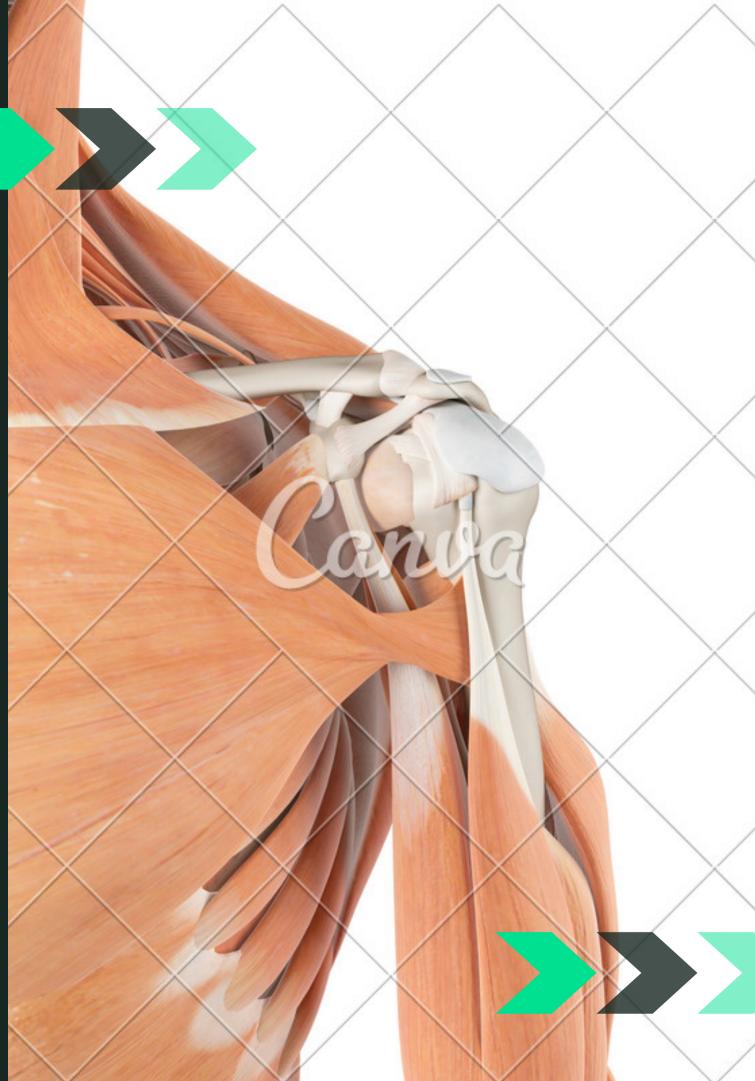
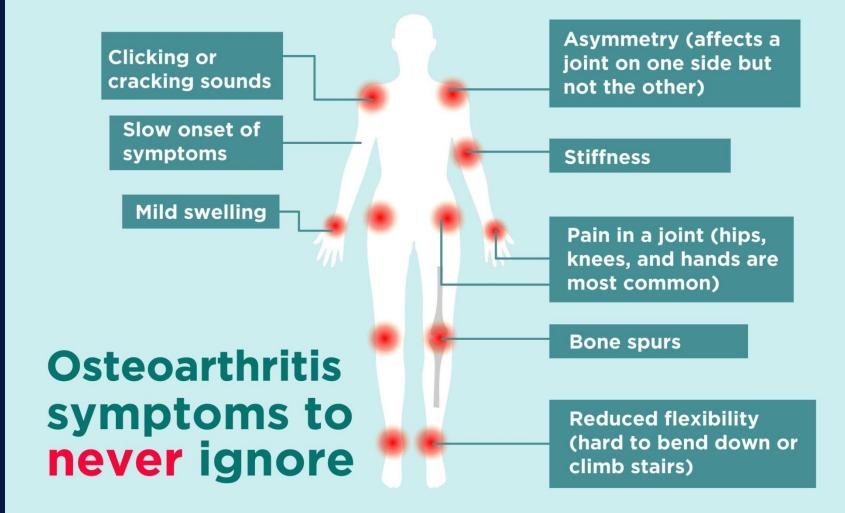
PASSION ACADEMIC TEAM **JU - MEDICINE** MUSCULOSKELETAL SYSTEM **Sheet#4 - Pharmacology** Lec. Title : Pharmacotherapy of Osteoarthritis Written By: Rahma Marie Abdallah AL-Qashi If you come by any mistake, please kindly report it to

shaghafbatch@gmail.com



# Pharmacotherapy of osteoarthritis

#### Dr. Romany Helmy Thabet, PhD



#### Distinction between rheumatoid arthritis and osteoarthritis

Feature	Rheumatoid arthritis	Osteoarthritis	
Primary joints affected	Metacarpophalangeal	Distal interphalangeal	
	Proximal interphalangeal	Carpometacarpal	
Heberden's nodes	Absent	Frequently	
Joint characteristics	Soft, warm, and tender	Hard and bony	
Stiffness	Worse after resting (eg, morning stiffness)	If present, worse after effort, may be described as evening stiffness	
Laboratory findings	Positive rheumatoid factor	Rheumatoid factor negative	
	Positive anti-CCP antibody	Anti-CCP antibody negative	
	Elevated ESR and C reactive protein	Normal ESR and C reactive protein	

CCP: cyclic citrullinated peptide

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Osteoarthritis must be differentiated from rheumatoid arthritis.

• Wear and tear of joists due to cartilage meaning down. This leads to bones rubbing against each other and cracking joints.

• Osteoarthritis mostly presents in females , obese. Focuses on large joints.

Rheumatoid : symmetrical

Osteoarthritis : asymmetrical

In hands, distal interphalangeal joints are NOT involved in rheumatoid but are in osteoarthritis.

movement worsens pain in osteoarthritis = night stiffness

Movement eases pain in rhomboid = morning stiffness

Soft joints = rheumatoid

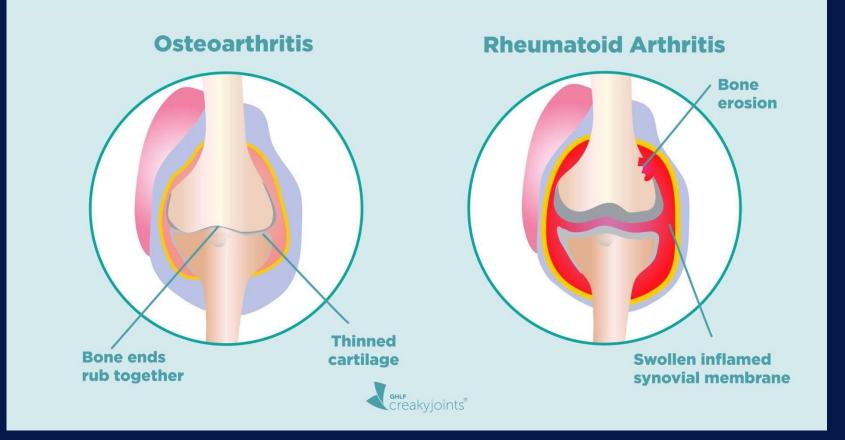
Hand points = osteoarthritis (due to bone on bone)

Check immune system to check it immune system has attacked the joints.

If yes = rheumatoid

If no = osteoarthritis

#### Osteoarthritis vs. Rheumatoid Arthritis in the Joint



→ Rheumatoid : synovial membrane undergoes hypertrophy leading to erosion.
→ Osteoarthritis : synovial membrane doesn't undergo hypertrophy.

# Management of OA

- Establish the diagnosis of OA on the basis of history and physical and x-ray examinations
- Decrease pain to increase function
- Prescribe progressive exercise to
  - Increase function
  - Increase endurance and strength
  - Reduce fall risk
- Patient education: Self-Help Course
  - Weight loss
  - Heat/cold modalities

The pain in o osteoarthritis is what needs to be addressed.

We deal with it using analgesics + nonpharmacologic therapy.

### Pharmacologic Management of OA

- Nonopioid analgesics
- Topical agents
- Intra-articular agents
- Opioid analgesics
- NSAIDs
- Unconventional therapies



Nonpharmacologic Therapy : 1. Exercise (low pressure like walking, swimming)

2. Cold packs for acute joint pain ( = vasoconstriction )

3. Warm packs for chronic joint pain

Irritant on irritated joint = no pain ( counter irritant topical creams )

Worn away cartilage (hyaluronic acid) can be replaced with injections that will act as nutrition for the cartilage. Not long-lasting. Needs frequent treatment.

There is no permanent treatment. These are only symptomatically treatment.

#### Nonopioid Analgesic Therapy

- First-line—Acetaminophen
  - Pain relief comparable to NSAIDs, less toxicity
  - Beware of toxicity from use of multiple acetaminophen-containing products
  - Maximum safe dose = 4 grams/day

Paracetamol is the first drug we use. Has no anti-inflammatory, nonsteroidal, non opioid.



First Line : paracetamol (acetaminophen) For Example : Panadol It works by inhibiting cox 2 (central) Might also inhibit cox 3 Has no peripheral effects in therapeutic doses. Has no anti-inflammatory effects Analgesic alone No over 4mg a day.

## Nonopioid Analgesic Therapy (cont'd)

#### NSAIDs

- Use generic NSAIDs first
- If no response to one may respond to another
- Lower doses may be effective
- Do not retard disease progression
- Gastroprotection increases expense
- Side effects: GI, renal, worsening CHF, edema
- Antiplatelet effects may be hazardous



Second Line : conventional NSAIDS.

- A. Ibuprofen
- B. naproxen
- They are non-selective.

They are harmful for GI, kidney, blood (anti platelet effect) so, we start at lower doses.

Does not retard disease progression!!!

### Nonopioid Analgesics in OA

- Cyclooxygenase-2 (COX-2) inhibitors
  - Pain relief equivalent to older NSAIDs
  - Probably less GI toxicity
  - No effect on platelet aggregation or bleeding time
  - Side effects: Renal, edema
  - Older populations with multiple medical problems not tested
  - Cost similar to generic NSAIDs plus proton pump inhibitor or misoprostol



Third Line : Cox-2 inhibitors

Different from paracetamols due to lower GIT toxicity so no need for proton pumps.

can harm kidneys (cox-2 inhibitors inhibit the cox-2 in kidneys)

# Nonopioid Analgesics in OA (cont'd)

#### Tramadol

- Affects opioid and serotonin pathways
- Nonulcerogenic
- May be added to NSAIDs, acetaminophen
- Side effects: Nausea, vomiting, lowered seizure threshold, rash, constipation, drowsiness, dizziness

# Sheet# 6

Fourth Line : Opioid analgesics

- 1. Tramadol is a weak opioid analgesic
- can be added with paracetamol
- has a lower seizure threshold

can cause constipation and fatigue, dizziness and depression only in very severe pain.

- 2. codeine and oxycodone are also weak opioid analgesics.
- 3. propoxyphene, a morphine
- 4. morphine and fentanyl patches (under the clavicle)

morphine and fentanyl are only given in cases of pain that stops daily activities.

# **Opioid Analgesics for OA**

- Codeine, oxycodone
  - Anticipate and prevent constipation
  - Long-acting oxycodone may have fewer CNS side effects
- Propoxyphene
- Morphine and fentanyl patches for severe pain interfering with daily activity and sleep

#### **Topical Agents for Analgesia in OA**

- Local cold or heat: Hot packs, hydrotherapy
- Capsaicin-containing topicals
  - Use well supported by evidence
  - Use daily for up to 2 weeks before benefit
  - Compliance poor without full instruction
  - Avoid contact with eyes
- Liniments = methyl salicylates
  - Temporary benefit

Capsaicin is an irritant (leads to counter irritant). Daily use. two weeks to work.

# **OA: Intra-articular Therapy**

- Intra-articular steroids
  - Good pain relief
  - Most often used in knees, up to q 3 mo
  - With frequent injections, risk infection, worsening diabetes, or CHF
- Joint lavage
  - Significant symptomatic benefit demonstrated

- Hyaluronate injections\*
  - Symptomatic relief
  - Improved function
  - Expensive
  - Require series of injections
  - No evidence of longterm benefit
  - Limited to knees



Intra- articular Therapy :

steroid injections can cause cartilage damage so only use it 3 times/year maximum. only after severe flair of disease. Can be systematically absorbed. no evidence of long-term benefits. needs frequent injections.

Hyaluronic injections can only be injected into the knees.

Joint lavage : doctor who cleans the bone fragments from the cartilage

# **OA: Unconventional Therapies**

- Polysulfated glycosaminoglycans—nutriceuticals
  - Glucosamine +/- chondroitin sulfate: Symptomatic benefit, no known side effects, long-term controlled trials pending
- Tetracyclines as protease/cytokine inhibitors
  - Under study
  - Have disease-modifying potential

Nutraceuticals : nutritional injections for the cartilage. For Example : glucosamine + Chondroitin sulfate. No proof of being 100% curative.

## OA: Management Summary

- First: Be sure the pain is joint related (not a tendonitis or bursitis adjacent to joint)
- Initial treatment
  - Muscle strengthening exercises and reconditioning walking program
  - Weight loss
  - Acetaminophen first
  - Local heat/cold and topical agents

# OA: Management Summary (cont'd)

- Second-line approach
  - NSAIDs if acetaminophen fails
  - Intra-articular agents or lavage
  - Opioids
- Third-line
  - Arthroscopy
  - Osteotomy
  - Total joint replacement

→ Osteotomy : they remove a piece of bone from the top or bottom bone to align the joints and relieve the pressure off destroyed areas.

			Discuss total joint replacement for osteoarthritis of the hip, knee, or shoulder if steps below are unsuccessful	
			Consider hyaluronic acid injection for persistent knee osteoarthritis	
			er corticosteroid injection for acute exacerbation e osteoarthritis	
		Consider opioid for dependence	therapy, but monitor carefully and abuse	
	Add combination glucosamine and chondroitin for moderate to severe knee osteoarthritis; discontinue if no change after three months, but continue if effect is noted			
	herapy, beginning with over-the re; use generics if possible	-counter ibuprofe	n or naproxen; switch to different NSAID if initial choice	
Begin with acetaminophen and continue if still effective, or step up to NSAID				
	rroughout treatment and encour erral for supervised exercise (land			
Mild osteoarthritis	Moderate osteoarthritis		Severe osteoarthritis	

