Intra-articular treatment

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Osteoarthritis

excessive use

wear and tear



Synovitis Osteoarthritis

acute injury





Anabolic / catabolic equilibrium disturbed



Intra-articular Injections - Advantages -

- Peak concentration faster and higher
- Lower doses needed
- Less systemic adverse effects

- Risks -

- Local adverse effects (rare)
 - Infection, flare, crystal induced synovitis

Intra-articular Injections - Therapeutic aim -

- Pain reduction
- Anti-inflammation
- Return to function
 - (not sound but functional?)
- Homeostasis
- Chondroprotection



Intra-articular Injections - Acceptance -

- Human
 Equine
 Canine
 +++
 +/-
 - Feline

Pressure from patient(-owner) Ease of technique

Intra-articular Injections - Agents -

- Corticosteroids
- Hyaluronic acid
- PSGAGs
- Local anesthetic

- Blood Products
- Stem Cells

- Polyacrylamide
- Bladder matrix
- Yttrium-90, Rehnium-186
- Botulinum neurotoxin

Corticosteroids - Advantages -

- Potent antiinflammatory agent
- Decrease of MMPs (catabolic reactions)
- Clinical success

- Risks -

- Increased aggrecanase activity
- Increased early catabolic response
- Short term effects

Corticosteroids

- Most studies (ca&eq) in vitro or OA model
- Contradictory results
- In dogs mainly used for end stage OA
 - Histologic and macroscopic positive outcomes in cclmodels. Lack of clinical evaluation
- In horses combined with HA (tradition)
 - Efficacy for approximately 8 weeks
- In human for acute rather than chronic
 - Pain reduction up to four weeks

Hyaluronic Acid - Aim -

- Restoring viscoelasticity
- Painreduction related to movement

- Suggestions -

- Anti-inflammatory, chondroprotective
- Promotion of HA synthesis
- Stimmulation of matrix synthesis
- Inhibition of MMPs

Hyaluronic Acid

- Most studies (ca&eq) in vitro or OA model
- Contradictory results
- In dogs mainly used for end stage OA
 - Not established
- In horses steroid combination (tradition)
- In human for acute rather than chronic
 - As second line treatment if steroids failed



TREATMENT OF OSTEOARTHRITIS OF THE KNEE

EVIDENCE-BASED GUIDELINE 2ND EDITION

Adopted by the American Academy of Orthopaedic Surgeons Board of Directors May 18, 2013

RECOMMENDATION 9

We cannot recommend using hyaluronic acid for patients with symptomatic osteoarthritis of the knee.

Strength of Recommendation: Strong

Description: Evidence is based on two or more "High" strength studies with consistent findings for recommending for or against the intervention. A **Strong** recommendation means that the quality of the supporting evidence is high. A harms analysis on this recommendation was not performed.

Implications: Practitioners should follow a **Strong** recommendation unless a clear and compelling rationale for an alternative approach is present.

Biologicals

Biological factors Cartilage **Synthesis Protein** ases IL-4 IL-1Ra IL-10 sTNFaR IL-13 IL-6 IL-8 LIF OSM IL-17 **IL-1** IL-18 TNFα **IGF-1 MMPs BMPs** Cartilage TGF-β3 **Degradation** PGE2

Biological idea

Situation





Effect



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Biologicals

- They are not all the same
- Variations from method to method
- Variations from patient to patient
- Variations from time to time

Autologous Blood Products

- Biological produced from patient's blood
 - Autologous, modified
- Biological produced by the veterinarian
 - Pharmaceutical Law, EMEA and FDA
- Medical device or standardised method
 - They are not all the same

ABPs indications

- Implantation & Reconstruction
- Joint disease
- Tendovaginitis, Bursitis
- Tendon & ligament injuries
- Radiculopathies

ABPs Taxonomy



Platelet Rich Plasma - PRP -

Content

- Non modified Plasma
 - Fibrinogen, Prothrombine
- Platelets
 - C > 4 x Baseline, (Marx 2001)
- Leucocytes
 - Variable concentrations

Concept

- Growthfactor release
 - At site of damage
- Growthfactors
 - PDGF, TGF, VEGF, FGF...
- Anabolic stimmulation
 - Stimulation of migration, cell devision, synthesis...

PRP - Clinical use

Pathologies with substantial loss/need

- Tendon lesions, fractures, implants, wounds
- Documented for canine equine and human medicine
- Joint pathologies
 - Used for OA, cave: coagulation, WBCs, RBCs
 - Lack of clinical documentation

Autologous (Conditioned) Plasma - A(C)P -

Ingredients

- Non modified Plasma
 - Fibrinogen, Prothrombine
- Platelets
 - No significant differnce to whole blood, (Stief 2011)
- Leucocytes
 - Depletion of WBCs

Concept

- Growthfactor release
 - At site of damage
- Growthfactors
 - PDGF, TGF, VEGF, FGF...
- Anabolic stimmulation
 - Stimulation of migration, cell devision, synthesis...

ACP - Preparation





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ACP - Clinical use

Same as PRP

Hardly any Platelet elevation \rightarrow low in GFs Depletion of WBCs \rightarrow necessary?

Autologous Conditioned Serum - ACS -

Ingredients

- Serum
 - No coagulation factors
- Growthfactors
 - Release of GF's during preparation (FGF, IGF...)
- Cytocines
 - De novo synthesis by WBCs (IL-1Ra, IL-10...)

Concept

- Direct application
 - Concentrated proteins
- Katabolic supression
 - IL-1 inhibition, antiinflammation
- Anabolic stimmulation
 - Stimulation of migration, cell devision, synthesis...

ACS - Preparation





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ACS - Clinical use

Pathologies of synovial structures

- OA, synovitis of joints, bursae and tendon sheaths
- Good documentation for equine and human use
- Caseseries for canine, studies in progress

Tendon pathologies

- Used for bowed tendon and core lesions
- Lack of clinical documentation, studies in progress

What for what and when?

	Joint	Tendon
Biological	ACS	PRP
Reason	Anti-inflammatory Anabolic No cells or clotting factors	Anabolic Clotting factors Cells
Timing	Acute and chronic, PO	Acute and proliferative phase

Everything depends on...

- Stage of disease
- Use of patient
- Owners compliance

Intra-articular injections for OA

- Safe if correct technique is applied
- Agent directly applied to pathology
- Selection of the right agent
- Selection of the right cases
- Multimodal treatment is essential
 - Surgery, weight management, physical exercise, nutritional supplementation

Vielen Dank!

