# Introduction to SPA

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Introduction

SpA disease information

> Pathogenesis

Clinical features



Miss diagnosed

We must (early diagnose)

# **AS Patient with Disappearance of the Lordosis of the Lumbar Spine**





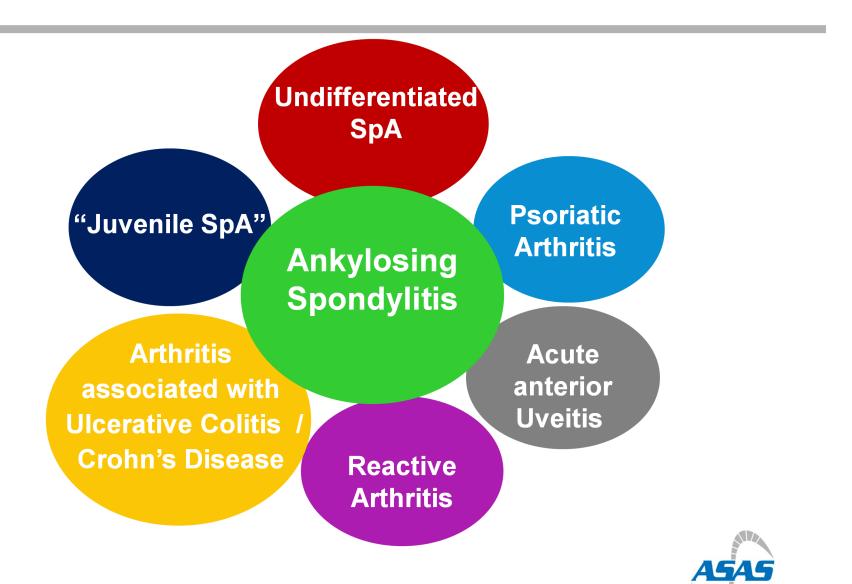
## Final Stage of AS with Severe Kyphosis of Thoracic and Cervical Spine

Unable to look ahead while walking (,patient cannot see the sun')





#### Spondyloarthritides (SpA)



### Reactive Arthritis

- Preceding enteric infection (i.e. diarrhea)
  - Salmonella of various serovars
  - Shigella, especially Shigella flexneri, but also Shigella dysenteriae and sonnei
  - Yersinia including Yersinia enterocolitica 0:3 and 0:9 and Yersinia pseudotuberculosis
  - Campylobacter especially Campylobacter jejuni
  - Clostridium difficile
- Preceding genitourinary infection (i.e. urethritis)
  - Chlamydia trachomatis

#### **Ankylosing Spondylitis - Prevalence**

Country	AS Prevalence
US <sup>1, 2</sup>	0.52%
The Netherlands <sup>3</sup>	0.1%
Germany⁴	0.55%
Norway <sup>5</sup>	1.1 – 1.4%
Haida Indians <sup>6</sup>	6.1%

HLA-B27 Prevalence	
6%	
8%	
9%	
14%	
50%	

<sup>&</sup>lt;sup>1</sup>Helmick CG et al. Arthritis Rheum 2008;58:15-25; <sup>2</sup> Reveille JD et al. Arthritis Rheum 2012;64:1407-11; <sup>3</sup>van der Linden S et al. Arthritis Rheum.1984;27:241-9; <sup>4</sup>Braun J et al. Arthritis Rheum 2005;52:4049-50; <sup>5</sup>Gran T et al. Ann Rheum Dis 1985;44:359-67; <sup>6</sup>Gofton JP et al. Ann Rheum Dis 1966;25:525-7



### The Prevalence of HLA-B27 in the Normal Population and in Patients with Axial Spondyloarthritis in Saudi Arabia

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# ASAS Classification Criteria for Axial Spondyloarthritis (SpA)

In patients with ≥3 months back pain and age at onset <45 years

Sacroiliitis on imaging\*

plus

≥1 SpA feature#

OR

HLA-B27
plus
≥2 other SpA features#

**#SpA features** 

- inflammatory back pain
- arthritis
- enthesitis (heel)
- uveitis
- dactylitis
- psoriasis
- Crohn's/colitis
- good response to NSAIDs
- family history for SpA
- HLA-B27
- elevated CRP

\*Sacroiliitis on imaging

- active (acute) inflammation on MRI highly suggestive of sacroiliitis associated with SpA
- definite radiographic sacroiliitis according to mod NY criteria

n=649 patients with back pain;

Sensitivity: 82.9%, Specificity: 84.4%

Imaging alone: Sensitivity: 66.2%, Specificity: 97.3%



# ASAS Classification Criteria for Spondyloarthritis (SpA)

In patients with ≥3 months back pain and age at onset <45 years

Sacroiliitis on imaging plus ≥1 SpA feature

OR

**HLA-B27** plus ≥2 other SpA features

#### SpA features

- inflammatory back pain (IBP)
- arthritis
- enthesitis (heel)
- uveitis
- dactylitis
- psoriasis
- Crohn's/colitis
- good response to NSAIDs
- family history for SpA
- HLA-B27
- elevated CRP

In patients with peripheral symptoms ONLY

Arthritis or enthesitis or dactylitis

plus

#### ≥1 SpA feature

- uveitis
- psoriasis
- Crohn's/colitis
- preceding infection
- HLA-B27
- sacroiliitis on imaging

#### OR

#### ≥2 other SpA features

- arthritis
- enthesitis
- dactylitis
- IBP ever
- family history for SpA



Sensitivity: 79.5%, Specificity: 83.3%; n=975

# Pathogenesis of spondyloarthritis

### Pathogenic mechanisms of new bone formation

- ➤ Genetic influences
- ➤ Microbes effects
- ➤ Biomechanical stress

### Pathogenic mechanisms of new bone formation

- The cause of AS is not completely understood
- Theory  $\rightarrow$  genetic mechanisms  $\rightarrow$  major role in to AS.
- Genome-wide studies have NOT revealed strong insights on the pathogenesis of new bone formation in AS

#### Genetic influences

- The major gene product associated with AS and the other forms of SpA is human leukocyte antigen (HLA)-B27.
- (HLA)-B27 gene was recognized in 1973.
- HLA-B27 is present in about 80 to 95 percent of patients with AS in most ethnic groups.
- 6 percent of the general population.
- Fewer than 5 % of HLA-B27 carriers in the general population develop disease

Brown MA, Kennedy LG, MacGregor AJ, Darke C, Duncan E, et al. Susceptibility to ankylosing spondylitis in twins 489, Page 6 of 9 Curr Allergy Asthma Rep (2015) 15:489 genes, HLA, and the environment. Arthritis Rheum. 1997;40:1823–8.

#### Genetic influences

Many genes other than HLA-B27.

■ First-, second-, and third-degree relatives of patients with AS have markedly increased risks of developing the disease (relative risks of 94, 25, and 4, respectively)





#### **Axial-SpA Features**



Neck Stiffness

Inflammatory Back Pain (IBP)

Good response to NSAIDs

Alternative Buttock Pain

X-Ray -Sacroiliitis

- Age at onset < 45 years</li>
- Family History of SpA
- Elevated CRP levels
- +ve HLA-B27



Crohn's/Colitis

**Psoriasis** 

Peripheral Arthritis

**Dactylitis** 

**Enthesitis** 

### ASAS Inflammatory Back Pain Criteria by Experts (Chronic Back Pain; n=648)

- age at onset < 40 years</li>
- insidious onset
- improvement with exercise
- no improvement with rest
- pain at night (with improvement upon getting up)

Sensitivity: 79.6%; Specificity: 72.4%

Inflammatory back pain present if <u>at least 4 out of 5</u> parameters are fulfilled.

# Acute Arthritis of the Right Knee in a Patient with Peripheral Spondyloarthritis

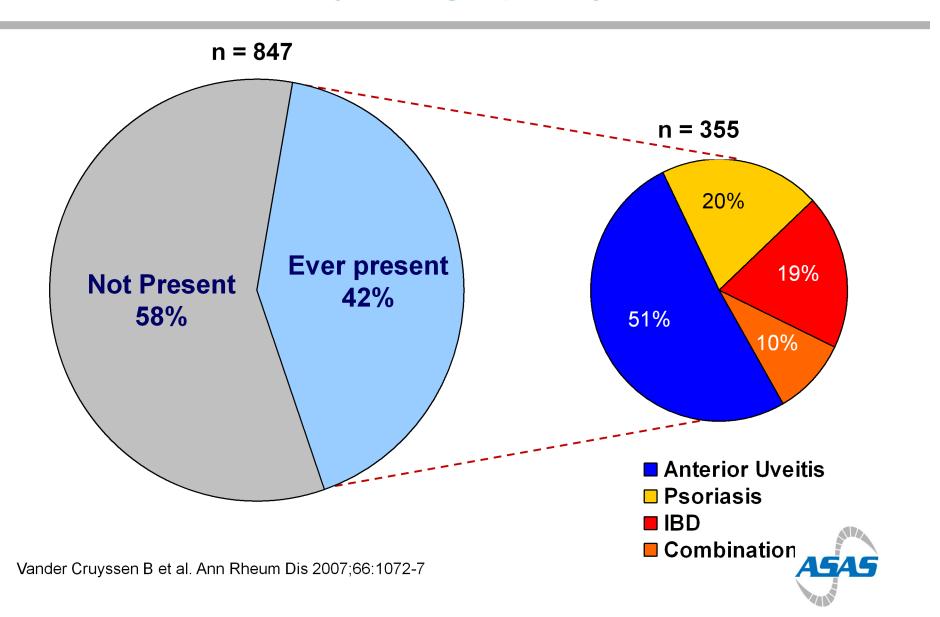




### Peripheral arthritis

- Predominantly involves the lower extremities.
- Arthritis is frequently asymmetrical and often affects only one to three joints.
- The severity ranges from mild to disabling.
- The presence of asymmetrical oligoarthritis is very suggestive of SpA, but its absence would not be helpful in excluding this possibility.

# **Extra-Articular Manifestations** in Ankylosing Spondylitis



# **Eye: Acute Anterior Uveitis** in Spondyloarthritis

- Acute onset
- Unilateral
- Anterior
- Spontaneous remission
- Recurrent
- Related to HLA B27





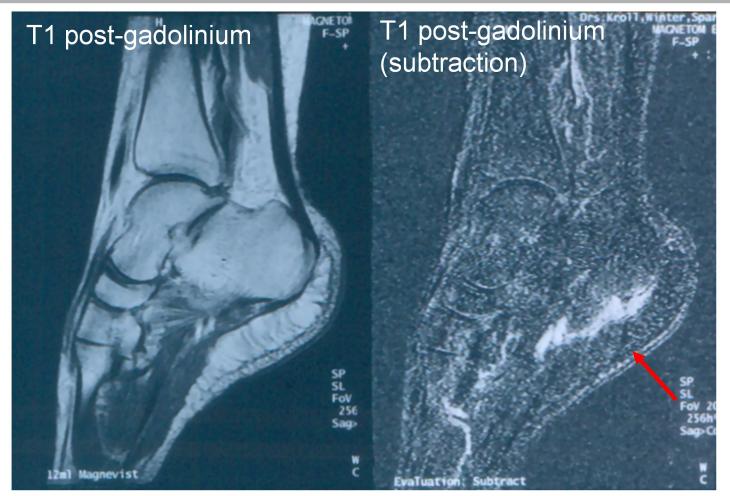
# Enthesitis (Insertion of Achilles Tendon at Calcaneus) Right Heel



#### **Enthesitis**

- Relatively specific to SpA
- The most common  $\rightarrow$  at the insertion of the Achilles tendon
- Plantar fascia ligament into the calcaneus
- severe pain and tenderness

#### **Enthesitis of the Plantar Fascia by MRI**





### **Dactylitis**



<sup>\*</sup> Psoriasis plaque lesions



### **Dactylitis**

- known sausage toe or sausage finger
  - Psoriatic arthritis
  - Occasionally reactive arthritis
- Unlike synovitis, in which swelling is confined to the joints, with dactylitis, the entire digit is swollen.
- Dactylitis is not specific for SpA and may also be seen
  - tuberculosis
  - syphilis
  - sarcoidosis
  - sickle cell disease
  - tophaceous gout

# **Skin Manifestations** in Spondyloarthritis

- Psoriasis
- Erythema nodosum
- Pyoderma gangrenosum
- Keratoderma blenorrhagicum



#### **Psoriasis**



Typical psoriasis with scaly patches and plaques



### Mild Nail Changes in Patient with Psoriasis





### Moderate to Severe Nail Changes in Patient with Psoriasis



#### **Psoriasis**





- Psoriasis is associated with all forms of SpA.
- Psoriasis is present in up to approximately 10 percent of patients with AS.

#### Palatal erosion in reactive arthritis



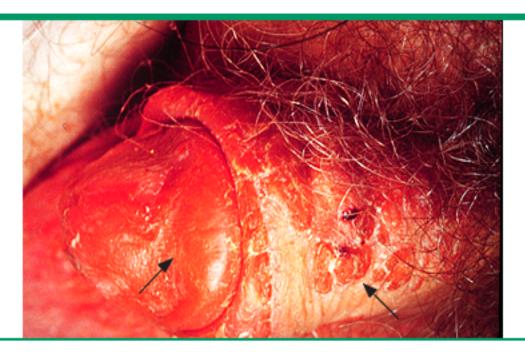
A sharply demarcated erosion of the hard palate is shown. This is among the more common of the oral manifestations of reactive arthritis.

#### Keratoderma Blenorrhagicum





#### Circinate balanitis



Circinate balanitis characterized by shallow ulcers on the glans penis and the shaft of the penis (arrows). The lesions are generally asymptomatic.

Courtesy of Professor Victor Newcomer, UCLA.



### SpA LAB

#### • HLA-B27

- 90 percent of patients with ankylosing spondylitis
- 50 to 70 percent of patients with other forms of SpA
- a positive HLA-B27 by itself is not diagnostic of SpA

#### Acute phase response

- Erythrocyte sedimentation rate (ESR) and levels of C-reactive protein (CRP)
- increased in between 35 and 50 percent of patients with axial SpA
- Elevated levels of CRP are also a predictor of radiographic progression and for a good response to tumor necrosis factor (TNF)-blocker therapy

#### Sacroiliitis Grade 0 (Normal)





#### Sacroiliitis Grade 2 Right, Grade 1 Left





#### **Sacroiliitis Grade 3 Bilaterally**





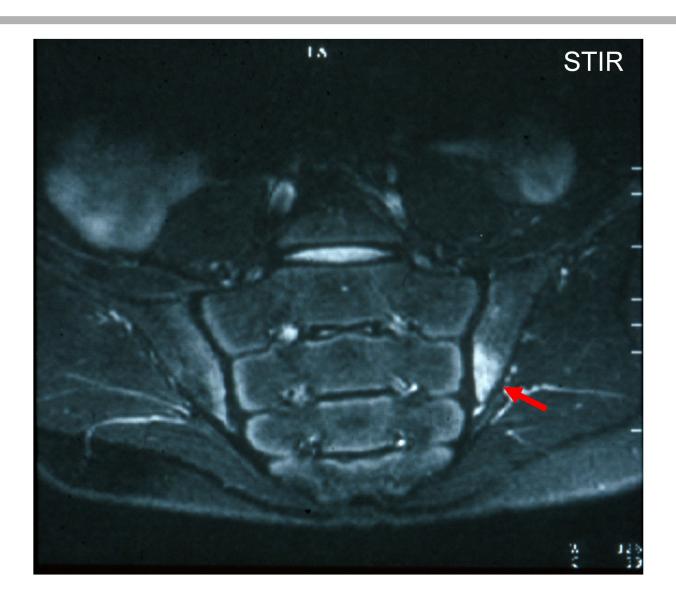
#### **Sacroiliitis Grade 4 Bilaterally**





### Early (limited) Sacroiliits

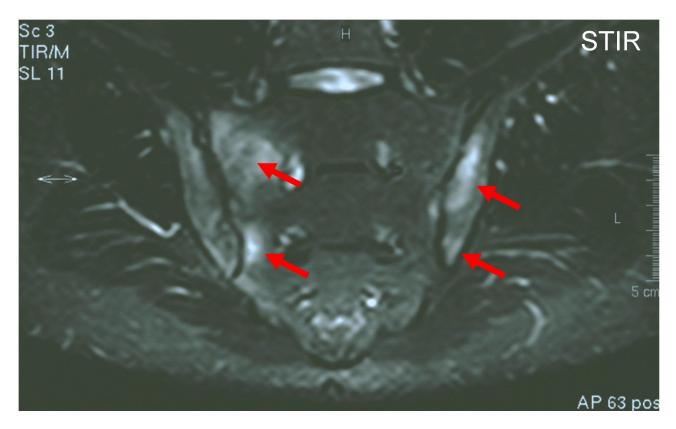
(Bone Marrow Edema)





#### **Definition of Positive MRI-SI Joint**

- subchondral bone marrow edema
- acute (bilateral) sacroiliitis





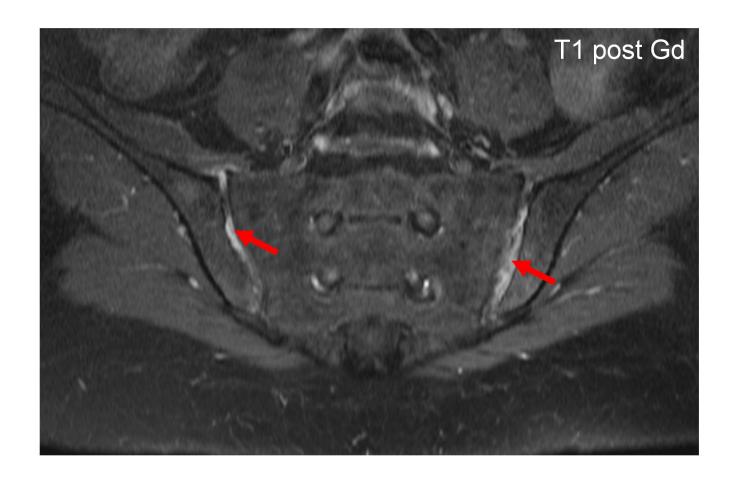
#### **Bone Marrow Edema (BME) / Osteitis**

Bone marrow edema may be associated with structural changes such as erosions.





#### **Synovitis of Sacroiliac Joint**





# ASAS/EULAR Recommendations for the Management of Ankylosing Spondylitis

**NSAIDs** Education, exercise, **Peripheral Axial** physical disease disease therapy, rehabilitation, patient Sulfasalazine associations, Local corticosteroids self help groups **TNF Blockers** 



