

# Osteoarthritis of the Knee

A PRIMARY CARE APPROACH

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# Disclosures

- ▶ Nothing to disclose

# Objectives

1. Describe the pathology of OA
2. Recognize exam findings consistent with diagnosis of OA
3. Describe non-surgical options for OA treatment

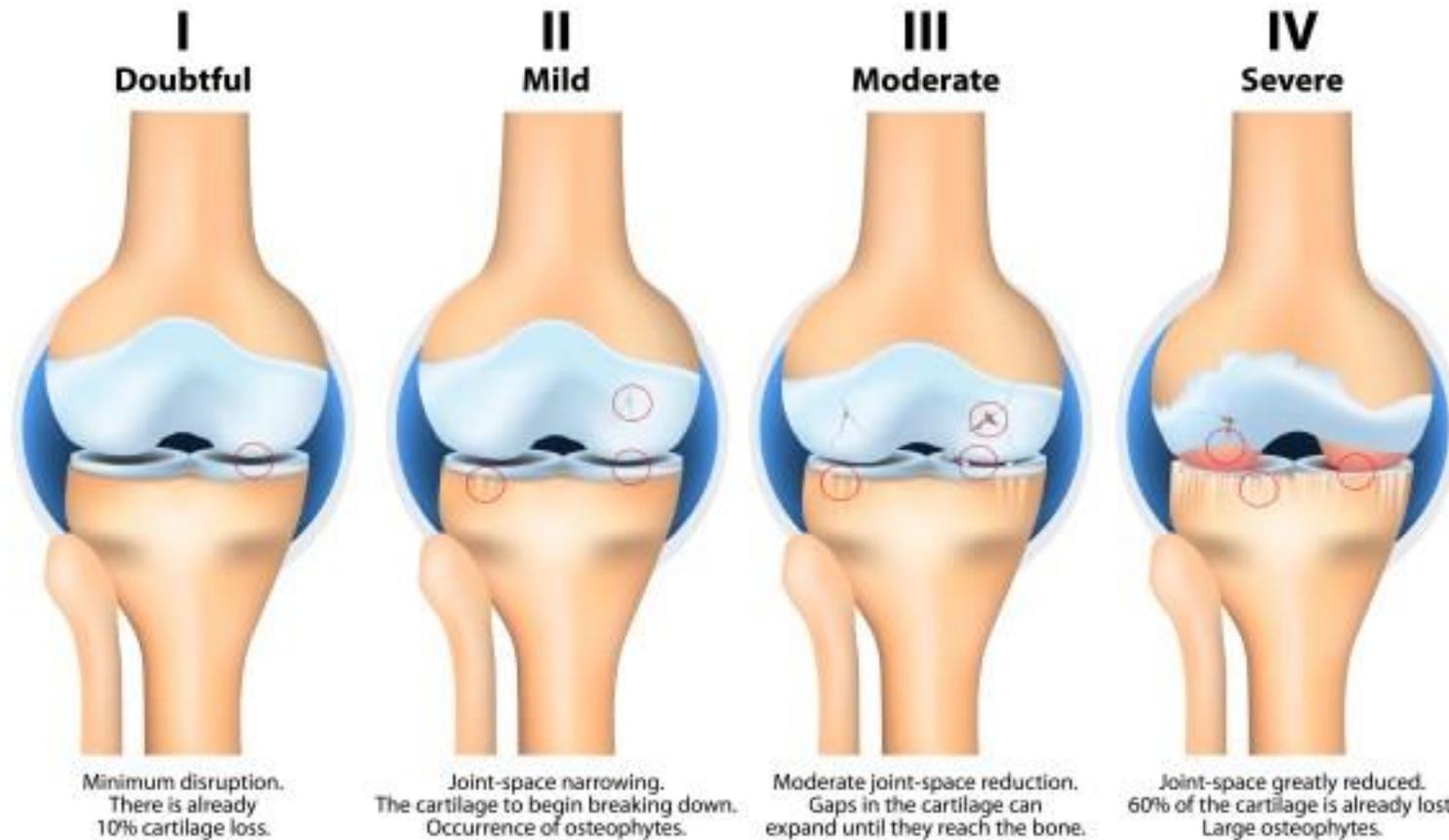
# Pathology

- ▶ Articular cartilage erosion
- ▶ Hypertrophy of bone margins (osteophytes)
- ▶ Subchondral sclerosis

# Pathology

- ▶ Osteoarthritis in name implies inflammatory disease
- ▶ Inflammatory cells may be present
- ▶ Considered intrinsic disease of cartilage
  - ▶ Biochemical and metabolic alterations occur resulting in breakdown
  - ▶ Age related changes in proteoglycans and collagen, decreasing tensile strength, shorten fatigue life
- ▶ Chondrocytes produce IL-1 and TNF- $\alpha$ 
  - ▶ Stimulate catabolic metalloproteinases
  - ▶ Inhibit synthesis of type 2 collagen and proteoglycans

## STAGE OF KNEE OSTEOARTHRITIS



Acute

OA, 1 grade

OA, 2 grade

OA, 3 grade

OA, 4 grade



# Epidemiology

- ▶ Affects >33% of adults 65 and older
- ▶ Nearly 27 million person in US affected
- ▶ Risk Factors:
  - ▶ Female gender
  - ▶ Age over 50
  - ▶ Obesity
  - ▶ History of joint injury
  - ▶ Family history
  - ▶ Occupation involving squatting, kneeling
  - ▶ Sport participation with repetitive impact (soccer, football)
  - ▶ Inflammatory joint disease

# Prevalence and limitations

Age (years)	Prevalence of physician-diagnosed OA (%)	Prevalence of activity limitation due to OA (%)
18 to 44	7.3	2.7
45 to 64	30	13
65 and older	50	22

# Clinical signs and symptoms

- ▶ Most common- pain worse with activity
- ▶ Stiffness up to 30 minutes
- ▶ Crepitus
- ▶ Swelling
- ▶ Limping
- ▶ Limited range of motion

# Diagnosis of OA-Clinical criteria

- ▶ Age over 50
- ▶ Bony enlargement
- ▶ Bony tenderness
- ▶ Crepitus
- ▶ Stiffness lasting less than 30 minutes
- ▶ No palpable warmth

# Diagnosis of OA-Labs and Imaging

- ▶ Laboratory Criteria
  - ▶ ESR <40 mm/hr
  - ▶ RF < 1:40
  - ▶ Synovial fluid: clear, viscous, WBC <2,000  $\mu$ L
- ▶ Radiographic Criteria
  - ▶ Presence of osteophytes

# Radiographs and OA

- ▶ Not required to diagnose OA in patients with risk factors and typical symptoms
- ▶ Do not always correlate well with symptoms
- ▶ One study showed 16% of patients with frequent pain had evidence of OA
- ▶ Another showed 21% of patients who met radiographic criteria of OA had frequent pain
- ▶ Typical findings
  - ▶ Osteophytes
  - ▶ Joint space narrowing
  - ▶ Subchondral sclerosis

# Radiographs and OA

- ▶ Radiographs can help determine severity for surgical planning
- ▶ MRI can detect joint abnormalities in 90% of patients 50+ y/o, even without joint pain



# Diagnostic Accuracy

Criteria	Sensitivity %	Specificity %
Knee pain +3 clinical criteria	95	69
Knee pain +5 clinical or laboratory criteria	92	75
Knee pain +5 clinical or laboratory criteria + osteophytes	91	86

# Treatments

- ▶ Physical Modalities and Exercise
- ▶ Pharmacological therapies
- ▶ Complementary and Alternative therapies
- ▶ Intraarticular injections
- ▶ Surgery

# Physical modalities and Exercise

- ▶ Physical therapy
- ▶ Exercise
- ▶ Weight loss
- ▶ Braces or shoe inserts

# Review of PT Interventions

- ▶ Exercise and weight loss reduce pain, improve function
- ▶ High and low-intensity exercise equal in improving pain, gait, function and aerobic capacity
- ▶ Water and land-based therapy reduce knee pain and physical disability
  - ▶ Aquatic therapy has small short-term benefits
- ▶ Small RCT shows Tai Chi 3x/week for 12 weeks effective for women > 50
- ▶ Insufficient evidence to support yoga

# Review of PT interventions

- ▶ 54 studies of patients with mild to moderate symptomatic OA
- ▶ Land-based exercise therapy included strengthening, functional training and aerobic conditioning
  - ▶ Exercise moderately reduced pain
  - ▶ Moderately improved physical functioning
  - ▶ Slightly improved quality of life
- ▶ Improvement sustained 2-6 months posttreatment for pain and physical function

# Strengthening therapy

- ▶ Quadriceps strengthening, lower limb strengthening, combined strengthening, walking programs
  - ▶ Improve pain and physical functioning
  - ▶ Combined strength and aerobic exercise programs reduced pain and improve function in the medium-term, reduce pain in the long term based on 4 RCT
- ▶ Home-based exercise and self-management program
  - ▶ Combine strength, agility, pain-coping
  - ▶ Beneficial effects in the short and medium-terms

# Physical Therapy

- ▶ American College of Rheumatology strongly recommends all patients with symptomatic OA be enrolled in exercise program that matches their ability to participate in required activities
  - ▶ No preference with aquatic vs. land-based therapy
- ▶ American Academy of Orthopedic Surgeons 2013 practice guidelines
  - ▶ Self-management program
  - ▶ Low-impact aerobic exercise
  - ▶ Weight loss if BMI > 25 kg/m<sup>2</sup>

# Physical Modalities

- ▶ Use of lateral heel wedge may reduce use of NSAIDs
  - ▶ Evidence for lateral wedge ineffective for medial knee OA
- ▶ Shoe inserts do not show any benefit on pain or function based on 8 RCTs
- ▶ Knee bracing has insufficient evidence to recommend use
- ▶ Knee taping may provide some short-term benefit

# Physical modalities

- ▶ TENS unit reduces pain in the short term
- ▶ Does not improve function or pain after 12 weeks
- ▶ Compared to sham controls



# Physical modalities

- ▶ Beneficial interventions for long-term (>26 weeks) benefit
  - ▶ Weight loss
  - ▶ Agility training
  - ▶ Combined exercise programs
  - ▶ Manual therapy (massage, acupressure, self-massage)

# Pharmacologic Treatments

- ▶ Topical
- ▶ Oral
- ▶ Intra-articular

# Topical Treatments

- ▶ Advantage of topical is avoidance of systemic adverse effects
- ▶ Review of placebo controlled trials of capsaicin cream 0.025%
  - ▶ Statistically more effective than placebo, less effective than topical NSAIDs
  - ▶ Associated with transient burning sensation
- ▶ Meta-analysis of RCTs comparing topical NSAIDs to placebo, oral NSAIDs
  - ▶ Topical NSAIDs superior to placebo, but only in first 2 weeks of treatment
  - ▶ Topical NSAIDs less effective than oral NSAIDs

# Oral Treatments

- ▶ Acetaminophen preferred in American College of Rheumatology guidelines
  - ▶ More effective than placebo
  - ▶ Low risk of liver toxicity
  - ▶ Use with caution in patients who consume alcohol daily
  - ▶ Regular kidney and liver monitoring in patients taking 3-4 g per day
- ▶ NSAIDs superior to Acetaminophen in moderate to severe pain
  - ▶ Diclofenac 150mg daily likely to be most effective, followed by Naproxen

# Oral Treatments

- ▶ Cyclooxygenase-2 inhibitors
  - ▶ Celebrex (celecoxib) only prototype available
  - ▶ Associated with increased risk of stroke and MI
  - ▶ Systematic review shows similar symptoms control as NSAIDs; does not reduce risk of serious GI adverse effects
  - ▶ Overall, no increased risk with COX-2 compared to NSAIDs
  - ▶ May not provide much benefit over NSAIDs

# Oral Treatments

- ▶ Duloxetine (Cymbalta)
  - ▶ Selective norepinephrine reuptake inhibitor
  - ▶ Approved for treatment of painful conditions
  - ▶ Most common side effect is nausea
  - ▶ NNT=7 for clinically significant pain reduction
  - ▶ NNH=6

# Oral Treatments

- ▶ Opioids
  - ▶ American College of Rheumatology support opioids when other treatments ineffective
  - ▶ American Geriatric Society recommends considering opioids in older individuals with moderate to severe pain, decreased function, diminished quality of life
  - ▶ Risks of NSAID use in older patients may exceed risk of addiction to opioids

# Opioids

- ▶ Tramadol (Ultram)
  - ▶ Decreases pain intensity
  - ▶ Relieves symptoms
  - ▶ Improves function
  - ▶ Increased seizure risk, especially with alcohol
  - ▶ 11 RCT report moderate improvement with treatment
  - ▶ NNT=6 for one person to report moderate improvement
  - ▶ NNH=8 for one person to stop taking it
- ▶ Because of similar harms and benefits to duloxetine, use only in selected patients

# Opioids

- ▶ Other oral and transdermal opioids have modest benefits that are of questionable clinical significance
- ▶ Due to significant adverse effects, long-term use is discouraged
- ▶ Daily dose recommendations
  - ▶ 50 mg or less of hydrocodone per day
  - ▶ 30 mg or less of oxycodone per day

# Complementary and Alternative Treatments

- ▶ Acupuncture
  - ▶ Benefit is not clear
  - ▶ Meta-analysis did not demonstrate clinically relevant improvement in pain or function score comparing acupuncture to sham acupuncture
  - ▶ Short-term (6 weeks) or long-term (6 months) acupuncture or sham acupuncture resulted in patients “feeling better” compared to usual treatment
  - ▶ 1 study of 6 months of traditional Chinese acupuncture
    - ▶ Decreased pain and increased functionality by 40 points on average

# Complementary and Alternative Treatments

- ▶ Glucosamine and Chondroitin
  - ▶ Marketed since the 1990's as disease modifying agents
  - ▶ 3 moderate-strength RCT of glucosamine with chondroitin
    - ▶ Little benefit with mild OA
    - ▶ Greater benefit in those with moderate to severe pain
    - ▶ Can improve medium-term outcomes (<26 weeks) but no benefit beyond
  - ▶ No evidence they slow cartilage loss
  - ▶ Glucosamine alone is safe, benefit is variable
  - ▶ Chondroitin alone does not decrease pain from hip or knee OA
  - ▶ Not regulated by FDA

# Complementary and Alternative Treatments

- ▶ S-adenosylmethionine (SAM-e)
  - ▶ Meta-analysis of RCT found SAM-e as effective as NSAIDs in reducing pain, disability
  - ▶ Better safety profile
- ▶ Ginger
  - ▶ 255 mg of ginger twice daily
  - ▶ Reduction in pain 63% compared to 50% with placebo
- ▶ Turmeric
  - ▶ Historically used for pain
  - ▶ No RCT

# Intra-Articular Injections

- ▶ Corticosteroid injections
  - ▶ Short-term symptomatic relief, do not improve overall quality of life
  - ▶ Low risk of adverse effects
  - ▶ 28 clinical trials show significant short-term pain reduction, improvement in patient self-assessment
  - ▶ Good evidence for long-term benefit lacking

# Corticosteroids

- ▶ Exact mechanism in joint is unknown
- ▶ Presumed to
  - ▶ Inhibit accumulation of inflammatory cell lines
  - ▶ Reduce prostaglandin synthesis
  - ▶ Inhibit leukocyte secretion from synovial cells
  - ▶ Decrease interleukin secretion by synovium



# Intra-Articular Injections

- ▶ Hyaluronic acid injections
  - ▶ Hyaluronic acid produced by synovium
  - ▶ Decreased in persons with OA
  - ▶ Exogenous supplementation thought to support and restore elastoviscous properties of the joint
  - ▶ No study shows these drugs alter disease course
  - ▶ Meta-analysis of studies show poor study design or were industry sponsored
  - ▶ No demonstrable clinical improvement in function

# Intra-articular injections

- ▶ Platelet-rich plasma (PRP)
  - ▶ Volume of autologous plasma with platelet concentration above average
  - ▶ Soft tissue healing enhancement occurs with concentrations  $>1,000,000$  platelets/ $\mu\text{L}$
  - ▶ Based on low strength of evidence from 4 RCTs
    - ▶ Reduce pain on the medium term (12-26 weeks) compared to placebo
    - ▶ Improves quality of life
  - ▶ No evidence to support soft tissue healing in OA
  - ▶ Associated with pain and stiffness in 44% of patients who received 2 injections in 1 study

# Surgical Treatments

- ▶ Arthroscopic surgery
  - ▶ Not appropriate unless loose bodies, mechanical symptoms
  - ▶ 2 well-designed RCT showed no benefit on OA
  - ▶ Not effective for patients with degenerative meniscal tears
- ▶ Total knee replacement
  - ▶ Last resort
  - ▶ AAOS- main indication for relief of pain when non-surgical treatments fail
  - ▶ Studies show improvement in pain and function with moderate OA compared to usual care
  - ▶ Outcomes from surgery similar between obese and non-obese

# Conclusion

Clinical recommendation	Evidence Rating
Radiography can confirm the diagnosis of OA and may be helpful before surgical referral, but findings tend NOT to correlate with symptoms	C
Exercise, PT, knee taping and tai chi are all beneficial for OA	B
Ineffective treatments for OA include braces, special shoes, lateral wedges for medial OA, hyaluronic acid injections	B
Medical therapy for OA should begin with full-strength acetaminophen and topical therapy, then proceed to NSAIDs and selectively to tramadol and other opioids. NSAIDs and opioids may reduce pain and improve function, but have significant potential harms	A
Joint replacement should be considered for patients with moderate to severe pain and radiographically confirmed OA	A
Corticosteroid injections may be helpful in the short-term, but evidence is mixed	B

# References

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