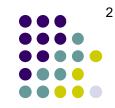


Graves aka Thyroid Eye Disease: True/False



• Graves orbitopathy is secondary to thyroid disease





Graves aka Thyroid Eye Disease: True/False

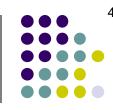


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What does it mean to say Graves is associated with thyroid dz, but not caused by it?

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What does it mean to say Graves is associated with thyroid dz, but not caused by it? Thyroid eye dz (TED) is an autoimmune dz. Research suggests that, moreso than those elsewhere in the body, orbital <u>cell type</u> are highly sensitive to circulating TSH-receptor antibodies (TSH-R Ab)--the same circulating antibodies implicated in many forms of autoimmune thyroid disease.

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Stimulation of orbital fibroblasts by TSH-R Ab has what effects on these cells?

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Stimulation of orbital fibroblasts by TSH-R Ab has what effects on these cells? It induces them to secrete glycosaminoglycans (GAGs), as well as pro-inflammatory cytokines (which attract inflammatory cells to the orbit). Stimulation even causes some fibroblasts to differentiate into adipocytes. Thus, much of the histopathology of TED (ie, an orbit full of ground substance, inflammatory cells, etc) can be traced directly to the effects of TSH-R Ab on orbital fibroblasts.

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But to the original question: Note that the above activities are **not** caused by what's going on in the thyroid gland itself. Thus, while TED often coincides with thyroid dysfunction, it does not **result** from it.

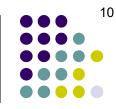
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Speaking of thyroid dysfunction...Is Graves more likely to be associated with hyperythoidism, or hypothyroidism?

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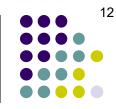


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What percent of Graves cases are associated with hyperthyroidism at presentation?

Graves aka Thyroid Eye Disease: True/False

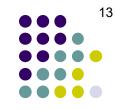


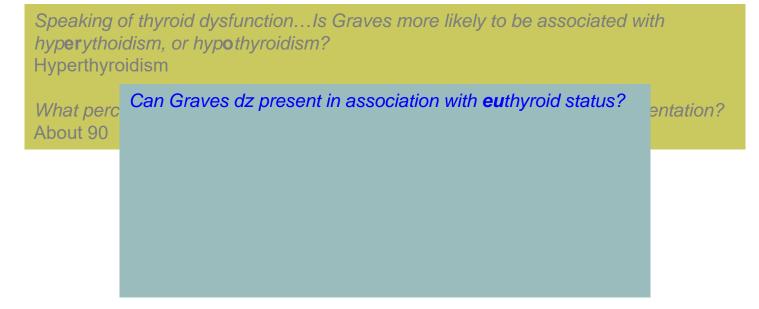
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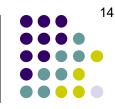
What percent of Graves cases are associated with hyperthyroidism at presentation? About 90

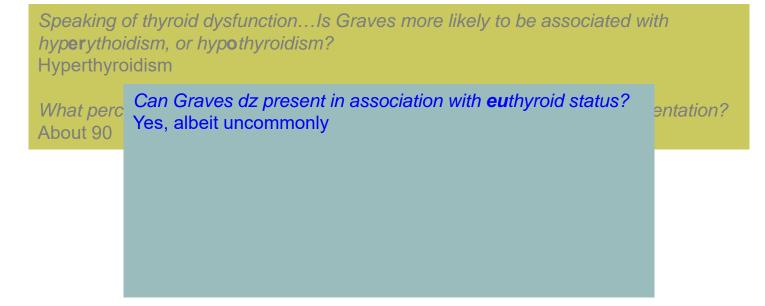
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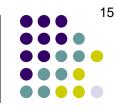


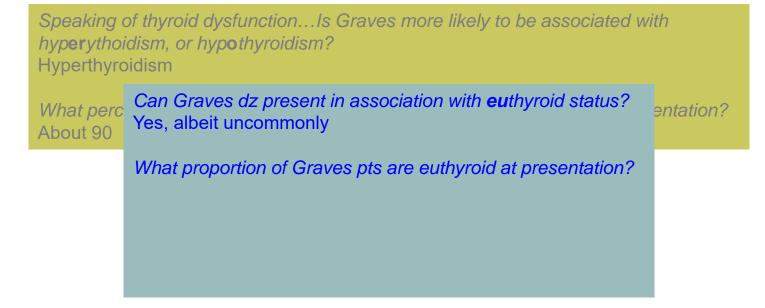
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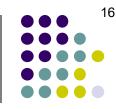


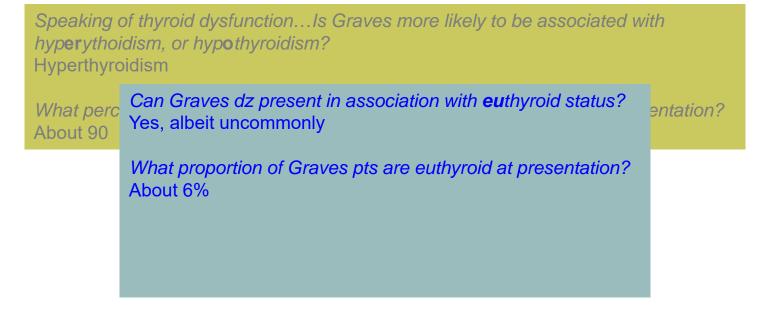
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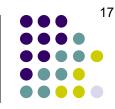


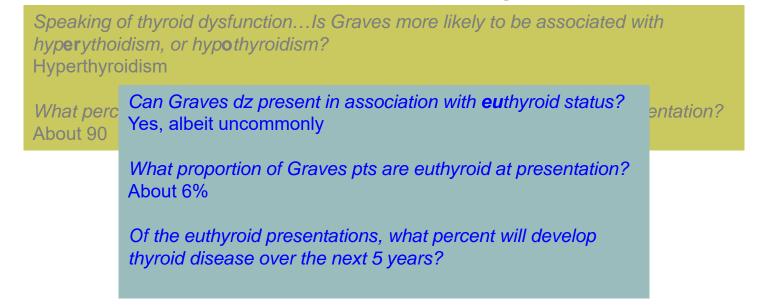
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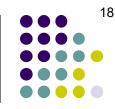


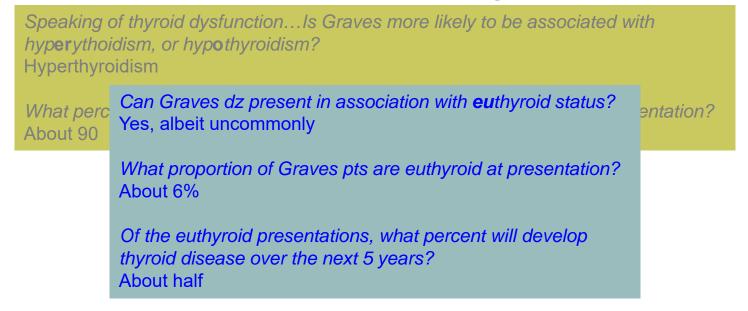
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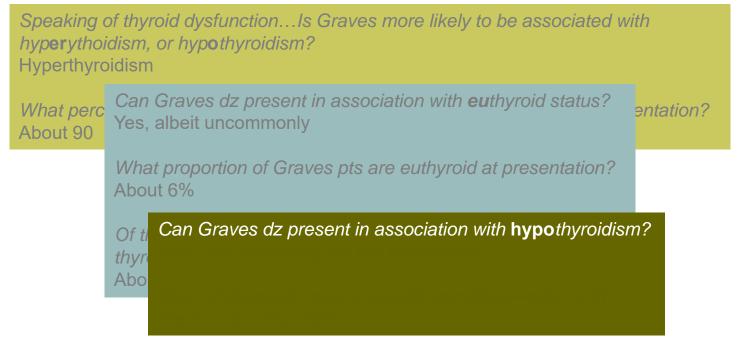
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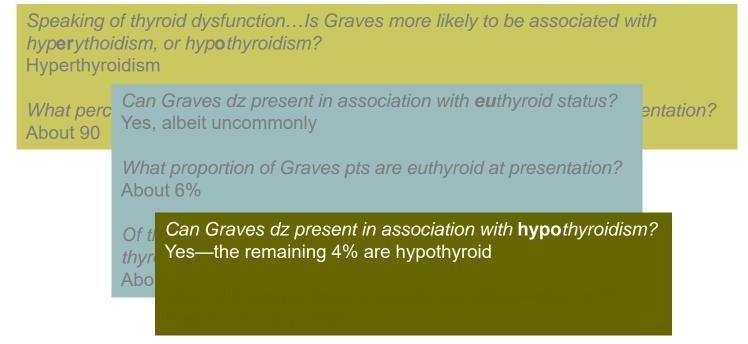
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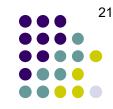


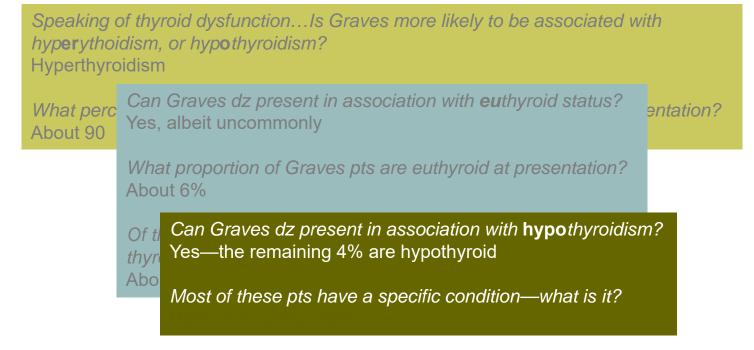
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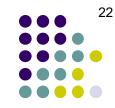


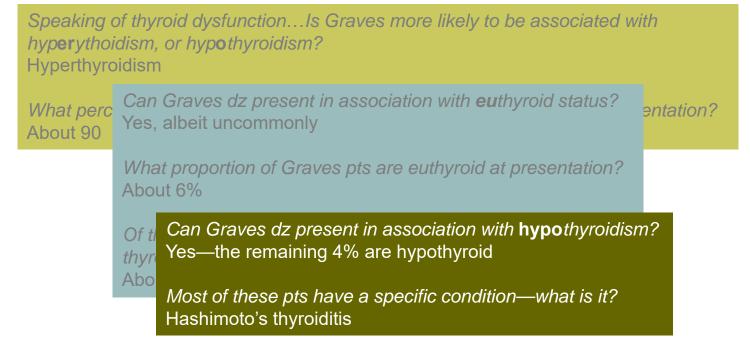
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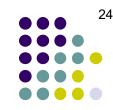
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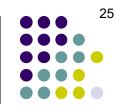


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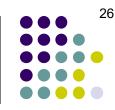
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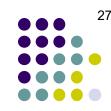
Smoking increases the likelihood of developing TED by how much?

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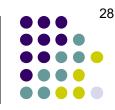
Smoking increases the likelihood of developing TED by how much? A smoker is 7x more likely to develop TED than a nonsmoker!



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- Graves is associated with MS (multiple sclerosis)

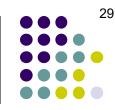
Q/A

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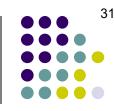
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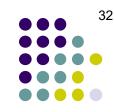
What does MG stand for in this context?

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What does MG stand for in this context? Myasthenia gravis



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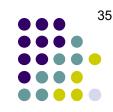




Esotropia

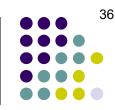
Hypotropia

TED: Strabismus



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A/Q



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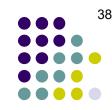
(all 4 rectus muscles)

Α



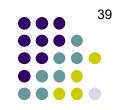
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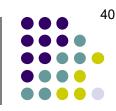


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• The me rectus to likely) is

nferior st to least

Д

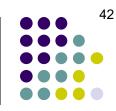
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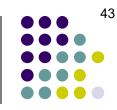
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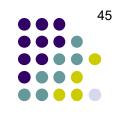
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What is/are the diagnostic criteria for TED?

disease that is strongly associated with thyroid dysfunction, but it is not *caused* by it

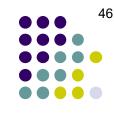
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- Men and women are at equal risk of TED Nope



What is/are the diagnostic criteria for TED? Pt must have at least two of the following:

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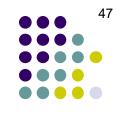


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 Autoimmune thyroid dz
 Typical orbital signs of TED
 Imaging findings c/w TED

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 disease that is not caused by it

 How is this defined?
 d by smoking True; Graves

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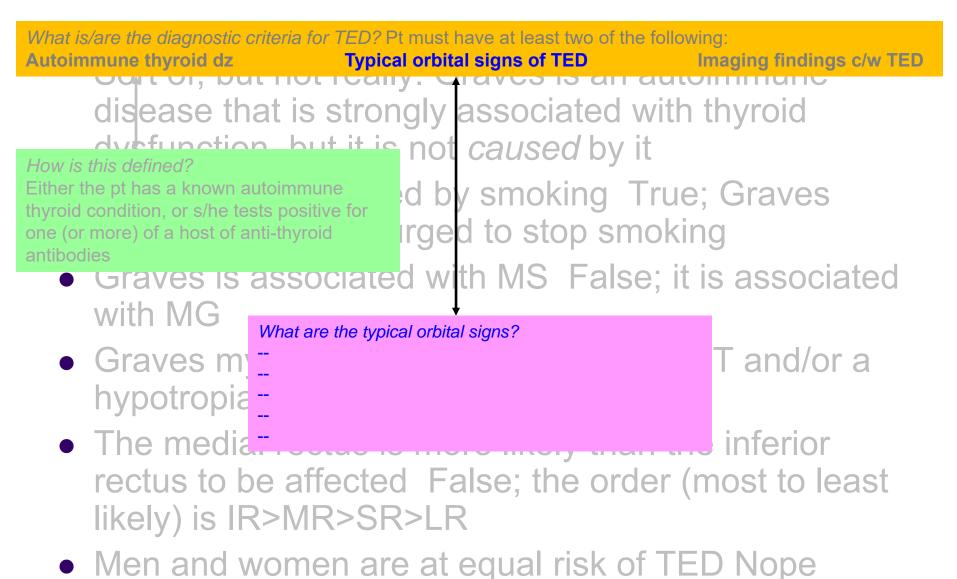
disease that is strongly associated with thyroid dynamics but it is not caused by it

How is this defined?

Either the pt has a known autoimmune thyroid condition, or s/he tests positive for one (or more) of a host of anti-thyroid antibodies d by smoking True; Graves rged to stop smoking

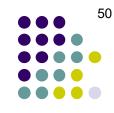
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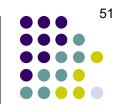


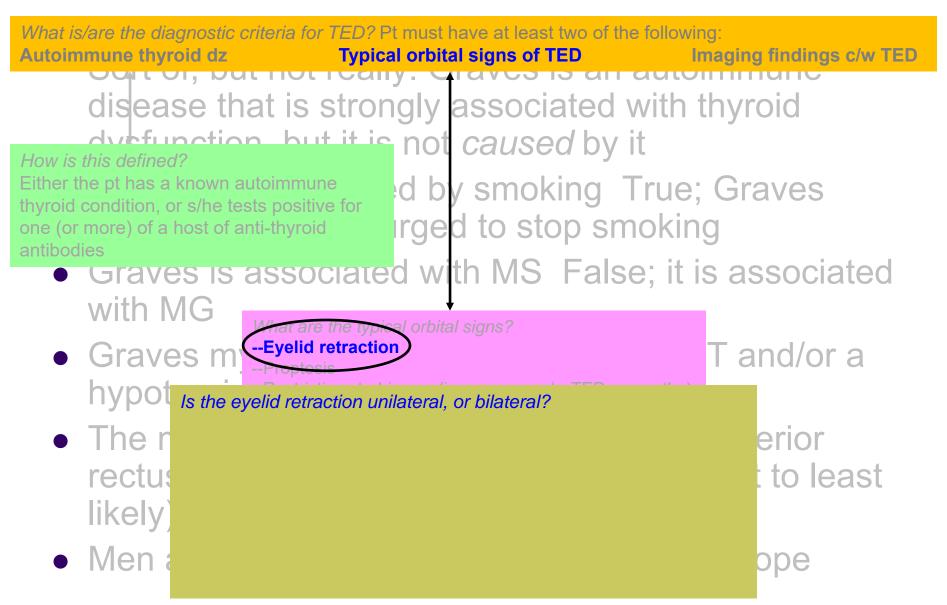
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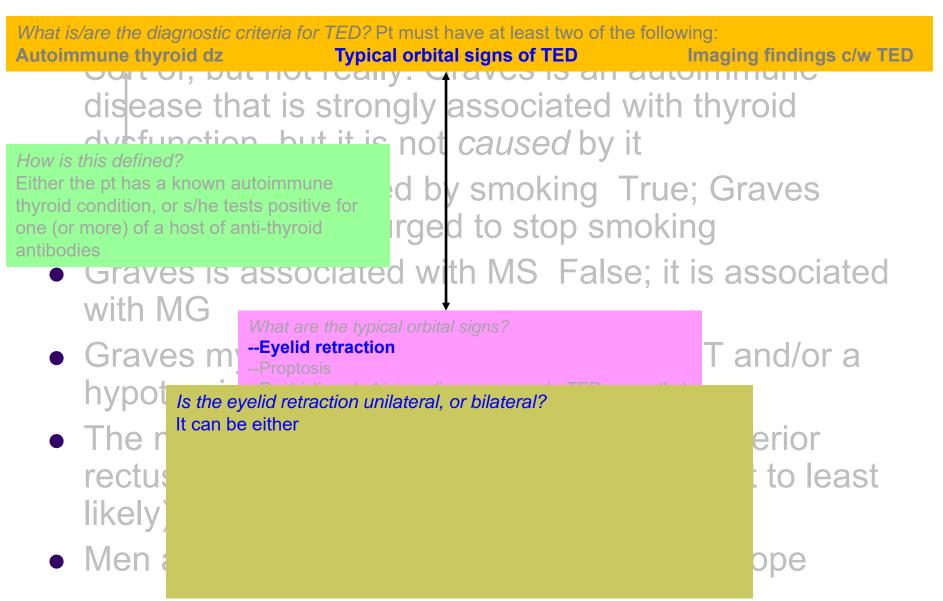
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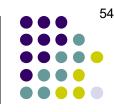
Unilateral*

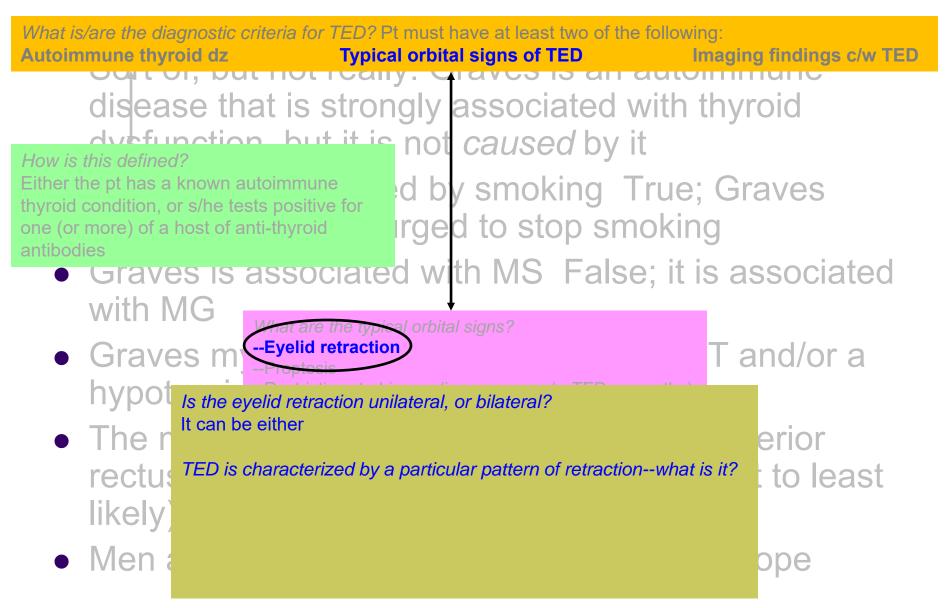


Bilateral

TED: Lid retraction

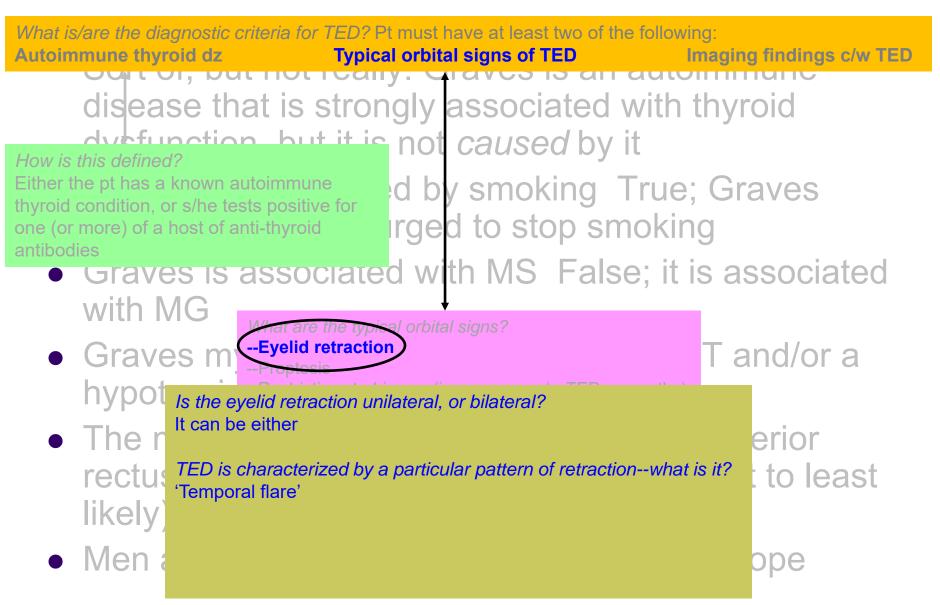
*Or highly asymmetric, at least



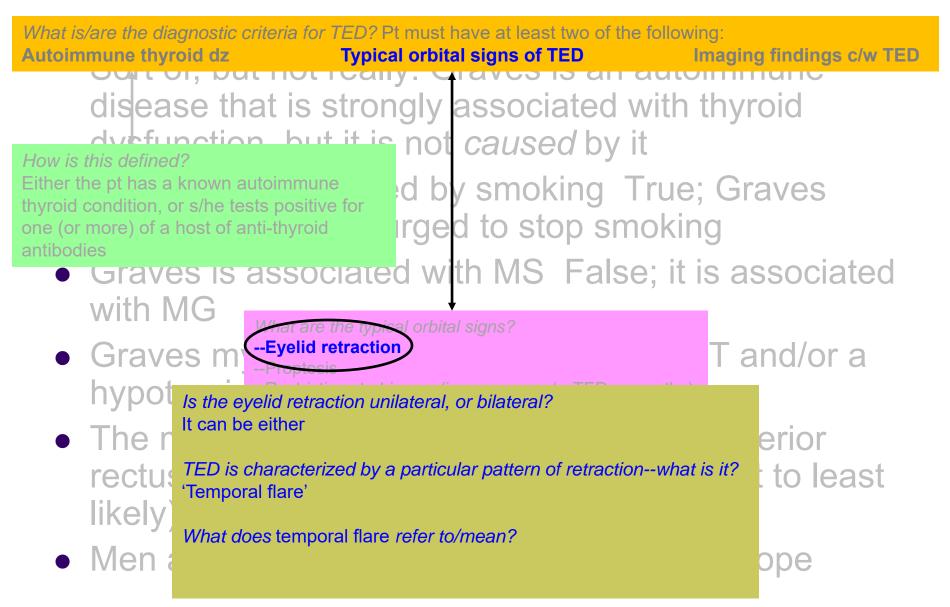




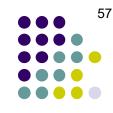


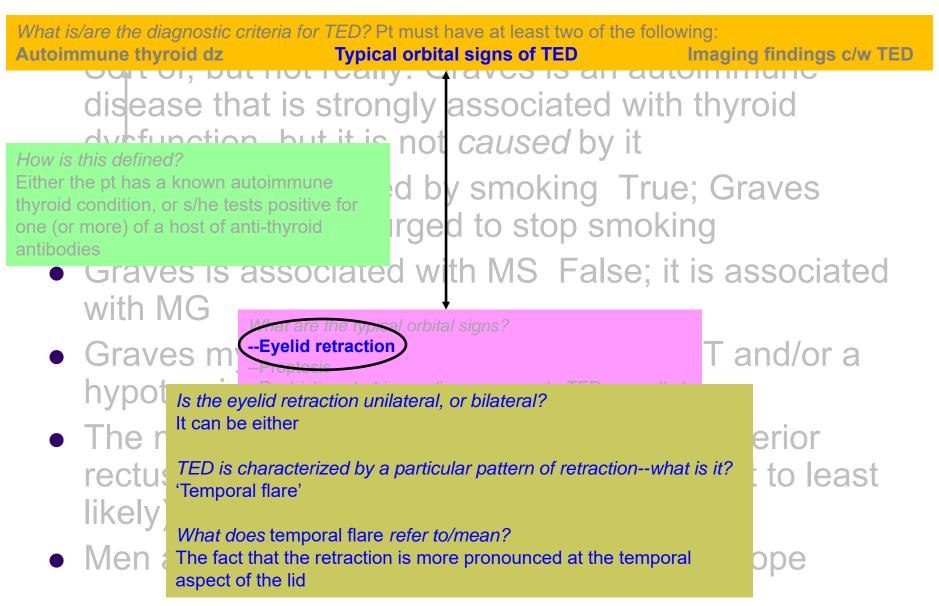








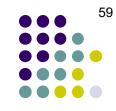






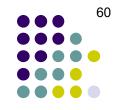
TED: Lid retraction with temporal flare





What is/are the diagnostic criteria for TED? Pt must have at least two of the follo Autoir What well-known brainstem syndrome is associated with lid retraction?		g findings c/w TED
	n thy	roid
one (or antibod	ing	aves ssociated
Graves m Graves m	Та	and/or a
 hypot Is the eyelid retraction unilateral, or bilateral? The n rectus likely TED is characterized by a particular pattern of retractionwh 'Temporal flare' What does temporal flare refer to/mean? 	at is it?	erior to least
• Men a the fact that the retraction is more pronounced at the tempor aspect of the lid	ral	ope





What is/are the diagnostic criteria for TED? Pt must have at least two of the follo Autoir What well-known brainstem syndrome is associated with lid retraction? Parinaud syndrome	wing: Imaging findings c/w TED thyroid
one (or antibod	e; Graves ing t is associated
Graves m Graves m	T and/or a
 hypot Is the eyelid retraction unilateral, or bilateral? The r recture likely Men a 	. 10 16431



What is Autoir	s/are the diagnostic criteria for TED? Pt must have at least two of the follo What well-known brainstem syndrome is associated with lid retraction? Parinaud syndrome What is the eponymous name for lid retraction in Parinaud syndrome?		g findings c/w TED Toid
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•	Graves m hypot is the eyelid retraction unilateral, or bilateral?	Та	nd/or a
•	The n rectuse likely	at is it?	erior to least
٠	What does temporal flare <i>refer to/mean?</i> The fact that the retraction is more pronounced at the temporal aspect of the lid	al	ope





What is/are the diagnostic criteria for TED? Pt must have at least two of the Autoir Autoir What well-known brainstem syndrome is associated with lid retraction Parinaud syndrome What is the eponymous name for lid retraction in Parinaud syndrome How is Either thyroid one (or	ion? Imaging findings c/w TE	D
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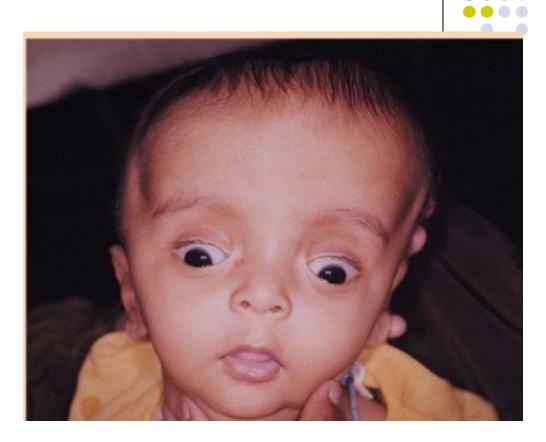
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What is/are the diagnostic criteria for TED? Pt must have at least two of the follow Autoir What well-known brainstem syndrome is associated with lid retraction? Parinaud syndrome What is the eponymous name for lid retraction in Parinaud syndrome? Online is associated with lid syndrome?	wing: Imaging findings c/w TED Official Control of the second sec
Light-near dissociation Impaired upgaze Convergence-retraction nystagmus	e; Graves ing t is associated
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 hypot is the eyelid retraction unilateral, or bilateral? The right can be either The right can be either TED is characterized by a particular pattern of retractionwho the temporal flare? What does temporal flare refer to/mean? Men a fact that the retraction is more pronounced at the temporal aspect of the lid 	. 10 16431



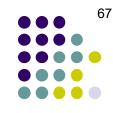


Parinaud syndrome. The combination of lid retraction + impaired upgaze gives rise to a characteristic appearance known as *setting sun sign*



What is/are the diagnostic criteria for TED? Pt must have at least two of the following: Autoimmune thyroid dz Typical orbital signs of TED Imaging findings c/w TED Our OI, but not really. Craves is an autoinmune disease that is strongly associated with thyroid dysfunction, but it is not caused by it How is this defined? Either the pt has a known autoimmune d by smoking True; Graves thyroid condition, or s/he tests positive for rged to stop smoking one (or more) of a host of anti-thyroid antibodies Graves is associated with MS False; it is associated with MG --Eyelid retraction T and/or a Graves m There is an important exception to the 'absence of lid retraction indicates it isn't Graves dz' contention -- in fact, such pts can present with ptosis. Under what circumstance might a Graves pt present with no lid retraction, or even frank ptosis? least 'Temporal flare' likely What does temporal flare refer to/mean? Men The fact that the retraction is more pronounced at the temporal pe aspect of the lid





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by smoking True; Graves d rged to stop smoking

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with MG

Graves m

--Eyelid retraction

T and/or a

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circumstance might a Graves pt present with no lid retraction, or even frank ptosis? If the pt has concurrent myasthenia gravis ... least

likelv

Men

Gotor 'Temporal flare'

What does temporal flare refer to/mean?

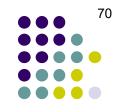
The fact that the retraction is more pronounced at the temporal aspect of the lid

pe



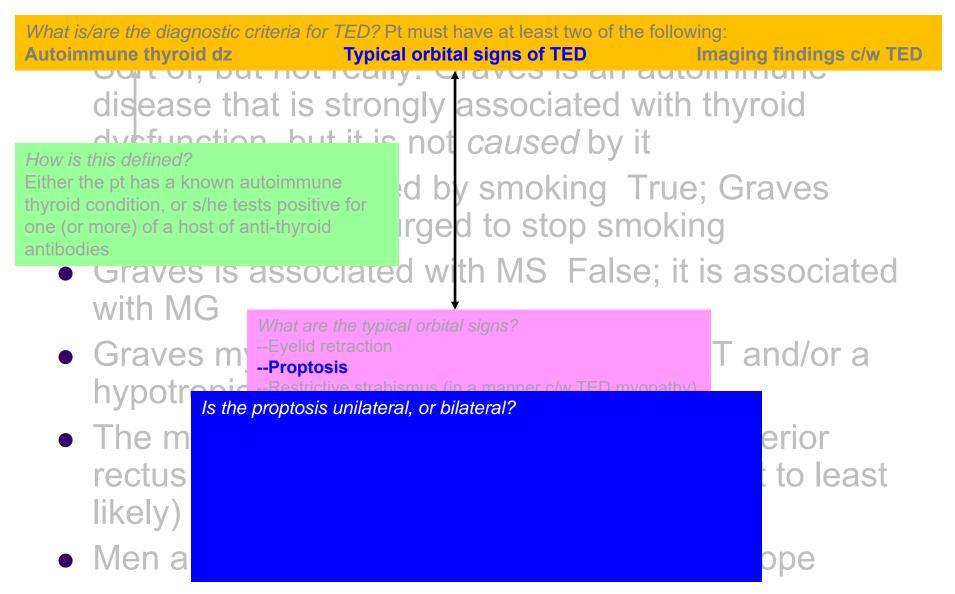
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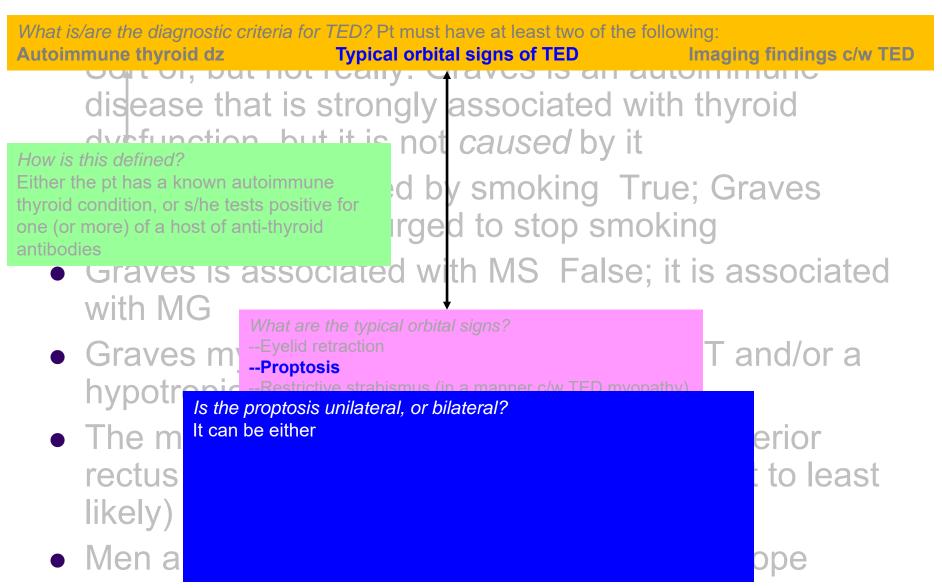
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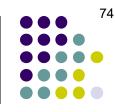
Unilateral*

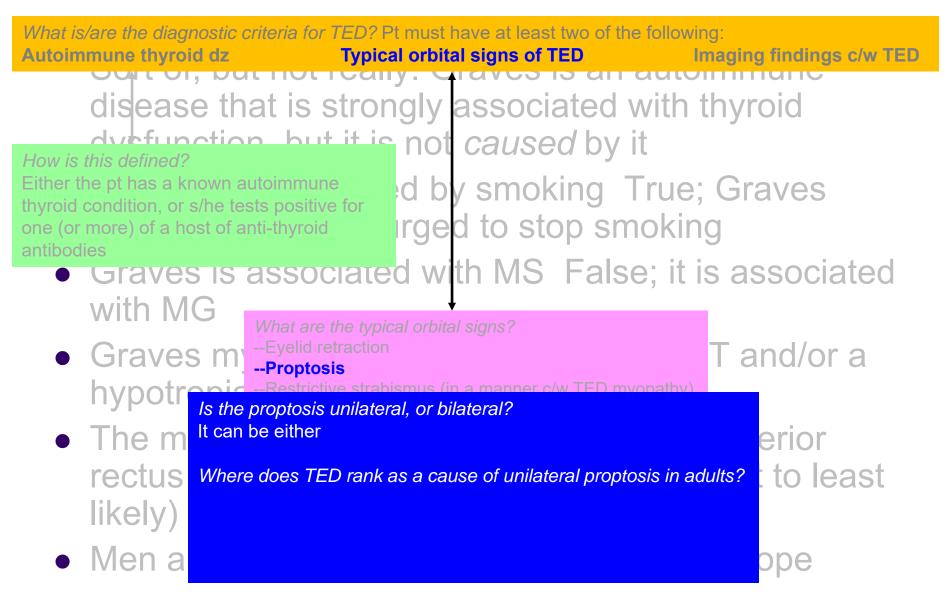


Bilateral

TED: Proptosis

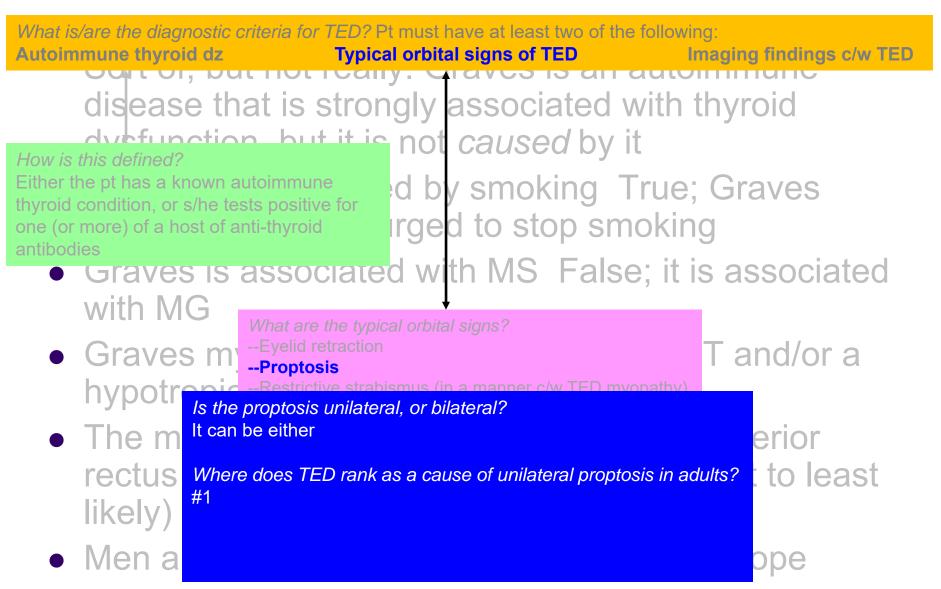
*Or highly asymmetric, at least

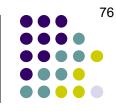


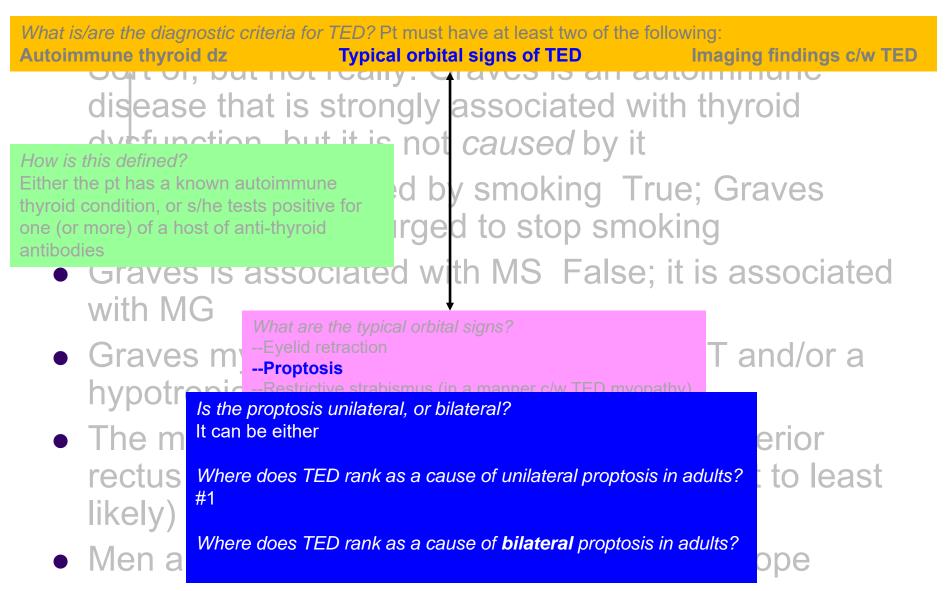






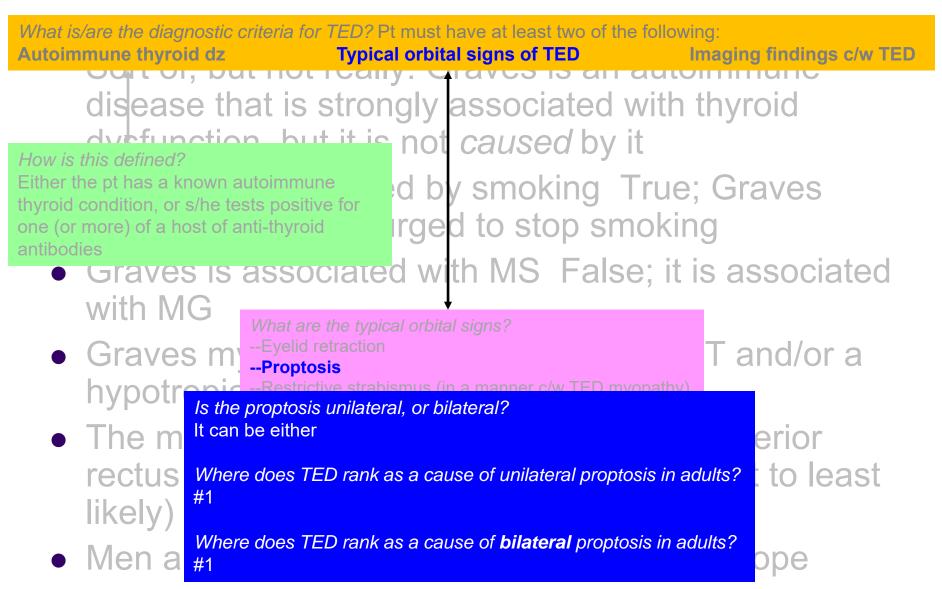


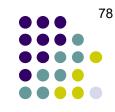


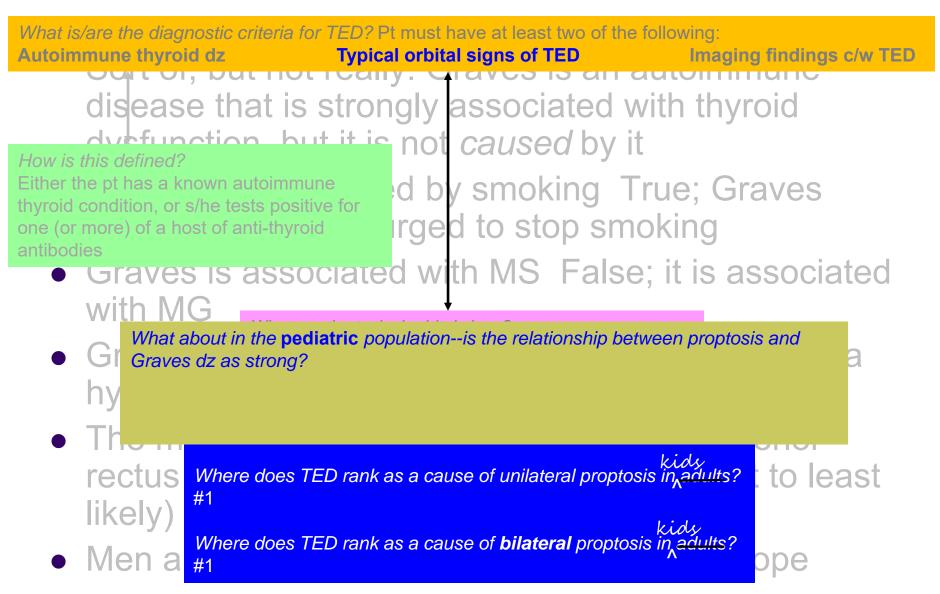






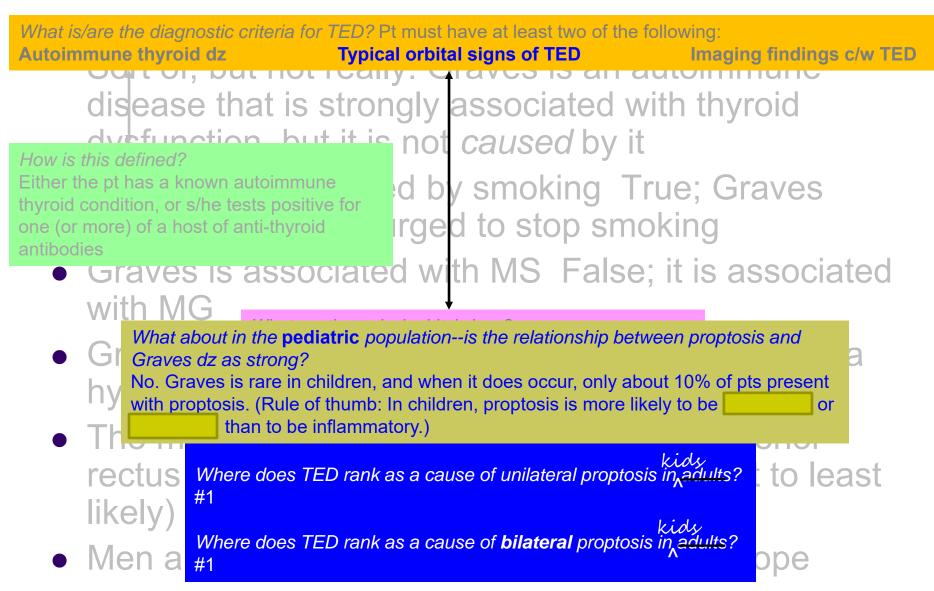




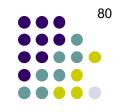


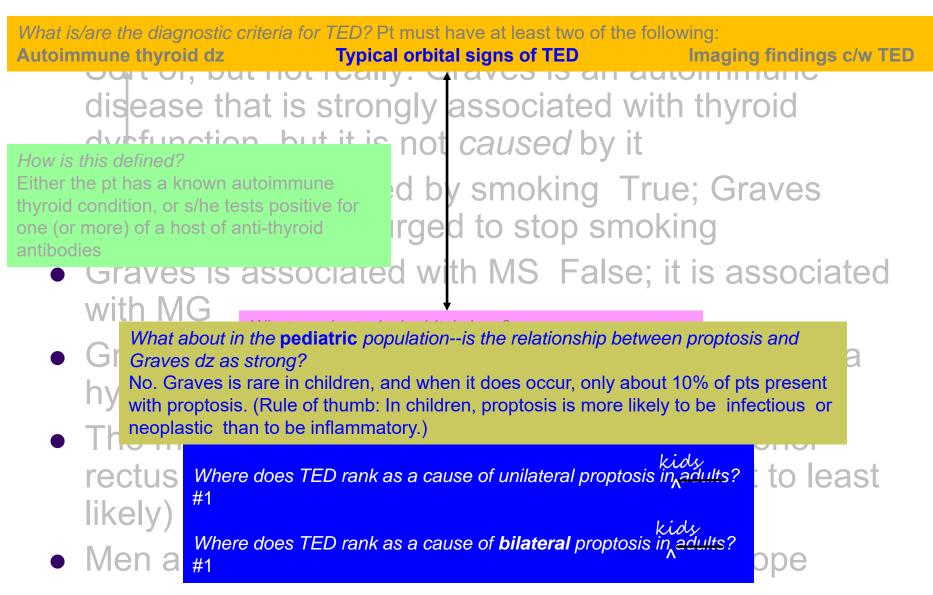














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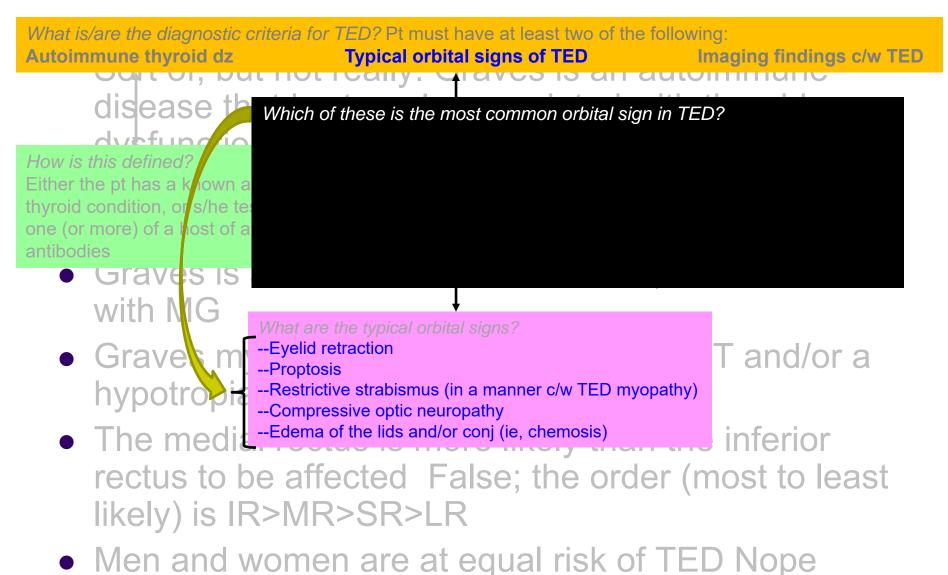
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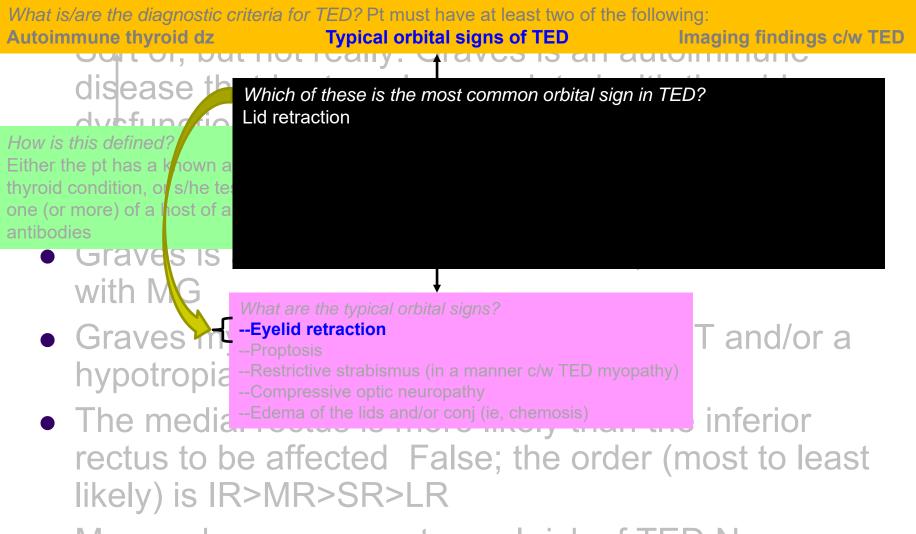
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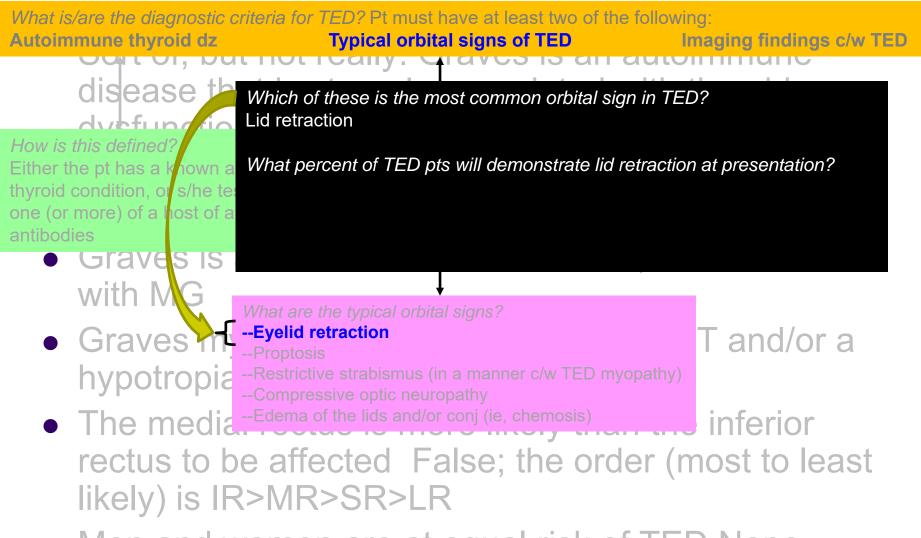






• Men and women are at equal risk of TED Nope





• Men and women are at equal risk of TED Nope

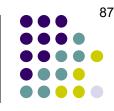


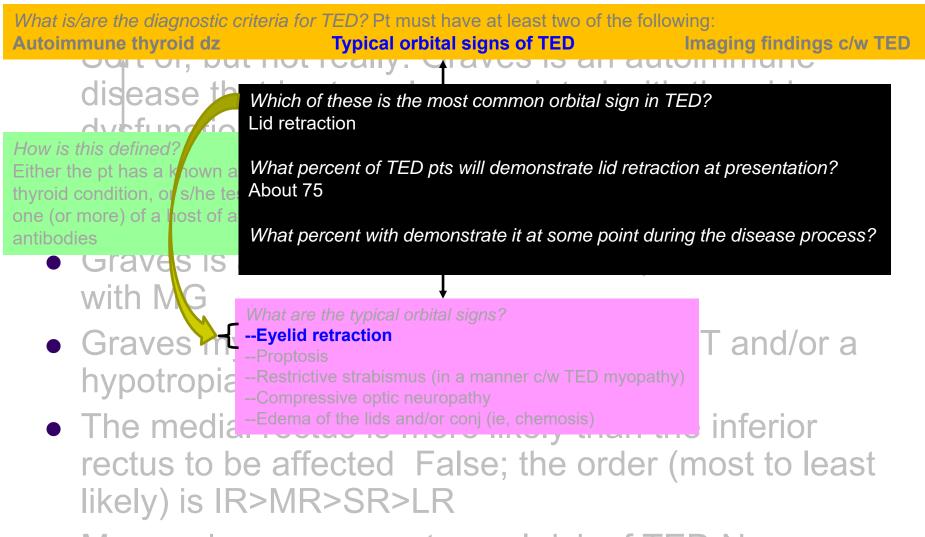


What is/are the diagnostic criteria for TED? Pt must have at least two of the following the following of the fo	maging findings c/w TED	
disease the Which of these is the most common orbital sign in TED? Lid retraction		
How is this defined? Either the pt has a known a thyroid condition, or s/he ter one (or more) of a host of a	n at presentation?	
 Graves IS 		
What are the typical orbital signs?	Tandlara	
Graves inEyelid retraction Proptosis hypotropia Restrictive strabismus (in a manner c/w TED myopathy) Compressive optic neuropathy	T and/or a	
• The mediaEdema of the lids and/or conj (ie, chemosis)	inferior	
rectus to be affected False; the order (most to least likely) is IR>MR>SR>LR		
 Mon and woman are at equal rick of TED Nana 		

• Men and women are at equal risk of TED Nope

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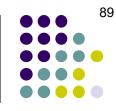
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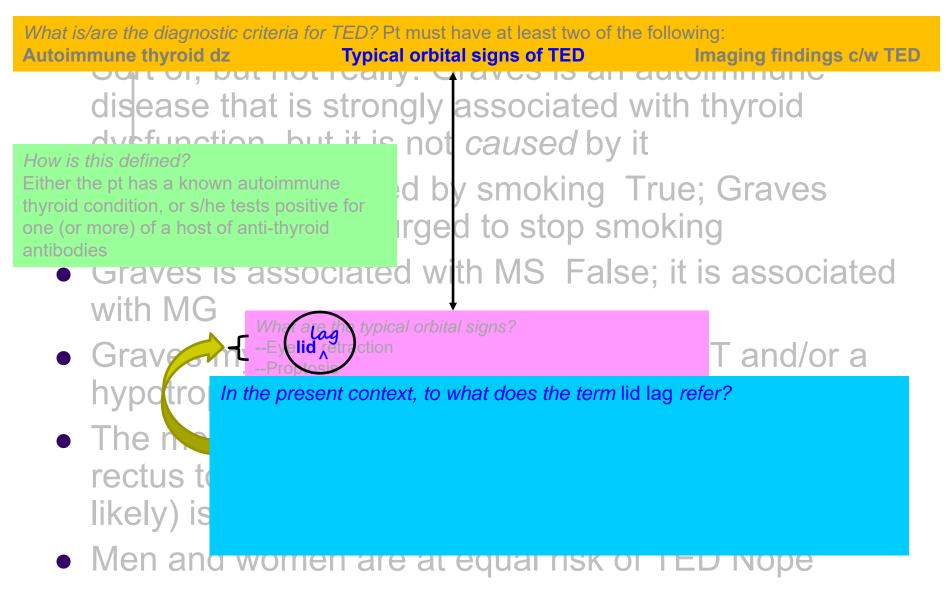




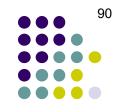
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disease the Which of these is the most common orbital sign in TED?		
How is this defined? Either the pt has a known a what percent of TED pts will demonstrate lid retraction at presentation? Thyroid condition, or s/he tere About 75		
one (or more) of a lost of a antibodies • Graves IS • Graves IS		
 What are the typical orbital signs? Graves in Proptosis 	T and/or a	
hypotropia Restrictive strabismus (in a manner c/w TED myopathy)		
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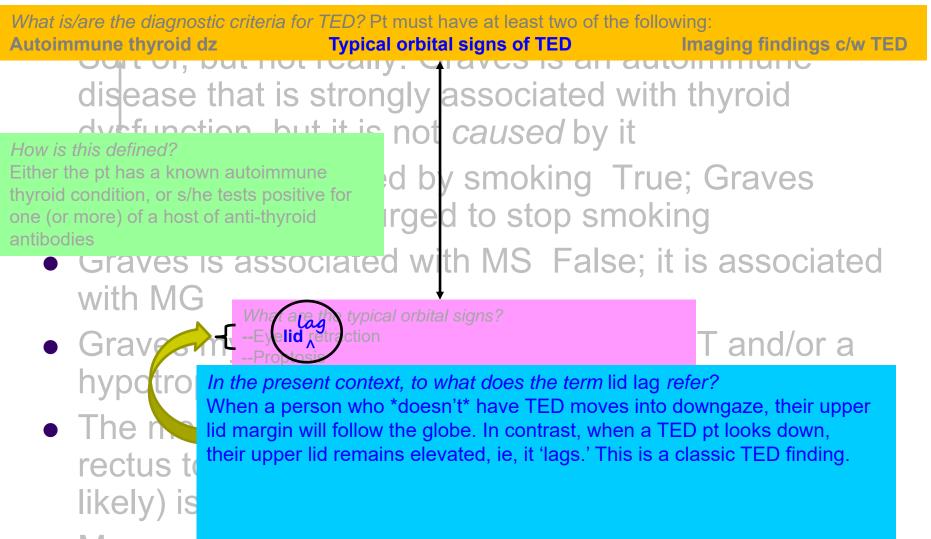
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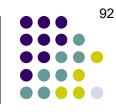


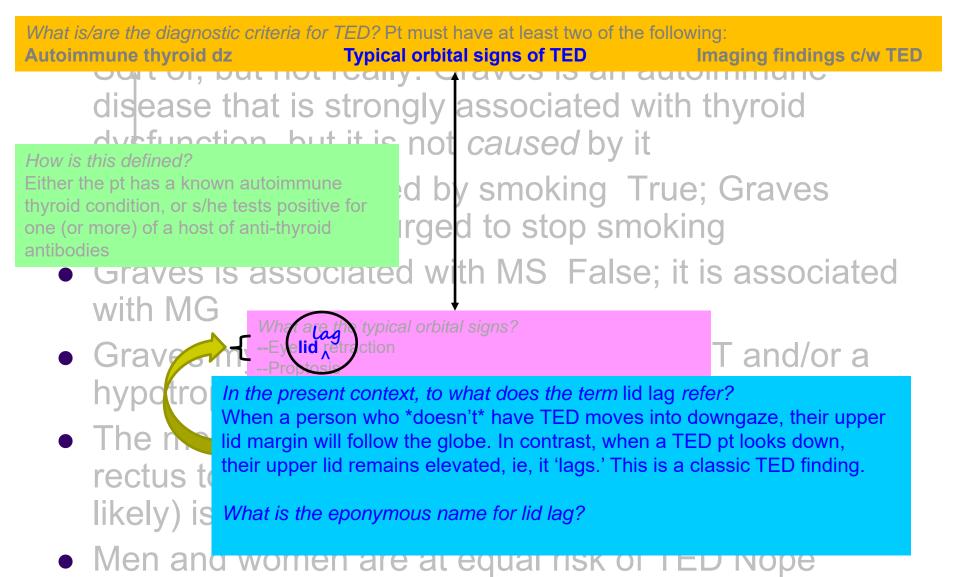


Unilateral. Note how the normal right upper lid 'follows' the eye into downgaze

Bilateral

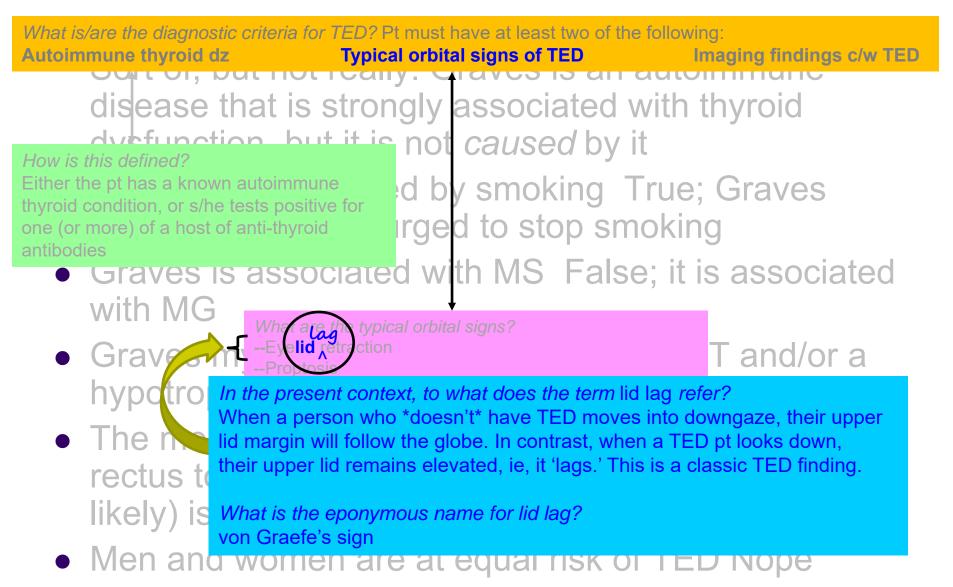
TED: Lid lag

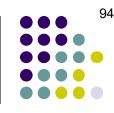


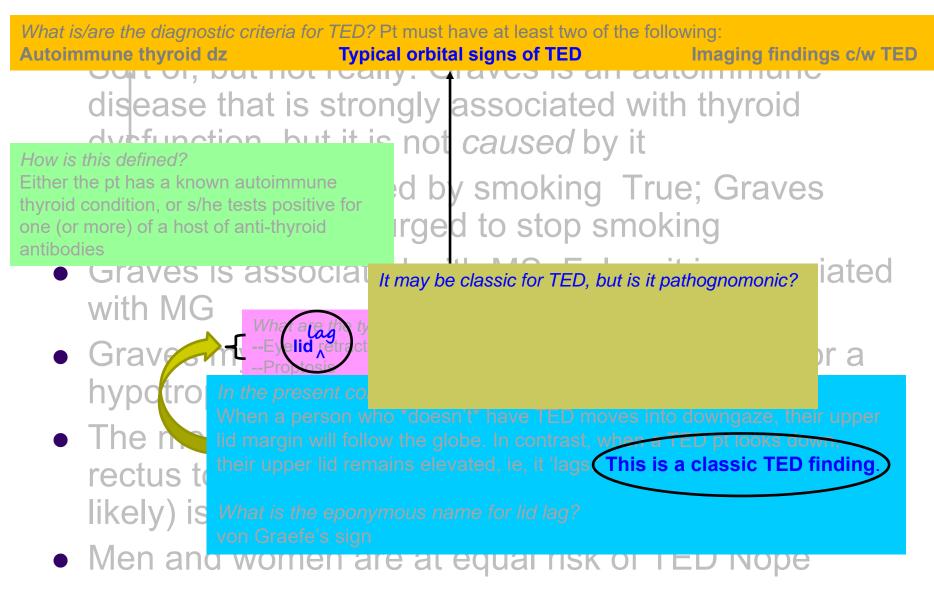






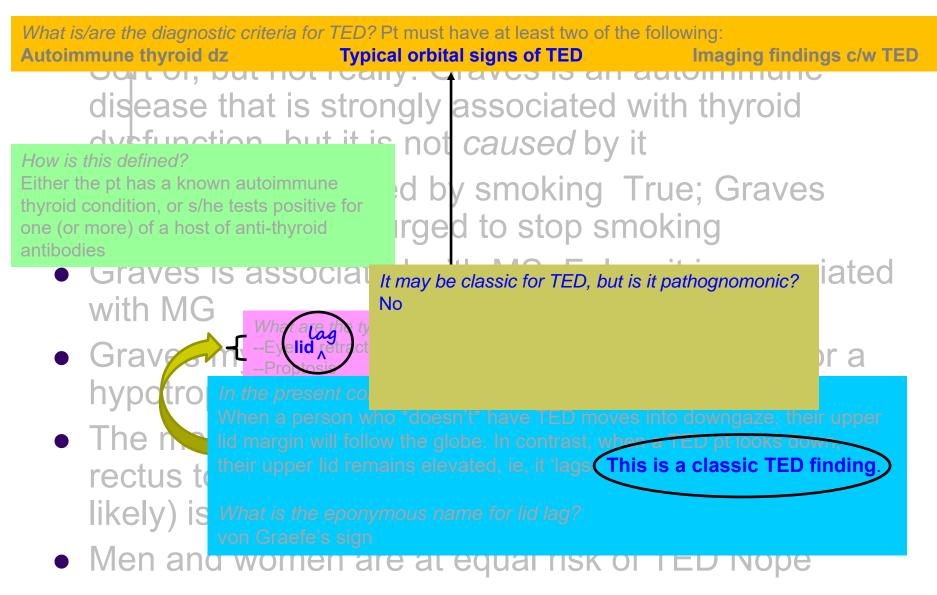




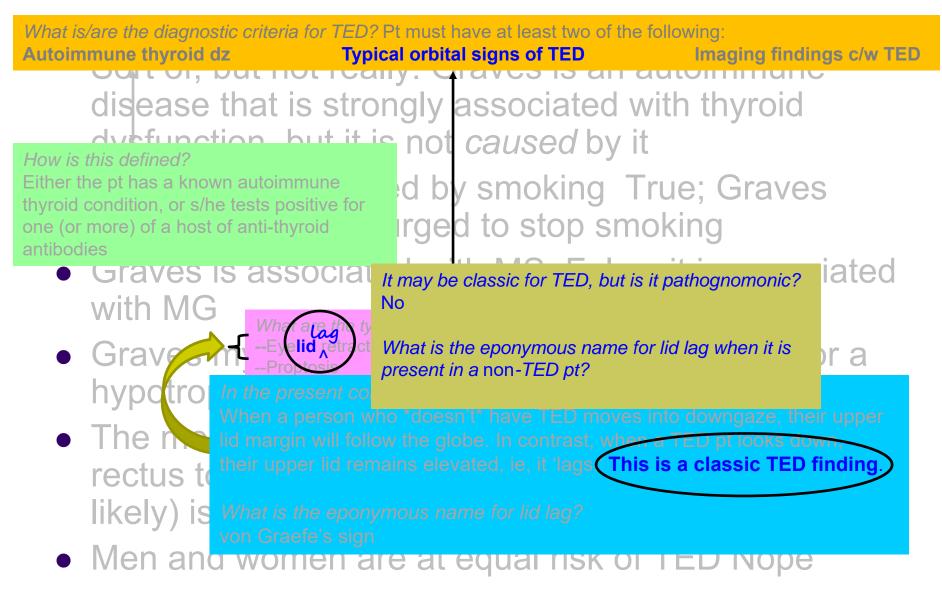






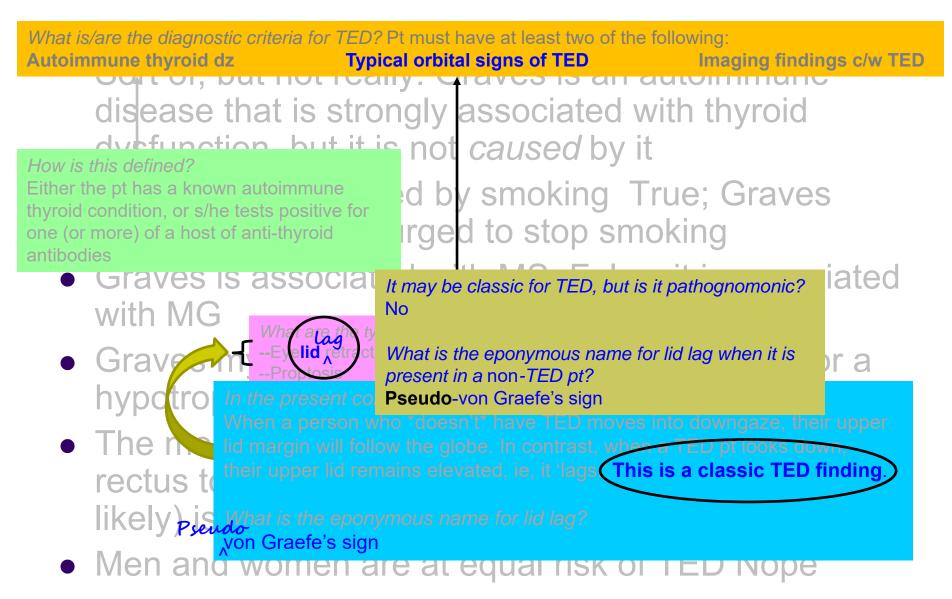








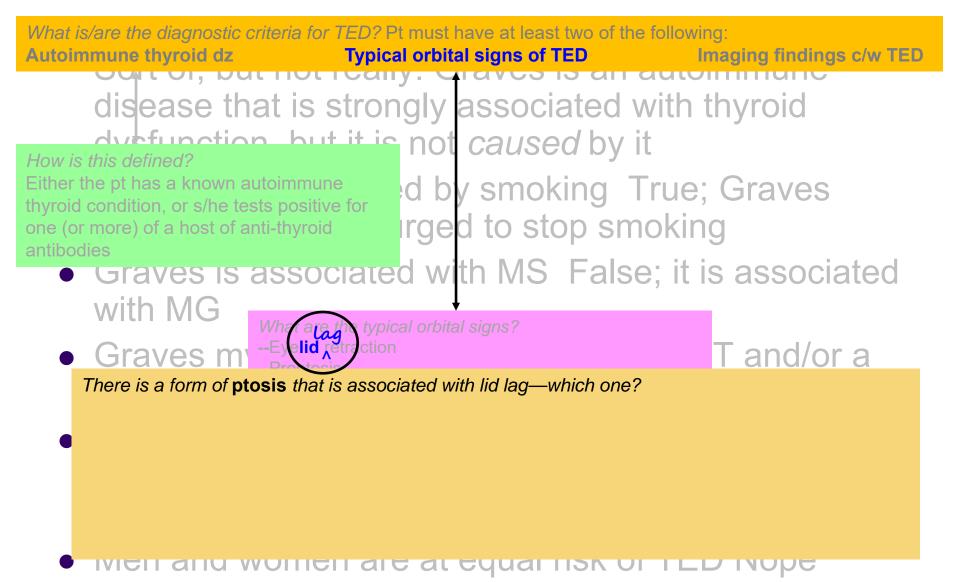




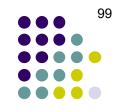
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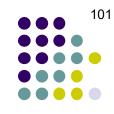
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What is/are the diagnostic criteria for TED? Pt must have at least two of the following: Autoimmune thyroid dz Typical orbital signs of TED Imaging findings c/w TED disease that is strongly associated with thyroid dyrefunction, but it is not caused by it How is this defined? Either the pt has a known autoimmune d by smoking. True: Graves

thyroid condition, or s/he tests positive for one (or more) of a host of anti-thyroid antibodies d by smoking True; Graves

- Graves is associated with MS False; it is associated with MG
- Graves m --Eyelid retraction

T and/or a

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In a nutshell, what is the pathogenesis of congenital myogenic ptosis? The levator fails to develop properly, with some or all of its muscle fibers replaced by fibrofatty tissue

OK, I can see how a lack of functioning levator leads to ptosis, but why do these pts have lid lag?

wen and women are at equal lisk of TED Nope





What is/are the diagnostic criteria for TED? Pt must have at least two of the following:Autoimmune thyroid dzTypical orbital signs of TEDImaging findings c/w TEDOut COT, DOT HOLT CATTY: Catty COT S and autoint for the following:

disease that is strongly associated with thyroid dynamics but it is not caused by it

How is this defined?

Either the pt has a known autoimmune thyroid condition, or s/he tests positive for one (or more) of a host of anti-thyroid antibodies d by smoking True; Graves

- Graves is associated with MS False; it is associated with MG
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 Protocia

T and/or a

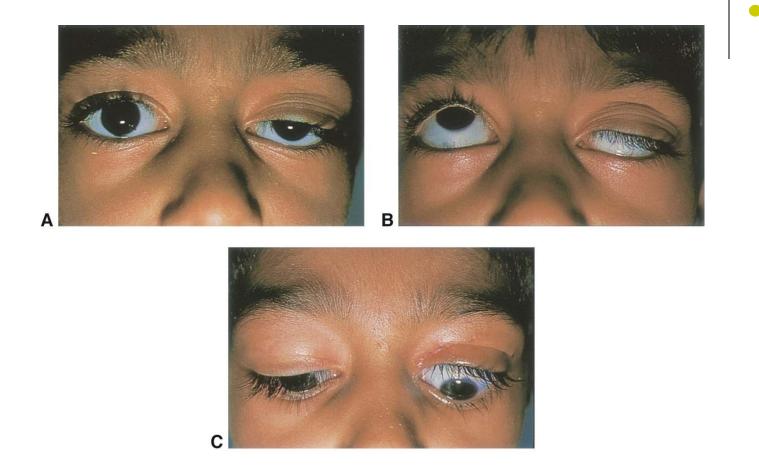
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In a nutshell, what is the pathogenesis of congenital myogenic ptosis? The levator fails to develop properly, with some or all of its muscle fibers replaced by fibrofatty tissue

OK, I can see how a lack of functioning levator leads to ptosis, but why do these pts have lid lag? Because the fibrofatty tissue can neither contract (causing ptosis) *nor* relax (causing lid lag)

wen and women are at equal lisk of TED hope

105



Congenital myogenic ptosis. A, ptosis. B, failed elevation in upgaze. C, lid lag in downgaze

Thursd Fue Diseas What well-known phenomenon associated with a cranial-nerve palsy frequently manifests with lid lag?

VVILI IVI

Wha --Eyelid retraction Graves m

typical orbital signs? aret



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What are the classic CN3 aberrant regeneration mis-connections?

--Attempted adduction or depression → eyelid retraction

--Attempted globe adduction, elevation or depression → miosis of the pupil

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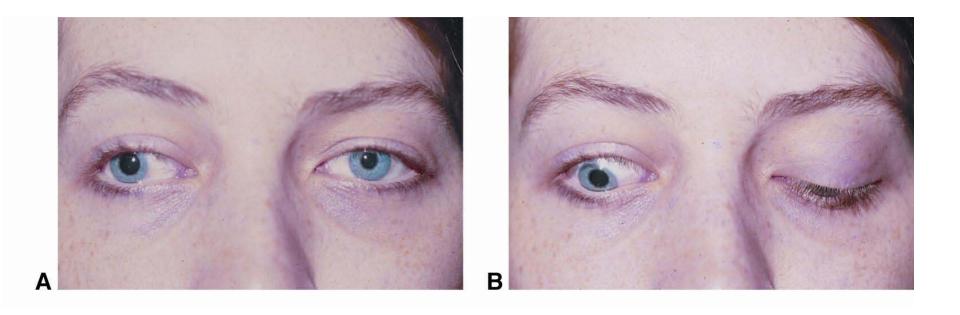
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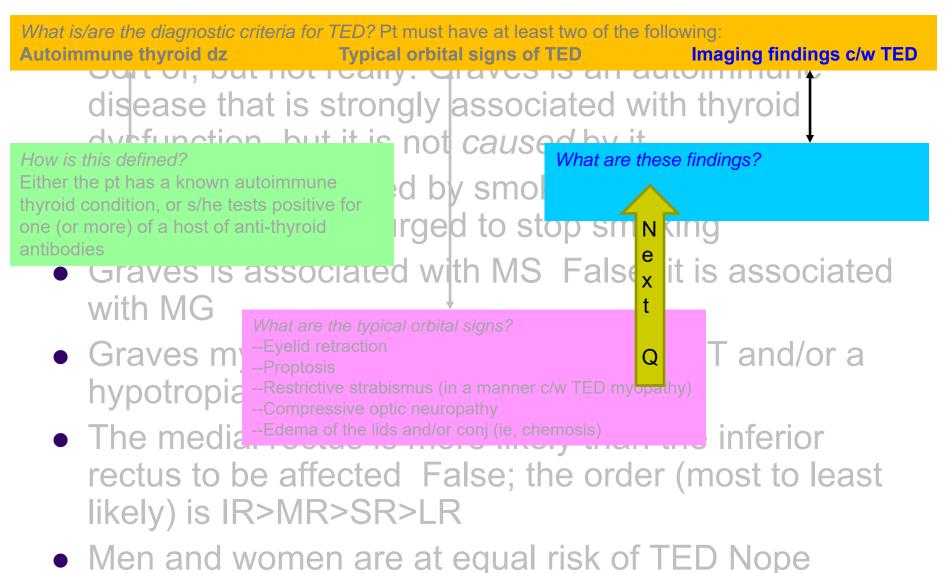
Aberrant regeneration of the right third nerve. A, In primary gaze, there is mild ptosis, pupillary mydriasis, and exotropia, all on the right.

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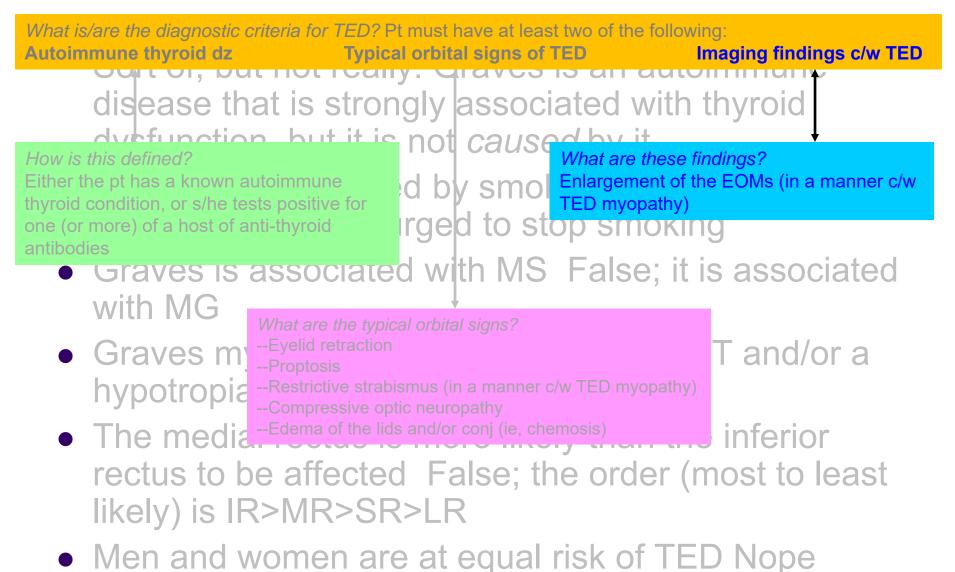
Aberrant regeneration of the right third nerve. A, In primary gaze, there is mild ptosis, pupillary mydriasis, and exotropia, all on the right. B, With attempted downward gaze, the right eyelid retracts as fibers of the right third nerve supplying the inferior rectus now also innervate the levator muscle



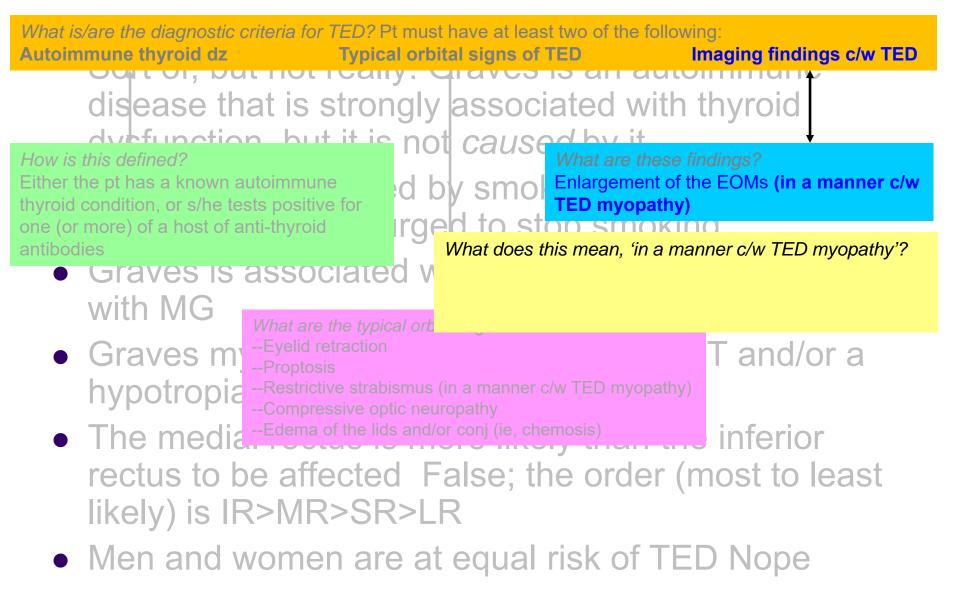




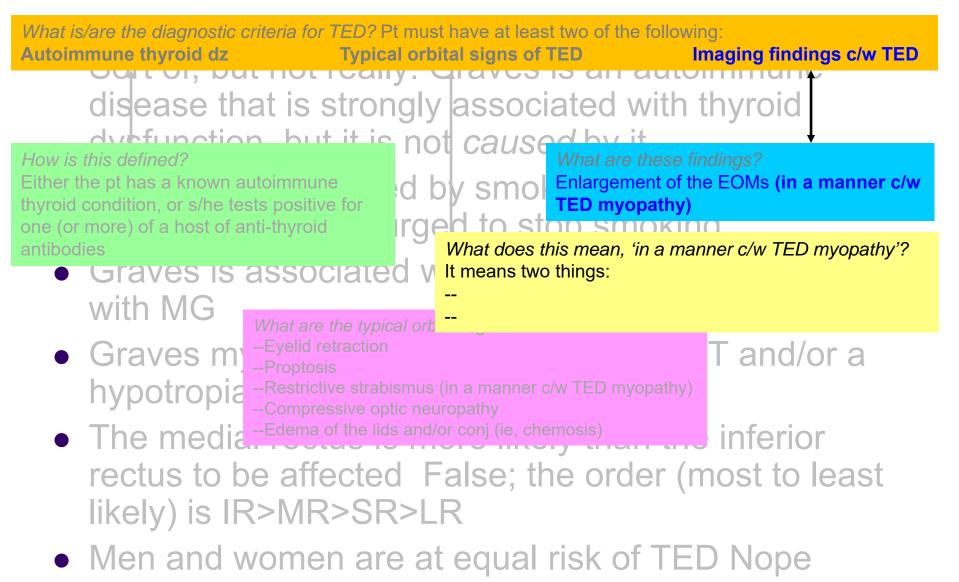






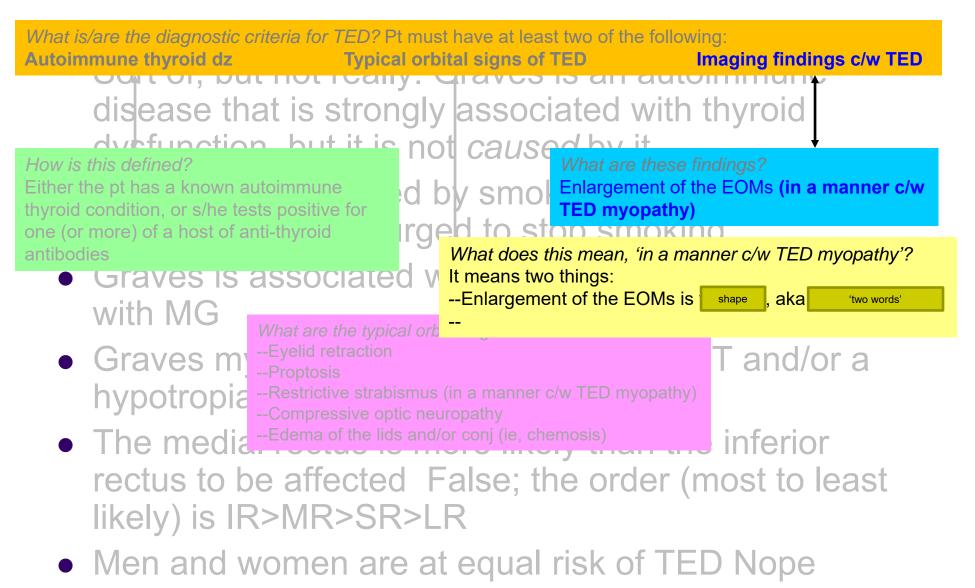






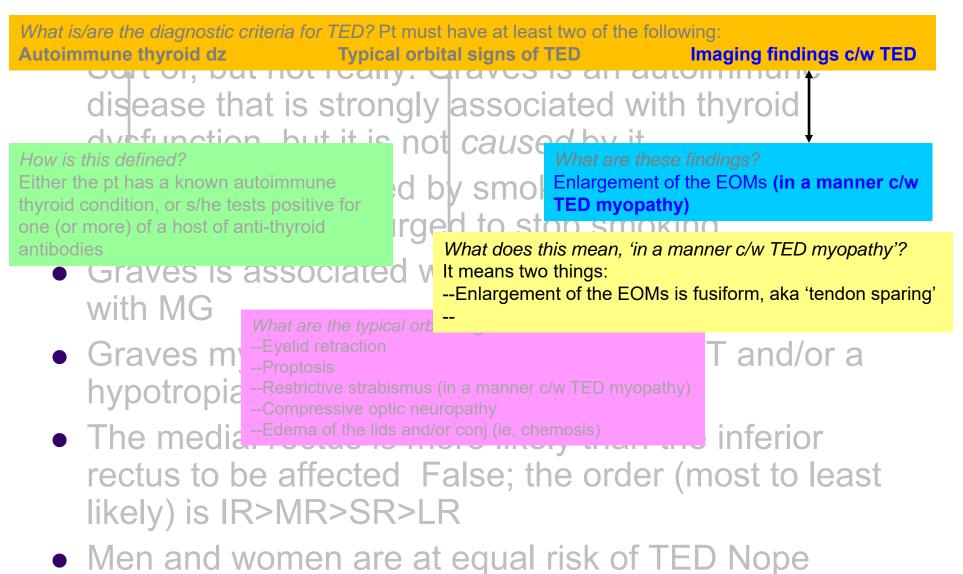










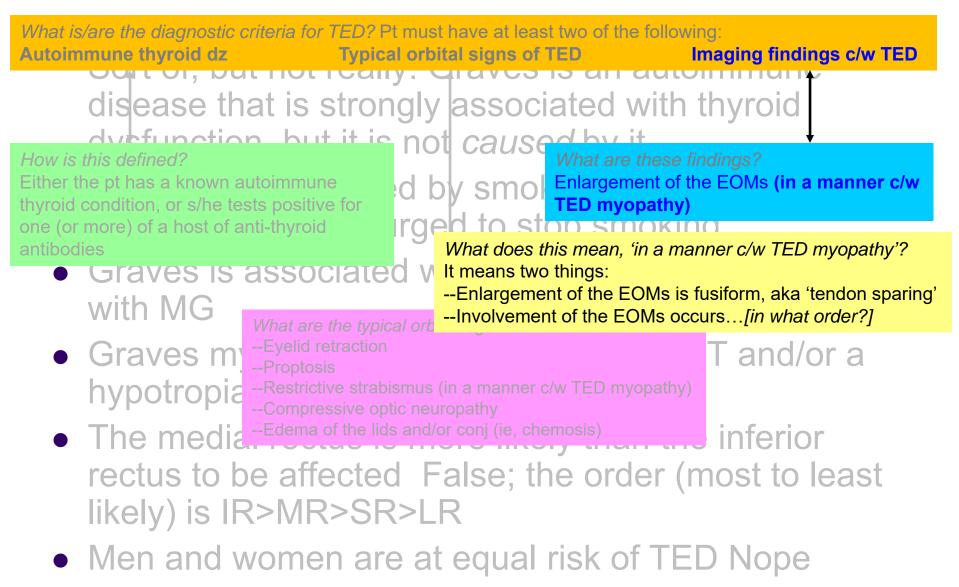






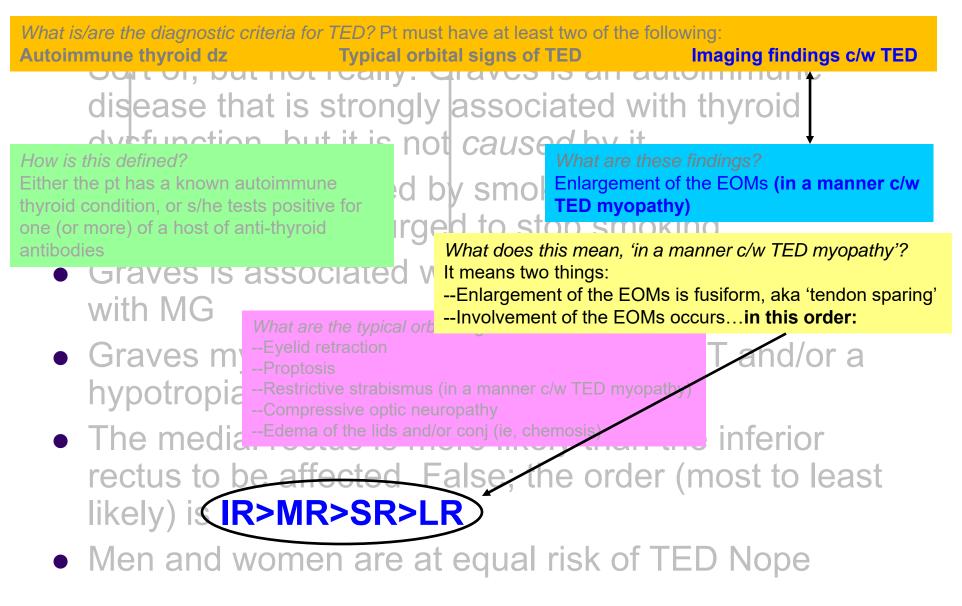
TED: Tendon-sparing EOM enlargement

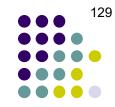


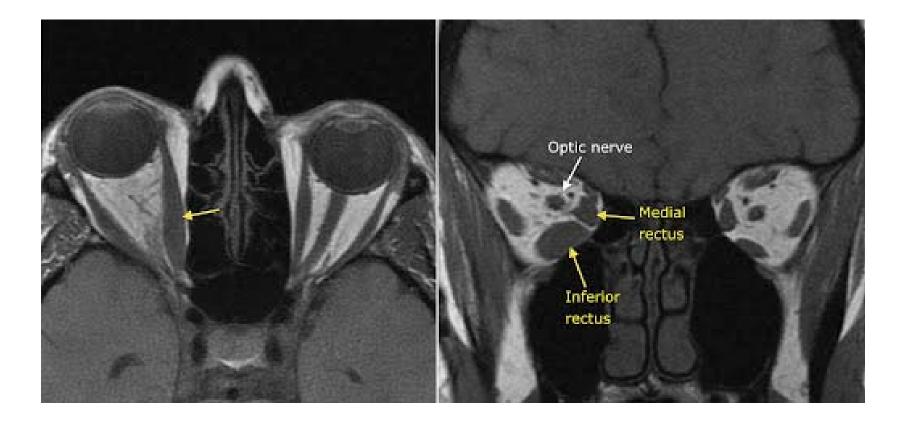






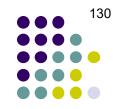






TED: Tendon-sparing EOM enlargement involving the IR > MR

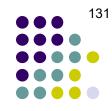




What is the natural history of TED?

- Graves myopathy usually results in an ET and/or a hypotropia True
- The medial rectus is more likely than the inferior rectus to be affected False; the order (most to least likely) is IR>MR>SR>LR
- Men and women are at equal risk of TED Nope





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It is a self-limited disease that tends to 'burn itself out' over time, but may leave disfiguring sequelae

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On average, how long does it take to burn itself out?

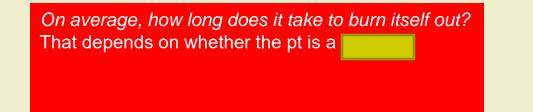
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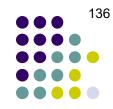
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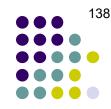
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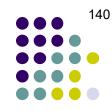
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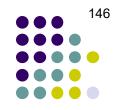




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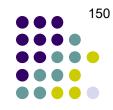




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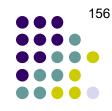


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Severe disease		and/or optic
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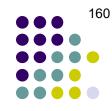
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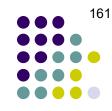
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If surgery during active TED is relatively contraindicated, when should it be considered? Once the pt is one word, and his/her orbital condition has been stable for at least amount of time





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If surgery during active TED is relatively contraindicated, when should it be considered? Once the pt is euthyroid, and his/her orbital condition has been stable for at least 6 months



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Suppose a pt meeting those criteria needs orbital decompression, has strabismus, and also has malpositioned eyelids. What should the overall surgical strategy be?

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Suppose a pt meeting those criteria needs orbital decompression, has strabismus, and also has malpositioned eyelids. What should the overall surgical strategy be?

Because orbital-decompression surgery will affect both EOM and lid position, the decompression surgery must precede strabismus and/or lid surgery. Likewise, because strabismus surgery will affect lid position, the strabismus surgery must precede the lid surgery.





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Because orbital-decompression surgery will affect both EOM and lid position, the decompression surgery must precede strabismus and/or lid surgery. Likewise, because strabismus surgery will affect lid position, the strabismus surgery must precede the lid surgery. <u>Putting it all together, the surgical order is as follows</u>:

- 1. Decompression
- 2. Strabismus
- 3. Eyelids



What is the natural history of TED? It is a self-limited disease that tends to 'burn itself out' over time, but may leave disfiguring sequelae

How is TED managed?

What about treating the concurrent thyroid disease?

--Severe disease involves signt-inreatening complications such as corneal decompensation, and/or optic neuropathy. This necessitates more aggressive steroid therapy, and consideration should be given to the use of steroid-sparing IMT and/or orbital radiation therapy.

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- What about treating the concurrent thyroid disease?
- Rendering the pt euthyroid is an important goal for TED pts (to be pursued by Endo/IM, of course). However, it is important to note that some treatments in this regard can
- --/ aggravate the TED by increasing immune activity against the TSH-receptor antigens that are responsible for TED.

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