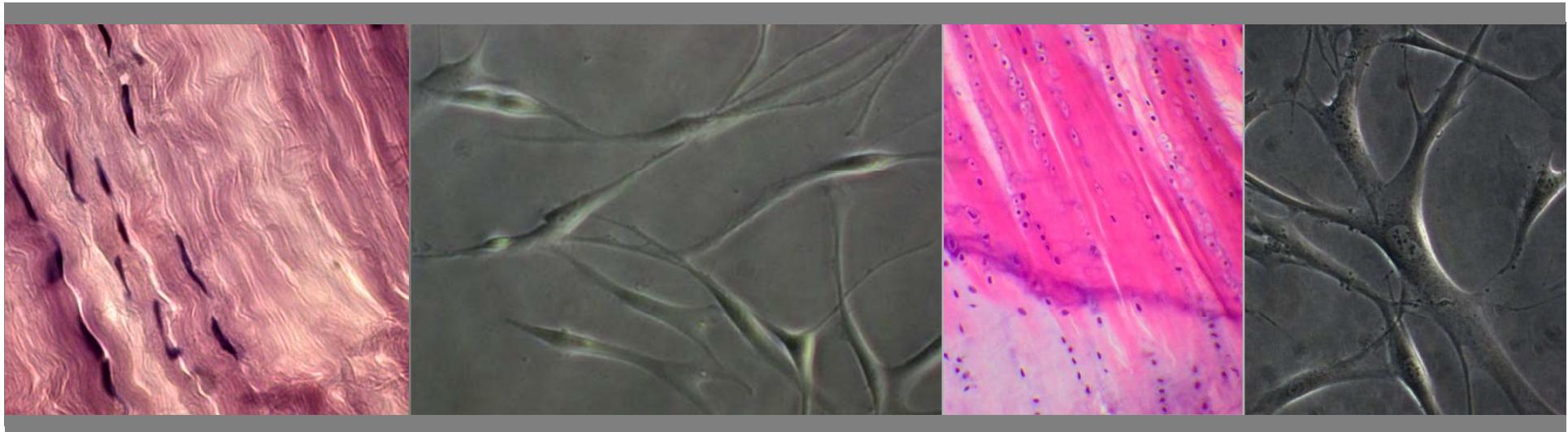


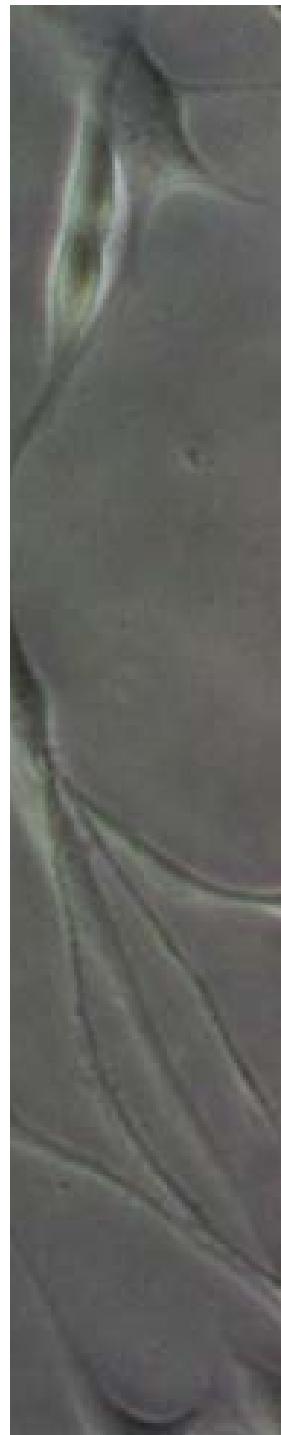
<sup>1,2</sup>Gundula Schulze-Tanzil, VMD

Research group „Bioreconstruction“



<sup>1</sup>*Institute of Anatomy, Paracelsus Medical University, Salzburg and  
Nuremberg, Germany*

<sup>2</sup>*Department of Orthopaedic, Trauma and Reconstructive Surgery, Charité-  
Universitätsmedizin Berlin, Campus Benjamin Franklin, Germany*



### **Education:**

- 1990-1996 School of veterinary medicine, Freie Universität Berlin
- 1999 VMD Degree, Freie Universität Berlin
- 1999 Postdoctoral Fellow, Institute of Anatomy, Freie Universität Berlin
- 2005 Scientific assistant, Centrum of Anatomy, Charité University of Medicine, CCM, Berlin
- 2005-2008 Dep. of Trauma and Reconstructive Surgery, CBF/Centrum of Anatomy, CCM, Charité University of Medicine, Berlin
- 2007 Leader of the research laboratory of experimental surgery, AG Bioreconstruction
- 2008-2014 Dep. of Trauma and Reconstructive Surgery, CBF, Charité University of Medicine, Berlin
- 2009 Habilitation: Anatomy and Cell Biology
- 2015 Professorship at the Institute of Anatomy, Paracelsus Medical University, Salzburg and Nuremberg, Germany

### **Teaching Activities:**

Dissection courses in gross anatomy, courses in histology, seminars and lessons in anatomy and cell biology

### **Scholarships:**

- 1996-1999 Doctoral scholarship, Freie Universität Berlin
- 2005-2008 Rahel-Hirsch habilitation Scholarship, Charité Berlin

# **Ad hoc reviewer for**

Acta Biomaterialia, Annals of Anatomy, Arthritis Rheumatism, Arthritis Research Therapy, Berliner Münchner Tierärztliche Wochenschrift, BMC Musculoskeletal Disorders, Biomaterials, Biotechnology Progress, Biotechnology Bioengineering, Bone, Cell Tissue Research, Cell Biology and Toxicology, Cells Tissues Organs, Clinical Medicine Insights: Arthritis Musculoskeletal Disorders, Clinical Experimental Immunology, Connective Tissue Research, Cytokine, Drug Dev reviews, European Cells Materials, European J Histochemistry, Experimental Biology and Medicine, Fibrogenesis and Tissue Repair, Folia Biologica, Gene Therapy, Histochemistry Cell Biology, Histology Histopathology, International J Molecular Science, J Biochips Tissue Chips, J Cellular Biochemistry, J Tissue Engineering Regenerative Medicine, J Biomedicine and Biotechnology, J Biomed Mater Res Part A, J Immunol Methods, J Orthop Surg Res, J Nanomedicine, Materials Letters, Medicinal Chemistry, Osteoarthritis Cartilage, Oxid Antioxid Med Sci, PLOSone, Review Material Letters, The Lancet, Tissue Engineering Part A.

# Scientific interests

- cartilage and tendon/ligament biology
- osteoarthritis (in vitro and in vivo models)
- tendon injury and healing (in vitro and in vivo models)
- interplay of anti- and proinflammatory cytokines in tendon and cartilage, co-culture models
- cell- and biomaterial based cartilage and tendon repair: cartilage and tendon tissue engineering
- tendon/ligament ageing
- comparative anatomy/cell biology of tendons and cartilage subtypes



# Techniques

## Cell biology

- characterization of human and animal-derived primary cells: chondrocytes, tenocytes, ligament cells, synovial fibroblasts, PBMCs and neutrophils
- three-dimensional cultures (pellet, organoid, spheroids, hydrogel and scaffold cultures, statical and dynamical seeding of biomaterials and self prepared cell-freed extracellular matrices
- interaction of musculoskeletal cells in co-cultures
- immunohistochemistry/-cytochemistry, immunofluorescence microscopy
- confocal laser scanning microscopy
- flow cytometry
- histology/histopathology of musculoskeletal tissues
- transmission-/scanning electron microscopy
- ELISA, assays (cytokines, proliferation, DMMB-assay, cell vitality, cytotoxicity, caspase activity, TUNEL...)
- SDS-PAGE and Western Blot Analyses
- RT PCR and Real Time (RTD)-PCR
- transfection of primary cells: adenoviral, lipofection, Electroporation
- gene silencing using siRNA

# Techniques



## Animal models

- matrix assisted autologous chondrocytes transplantation in the minipig and rabbit (chondral vs. Osteochondral defects)  
*Lohan A et al., 2014*
- partial Achilles tendon defect model in the rabbit (Stoll C et al., 2011, Biomaterials)
- nude mice xenograft model: *in vivo* Chondrogenesis and tendogenesis  
*Lohan A et al., 2011: Histochem Cell Biol*
- osteoarthritis model in the rat (surgically induced: MMT)

## gross anatomical preparation techniques of human and animal-derived samples

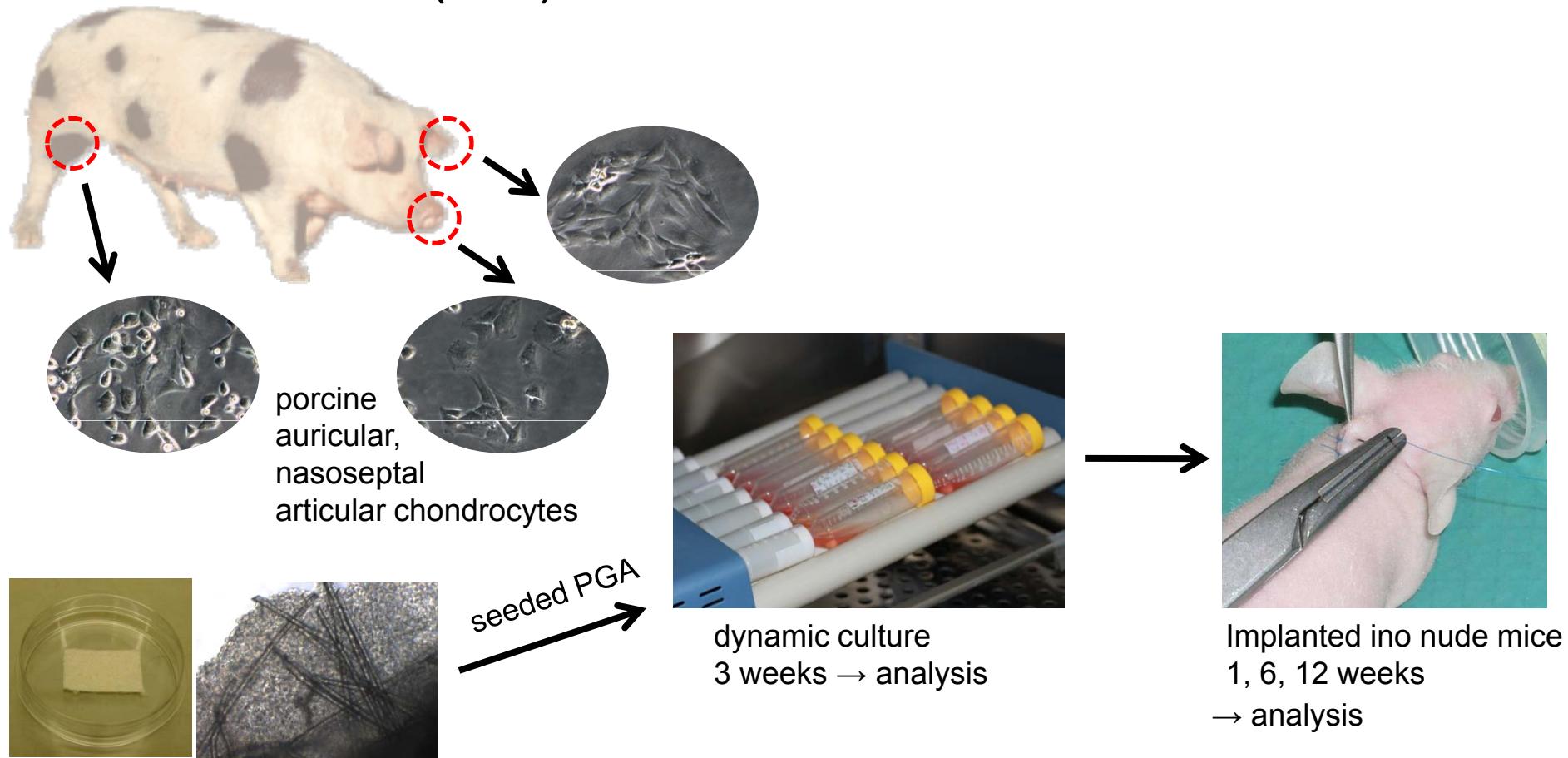


# Some recent publications (2014)

- Mrosewski I, Jork N, Gorte K, Conrad C, Wiegand E, Kohl B, Ertel W, John T, Oberholzer A, Kaps C, Schulze-Tanzil G (2014). Regulation of OA associated key mediators by TNF $\alpha$  and IL-10: Effects of IL-10 overexpression in human synovial fibroblasts and a synovial cell line. **Cell Tissue Res** 357(1):207-23.
- The Phantom 5 Consortium (2014). A promoter level mammalian expression atlas. **Nature** 27;507(7493):462-70.
- Gröger D, Kerschnitzki M, Weinhart M, Schneider T, Kohl B, Wagermaier W, Schulze-Tanzil G, Fratzl P, Haag R (2014). Selectivity in Bone Targeting with Different Polyanionic Dendritic Dye Conjugates. **Advanced Healthcare Materials** 3(3):375-85.
- Lohan A, Marzahn U, El Sayed K, Haisch A, Müller RD, Kohl B, Stölzel K, Ertel W, John T, Schulze-Tanzil G (2014). Osteochondral cartilage defect repair using autologous orthotopic and heterotopic chondrocytes in the rabbit model. **Ann Anat.** 196(5):317-26.
- Girke G, Kohl B, Busch C, John T, Godkin O, Ertel W, Schulze-Tanzil G (2014). Tenocyte activation and regulation of complement factors in response to in vitro cell injury. **Mol Immunol.** 60(1):14-22.
- Stölzel K, Schulze-Tanzil G, Olze H, Schwarz S, Feldmann EM, Rotter N (2014). Immortalised human mesenchymal stem cells undergo chondrogenic differentiation in alginate and PGA/PLLA scaffolds. **Cell Tissue Bank.** 2014 May 16
- Hoyer M, Meier C, Breier A, Hahner J, Heinrich G, Drechsel N, Meyer M, Rentsch C, Garbe LA, Ertel W, Lohan A, **Schulze-Tanzil G**. In vitro characterization of self-assembled anterior cruciate ligament cell spheroids for ligament tissue engineering. **Histochem Cell Biol.** 2014 Sep 26.
- Jagielski M, Wolf J, Marzahn U, Völker A, Lemke M, Meier C, Ertel W, Godkin O, Arens S, **Schulze-Tanzil G** (2014). The influence of IL-10 and TNF $\alpha$  on chondrogenesis of human mesenchymal stromal cells in three-dimensional cultures. **Int J Mol Sci.** 15(9):15821-44.
- Hoyer M, Drechsel N, Meyer M, Meier C, Hinüber C, Breier A, Hahner J, Heinrich G, Rentsch C, Garbe LA, Ertel W, **Schulze-Tanzil G**, Lohan A. (2014). Embroidered polymer-collagen hybrid scaffold variants for ligament tissue engineering. **Mater Sci Eng C Mater Biol Appl.** 1;43:290-9.

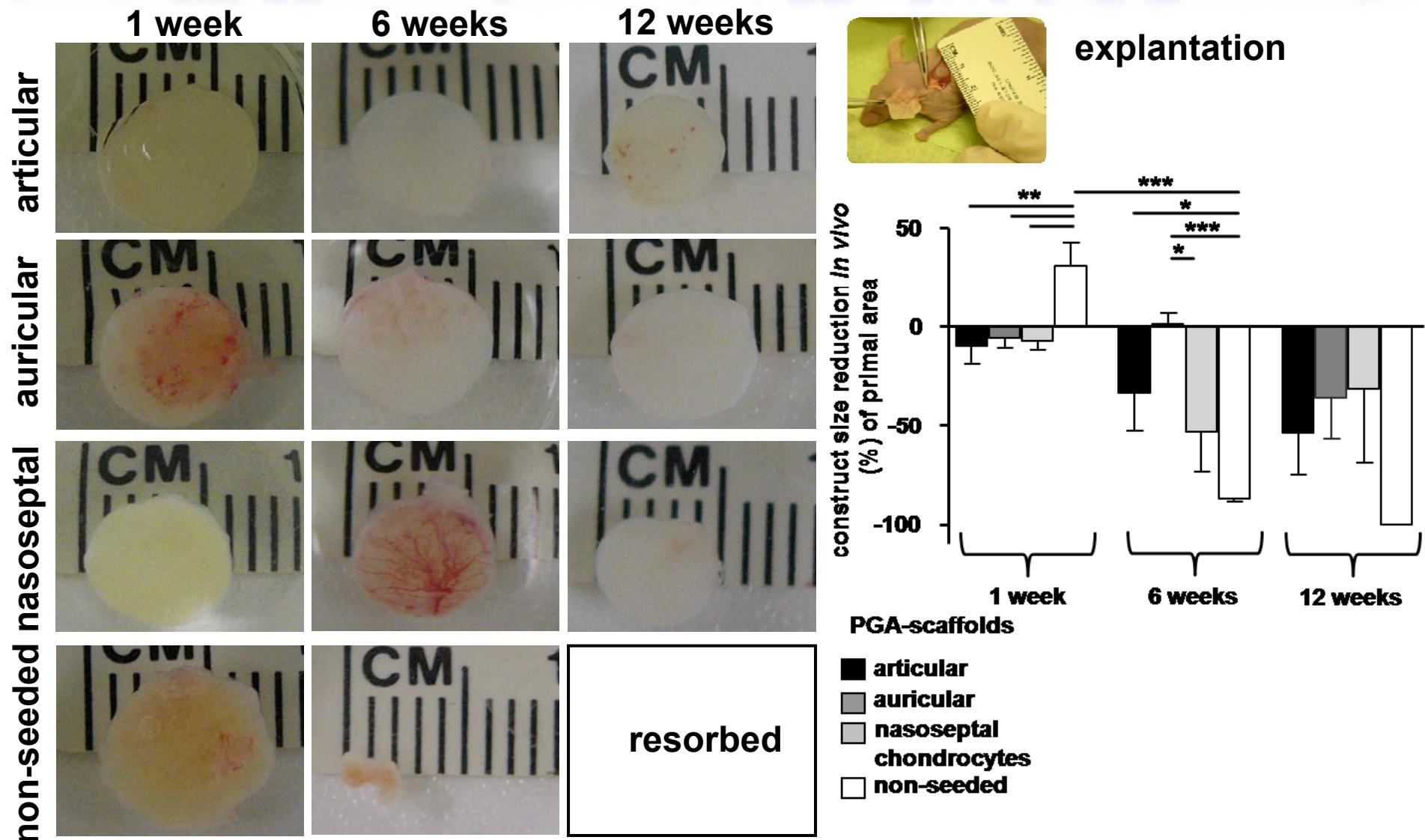
# Cartilage tissue engineering

to characterize /compare the quality of tissue engineered cartilage produced by heterotopic chondrocytes seeded on polyglycolic acid (PGA) scaffolds *in vitro* and *in vivo*

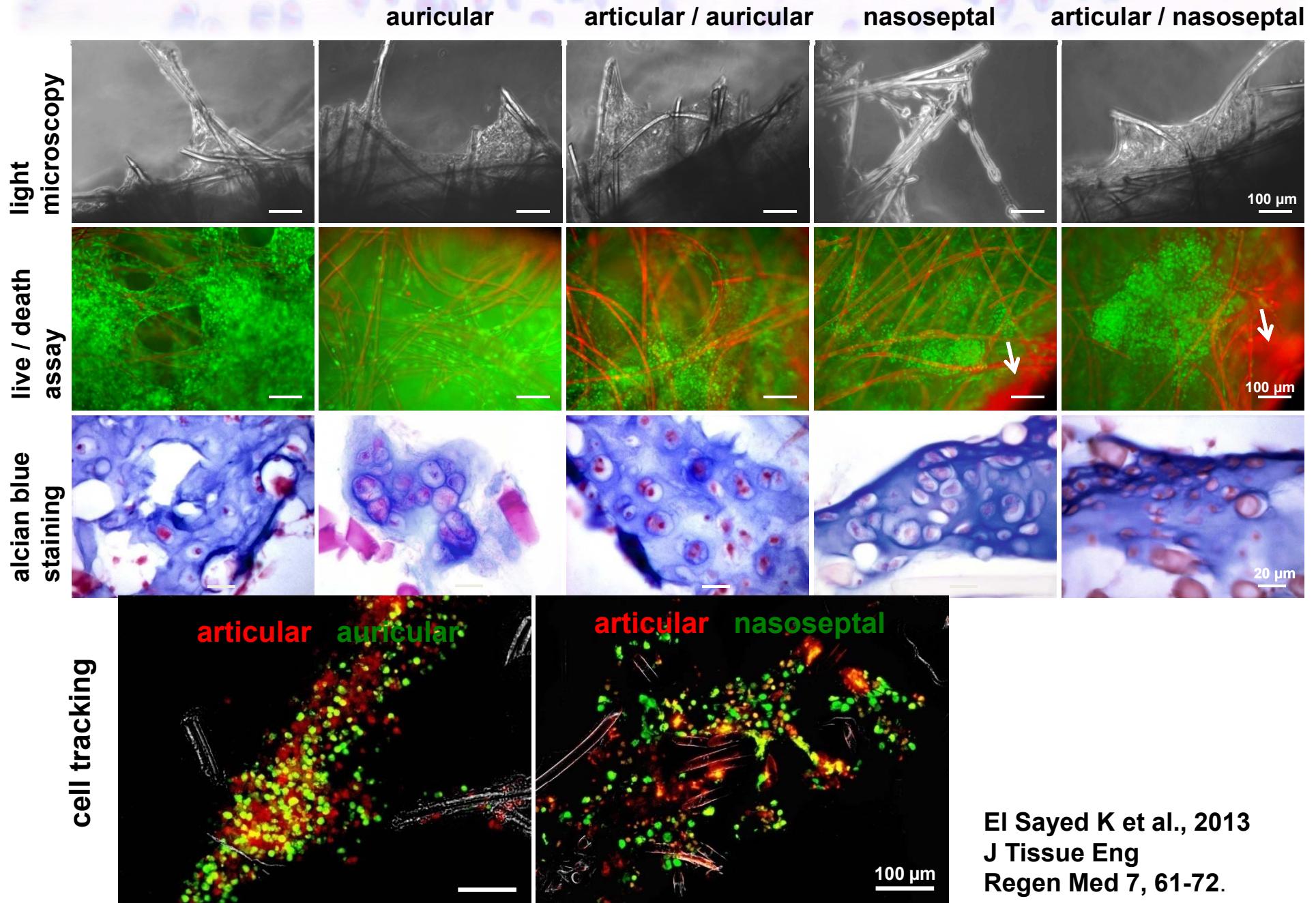


Lohan A et al., 2011  
Histochem Cell Biol. 136(1):57-69

# Chondrogenesis by heterotopic chondrocytes *in vivo*

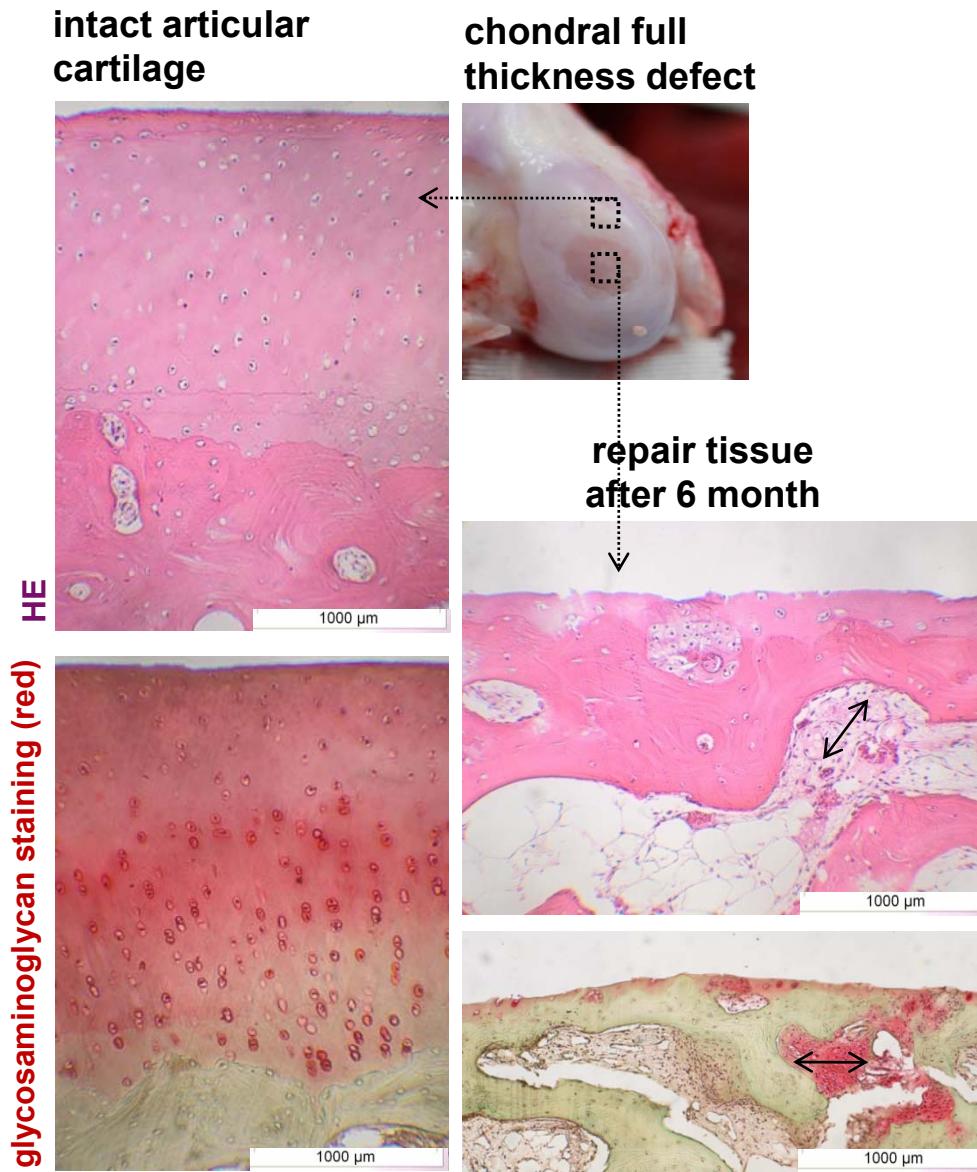


# Chondrogenesis by heterotopic chondrocyte mono- and cocultures



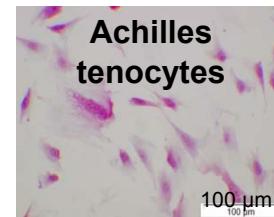
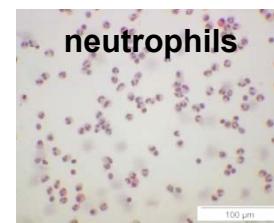
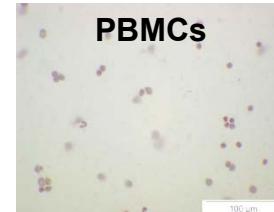
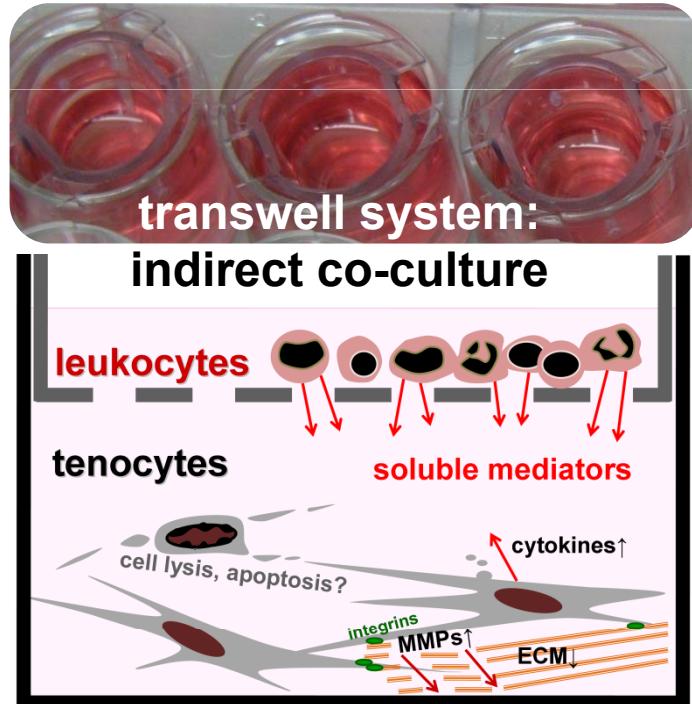
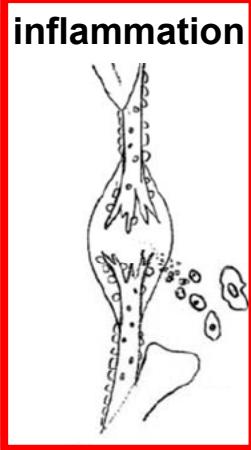
EI Sayed K et al., 2013  
J Tissue Eng  
Regen Med 7, 61-72.

# Cartilage repair *in vivo*



Lohan A et al., 2013, 195(5):488-97.

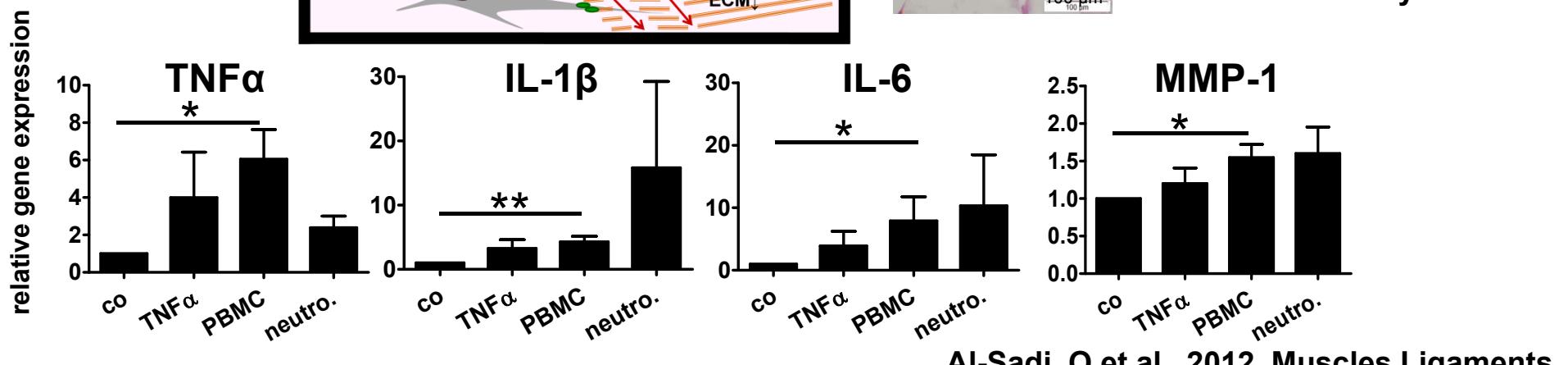
# influence of autologous leukocytes on tenocytes?



## tenocyte response

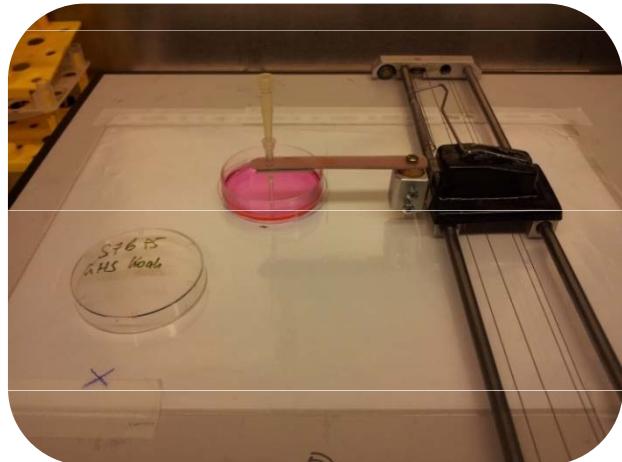
- to cytokines?
- to leukocytes?
- to complement activation?
- to cell injury?

10 ng/mL TNF $\alpha$ , 24 h  
4x10<sup>6</sup> rabbit leukocytes  
10<sup>5</sup> rabbit tenocytes

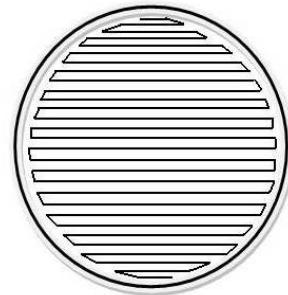


Al-Sadi O et al., 2012. Muscles Ligaments Tendons J. 2012 1(3):68-76.

# tenocyte response to cell injury

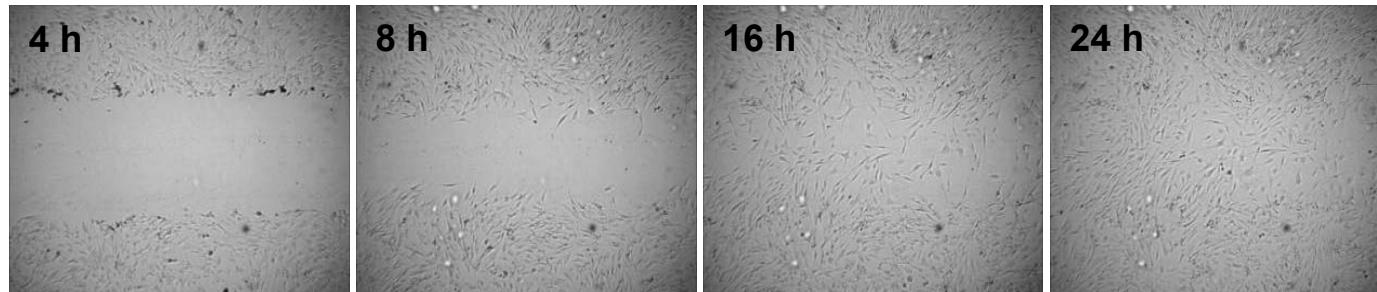


scratch assay

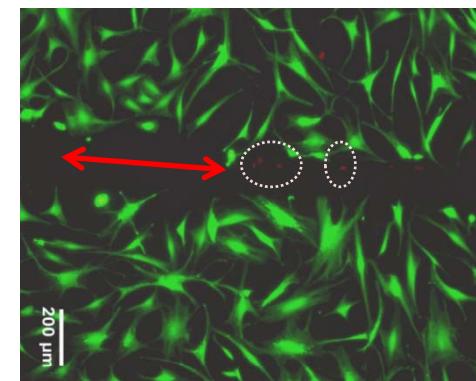
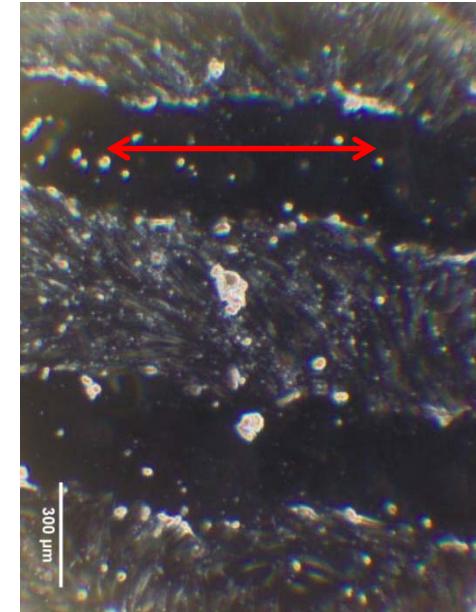


computer guided reproducible plotting system

„healing“ after injury

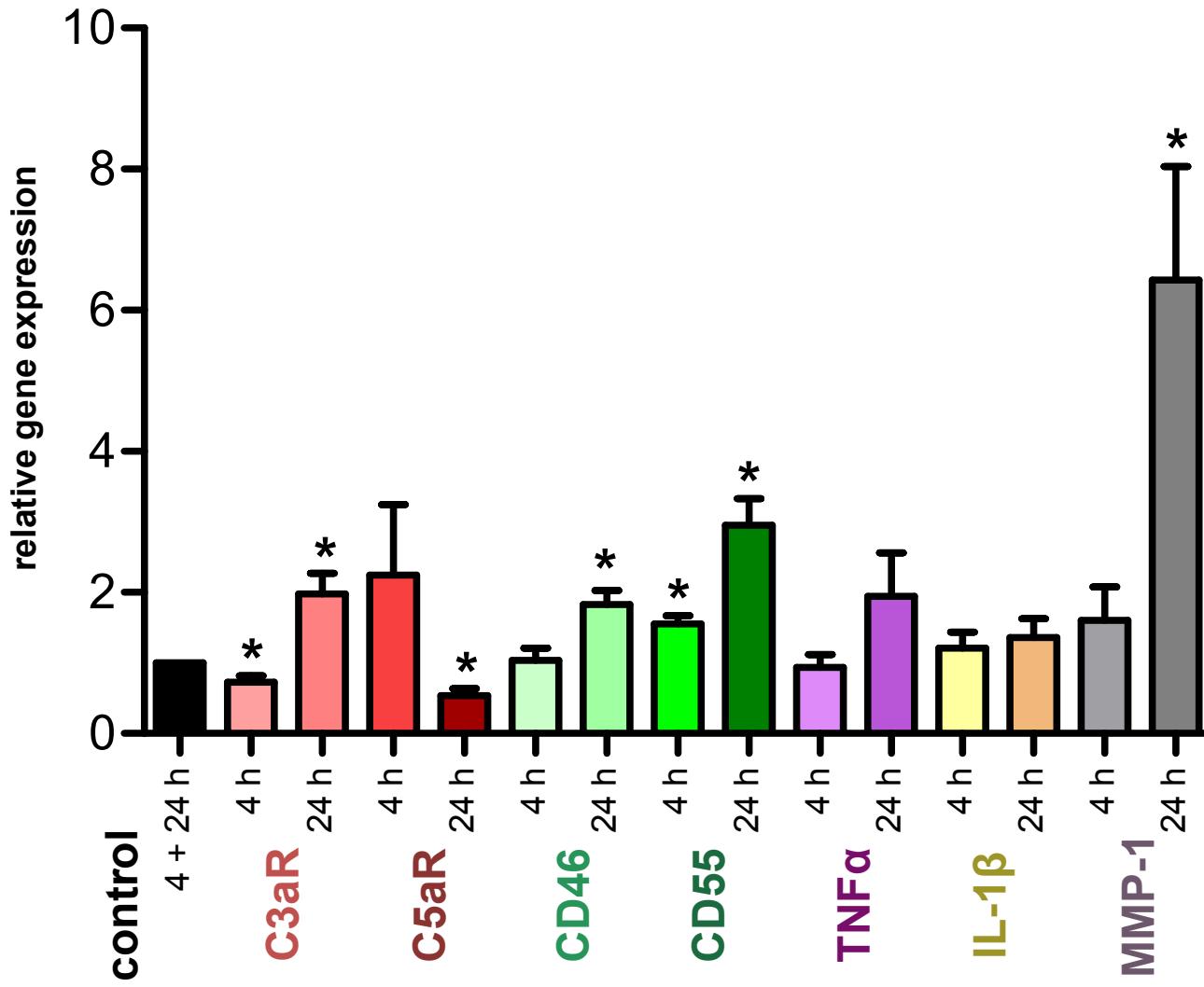


human Hamstring tenocytes



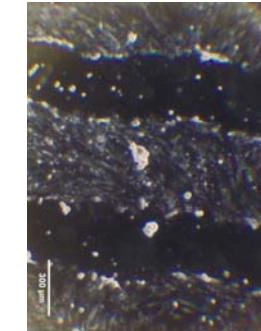
green: vital, red: dead cells

# tenocyte response to cell injury



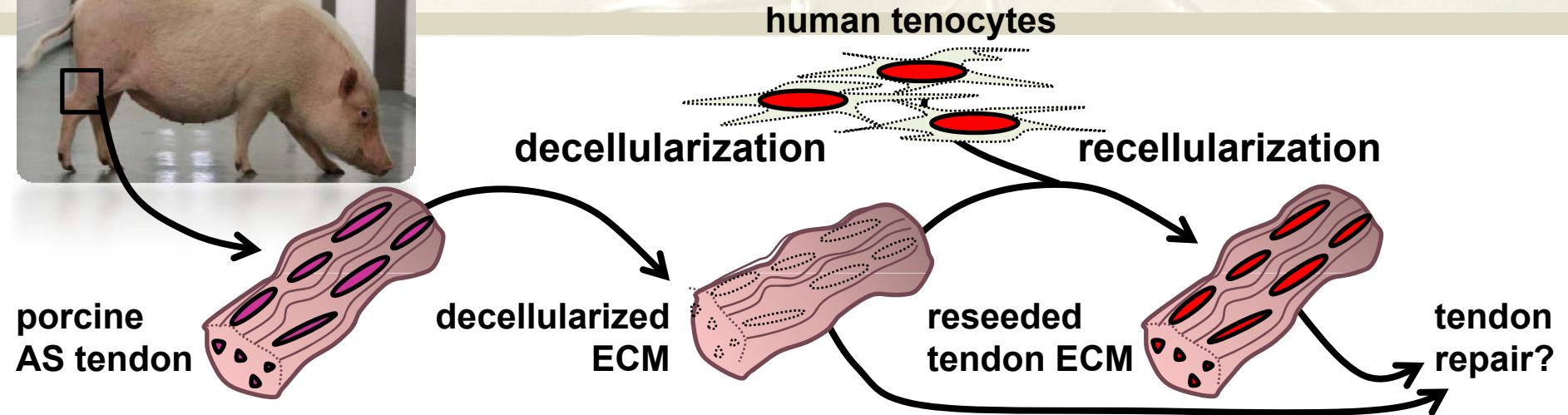
**tenocyte response**

- to cytokines?
- to leukocytes?
- to complement activation?
- to cell injury?



human Hamstring tenocytes

# decellularized tendon ECM scaffolds

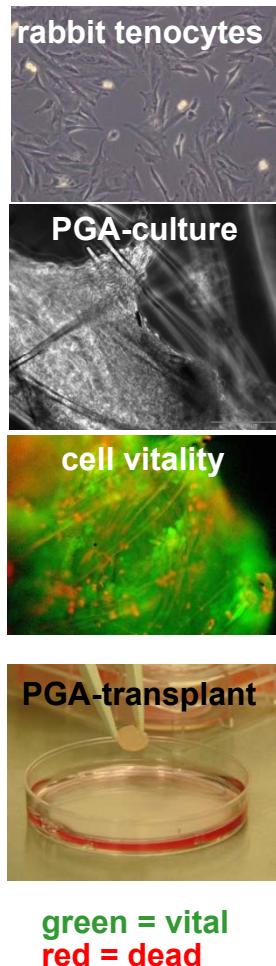
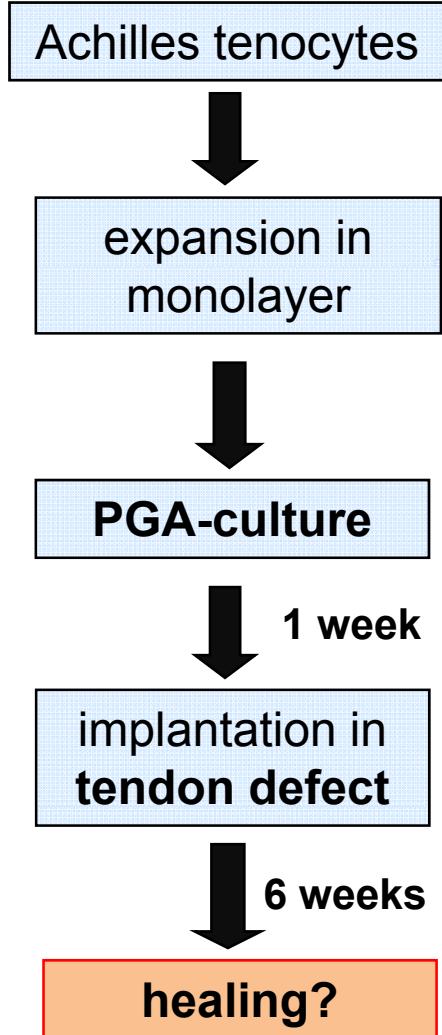


HE

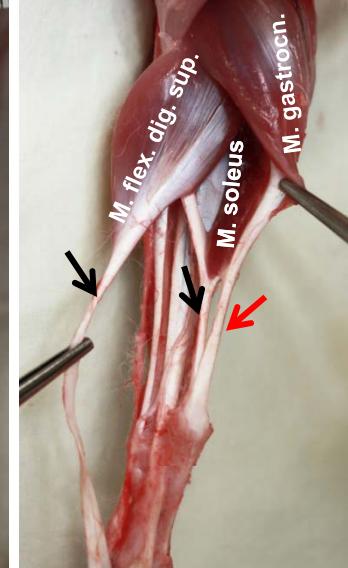


Lohan et al., 2013, Schulze-Tanzil G et al., 2012  
Connect Tissue Res. 54(4-5):305-12. Cells 1:1010.

# tenocyte implantation in partial tendon defects (rabbit)



rabbit Achilles tendon



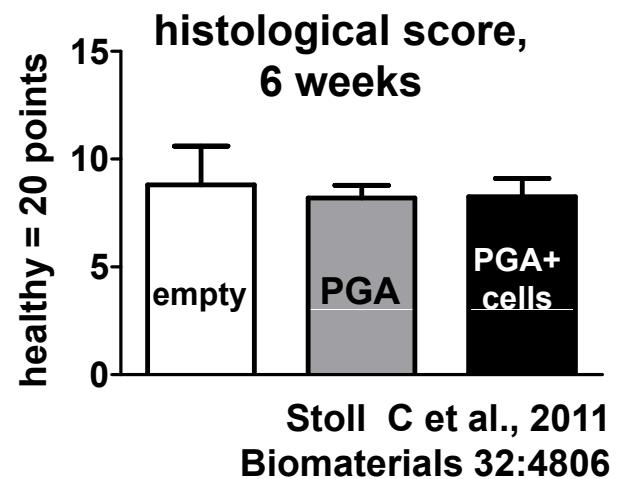
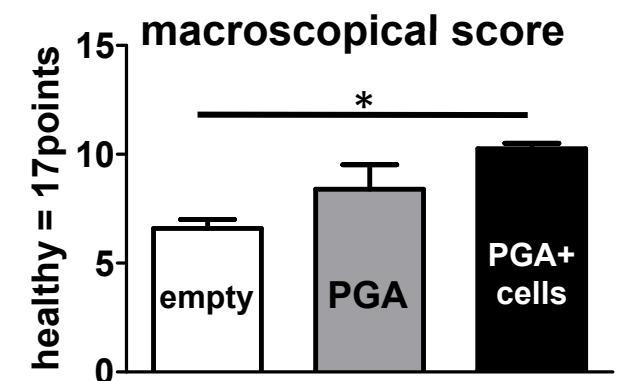
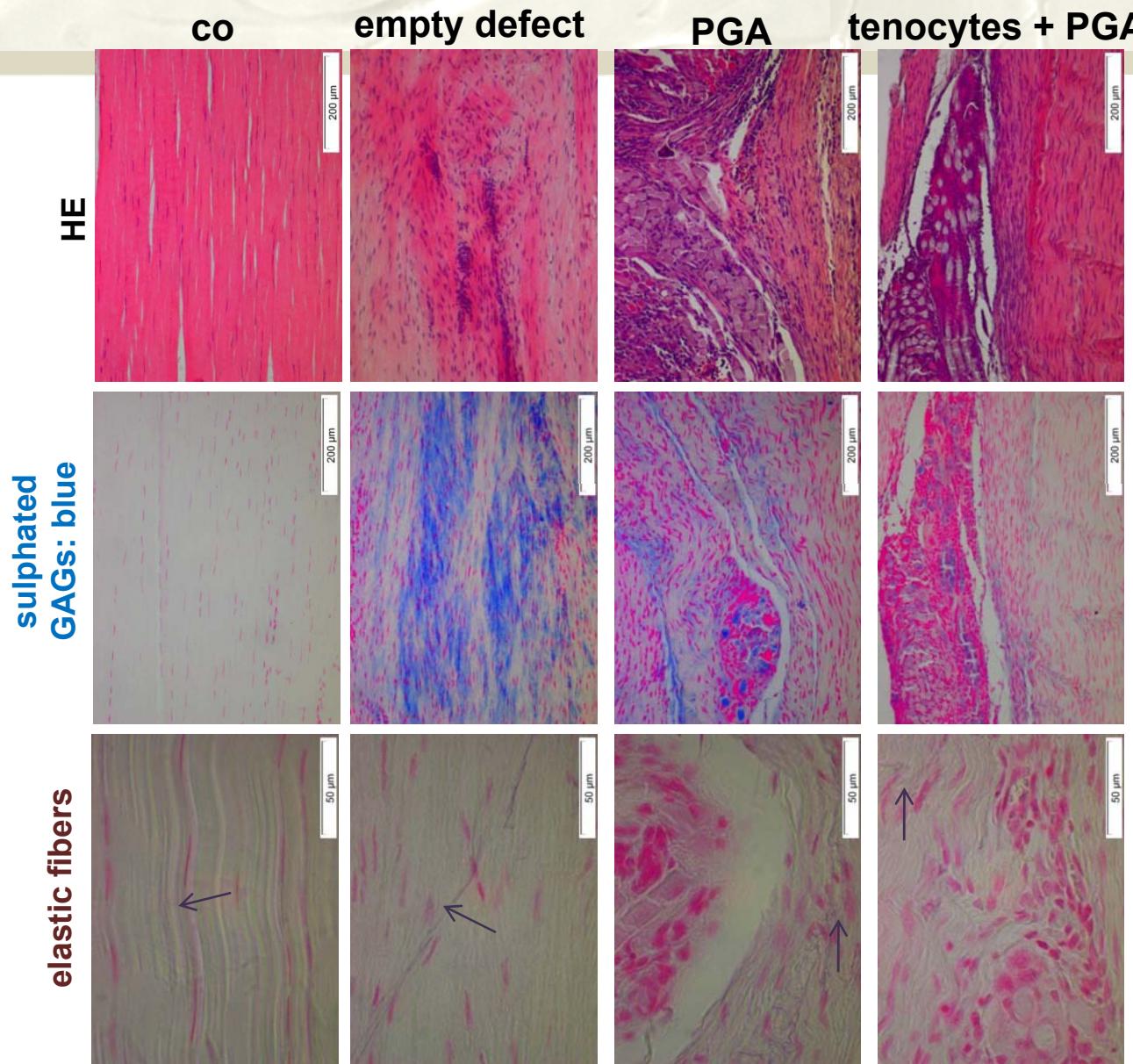
medial *M. gastrocnemius* tendon



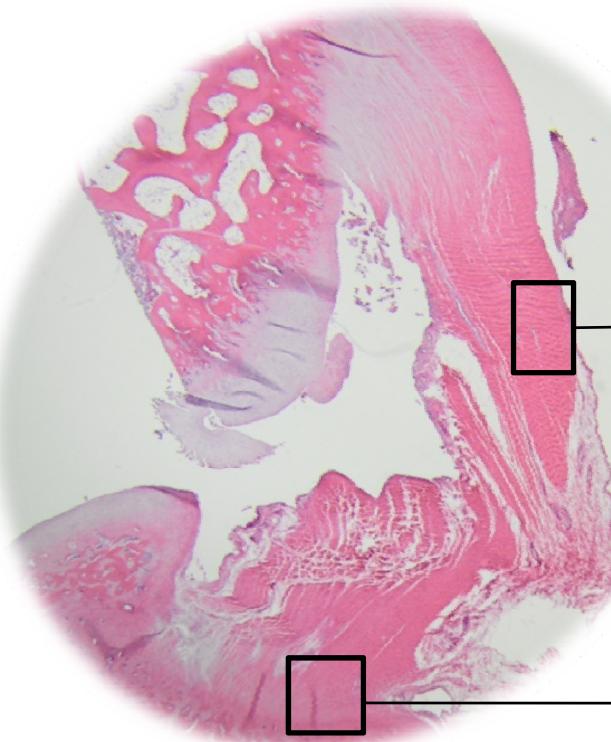
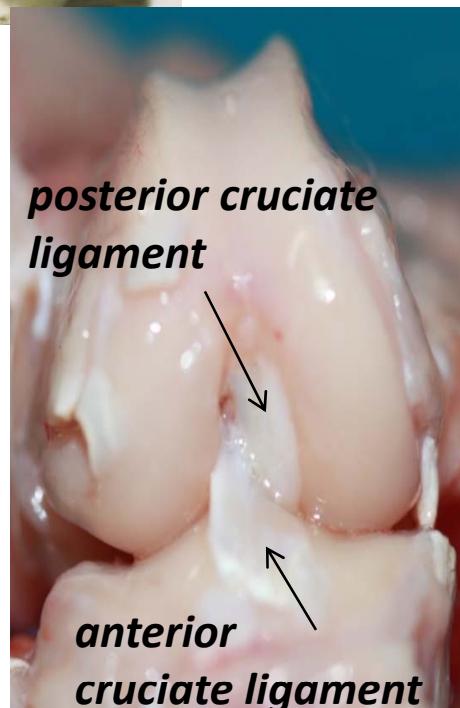
macroscopical and histological scoring

Stoll C et al., 2011  
Biomaterials 32:4806

# tenocyte implantation in partial tendon defects (rabbit)

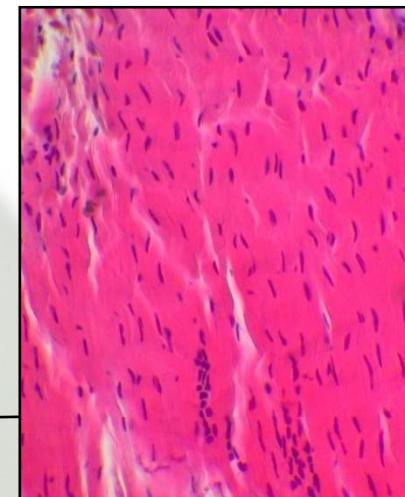


# Lapine anterior cruciate ligament (ACL)

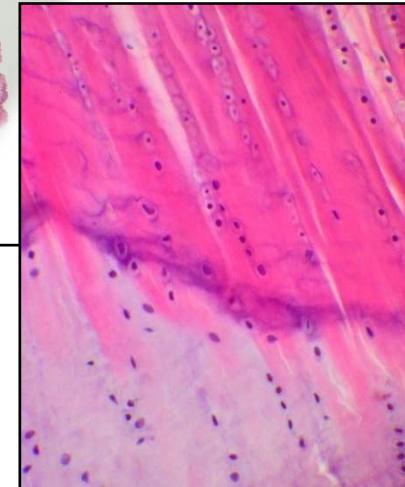


**Dimensions:**  
 $15 \text{ mm} \times 3 \text{ mm} \times 4 \text{ mm}$

mean substance

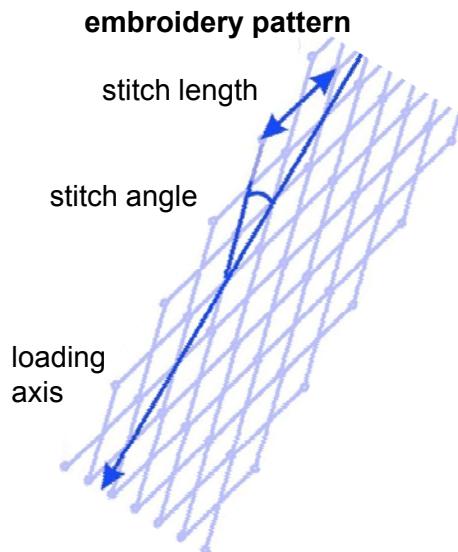


enthesis



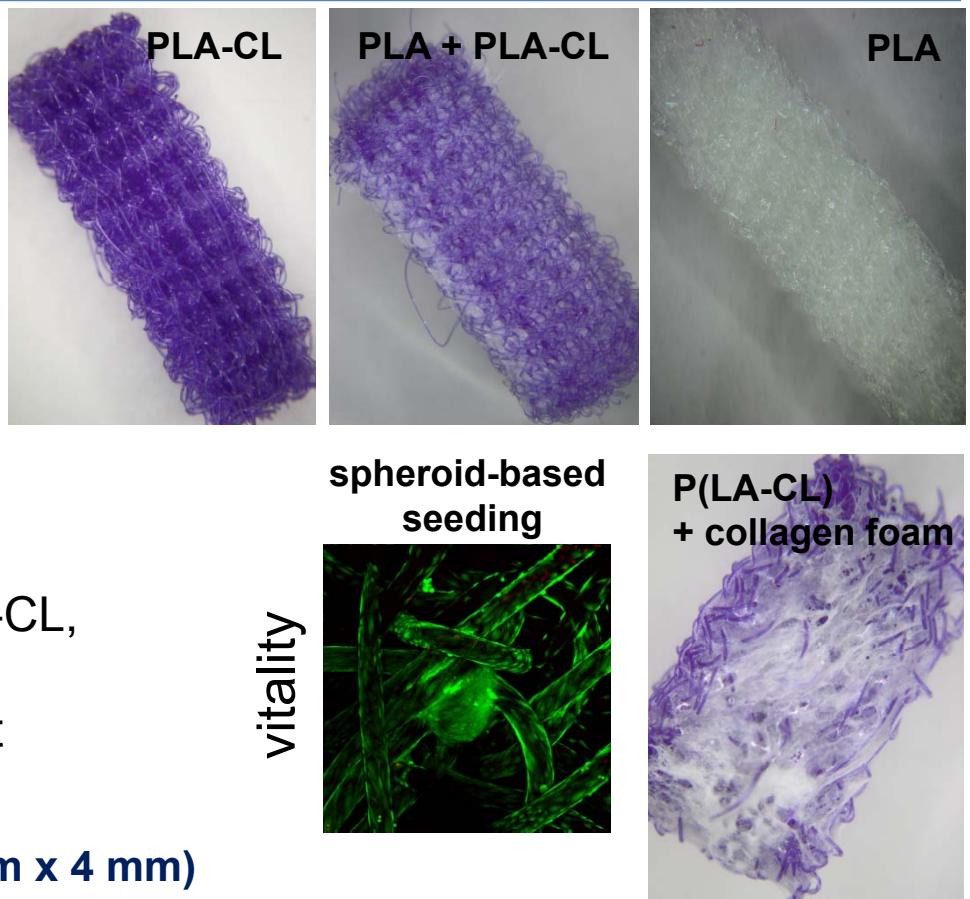
# embroidered scaffolds with collagen

- anterior cruciate ligament (ACL) tissue engineering using embroidered scaffolds supplemented with collagen

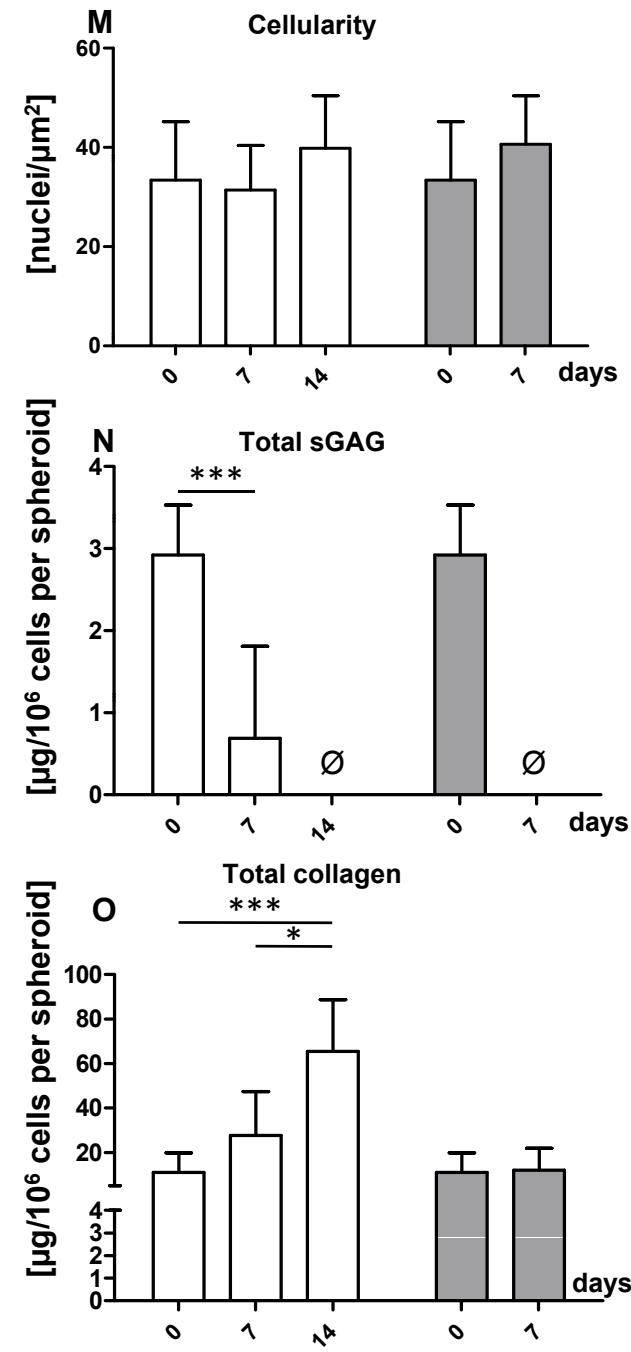
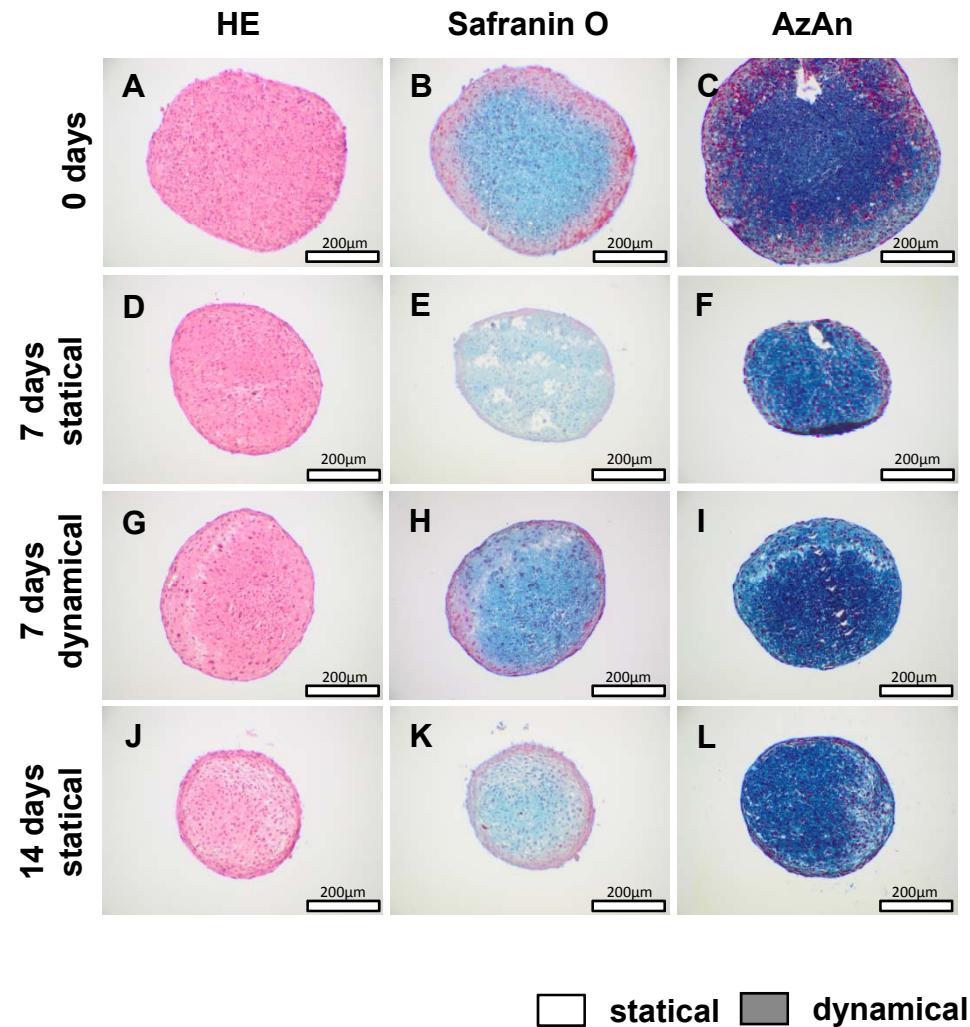


- Poly(lactic-co- $\epsilon$ -caprolacton) = PLA-CL, monofilament
- Poly-L-lactic acid = PLA, multifilament
- both materials combined

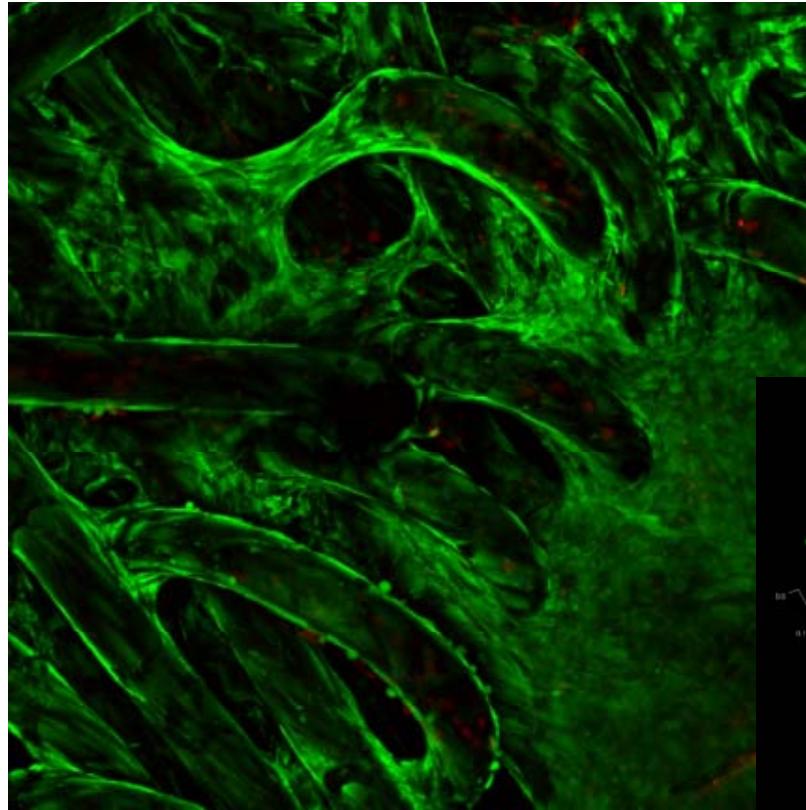
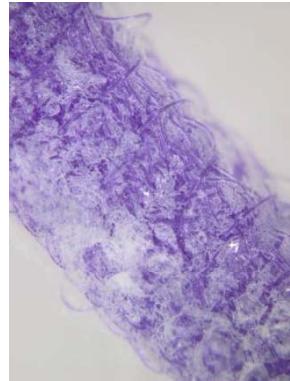
Dimensions of the lapine ACL (15 mm x 3 mm x 4 mm)



# ACL spheroid characterization



# Collagen scaffolds seeded with ligament cells



7 d

green: vital, red: dead cells

