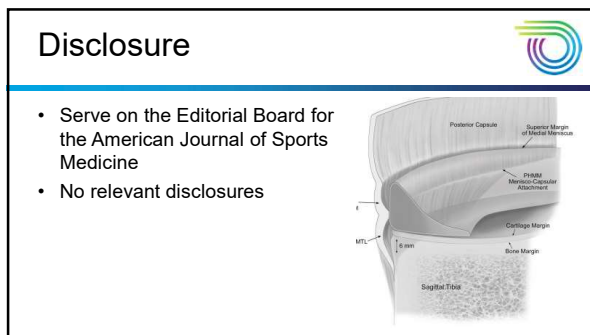




1



2



3

Learning Objectives



- Review boney and cartilaginous knee anatomy
- Discuss treatment options for meniscus injuries
- Review evidence of knee injectables

4

Outline



- The Meniscus
- Meniscus Tear Types
- Evaluation of Knee Pain
- Treatments Options
- Injections

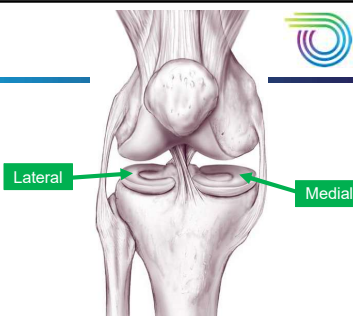


5

Anatomy



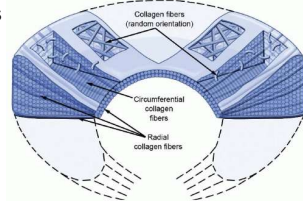
- Knee Joint
 - 4 bones
 - 3 "articulations"
 - Cartilage surface
 - Ligaments
 - Meniscus



6

The Meniscus

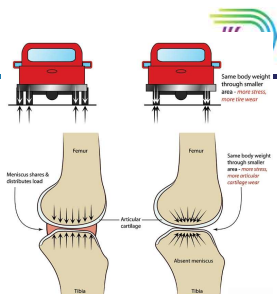
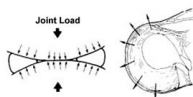
- Type I collagen fibers
 - Circumferential fibers
 - Radial fibers



7

The Meniscus

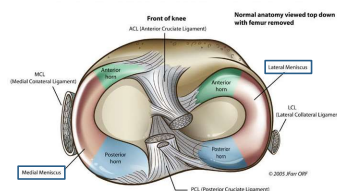
- Function
 - Dissipate forces across the knee joint
 - Shock absorption
 - Increased congruency



8

The Meniscus

- Medial meniscus
 - C-shaped
 - Less mobile
- Lateral meniscus
 - Circular
 - More mobile
 - Covers more of the cartilage surface



9

The Meniscus

- Attachment sites
 - Coronary ligaments peripherally
 - Root attachments

10

The Meniscus

- Attachment sites
 - Coronary ligaments peripherally
 - Root attachments
- Nutrition/Blood supply
 - Only the peripheral 1/3 is well vascularized
 - Inner 2/3 – diffusion

11

Meniscus Tear Types

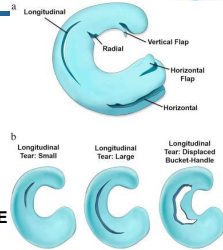
- Tear types
 - Radial tear
 - Oblique/Flap tear
 - Peripheral longitudinal
 - Bucket-handle tear
 - Horizontal cleavage
 - Root tear

12

Meniscus Tear Types

• Tear types

- Radial tear
- Oblique/Flap tear
- Peripheral longitudinal
- Bucket-handle tear
- Horizontal cleavage
- Root tear
- **ANY COMBINATION OF THE ABOVE**



13

Evaluation of Knee Pain

• Presentation

- When?
- Associated with an acute event?
- Where is the pain?
- Aggravating factors?
- Associated symptoms?



14

Evaluation of Knee Pain

• Physical examination

- Evaluate for effusion
 - Presence of an effusion should raise clinical concern of acute process



• 3 possibilities

- Acute injury (ligament tear, loose body, meniscus tear)
- Inflammatory process (crystalline arthropathy, etc)
- Infectious process (septic arthritis)

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Evaluation of Knee Pain




- Physical examination
 - Evaluate for effusion
 - Check range of motion



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Evaluation of Knee Pain


- Physical examination
 - Evaluate for effusion
 - Check range of motion
 - Assess for stability
 - ACL and PCL
 - MCL and LCL

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Evaluation of Knee Pain

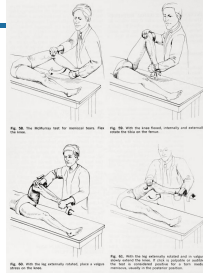
- Physical examination
 - Evaluate for effusion
 - Check range of motion
 - Assess for stability
 - Evaluate the meniscus
 - Joint line tenderness



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Evaluation of Knee Pain

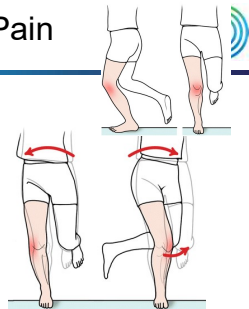
- Physical examination
 - Evaluate for effusion
 - Check range of motion
 - Assess for stability
 - Evaluate the meniscus
 - Joint line tenderness
 - McMurray's test



19

Evaluation of Knee Pain

- Physical examination
 - Evaluate for effusion
 - Check range of motion
 - Assess for stability
 - Evaluate the meniscus
 - Joint line tenderness
 - McMurray's test
 - Thessaly's test



20

Evaluation of Knee Pain

- Physical examination
 - Evaluate for effusion
 - Check range of motion
 - Assess for stability
 - Evaluate the meniscus
 - Joint line tenderness
 - McMurray's test
 - Thessaly's test
 - Apley's compression



FIG. 6
The grinding test

21

Treatment

- Conservative treatment
 - More to come
- Operative treatment
 - Acute injury
 - Presence of an effusion
 - Experiencing mechanical symptoms
 - Have failed appropriate conservative treatment measures

22

Treatment

- Meniscal repair
 - Best chances for successful repair
 - Acute injury
 - Young patient (better blood supply)
 - Relatively simple tear
 - Location close to blood supply (red-red zone)
 - BMI with certain tear types

When all these conditions are met healing rates can reach 80-90%

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Treatment

- Meniscal repair
 - CM 17 yo M, acute football injury
 - Displaced bucket-handle tear lateral meniscus

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Treatment



• Partial Meniscectomy

• Indications

- Acute injury AND/OR
- Significant mechanical symptoms
- Irreparable meniscus tear
- Failure of conservative management

- The most common type of knee surgery performed

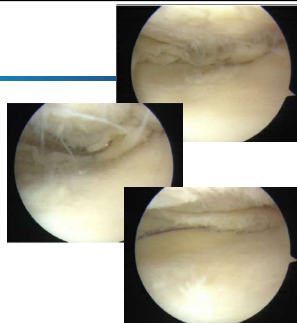


25

Treatment

• Partial Meniscectomy

- VF 56 yo F, gradual onset of L knee pain, ↑ BMI
 - Mild-mod knee OA
 - Irreparable complex tear type
 - Failed 6 weeks prescription NSAID, PT, and corticosteroid injection



26

Treatment



• Is partial meniscectomy 1st-line treatment?

- 2013 FIDELITY trial
- Multicenter, RCT, double-blind
- 146 patients w/ degenerative meniscus tears randomized
 - 70 partial meniscectomy
 - 76 sham surgery

ORIGINAL ARTICLE

Arthroscopic Partial Meniscectomy versus Sham Surgery for a Degenerative Meniscal Tear

Raine Sihvonen, M.D., Mikko Pajuoja, M.D., Ph.D., Antti Mäkinen, M.D., Ph.D., Ari Heliö, M.D., Ph.D., Antti Jousimäki, M.D., Ph.D., Heikki Nurmio, M.D., Jukka Kallio, M.D., and Tero L. Järvenpää, M.D., Ph.D., for the Finnish Degenerative Meniscal Lesion Study (FIDELITY) Group

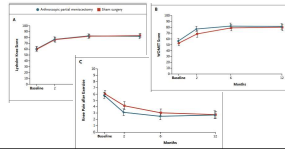
For the sham surgery, a standard arthroscopic partial meniscectomy was simulated. To mimic the sensation and sounds of a true arthroscopic partial meniscectomy, the surgeon asked for all instruments, manipulated the knee as if an arthroscopic partial meniscectomy was being performed, pushed a mechanized shaver (without the blade firmly against the patella [outside the knee], and used suction. The patient was also kept in the operating room for the amount of time required to perform an actual arthroscopic partial meniscectomy.

27

Treatment

- Is partial meniscectomy 1st-line treatment?
- 2013 FIDELITY trial
- Multicenter, RCT, double-blind
- 146 patients w/ degenerative meniscus tears randomized
 - 70 partial meniscectomy
 - 76 sham surgery
- No significant difference in outcomes

ORIGINAL ARTICLE
Arthroscopic Partial Meniscectomy versus Sham Surgery for a Degenerative Meniscal Tear
Raine Sihvonen, M.D., Mika Payola, M.D., Ph.D., Antti Malmivaara, M.D., Ph.D., Ari Heliö, M.D., Ph.D., Antti Jouskainen, M.D., Ph.D., Heikki Huuhtaniemi, M.D., Jukka Kallala, M.D., and Teppo L. N. Järvenpää, M.D., Ph.D., for the Finnish Degenerative Meniscal Lesion Study (FIDELITY) Group



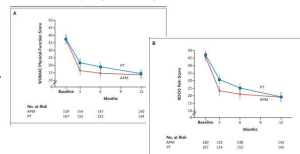
28

Treatment

- Is partial meniscectomy 1st-line treatment?
- 2013 METEOR trial
- Multicenter, RCT
- 351 patients w/ degenerative meniscus tears randomized
 - 174 partial meniscectomy w/ PT
 - 177 physical therapy
- No significant difference

Surgery versus Physical Therapy for a Meniscal Tear and Osteoarthritis

Jeffrey N. Katz, M.D., Robert N. Roodyn, M.D., Christine E. Chiu, M.P.H., Leigh de Chazeau, P.T., D.C., Bruce J. Cole, M.D., M.Sc., Peter J. Doherty, M.D., David A. D'Amato, M.D., M.P.H., M.Sc., Ph.D., Kenneth E. Fong, M.A., Morgan H. Jones, M.D., M.P.H., David A. Lenz, M.D., Lisa A. Mandel, M.D., M.P.H., David B. Reame, M.D., Robert C. Stein, M.D., Anthony M. Stone, M.D., William J. Murray, M.D., Joseph P. McNamee, M.P.H., Emily A. Reade, Ph.D., Robert E. Richardson, P.T., M.S., D.C., L.C.C., Benjamin B. Stone, M.D., David E. Lohrke, P.T., M.D., D.C., L.C.C., Jeffrey J. Chiu, M.D., M.P.H., David H. Solomon, M.D., M.P.H., Matthew J. Smith, M.D., Amy T. Spindler, M.D., M.P.H., Stuart M.D., John Wright, M.D., Rick W. Wright, M.D., and Glen Levine, Ph.D.



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Treatment

- Conservative Treatment
- Don't meet indications for meniscus repair
 - No effusion (chronic issue?)
 - Older patients (decreased blood supply)
 - Degenerative/complex tear types
 - No mechanical symptoms
 - Mainly pain-based complaints

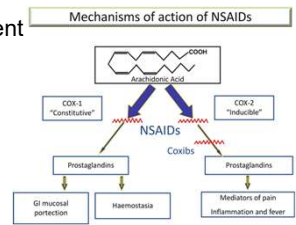


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Treatment

• Conservative Treatment

- Oral analgesics
 - Acetaminophen
 - NSAIDs
 - COX-2 inhibitors



31

Treatment

• Conservative Treatment

- Oral analgesics
- Physical therapy/home-exercises
 - Mechanics/biomechanics minimize loading
 - Aquatic therapy



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Treatment

• Conservative Treatment

- Oral analgesics
- Physical therapy/home-exercises
- Injections
 - Corticosteroid
 - Viscosupplementation
 - Orthobiologics



33

Treatment



- When to perform an injection?
 - When patients are not improving with other appropriate conservative treatment options (i.e. oral medications, PT/HEP, etc)
 - AND/OR
 - At an appropriate position on the "Misery Scale"



34

Treatment



- Misery Scale

No Injection

Injection

Mild Issue

- Mild pain
- Intermittent symptoms
- Not affecting ADLs
- Haven't tried any other treatment modalities

Severe Issue

- Severe pain
- Constant symptoms
- Affecting ADLs and QOL
- Have already tried other treatment modalities

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Intraarticular Knee Injection



- Site preparation and technique
 - No consensus
 - Skin antiseptics
 - Sterile vs non-sterile gloves
 - Anesthetic spray/injection
 - Sterile field



36

What to Inject?



• Corticosteroids

- No **strong** evidence to suggest one corticosteroid is better than another
- JAAOS 2009 Systematic Review suggested triamcinolone was most effective
- American College of Rheumatology study noted providers chose medication based on "availability" or out of "habit"

Table 1
Common Injectable Corticosteroids, Concentrations, and Dosage

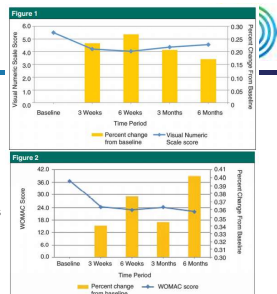
Injectable Corticosteroids	Concentration (mg/mL)	Common Dose for Large Joint (mg)
Betamethasone (often mixture of betamethasone acetate and betamethasone sodium phosphate)	6	6-12
Methylprednisolone acetate	20, 40, or 80	20-80
Triamcinolone acetonide	10 or 40	10-40
Triamcinolone diacetate	40	20-40
Triamcinolone hexacetonide	20	10-20
Dexamethasone sodium phosphate	4 or 8	2-4

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What to Inject?

• Corticosteroids

- Fairly poorly studied compared to other tx modalities
- Matzkin et al 2017 – 100 pts
 - Triamcinolone
 - Statistically sig improvements in Visual Numeric Pain and total WOMAC Scores

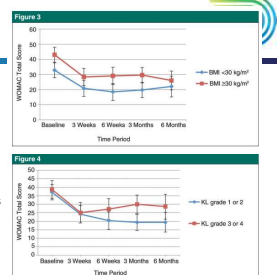


38

What to Inject?

• Corticosteroids

- Fairly poorly studied compared to other tx modalities
- Matzkin et al 2017 – 100 pts
 - Triamcinolone
 - Statistically sig improvements in Visual Numeric Pain and total WOMAC Scores
 - Those with BMI > 30 and worse OA did not see as much improvement

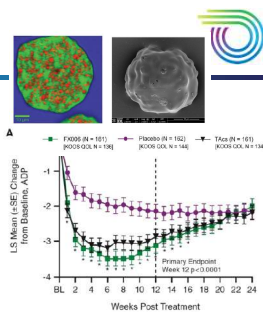


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What to Inject?

• Corticosteroids

- Conaghan et al 2018 – 484 pts
 - ER triamcinolone (Zilretta)
 - Industry-sponsored phase III trial – level I study
 - Average Daily Pain (ADP)
 - Zilretta >> placebo
 - Zilretta = triamcinolone



40

What to Inject?

• Corticosteroids

- Adverse events/Complications
 - Similar systemic effects as oral route, but less severe
 - Hyperglycemia typically occurs in the hours and days following injection
 - Risk of infection less than 0.05%
 - Post-injection "flare"

Table 3

Possible Adverse Effects and Complications of Intra-articular Corticosteroid Injections

Adverse Effects and Complications

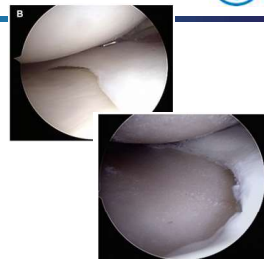
Skin pigmentation changes
 Fat or skin atrophy
 Residual injection site pain
 Facial flushing
 Hypothalamic-pituitary-adrenal axis suppression
 Increased blood glucose
 Septic arthritis
 Direct injury to the cartilage with a needle

41

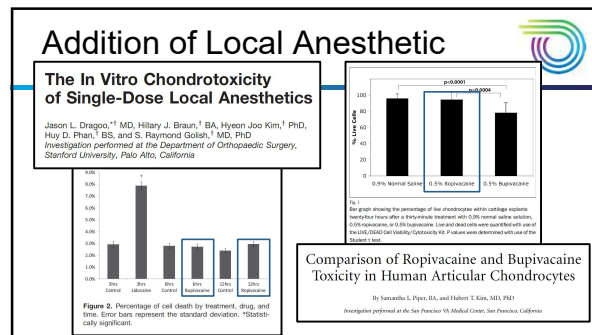
What to Inject?

• Corticosteroids

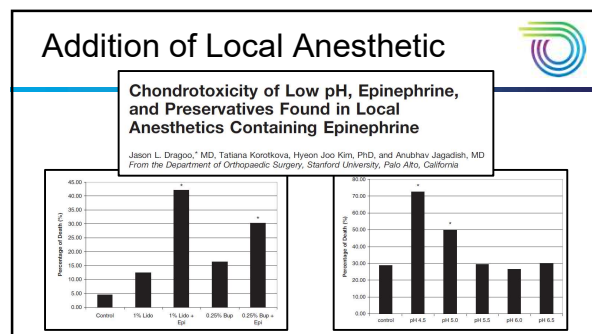
- Addition of local anesthetic
 - Somewhat controversial
 - Case reports of chondrolysis after use of local anesthetics in post-op infusion pumps
 - No high-quality clinical studies show this phenomenon following single-dose injection



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44

What to Inject?

- **Corticosteroids**
 - What I do:
 - Betadine swab, Non-sterile gloves, No sterile field, Ethyl Chloride spray
 - Triamcinolone (40mg/mL)
 - No local anesthetic
 - OK to reinject every 3-4 months, if necessary

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What to Inject?

- Viscosupplementation
 - Hyaluronic acid
 - Approved for use in the US in 1997
 - Viscoelastic properties
 - Anti-inflammatory
 - Anti-nociceptive



46

What to Inject?

- Viscosupplementation

Product Name	Active Ingredient	Molecular Weight (kDa)	Cross-Linked	No. of Injections*	Cost Per Syringe† (\$)
Hyalgan	Sodium hyaluronate	500 to 720	No	3 to 5	228.00
Symvisc	Hylan G F 20 (80:20 ratio of hylan A to hylan B)	5,000 to 6,000	Yes	3	473.88
Supartz, Supartz FX	Sodium hyaluronate	620 to 1,200	No	3 to 5	276.36
Orthovisc	Sodium hyaluronate	1,000 to 2,900	No	3 to 4	626.40
Nuflexa, Nuflexa	Sodium hyaluronate	2,400 to 3,600	Yes	3	388.48
Symvisc One	Hylan G F 20 (80:20 ratio of hylan A to hylan B)	5,000 to 6,000	Yes	1	1,421.65
Gel-One	Hyaluronican hydrogel	N/A	Yes	3	1,170.00
Monoject	Sodium hyaluronate	1,000 to 2,900	Yes	1	1,252.80

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What to Inject?

- Viscosupplementation
 - Adverse events/Complications
 - Similar to those undergoing corticosteroid injections
 - Localized pain and effusion
 - Risk of infection less than 0.05%
 - Pseudoseptic reaction



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What to Inject?



- Viscosupplementation
 - Bellamy et al 2006 - Cochrane Review
 - Beneficial effects on pain and physical function most pronounced at 5-13 weeks after injection
 - Campbell et al 2007 - review of 6 meta-analyses
 - Concluded that viscosupplementation results in improvements in physical function and pain reduction with a low risk for harm

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What to Inject?



- Viscosupplementation
 - Bellamy et al 2006 - Cochrane Review
 - Beneficial effects most pronounced
 - Campbell et al 2007
 - Concluded that improvements in reduction with a

Declarations of Interest

The original review was partially supported by Genzyme Biologics (formerly Biogen, Inc) and Hyaluronate, an unreviewed educational grant. Genzyme supported research and to review the treatment area for its products, and just that manufactured by Genzyme Biologics (formerly Biogen, Inc). The interpretation of the results are those of the reviewers who retain the right to publish.

Dr. Viscosupplementation and Dr. Viscosupplementation participated in the Reynolds (Reynolds 2002) trial. Dr. Bellamy was a co-investigator on the Steering Committee of the Reynolds (Reynolds 2002) trial, and previously provided consulting services to Biogen and Genzyme Inc.

Dr. Bousquet was a clinical investigator in the Reynolds (Reynolds 2002) trial.

Product label analyses were conducted for each respective manufacturer prior to finalization of the original review to permit any factual errors to be addressed. Comments were received from some but not all manufacturers.

50

What to Inject?



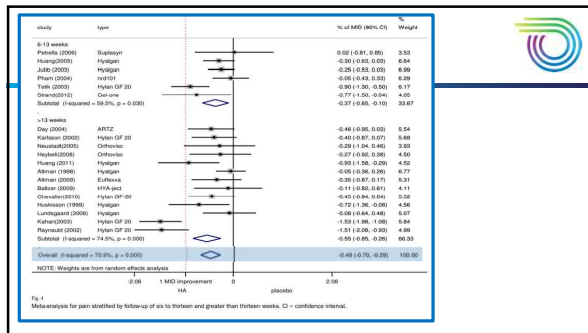
- Viscosupplementation
 - Miller et al 2013
 - Systematic review and meta-analysis

US-Approved Intra-Articular Hyaluronic Acid Injections are Safe and Effective in Patients with Knee Osteoarthritis: Systematic Review and Meta-Analysis of Randomized, Saline-Controlled Trials

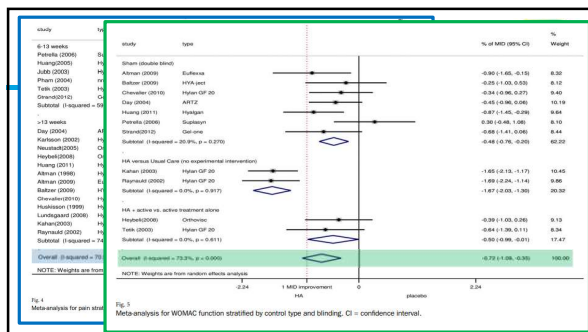
Larry E. Miller¹ and Jon E. Block²
¹Miller Scientific Consulting, Inc., Arroyo, CA, USA; ²The Joint Group, San Francisco, CA, USA
 Corresponding author email: jblock@jointgroup.com

Abstract: We conducted a systematic review and meta-analysis of randomized saline-controlled trials to determine the safety and efficacy of US-approved intra-articular hyaluronic acid (IAHA) injections for symptomatic knee osteoarthritis. A total of 29 studies representing 4,866 unique subjects (IAHA: 2,673; saline: 2,193) were included. IAHA injection resulted in very large treatment effects between 4 and 26 weeks for knee pain and function compared to pre-injection values, with standardized mean difference (SMD) values ranging from 1.07-1.37 (all $P < 0.001$). Compared to saline controls, SMDs with IAHA ranged from 0.38-0.43 for knee pain and 0.32-0.34 for knee function (all $P < 0.001$). There were no statistically significant differences between IAHA and saline controls for any safety outcome, including serious adverse events (SAEs) ($P = 0.12$), treatment-related SAEs ($P = 1.0$), study withdrawal ($P = 1.0$), and AE-related study withdrawal ($P = 0.46$). We conclude that intra-articular injection of US-approved HA products is safe and efficacious in patients with symptomatic knee osteoarthritis.

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What to Inject?

• Viscosupplementation

• What I do:


- Utilized for those patients with mild-moderate knee OA or degenerative meniscus tears
- Failed other appropriate conservative tx options
- Same technique as steroid injection
- Seems to either "really work" or "didn't do anything"
- Still approved by insurance carriers



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What to Inject?

- Orthobiologics
 - Platelet-rich plasma (PRP)
 - Mesenchymal "stem" cells



58


What to Inject?

- Orthobiologics
 - Platelet-rich plasma (PRP)
 - Mesenchymal "stem" cells

★ These treatments are NOT FDA-approved for the treatment of knee OA or degenerative meniscus tears

AND

They are NOT typically covered by insurance carriers




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What to Inject?

- Platelet-rich plasma (PRP)
 - Venous blood draw
 - Centrifugation
 - Platelet concentration "above baseline"
 - Release of growth factors

★

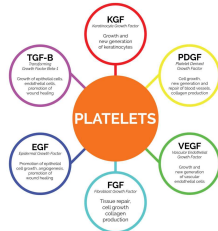


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What to Inject?

• Platelet-rich plasma (PRP)

- Venous blood draw
- Centrifugation
- Platelet concentration "above baseline"
 - Release of growth factors



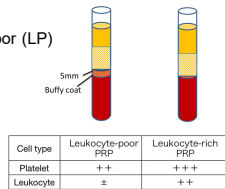
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What to Inject?

• Platelet-rich plasma (PRP)

- Leukocyte-rich (LR) vs Leukocyte-poor (LP)
 - LR-PRP results in much higher platelet concentrations but necessitates the inclusion of leukocytes
 - LP-PRP results in moderately elevated concentration of platelets without inclusion of leukocytes



*

62

What to Inject?

• Platelet-rich plasma (PRP)

- Meheux et al 2016
 - 6 level I randomized controlled trials comparing PRP to HA (visco)

Table 1

	WOMAC Score (Meheux et al)		
	PRP	HA	p-value
Baseline	52.16	52.09	<0.02
12-26 weeks	38.5	43.4	<0.0008
26-52 weeks	22.8	38.1	<0.0002

WOMAC
Score

*

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What to Inject?

• Platelet-rich plasma (PRP)

• What I do:

- Starting to offer more frequently
- Those patients with mild-moderate knee OA or degenerative meniscus tears
- Failed other appropriate conservative tx options
- Clearly notify patients that this is NOT covered by insurance
- Utilize LP-PRP



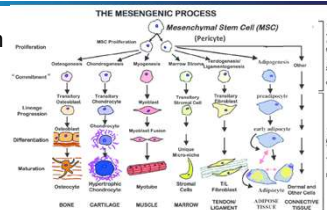
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What to Inject?

• Mesenchymal stem cells (MSCs)

- Originally described by Caplan in 1991



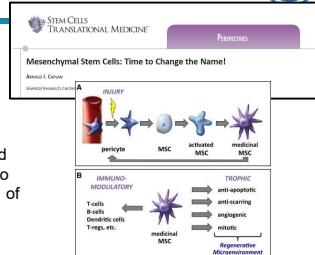
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What to Inject?

• Mesenchymal stem cells (MSCs)

- Originally described by Caplan in 1991
- Caplan 2017 – suggested changing nomenclature to reflect our understanding of how these cells actually function



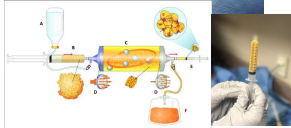
*

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What to Inject?

- MSC injection
 - Dai et al 2020 – Systematic Review and meta-analysis
 - 13 RCTs included
 - No significant difference when compared to placebo
 - VAS Pain
 - WOMAC Pain
 - Function
 - Stiffness

*

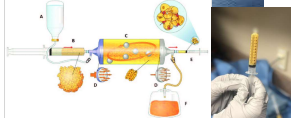


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What to Inject?

- Mesenchymal stem cells (MSCs)
 - What I do:
 - We do not offer this service
 - Cost – \$\$\$\$
 - Not necessarily against autologous MSCs

*



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What to Inject?

- Mesenchymal stem cells (MSCs)
 - What I do:
 - We do not offer this service
 - Cost – \$\$\$\$
 - Not necessarily against autologous MSCs
 - AVOID allogeneic stem cell products

*

Nebraska, Iowa attorneys general announce lawsuits against stem cell therapy clinics

December 16, 2019 | 2 min read
Health officials issue safety alert on unapproved stem cell products after Nebraska outbreak

[ADD TOPIC TO EMAIL ALERTS](#)

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Summary



- Appropriate H&P
 - Specific PE tests for meniscus tears are unreliable
- Many different types of meniscus tears
 - Treatment recommendations differ based upon these as well as patient factors
- Many meniscus tears are treated conservatively initially
- Injections are a common conservative treatment modality
 - Ensure proper injection technique
- Injectables
 - Corticosteroid
 - Viscosupplementation
 - Orthobiologics ***

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References



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