#### Spondyloarthropathies

Inflammatory Arthritis







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### Disclosures

NONE

# What is meant by spondyloarthropathy?

 Literally translated to "inflamed spine growing together"

 The first documented ankylosing spondylitis case was reported in 1691, although it may have been present in ancient Egyptians

### Seronegative SpA

- Inflammatory axial spine involvement
  - Spondylitis, sacroilitis
- Asymmetric peripheral arthritis
  - DIP involvement, dactylitis ("sausage digits")
- Inflammatory eye disease
- Mucocutaneous features
- Enthesopathies:
- Achilles tendinitis
- Plantar fasciitis
- Tenosynovitis
- Familial aggregation
- HLA-B27 association

### SpA's include

- Ankylosing spondylitis
- Reactive arthritis
- Psoriatic arthritis/spondylitis
- Arthritis/spondylitis of inflammatory bowel dz
- Juvenile spondyloarthropathy
- Undifferentiaed spondyloarthropathies
- ?Whipple dz, Bechet's dz, Celiac Dz

#### Prevalence

- 2.7 million adults in the United States have seronegative SpA
  - Rheumatoid arthritis only affects 1.3 million adults, down from the previous estimate of 2.1 million
  - Ankylosing spondylitis is the most prevalent of the classic spondyloarthropathies.
- Prevalence AS is 0.1-1% overall, but is higher in certain Native American populations and lower in African Americans
  - highest prevalence in northern European countries and the lowest in sub-Saharan Africa

#### Who's affected?

- Male-to-female ratio of AS is 3:1
- Male-to-female ratio of PsA is 1:1
- Age of onset for AS is from late teens to 40 yo.
  - Approximately 10%-20% of all pts have onset of symptoms before 16 yo

## Relationship with HLA-B27

Strong association with HLA-B27

	Population or Disease Entity	HLA-B <sub>2</sub> 7 +
	Healthy whites	8%
	Healthy African Americans	4%
	Ankylosing spondylitis (whites)	92%
	Ankylosing spondylitis (African Americans	5)50%
	Reactive arthritis	60-80%
	Psoriasis associated with spondylitis	60%
٠	IBD associated with spondylitis	60%
٠	Isolated acute anterior uveitis	50%
٠	Undifferentiated spondyloarthropathy	20-25%

## Spondyloarthropathies (SpA)

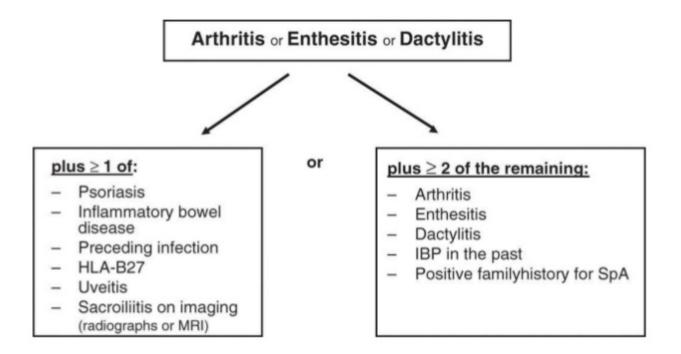
- Newer diagnostic criteria (2.7 million patients)
- SIJ MRI (more sensitive for sacroiliitis than plain radiographs)
- Ustekinumab (Stelara) IL12/IL23 targets; approved for PsA 9/2013, AS '16
- Apremilast (Otezla) phosphodiesterase-4 (PDE4) inhibitor. Oral Rx. No routine blood work, nausea, weight loss (5-10#), depression
- TNF alpha inhibitors (patients feel great on these)
- Secukinumab (IL-17 inhibitor) Cosentyx now available for AS 1/2016

#### CASPAR for PsA

- 3 points or greater
- Current Psoriasis (2)
- H/o Psoriasis (1)
- Family history (1)
- Dactilytis (1)
- Juxta-articular bone formation (1)
- RF neg (1)
- Nail dystrophy (1)

Sacroiliitis on imaging <sup>A</sup>	OR	HLA-B27	
Plus ≥I SpA feature <sup>B</sup>		Plus ≥2 other SpA features <sup>B</sup>	
Active (acute) inflammation		<ul> <li>Inflammatory back pain</li> </ul>	
on MRI highly suggestive of		Arthritis	
sacroiliitis associated with SpA		• Enthesitis (heel)	
<ul> <li>Definite radiographic sacroiliitis according to modified New York</li> </ul>		• Uveitis	
		Dactylitis	
criteria		Psoriasis	
		<ul> <li>Crohn's/ulcerative colitis</li> </ul>	
		<ul> <li>Good response to NSAIDS</li> </ul>	
		<ul> <li>Family history of SpA</li> </ul>	
		<ul> <li>HLA-B27 positivity</li> </ul>	
		Elevated CRP	

Final set of classification criteria for peripheral spondyloarthritis (SpA) (set 2D) selected by Assessment of SpondyloArthritis international Society (ASAS).



Rudwaleit M et al. Ann Rheum Dis 2011;70:25-31



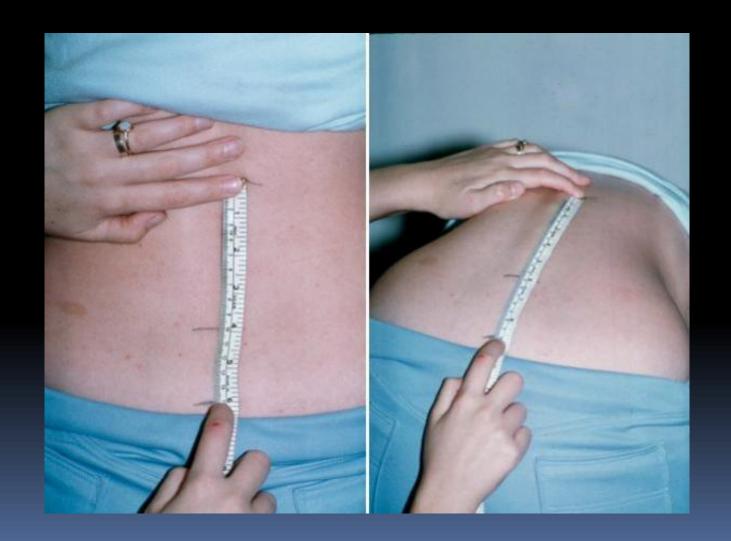
#### Axial Spine

- Schober Test: expect greater than 5 cm change when pt bends forward, if <5 cm change = reduced lumbar mobility
- Spinal fusion results in irreversible impairments, but reductions in mobility also can be induced by pain or muscle spasm so may vary between visits
- With advanced disease, lead to loss of lumbar lordosis, exaggeration of thoracic kyphosis, an inability to extend the neck, and compensatory hip flexion deformities
- Costovertebral and costochondral joints commonly lead to impaired chest expansion; slight 
   ✓ in VC & TLC

## Physical examination tests to assess spinal disease in AS

- Occiput-to-wall-test
- Tragus-to-wall-test
- Chest expansion
- Schober's test
- Sacro-iliac stress test

## Schober Test



### Lab Testing

- Anaemia
- secondary to chronic disease
- blood loss
- -? drugs
- ESR/CRP
- Renal function, urinanalysis,
- Serum IgA elevation
- Negative RF
- HLA B<sub>27</sub>

#### Other manifestations

- Ocular: acute anterior uveitis-unilateral eye pain, photophobia, blurred vision, and increased lacrimation
- Osteoporosis-d/t increased inflammation, syndesmophytes can lead to falsely elevated DEXA score
- GI: Small or large bowel inflammation
- Cardiac: ascending aortitis, AR, conduction abnormalities, and myocardial disease
- Pulmonary: apical fibrobullous disease
- Neuro: spinal fracture, cauda equina syndrome
- Rare: secondary amyloidosis, retroperitoneal fibrosis

## Spondyloarthropathies

Iritis (Uveitis)





#### Likelihood of Iritis

<u>Disease</u>	Percent	
·Ank spondylitis	20-30	
•Reiter's	12-37	
•Psor Spondylitis	7-16	
•IBD	2-9	
•Undiff SA	ND	

## Radiographic Testing

- X-ray of SI joints:
  - □ Iliac erosions (postage stamp serrations) → erosions become more prominent and produce "pseudowidening" of the SI joint → fusion → complete obliteration of the SI joint by bone and fibrous tissue. The pattern of sacroiliac joint involvement is bilaterally symmetric

## AS pelvis X ray

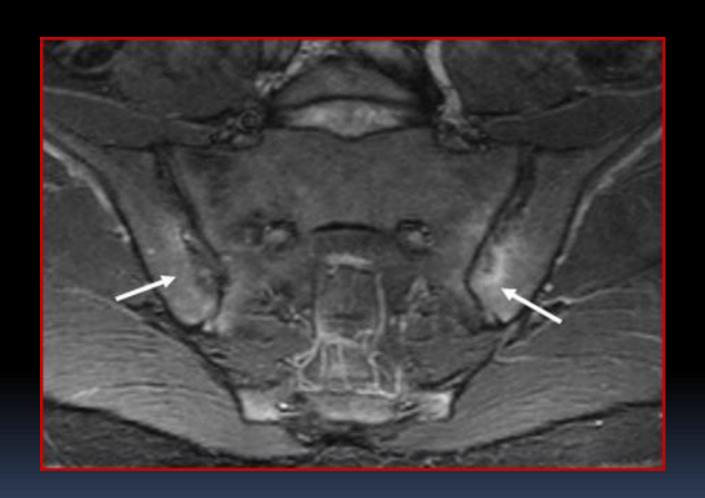






#### MRI

- Acute sacroiliitis, spondylitis, and spondylodiscitis
- Acute inflammation of the entheses, bone, and synovium
- Detect early inflammation and accurately visualize cartilaginous and enthesal lesions
- If X-ray negative and high index of suspicion then f/u with an MRI



#### AS

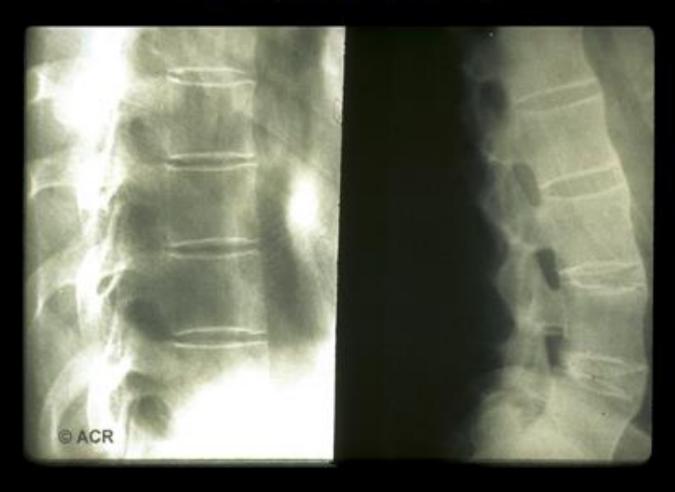
- Most common cause of inflammatory back pain in young adults
- Prevalence: 150 per 100,000
- Age of onset: < 40 years of age (average 26 yrs)</li>
- M:F ratio 3:1
- Back pain (insidious onset)
- Duration longer than 3 months
- Associated with morning stiffness
- Improves with exercises
- Decreased spinal movements
- Enthesopathies

## Ankylosing spondylitis

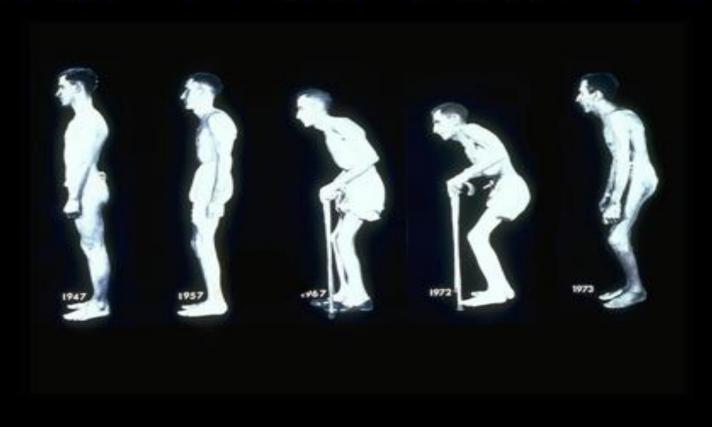
- Sacroiliitis and spondylitis
- Bamboo spine- ossification of the annulus fibrosus
  - Occurs from the lumbosacral region proximally
- Enthesitis
- Peripheral arthropathy

Do not fit into another category

## AS spine X ray



#### Progressive spinal changes in AS



## Ankylosing spondylitis

- 🔼 Aortic insufficiency, ascend. aortitis, conduction defects, etc,
- Neurologic: atlantoaxial subluxation & cauda equinal syndrome
- K Kidney: secondary amyloidosis
- 🧲 Spine: cervical fracture, spinal stenosis:
- Ocular: anterior uveitis (25-30% of pts)
- Pulmonary: upper lobe fibrosis, restrictive changes.
- Nephropathy (IgA)
- Discitis D

In addition, 30-60% of pts have asymptomatic -colitis

## Atypical fibrosis on CXR



#### Reactive Arthritis

- Autoimmune process that develops 1-3 weeks after bacterial infection
- Associated w/ C. trachomatis, gastroenteritis-SSYC
- "can't see, can't pee, can't climb a tree"
- Tx underlying infection, if continues may need analgesics, steroids or immunosuppressants
- PE-Keratoderma blennorhagica, oral or genital ulcers, conjunctivitis

# Reiter's syndrome and Reactive arthritis (ReA)

Infectious trigger factor:

- Enterogenic: Shigella

Champylobacter

Yersinia

Salmonella

- Genito-urinary: Chlamydia

Gonococcal

- Others: Ureaplasma

Clostridium difficile

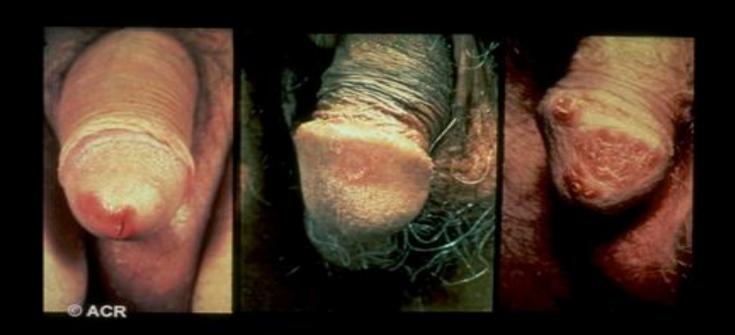
## Reiter's syndrome and Reactive arthritis (ReA)

- Prevalence: 16 per 100,000
- Primarily young adults, aged 20-40 years.
- M:F ratio 1:1 for enterogenic reactive arthritis
- M>F ratio for urogenital reactive arthritis.
- Rare in children and uncommon in Blacks

#### Arthritis

- Mono-or oligo-arthritis (< 5 joints)</li>
- Sacroilitis
- Spondylitis
- Enthesitis
- 80% resolve within 12 months

#### **Balantitis**



## Keratoderma Blenorhagicum



#### Psoriatic Arthritis

- May be present with or without skin findings
  - Look for hidden psoriasis-behind ears, hairline, nail pitting
  - May precede skin findings
- Arthritis does NOT correlate with skin findings
- May cause RA-like arthritis
- Dactylitis common

- Prevalence: 100 per 100,000 (1/20 people with Ps)
- Psoriasis precedes arthritis: 67%
- Arthritis precedes psoriasis or occurs simultaneously: 33%
- Feature of HIV infection

#### Psoriatic Arthritis

- 5 presentation types
- Asymmetrical oligoarticular arthritis
  - dactylitis
- Symmetrical polyarthritis
- Distal interphalangeal arthropathy
  - Nail involvement
- Arthritis mutilans
  - Telescoping motion, opera hands
- Spondylitis with or without sacroiliitis

# **Psoriatic hands**



# Asymmetrical arthritis



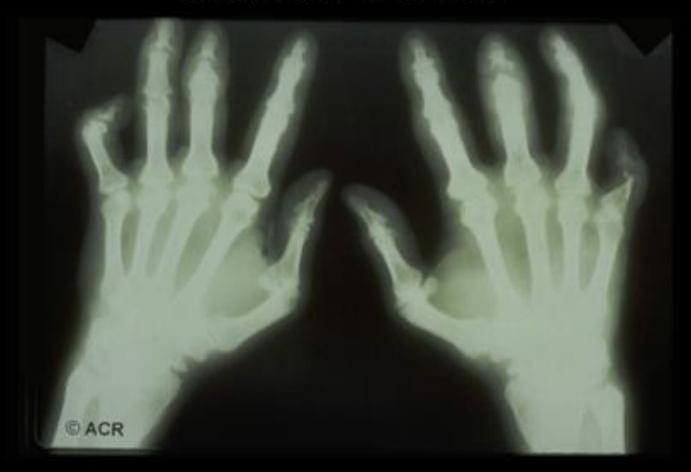
## **Arthritis mutilans**

"telescoping' of digits"



## **Arthritis mutilans**

osteolysis of involved joints



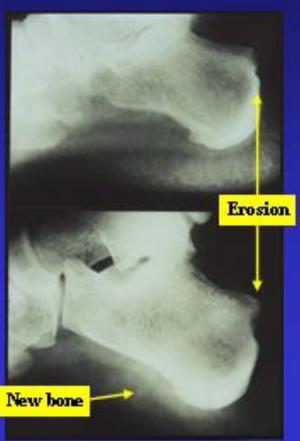
# "Sausage digits"



## **Spondyloarthropathies**

Enthesopathy





### Enteropathic arthritis

- What bowel diseases are associated?
  - Idiopathic, inflammatory bowel disease (UC, CD)
  - Microscopic colitis and collagenous colitis
  - Whipple's disease
  - Gluten-sensitive enteropathy (celiac disease)
  - Intestinal by-pass arthritis

## IBD-related SpA

- Hx of abdominal pain, bloody diarrhea
- Have an axial form and a peripheral arthritis form

### Enteropathic arthritis

- Prevalence: 5 per 100,000
- M:F ratio 1:1
- Children= adults

- Arthritis: acute onset, migratory, asymmetric, oligo-articular (< 5 joints)</li>
- Extra-articular: uveitis, erythema nodosum, aphthous stomatitis, pyoderma gangrenosum

# Erythema nodosum



#### Treatment

- NSAIDs
- Sulfasalizine
  - SE: rash, nausea, diarrhea, and agranulocytosis (rarely)
- TNF- $\alpha$  antagonists
  - Etanercept, infliximab, adalimumab, golimumab
  - Screen for latent TB, hepatitis B, and HIV infection prior to starting therapy
- Corticosteroids-short-term or intra-articular

- Methotrexate, azathioprine, cyclophosphamide, and cyclosporine have been used in SpA.
- Methotrexate is of questionable benefit in ankylosing spondylitis, as various studies have shown conflicting results. Useful in PsA.
- Leflunomide was evaluated in a randomized, double-blind, placebo-controlled study in active ankylosing spondylitis but was not found to be effective.

### Follow-Up

- F/U includes assessment of fatigue, back pain, mobility, synovitis and enthesitis
- Surgery is occasionally useful to correct spinal deformities or to repair damaged peripheral joints.
- Physical therapy, including an exercise program and postural training, is important to maintain function.

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