

# TREATMENT OF CARTILAGE INJURIES IN THE KNEE

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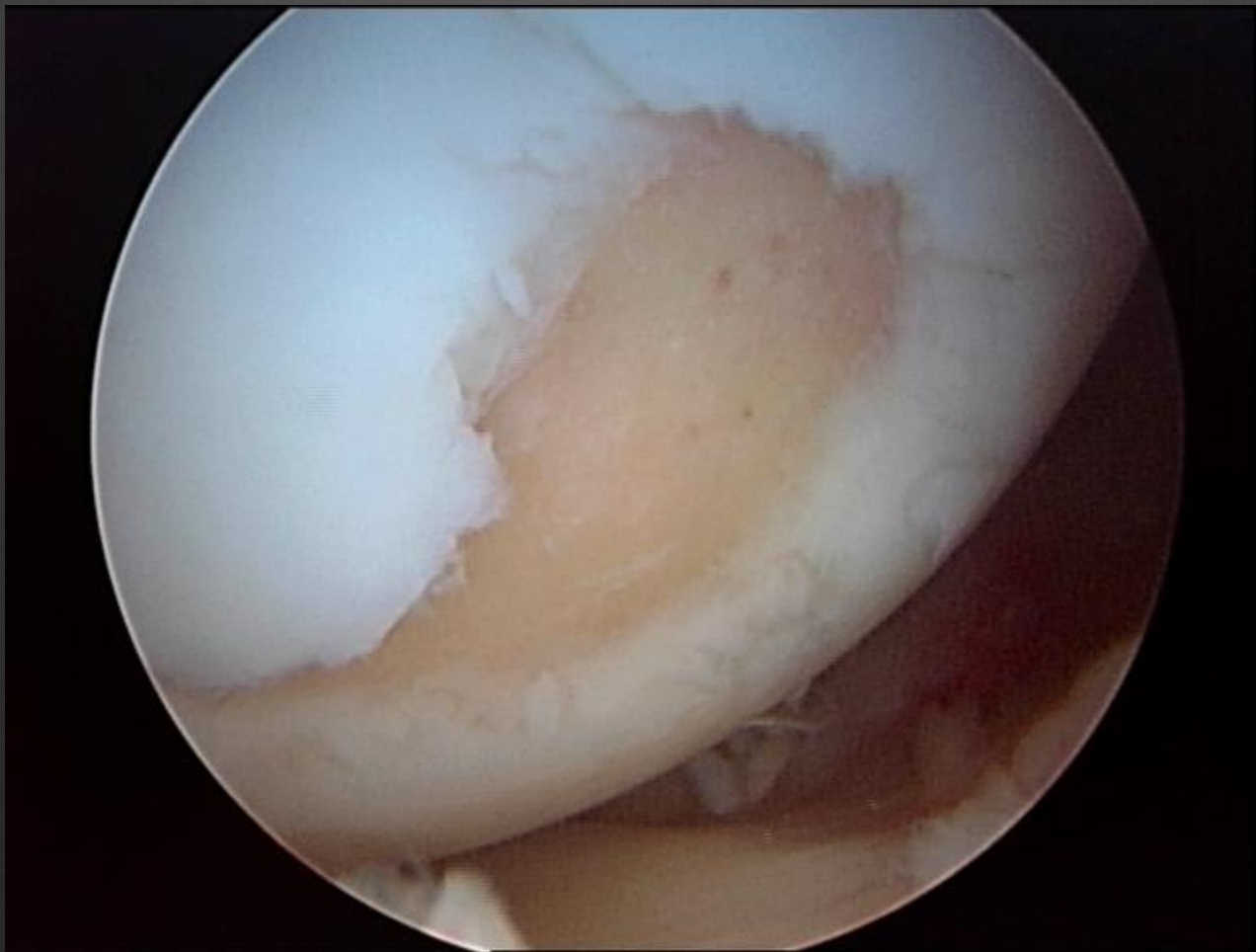


# WHAT I HOPE YOU GET FROM THIS TALK.....

- ❁ DEVELOP UNDERSTANDING OF WHY CARTILAGE REPAIR IS DIFFICULT TO MANAGE
- ❁ SEE THE MANY MANY OPTIONS THAT ARE USED TODAY
- ❁ FUTURE STRATEGIES IN CARTILAGE REPAIR
- ❁ RETURN TO PLAY AFTER SURGERY
- ❁ UNDERSTAND WHY SOME PEOPLE DON'T DO AS WELL WITH THERAPY AFTER A KNEE SCOPE



THIS CARTILAGE LESION NEEDS A SOLUTION....



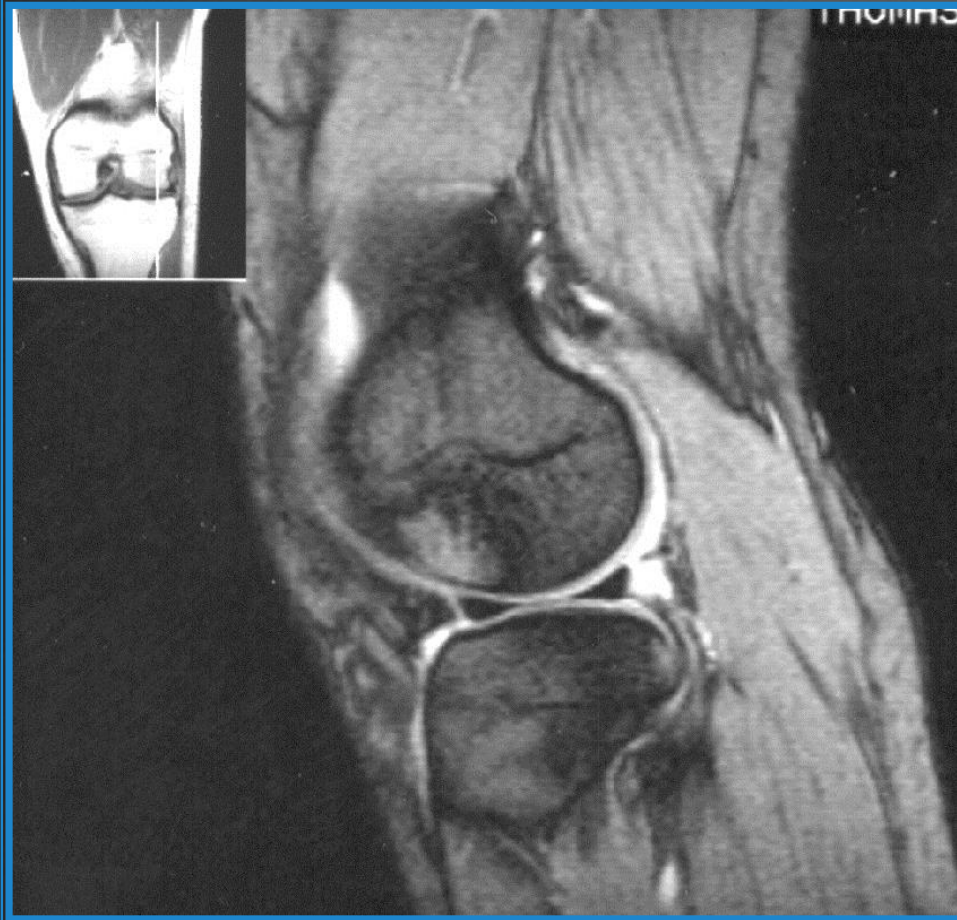
Courtesy of Brian Cole, MD

C-00029.A 08/2006

# SOLUTION TO THIS .. Weight loss and therapy



# The “Bone Bruise”



OCCURS WITH ACL  
INJURIES

VERY UNPREDICTABLE

WHY YOU CANT TREAT  
ALL ACL INJURIES THE  
SAME

# CARTILAGE

- ❁ NO NERVES
  - ❁ NO BLOOD SUPPLY
  - ❁ NO LYMPHATIC SYSTEM
  - ❁ FEW CELLS
- 
- ❁ AND WE WONDER WHY ITS HARD TO TREAT!!!!!!!

# CARTILAGE

- ❁ FUNCTIONS TO PROTECT SUBCHONDRAL BONE
- ❁ LOW FRICTION SURFACE DISTRIBUTES LOAD
- ❁ COMPOSED OF CHONDROCYTES SURROUNDED BY EXTRACELLULAR MATRIX
  - ❁ COLLAGEN TYPE 2
  - ❁ PROTEOGLYCAN
  - ❁ WATER



# ARTICULAR CARTILAGE: LAYERS

## SUPERFICIAL

- Flat cells
- High collagen content
- High water content
- High tensile strength

## TRANSITIONAL

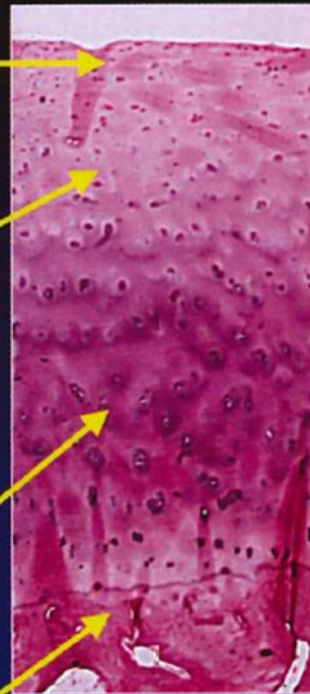
- Spherical cells
- More proteoglycans
- Low water content
- Fibrils bent to form arcades

## RADIAL

- Active groups of cells
- Perpendicular fiber orientation

## CALCIFIED CARTILAGE

- Anchor point of collagen



# CLASSIFICATION SYSTEM

**Table 1 Classification of articular lesions by severity**

Grade	Outerbridge	Modified outerbridge	ICRS
0	Normal cartilage	Intact cartilage	Intact cartilage
I	Softening and swelling	Chondral softening or blistering with intact surface	Superficial (soft indentation or superficial fissures and cracks)
II	Fragmentation and fissures in area less than 0.5 inch in diameter	Superficial ulceration, fibrillation, or fissuring less than 50% of depth of cartilage	Lesion less than half the thickness of articular cartilage
III	Fragmentation and fissures in area larger than 0.5 inch in diameter	Deep ulceration, fibrillation, fissuring or chondral flap more than 50% of cartilage without exposed bone	Lesion more than half the thickness of articular cartilage
IV	Exposed subchondral bone	Full-thickness wear with exposed subchondral bone	Lesion extending to subchondral bone

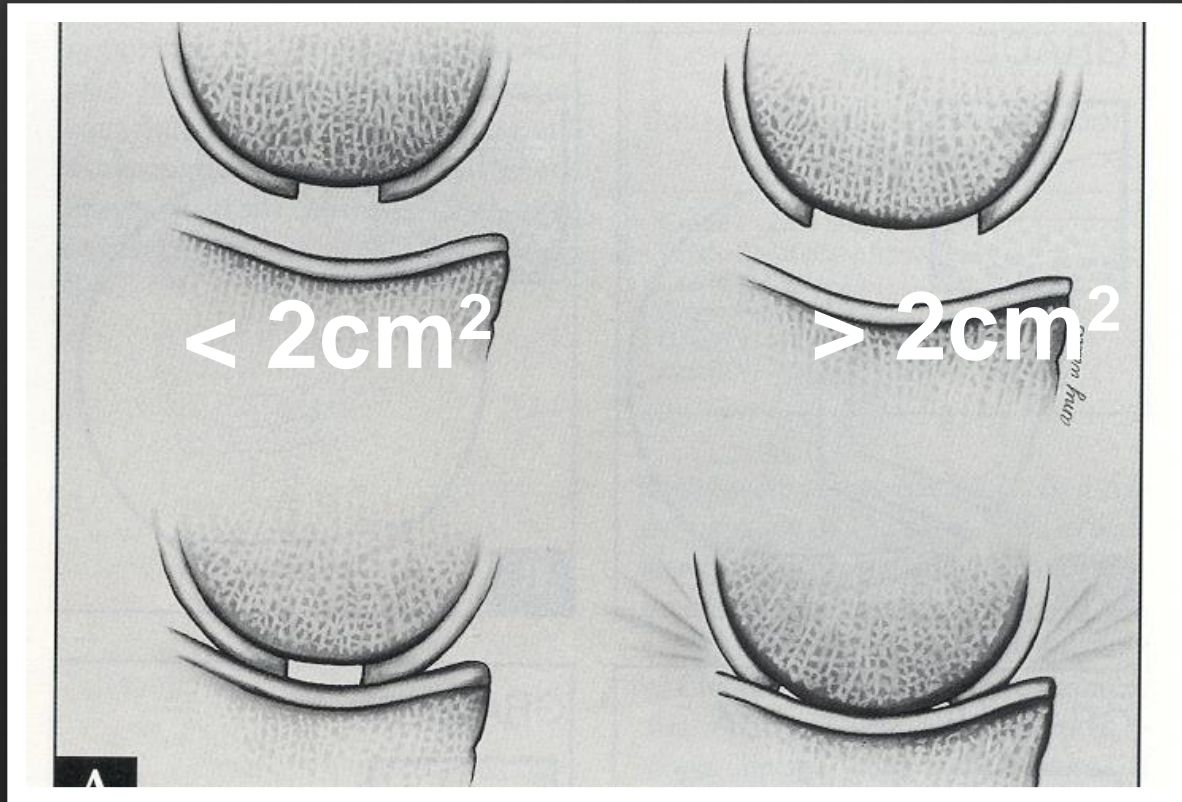
ICRS: International Cartilage Repair Society.

# HOW I APPROACH CARTILAGE REPAIR..

- ⊗ DEFECT SPECIFIC FACTORS
- ⊗ PATIENT SPECIFIC FACTORS
- ⊗ JOINT SPECIFIC FACTORS

# DEFECT SPECIFIC FACTORS

- ❁ SIZE
- ❁ DEPTH
- ❁ LOCATION
- ❁ DEGREE OF CONTAINMENT



SIZE , DEPTH, CONTAINMENT  
65% more pressure on surrounding  
cartilage if  $> 1 \text{ cm}^2$

# PATIENT SPECIFIC FACTORS

- ⊗ AGE – BETTER OUTCOMES IF LESS THAN 30Y/O
- ⊗ ACTIVITY LEVEL
- ⊗ EXPECTATIONS – 3 YEAR RESULTS BETTER THAN 10 YEAR OUTCOMES
- ⊗ WEIGHT- WORSE IF BMI > 30 KG/M2



# JOINT SPECIFIC FACTORS

- ⊗ ALIGNMENT – WILL NEVER WORK IF VARUS AND VALGUS NOT CORRECTED
- ⊗ STABILITY- WILL FAIL IF ACL DEFICIENT KNEE
- ⊗ STATUS OF MENISCUS- IF LARGE MENISCECTOMY  
..INCREASES CONTACT FORCES ON CARTILAGE  
AND ADDS INSTABILITY



# Meniscectomy Vs. Repair



# CARTILAGE OPTIONS

- ⦿ ABRASION ARTHROPLASTY
- ⦿ MICROFRACTURE
- ⦿ ACELLULAR MATRIX INDUCED MICROFRACTURE
- ⦿ OATS – OSTEOCHONDRAL AUTOGRAFT
- ⦿ OSTEOCHONDRAL ALLOGRAFT
- ⦿ ACI- AUTOLOGOUS CHONDROCYTE IMPLANTATION
- ⦿ MACI

**Table 2 Treatment options for articular cartilage lesions**

<b>Procedure</b>	<b>Indications</b>	<b>Outcome</b>
Arthroscopic debridement and lavage	Minimal symptoms	Palliative
Marrow stimulation	Smaller lesions, low-demand patient	Reparative
Osteochondral autograft	Smaller lesions, low-or high-demand patients	Restorative
Osteochondral allograft	Larger lesions with bone loss, low-or high-demand patients	Restorative
Autologous chondrocyte implantation	Small and large lesions with and without bone loss, high-demand patients	Restorative
Genetic engineering	Investigational	Restorative

From Garrick JG, editor: Orthopaedic knowledge update: sports medicine, 3<sup>rd</sup> ed, Rosemont, IL, 2004, American Academy of Orthopaedic Surgeons.

## Özmeriç A *et al.* New treatment algorithm in cartilage injuries

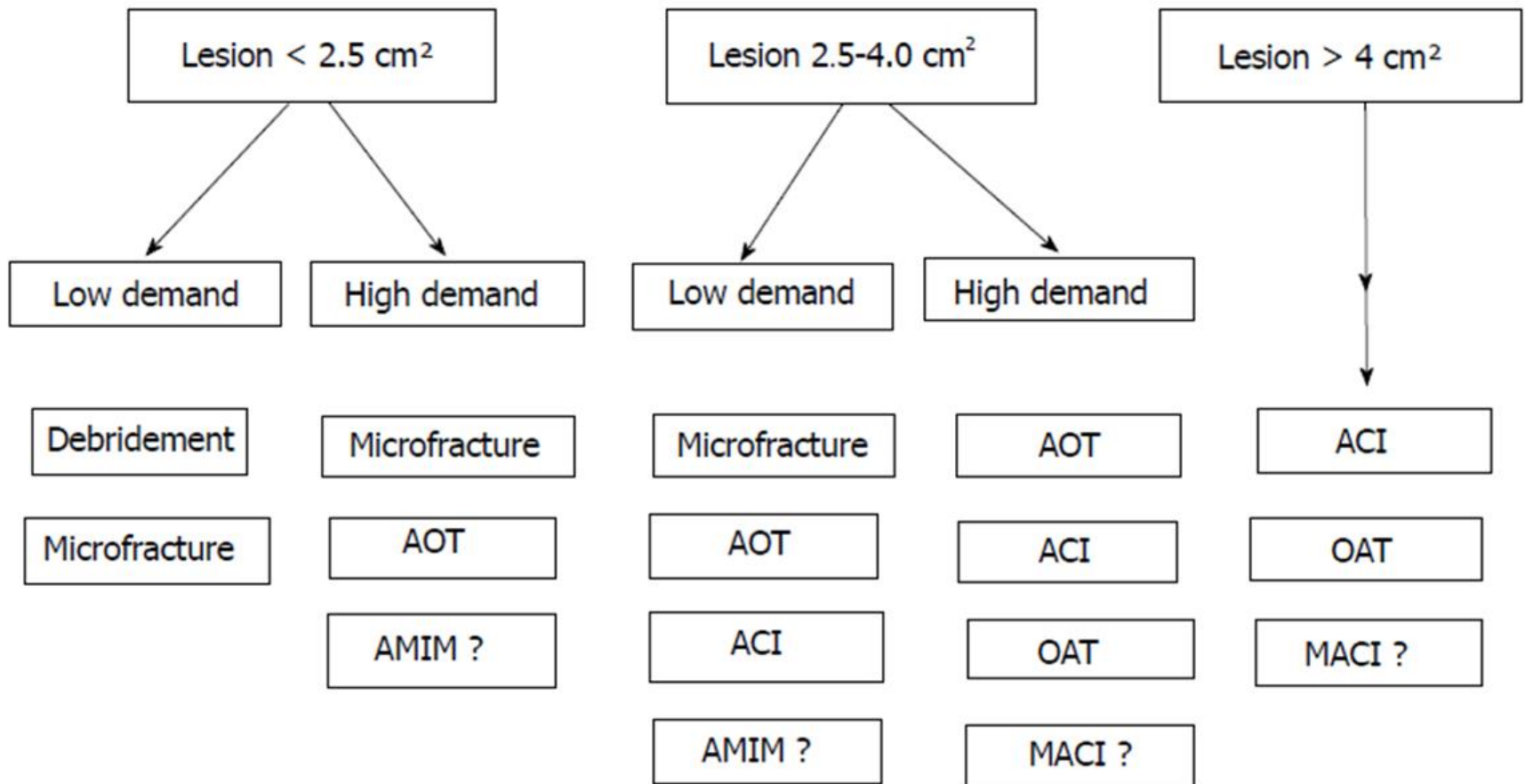
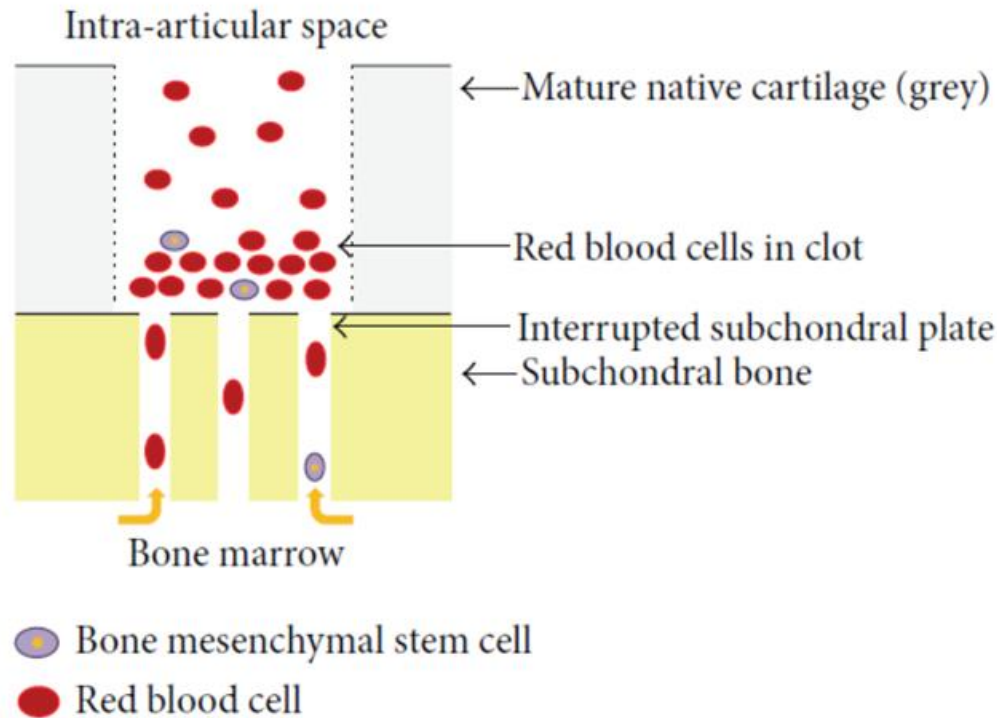


TABLE 1: Summary of techniques.

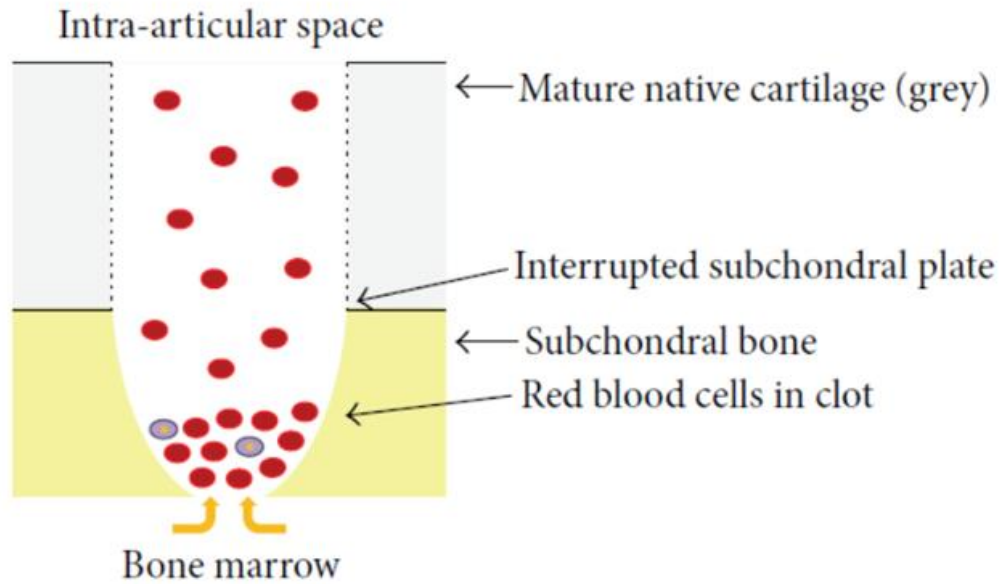
Category/technique	Schematic representation	Notes	Commercial products
Surgical bone marrow access			
Pridie drilling	<p>Intra-articular space</p> <p>← Mature native cartilage (grey)</p> <p>← Red blood cells in clot</p> <p>← Interrupted subchondral plate</p> <p>← Subchondral bone</p> <p>Bone marrow</p> <p>● Bone mesenchymal stem cell</p> <p>● Red blood cell</p>	<p><i>Cell source:</i> autologous bone marrow constituents</p> <p><i>Scaffold:</i> none</p> <p>(i) 1 stage</p> <p>(ii) Open procedure</p> <p>(iii) 2- to 2.5-mm drill holes to access bone marrow</p> <p>(iv) Inconsistent results</p> <p>(v) Long recovery</p> <p>(vi) High complication rate</p>	

Microfracture



*Cell source:* autologous bone marrow constituents  
*Scaffold:* none  
(i) 1 stage  
(ii) Arthroscopic procedure  
(iii) 0.5- to 1-mm drill holes to access bone marrow  
(iv) Less impact than Pridie drilling on biomechanics of underlying subchondral bone

Abrasion  
chondroplasty



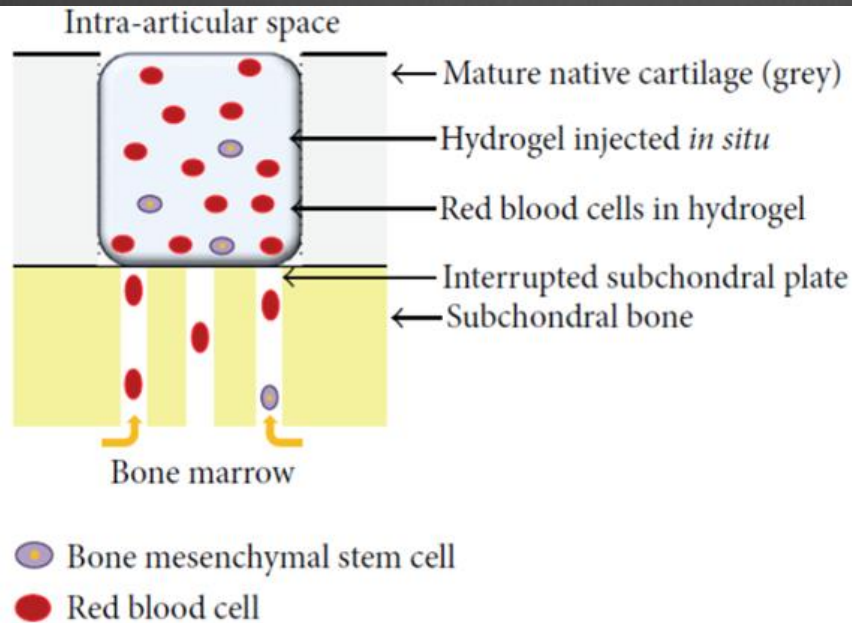
- Bone mesenchymal stem cell
- Red blood cell

*Cell source:* autologous  
bone marrow constituents  
*Scaffold:* none  
(i) 1 stage  
(ii) Arthroscopic procedure  
(iii) Irreproducible,  
unreliable  
(iv) Loss of underlying  
subchondral mechanical  
support

Category/technique	Schematic representation	Notes	Commercial products
Mosaicplasty	<p>Intra-articular space</p> <p>← Mature native cartilage (grey)</p> <p>← Implanted osteochondral autografts</p> <p>← Interrupted subchondral plate</p> <p>← Subchondral bone</p> <p>Bone marrow</p> <p>● Red blood cell</p>	<p><i>Cell source:</i> osteochondral autograft and autologous bone marrow constituents</p> <p><i>Scaffold:</i> N/A</p> <p>(i) 1 stage</p> <p>(ii) Morbidity at harvest site</p> <p>(iii) Osteochondral plugs 15–20 mm deep</p> <p>(iv) Blood clot in interspaces</p>	



Autologous  
matrix-induced  
chondrogenesis



Cell source: autologous  
bone marrow constituents

Scaffold: hydrogel

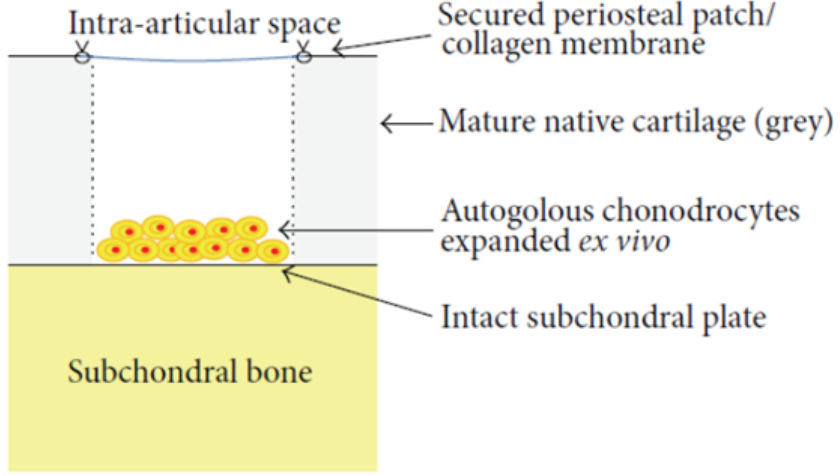
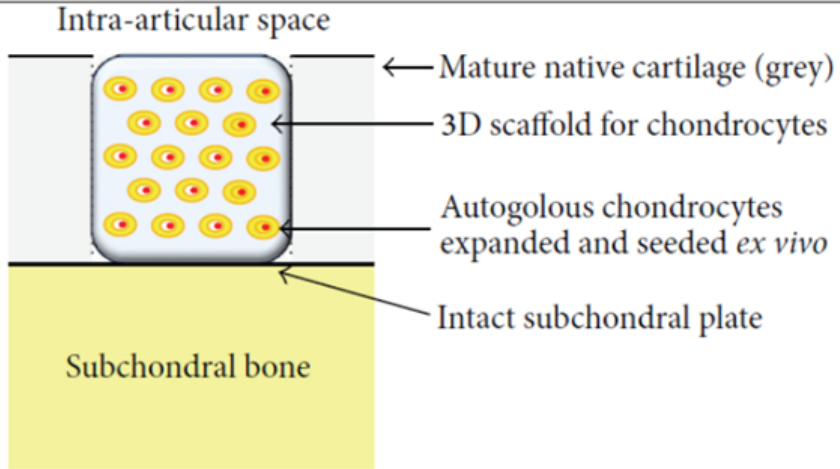
(i) 1 stage

(ii) Arthroscopic procedure ChonDux

(iii) 0.5- to 1-mm drill holes to access bone marrow CART-PATCH  
BST-CarGel

(iv) Less impact than Pridie  
drilling on biomechanics of  
underlying subchondral  
bone

Chondrocyte implantation

First generation	 <p>Intra-articular space</p> <p>Secured periosteal patch/ collagen membrane</p> <p>Mature native cartilage (grey)</p> <p>Autologous chondrocytes expanded <i>ex vivo</i></p> <p>Intact subchondral plate</p> <p>Subchondral bone</p>	<p><i>Cell Source:</i> autologous chondrocytes  <i>Scaffold:</i> none          (i) 2 stages          (ii) Periosteal patch or collagen membrane          (iii) Secured by sutures and/or fibrin glue          (iv) Greatest clinical experience</p>	Chondro-Gide Carticel
Second generation	 <p>Intra-articular space</p> <p>Mature native cartilage (grey)</p> <p>3D scaffold for chondrocytes</p> <p>Autologous chondrocytes expanded and seeded <i>ex vivo</i></p> <p>Intact subchondral plate</p> <p>Subchondral bone</p>	<p><i>Cell source:</i> autologous chondrocytes  <i>Scaffold:</i> hydrogel, fibrous scaffold, decellularized ECM, or composite          (i) 1 or 2 stages          (ii) ±Cells expanded and seeded in scaffold or matrix          (iii) Also known as matrix-induced autologous chondrocyte implantation (MACI)</p>	<p>Hyalograft C          BioSeed-C          Histogenics          NeoCart          CaReS          Cartilage Autograft Implantation System</p>

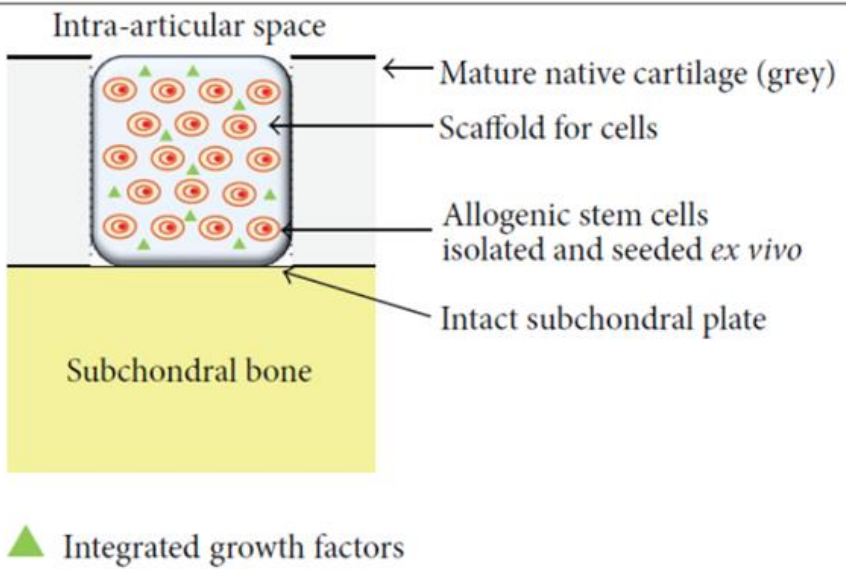
Category/technique

Schematic representation

Notes

Commercial products

Third generation



Cell source: allogenic stem cells, autologous stem cells  
Scaffold: hydrogel, fibrous scaffold, decellularized ECM, or composite  
(i) 1 stage  
(ii) Differentiation of pluripotent stem cells induced by environment  
(iii) Least clinical experience

DeNovo ET  
DeNovo NT

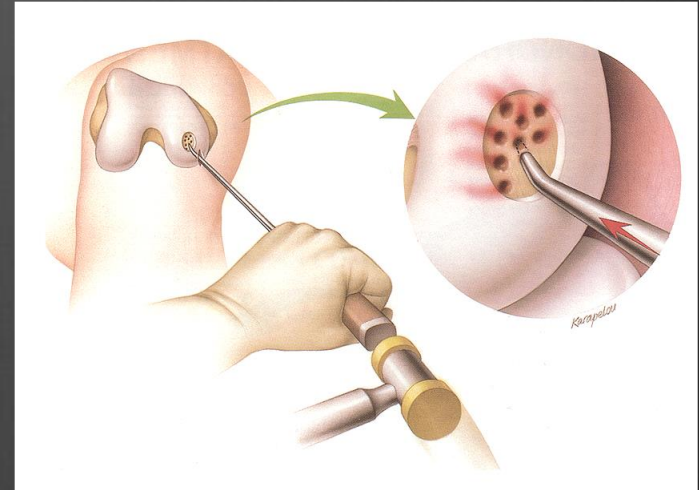
# Debridement & Lavage

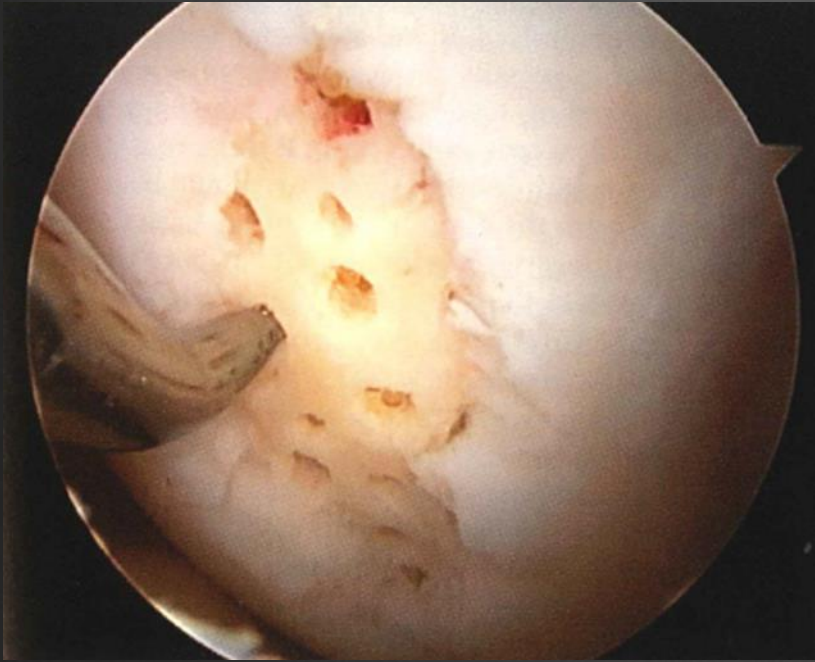
- ⊗ Strengths:
  - ⊗ Arthroscopic
  - ⊗ Does not violate sub-chondral bone
  - ⊗ Temporary pain relief
- ⊗ Limitations:
  - ⊗ Low prospects for long term result
  - ⊗ Fibro-cartilage



# MicroFracture

- ⊗ Strengths:
  - ⊗ Arthroscopic
  - ⊗ Relatively simple procedure
- ⊗ Limitations:
  - ⊗ Creates fibro-cartilage / poor wear characteristics
  - ⊗ More effective on smaller defects
  - ⊗ 6-8 weeks non-weight-bearing and CPM required to optimize results





## Table 1. Key Rehabilitation Points

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### Lesions of the femoral condyle or tibial plateau

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Immediate continuous passive motion, 8 hours daily for 8 weeks; 1 cycle per minute at 30° to 70°

No brace

Touch-down (20%-30%) crutch walking for 8 weeks

Cycling (light resistance): start 2 weeks postoperatively

Deep water exercise: start 2 weeks postoperatively

After 8 weeks, full weightbearing and active range of motion

No cutting, turning, or jumping for at least 4 to 9 months depending on the patient

May be longer for competitive or larger patients

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### Patellofemoral lesions

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Immediate continuous passive motion, 8 hours daily for 8 weeks at 0° to 50°

Brace locked at 0°; full weightbearing at 2 weeks

Stationary bike (light resistance): start 2 weeks postoperatively

Water program (no impact): start 2 weeks postoperatively

After 8 weeks, begin walking with a brace

Treadmill at 7° incline starting at 12 weeks postoperatively

Biking and water program: increase intensity at 8 to 12 weeks

Elastic resistance program with 0° to 30° knee bends starting at 12 weeks

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# MICROFRACTURE

- ❁ PERFORATE SUBCHONDRAL BONE
- ❁ PROMOTES MIGRATION OF MESENCHYMAL STEM CELLS
- ❁ HYALINE LIKE CARTILAGE EARLY THEN FIBROCARILAGE - DETERIORATES AFTER 3 YEARS



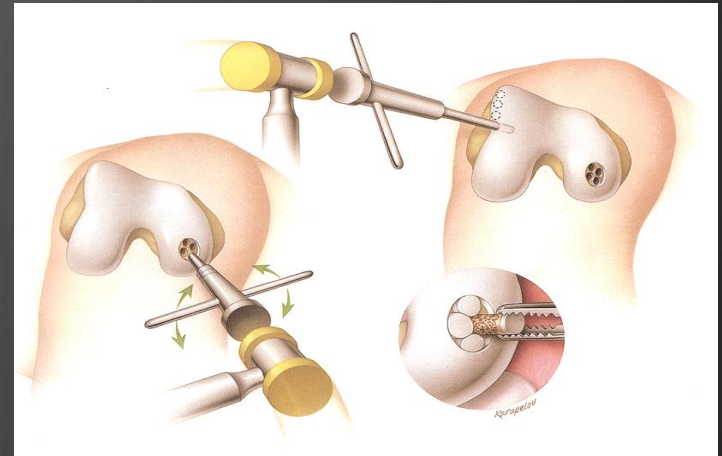
# Osteochondral Autografting

## ⊗ Strengths:

- ⊗ May be performed arthroscopically
- ⊗ Fills defect with native cartilage

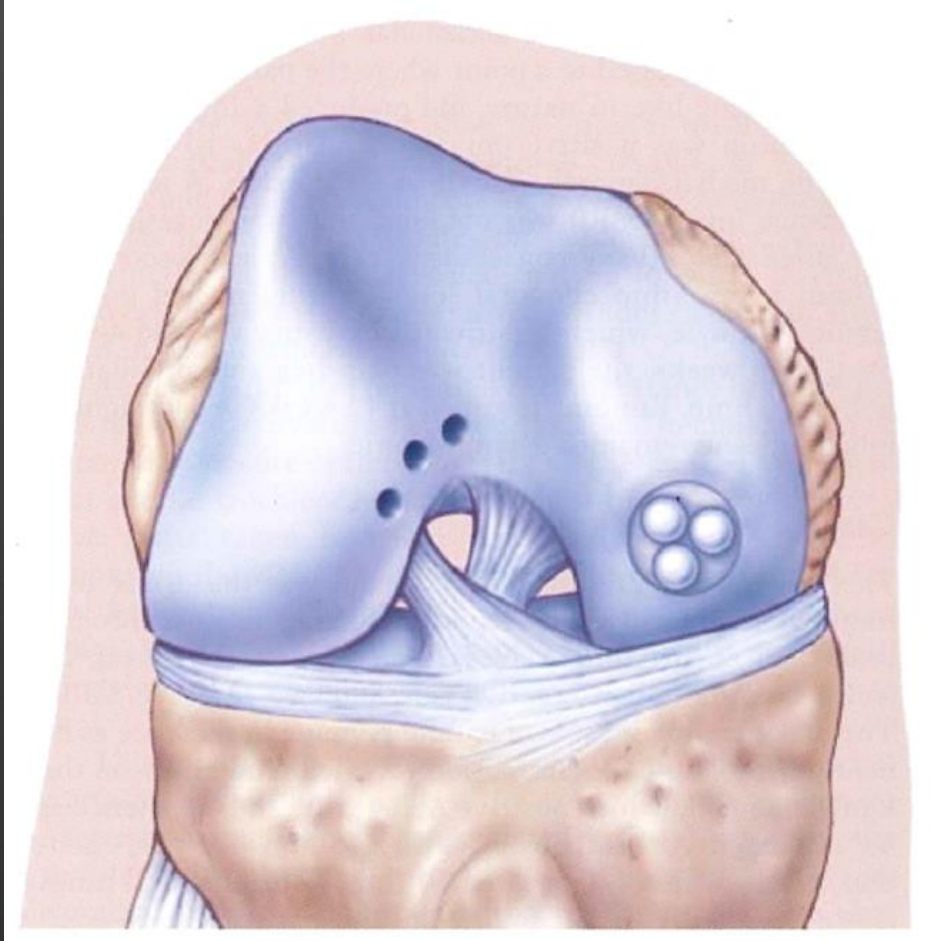
## ⊗ Limitations:

- ⊗ Limited to smaller defects
- ⊗ Donor site morbidity
- ⊗ No lateral integration
- ⊗ Congruity of joint difficult to re-produce with multiple plugs



Courtesy of Brian Cole, MD

C-00029.A 08/2006

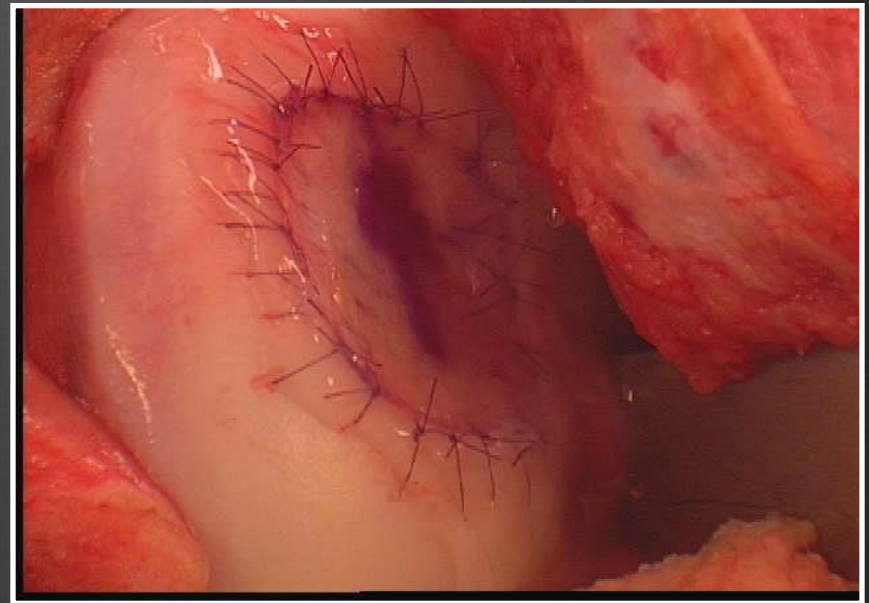


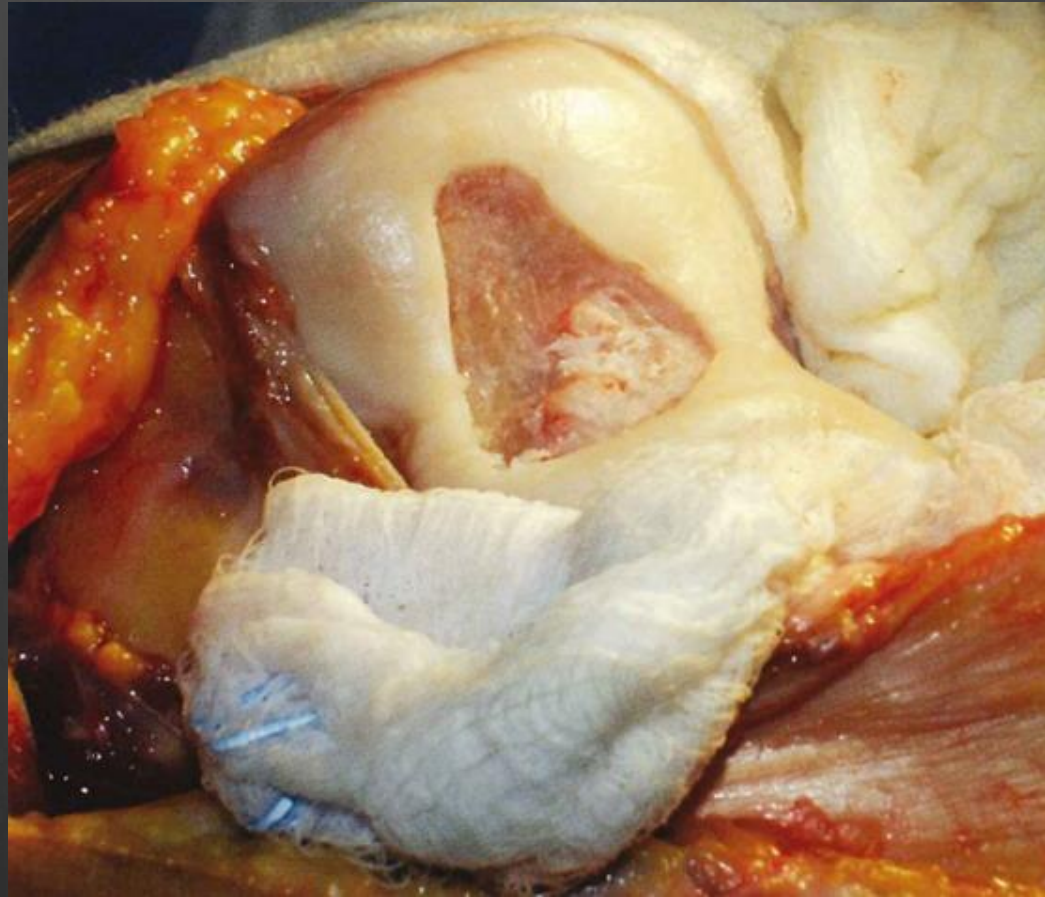
# OSTEOCHONDRAL ALLOGRAFT

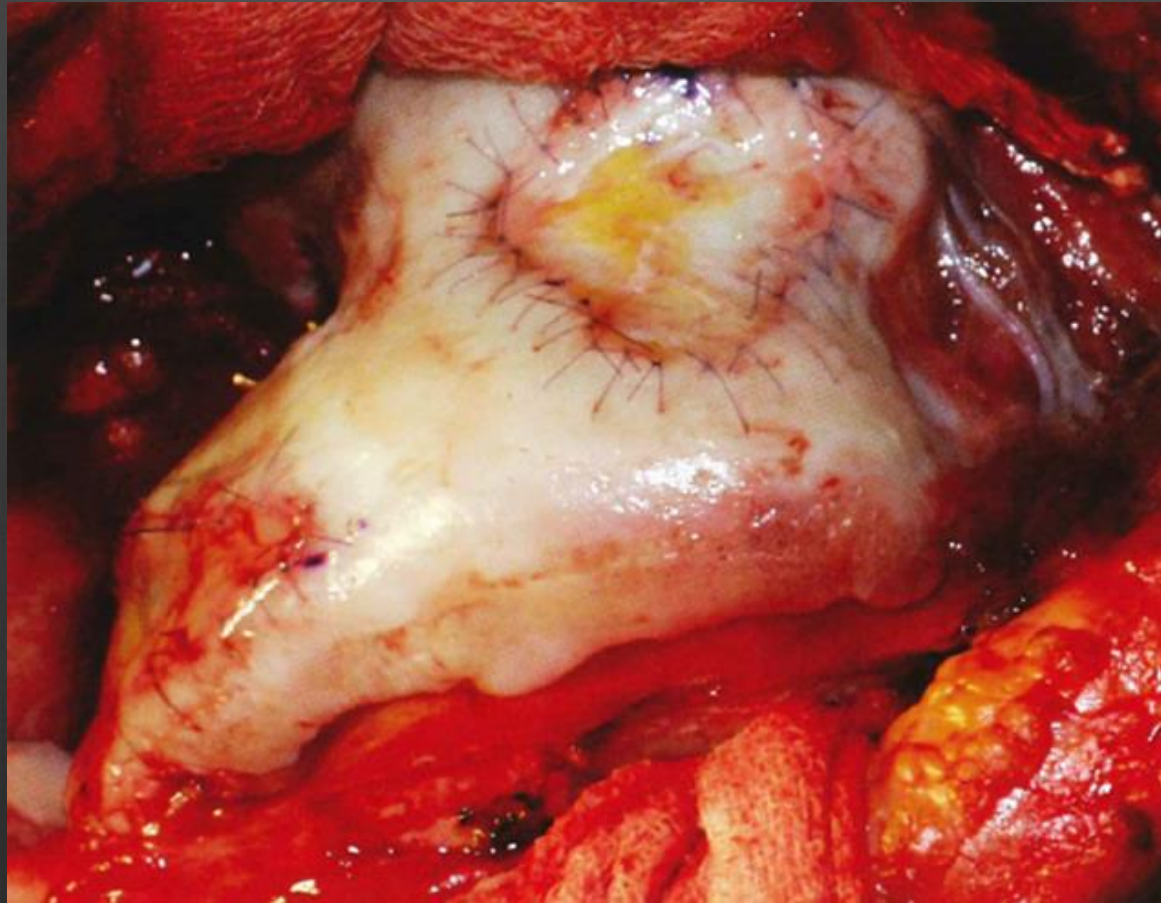
- ⊗ USED WITH LARGE LESIONS FOR SALVAGE PROCEDURES
- ⊗ MUST RADIATE TO DECREASE IMMUNOGENICITY WHICH KILLS CHONROCYTES
- ⊗ NEWER METHODS ( CRYOPRESERVED, FRESH, FRESH FROZEN ) HAVE IMPROVED RESULTS
  
- ⊗ DIFFICULT TO OBTAIN

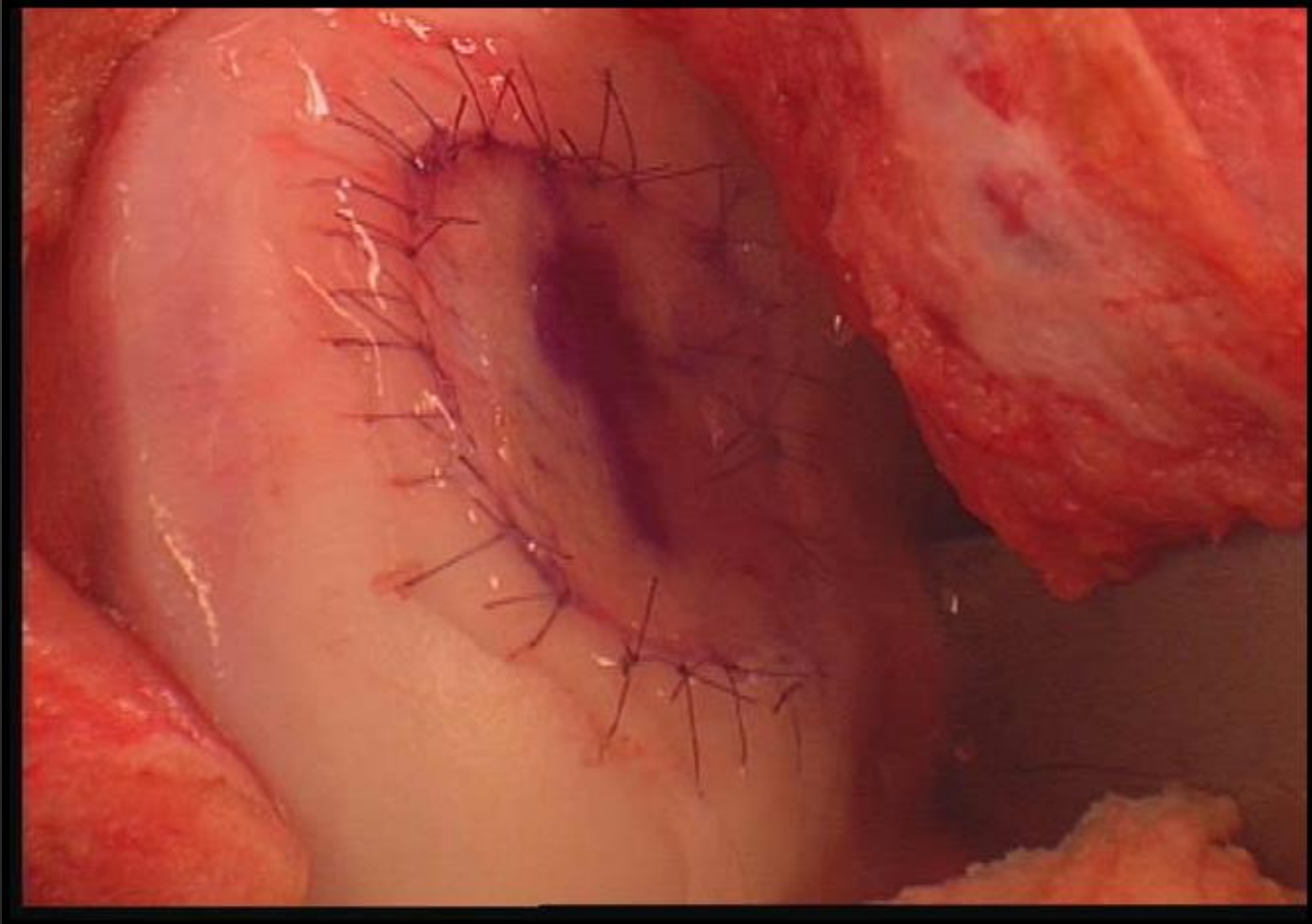
# Autologous Chondrocyte Implantation / Carticel®

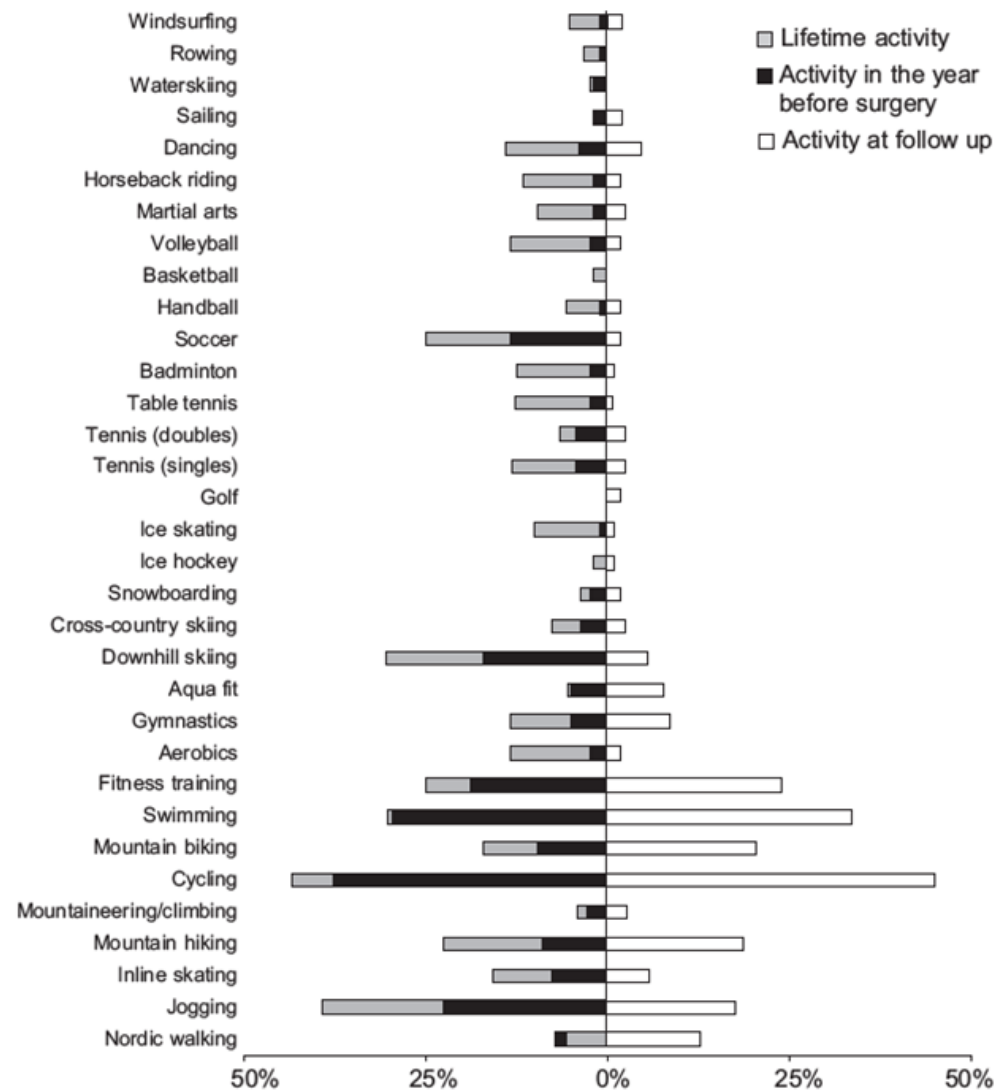
- ⊗ Strengths:
  - ⊗ Can produce hyaline-like cartilage
  - ⊗ No correlation with defect size and clinical results
  - ⊗ Can reduce symptoms and improve function in patients with large defects that are significantly disabled
- ⊗ Limitations:
  - ⊗ Invasive
  - ⊗ Surgical time / Complexity of delivery
  - ⊗ Expense
  - ⊗ Subsequent surgical procedures
  - ⊗ Longer recovery – return to sports











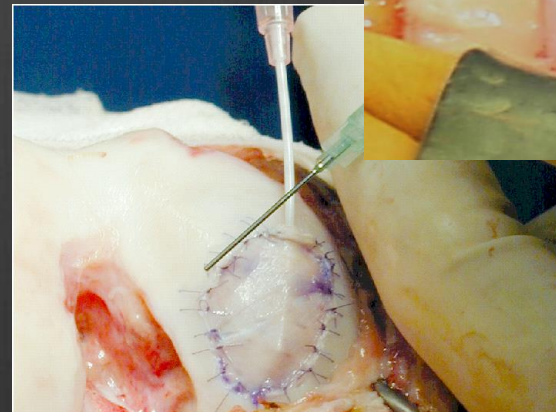
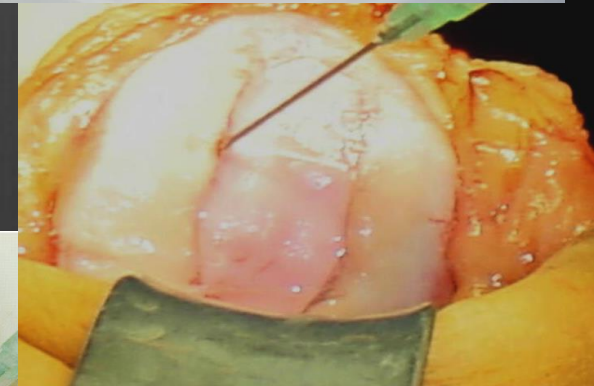
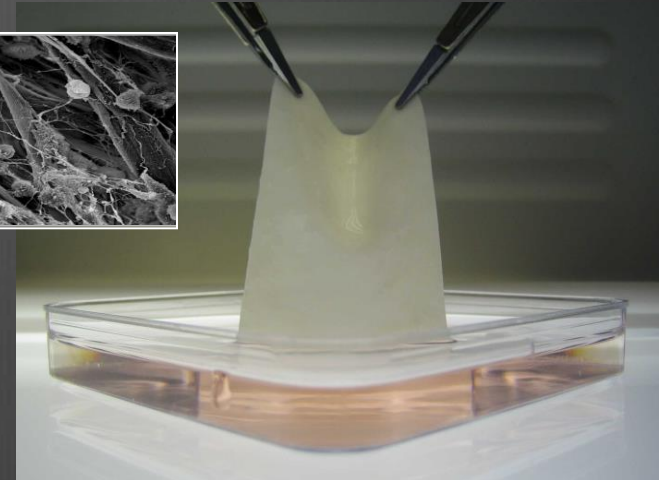
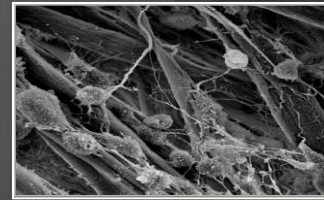
**Figure 1.** Percentage of study patients participating in sports activities throughout lifetime, 1 year before autologous chondrocyte implantation (ACI), and at the time of the follow-up survey.



# MACI

- ❁ “ ON DECEMBER 14,2016, THE FDA HAS APPROVED MACI – AUTOLOGOUS CULTURED CHONDROCYTES ON PORCINE COLLAGEN MEMBRANE – FOR REPAIR OF SINGLE OR MULTIPLE FULL THICKNESS CARTILAGE DEFECTS

# Matrix-Induced Autologous Chondrocyte Implantation MACI<sup>®</sup>

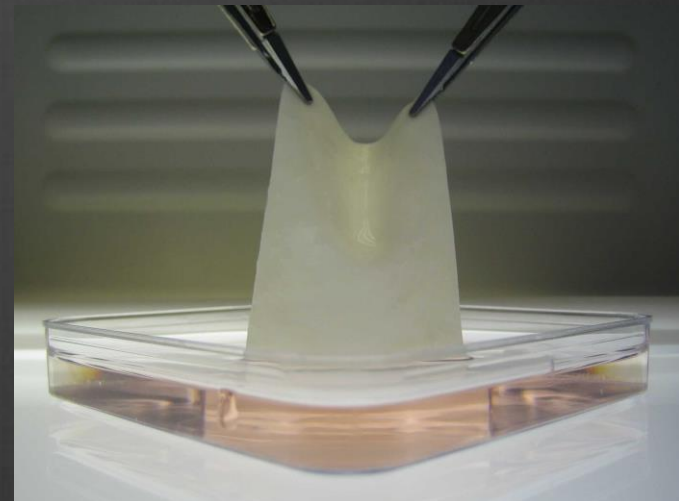
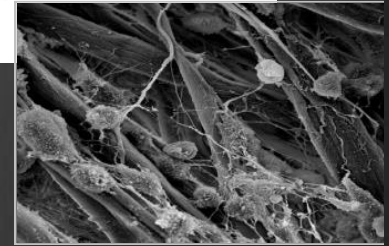
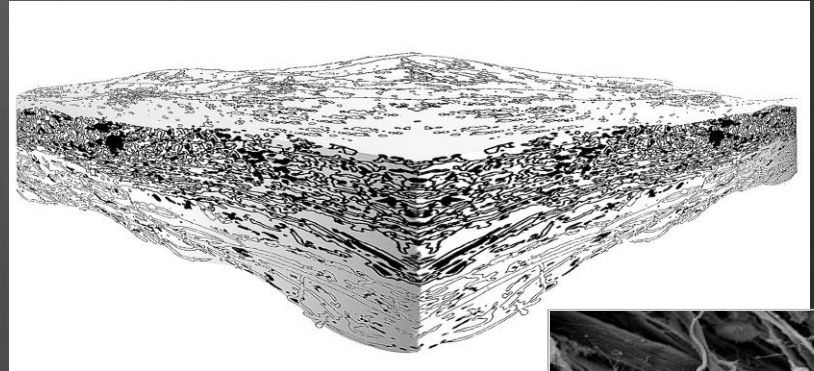


- ⊗ Addresses Cartilage adoption barriers:
  - ⊗ Less morbid delivery
    - ⊗ Arthroscopic in some cases
    - ⊗ Eliminates periosteum and suture fixation
  - ⊗ Reduces operative time
- ⊗ Similar clinical results
  - ⊗ Less post-operative complications
- ⊗ Limitations:
  - ⊗ 2-stage procedure
  - ⊗ Custom product
  - ⊗ Expense

# MACI®

(Matrix-induced Autologous Chondrocyte Implantation)

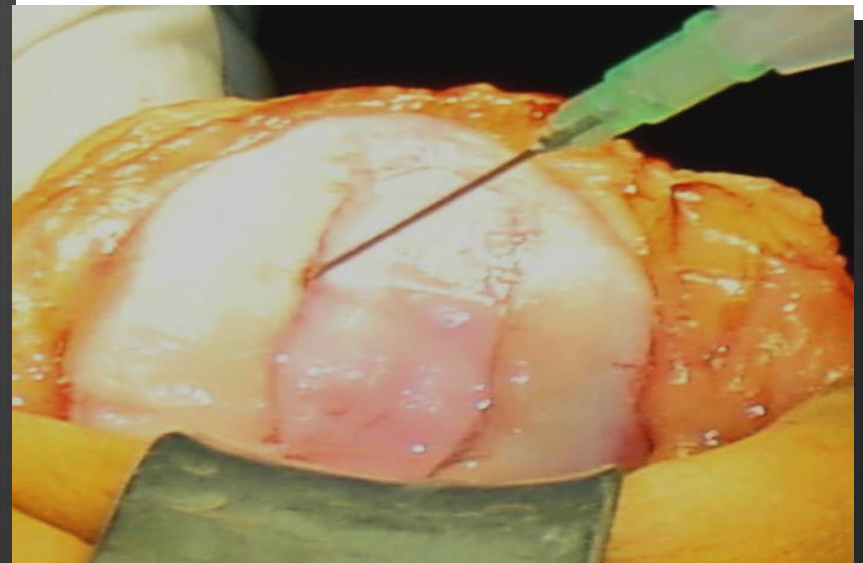
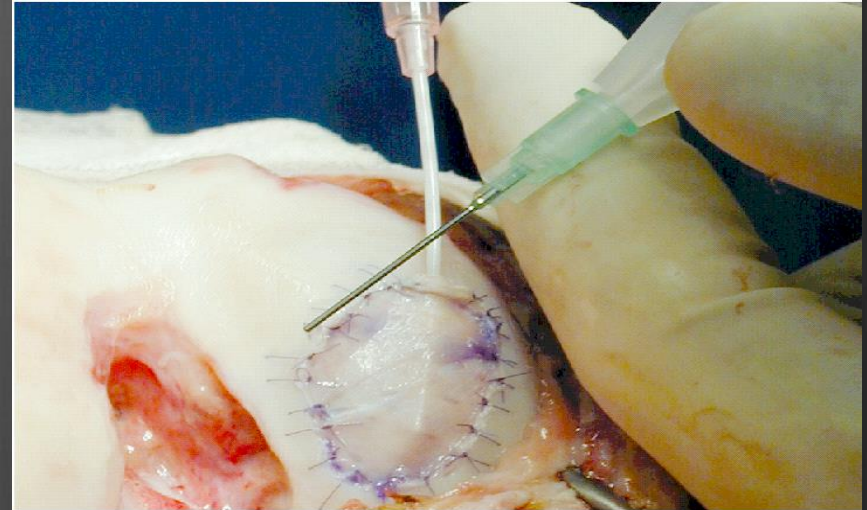
- ❶ Autologous cultured chondrocytes (manufactured similar to Carticel)
- ❷ Seeded on a proprietary, single-ply, Type I & III collagen membrane
- ❸ The *ACI-MAIX*<sup>TM</sup> membrane, is manufactured by Matricel GmbH of Herzogenrath, Germany

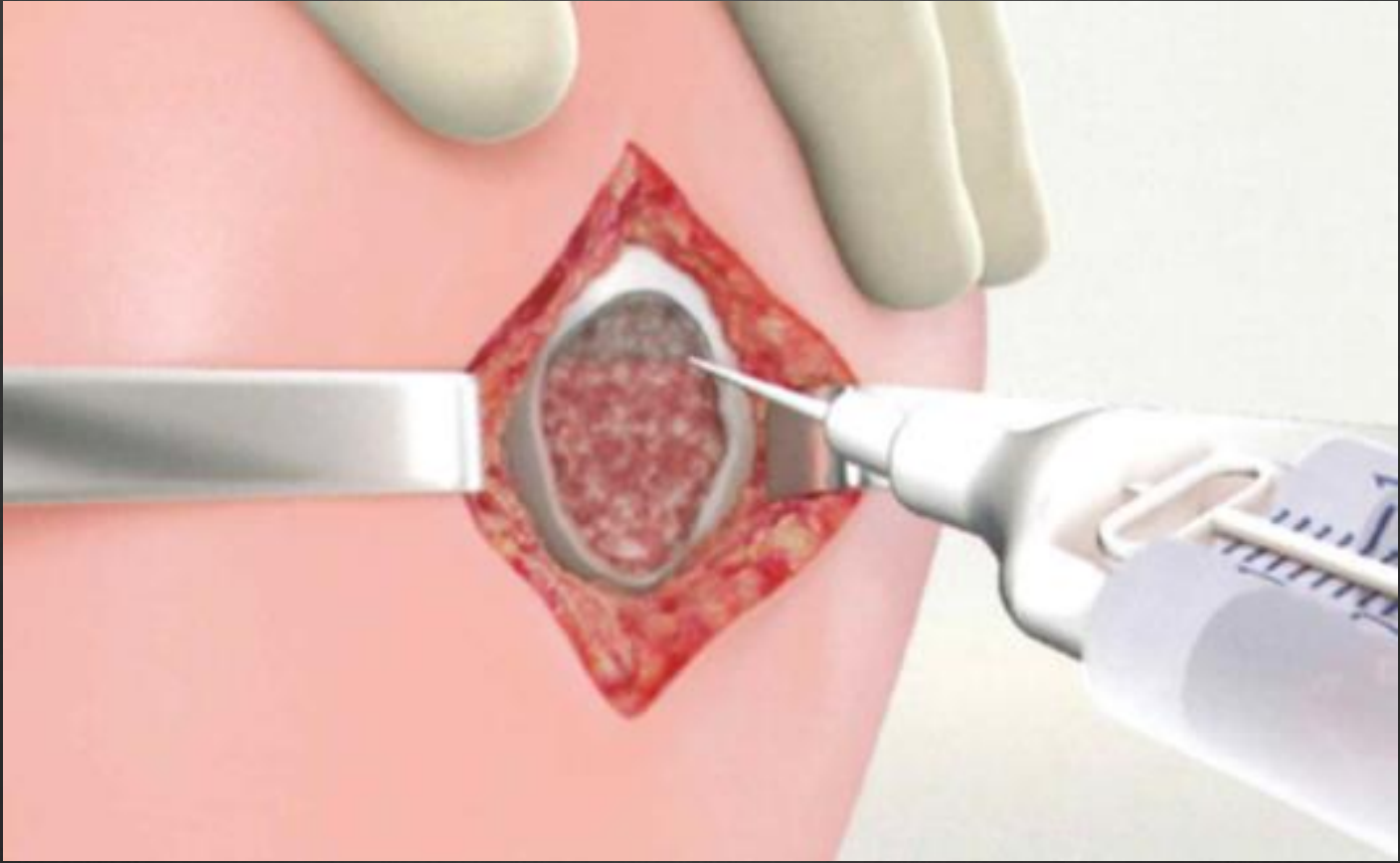


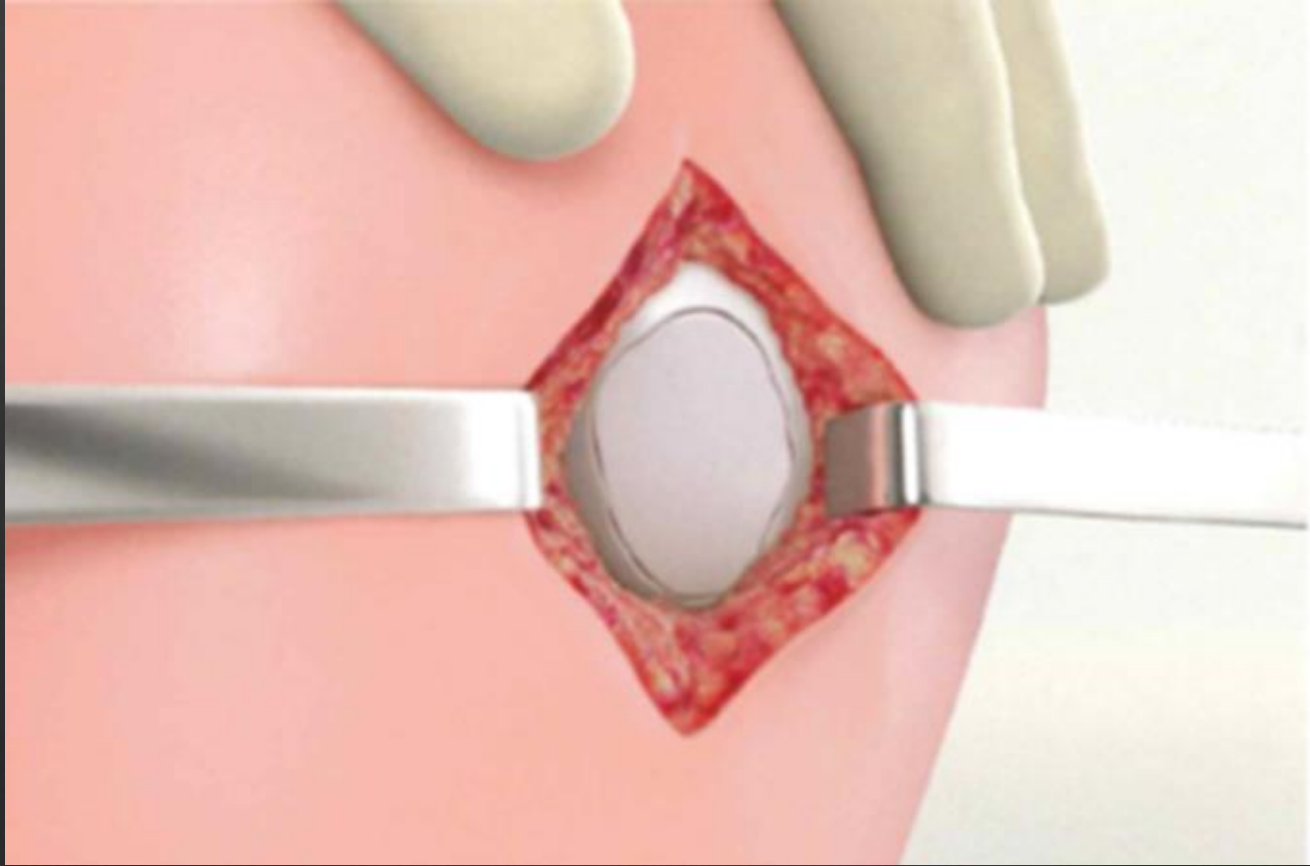
# MACI®

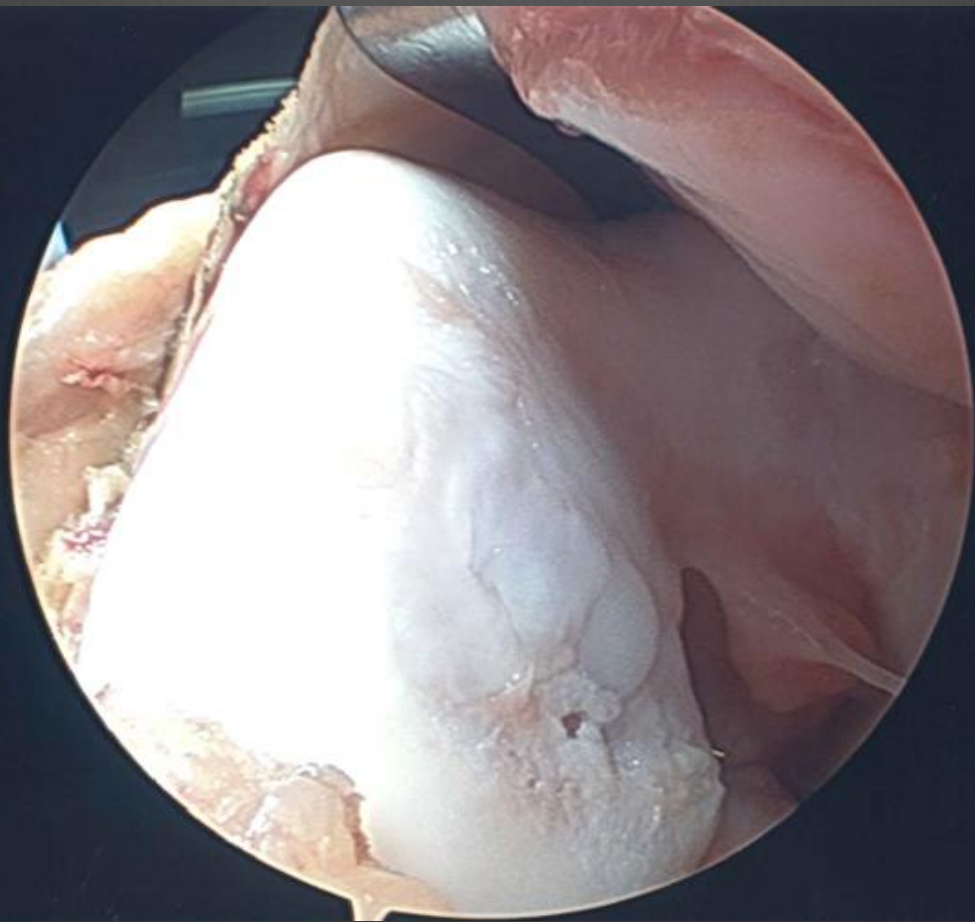
(Matrix-induced Autologous Chondrocyte Implantation)

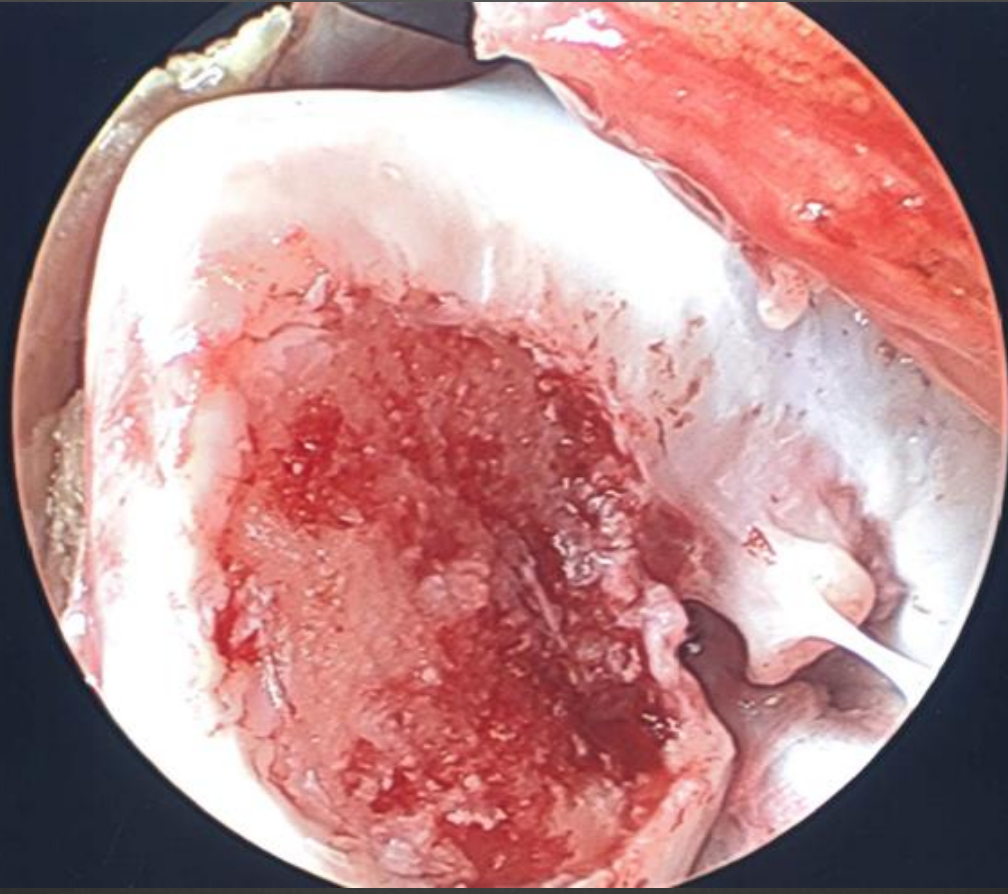
- 2nd generation ACI addresses the many of the limitations / delivery complexities of Carticel:
  - No periosteum harvest
  - No suturing
  - Significantly reduced OR time
- Smaller incision → less patient morbidity
  - Likely to have fewer surgical complications, e.g. tissue hypertrophy, adhesions
- Over 4,000 patients treated since 2001 in Europe & Australia



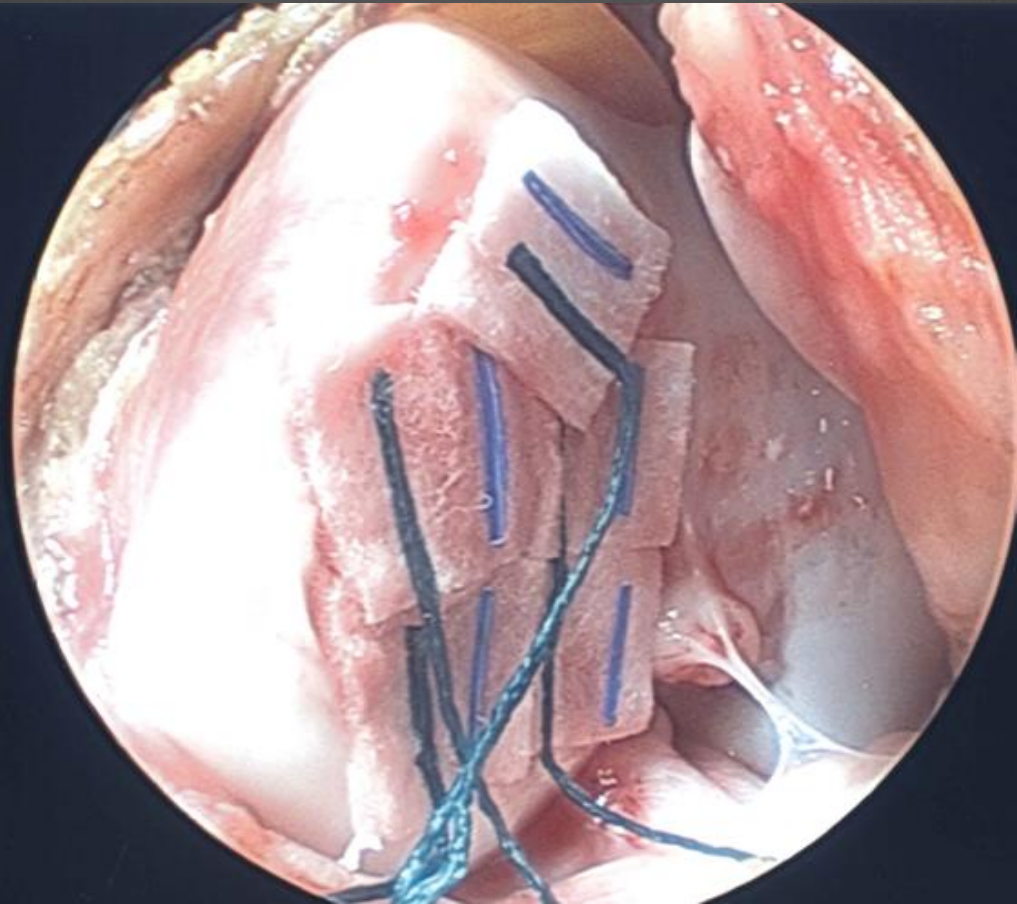


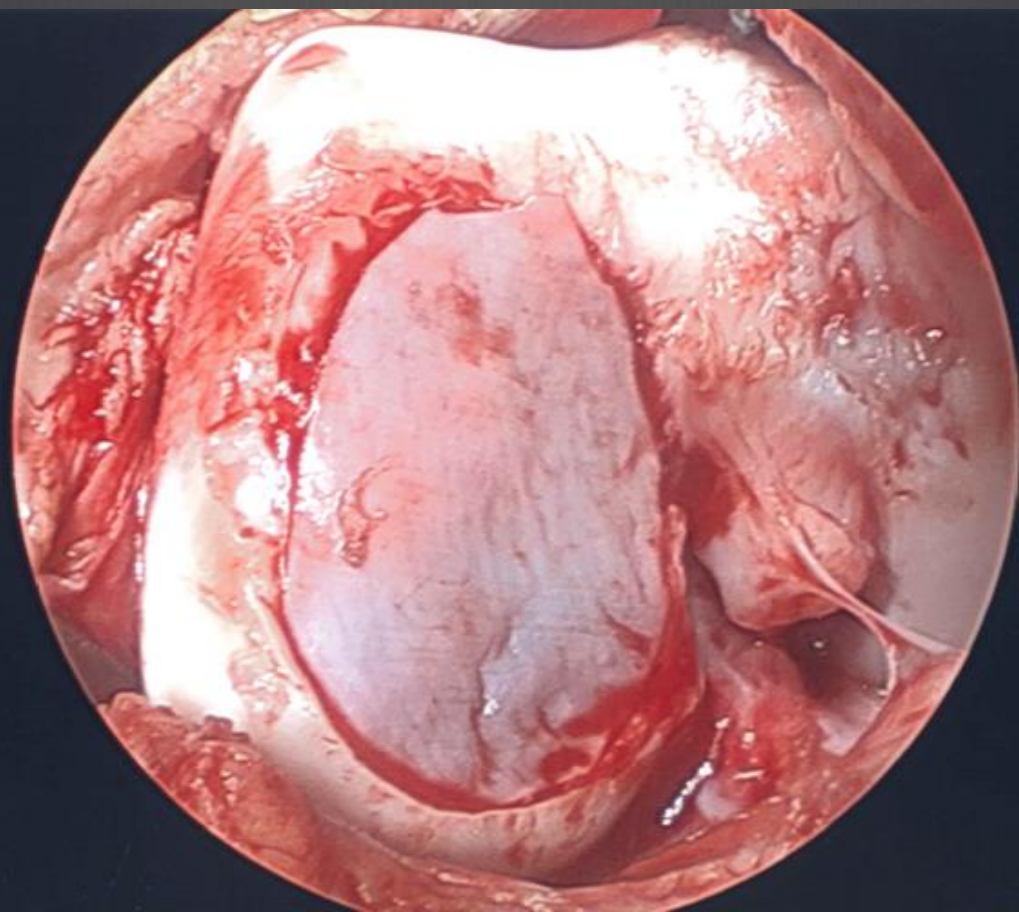


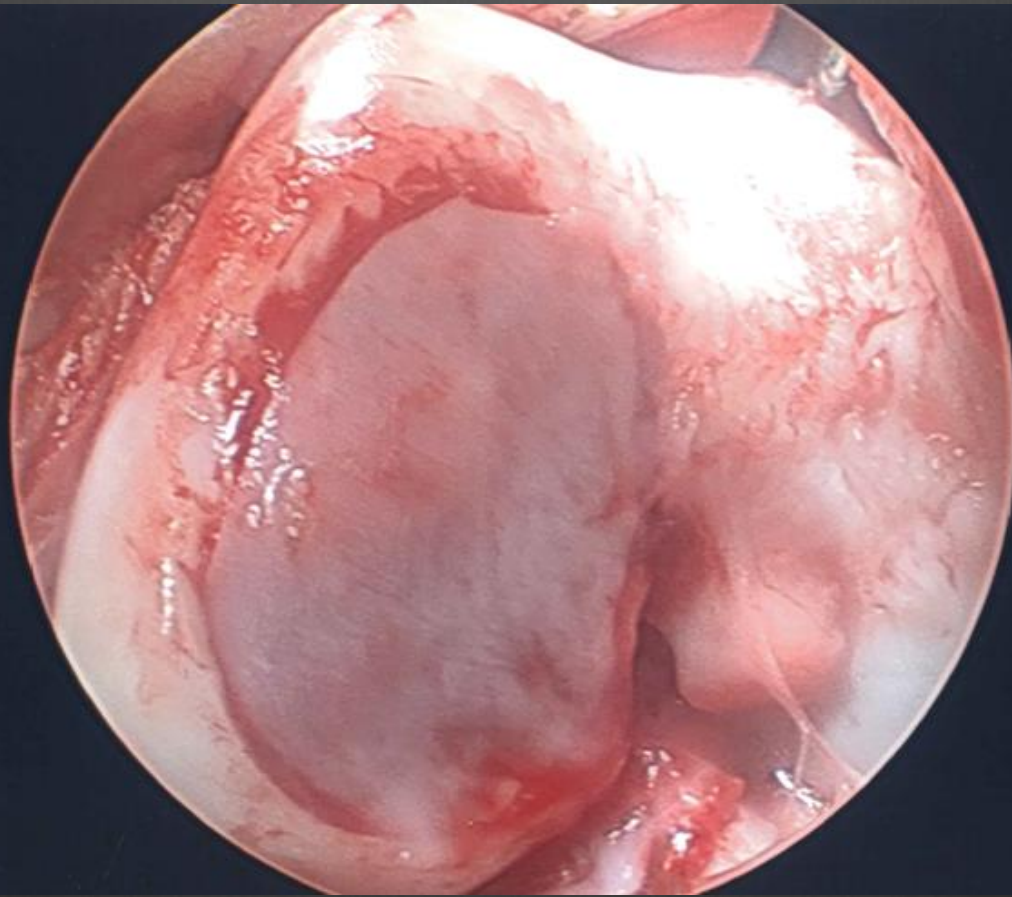






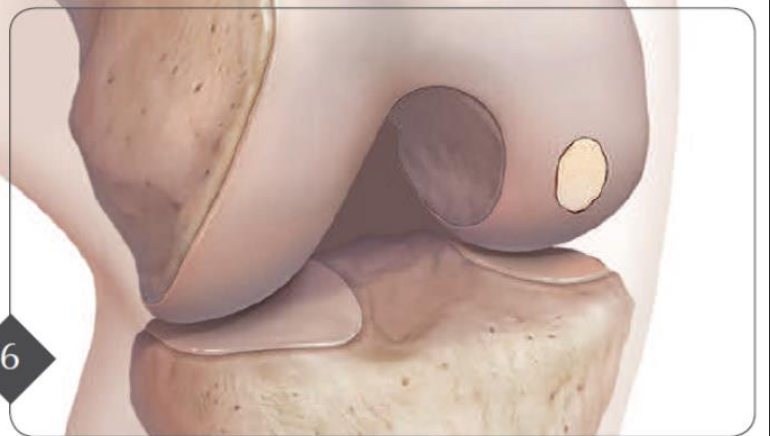
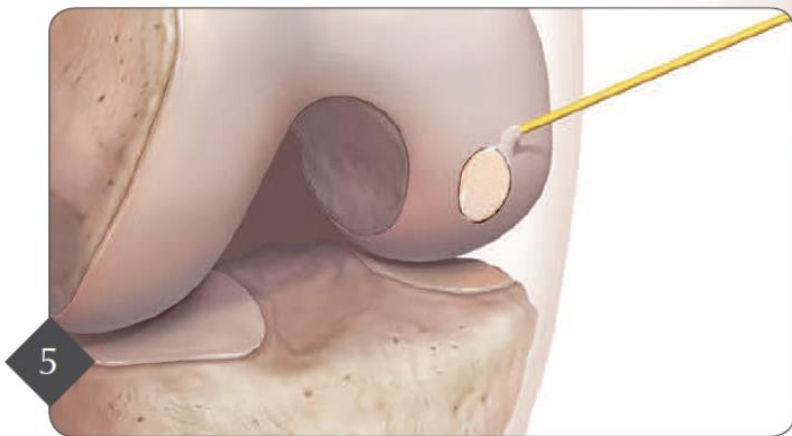
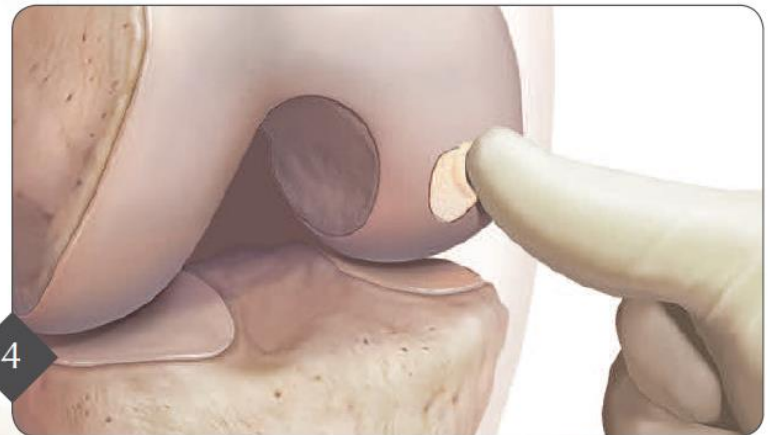
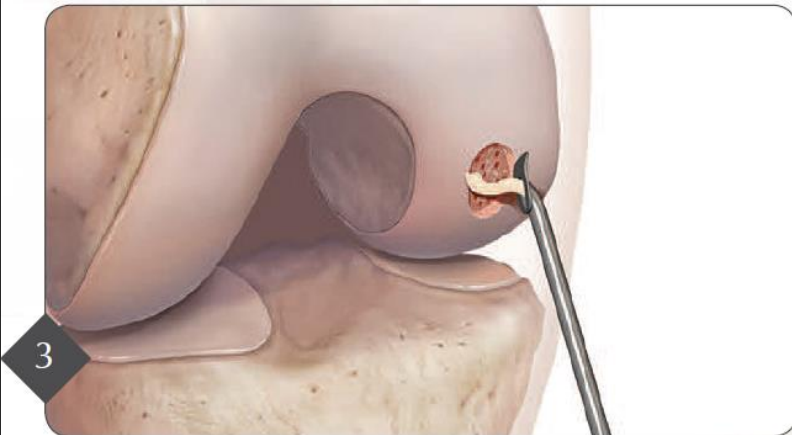
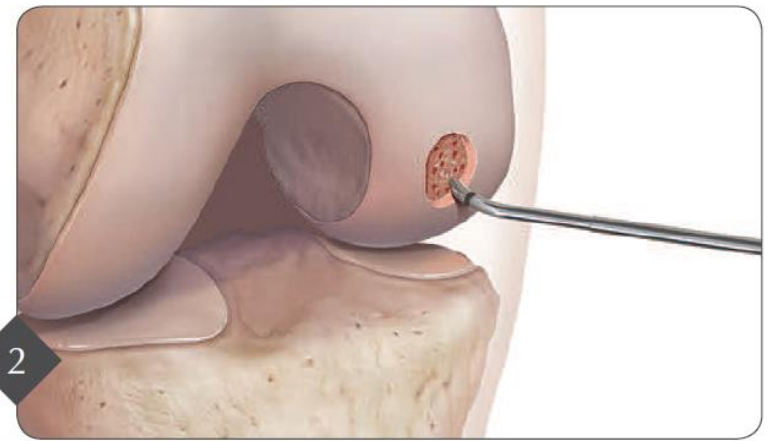
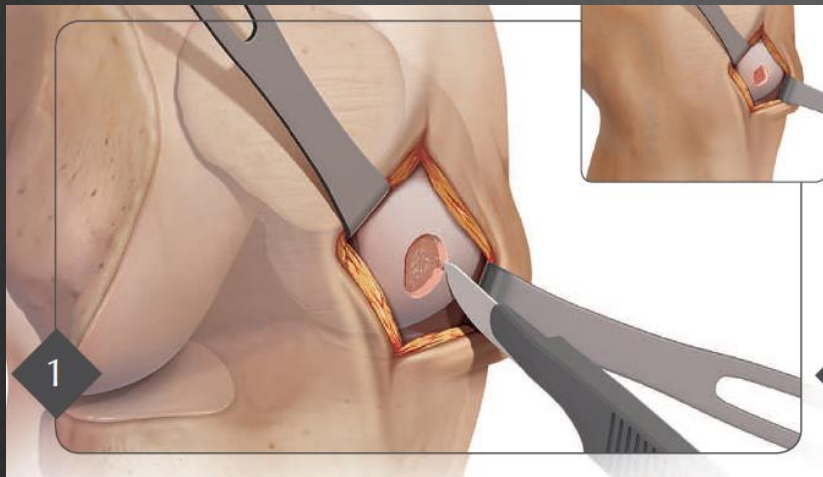






# ACELLULAR MATRIX INDUCED MICROFRACTURE

- ⊗ SCAFFOLD WITH MICROFRACTURE
- ⊗ SINGLE STAGE
- ⊗ KEEPS MESENCHYMAL STEM CELLS IN PLACE
- ⊗ PROMISING RESULTS
- ⊗ LITTLE EXPENSE
- ⊗ ONE STAGE



# RESULTS

# OATS VS MICROFRACTURE

## ⊗ GUDAS ET AL –

- ⊗ AFTER 37 MONTHS – 96% GOOD TO EXCELLENT RESULTS WITH OATS AND 53% GOOD TO EXCELLENT RESULTS WITH MICROFRACTURE
- ⊗ BIOPSY AT 12 MONTHS SHOWED GOOD REPAIR TISSUE IN OATS IN 84% AND GOOD REPAIR TISSUE IN MICROFRACTURE IN 57%

# OATS VS MICROFRACTURE

- ⊗ RETURN TO FOOTBALL
  - ⊗ OATS 91%
  - ⊗ MICROFRACTURE 68%
- ⊗ BOTH HAVE DIMINISHING RESULTS AT 3-10 YEARS
- ⊗ POOR PROGNOSIS IN NBA PLAYERS-MICROFRACTURE



# RETURN TO SPORT

## PERCENT

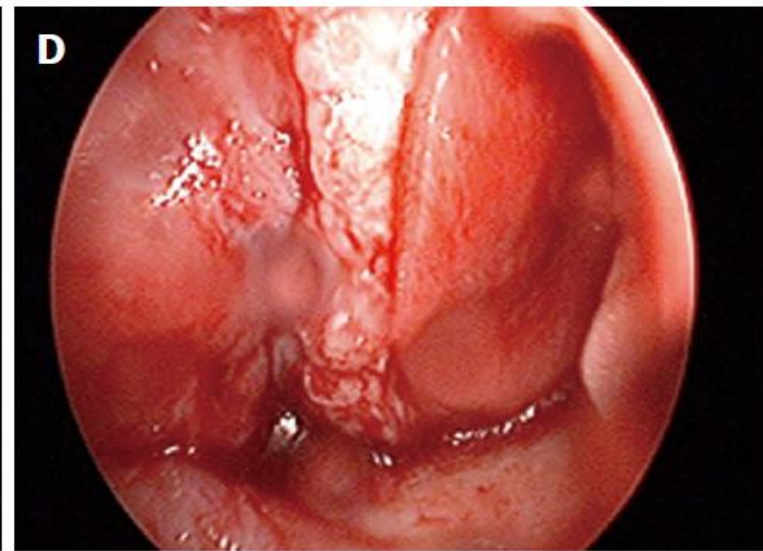
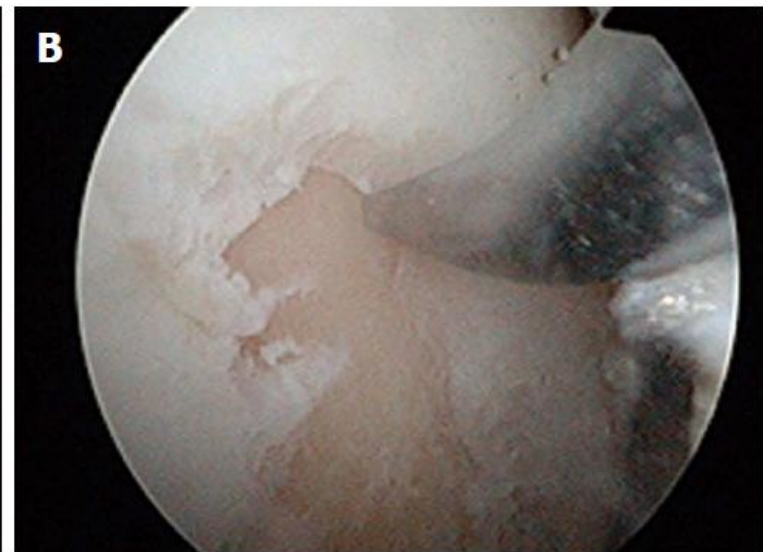
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- OVERALL – 76%
- MFX – 58%
- OATS- 93%
- ACI- 83%
- OCA – 88%

## TIME

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- OVERALL – 9 MONTHS
- MFX – 9.1 MONTHS
- OATS – 5.2 MONTHS
- ACI – 11 MONTHS
- OCA – 9.6 MONTHS



LOOK I KNOW ITS RUPTURED DR. ....  
..... BUT SORELY I'LL STILL BE ABLE TO  
RUN IN THE MARATHON TOMORROW?!!



ANY QUESTIONS.

THE END