

INTEGRATED RHEUMATOLOGY

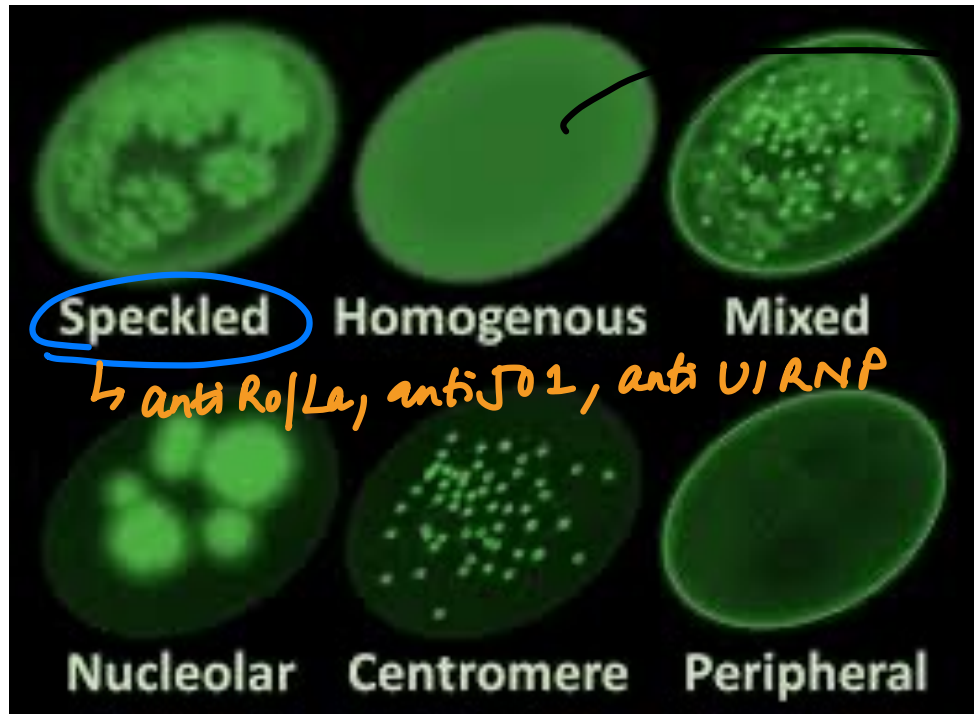
Auto-Antibodies

- ANA → most sr SLE → entry criteria (>1:80)
- Anti-C1q > Anti-dsDNA : correlate \bar{c} disease activity
- Anti-histone DRUG induced lupus → SHIP
- Anti-Sm -most specific
- Antineuronal Ab/ Antiglutamate receptor 2 : CNS Lupus
- Anti-ribosomal P : psychiatric manif SLE
- Anti-U1 RNP : MCTD
- Anti-Ro/SS-A } Sjogren's D → Cong & blocks risk ↑
- Anti-La/SS-B }
- Anti-Jo1 (Anti-synthetase) → Antisynthetase Sx
- Anti-Mi2 : Dermatomyositis → good prognosis
- Anti-topoisomerase 1 / Anti-scl-70/anti-RNA polymerase III : Diffuse scleroderma
- Anti-centromere : Limited SSc = CREST mechanic hands
- p-ANCA : MPA, microscopic polyangiitis, Churg Strauss, UC
- c-ANCA : Wegener's granulomatosis (GPA) → EGPA
- RF : IgM against Fc-IgG RA: most sp-anti-CCP
- Anti-endothelial cell Ab → Kawasaki
- Anti-alpha-enolase Ab → Behcet's



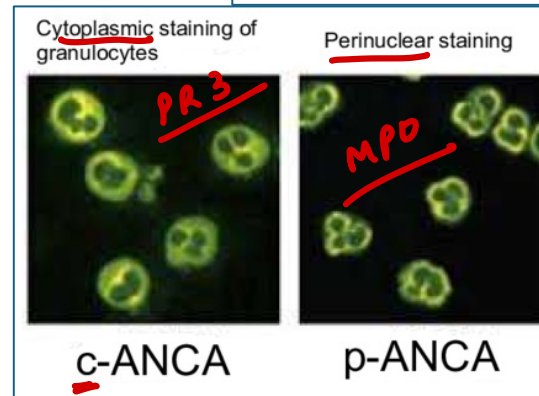
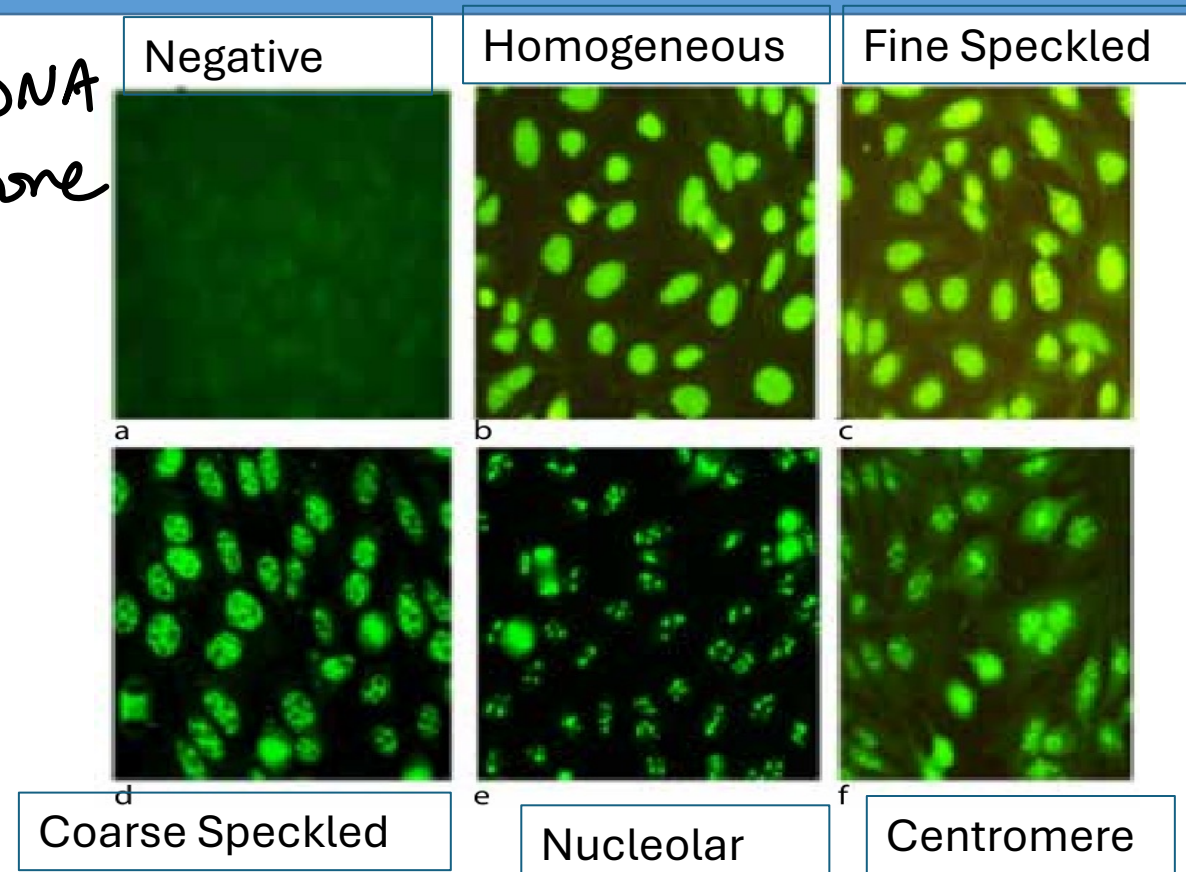
mechanic hands

Auto-Antibodies-Fluorescence pattern



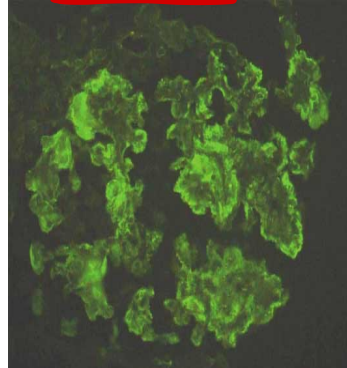
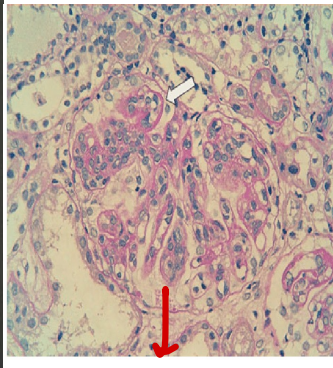
Anti-dsDNA
 Anti-histone

Anti-topo / Scl-70 scleroderma
 CREST
 anti-Sm



New EULAR/ACR criteria for the classification of SLE >10

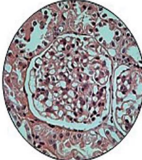
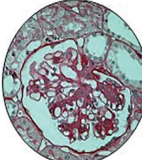
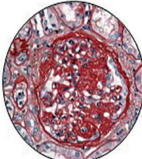
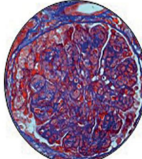
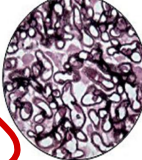
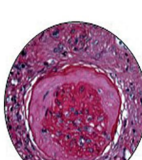
ANA +

Clinical domains	Points	Immunologic domains	Points	
Constitutional domain Fever	2	Antiphospholipid antibody domain Anticardiolipin IgG > 40GPL or anti-B2GP1 IgG > 40 units or lupus anticoagulant	2	
Cutaneous domain Non-scarring alopecia Oral ulcer Subacute cutaneous or discoid lupus	2	Complement proteins domain Low C3 or Low c4 Low C3 and Low c4	3	
	2		4	
Arthritis domain "Jaccoud" Synovitis tenderness in a at least 2 joints non-erosive	6	Highly specific antibodies domain Anti- dsDNA antibody ✓ Anti- Sm antibody ✓	6	
Neurologic domain Delirium Psychosis Seizure	2 3 5			
Serositis domain Pleural or pericardial effusion Acute pericarditis	5 6			
Hematologic domain Leukopenia Thrombocytopenia Autoimmune hemolysis Iga "warm" DCT (+)	3 4 4	IgG, IgM, IgA, C3, C1q	FULL HOUSE effect	
	Renal domain Proteinuria >0.5g/24hr Class II or V lupus nephritis Class III or IV lupus nephritis			4 8 10

- False- positive VDRL/RPR anti-cardiolipin
- Prolonged PTT that is not corrected by the addition of normal platelet-free plasma
↳ Lupus anticoagulant

Steroids + MMF/Cyclophosphade

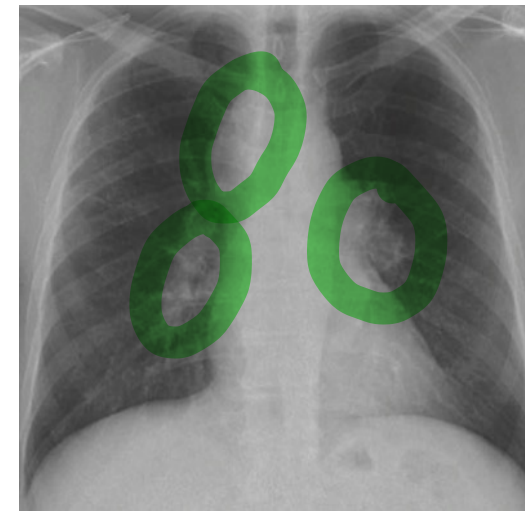
HISTOPATHOLOGICAL CLASSIFICATION OF LUPUS NEPHRITIS

 <p>Class I Minimal Mesangial Lupus Nephritis Deposition of immune complexes detectable by immunofluorescence techniques.</p>	 <p>Class II Mesangial Proliferative Lupus Nephritis Mesangial hypercellularity of any degree or mesangial matrix expansion with immune deposits detectable by light microscopy.</p>
 <p>Class III QQ Focal Lupus Nephritis Active or inactive focal, segmental or global endo/extracapillary glomerulonephritis involving <50% of all glomeruli. Manifestations include active lesions (A), chronic inactive lesions (C) or active and chronic lesions (A/C)</p>	 <p>Class IV QQ Diffuse Lupus Nephritis Active or inactive diffuse, segmental or global endo/extracapillary glomerulonephritis involving ≥50% of all glomeruli. Subendothelial diffuse immune deposits, with or without mesangial alterations, are common. This class is also divided in: diffuse segmental (IV-S), when ≥ 50% of the involved glomeruli have segmental lesions, and diffuse global (IV-G), when ≥ 50% of the involved glomeruli have global lesions. It can also manifest A, C or A/C lesions.</p>
 <p>Class V Membranous Lupus Nephritis Global or segmental subepithelial immune deposition or their morphologic sequelae detectable by light, immunofluorescence or electron microscopy, with or without mesangial alterations. It can occur in combination with class III or IV and it can manifest advanced sclerosis.</p>	 <p>Class VI Advanced Sclerosis Lupus Nephritis Lupus Nephritis with terminal prognosis. 90% of the glomeruli in global sclerosis.</p>

wire loop lesions

SLE Nephritis

Sarcoidosis



1-2-3 /
Garland
sign /
Lambda
sign (Ga)

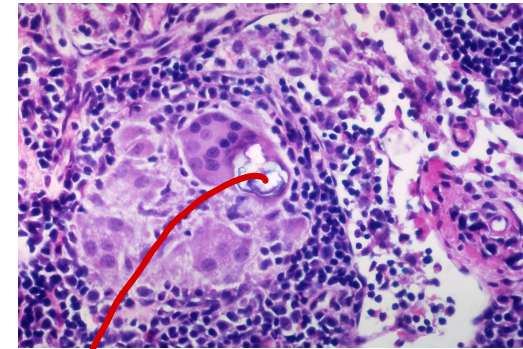


EN



Lupus pernio

Id OK (+)



NC granulomas
Schaumann
Asteroid bodies



Ga scan
PANNA

Scadding

Stage 0	Normal
Stage 1	Bilateral hilar adenopathy, normal lung parenchyma
Stage 2	Hilar adenopathy with pulmonary infiltration
Stage 3	Pulmonary infiltration without hilar adenopathy
Stage 4	Pulmonary fibrosis

Lab:
ACE- (↑) → granulomatous
Calcium- (↑) / hypercalcemia
Lymphopenia (N): 2:1
BAL: CD4: CD8 → 3.5-5:1
Kveim test

HIV INFECTION	SJOGREN'S SYNDROME	SARCOIDOSIS
BLEL <small>Benign Lymphoepithelioid Lesions</small>		
Lack of antibodies	Presence of autoantibodies	Lack of autoantibodies
Lymphoid infiltrate of salivary glands by CD8+ T lymphocytes	Lymphoid infiltrate of salivary glands by CD4+ T lymphocytes	Granulomas in salivary glands
Association with HLA-DR5	Association with HLA-DR3 and DRw52	Unknown
Positive serologic test	Negative serologic test for HIV	Negative serologic test for HIV

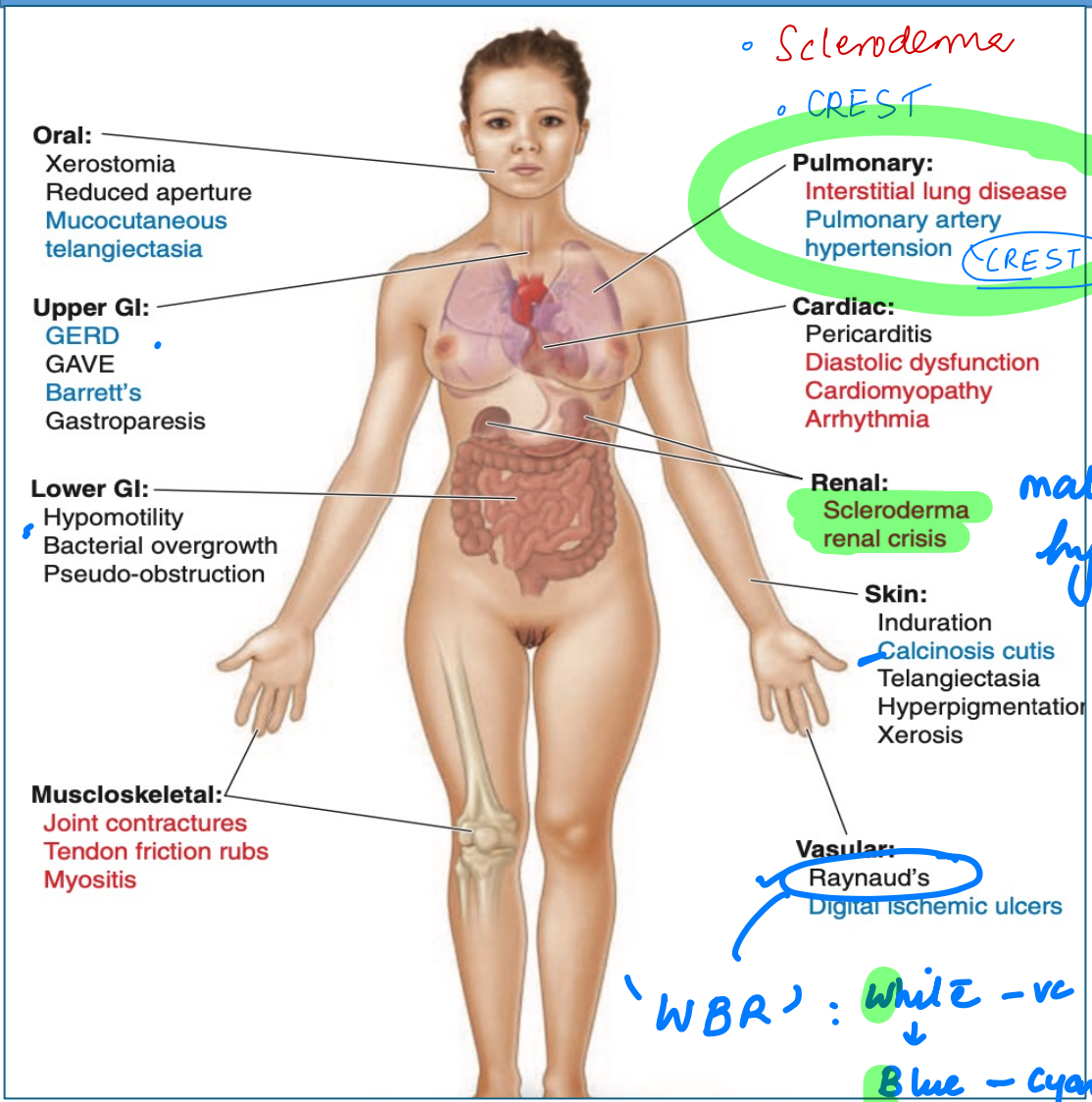
Sialosis
↓
alcoholics

Lofgren syndrome: LN + EN

Heerfordt syndrome: uveoparotid fever + 7M CN palsy

Scleroderma-CREST

Raynaud's
CREST
Esophageal



Calcinosis cutis



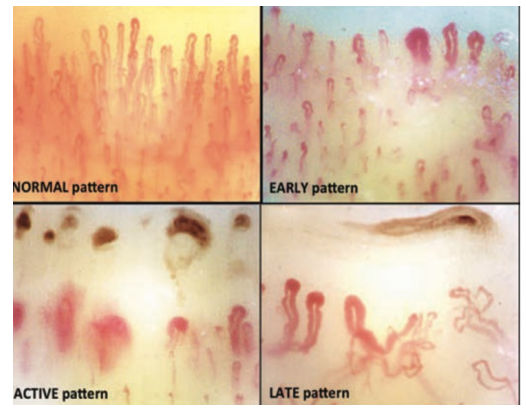
sclerodactyly



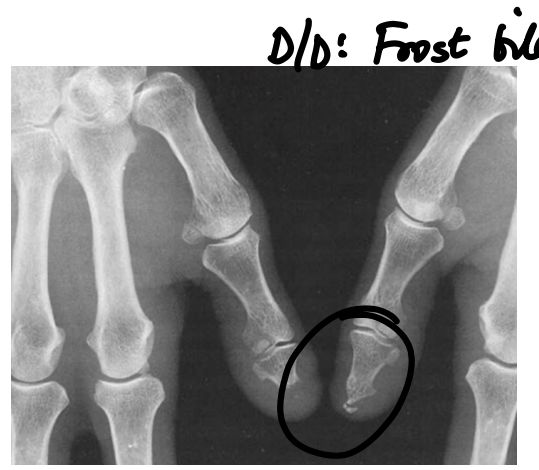
telangiectasia

malignant
hyp-hyp bitten
R-ACE ⊖

'WBR': White - ve
↓
Blue - cyanosis
↓
Red - vd



nail fold capillaroscopy



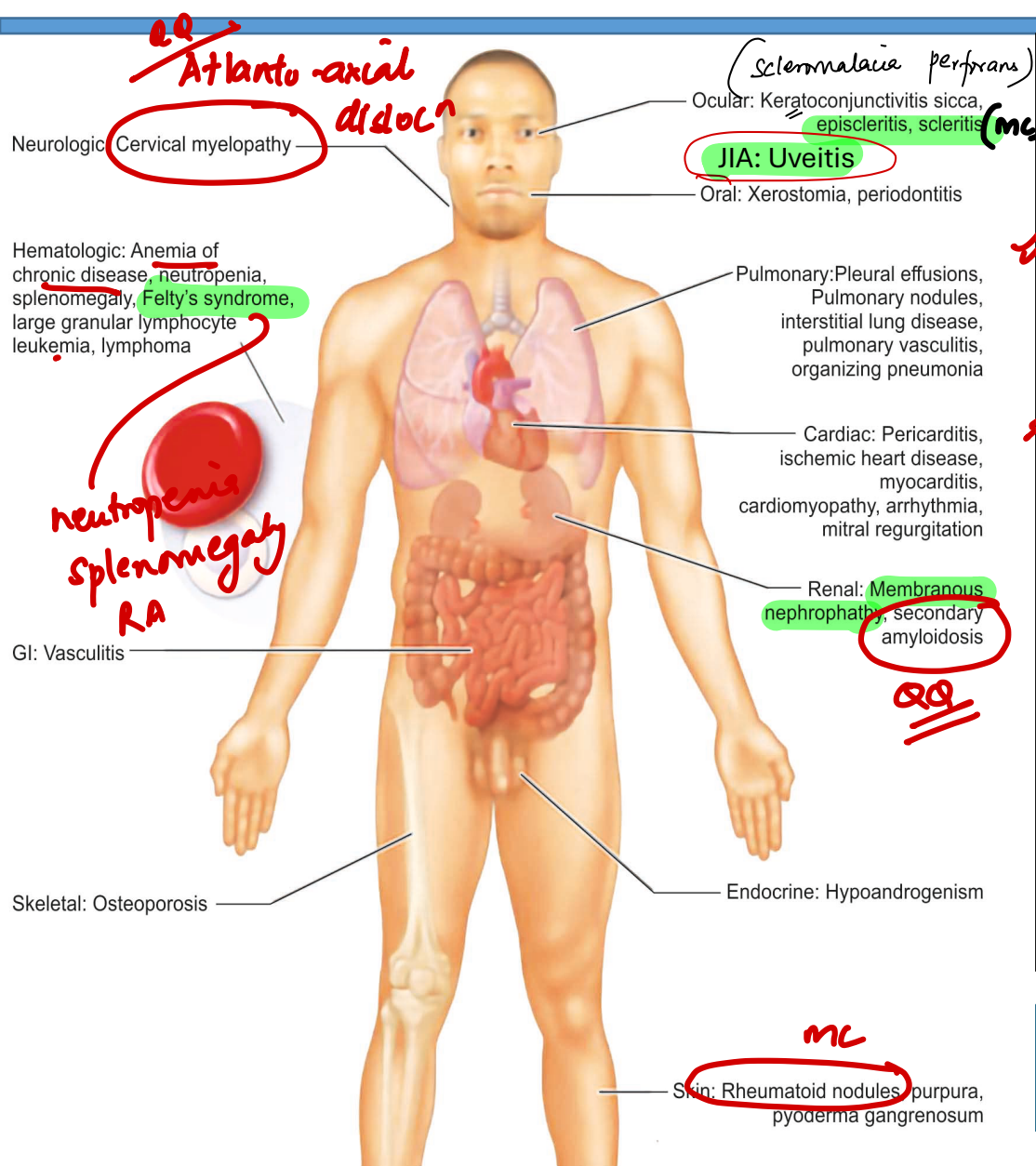
acro-osteolysis

D/D: Frost bite

Anti-fibrillar / Anti-U3 RNP: prognostic → ↑ RPGN / PAH / ILD

Rheumatoid Arthritis

JIA → < 16 yrs



RA: NSAIDs → ~~x~~ → C-DMARDs → ~~x~~ → Biologicals

DOC (circled) → Mtx (circled) + folic acid

DOC in pregnancy: Leftunonide

HCO: Irreversible retinal damage, cardiotoxicity ^{slc}

Sulfasalazine: Male infertility, CI in G6PD

BIOLOGICALS in RA

TNF- α inhibitors:

- Adalimumab
- Certolizumab
- Etanercept: fusion protein
- Infliximab
- Golimumab

IL-1R Antagonist: Anakinra

IL-6 inhibitor: Tocilizumab, Sarilimumab

B-cell depletor/CD20-: Rituximab (also Hep B)

Co-stimulation inhibitor/CTLA4-: Abatacept

JAK inhibitors: Tofacitinib, Baricitinib, Upadacitinib

2/0 - (TB) exposure, Mantoux test / PPD, Quantiferon GOLD = (IGRA (IFN- γ release assay) BLOOD test)

FLARES (circled) ↓

Ank spondylitis: xx DMARDs

NSAIDs → Biologicals

DOC

GOUT-PSEUDOGOUT

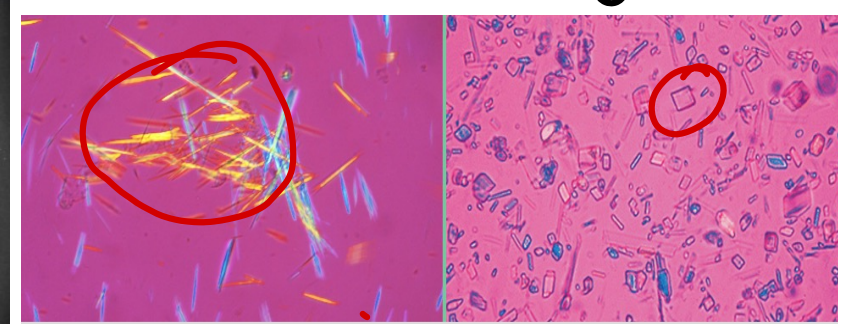
S. aureus

	Normal	OA	Inflammatory arthritis RA	Septic arthritis
Appearance	Clear	clear	turbid	turbid
Viscosity "stringing"	Normal	↓	↓	↓
WBCs	<200	200 - 5k	5k - 50k	>50k
PMNs	<25%	<25%	25-50%	>50%

Acute Gout
 NSAIDS DOC except *aspirin*
 Colchicine *most effective*
 Chronic gout DOC: **ALLOPURINOL**
 HLA B5*801 *hsv*
 AVOID WITH: **GMP, Azathioprine**

side: hepatotoxic, chemolaxis, diarrhea

Polyarthralgia + Tenosynovitis + Dermatitis : *Pustules* **Disseminated gonorrhea**



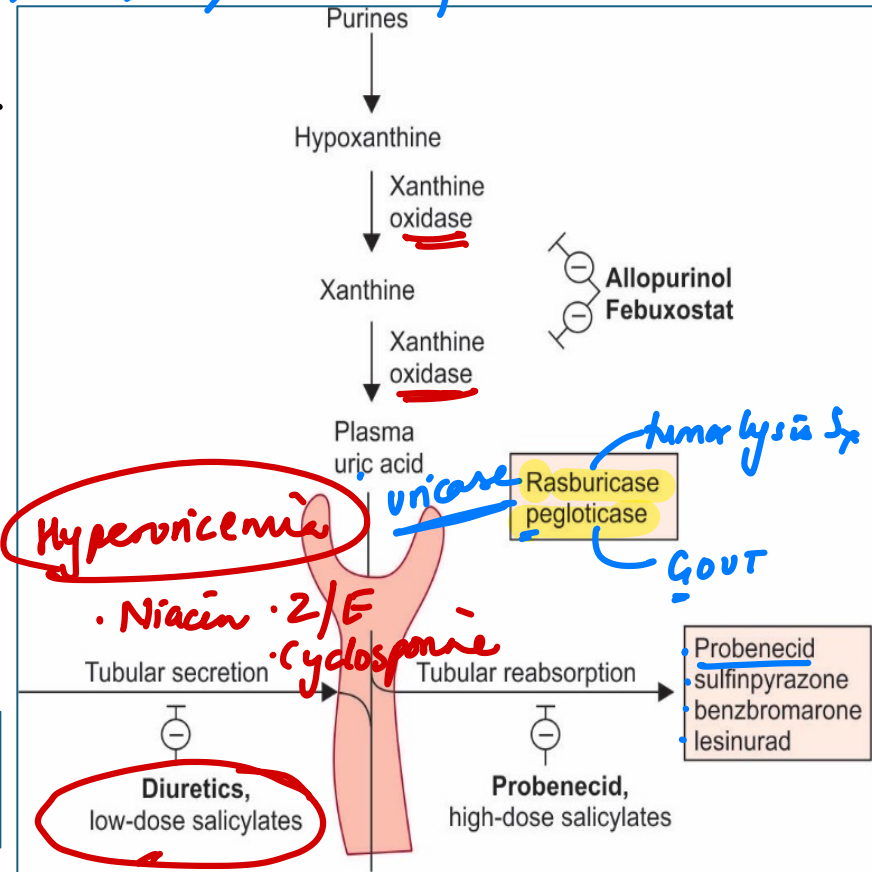
pol: Uric acid
u: negatively birefringent
 needle shaped

CPPD
 +ve birefr
 Rhomboid

GOUT MC: 1st MTP
 Rat bite erosions
 Mantel & sign

Pseudogout
CPPD
 mc knee

Hook shaped metacarpals:
Hemochromatosis



Hyperuricemia

- Diuretics
- Low dose aspirin
- Niacin
- $\textcircled{Z} / \underline{\underline{E}}$
- Cyclosporine

Chronic Arthritis

Inflammatory

Pain, swelling, improves with activity, Morning stiffness > 1hr

Rheumatoid arthritis

- HLA DR4
- MC- MCP
- Sparing- DIP, 1st CMC
- Symmetrical joint space reduction
- Osteopenia, Erosions
- Sea-gull sign: Erosive OA



- DIP, PIP
- Pencil-in-cup, telescoping of digits, Arthritis mutilans
- Skin and Nail

SLE = Jaccoud

- No erosions
- Deformity +



Non-inflammatory

Pain after weight bearing, improve with rest

Osteoarthritis

- Hip, Knee, DIP, PIP, 1st CMC
- MCP sparing
- Asymmetric joint space reduction
- Osteophytes, Subchondral sclerosis, cyst



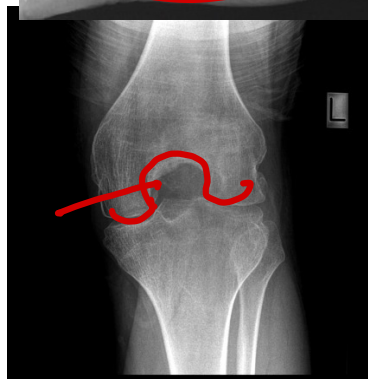
Neuropathic jt = CHARLOT'S

- 5Ds-Disorganisation, Density, Destruction, Debris, Distension
- Diabetes/ neuropathy
- Looks bad, doesn't feel bad



Hemophilia

- Child
- Knee-Squared patella, Intercondylar notch widening
- Pseudotumor

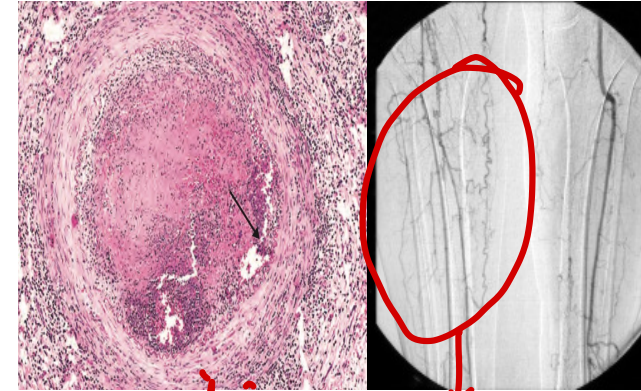
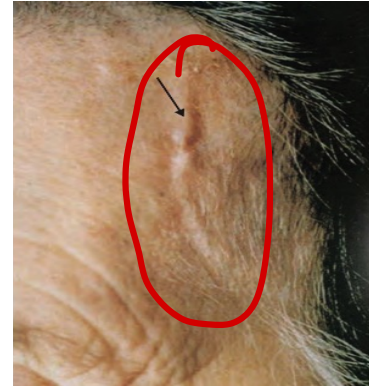


Vasculitis

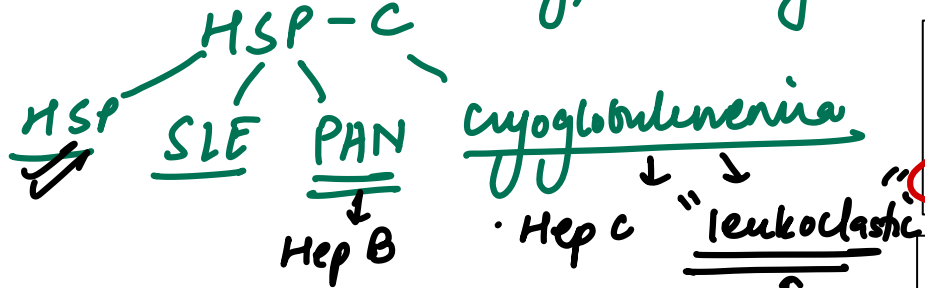
CHAPEL-HILL

Granulomas

- GCA
- Takayasu
- WG = GPA
- Churg Strauss = EGPA



IMMUNE COMPLEX = type III hypers



>50yr, Jaw claudication, **PMR**

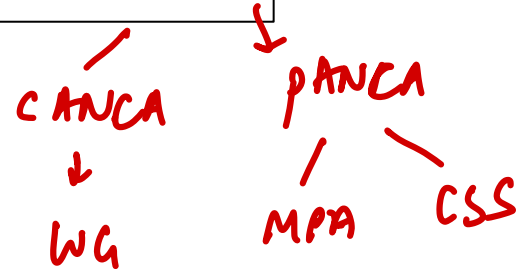
<40yr, BP difference between UL, RAS/MI

MC artery: **STA**
Next step: **steroids**
Gold standard: **Bopsy: >1cm**

MC artery: **LT SCA**

Neutrophilic abscesses
fibrin/radial A
<35yr, smoker, severe limb pain: **A/V/N**

ANCA

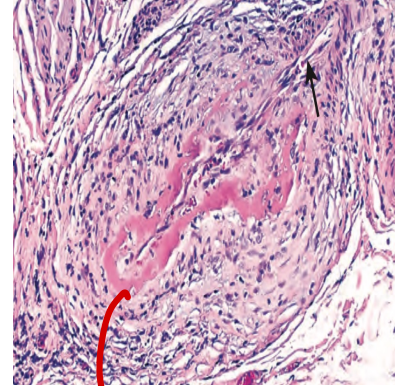


∴ skip
Giant cell / prevent vision loss
temporal arteritis

Takayasu = Aortic arch Sx

Corkscrew collaterals
BUERGER'S = TAO

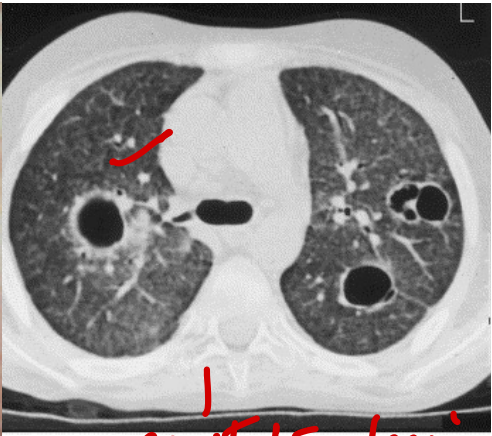
Berger's - IgA nephritis



fibrinoid necrosis



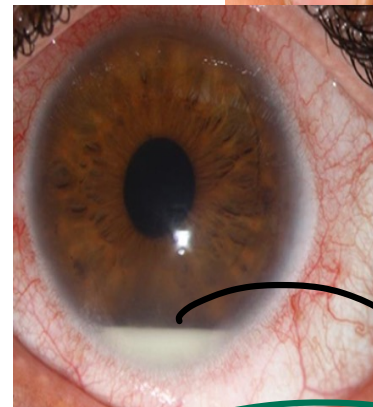
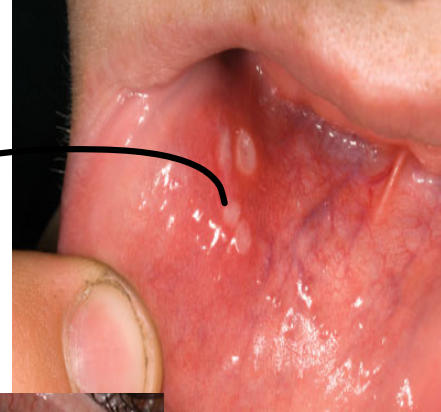
saddle nose



VRT + cavitary lesions



ORAL / genital ulcers



Palmery test (+)

uveitis

Hep B+ 30%
Hypertension

Hematuria-Crescentic
RPGN

WG = GPA

pauci-immune

HSP

Not involved:
PA NO
Glomeruli NO

Triad:
VAT / LRT / C
Granulomas = (X)

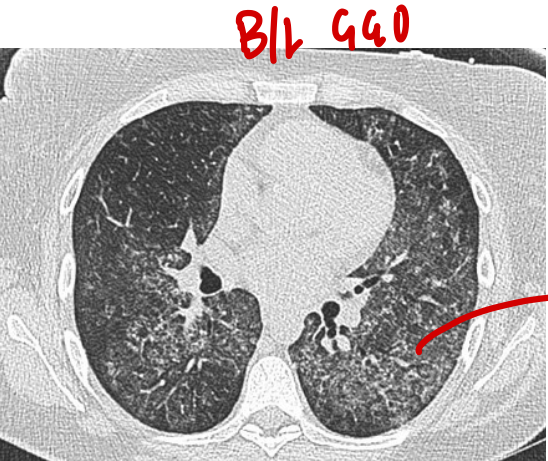
Description
Purpura or petechiae with lower limb predominance

1. Diffuse abdominal pain with acute onset. Submucosal hematoma
2. Histopathology showing leukocytoclastic vasculitis or proliferative glomerulonephritis with predominant immunoglobulin A deposits.
3. Arthritis or arthralgia of acute onset.
4. Renal involvement in the form of proteinuria or haematuria.

intessu

Behcet's

HLA: B51
Ab: Anti-d enolase
Biopsy: neutrophilic vasculitis



B/L 940

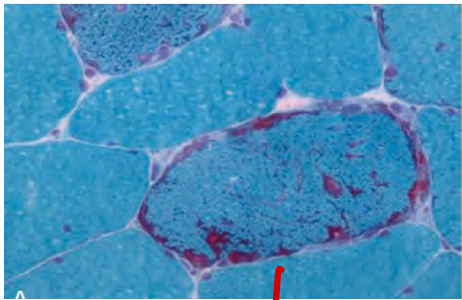
Hemoptysis + Hematuria

pulm alveolar hge
MPA
Goodpasture
WG

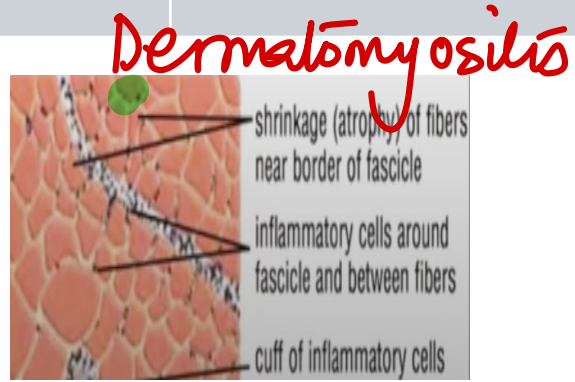
DOC: steroids

Approach to myopathies

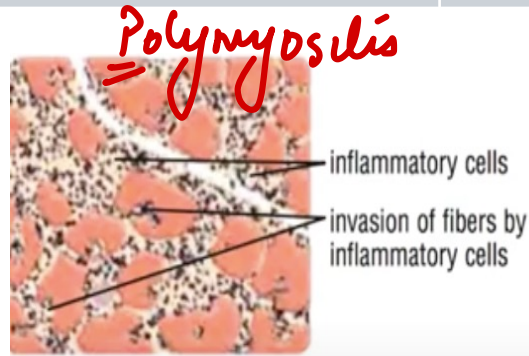
	Polymyalgia Rheumatica	Inflammatory	Steroid	Statin	Hypothyroid
ESR	Raised	Raised	N	N	N
CK	N	Raised	N	Raised	Raised
	Shoulder, hip pain Morning Stiffness >50yrs GGO	• Proximal pain and weakness	• Proximal weakness • No pain	• Pain • No weakness	• Proximal Pain and weakness • Delayed reflexes



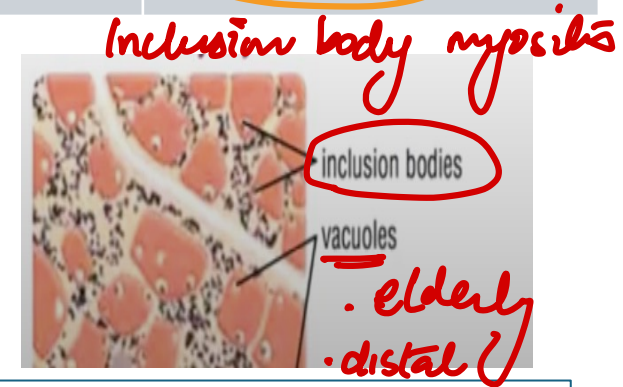
• Red ragged fibres
• mitochondrial accum
MERRF



Perimysial, perivascular Inflammation (CD4) + Perifascicular atrophy

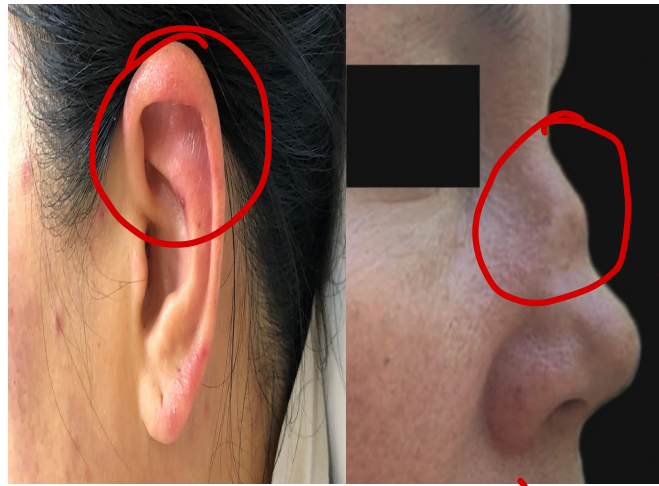
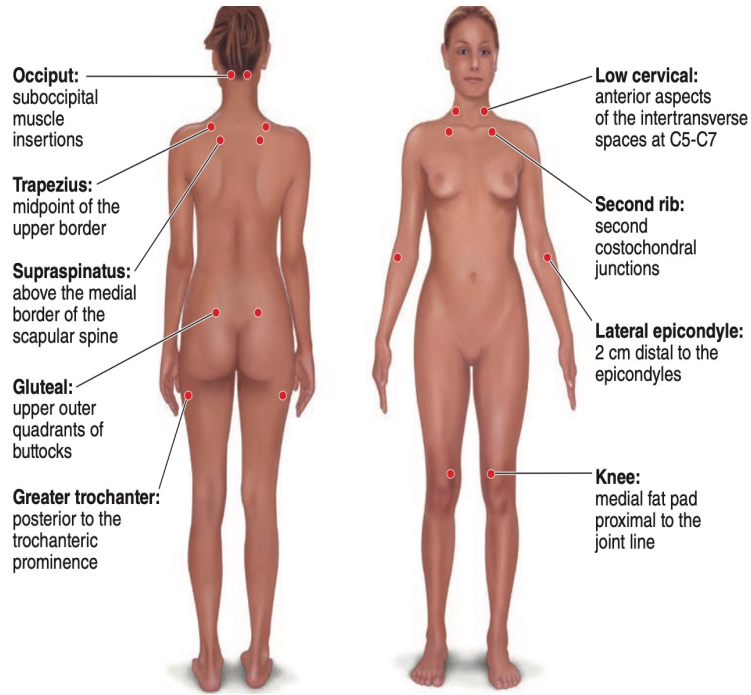


Endomysial, Perivascular inflammation (CD8)



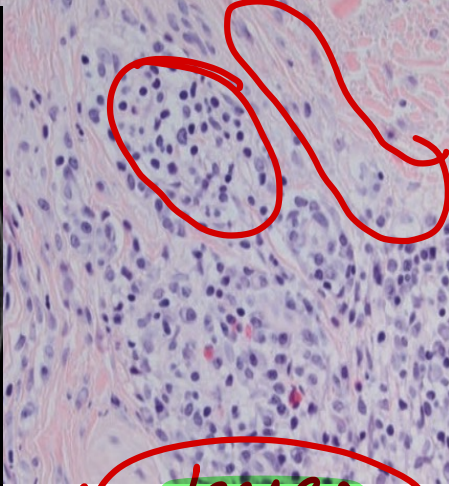
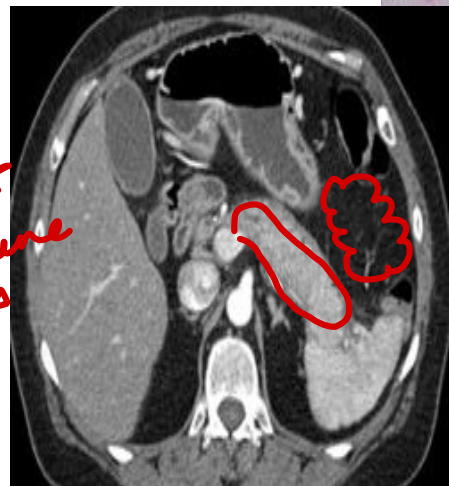
Endomysial, perivascular inflammation (CD8) + Rimmed vacuoles

Miscellaneous



hyaline elastic
Relapsing polychondritis

AIP
autoimmune
panc'itis



sausage shaped IgG4R

Fatigue *Fibromyalgia*
Sleep dysfunction
Tension headache
ESR: (N)
>11/18 tender points
TOC: *Behavioural therapy*
SSRI

Criterion <u>H</u> -Histology (at least one of the following)	<ol style="list-style-type: none"> 1. <u>Periductal lymphoplasmacytic infiltrate</u>, <u>obliterative phlebitis</u>, <u>storiform fibrosis</u> 2. Lymphoplasmacytic infiltrate, storiform fibrosis, abundant IgG4+ cells (≥ 10 HPF)
Criterion <u>I</u> -Imaging of pancreas	<ol style="list-style-type: none"> 1. Typical-diffusely enlarged gland with delayed (rim) enhancement; diffusely irregular, attenuated main pancreatic duct 2. Others-Focal pancreatic mass/enlargement; focal pancreatic duct stricture; pancreatic atrophy; pancreatic calcification; pancreatitis
Criterion <u>S</u> -Serology	<u>Elevated serum IgG4</u> (normal: 8-140 mg/dL)
Criterion <u>O</u> -Other organ involvement (can be confirmed by biopsy or resolution/improvement with steroid therapy)	<u>Hilar/intrahepatic biliary strictures</u> ; persistent distal biliary stricture; parotid/lacrimal gland involvement; mediastinal lymphadenopathy; retroperitoneal fibrosis
Criterion <u>R</u> -Response to steroid therapy	Resolution or marked improvement of pancreatic/extrapancreatic manifestation with steroid therapy

Mayo Clinic