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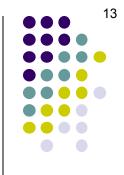
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What proportion of diabetics receive screening eye exams at recommended intervals? Only about 60%



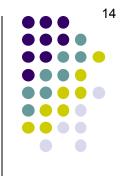


Worldwide, what proportion of diabetics have retinopathy?





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Of those diabetics with retinopathy, what proportion have vision-threatening retinopathy?





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Diabetic Retinopathy: The Basics



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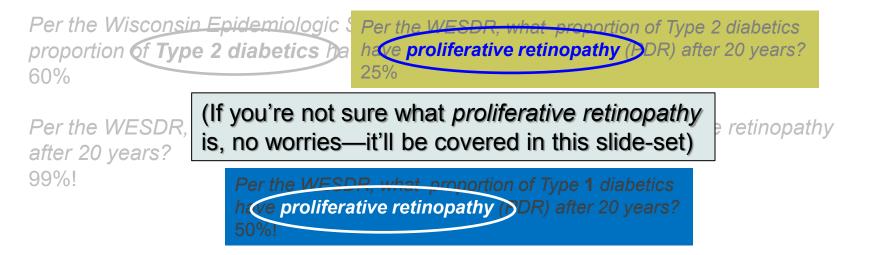
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Diabetes also causes CAD, CVA, nephropathy, and neuropathy. (Thanks Captain Obvious.) However, only one has on onset that is strongly correlated with the onset of retinopathy. Which one?





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Diabetes also causes CAD, CVA, nephropathy, and neuropathy. (Thanks Captain Obvious.) However, only one has on onset that is strongly correlated with the onset of retinopathy. Which one? Nephropathy—when a pt develops one of the two, it's a lock s/he's going to develop the other very soon

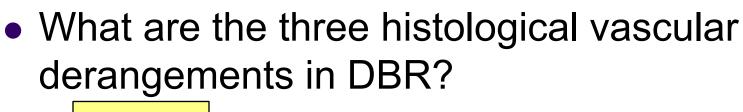


What are the three histological vascular derangements in DBR?
 1)
 2)

3)











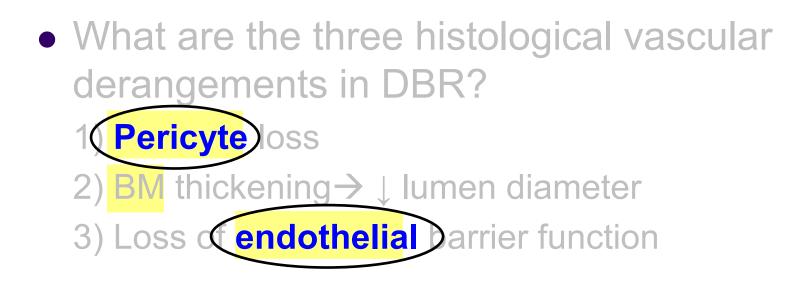


 What are the three histological vascular derangements in DBR?

- 1) Pericyte loss BM = Basement membrane
- 2) BM thickening $\rightarrow \downarrow$ lumen diameter
- 3) Loss of endothelial barrier function



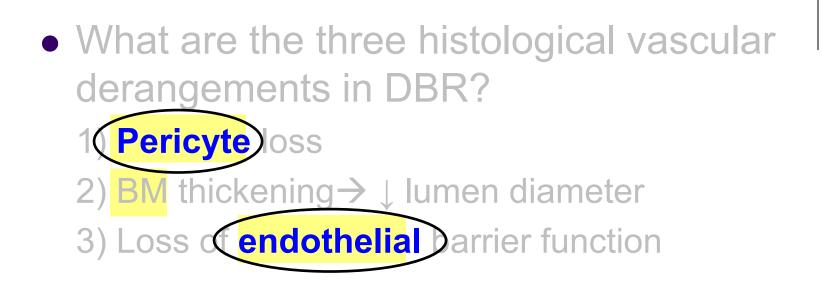




With respect to the structure of retinal arterioles and capillaries, how are pericytes and endothelial cells related to one another?





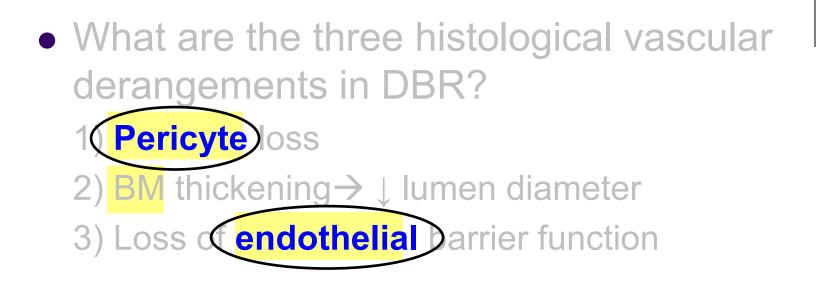


With respect to the structure of retinal arterioles and capillaries, how are pericytes and endothelial cells related to one another?

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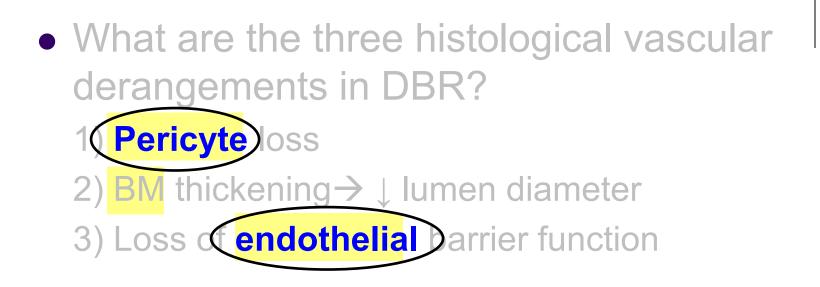




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With respect to the structure of retinal arterioles and capillaries, how are pericytes and endothelial cells related to one another? The endothelial cells line the lumen of the vessel. They are surrounded by their BM. They are <u>fenestrated or non-</u>.



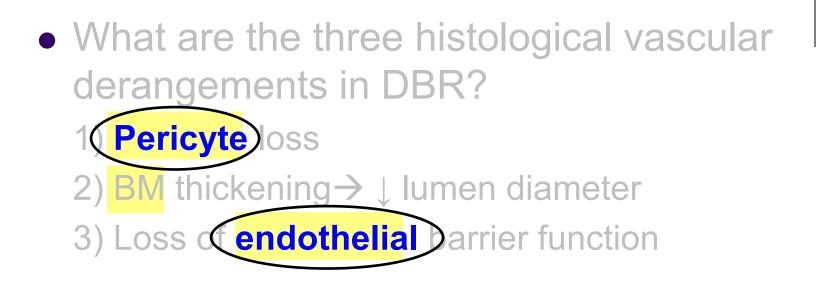




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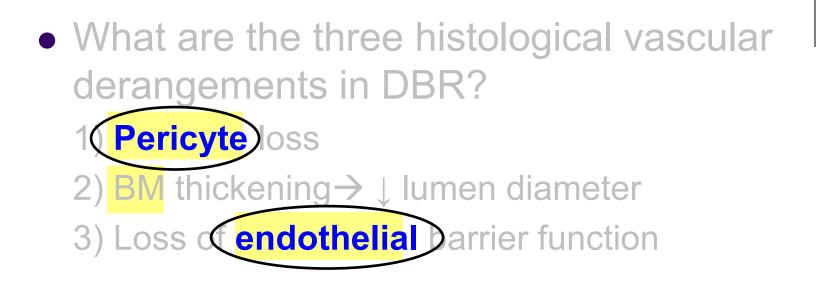


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four words







With respect to the structure of retinal arterioles and capillaries, how are pericytes and endothelial cells related to one another? The endothelial cells line the lumen of the vessel. They are surrounded by their BM. They are nonfenestrated . Tight junctions between cells form the so-called *inner blood-retina barrier*.







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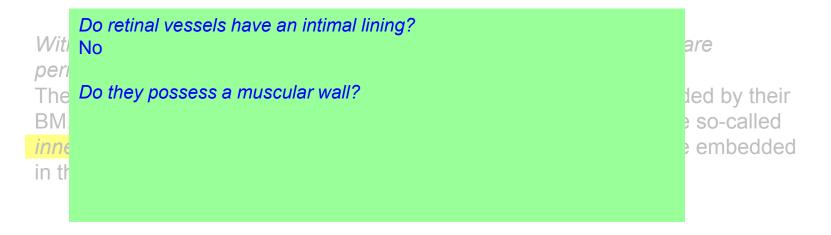




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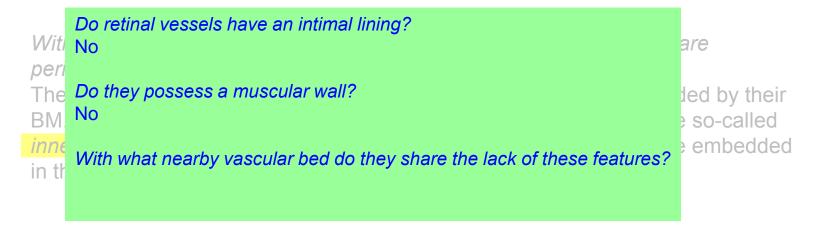




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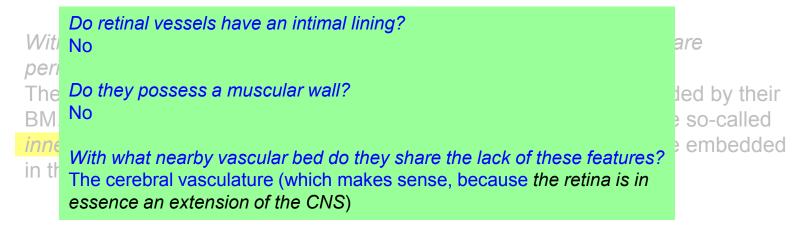
















What are the three histological vascular derangements in DBR?
1) Pericyte loss
2) BM thickening -> | lumen diameter
3) Loss of endothelial barrier function

That this is known as the inner blood-retine barrier implies the existence of what?

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BM. They are nontenestrated . Tight junctions between cells form the so-called **inner blood-retina barrier**. The pericytes surround the vessel, and are embedded in the BM of the endothelial cells.





An outer blood-retina barrier

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Yup. What forms the outer blood-retina barrier?

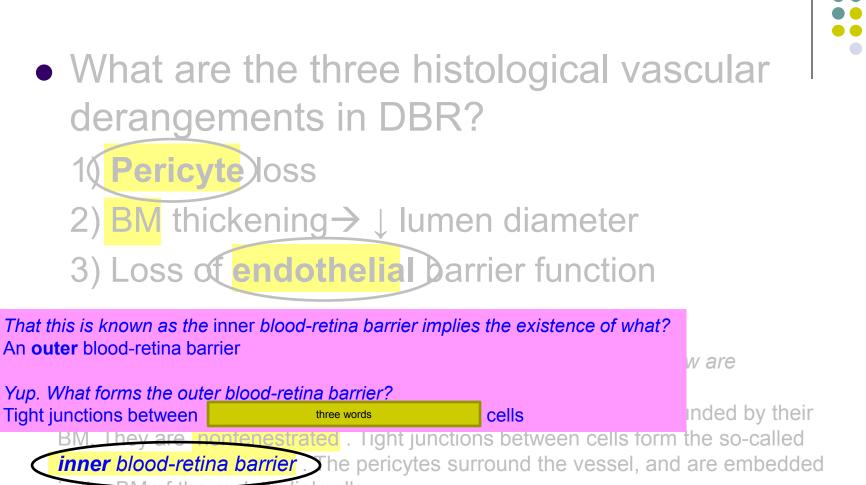
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Diabetic Retinopathy: The Basics



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That this is known as the inner *blood-retina barrier implies the existence of what?* An **outer** blood-retina barrier

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Yup. What forms the outer blood-retina barrier? Tight junctions between retinal pigment epithelium (RPE) cells

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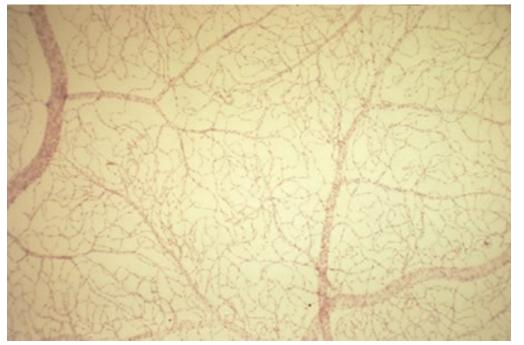


1) Pericyte loss

2) BM thickening $\rightarrow \downarrow$ lumen diameter

Which occurs first? Pericyte loss

3) Loss of endothelial barrier function

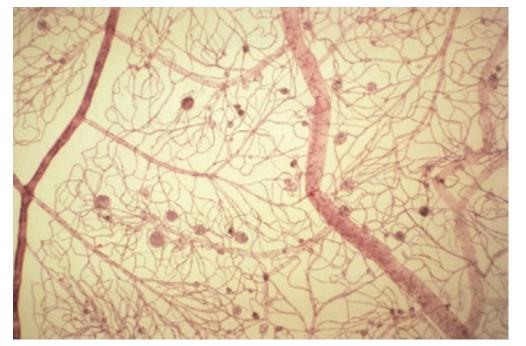




Trypsin mount of normal retina-low and high mag

The dark nuclei belong to pericytes; the lighter, to endothelial cells. *Note that the ratio between them is roughly 1:1.*



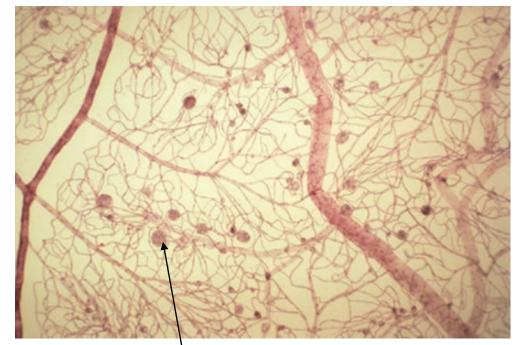




Trypsin mount of DBR retina-low and high mag

But in a retina with damage 2ndry to diabetes, the ratio of endothelial cells to pericytes is many-to-one.



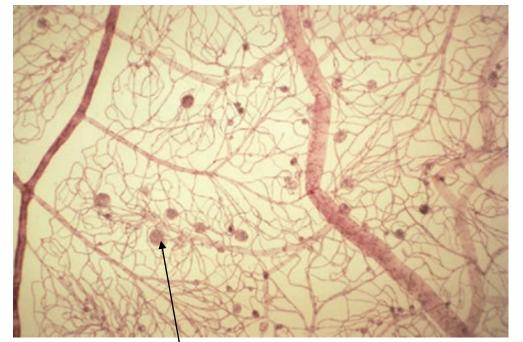




Trypsin mount of DBR retina-low and high mag

What are these things?

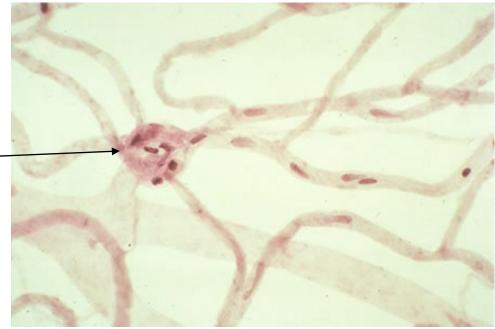






Trypsin mount of DBR retina-low and high mag

What are these things? Microaneurysms





 What are the three histological vascular derangements in DBR?
 Derivate lass

1) Pericyte loss

2) BM thickening → ↓ lumen diameter

What pathological state is the endpoint of decreasing lumen diameter?





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What pathological state is the endpoint of decreasing lumen diameter? Occlusion of the retinal vessel





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Vessel occlusion leads to what pathological event?





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Retinal ischemia leads to what pathological state?





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Hypoxic retinal cells release a signaling molecule that is central to the pathogenesis of DBR. What is that signaling molecule?





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Hypoxic retinal cells release a signaling molecule that is central to the pathogenesis of DBR. What is that signaling molecule? **VEGF** (we will have much more to say about VEGF shortly)





What are the three histological vascular derangements in DBR?
 1) Pericyte loss
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Loss of endothelial barrier function leads to what pathologic event?





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Loss of endothelial barrier function leads to what pathologic event? Leaching of serum into the retina





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Loss of endothelial barrier function leads to what pathologic event? Leaching of serum into the retina

Leaching of serum into the retina leads to what pathological state?





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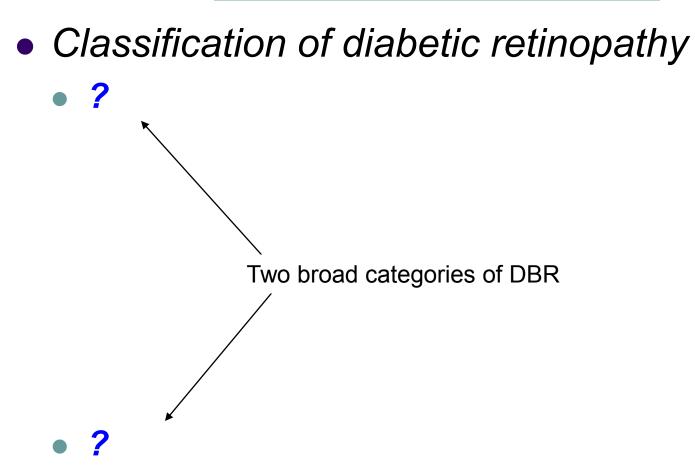
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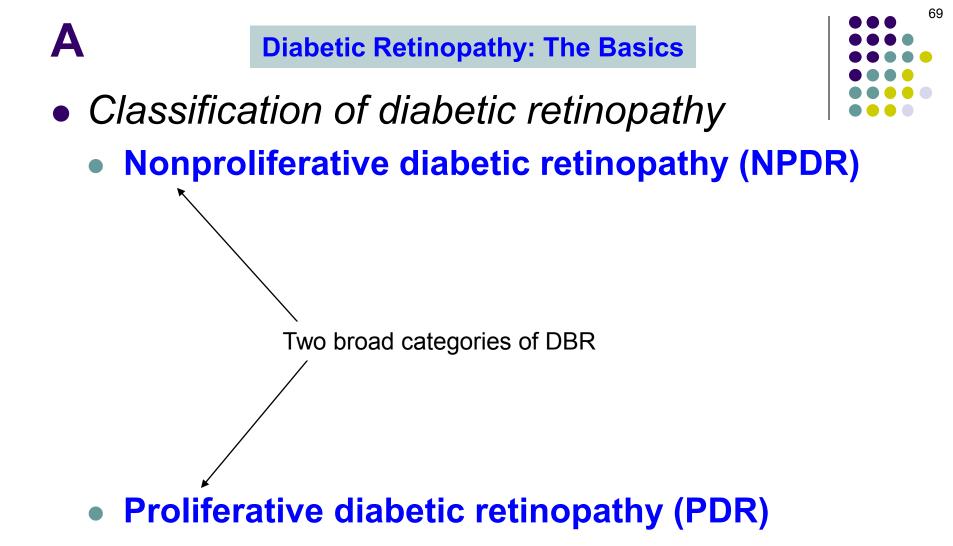
Leaching of serum into the retina leads to what pathological state? Retinal edema

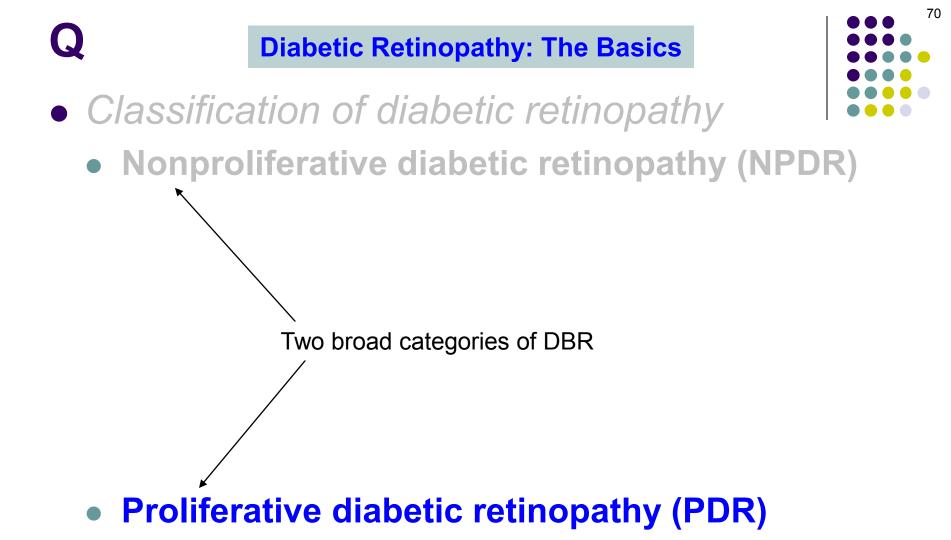












What is the histological definition of proliferation in this context?



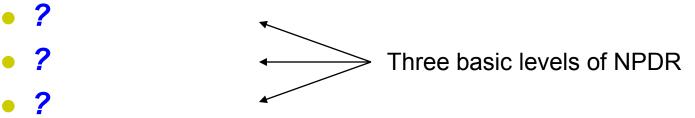
Proliferative diabetic retinopathy (PDR)

What is the histological definition of proliferation in this context? Retinal neovascularization that breaks through the internal limiting membrane (ILM)



• Classification of diabetic retinopathy

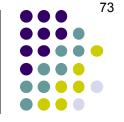




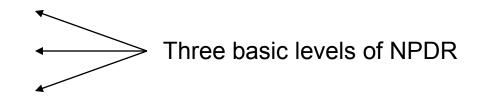
• Proliferative diabetic retinopathy (PDR)



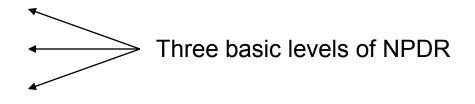




- Nonproliferative diabetic retinopathy (NPDR)
 - Mild
 - Moderate
 - Severe



- Classification of diabetic retinopathy
 - Nonproliferative diabetic retinopathy (NPDR)
 - Mild
 - Moderate
 - Severe

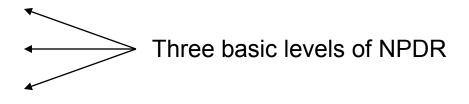


? One more level (not universally used)





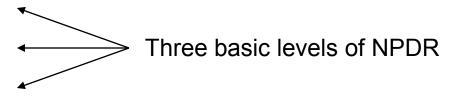
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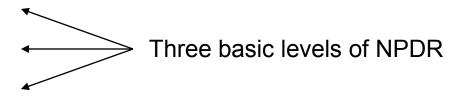


- Proliferative diabetic retinopathy (PDR)
 - ? One level of concern





- Classification of diabetic retinopathy
 - Nonproliferative diabetic retinopathy (NPDR)
 - Mild
 - Moderate
 - Severe



- Proliferative diabetic retinopathy (PDR)
 - *High-risk PDR* ← One level of concern





- Classification of diabetic retinopathy
 - Nonproliferative diabetic retinopathy (NPDR)
 - Mild
 - Moderate
 - Severe
 - Very severe

What landmark clinical trial provided this system of DBR classification?

- Proliferative diabetic retinopathy (PDR)
 - High-risk PDR



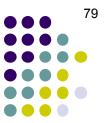


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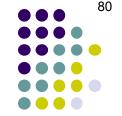
What landmark clinical trial provided this system of DBR classification? The Early Treatment of Diabetic Retinopathy Study. Note that the ETDRS is one of the studies you are expected to be familiar with by name.

Proliferative diabetic retinopathy (PDR)

• High-risk PDR



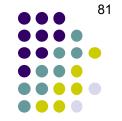




- Nonproliferative diabetic retinopathy (NPDR)
 - *Mild How are mild and moderate NPDR defined?*
 - Moderate
 - Severe
 - Very severe

- Proliferative diabetic retinopathy (PDR)
 - High-risk PDR





Nonproliferative diabetic retinopathy (NPDR)

How are mild and moderate NPDR defined?

With respect to the standard photographs

- Mild
- Moderate
- Severe
- Very severe

Proliferative diabetic retinopathy (PDR)

employed in the DRS

• High-risk PDR





- Nonproliferative diabetic retinopathy (NPDR)
 - Mild:
 definition
 - Moderate
 - Severe
 - Very severe

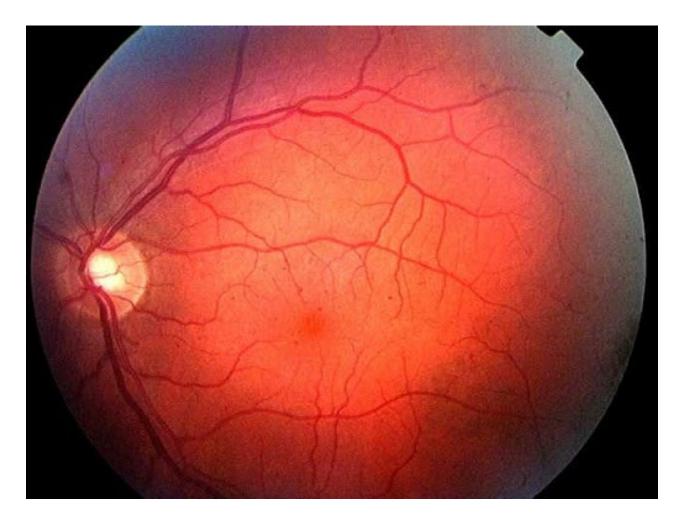
- Proliferative diabetic retinopathy (PDR)
 - High-risk PDR





- Nonproliferative diabetic retinopathy (NPDR)
 - *Mild:* Any DBR < *moderate*
 - Moderate
 - Severe
 - Very severe

- Proliferative diabetic retinopathy (PDR)
 - High-risk PDR



Mild nonproliferative diabetic retinopathy







Nonproliferative diabetic retinopathy (NPDR)

definition

- Mild: Any DBR < moderate</p>
- Moderate:
- Severe
- Very severe

- Proliferative diabetic retinopathy (PDR)
 - High-risk PDR





- Nonproliferative diabetic retinopathy (NPDR)
 - Mild: Any DBR < moderate</p>
 - Moderate: DBR > mild but < severe</p>
 - Severe
 - Very severe

- Proliferative diabetic retinopathy (PDR)
 - High-risk PDR





Moderate nonproliferative diabetic retinopathy





Nonproliferative diabetic retinopathy (NPDR)

definition

- Mild: Any DBR < moderate</p>
- Moderate: DBR > mild but < severe</p>
- Severe:



- Proliferative diabetic retinopathy (PDR)
 - High-risk PDR



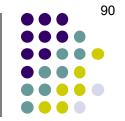


- Nonproliferative diabetic retinopathy (NPDR)
 - Mild: Any DBR < moderate</p>
 - Moderate: DBR > mild but < severe</p>
 - Severe: Presence of any 1 of the 4:2:1 rule
 - Very severe

- Proliferative diabetic retinopathy (PDR)
 - High-risk PDR



• Classification of diabetic retinopathy



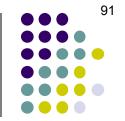
- Nonproliferative diabetic retinopathy (NPDR)
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chance of bigh rick DDD within 1 w What is the 4:2:1 rule?

• Very severe



• Classification of diabetic retinopathy



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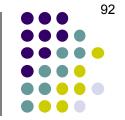
• Very severe --4 retinal quadrants of...

--2 retinal quadrants of...

--1 retinal quadrant of...



• Classification of diabetic retinopathy

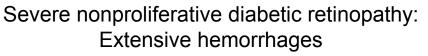


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- Very severe --4 retinal quadrants of...extensive retinal hemorrhages
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 -1 retinal quadrant of...
- Proliferative diabetic retinopathy (PDR)

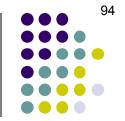








• Classification of diabetic retinopathy



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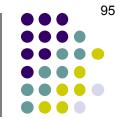


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• Classification of diabetic retinopathy



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• Very severe --4 retinal quadrants of...extensive retinal hemorrhages --2 retinal quadrants of...venous beading

--1 retinal quadrant of...

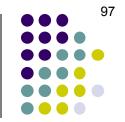


Severe nonproliferative diabetic retinopathy: Venous beading





• Classification of diabetic retinopathy



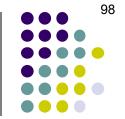
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chance of bigh rick DDD within 1 What is the 4:2:1 rule?

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• Classification of diabetic retinopathy



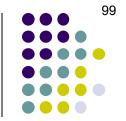
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chance of bigh rick DDD within 1 What is the 4:2:1 rule?

Very severe --4 retinal quadrants of...extensive retinal hemorrhages
 --2 retinal quadrants of...venous beading
 --1 retinal quadrant of...IRMA



• Classification of diabetic retinopathy



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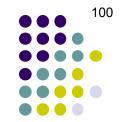
What does **IRMA** stand for?

D within 1 w

ts of ...extensive retinal hemorrhages ts of ...extensive beading t of ...IRMA



• Classification of diabetic retinopathy



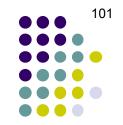
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What does **IRMA** stand for? Intraretinal microvascular anomalies

rule? Its of...extensive retinal hemorrhages Its of venous beading It of ...IRMA



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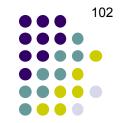
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• Classification of diabetic retinopathy



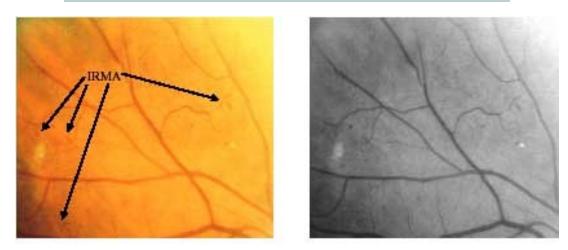
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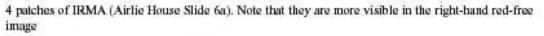
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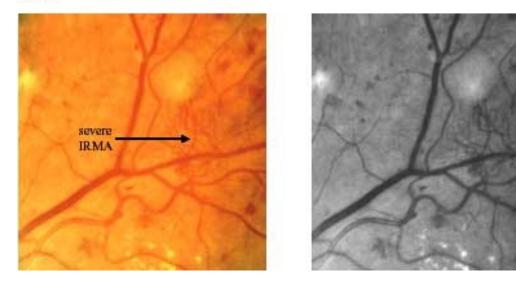
What does that mean? Think of it as neovascularization that has **not** broken through the ILM rule? Its of...**extensive retinal hemorrhages** Its of venous beading It of ...IRMA

• Proliferative diabetic retinopathy (PDR)

IRMA What is the histological definition of ^Vproliferation in this context? Retinal neovascularization that, breake through the internal limiting membrane (ILM) hasn't broken



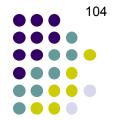




Severe nonproliferative diabetic retinopathy: IRMA





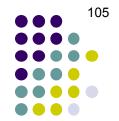


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definition

- Proliferative diabetic retinopathy (PDR)
 - High-risk PDR





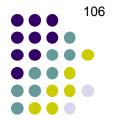
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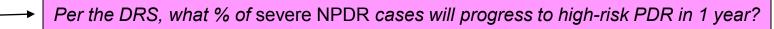


Next Q

Classification of diabetic retinopathy



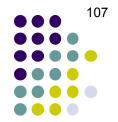
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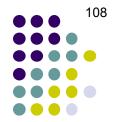
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- Proliferative diabetic retinopathy (PDR)
 - High-risk PDR



Next Q

Classification of diabetic retinopathy

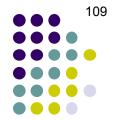


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What % of **very** severe NPDR cases will progress to high-risk PDR in 1 year?

- Proliferative diabetic retinopathy (PDR)
 - High-risk PDR





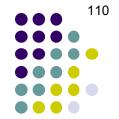
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45% chance of high-risk PDR within 1 year

- Proliferative diabetic retinopathy (PDR)
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Q

Classification of diabetic retinopathy



- Nonproliferative diabetic retinopathy (NPDR)
- Mild: How should NPDR be managed?





- Nonproliferative diabetic retinopathy (NPDR)
 - Mild: Any DBR < moderate</p>

How should NPDR be managed? There is a clear role for controlling three systemic risk factors: Α

Classification of diabetic retinopathy



- Nonproliferative diabetic retinopathy (NPDR)
 - Mild: Any DBR < moderate</p>
 - How should NPDR be managed? There is a clear role for controlling three systemic risk factors:
 - --Blood glucose --Blood pressure --Lipid profile

• Pr

- Classification of diabetic retinopathy
 - Nonproliferative diabetic retinopathy (NPDR)

113

- Mild: Any DBR < moderate</p>
- *How should NPDR be managed?* There is a clear role for controlling three systemic risk factors:
- --Blood glucose
 --Blood pressure

- --Lipid profile
- What's **less** clear (at the time of this writing) is the role of two modalities that have shown considerable potential:





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- Pr



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 - -Blood glucose
 - --Blood pressure
 - --Lipid profile

- What's **less** clear (at the time of this writing) is the role of two modalities that have shown considerable potential:
 - --Intravitreal anti-VEGF injections
 - --Intravitreal steroids
- There is good clinical-trial data demonstrating that these interventions can lessen the severity of NPDR--substantially so in some cases. What is uncertain at this time is *whether the cost/benefit ratio of these interventions is favorable enough to warrant mandating their use.* (Trials addressing this issue are ongoing.)





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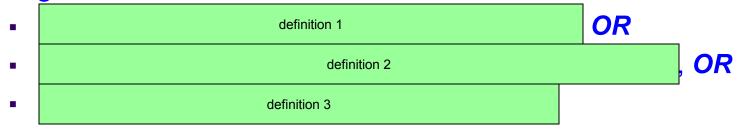
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Proliferative diabetic retinopathy (PDR)

• High-risk PDR







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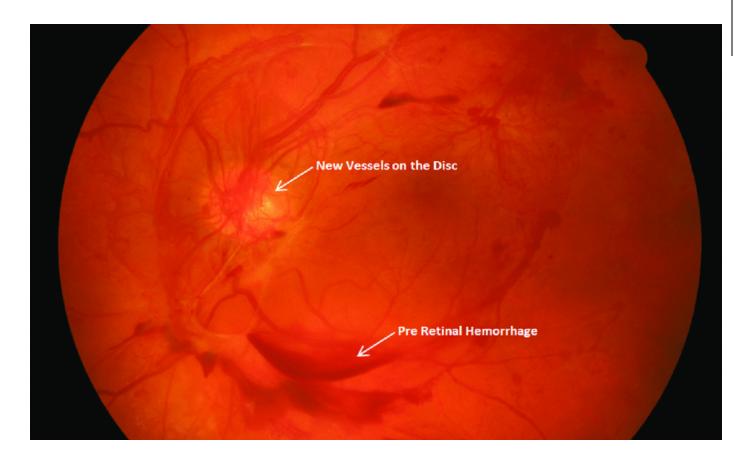
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Proliferative diabetic retinopathy (PDR)

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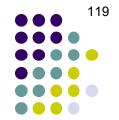
NVD = Neovascularization of the disc	Any NVD associated with vitreous heme (VH), OR		_
	definition 2		, O R
	definition 3		-





High-risk proliferative diabetic retinopathy: NVD + vitreous hemorrhage





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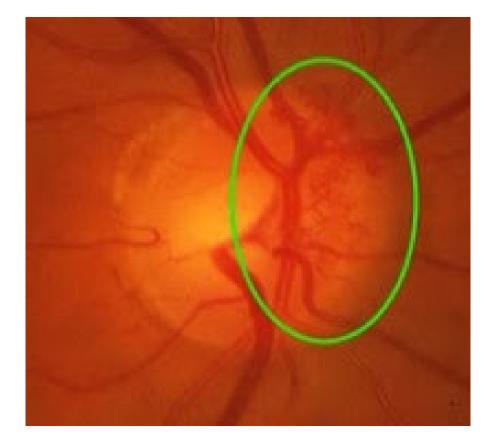
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Proliferative diabetic retinopathy (PDR)

- High-risk PDR
 - Any NVD associated with vitreous heme (VH), OR
 - Large (at least ¼ DD) area of NVD with or without VH, OR

DD = Disc diameter

definition 3



High-risk proliferative diabetic retinopathy: Extensive NVD







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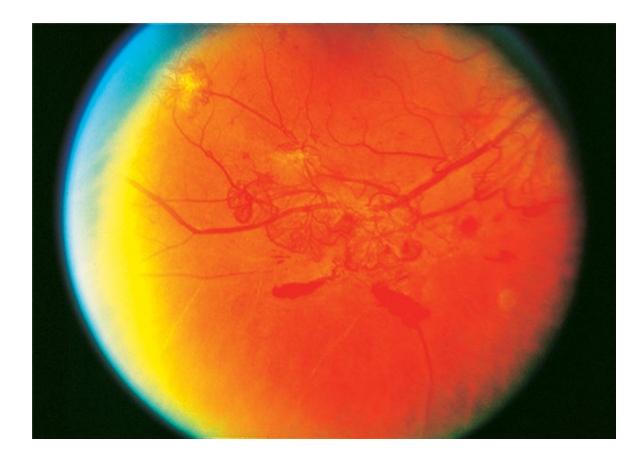
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Proliferative diabetic retinopathy (PDR)

- High-risk PDR
 - Any NVD associated with vitreous heme (VH), OR
 - Large (at least ¼ DD) area of NVD with or without VH, OR
 - Large (at least ½ DD) area of NVE with VH

NVE = Neovascularization elsewhere (ie, anywhere but the disc)



High-risk proliferative diabetic retinopathy: Large area NVE + associated vitreous heme





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(at least ½ DD)a

How big is a DD in microns?







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How big is a DD in microns? Jt V 1500 (1.5 mm)



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Circling back for a minute...We said that PDR consists of retinal neovascularization. What sequence of events leads to retinal neovascularization?



is the histological definition of proliferation in this context?



Classification of diabetic retinopathy

What pathological state is the endpoint of decreasing lumen diameter? Occlusion of the retinal vessel (No question—proceed when ready)

Vessel occlusion leads to what pathological event? Ischemia of the retinal area serviced by the vessel

Retinal ischemia leads to what pathological state? Hypoxia of the affected retinal cells

Hypoxic retinal cells release a signaling molecule that is central to the pathogenesis of DBR. What is that signaling molecule? **VEGF** (we will have much more to say about VEGF shortly)

Circling back for a minute...We said that PDR consists of retinal neovascularization. What sequence of events leads to retinal neovascularization? The answer can be found in this set of questions/answers from earlier in the slide-set:



Retinal neovascularization that breaks through the internal limiting membrane (ILM).





Classification of diabetic retinopathy

What pathological state is the endpoint of decreasing lumen diameter? Occlusion of the retinal vessel

(No question—proceed when ready)

127

Vessel occlusion leads to what pathological event?

To summarize: Occlusive vasculopathy secondary to diabetic derangements produces retinal ischemia.

VEGF we will have much more to say about VEGF shortly)

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• Classification of diabetic retinopathy

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128

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Classification of diabetic retinopathy

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129

Vessel occlusion leads to what pathological event?

To summarize: Occlusive vasculopathy secondary to diabetic derangements produces retinal ischemia. In a desperate attempt to recruit a blood supply, hypoxic retinal cells release **VEGF**, which diffuses throughout the vitreous cavity promoting neovascularization. Unfortunately, the resulting new fibrovascular tissue is highly abnormal—it is prone to bleeding and contraction, leading to vitreous hemorrhages and/or tractional retinal detachment.

VEGF we will have much more to say about VEGF shortly)

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To summarize: Occlusive vasculopathy secondary to diabetic derangements produces retinal ischemia. In a desperate attempt to recruit a blood supply, hypoxic retinal cells release **VEGF**, which diffuses throughout the vitreous cavity promoting neovascularization. Unfortunately, the resulting new fibrovascular

Obviously, VEGF plays a central role in the pathogenesis of DBR. Let's take a closer look at it.

VEGF (we will have much more to say about VEGF shortly)

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What does **VEGF** stand for?







What does **VEGF** stand for? Vascular endothelial growth factor







What does **VEGF** stand for? Vascular endothelial growth factor

Broadly speaking, what is it?







What does **VEGF** stand for? Vascular endothelial growth factor

Broadly speaking, what is it? An extracellular signaling protein involved in vascular development



Q

Diabetic Retinopathy: The Basics

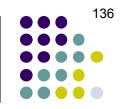


What does **VEGF** *stand for?* Vascular endothelial growth factor

Broadly speaking, what is it? An extracellular signaling protein involved in vascular development Does VEGF do anything besides grow new blood vessels?







What does **VEGF** *stand for?* Vascular endothelial growth factor

Broadly speaking, what is it? An extracellular signaling protein involved in vascular development *Does VEGF do anything besides grow new blood vessels?* Yes, it also is a potent vasodilator (it was known originally as *vascular permeability factor*)

VEGF-A₁₆₅

Q

Diabetic Retinopathy: The Basics



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How potent?







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How potent? About 10,000x more potent than histamine!





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Broadly speaking, what is it? An extracellular signaling protein involved in vascular development *Does VEGF do anything besides grow new blood vessels?* Yes, it also is a potent vasodilator (it was known originally as *vascular permeability factor*)

low potent?

About 10,000x more potent than histamine!



This property accounts for VEGF's role in the development of diabetic macular edema, and explains why anti-VEGF meds can treat this condition!

(Diabetic macular edema is addressed in slide-set R32)

Q

Diabetic Retinopathy: The Basics



What does **VEGF** *stand for?* Vascular endothelial growth factor

Broadly speaking, what is it? An extracellular signaling protein involved in vascular development

How does VEGF work?







What does **VEGF** stand for? Vascular endothelial growth factor

Broadly speaking, what is it? An extracellular signaling protein involved in vascular development

How does VEGF work? Extracellular VEGF binds to VEGF receptors (VEGFR), which are transmembrane receptor tyrosine kinase (RTK) structures.



Q



What does the A signify?

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VEGF-A₁₆₅



VEGF-A₁₆₅



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How does VEGF work? Extracellular VEGF binds to VEGF receptors (VEGFR), which are transmembrane receptor tyrosine kinase (RTK) structures. What does the **A** signify?

VEGF is not a single entity—a number of similar-but-different proteins comprise the 'VEGF family.' These are differentiated as VEGF-A through VEGF-F. (One family member, *placental growth factor* [PIGF], is the exception to the naming rule.) When the term *VEGF* is used in the ophthalmology literature without a subfamily designation, it is understood to mean VEGF-A.



VEGF-A₁₆₅



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What does **165** signify?



VEGF-A₁₆₅



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What does **165** signify?

VEGF-A is not a single entity either. At least 4 isoforms exist; these differ in the number of peptides they contain, and that number is used as a subscript to identify specific isoforms.

VEGF-A₁₆₅



What does **VEGF** stand for? Vascular endothelial growth factor

Broadly speaking, what is it? An extracellular signaling protein involved in vascular development

How does VEGF work? Extracellular VEGF binds to VEGF receptors (VEGFR), which are transmembrane receptor tyrosine kinase (RTK) structures. What does the **A** signify?

VEGF is not a single entity—a number of similar-but-different proteins comprise the 'VEGF family.' These are differentiated as VEGF-A through VEGF-F. (One family member, *placental growth factor* [PIGF], is the exception to the naming rule.) When the term *VEGF* is used in the ophthalmology literature without a subfamily designation, it is understood to mean VEGF-A.

What does **165** signify?

VEGF-A is not a single entity either. At least 4 isoforms exist; these differ in the number of peptides they contain, and that number is used as a subscript to identify specific isoforms.

Why focus on isoform 165?



VEGF-A₁₆₅



What does **VEGF** stand for? Vascular endothelial growth factor

Broadly speaking, what is it? An extracellular signaling protein involved in vascular development

How does VEGF work? Extracellular VEGF binds to VEGF receptors (VEGFR), which are transmembrane receptor tyrosine kinase (RTK) structures. What does the **A** signify?

VEGF is not a single entity—a number of similar-but-different proteins comprise the 'VEGF family.' These are differentiated as VEGF-A through VEGF-F. (One family member, *placental growth factor* [PIGF], is the exception to the naming rule.) When the term *VEGF* is used in the ophthalmology literature without a subfamily designation, it is understood to mean VEGF-A.

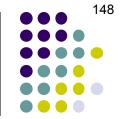
What does 165 signify?

VEGF-A is not a single entity either. At least 4 isoforms exist; these differ in the number of peptides they contain, and that number is used as a subscript to identify specific isoforms.

Why focus on isoform 165?

It seems to be the most important with respect to pathologic angiogenesis in the human eye.

Classification of diabetic retinopathy



- Nonproliferative diabetic retinopathy (NPDR)
 - Mild: Any DBR < moderate</p>

What landmark clinical trial provided this system of PDR classification?

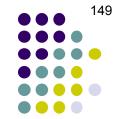
Proliferative diabetic retinopathy (PDR)

• High-risk PDR

- Large (at least ¼ DD) area of NVD with or without VH, OR
- Large (at least ½ DD) area of NVE with VH

Α

Classification of diabetic retinopathy

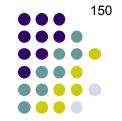


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 - What landmark clinical trial provided this system of PDR classification? The **Diabetic Retinopathy Study** (DRS)

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What question did the DRS seek to answer?

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What question did the DRS seek to answer? 'Is PRP effective in treating PDR/severe NPDR?'



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Classification of diabetic retinopathy



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What landmark clinical trial provided this system of PDR classification?

The Diabetic Retinopathy Study (DRS)

What does PRP stand for in this context?

Proliferative diabetic retinopathy (PDR)

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Classification of diabetic retinopathy



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 - Mild: Any DBR < moderate</p>

What landmark clinical trial provided this system of PDR classification?

The Diabetic Retinopathy Study (DRS)



What does PRP stand for in this context? Panretinal photocoagulation

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Classification of diabetic retinopathy



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What question did the DRS seek to answer? 'Is PRP effective in treating PDR/severe NPDR?'

And the answer was...?

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Classification of diabetic retinopathy



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And the answer was...? We'll get to that in a few slides

Proliferative diabetic retinopathy (PDR)

• High-risk PDR

Δ

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Classification of diabetic retinopathy



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 - What landmark clinical trial provided this system of PDR classification?
 - The Diabetic Retinopathy Study (DRS)

What guestion did the DRS seek to answer?

PRP ffec Let's drill down on PRP for a minute...

And the answer was...? We'll get to that in a few slides

Proliferative diabetic retinopathy (PDR)

- High-risk PDR
 - Any NVD associated with vitreous heme (VH), OR
 - Large (at least ¼ DD) area of NVD with or without VH<mark>, OR</mark>
 - Large (at least ½ DD) area of NVE with VI

First, let's talk about laser-tissue interaction ...





?

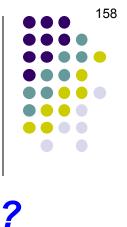
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Diabetic Retinopathy: The Basics

?

What are the five modes of laser-tissue interaction?

?





What are the five modes of laser-tissue interaction?

Thermal

chemical aka photoactivation

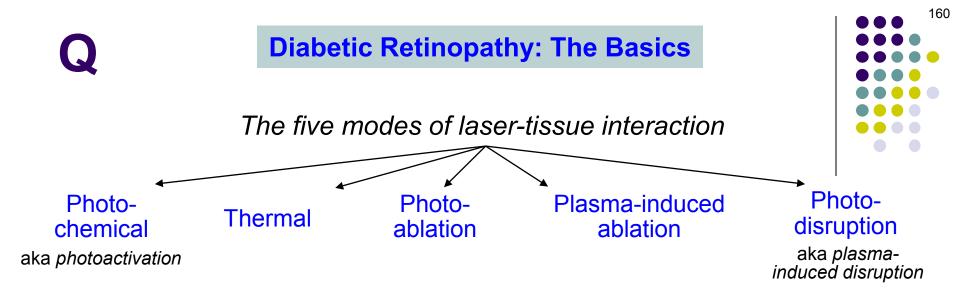
Photo-

Photoablation Plasma-induced ablation

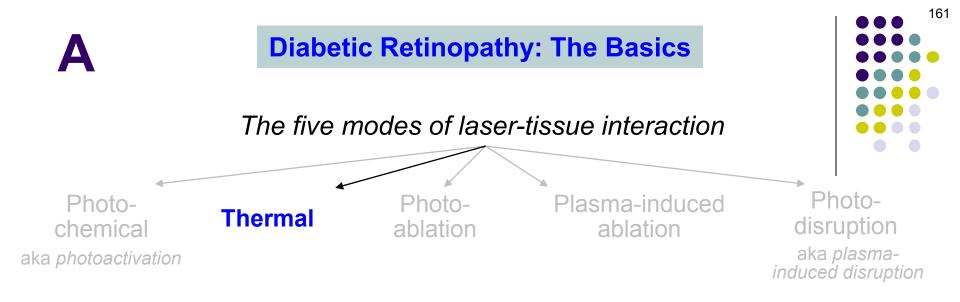
Photodisruption

aka plasmainduced disruption

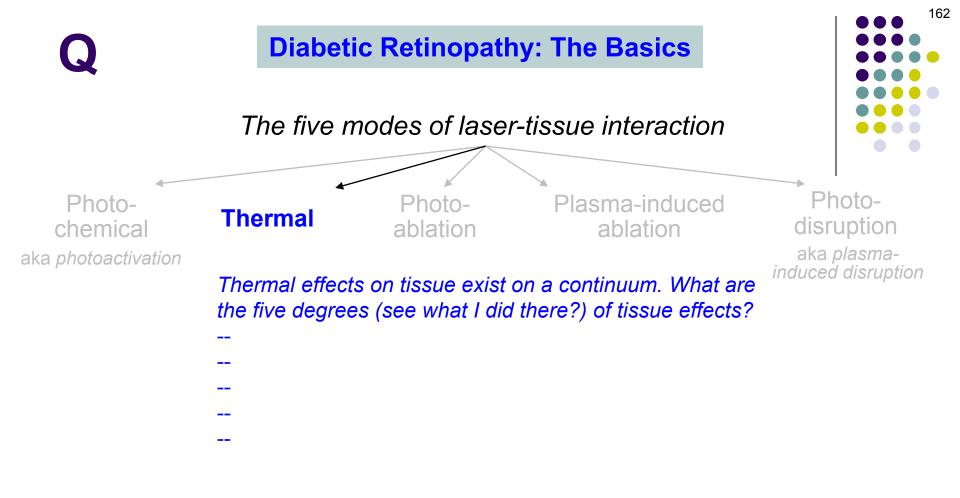


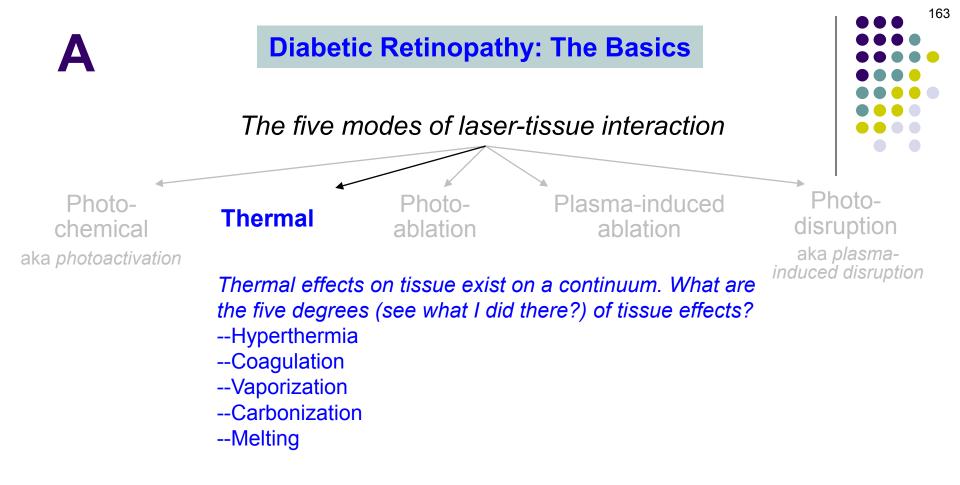


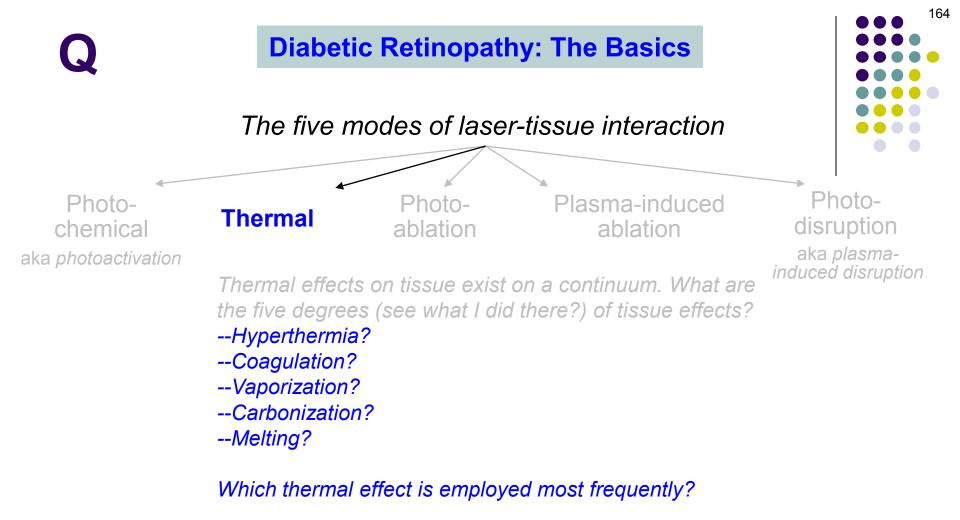
Which mode is PRP an exemplar of?

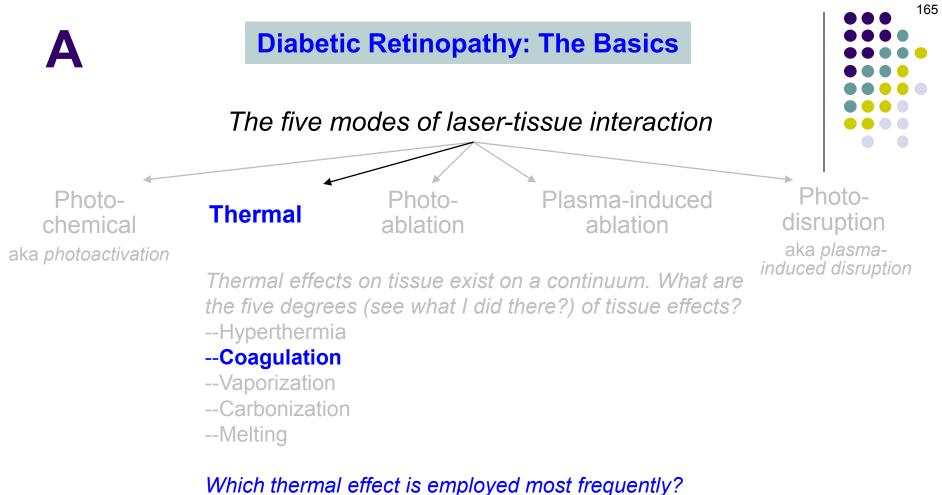


Which mode is PRP an exemplar of? **Thermal**

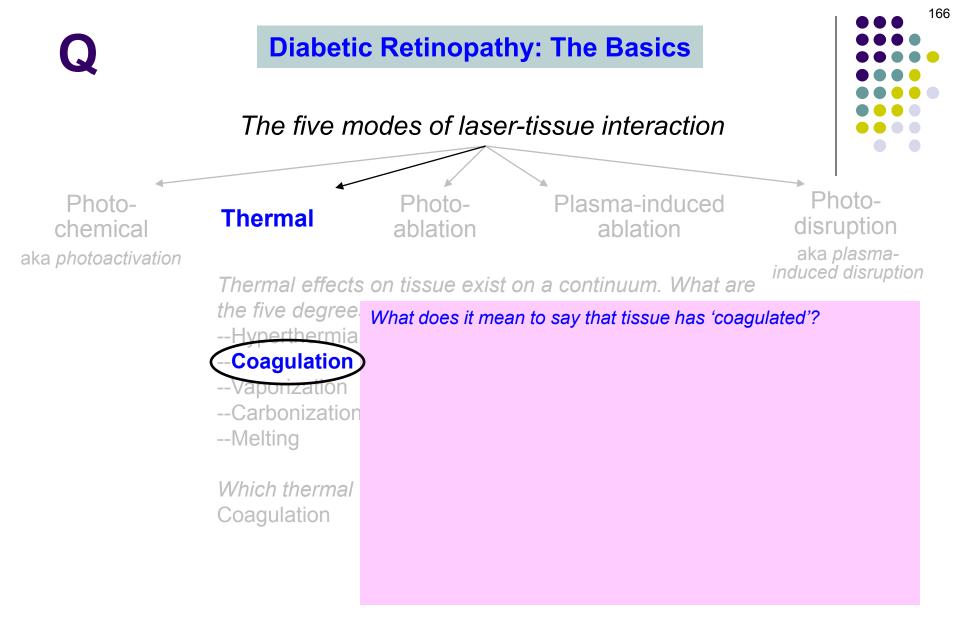


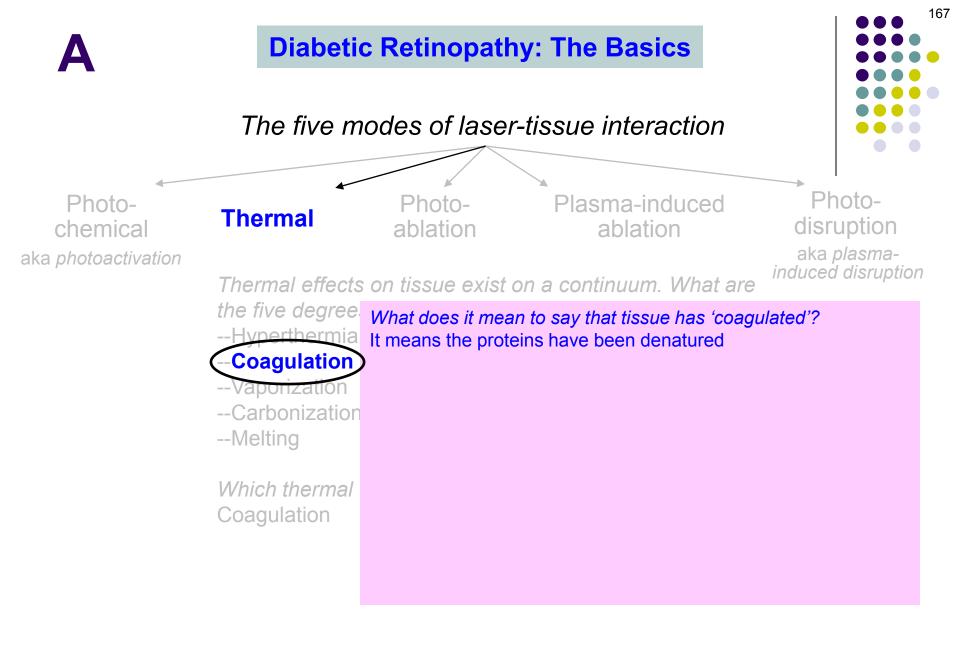


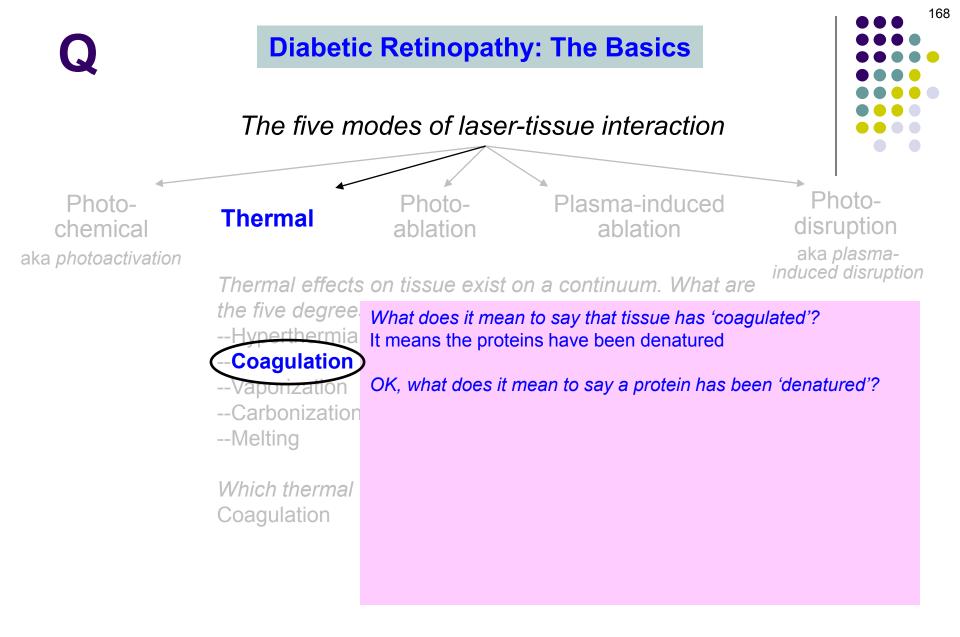


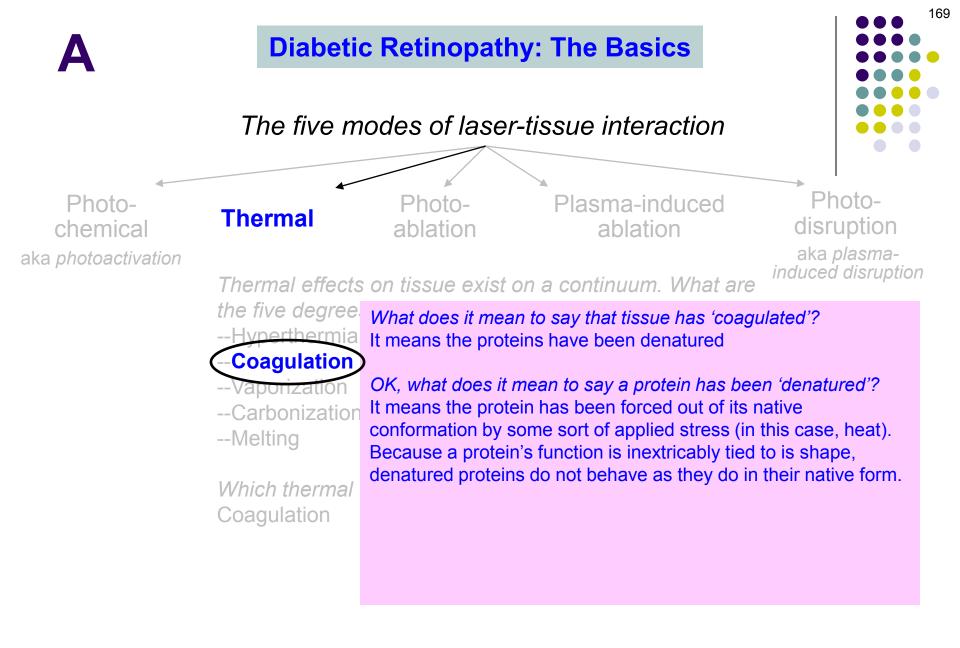


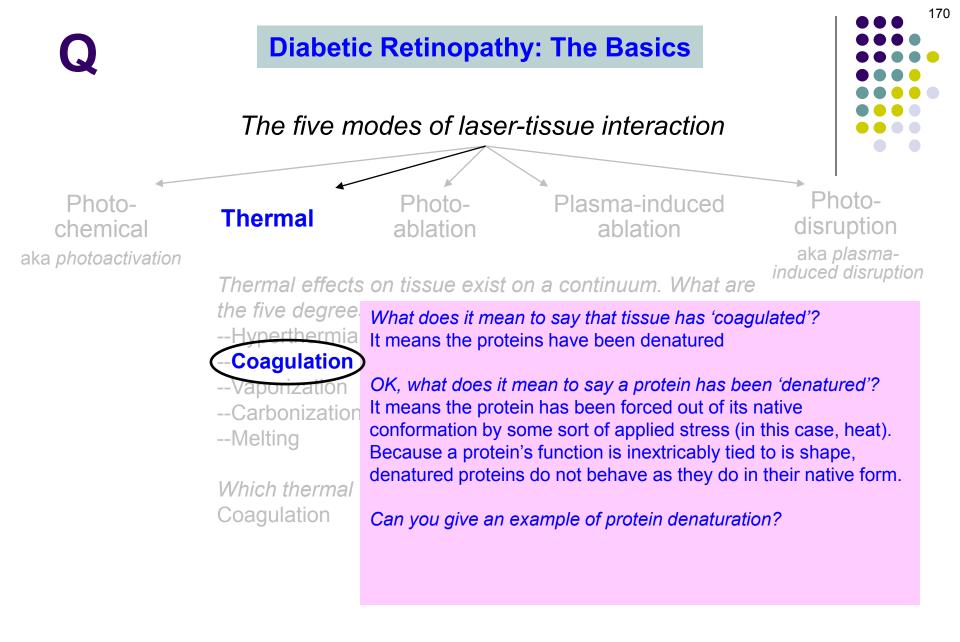
Coagulation

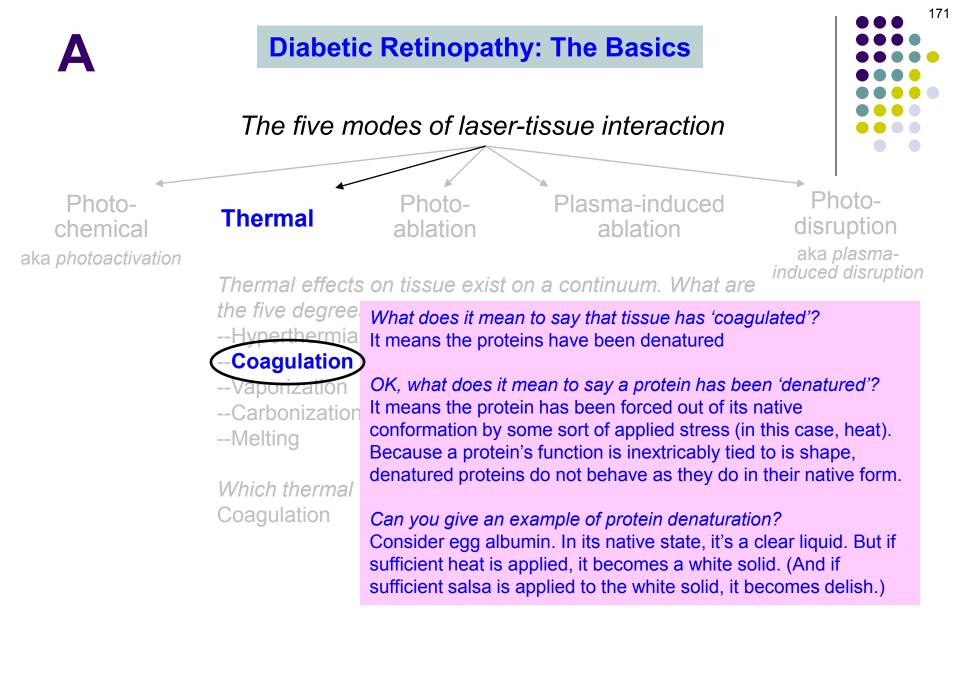


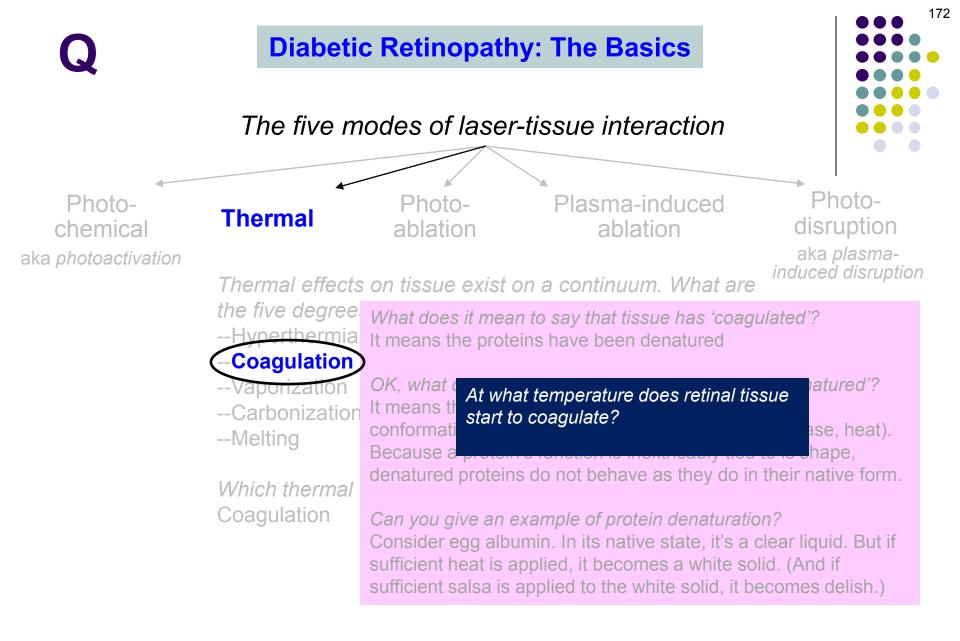


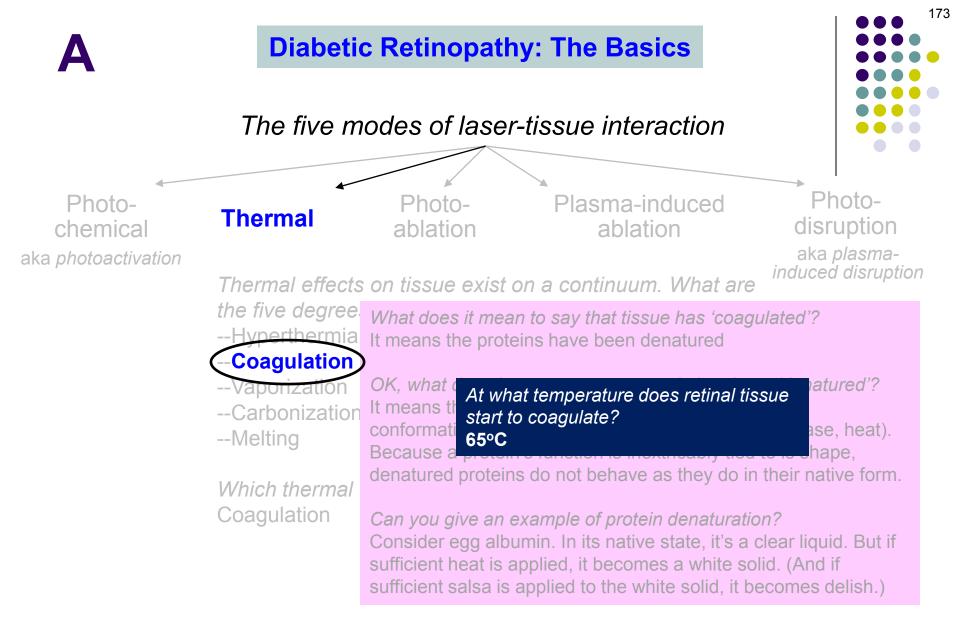


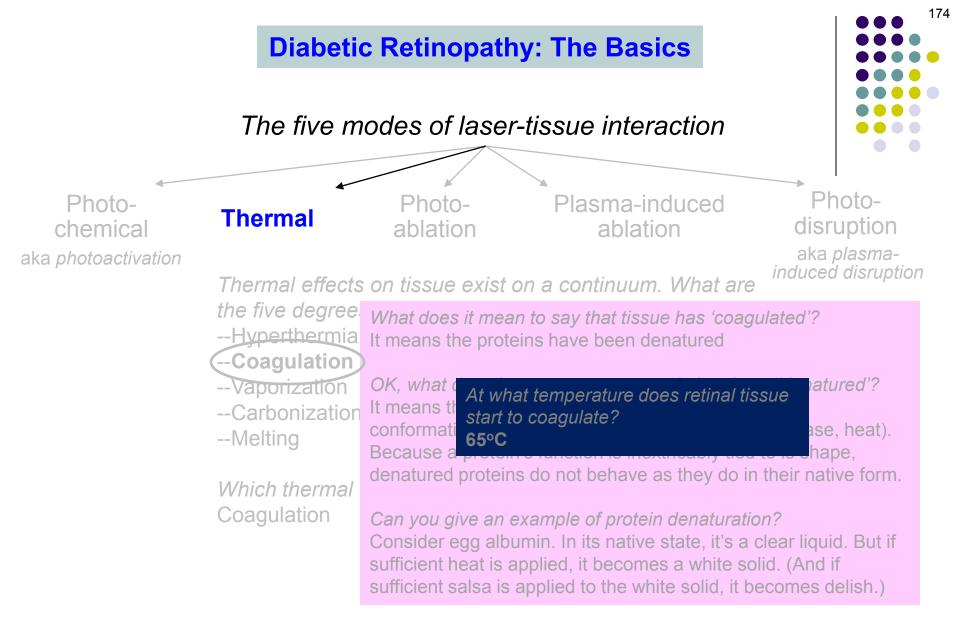












For more info on lasers, see slide-set FELT26



• Which laser is used to perform PRP?





 Which laser is used to perform PRP? Argon green or blue-green





- Which laser is used to perform PRP? Argon green or blue-green
- How many shots constitute a full compliment of PRP?





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- Which laser is used to perform PRP? Argon green or blue-green
- How many shots constitute a full compliment of PRP? About 1200-1400
- What spot size should be used? 500 μm
- How much power?





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color

or

- Which laser is used to perform PRP? Argon green or blue-green
- How many shots constitute a full compliment of PRP? About 1200-1400
- What spot size should be used? $500 \ \mu m$
- How much power? Enough to produce a light coor -colored burn



- Which laser is used to perform PRP? Argon green or blue-green
- How many shots constitute a full compliment of PRP? About 1200-1400
- What spot size should be used? 500 μm
- How much power? Enough to produce a gray or light cream-colored burn





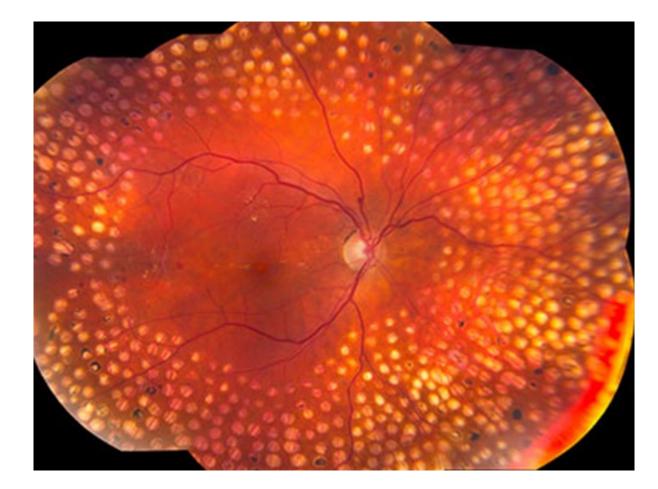
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- How much distance between burns?





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- How many shots constitute a full compliment of PRP? About 1200-1400
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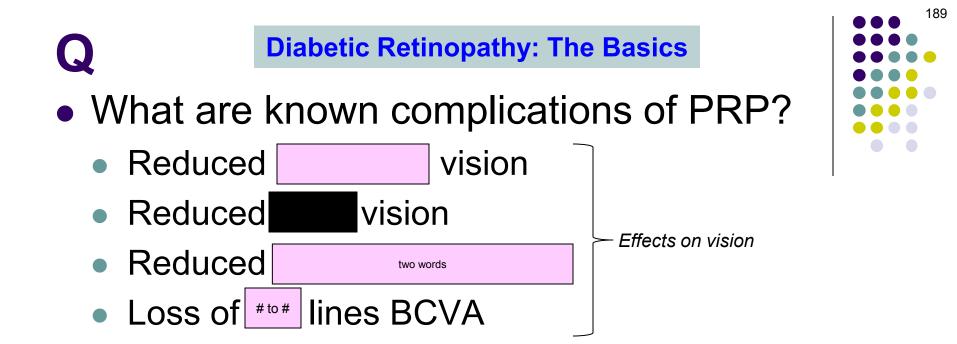
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- Should it be done in one, or multiple sessions?

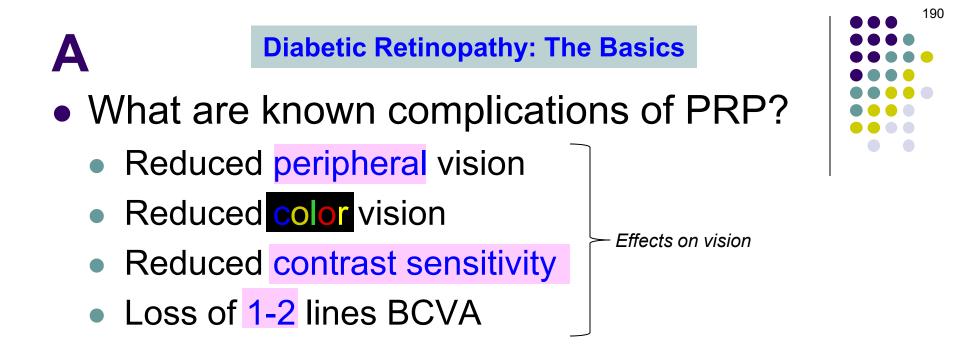




- Which laser is used to perform PRP? Argon green or blue-green
- How many shots constitute a full compliment of PRP? About 1200-1400
- What spot size should be used? 500 μm
- How much power? Enough to produce a gray or light cream-colored burn
- How much distance between burns? About half a burn's width
- Should it be done in one, or multiple sessions?
 It doesn't matter









- What are known complications of PRP?
 - Reduced peripheral vision
 - Reduced color vision
 - Reduced contrast sensitivity
 - Loss of 1-2 lines BCVA
 - Decreased
 - Decreased

parasympathetic function

two words





- What are known complications of PRP?
 - Reduced peripheral vision
 - Reduced color vision
 - Reduced contrast sensitivity
 - Loss of 1-2 lines BCVA
 - Decreased accommodation
 - Decreased corneal sensitivity





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What do accommodation and corneal sensitivity have in common?





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What do accommodation and corneal sensitivity have in common? Both are mediated by the two words nerves





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What do accommodation and corneal sensitivity have in common? Both are mediated by the long ciliary nerves

OK, but what do the long ciliary nerves have to do with PRP?





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 - Loss of 1-2 lines BCVA
 - Decreased accommodation
 - Decreased corneal sensitivity

What do accommodation and corneal sensitivity have in common? Both are mediated by the long ciliary nerves

OK, but what do the long ciliary nerves have to do with PRP? The long ciliary nerves run pretty deep (ie, just under the choroid) in the meridian.





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What do accommodation and corneal sensitivity have in common? Both are mediated by the long ciliary nerves

OK, but what do the long ciliary nerves have to do with PRP? The long ciliary nerves run pretty deep (ie, just under the choroid) in the horizontal meridian.





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What do accommodation and corneal sensitivity have in common? Both are mediated by the long ciliary nerves

OK, but what do the long ciliary nerves have to do with PRP? The long ciliary nerves run pretty deep (ie, just under the choroid) in the horizontal meridian. Because of their location, they are vulnerable to damage during PRP.





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OK, but what do the long ciliary nerves have to do with PRP? The long ciliary nerves run pretty deep (ie, just under the choroid) in the horizontal meridian. Because of their location, they are vulnerable to damage during PRP.

How can one minimize the risk to the long ciliary nerves?





- What are known complications of PRP?
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What do accommodation and corneal sensitivity have in common? Both are mediated by the long ciliary nerves

OK, but what do the long ciliary nerves have to do with PRP? The long ciliary nerves run pretty deep (ie, just under the choroid) in the horizontal meridian. Because of their location, they are vulnerable to damage during PRP.

How can one minimize the risk to the long ciliary nerves? By avoiding the horizontal meridian during PRP





- What are known complications of PRP?
 - Reduced peripheral vision
 - Reduced color vision
 - Reduced contrast sensitivity
 - Loss of 1-2 lines BCVA
 - Decreased accommodation
 - Decreased corneal sensitivity
 - Macular
 - Inadvertent
 burn





- What are known complications of PRP?
 - Reduced peripheral vision
 - Reduced color vision
 - Reduced contrast sensitivity
 - Loss of 1-2 lines BCVA
 - Decreased accommodation
 - Decreased corneal sensitivity
 - Macular edema
 - Inadvertent foveal burn





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 - Macular edema
 - Inadvertent foveal burn
 - Choroidal





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 - Decreased corneal sensitivity
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 - Choroidal detachment
 - latrogenic break in Bruch's \rightarrow





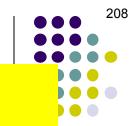
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 - Decreased corneal sensitivity
 - Macular edema
 - Inadvertent foveal burn
 - Choroidal detachment
 - Iatrogenic break in Bruch's → CNVM

(CNVM = Choroidal neovascular membrane)





What does the term high-risk PDR mean? High risk of what?



• Proliferative diabetic retinopathy (PDR) High-risk PDR

- Any NVD associated with vitreous heme (VH), OR
- Large (at least ¼ DD) area of NVD with or without VH, OR
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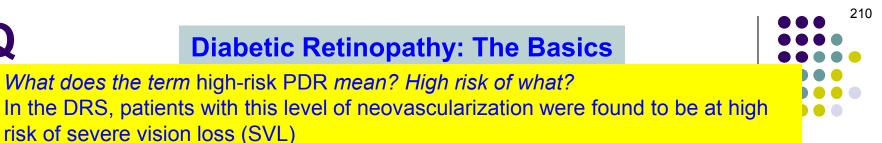


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What does the term high-risk PDR mean? High risk of what? In the DRS, patients with this level of neovascularization were found to be at high risk of severe vision loss (SVL)

Proliferative diabetic retinopathy (PDR) High-risk PDR

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What was the definition of SVL in the DRS?



- Any NVD associated with vitreous heme (VH), OR
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What does the term high-risk PDR mean? High risk of what? In the DRS, patients with this level of neovascularization were found to be at high risk of severe vision loss (SVL)

What was the definition of SVL in the DRS? Snellen acuity $\leq 5/200 (20/800)$



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What does the term high-risk PDR mean? High risk of what? In the DRS, patients with this level of neovascularization were found to be at high risk of severe vision loss (SVL)

What was the definition of SVL in the DRS? Snellen acuity $\leq 5/200 (20/800)$

Why was this level of vision chosen as the benchmark?



- Any NVD associated with vitreous heme (VH), OR
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What does the term high-risk PDR mean? High risk of what? In the DRS, patients with this level of neovascularization were found to be at high risk of severe vision loss (SVL)

What was the definition of SVL in the DRS? Snellen acuity ≤ 5/200 (20/800)

Why was this level of vision chosen as the benchmark? At or below 5/200, visually-guided ambulation becomes problematic



- Any NVD associated with vitreous heme (VH), OR
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What is the clinical implication of finding high-risk PDR in a patient?



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Α

Diabetic Retinopathy: The Basics

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What was the definition of SVL in the DRS? Snellen acuity ≤ 5/200 (20/800)

Why was this level of vision chosen as the benchmark? At or below 5/200, visually-guided ambulation becomes problematic

What is the clinical implication of finding high-risk PDR in a patient? High-risk PDR is the formal justification for performing PRP (I say 'formal' because many clinicians will offer PRP at lesser levels of DBR if they feel it is warranted)



- Any NVD associated with vitreous heme (VH), OR
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Q

Diabetic Retinopathy: The Basics

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Why was this level of vision chosen as the benchmark? At or below 5/200, visually-guided ambulation becomes problematic

What is the clinical implication of finding high-risk PDR in a patient? High-risk PDR is the formal justification for performing PRP (I say 'formal' because many clinicians will offer PRP at lesser levels of DBR if they feel it is warranted)

To answer an earlier question: Per the DRS, is PRP effective at preventing SVL?



- Any NVD associated with vitreous heme (VH), OR
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Q/A

Diabetic Retinopathy: The Basics

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What does the term high-risk PDR mean? High risk of what? In the DRS, patients with this level of neovascularization were found to be at high risk of severe vision loss (SVL)

What was the definition of SVL in the DRS? Snellen acuity $\leq 5/200 (20/800)$

Why was this level of vision chosen as the benchmark? At or below 5/200, visually-guided ambulation becomes problematic

What is the clinical implication of finding high-risk PDR in a patient? High-risk PDR is the formal justification for performing PRP (I say 'formal' because many clinicians will offer PRP at lesser levels of DBR if they feel it is warranted)

To answer an earlier question: Per the DRS, is PRP effective at preventing SVL? Indeed it is—it reduces the risk by at 5 years post-treatment



- Any NVD associated with vitreous heme (VH), OR
- Large (at least ¼ DD) area of NVD with or without VH, OF
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Α

Diabetic Retinopathy: The Basics

What does the term high-risk PDR mean? High risk of what? In the DRS, patients with this level of neovascularization were found to be at high risk of severe vision loss (SVL)

What was the definition of SVL in the DRS? Snellen acuity ≤ 5/200 (20/800)

Why was this level of vision chosen as the benchmark? At or below 5/200, visually-guided ambulation becomes problematic

What is the clinical implication of finding high-risk PDR in a patient? High-risk PDR is the formal justification for performing PRP (I say 'formal' because many clinicians will offer PRP at lesser levels of DBR if they feel it is warranted)

To answer an earlier question: Per the DRS, is PRP effective at preventing SVL? Indeed it is—it reduces the risk by 50% at 5 years post-treatment



- Any NVD associated with vitreous heme (VH), OR
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- Large (at least ½ DD) area of NVE with VH

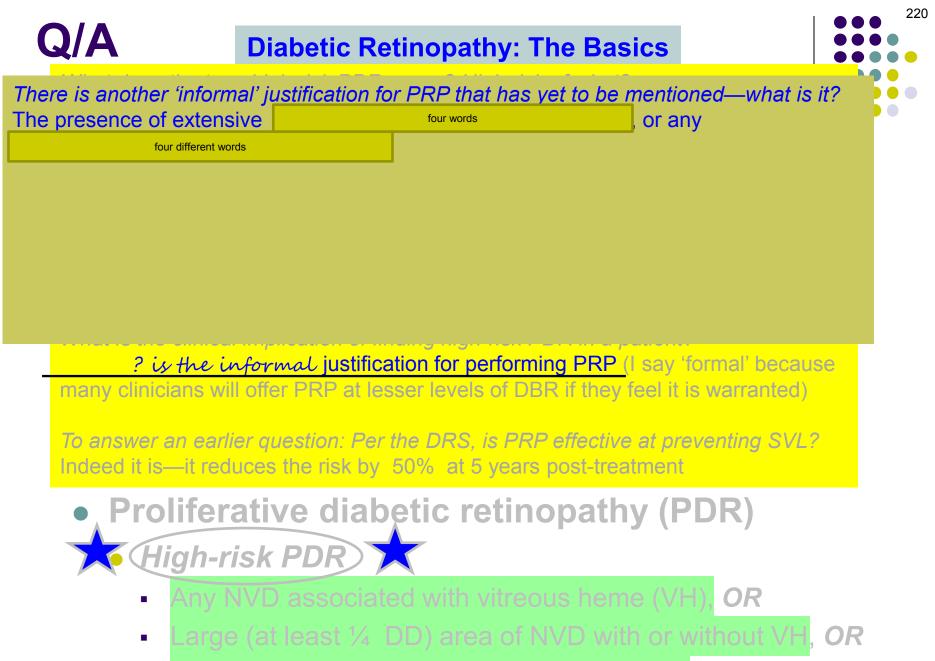
There is another 'informal' justification for PRP that has yet to be mentioned—what is it?

? is the informal justification for performing PRP (I say 'formal' because many clinicians will offer PRP at lesser levels of DBR if they feel it is warranted)

To answer an earlier question: Per the DRS, is PRP effective at preventing SVL? Indeed it is—it reduces the risk by 50% at 5 years post-treatment



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- Large (at least ½ DD) area of NVE with VF



Large (at least ½ DD) area of NVE with VF



There is another 'informal' justification for PRP that has yet to be mentioned—what is it? The presence of extensive **neovascularization of the iris** (NVI), or any **neovascularization of the angle** (NVA)

NVI/NVA is the informal justification for performing PRP (I say 'formal' because many clinicians will offer PRP at lesser levels of DBR if they feel it is warranted)

To answer an earlier question: Per the DRS, is PRP effective at preventing SVL? Indeed it is—it reduces the risk by 50% at 5 years post-treatment



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Extensive NVI



223

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two words

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227

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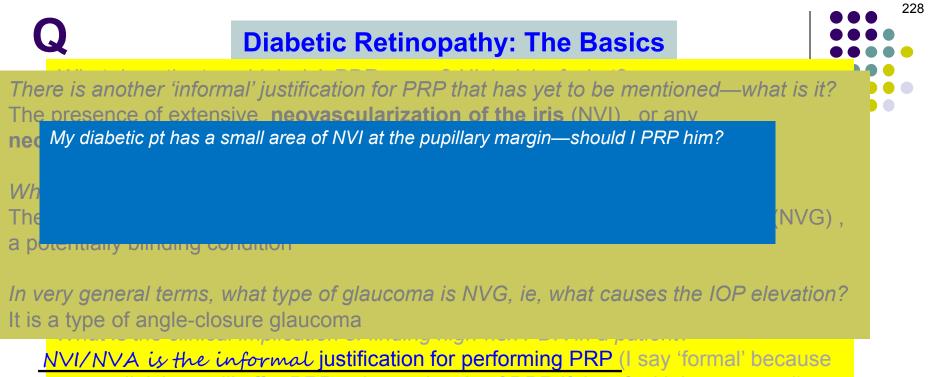
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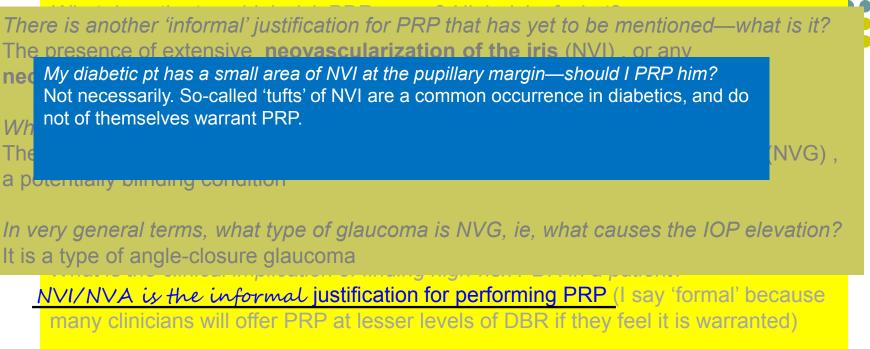
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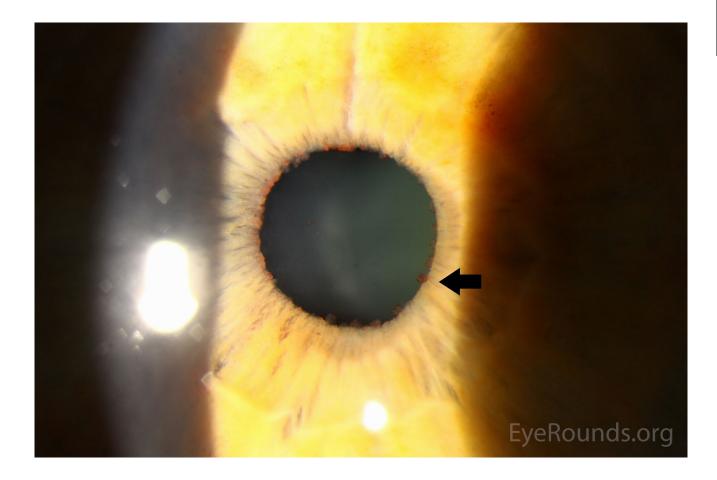
Diabetic Retinopathy: The Basics



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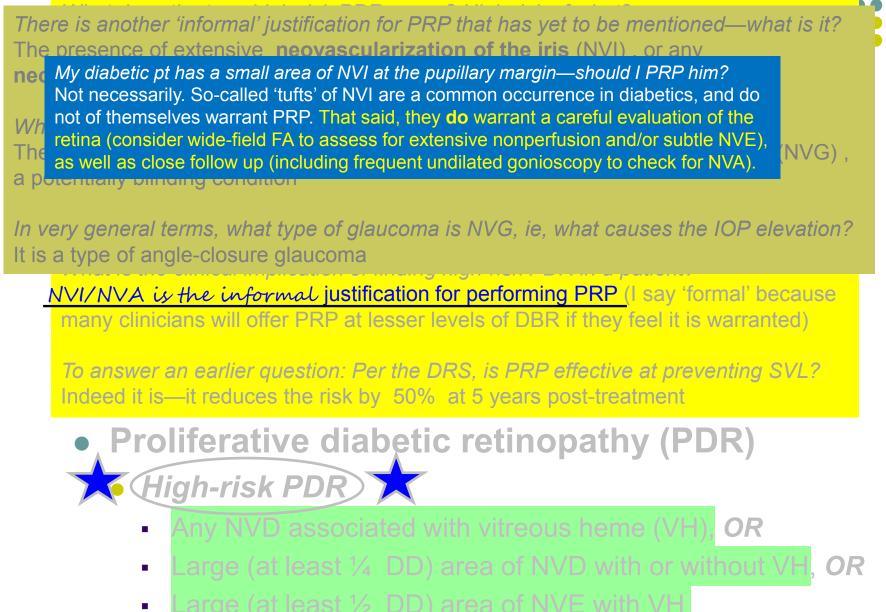
- OR



Pupillary vascular tufts









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There is another 'informal' justification for PRP that has yet to be mentioned—what is it? The presence of extensive **neovascularization of the iris** (NVI) , or any

Net My diabetic pt has a small area of NVI at the pupillary margin—should I PRP him?

I'm confused—if extensive NVI or NVA is concerning for the development of angle-closure glaucoma, shouldn't I be performing laser peripheral iridotomy (LPI) rather than PRP?

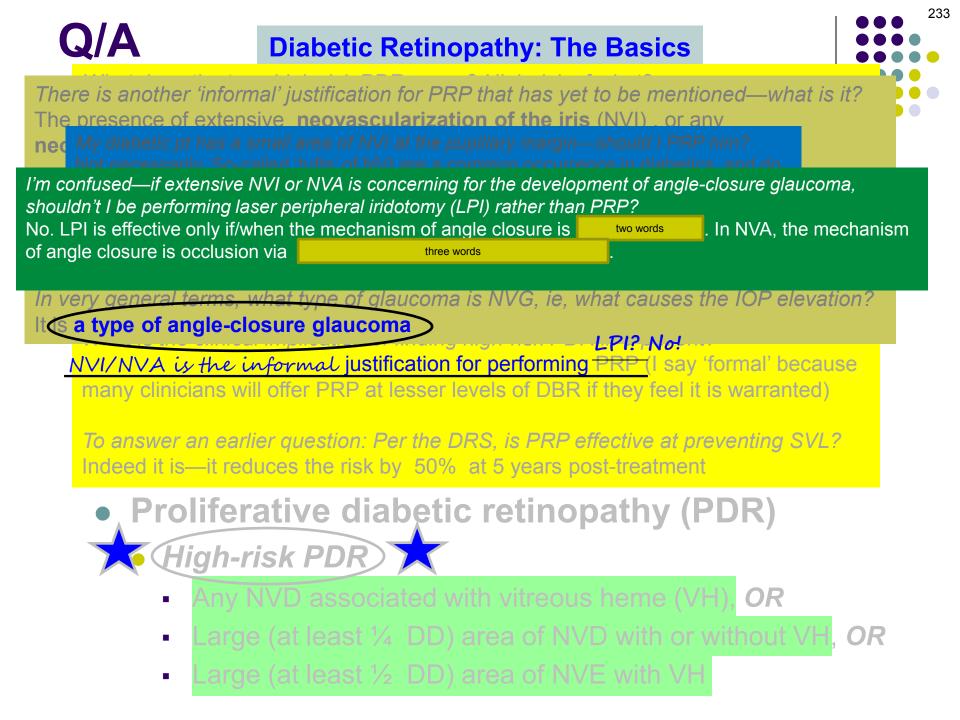
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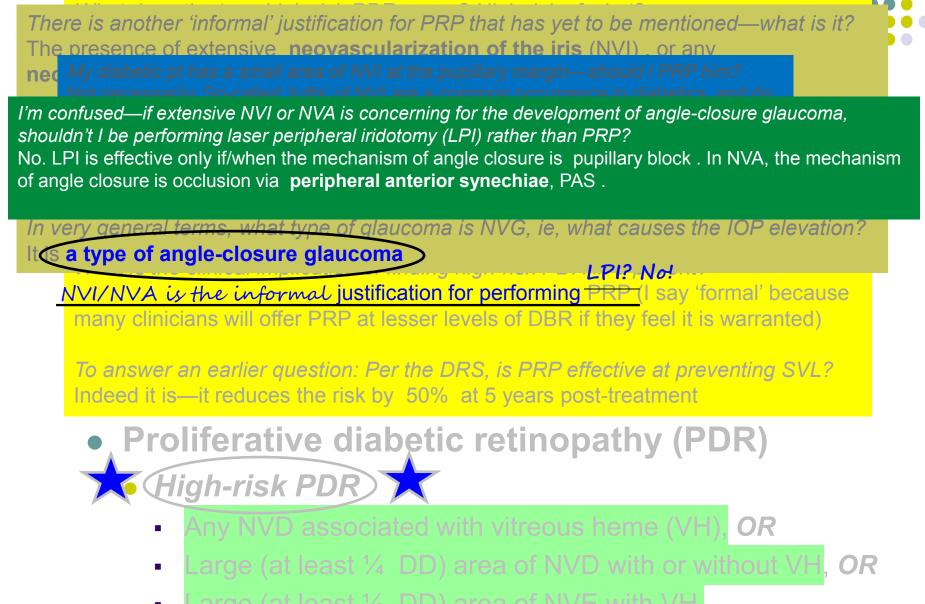
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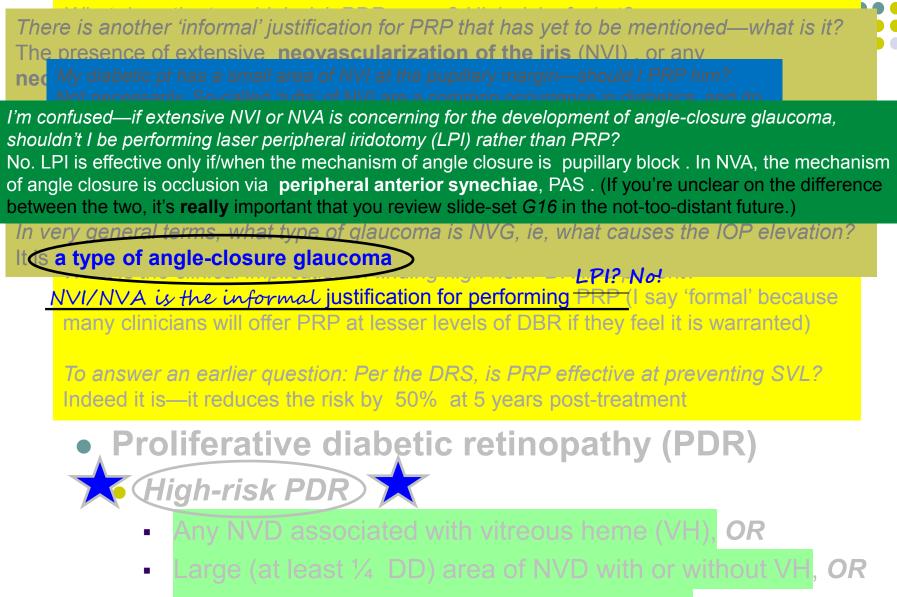




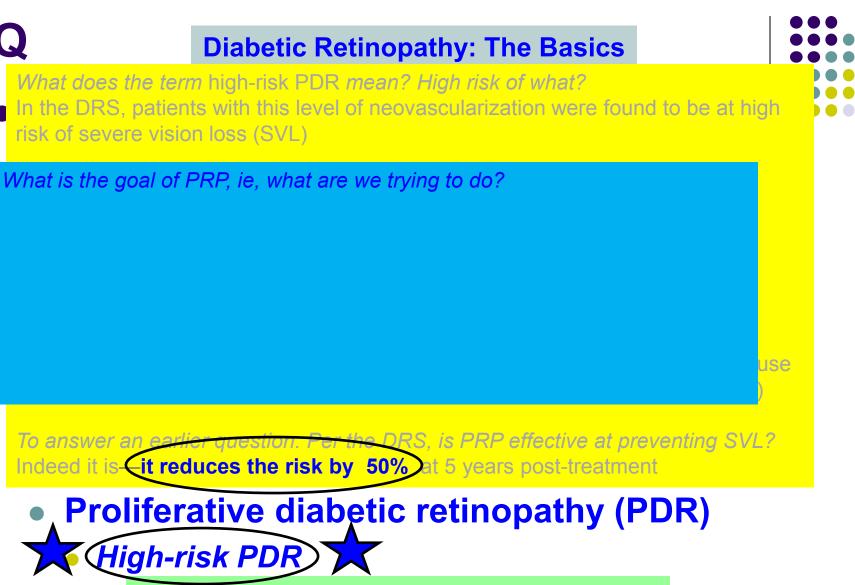




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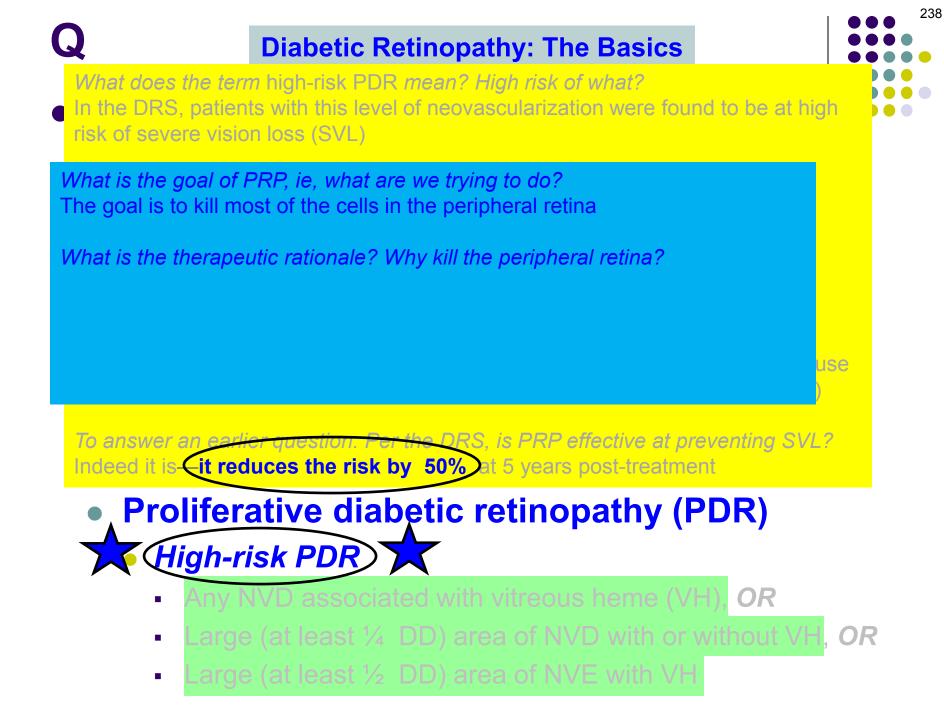


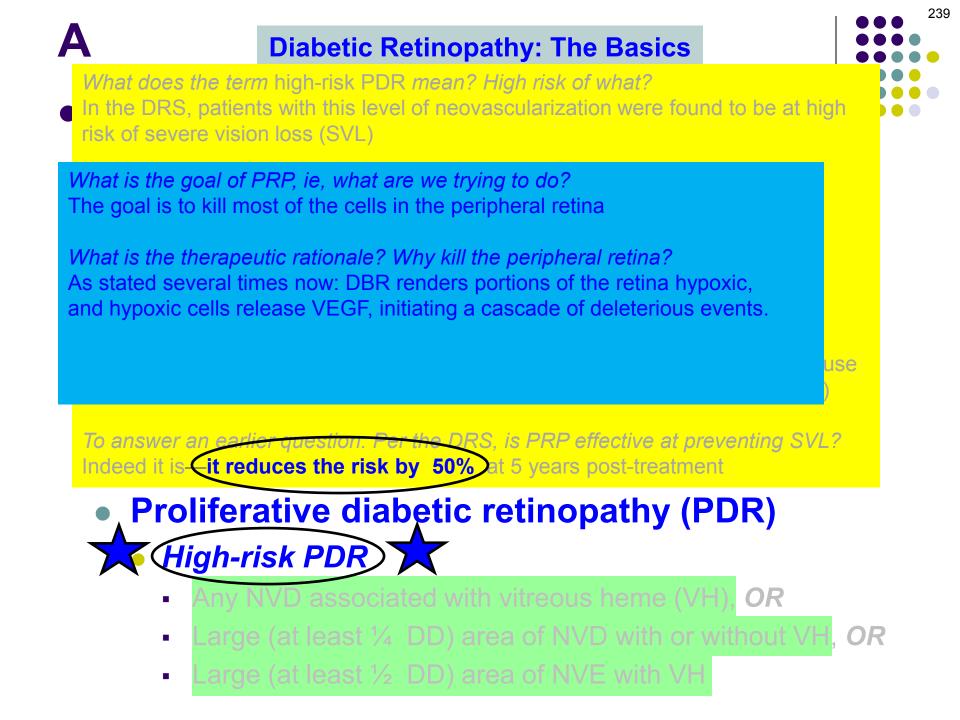
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What does the term high-risk PDR mean? High risk of what? In the DRS, patients with this level of neovascularization were found to be at high risk of severe vision loss (SVL)

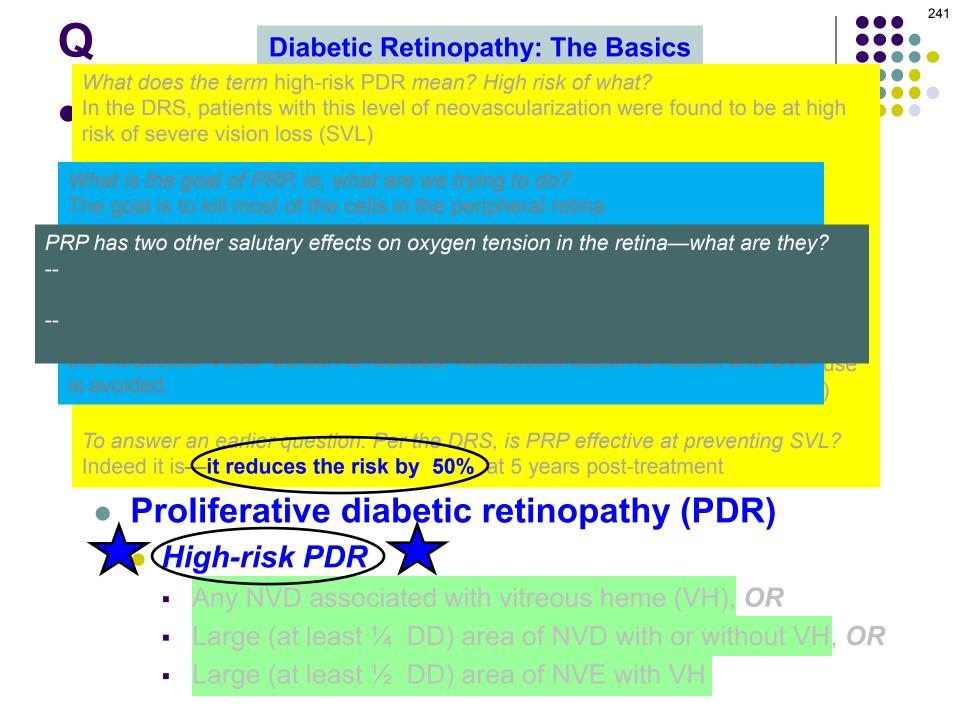
What is the goal of PRP, ie, what are we trying to do? The goal is to kill most of the cells in the peripheral retina

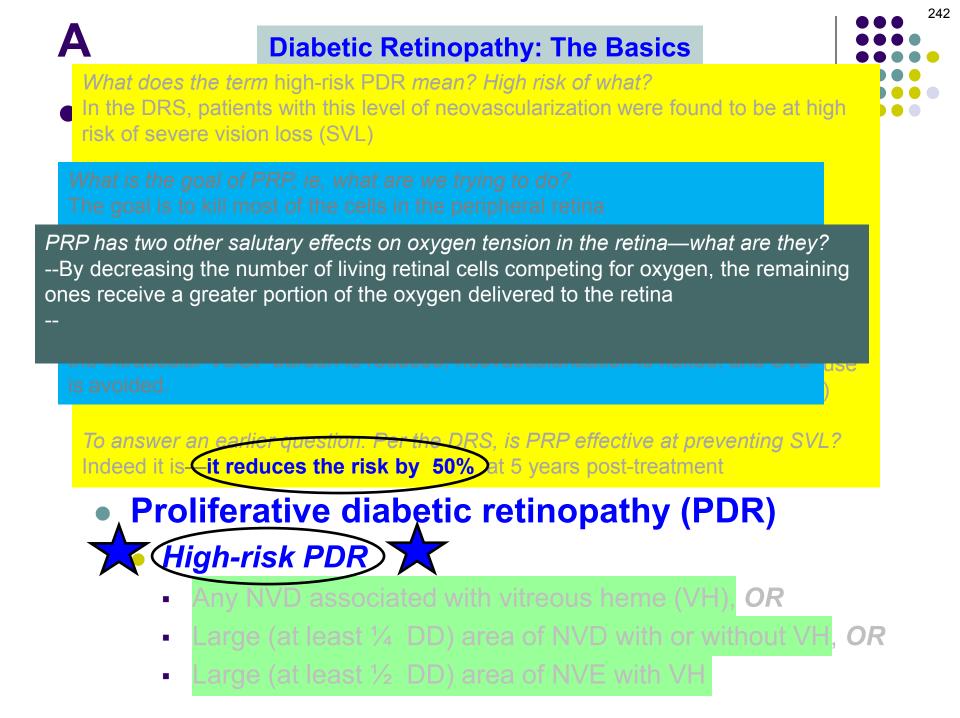
What is the therapeutic rationale? Why kill the peripheral retina? As stated several times now: DBR renders portions of the retina hypoxic, and hypoxic cells release VEGF, initiating a cascade of deleterious events. <u>OTOH, dead cells do not release VEGF</u>. So by euthanizing the hypoxic retina, the intraocular VEGF burden is reduced, neovascularization is halted, and SVL is avoided.

To answer an earlier question. Per the DRS, is PRP effective at preventing SVL? Indeed it is **- it reduces the risk by 50%** at 5 years post-treatment

Proliferative diabetic retinopathy (PDR)
High-risk PDR

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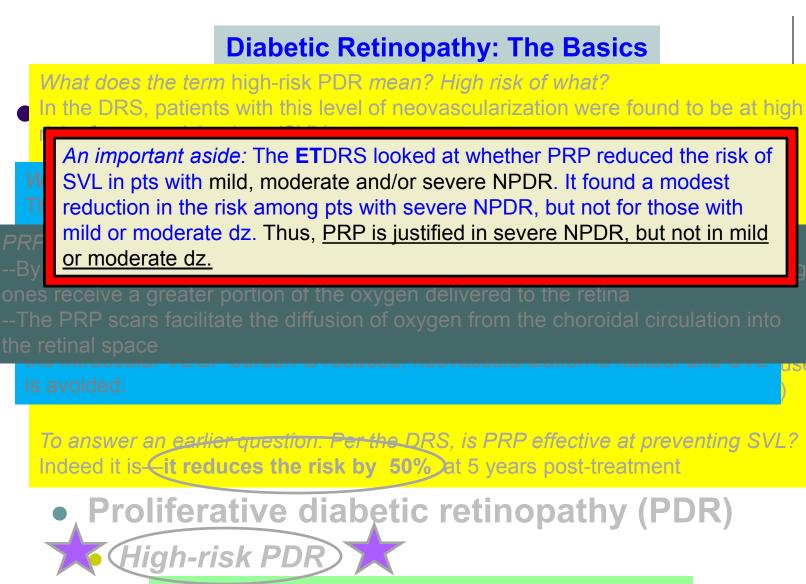
243

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What is the goal of PRP, ie, what are we trying to do? The goal is to kill most of the cells in the peripheral retina

PRP has two other salutary effects on oxygen tension in the retina—what are they? --By decreasing the number of living retinal cells competing for oxygen, the remaining ones receive a greater portion of the oxygen delivered to the retina --The PRP scars facilitate the diffusion of oxygen from the choroidal circulation into the retinal space





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Classification of diabetic retinopathy



- Nonproliferative diabetic retinopathy (NPDR)
 - Mild?Any DBR < moderate</p>
 - Moderate?DBR > mild but < severe</p>
 - Severe? Presence of any 1 of the 4:2:1 rule

The **ET**DRS looked at whether PRP for mild, moderate and/or severe NPDR reduced the risk of SVL. What did it find in this regard?

ET early Per the DRS, is PRP effective at preventing SVL?

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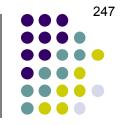
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Per the DRS, is PRP effective at preventing SVL?

It is for severe NPDR (but not mild or moderate)

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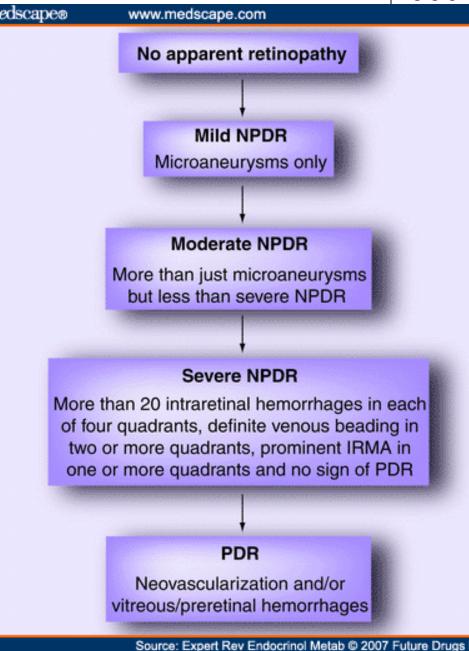
The **ET**DRS looked at whether PRP for mild, moderate and/or severe NPDR reduced the risk of SVL. What did it find in this regard? It found that PRP resulted in a modest reduction of SVL in severe NPDR (especially in pts with Type 2 DM), but not in mild or moderate dz

Per the DRS, is PRP effective at preventing SVL?

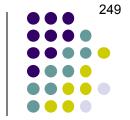
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Take note: DBR is a **progressive** condition, one that passes through a well-defined series of stages on its way to blinding a pt. If DBR is identified at an early stage, the pt has a chance to enact lifestyle modifications that will lead to its resolution. If it is recognized at a later (but pre-SVL) stage, treatment can be performed that may prevent it from blinding the pt. This is why we screen DM pts on the reg.



Q

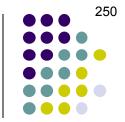


- Classification of diabetic retinopathy
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 - Se And finally: With respect to DBR, what does DME stand for?



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Classification of diabetic retinopathy

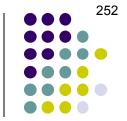


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Where does DME fit into this classification scheme?

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DME is addressed in detail in its own slide-set

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