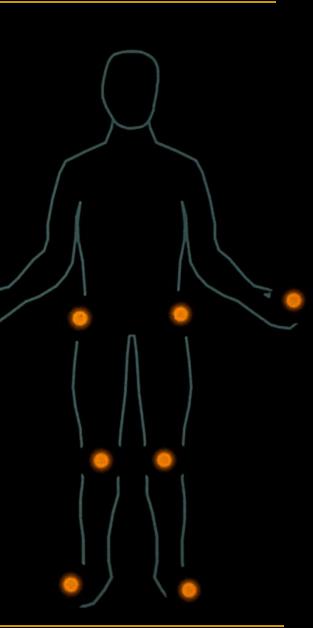
APPROACH TO THE RHEUMATOLOGICAL PATIENT

Catalina Orozco, MD Rheumatology Associates September 23rd 2013

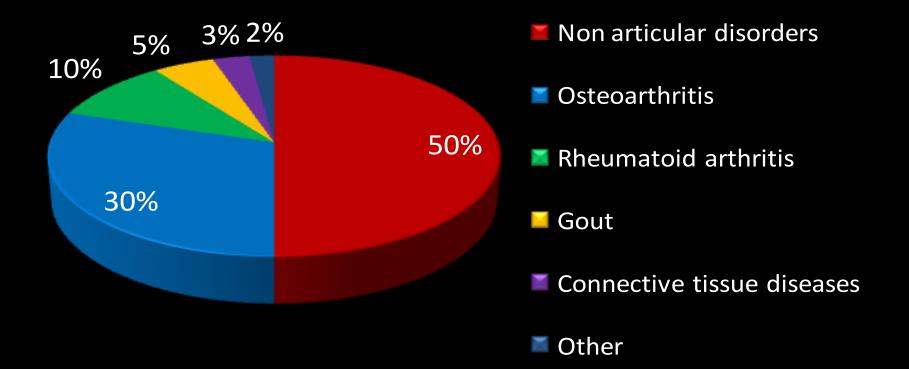
OUTLINE

- Introduction
- Approach to the patient with a musculoskeletal complaint
- Approach to the patient with a connective tissue disease
- Approach to the patient with vasculitis
- Take home messages

- More than 100 rheumatic conditions
- Overlap in clinical presentations
- Initial presentation may not lead to a precise diagnosis in up to 50% of cases
- Over time most patients will have characteristic features of a disease



Relative Prevalence of Rheumatic Diseases



COMPLETE HISTORY

COMPLETE PHYSICAL EXAM

LABORATORIES

IMAGING AND OTHER STUDIES

A DIAGNOSIS CAN BE MADE BASED ON HISTORY AND PHYSICAL EXAM FINDINGS 80-90% OF THE TIME

DIAGNOSTIC APPROACH

<u>History:</u>

- Establish the demographics of the patient: Age Gender Ethnicity Family History
- Characterize joint pain and ask about associated features
- Characterize back pain: Inflammatory vs. Non inflammatory
- Constitutional symptoms

DIAGNOSTIC APPROACH

<u>History:</u>

Organ specific symptoms

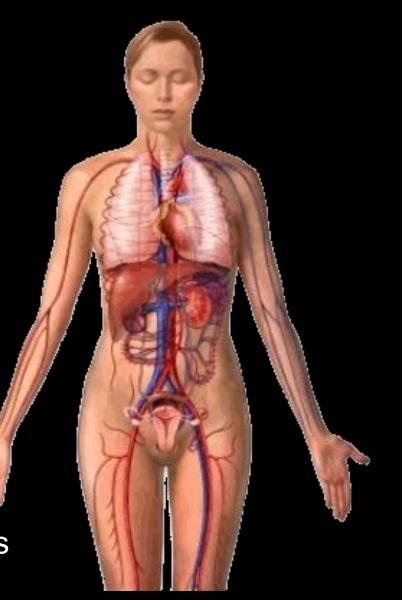
 Eye: Pain, redness, dryness, vision changes
 Heart: Chest pain, palpitations, orthopnea, PND
 Lungs: Dyspnea, cough
 Kidneys: Hematuria, edema
 GI: GERD, dysphagia, GIB, bowel habit changes
 Skin: Ulcer, photosensitivity, rashes, alopecia, nail abnlity
 Neuro: CNS changes, neuropathy, CN abnormalities
 ID: recent infections

DIAGNOSTIC APPROACH

Physical Exam:

Head to toe evaluation

Eye redness Mouth dryness/ulcers Rales, effusions, murmurs, rubs Hepatosplenomegaly LAD Motor strength, sensation, DTR's

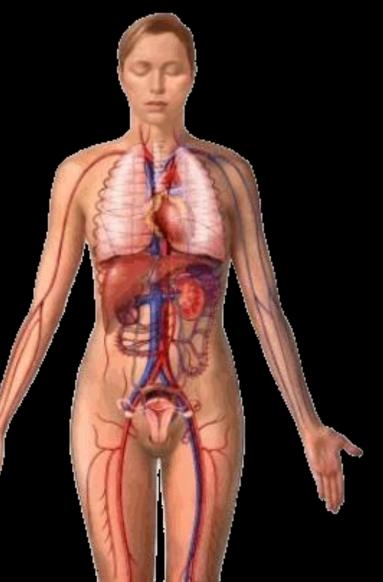


DIAGNOSTIC APPROACH

Physical Exam:

Head to toe evaluation

Rashes, telangiectasis, nail changes, pigmentation changes Peripheral pulses, bruits Back exam Joint exam



DIAGNOSTIC APPROACH

Laboratories:

- Results must be interpreted in light of the clinical findings
- Three areas of interest: Blood, Urine, Synovial fluid

DIAGNOSTIC APPROACH

<u>Blood:</u>

- CBC: Anemia, leukopenia, thromobocytopenia
- Chemistries: renal insufficiency, elevated LFT's, uric acid
- ESR/CRP: non specific
- Autoantibodies: RF, ANA, ENA, dsDNA, ANCA
- HLA B-27, HLA B-51
- ASO
- Ferritin
- Lyme titer

DIAGNOSTIC APPROACH

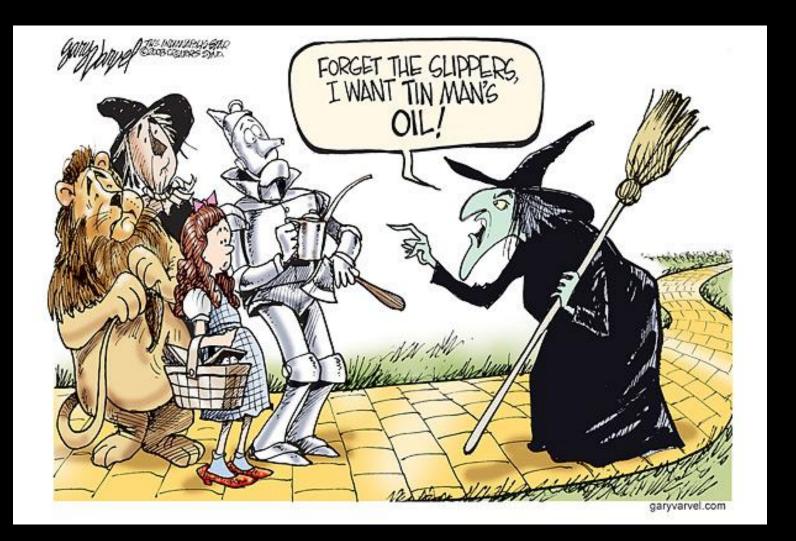
<u>Urine:</u>

- Proteinuria
- Hematuria
- Active sediment

DIAGNOSTIC APPROACH

Synovial fluid:

- Cell count
- Gram stain and culture
- Crystal analysis





CC: Arthralgia in hands and feet

HPI: 50 yo WW presents with 6 weeks of arthralgia in hands and feet. Swelling over MCP's, PIP's. Has tried naproxen with partial relief

ROS, PMH, PSH, Social HX: Negative

FH: mother with rheumatoid arthritis

PE: Tenderness & synovitis of wrists, R MCP 2-4 and B PIP 3

Labs: CRP: 1.3 mg/dl, RF -, CCP 25 (nl < 20 u/ml), normal Xray

EVALUATION OF PATIENTS WITH MUSCULOSKELETAL COMPLAINTS

Goals

Accurate diagnosis Timely provision of therapy Avoidance of unnecessary diagnostic testing

Harrison's Principles of Internal Medicine - 17th Ed. (2008)

DIAGNOSTIC APPROACH

Articular Vs. Non Articular:

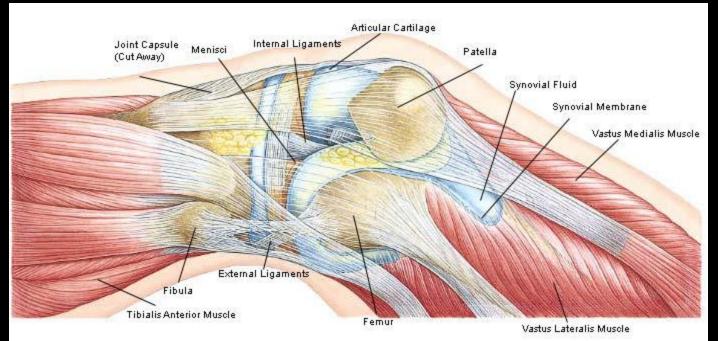
Articular structures



DIAGNOSTIC APPROACH

Articular Vs. Non articular:

Non articular structures



DIAGNOSTIC APPROACH

History and physical exam pearls:

 Articular: Pain is diffuse and deep Pain with active and passive range of motion Swelling Crepitation, locking, instability or deformity

DIAGNOSTIC APPROACH

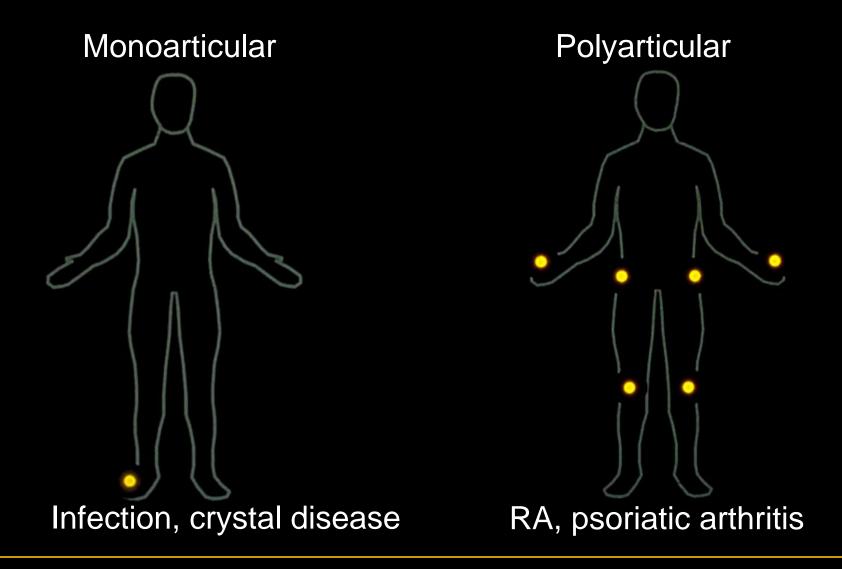
History and physical exam pearls:

• Non articular: Pain with active but not passive ROM Tenderness in adjacent structures Other physical findings in remote areas

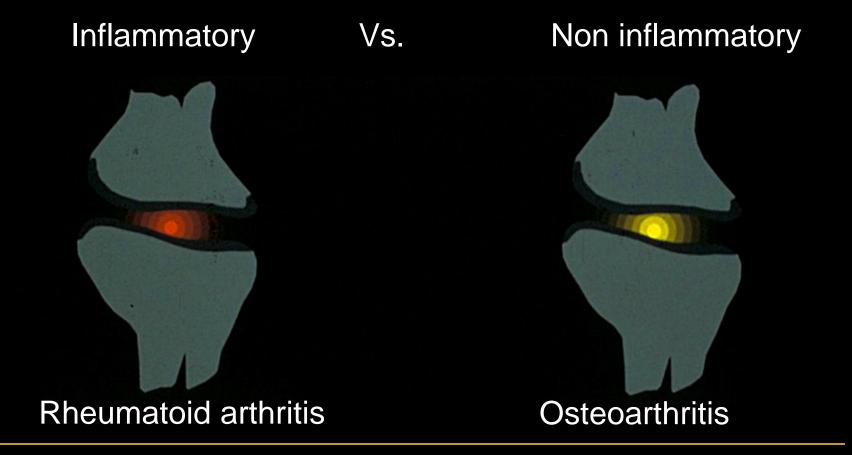
DIAGNOSTIC APPROACH

Monoarticular Vs. Polyarticular

- Monoarticular: 1 joint involved
- Oligoarticular: 2-4 joints involved
- Polyarticular: >4 joints involved



DIAGNOSTIC APPROACH



DIAGNOSTIC APPROACH

History and Physical exam Pearls:

• Inflammatory:

Erythema Warmth Pain Swelling Tenosynovitis Stiffness after prolonged rest Fatigue

DIAGNOSTIC APPROACH

History and Physical exam Pearls:

 Non Inflammatory: Pain without swelling Gel phenomenon Pain is aggravated with activity

Inflammatory arthritis



Inflammatory arthritis

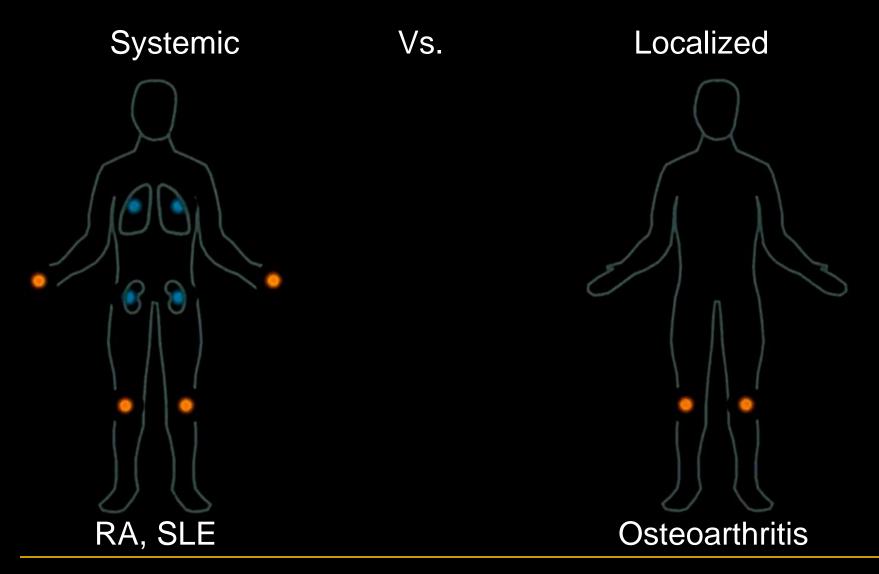


Inflammatory arthritis

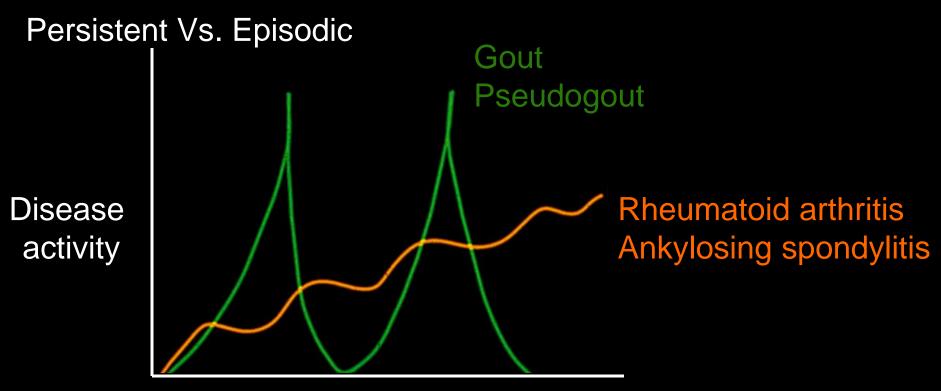


Non Inflammatory arthritis



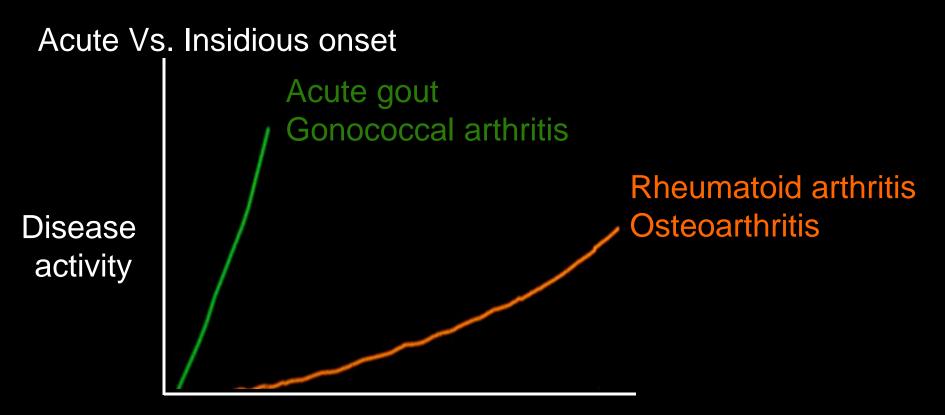


DIAGNOSTIC APPROACH



Time (weeks)

DIAGNOSTIC APPROACH



Time (weeks)

DIAGNOSTIC APPROACH

Physical exam:

Look for clues in other organs that may lead to a diagnosis



Rheumatoid arthritis



Psoriasis



Gonoccocal arthritis



Ankylosing Spondylitis



Gout





Sarcoidosis

IBD related arthritis

DIAGNOSTIC APPROACH

Laboratories:

 CBC: Anemia, thrombocytosis
 Neutropenia, thrombocytopenia
 Felty's syndrome Hemolysis, thrombocytopenia
 SLE

Elevated ESR, CRP inflammation

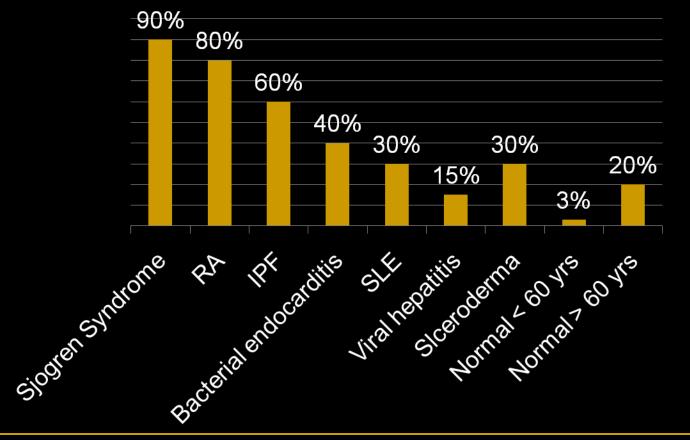
• CCP antibodies: Positive in 40% of RF negative RA, CCP is far more specific for RA than RF is, but a negative result does not exclude the diagnosis of RA

Van Venrooij WJ, Autoimmun Rev 2006; 6(1):37-41

DIAGNOSTIC APPROACH

Laboratories:

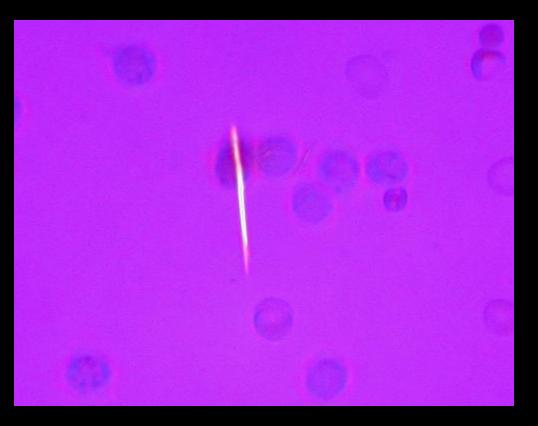
Ocurrence of positive RF



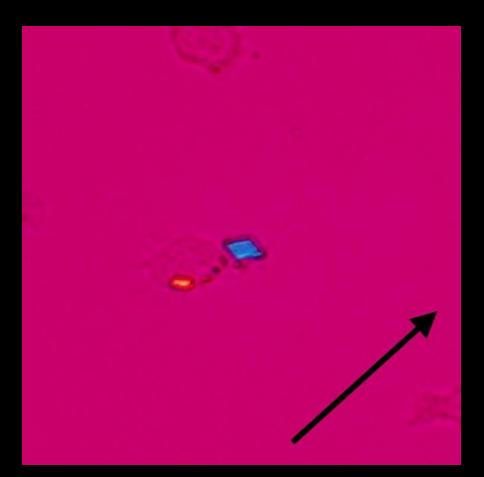
DIAGNOSTIC APPROACH

Synovial fluid analysis

Condition	Appearance	WBC count	PMN%	Other
Normal	Clear/straw	< 200	< 25%	
Trauma	Clear/bloody	< 2000	< 25%	
OA	Clear/slightly cloudy	< 2000	< 25%	Occasional cartilage fragments
Infection	Cloudy/purulent	> 50000	> 80%	Organisms on gram stain
RA	Cloudy/light yellow	< 10000	75%	Occasional cholesterol crystals
Gout	Cloudy/white or yellow	> 10000	75%	MSU crystals
Pseudogout	Cloudy/white or yellow	> 10000	75%	CPPD crystals



Monosodium urate crystal



Calcium pyrophosphate crystal



CASE 2

- CC: Arthralgia and muscle weakness
- HPI: 46 yo AAW who presents with 6 months of arthralgia in shoulders, hands, knees and feet. Swelling wrists, and PIP's. She also has difficulty getting up from a chair, and combing her hair.
- ROS: Easy bruisability, Raynaud's phenomenon, hair loss PMH, PSH, social history and family history: NC PE: Diffuse hair loss, TJC 26, SJC 9. Weakness in proximal arms, legs and neck flexors
- Labs: Leukopenia, elevated AST, CPK 1000, aldolase 12, UA: nl, ANA 1:640, RNP pos, C3 120, C4 8

Diagnostic approach:

History pearls:

• Multisystem organ involvement is common

Constitutional: Fatigue, fever, weight loss Eyes: Sicca, redness, pain ENT: Sicca, oral/nasal ulcers, dysphagia, sore throat Heart: Chest pain, orthopnea, PND Lungs: Pleuritic pain, dyspnea, cough Abdomen: GERD, bowel changes, GIB Joints: Pain, swelling, stiffness

Diagnostic approach:

History pearls:

Skin: Photosensitivity, hair loss, rashes, Raynaud's, nail changes Neurological: Neuropathy, muscle weakness, mental status changes, headache, seizures Psychiatric: Depression, psychosis GU: genital ulcers

Diagnostic approach:

Physical exam pearls:

Head to toe examination



Relapsing polychondritis, granulomatosis with polyangiitis



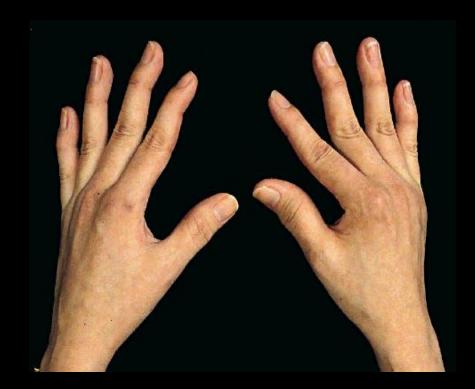
SLE



SLE, Behcet's disease



Systemic sclerosis



SLE



Systemic sclerosis



Systemic sclerosis, dermatomyositis





Systemic sclerosis, SLE



Dermatomyositis





Dermatomyositis



SLE



Chronic cutaneous lupus



SLE, antiphospholipid syndrome, Sjogren syndrome

	Systemic lupus erythematosus	Sjögren's syndrome	Systemic sclerosis	Idiopathic inflammatory myositis
Constitutional signs and symptoms (eg, fever, fatigue, and weight loss)	Common	Fatigue (severe)	Less prominent primary symptoms	Common, mainly fatigue, usually without fever
Arthritis or arthralgia	Common, typically nonerosive	Common, nonerosive	Arthralgia common, synovitis rare	Polyarticular, and mild, erosive, and deforming with anti-Jo-1
Muscle symptoms	Myalgia, myositis (1–4%)	Myalgia	Myalgia, disuse atrophy, myositis (rare)	Substantial weakness, occasional pain
Mucocutaneous manifestations	Malar, discoid rashes, photosensitivity, mouth ulcers	Dryness, oral infections, hypergammaglobulinaemia purpura	Skin fibrosis, sclerodactyly, calcinosis, telangiectasia†	Rashes with dermatomyositis (eg, Gottron's papules, heliotropic rash eyes, V signs, and shawl signs)
Raynaud's syndrome	Yes (about 20% of patients)	Yes	Frequent, severe	Yes
Sicca syndrome	10–20% of patients	Prominent mouth and eyes	Occasional	<10% of patients
Cardiovascular disease	Pericarditis, early cardiovascular disease, and Libman-Sacks endocarditis	Uncommon	Right heart failure and secondary pulmonary hypertension	Arrhythmias, valvular heart disease, and ischaemia
Pulmonary symptoms	Serositis, pulmonary embolism, interstitial lung disease, pulmonary hypertension, shrinking lung	Chronic cough and lymphoproliferative disorders	Interstitial lung diseas and pulmonary hypertension (10–50% of patients)	Dry cough, shortness of breath, respiratory muscle weakness, interstitial lung disease, pulmonary hypertension, and bronchiolitis obliterans organising pneumonia
Gastrointestinal symptoms	Mesenteric vasculitis	Dysphagia and primary biliary cirrhosis	Oesophageal dysfunction, gastro- oesophageal reflux disease, diarrhoea, and faecal incontinence	Dysphagia most common, gastro- oesophageal reflux disease
Renal symptoms	Yes (30–50% of patients), glomerulonephritis	Interstitial lymphocytic nephritis, distal tubular acidosis, and interstitial cystitis	Renal crisis (diffuse systemic sclerosis), more common mild dysfunction	Very rare
Neurological symptoms	Headache, mood, cognitive disorders (20–30% of patients), other more severe and rare	CNS, peripheral nervous system, autonomic nervous system; Adie's pupil, orthostatic intolerance	Very infrequent, includes cranial neuropathies	CNS very uncommon
Haematological symptoms	Common, decreased white cell count, platelets, and haemoglobin	Decreased white cell count, anaemia (<10% of patients)	Anaemia secondary to gastrointestinal blood loss	Very rare

*Clinical features of systemic vasculitides in table 2. †Dependent on whether disease subtype is limited or diffuse.

Diagnostic approach:

Laboratories:

- CBC: Hemolysis, thrombocytopenia, leukopenia, lymphopenia
- CMP: Renal insufficiency, elevated LFT's, hypokalemia
- Urinalysis: Hematuria, proteinuria, active sediment, inability to acidify the urine

Diagnostic approach:

Laboratories:

• ANA: Antibodies against nuclear specificities (e.g. DNA, snRNP)

• ANA can be seen in the normal population:

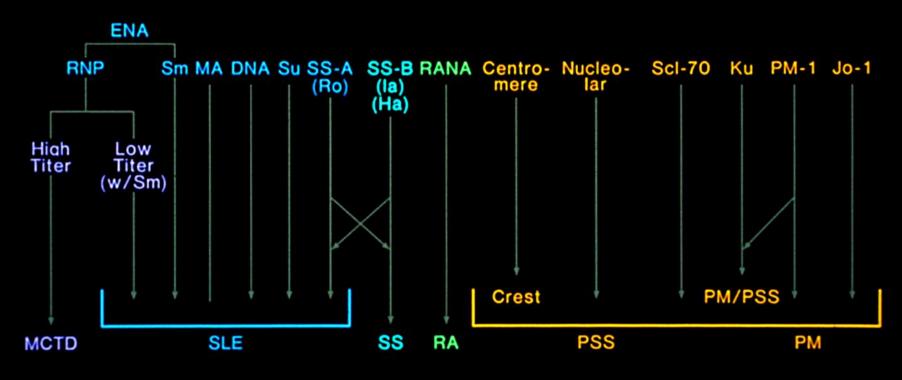
≥ 1:40: 20-30%
≥ 1:80: 10-12%
≥ 1:160: 5%

≥ 1:320: 3%

Causes of Positive ANA	%
SLE	~99
Systemic sclerosis	97
PM/DM	40-80
Sjogren syndrome	48-96
Drug induced lupus	100
MCTD	100
Autoimmune hepatitis	100
Juvenile arthritis	20-50
APS	40-50
Raynaud's phenomenon	20-60

Diagnostic approach:

Laboratories:



Diagnostic approach:

Laboratories:

In clinical use, ANA is insufficient to establish or refute a diagnoses

• ANA results add weight to diagnoses that throughout the evaluation should rely heavily on other clinical information

Diagnostic approach:

Imaging studies:

Systemic sclerosis



Diagnostic approach:

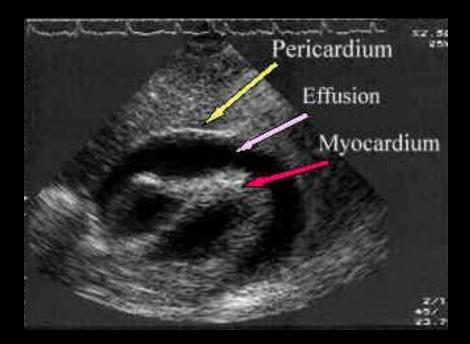
Imaging studies:



Systemic sclerosis

Diagnostic approach:

Imaging studies:



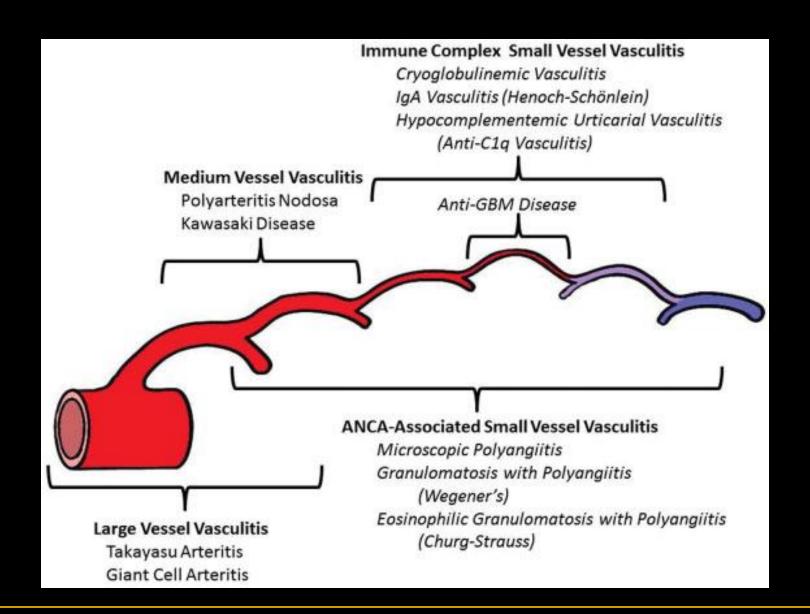
SLE



He's got a wierd disorder called acronymophobia.Cracked up on the BAFF batch.

CASE 3

- CC: Cough and dyspnea
- HPI: 50 yo WM who presents with 2 months of worsening
- cough, productive of yellow sputum, intermittent fever, DOE at 50 ft, sinus pressure and arthralgia.
- ROS: Loss of hearing
- PMH: Scleritis 1990, pneumonia 2012
- PSH, social history and family history: NC
- PE: Tenderness of paranasal sinuses, rhonchi in B lung fields Labs: Normocytic anemia, creatinine 1.4 mg/dl, CRP 3 mg/dl, ESR 50 mm/hr, ANA -, ANCA +, PR 3 +, UA: blood +, 20 RBC CXR: bilateral alveolar infiltrates and cavitary lesion on the left lower lobe



Jennette, JC. Arthritis & Rheumatism 2013 65;1: 1-11

Diagnostic approach:

History pearls:

- Constitutional symptoms are common
- Hearing loss, eye pain, redness
- Upper respiratory symptoms: sinusitis, purulent drainage, epistaxis, nasal ulceration, stridor
- Lower respiratory symptoms: dyspnea, cough, hemoptysis, asthma

Diagnostic approach:

History and Physical exam pearls:

- Neuropathic symptoms, muscle weakness
- History of hepatitis B or C, HIV, other viruses
- Skin rashes, ulcers, nodules or vesicles
- Abdominal pain, GIB

Diagnostic approach:

Physical exam pearls:

GPA Relapsing polychondritis



Diagnostic approach:

Physical exam pearls:





Diagnostic approach:

Physical exam pearls:

Eosinophilic granulomatosis with polyangiitis



Diagnostic approach:

Physical exam pearls:



Polyarteritis nodosa

Diagnostic approach:

Physical exam pearls:



Henoch-Schonlein purpura

	Giant cell arteritis	Polyarteritis nodosa	Granulomatosis with polyangiitis	Eosinophilic granulomatosis with polyangiitis	Microscopic polyangiitis
Predominant blood vessel type	Large	Muscular arteries (medium or small arteries)	Arterioles-venules, sometimes arteries, veins	Arterioles-venules, sometimes arteries, veins	Arterioles-venules, sometimes arteries, veins
Granulomatous inflammation	Yes	No	Yes	Yes	No
ANCA positivity	No	No	80–90% (cytoplasmic ANCA/ antiproteinase 3)	50% (perinuclear ANCA/ myeloperoxidase)	75% (perinuclear ANCA/ myeloperoxidase)
Upper respiratory tract symptoms	No	No	Sinusitis, epistaxis, nasal septal perforation, saddle nose deformity, nasal crusting, and subglottic stenosis	Nasal polyps and allergic rhinitis	Usually absent or mild
Lung	No	Asthma	Nodules, infiltrates or cavitary lesions, and haemoptysis	Asthma, infiltrates, and haemoptysis	Alveolar haemorrhage
Glomerulonephritis	No	No	Yes	Yes	Yes
Renal hypertension	No	Yes	No	No	No
Gastrointestinal	No	Yes	No	Yes (eosinophilic gastroenteritis); pain, diarrhoea, and bleeding	Yes
Cardiac	Uncommon, coronary artery involvement, ischaemic heart disease, congestive cardiac failure	Yes (myocarditis)	Yes (valvular infarction)	Yes (eosinophilic myocardial infiltration, cardiomyopathy in 50% of patients)	Uncommon (congestive cardiac failure)
Skin	No	Nodules, livido reticularis painful serpiginous rash	Yes	Yes (tender nodules)	Yes
Mononeuritis multiplex	Yes	Common	Occasional	Common	Common
Ocular	Yes (vasculitic optic neuritis)	Yes (scleritis)	Yes (including retro-orbital granulomata)	Uncommon (scleritis and uveitis)	Yes (typically mild)
Distinguishing features	Blindness, headaches, scalp tenderness, jaw claudication Often associated with polymyalgia rheumatica	ANCA-negative, testicular involvement, painful rash	Destructive upper airways disease, granulomatous inflammation	Asthma, allergy, eosinophilia, granulomatous infiltrates with abundant eosinophils	No granulomatous inflammation

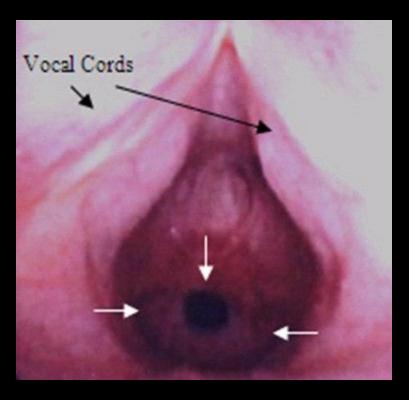
Diagnostic approach:

Labs:

- CBC: Anemia
- Elevated ESR/CRP
- CMP: Renal insufficiency, hypoalbuminemia
- Immunological: ANA, ANCA, cryoglobulins, RF, complements
- Other: Hepatitis serologies
- Urinalysis: Hematuria, proteinuria, cellular casts

Diagnostic approach:

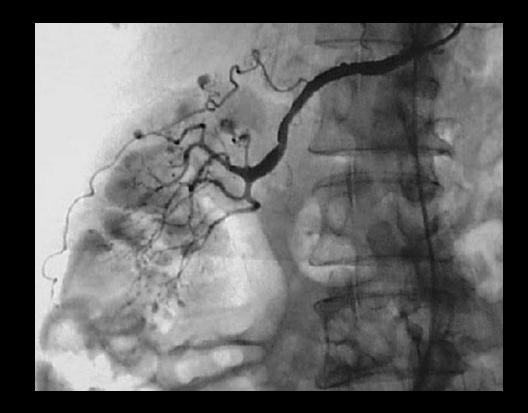
Imaging:





Diagnostic approach:

Imaging:

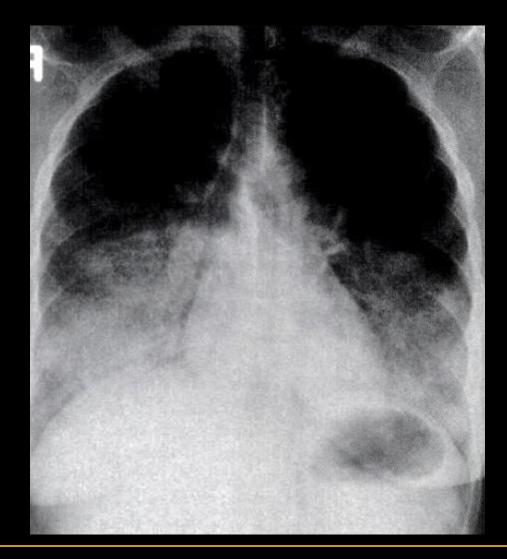


Polyarteritis nodosa

Diagnostic approach:

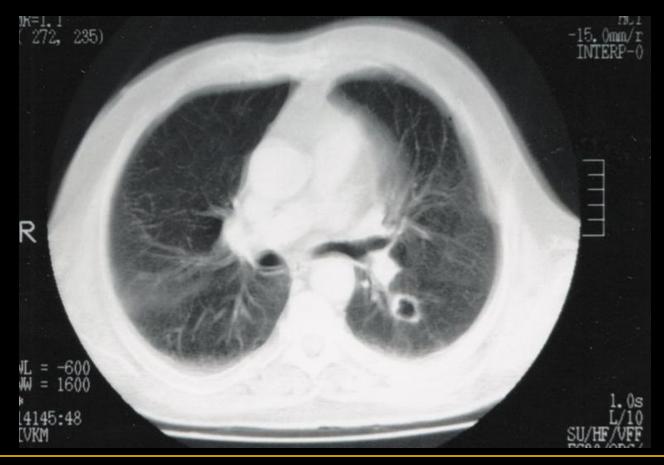
Imaging:

Goodpasture's syndrome GPA Microscopic polyangiitis



Diagnostic approach:

Imaging:



GPA

TAKE HOME MESSAGES

• A full history and physical exam can lead to a diagnosis 80-90% of the time

• In patients with a musculoskeletal symptom characterize the pain: Is it articular?, how many joints are involved?, is it inflammatory?, is it a systemic process?, What is the duration of symptoms?

• ANAs can be positive in multiple conditions and even in the normal population. Think of the pre-test probability for a connective tissue disease before ordering it



