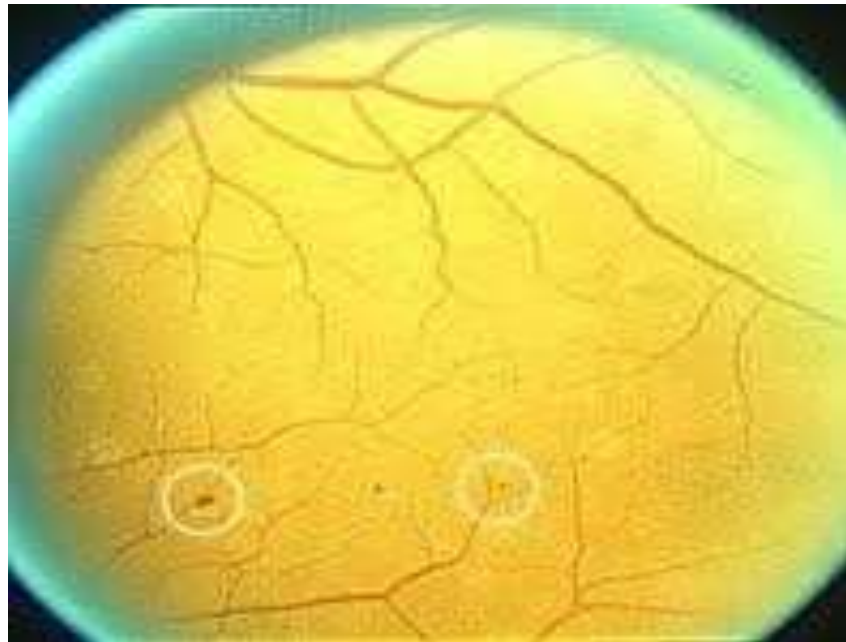


Dilated Ophthalmoscopy Findings	Proposed Disease Severity Level
No abnormalities	<u>No apparent DR</u>
Microaneurysms only	Mild NPDR
More than "mild" but less than "severe"	Moderate NPDR
Any of the following: <i>4 2 1 rule</i> 20 or more microaneurysms in 4 quadrants Definite venous beading in 2 or more quadrants Prominent IRMA in 1 or more quadrants and no neovascularization	Severe NPDR
1 or more of the following: Definite neovascularization Preretinal or vitreous hemorrhage	<u>PDR</u>

**Table 7. International Clinical Diabetic Retinopathy Disease Severity Scale.**

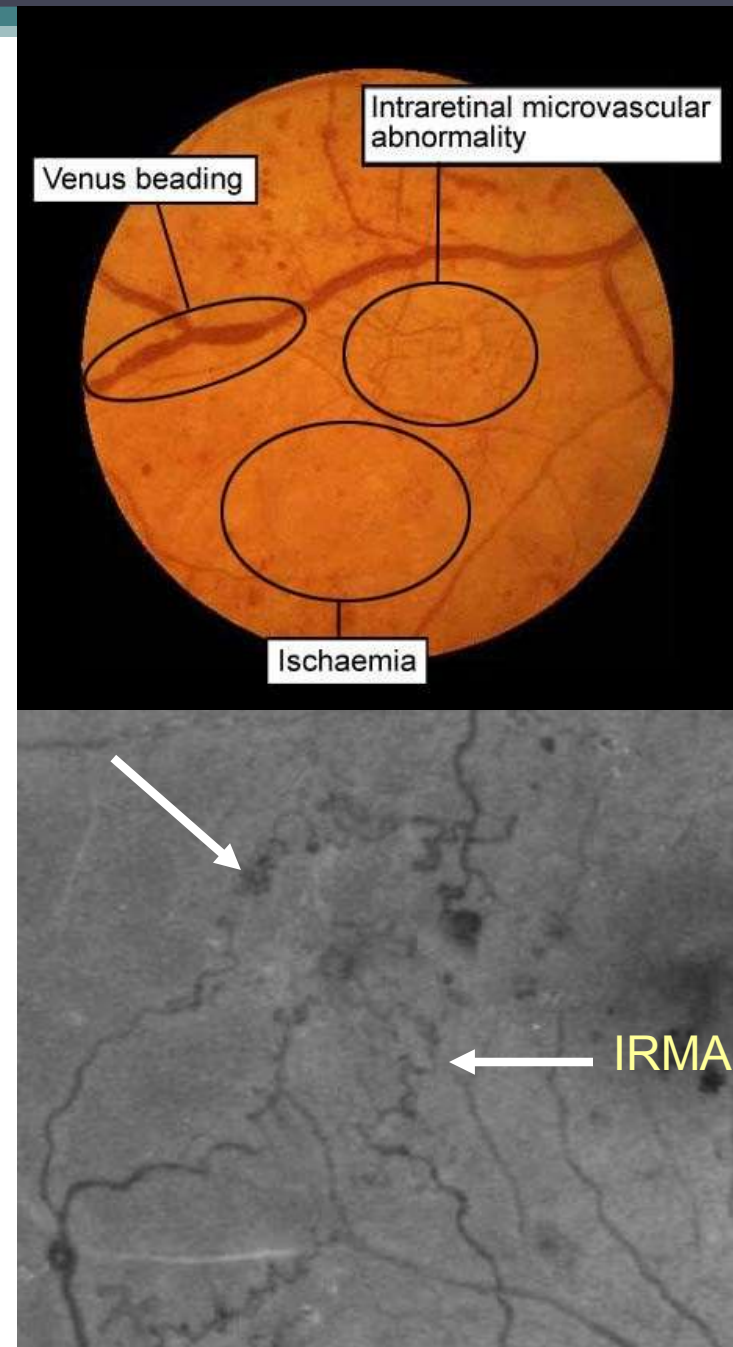
# Mild NPDR

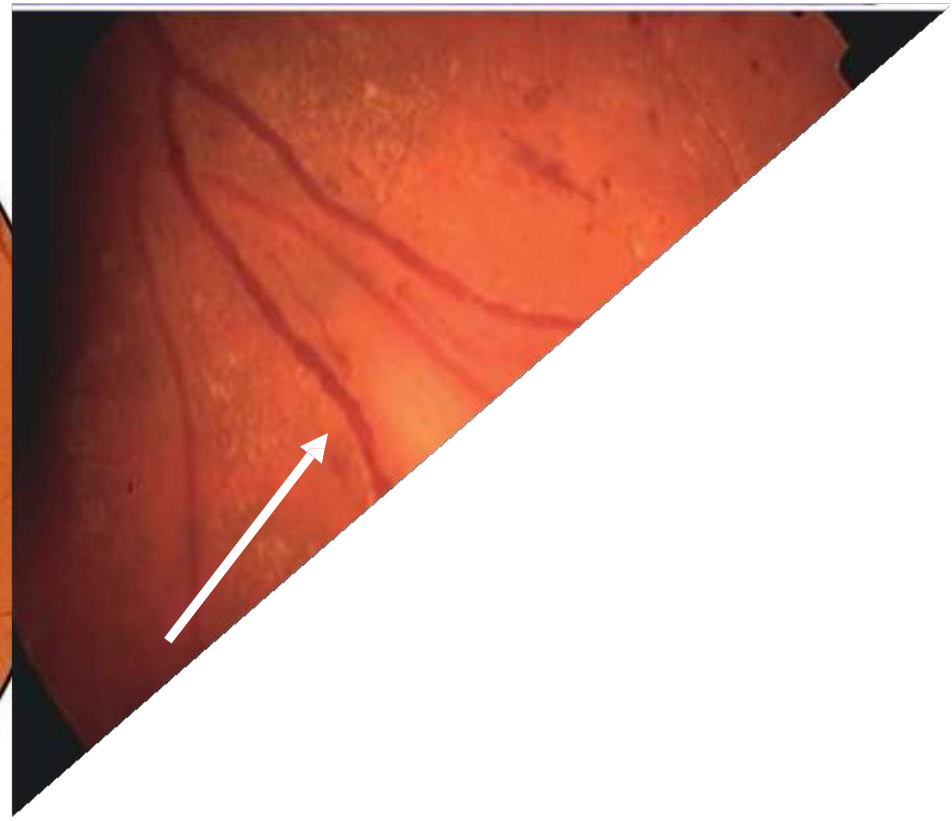
- **Microaneurisms only**
- Earliest clinically detectable lesion



# Moderate NPDR

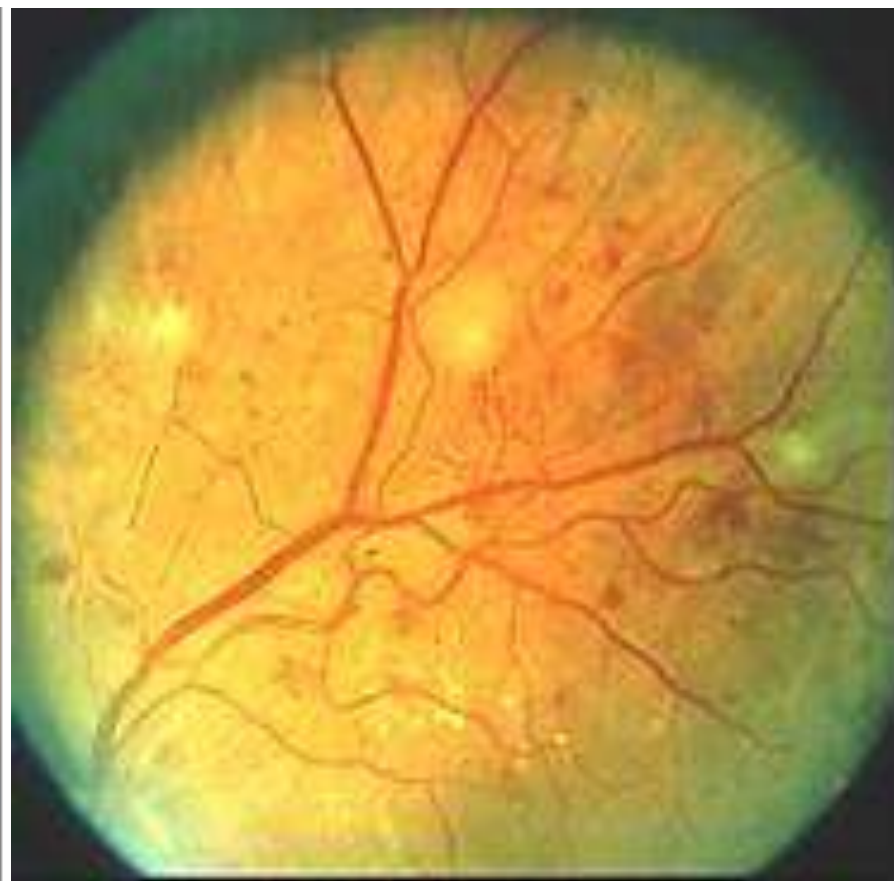
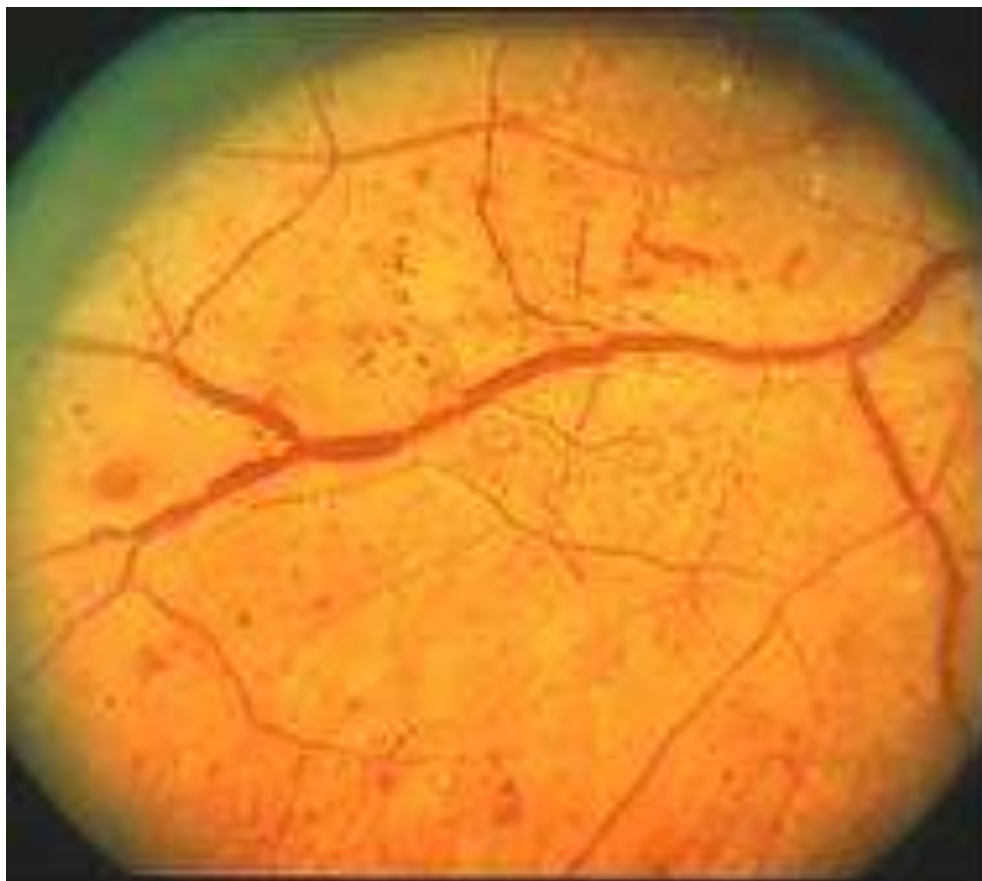
- Microaneurysms and/or dot and blot hemorrhages in more than 1 quadrant.
- Soft exudates (Cotton wool spots).
- Venous beading in one quadrant.





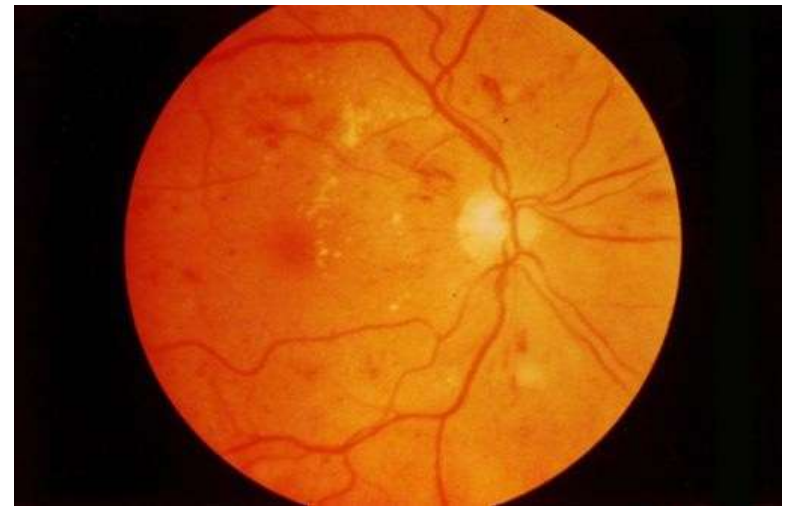
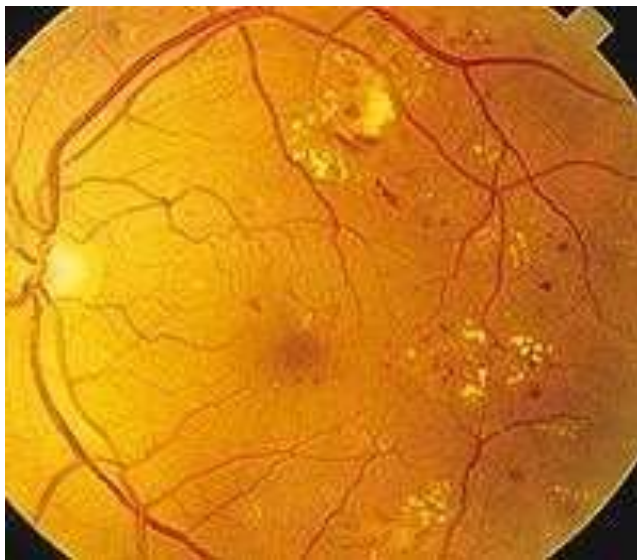
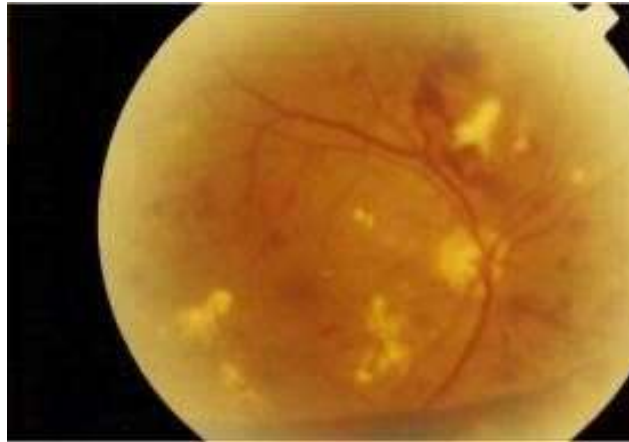


# Mild vs Moderate NPDR



# Severe NPDR

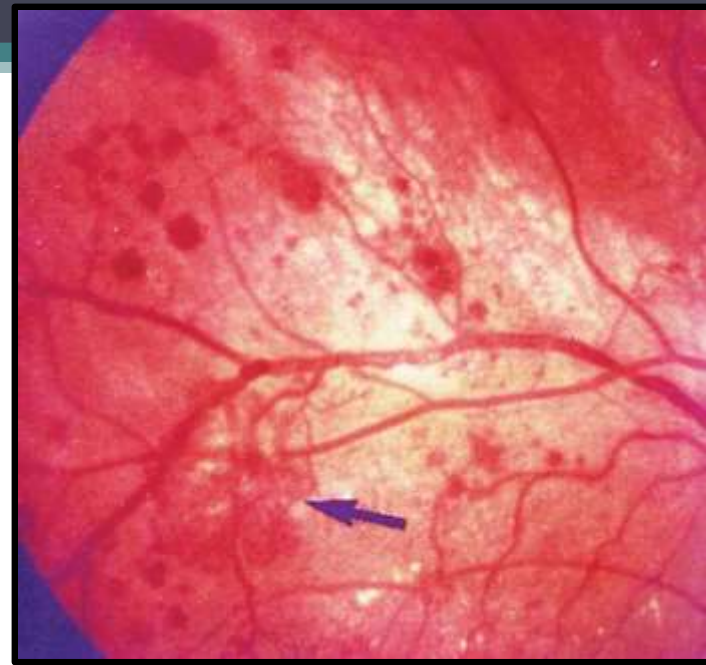
Micro anueysms in 4 quadrants, venous changes in 2 quadrants or IRMA in one quadrant



# Proliferative DR

Characterized by  
Proliferation of new  
vessels from retinal  
veins

- New vessels on the optic disc
- New vessels elsewhere on the retina



## TABLE. RECOMMENDED FOLLOW-UP SCHEDULE FOR DIABETIC PATIENTS

Severity of Retinopathy	Examination Criteria	Follow-up (month)
Normal	No retinopathy	12
Mild NPDR	Microaneurysms only	9
Moderate NPDR	More than microaneurysms but less than severe	6
Severe NPDR	Any of the following (4-2-1 rule)	
• Intraretinal hemorrhages in all four quadrants		4
• Venous beading in two or more quadrants		4
• IRMA in one or more quadrants		4
Proliferative DR	Neovascularization of disc or elsewhere	Refer to retina specialist
Macular edema	Macular thickening and/or cystic edema	Refer to retina specialist

*Abbreviations: NPDR, nonproliferative diabetic retinopathy; IRMA, intraretinal microvascular abnormalities; DR, diabetic retinopathy.*

# Treatment

Indications? PDR or clinically significant macular edema or both + complications

## ❖ Mild & Moderate NPDR

- No specific treatment for retinopathy
- Good diabetic control to delay progression
- Control of associated Hypertension, Anemia and Renal failure

## ❖ Severe NPDR

- Close follow up by Ophthalmologist

## ❖ Clinically Significant Macular Edema

- Intra-vitreous anti-VEGF.
- **Laser photocoagulation** to minimize risk of visual loss.

We aim the laser at the points of leakage, the exudate is often seen as to be in a circular or circinate pattern, with the focus of leakage or microaneurysm in the middle. If the treatment is effective, the retinal edema and exudate will resorb, although this may take some months.

# Circinate retinopathy - Hard exudates in a ring around leaking aneurysms



## ❖ Proliferative DR

↪ *mostly used*  
—Retinal **laser photocoagulation** as per the judgment of ophthalmologist (in high risk eyes) , it improves retinal circulation and decreases production of vasoproliferative factors (by ablating areas of ischemic retina).

Our aim here is scattered laser burns to the entire retina (pan-retinal laser pr PRP), leaving an untreated area around the optic disc and around the central region of the macula, to preserve vision.

—**Anti-VEGF** , shrinks neovascularization and decrease leakage, given as intravitreal injection, like avastin.





Leaking  
blood vessel

© JirehDesign.com

Diabetic retinopathy typically presents no symptoms during the early stages.

The condition is often at an advanced stage when symptoms become noticeable. On occasion, the only detectable symptom is a sudden and complete loss of vision.

DR usually affects both eyes.

The only way people with diabetes can prevent DR is to attend every eye examination scheduled by their doctor.

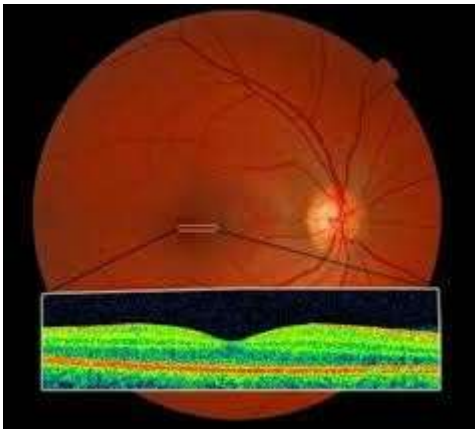
# Symptoms of diabetic retinopathy may include:

- Blurred vision
- The impairment of color vision
- Floaters, or transparent and colorless spots and dark strings that float in the patient's field of vision
- Patches or streaks that block the person's vision
- Poor night vision
- Sudden and total loss of vision

# Investgations

HbA1c, blood sugar

OCT, to determine the thickness, presence of swelling, to diagnose macular edema or CSME



Fluorescein angiography



# Complications of Diabetic Retinopathy

- Vitreous hemorrhage
- Tractional retinal detachment
- Rubeosis Iridis
- Glaucoma
- Blindness

# Neovascular Glaucoma

- Complication of rubeosis iridis
- New vessels cause angle closure
- Mechanical obstruction to aqueous outflow
- Intra ocular pressure rises
- Pupil gets distorted as iris gets pulled.
- Eye becomes painful and red
- Loss of vision

# Blindness

- Non-clearing vitreous hemorrhage
- Neovascular glaucoma
- Tractional retinal detachment
- Macular ischemia

# Prevention of Complications

- 1- By early institution of appropriate treatment
- 2- Early detection of DR in its asymptomatic treatable condition
3. Routine fundus examination of all diabetics (at least yearly)
4. Appropriate referral to ophthalmologist