

Psoriatic Arthritis

Psoriatic arthritis (PsA)

Inflammatory arthropathy associated with psoriasis, usually seronegative and without rheumatoid nodules.

Output the second se by clinical changes, including arthralgia, hyperemia, swelling, often with prolonged morning stiffness and characteristic radiological manifestations, occurring in patients with psoriasis or may be precursor to psoriasis.

PsA

- The association between arthritis and psoriasis was observed in XIX-th centure.
- In the 1960s, based on epidemiological and clinical studies, it became clear that unlike RA, arthritis associated with psoriasis is usually seronegative, often involving distal interphalangeal joints of the fingers, vertebrae and sacroiliac, has distinctive radiological features and considerable family aggregation. In the 1970s..., PsA was included in the Seronegative
 - *spondilarthritidies* (SNSA) category due to the similar events with those of AS and ARe.

Epidemiology

- The prevalence of PsA is estimated to be 5-30% among patients with psoriasis.
- In Caucasians psoriasis has a prevalence of approx. 1-3%.
- Psoriasis and PsA are less common in other ethnic groups.
- First-degree relatives of patients with PsA have an increased risk of disease (up to 30%) for both PsA and other forms of SpA.
- In monozygotic twins, the concordance for psoriasis varies from 35 to 72%, and for APs 10-30%.

PsA

- The prevalence of psoriasis in the general population: 0.1-2.8%.
- The prevalence of psoriasis in arthritis patients: 2.6-7.0%.
- The prevalence of arthritis in the general population : 2-3%.
- The prevalence of arthritis in patients with psoriasis : 6-42%.

PsA

It evolves with erosive and destructive articular modifications in \approx 40-60% pts. The male-female ratio 1:1 OPSOriatic arthritis develops mainly in people aged 35-55. In juvenile form, the age of onset is 9-11 years.

PsA. Etiopathogeny

Genetic
Trigger factor
Immunological abnormalities

Trigger factors Mechanical trauma Infection (Streptococci..., HIV etc.) Stress



Psoriatic Arthritis

Psoriatic arthritis. Etiological factors of psoriasis



Genetic factors

Sevidence: family aggregation and human genome scanning - 1p, 4q, 6p, 17q

Ohromosome 6p:

- HLA B13, B17, B39, C*0602: Ps & PsA
- HLA B27: male, axial involvement
- HLA DR4: erosive, polyarticular form
- HLA B27, B39, B57, DQ3: progressive
- HLA Cw6: psoriasis, juvenile onset
- MICA-002: PsA
- polymorphism promoter TNF-α

Other:

- Polymorphism of the HCP5 locus related to HLA-B27 locus, and IL-23R, IL-12B (chromosome 5q31)

Immunological abnormalities Angiogenesis (VEGF, Angiopoietin) Inflammatory Infiltration Ocitokines Ortilage and bone degradation () RANKL-RANK, ↓ OPG, BMP)

• Fibrosis and ankylosis (PDGF, TGFβ)

VEGF Vascular endothelial growth factor PTGF Platelet-derived growth factor TGF Transforming growth factor BMP Bone morphogenetic proteins OPG Osteoprotegerin

Cells and pathways in PsA



Pathogenesis

- PsA is a immune-mediated disease sharing pathogenetic mechanisms with psoriasis
- The PsA synovium presents with infiltration of T cells, B cells, macrophages and cells expressing NK receptors.
- Expansion of CD8+ LT clones is common in PsA.
- In Plasmacitoid dendritic cells probably play a key role in psoriasis, as well as PsA.
- Synovia is abundantly infiltrated by pro-inflammatory cytokines. IL-2, IFNγ, TNF-α, IL-1β, IL-6, IL-8, IL-10, IL-12, IL-13, and IL-15 are found in synovial fluid of patients with PsA.
- Over Cytokines derived from LTh17 are important in PsA, with genetic association with IL-12 / IL-23 axis genes.
- Osteoclast precursor is increased in peripheral blood of patients with PsA, as well as increased RANKL activity in synovia.

Pathogenesis of psoriatic arthritis



HLA-B27 misfolding and IL-23 / IL-17 axis. Possible consequences of HLA-B27 misfolding



IL-23 and enteseal LT promote enthesitis and bone proliferation



Inflammation and new-bone formation: differences between rheumatoid arthritis and SNSA



Clinical classification of PsA

- Oistal predominant (distal interphalangeal joint affected). Oligoarticular (<5 joints) more</p> often with asymmetric distribution. Olyarticular (>5 joints) (pseudorheumatoid). •Axial.
- "Arthritis Mutilans".

Joints affected by psoriatic arthritis



Clinical types of PsA



- Wright V. Psoriasis and arthritis. Ann Physical Med 1959 1. 2.
 - Moll, Wright, Psoriatic Arthritis, Semin Arthritis Rheum 1973

PsA, clinical picture

- In 60–70% of cases, psoriasis precedes joint damage.
- In 15–20% both manifestations occur in an interval of 1 year from each other.
- In approx. 15–20% arthritis precedes the onset of psoriasis, which is a diagnostic problem.
 M:F ratio 1:1.
- The onset of the typical disease is in the young adult with the average age of 37 years.
 The disease can also start in childhood.

PsA, clinical picture

 Psoriatic arthritis manifests with arthralgia, hyperemia and swelling of the joints, disturbance of their function, often with morning stiffness. **Distinctive features of PsA, (and SpA):** • enthesopathies (inflammation of the areas of insertion) of the tendon, ligament and joint capsule); dactylitis (swelling of fingers, fingers "in sausages"); asymmetric joint damage, sometimes oligoarticular involvement; Iritis/uveitis.

Distal interphalangeal arthritis A rarer form of PsA (8-16%) It is usually associated with nail psoriasis (it may be the only localization of the pathological process). Oistal interphalangeal joint injury is considered a classic and unique feature for psoriatic arthritis, it appears primarily in men.

Nail affection

OPitting (pitting edema), Output A strips, Onycholysis, Yellowish discoloration of the edges of the nails, Oystrophic hyperkeratosis, Combinations of disorders.



Distal interphalangeal arthritis



Asymmetric oligoarthritis

- The most common form of APs (about 50%).
 It is manifested by the asymmetric inflammation of distal and proximal interphalangeal joints of the hands and plants, and metacarpophalangeal joints.
- The association of tenosynovitis of the flexors achieves the appearance of the finger *"in sausages"*, concomitantly being affected less than 5 joints.

Asymmetric oligoarthritis



Symmetrical polyarthritis

- It is the clinical variant that mimics appearance of RA.
 It evolves with sacroiliitis (a particular feature of this form).
- About 25-40% of PsA patients have symmetrical polyarthritis.
- Affected joints radio-carpal, radio-ulnar, proximal and distal interphalangeal, with a tendency towards ankylosis.
- It is less extensive and less deforming than seropositive RA.

Symmetrical polyarthritis



Polyarticular pattern



Arthritis mutilans

- It is characterized by osteolytic lesions of the phalanges and metacarpal joints, which are often associated with sacroiliitis and extensive skin lesions.
- Sually fingers, toes, hands, and feet are affected. Fingers or toes that shorten and bunch together, which is called telescoping.
- Apr. 5% of PsA patients.
- Sometimes with ankylosis and contractures in other fingers.
- Oulike RA, many patients have temporary remissions.
- Erosive disorders with progressive evolution towards deformation, osteolysis and disability, induce a significant increase in mortality, compared with the rest of the population.

Arthritis mutilans





Arthritis mutilans

"Telescoped" fingers



Axial form

- Involvement in the process of the spine and/or sacroiliac joints.
- Sacroiliitis (often unilateral).
- May dominate the clinical picture.
- Olinical picture similar to AS, but associated with a serious evolution of psoriasis.
- Clinical signs of spondylitis and/or sacroiliitis may be associated with other clinical forms of psoriatic arthritis.

Axial form

- Spondylitis may manifest without radiological signs of sacroiliitis or with radiological signs, but without morning stiffness.
- There is no correlation between the clinical and radiological signs of sacroiliitis.
- The vertebrae are affected asymmetrically with the possible involvement of the atlantoaxial joint with odontoid erosion and subluxation.
- Radiologically, asymmetrical nonmarginal syndesmophytes, paravertebral ossification and, vertebral ankylosis with calcification of the disc can be detected.
- Axial arthropathy without peripheral involvement in 5% of patients with PsA.

Psoriatic spondyloarthritis







Juvenile psoriatic arthritis

- Approx. 8-20% of all arthritis in children.
- Usually, monoarticular at onset (girls 9-10 years).
- The disease develops slowly, sometimes can be severe, with high activity, progressing to adulthood.
- In more than 50% of patients, arthritis is monoarticular.
- 35-40% affecting distal interphalangeal joints.
- 30% tenosinovitis
- ●63-71% nail damage "pitting" (point erosion),
- In 39-47% pts there are bone growth disorders and, as a result, the shortening of the bones due to the affect of the epiphysis of the bone by inflammatory process.
- Sacroiliitis occurs in 28% pts and is associated with HLA-B27.
 The presence of HLA-B8 may serve as a marker of disease severity.

Enthesitis and enthesopathy

Enthesitis, an important feature of PsA, represents inflammation at the place of bone insertion of the tendons, ligaments and capsules of the joints.


Dactylitis, tendinitis and tenosynovitis

In the etymological sense, the term dactylitis (gr. Dactylos- finger, itis- inflammation) defines the inflammation localized at the level of the fingers of the hands or feet. • Dactylitis in PsA is common (22%). Specificity 82.4%, sensitivity 84.9. Item in the CASPAR (Classification Criteria for Psoriatic Arthritis) criteria for the diagnosis of PsA.

Systemic manifestations of PsA

- Lymphadenopathy,
- Heart disease (most commonly aortic regurgitation, aortitis),
- Renal impairment (proteinuria, hematuria, cylindruria),
- Ocular impairment (conjunctivitis, iritis),
- Neuromuscular impairment,
- Vascular affection (Raynaud's Syndrome) etc.
- 10-15% of patients with APs have ocular damage through anterior uveitis, iritis, iridocyclitis, sometimes relapsing.

Impairment of the cardiovascular system

- Possible in 18-22% of cases of PsA.
- Clinically manifested by dyspnoea, heart palpitations, pain in the heart region of different intensity, caused by the development of aortitis, myocarditis, pericarditis or myocardiodistrophy.
- Oifferent rhythm and cardiac block disorders.
- Cases of severe pericarditis with progressive heart failure, complete atrioventricular block are described.
- A ortic valve regurgitation may develop (7% of cases).

Pulmonary involvement

It manifests as a result of limiting the respiratory excursion of the thoracic box with the development of pulmonary emphysema as a consequence of kyphosis and affecting the costovertebral joints in the axial variant of PsA. As in ankylosing spondylitis, apical
 As in ankylosing spondylitis.
 As in ankylosing spondyli pneumofibrosis is characteristic, which is extremely rare (1-2%) and which must be differentiated from specific lung conditions.

Renal impairment

The kidneys are most often involved in the systemic pathological process in APs (approx. 27-36%).

- According to some authors, this is caused by
 - hypergamaglobulinemia and, in particular, by the increased level of IgA with the development of IgA-nephropathy.
- At the same time, edema, renal hypertension, secondary anemic syndrome and renal failure develop in the final stages of the disease.
- Secondary renal amyloidosis(14-17%).
- Often, the cause of renal impairment may be prolonged administration of NSAIDs, manifested by urinary syndrome with proteinuria and microhematuria.

Affection of the peripheral nervous system

- Caused by root damage in the cervicothoracic and lumbosacral segments of the spine.
- Caused by joint damage in the cervicothoracic and lumbosacral segments of the spine.
- At minimal trauma, in the presence of disorganization of the transverse ligament of the atlanta, the atlantoaxial subluxation may develop (2-3%).
- Ser rarer "cauda equina" syndrome from chronic epiduritis, accompanied by impotence and urinary incontinence.

Laboratory diagnosis

Tests to determine the activity of the disease and to monitor the evolution:

- OBC
- ESR
- OC-reactive protein
- Fibrinogen
- In <10% of patients positive with RF, ANA and anti-CCP.
- In patients with extensive psoriasis - hyperuricemia

For differential diagnosis Rheumatoid factor
 ● HLA-B27 Ourine analysis Biochemistry (ALT, AST, bilirubin,
 alkaline phosphatase, urea, creatinine, proteins) Joint aspiration (in case of synovitis) with bacteriological examination (molecular-biological by polymerization chain reaction to the DNA of the provocative agent) and clinical of the synovial fluid

Radiological joint examination

- The hallmark of PsA is the combination of erosive change with bone proliferation, in a predominantly distal distribution (e.g. interphalangeal more than MCP joints). Terminal narrowing of phalanges with acroosteolysis and cup deformation of proximal portions ("pencil in cup"). • Arthritis mutilans: marked bony resorption and the consequent collapse of soft tissue; when this affects the hands, it can cause a phenomenon sometimes referred to as "telescoping fingers".
- Our Output of the second se
- Psoriatic arthritis associates enthesopathy, as a characteristic feature, absent in rheumatoid arthritis.

"Pensil in cup"



X-ray examination

inflammatory and destructive changes in metatarsal-phalangeal and interphalangeal

Marginal bone erosions with adjacent bone proliferation





increased bone density and proliferation of the distal phalanx ("ivory phalanx")

PsA acro-osteolizis





Dactilitis



"Licked candy stick appearance"







X-ray exam

Paravertebral ossifications in comma form

Anteroposterior sacral X-ray: bone erosion and narrowing art. sacroiliac with partial fusion.

Bone sclerosis and inflammatory enthesopathy

CT of sacroiliac joints

PsA: MRI

Erosion at the base of the middle phalanx

(a)Until the contrast is introduced(b)After contrast

MRI: sacroiliac joints

PET/CT in PsA diagnosis

CT image

Combined PET/CT image

PET image

PsA diagnosis

- To establish the diagnosis of PsA it is necessary to go through 2 stages : • Establishing the diagnosis of seronegative spondyloarthropathy (Criteria for spinal inflammatory pain, ESSG criteria, Amor, ASAS)
- Classification of SNSA established as psoriatic arthritis (CASPAR criteria)

Criteria set for inflammatory back (spinal) pain (IBP)

Calin Criteria for IBP (1997)	Berlin Criteria for IBP (2005)	ASAS criteria for IBP, (mnemonic "iPAIN") (2009)
 Age at onset <40 yrs. Back pain duration >3 months Insidious onset Morning stiffness Improvement with exercise 	 Morning stiffness >30 min Improvement with exercise, not with rest Awaking at the second part of night because of pain Alternating buttock pain 	 Insidious onset Pain at night (with improvement after getting up) Age at onset <40 yrs. Improvement with exercise No improvement with rest
IBP if 4/5 are present	Sensibility - 70% Specificity - 81% if 2/4 are present	Sensibility - 77.0% Specificity - 91.7% if 4/5 are present

Amor	classification	criteria	for	SpA	(1990)
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A. Clinical Symptoms/History	Score
1. Pain at night (spine) or morning stiffness	1
2. Asymmetrical oligoarthritis	2
3. Gluteal (buttock) pain (any)	1
or	
alternating gluteal pain	2
4 Sausage like digit or toe (dactylitis)	2
5. Enthesitis (heel)	2
6. Uveitis	2
7. Urethritis/Cervicitis within 1 month before onset of arthritis	s 1
8. Diarrheae within 1 month before onset of arthritis	1
9. Psoriasis, balanitis or inflammatory bowel disease	2
B. X-rays	
10. Sacroiliitis (grade 2 bilaterally or grad 3 unilaterally)	з
C. Genetical background	
11. HLA-B27 positive or positive family history for AS.	
ReA, uveitis, psoriasis or inflammatory bowel disease	2
D. Good response to NSAIDs	
12. NSAIDs show a good response within 48 hours, or	
relapse within 48 hours after NSAID are stopped	2

CASPAR Criteria (CLASsification criteria for Psoriatic ARthritis) Inflammatory articular disease (joint, spine, or enthesis) AND at least 3 points from the following categories:

- Current psoriasis (2 points), a personal history of psoriasis (1 point), or a family history of psoriasis (1 point)
- 2. Typical psoriatic nail dystrophy (1 point)
- 3. Negative rheumatoid factor test (1 point)
- Dactylitis: current or previous episode noted by a rheumatologist (1 point)
- Juxta-articular new bone formation on hand or foot radiographs (1 point)

Differential diagnosis

Skin lesions - seborrheic dermatitis Nail injuries - fungal infections Output Articular impairment: polyosteoarthritis (Heberden, Bouchard nodules), rheumatoid arthritis, septic arthritis, gout, seronegative spondyloarthritis (AS, ReA, arthritis in inflammatory bowel disease, chronic juvenile arthritis)

Differential diagnosis of PsA

Manifestation	PsA	AS	RA	ReA
M:F	1:1	3:1	2:1	8:1
RF	<10%	negative	80%	negative
DIP involvement	30-50%		Noncharacteristic	
Peripheral arthritis	90-95%	40%	100%	90%
	Asymmetric,	Asymmetric,	Symmetric,	Asymmetric
	Large & small joints	Small joints	Large joints	
Sacroiliac/axial	35%,	100%	Cervical spine at	20%
ioints	any level		late stages	Lumbar
				predominance
Others	Enthesitis	Enthesitis	Muscular atrophy	Enthesitis
	Dactilitis			Keratoderma
	Periarticular erythema			Balanitis circinata
Extraarticular	Skin eruptions	Uveitis	Nodules	Uveitis
features	Nail dystrophy	Cardiac	Sjögren syndrome	
			Vasculitis	
HLA-B27	10-25%	90%	Negative	40%
X-ray	Bone erosions, DIP	Syndesmophytes,	Periarticular	Periostitis /
	Periostitis /	osteitis, bone	osteopenia	osteopenia
	proliferation	erosions,	Erosions	Asymmetric
	Syndesmophytes	Sacroiliitis		sacroiliitis

Tools for evaluation and monitoring of PsA

GRAPPA (Group for Research) and Assessment of Psoriasis and **Psoriatic Arthritis)** OMERACT (Outcome Measures) in Rheumatology)

Fields of evaluation OMERACT 8.

Rheumatological

- Peripheral joints
- Dactylitis
- Enthesitis
- Axial involvement

Functional

- Life quality
- Fatigability

Dermatological

- Skin
- Nails

Radiological

- Modified Sharp score
- Erosions score
- Joint space
 narrowing

- Tender and swollen joint count,
- The area of skin involvement (Psoriasis Area and Severity Index (PASI),
- Nail affection: (Nail Psoriasis Severity Index (NAPSI) or Modified Nail Psoriasis Severity Index (mNAPSI),
- Enthesitis: Mander/Newcastle Enthesitis Index (MEI), Leeds Enthesitis Index (LEI), Spondyloarthritis Research Consortium of Canada (SPARCC), Maastricht Ankylosing Spondylitis Enthesis Score (MASES),
- Dactylitis: Leeds Dactylitis Index (LDI),
- Patient Global for Psoriatic Arthritis,
- Dermatology Life Quality Index (DLQI),
- Psoriatic Arthritis Quality of Life (PsAQOL),
- Functional Assessment of Chronic Illness Therapy Fatigue (FACIT-F),
- Psoriatic Arthritis Response Criteria (PsARC),
- Psoriatic Arthritis Joint Activity Index (PsAJAI),
- Disease Activity in Psoriatic Arthritis (DAPSA),
- Composite Psoriatic Disease Activity Index (CPDAI)

Rheumatological evaluation

Peripheral joint involvement Axial involvement Enthesitis Dactylitis

Minimal disease activity score (MDA)

- The patient is considered to have achieved MDA if he meets 5 of the following 7 criteria :
- Number of tender joints ≤1
- Number of swollen joints ≤1
- ●PASI (Psoriasis Activity and Severity Index) ≤ 1 or
 ↓
 - the affected body surface ≤3%
- ●Patient pain VAS ≤15 mm
- Global patient activity VAS ≤20 mm
- ●HAQ (Health Assessment Questionnaire) ≤0.5
 ●Painful entheses sites ≤1

Tender/swollen joints

 British Society for Rheumatology (BSR) recommends count of 66 swollen and 68 tender joints.

Global disease activity assessment: patient / physician

Visual analog scale (VAS) (10 cm/100 mm)

The Likert scale can be used.

Patient pain score

Visual analog scale (VAS) (10 cm/100 mm)

The respondent is asked to place a line perpendicular to the VAS line at the point that represents their pain intensity for the last 24 hours

Dermatology Quality Life Index (DLQI)

- DLQI is a measure of quality of life (QoL) taking into account skin disease and aspects of psoriasis and PASI.
- The DLQI contains 10 simple questions related to how the skin disease affects the patient's life. The quality of life for the last week is evaluated. It is completed by the patient and lasts approx. 2 minutes.

DLQI is calculated by summing the scores for each question Maximum score - 30; Minimum score - 0. Higher scores indicate greater impairment of quality of life.

DLQI Score

0-1 = without effects on the patient's life 2-5 = small effects on quality of life 6-10 = moderate effects on quality of life If DLQI >10, the quality of life is affected, for which intervention is required.

11-20 = quality of life greatly affected 21-30 = the quality of life extremely affected

Health Assessment Questionnaire (HAQ)

Two-pages questionnaire contains : • HAQ Disability Index (HAQ-DI) – assesses the patient's functional ability •HAQ VAS pain score – assesses the presence or absence of joint pain and its severity • HAQ VAS patient global score – assesses the quality of life in general
Health Assessment Questionnaire (HAQ) HAQ DI Category Devices used for help **Dressing & Grooming** Dress yourself, including shoelaces and buttons? Shampoo your hair? Arising Stand up from a straight chair? Get in & out of bed? Eating Cut your own meat? Lift a full cup or glass to your mouth? Open a new milk carton? Walking Walk outdoors on flat ground? Climb up five steps? Wash and dry your body? Take a tub bath? Get on Hygiene and off the toilet? Reach and get down a 5 pound object (such as a Reach bag of sugar) from above your head? Bend down to pick up clothing from the floor? Grip Open car doors? Open previously opened jars? Turn faucets on and off?

ou usually use for any the above activities:

aised toilet seat

Bathtub seat

Long-handled appliances n bathroom

ong-handled appliances for reach

Jar opener (for jars previously opened)

Assessment of PsA patient with axial involvement

Domain	Tools
Patient global assessment	VAS for the last week
Back pain	The average value of VAS for the last week:
	1) Pain caused from SpA
	2) Night pain
Spinal stiffness	VAS morning stiffness
Spinal mobility	Chest expansion
	Modified Schöber test
	Occiput – wall distance
	Cervical rotation
	Lateral spinal flexion or BASMI
Physical ability	BASFI or Dougados Functional Index
Peripheral joints and entheses	Swollen joint count (count of 44 joints).
	Enthesitis scores MASES, San Francisco, Berlin
Fatigability	The fatigue question from BASDAI
Acute phase indicators	ESR or CRP
Imagistics	Lateral lumbar, cervical vertebral and sacral X-ray (sacroiliace & coxofemoral)

Dactylitis

• sausage finger" = flexors tenosynovitis + interphalangeal synovitis • Acute (swelling + redness + pain) / chronic Observation of the second s LDI (Leeds Dactylitis Instrument)

Enthesitis

- Newcastle Enthesitis Index (NEI) 66 pct
- Maunder Entheses Index (MEI) 66 pct; 0–3 p. palpatory pain
- Maastricht Ankylosing Spondylitis Entheses Score (MASES)
 - 13 pct
- Spondyloarthritis Research Consortium of Canada (SPARCC) – 8 pct
- Leeds Enthesitis Index (LEI) 6 pct:
- lateral epicondyle 2
- medial epicondyle 2
- Achilles tendon insertion 2

MASES:

Maastricht Ankylosing Spondylitis Enthesitis Score



- 13 sites
- Easy to locate
- No grading
- Score from 0 to 13.

Costochondral 1 ri/le Costochondral 7 ri/le Spina iliaca anterior superior ri/le Crista iliaca ri/le Spina iliaca posterior ri/le Proc. spin L5 Achilles tendon prox. insertion ri/le



Evaluation of response to treatment

PSARCACR

Psoriatic Arthritis Response Criteria (PsARC)

Clegg 1996

Criteria TJC SJC Patient global assessment Physician global assessment

PsARC response

Improvement of at least two of the following 4 criteria (one of which must be TJC/SJC) without worsening any criterion:

- 20% or more improvement in the overall assessment of the disease activity by the physician
- 20% or more improvement in the overall assessment of the disease activity by the patient
- 30% or more TJC improvement
- 30% or more SJC improvement

ACR response criteria

Parameters

TJC SJC Pain (VAS 0-10 cm) Global patient assessment Global physician assessment ESR CRP HAQ

ACR response

20%; 50%; 70% improvement in TJC, SJC 20%; 50%; 70% improvement in 3 from 5

Treatment of PsA The purpose of APs treatment is to relieve pain, reduce inflammation, maintain joint flexibility and normal posture, reduce functional limitations, prevent ankylosis; OPatient education: exercises, kinetotherapy, physiotherapy, balneotherapy

Treatment of PsA

- Medication:
- NSAIDs;
- •analgesics;
- Iconstruction of the second second
- Remissive treatment with DMARDs (Methotrexate,
 - Leflunomid, Sulfasalazine) low efficacy, multiple adverse reactions.
- Biologic remissive treatment (anti-TNFα etc.).
 Surgical treatment (orthopedic) if necessary;
 Rehabilitation treatment.

NSAIDs

NSAIDs are useful and effective in most patients with PsA. They reduce pain and signs of joint inflammation, the main mechanism of action being the inhibition of prostaglandin synthesis :

- oreduce pain and inflammation;
- does not change the progression of joint erosions;
 does not influence the occurrence of extra-articular manifestations;
- It is the symptomatic effect and manifest only during the treatment.

Posology A. Non-selective inhibitors COX-1, COX-2. • Diclofenac - 150 mg/day (75 mg b.i.d.) Ibuprofen - 400 mg q.t.d. B. Selective inhibitors COX-2. Meloxicam - 15-7,5 mg/day, q.d. Nimesulide - 100 mg b.i.d. C. Ultra-selective inhibitors COX-2 (Coxibs). Celecoxib - 100 -200 mg/day 1 q.d. Etoricoxib – 30-120 mg/day 1-2 time/day

Glucocorticoids

• Caution (exacerbations of skin manifestations upon discontinuation of therapy).

Indications:

• Active forms that do not respond to NSAID administration.

- Prednison 10-20 mg/day with gradual dose tapering.
- Local corticosteroids (mono-, oligoarthritis) intra-articular or intra-tendon (in a single joint no more than 3-4 infiltrations per year).
- Ocrtisone preparations may be indicated locally and as an ointment, active on skin manifestations.
 Ocrtisone preparations and a second se

Remissive treatment

- Methotrexate is effective in the treatment of skin and joint manifestations.
- •7,5-25 mg/week.
- Methotrexate is indicated in mild to severe forms of PsA, which do not respond to NSAIDs treatment, being preferred to corticosteroids
- Mtx effectiveness level C of recommendations.

Remissive treatment

- Sulfasalazine initially in doses of 500 mg/day, then increasing the weekly dose by 500 mg, extending to 3-4 g/day, which is continued for a long time.
- The therapeutic effect occurs over 4-6 weeks.
 In clinical trials SZA has not shown high efficacy in the treatment of PsA. It can be used as an alternative medicine in the peripheral forms of PsA.

Leflunomide

- It inhibits dihydro-orotate-dehydrogenase (DHODH), a mitochondrial enzyme required for de novo synthesis of pyrimidine nucleotides. Thus, the proliferation of lymphocytes is blocked.
- It is given in a dose of 20 mg/day. It can be used in both early and late forms of the disease. The therapeutic response is rapidly established, being maximal after 4 weeks (initial dose of 100 mg/day for first 3 days).
- The efficacy is maintained after 2 years of treatment.
- IF slows the radiological progression of the disease in all its stages, more than placebo, MTX or SZA.
- LF is recommended in the treatment of peripheral forms of PsA (recommendation level A)

Cytostatics

- Cyclosporine A,
 Cyclophosphamide,
 Azathioprine,
 D popicillomine
- D-penicillamine
- are not recommended for wide administration in the treatment of APs!

Biologic therapy • Biological agents are substances that interact with the specific components of inflammation, the following groups of biological agents being highlighted : Monoclonal antibodies (MAB). Receptor antagonists. Soluble receptors.

Biologic therapy

- Infliximab (Remicade[®]): 5 mg/kg/, i.v., week 0, 2, 6 and then every 8th week. It is effective in patients with moderate to severe active disease.
- Adalimumab (Humira[®])*: s.c., 40 mg every second week. Effective in high activity forms.
- Etanercept (Enbrel®)*: s.c., 25 mg x 2/week or 50mg 1/week.
 Infliximab, Adalimumab, Etanercept have similar efficacy and
 - adverse reaction profile in PsA and are recommended in the treatment of active disease in patients with failure or intolerance in the treatment with at least 2 DMARDs.
 - **Note**: Before initiation of treatment it is mandatory: screening for tuberculosis: PPD / quantiferon test and chest x-ray; viral infection screening: AgHBs, anti-HCV, HIV; exclusion of neoplasia, demyelinating diseases.

Biologic therapy

- Golimumab (Simponi[®]) 50 mg s.c. monthly
 Certolizumab pegol (Cimzia[®]) 200 mg s.c. weeks 0, 2 & 4, then 200 mg every 2 weeks or 400 mg monthly.
 Ustekinumab (Stelara[®]) (anti-IL-12 & anti-IL-23 antibodies) 45 mg for pts with body weight <100 kg and 90 mg to them >100 kg administered in weeks 0 & 4 and then every 12 weeks. For severe psoriasis.
- *Note*: Anti-TNF treatment is required for patients with active disease (at least 3 tender and swollen joints) who have failed at least two DMARDs. Level of evidence A.

Therapeutic Landscape for PsA Is Expanding



Abatacept, a selective co-stimulation modulator, has been approved by FDA, not yet by EMA

GRAPPA Treatment Guidelines for Psoriatic Arthritis



Kavanaugh A et al. J Rheumatol 2006;33:1417-21 (with permission)

Surgical treatment

In the early stages - synovectomy, interventions for carpal canal syndrome, tendon ruptures, atlantoaxial subluxation, ruptured Baker cyst. • At late stages - arthroplasty with total

prosthesis of the respective joint.

Balneophysical treatment

Kineto- and hydrotherapy, which can:

- relieves pain
- oreduce inflammation
- In the strengthen muscles
- oprevents osteoporosis
- oprevents muscle atrophy
- Specific exercises: wide breathing, extension mobilization of the spinal column (to avoid spinal kyphosis),
- movements in all planes.
- Swimming is recommended sports.

Poor prognostic factors

- Peripheral polyarticular involvement (arthritis mutilans, symmetrical arthritis)
- In High inflammatory biological syndrome at onset
- Genetic factors:
 - Family history of PsA;
 - HLA-B27, B39 in presence of HLA-DR7 \rightarrow SEVER!
 - HLA-DQw3, B22 in presence of HLA-DR7 \rightarrow GOOD!
- Patients with severe forms
- Young age at the onset of arthritis
- Female
- Acute onset of arthritis
- Extensive skin damage
- Lack of response to NSAIDs
- Association with HIV/AIDS

Mulţumesc pentru atenţie!

