

Clinical Need and Design Inputs

Osteoarthritis (OA) affects ~32.5 million people in the US annually¹



Healthy knee joint Osteoarthritis



ECM Cells



Cell death
ECM degradation

Specific Need

- Study cartilage cells and extracellular matrix (ECM) to treat OA
- Mimic compression on cartilage; study **cellular response**

Challenges

- Requires **high cell numbers** to maintain viability
- Ensuring load transmission onto cells

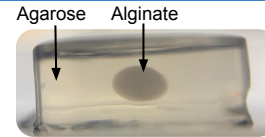
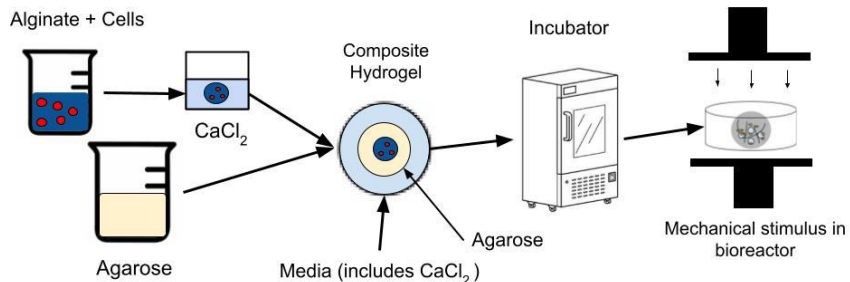
Key Requirements

- R1** Gel containing cells (alginate bead) must remain solid in polymerized state²
- R2** Composite gel must remain stable >20% strain³
- R3** Compressive load applied to entire gel must result in >10% deformation of bead³
- R4** >80% viability after 1 week in culture⁴

Objective and Solution

Objective

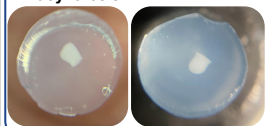
Synthesize a hydrogel that allows compression of a **small cell number** to study the effect of **mechanical loading** on cartilage ECM



Testing and Results

V1 Alginate Polymerization

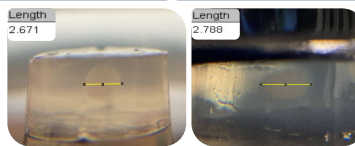
At synthesis 72 hours



Results: **Pass**

- Alginate stays polymerized
- P value = 0.347
- No significant change in diameter

V3 Load Sharing



Results: **Pass**

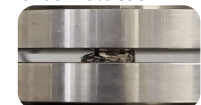
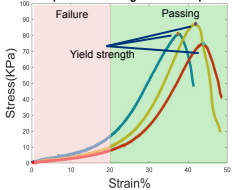
- Change in diameter 14.07% ± 4.6%
- n=3

V2 Mechanical Stability

Results: **Pass**

- Composite does not break under 20% strain

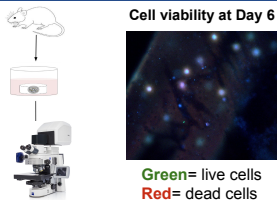
Compression testing on 3% Composite



Compression to failure on MTS

V4 Cell Viability

Cell viability at Day 6



Results: **Inconclusive**

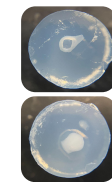
- Proof of *some* viable cells
- No accurate method to count cells yet

Revisions and Future Work

Revisions

Agarose formulation modified to 3% from 2%

CaCl₂ supplemented to culture media



Future Work

Measure cell viability and gene expression with mechanical loading in bioreactor



Impact

- Further understanding of OA upon observing cell mechanosensing in vitro
- Solution may be implemented for other cell types (ex. cancer cells)

References

¹The University of North Carolina at Chapel Hill. (2022, October 12). On Prevalence and Burden. Osteoarthritis Action Alliance. Retrieved October 12, 2022, from <https://www.osteoarthritisalliance.org/prevalence-and-burden>
²Wang, Y., Liu, T., Cui, L., Zhu, W., Li, H., Jiang, R., Yu, S., & Cui, C. (2020). Promotes Viability and Differentiated Phenotype of Cultured Chondrocytes With Low-Level Laser Irradiation. *Polymer Degradation and Stability*, 4, 458. <https://doi.org/10.1080/23802381.2020.1804608>
³Wang, Y., Liu, T., Cui, L., Zhu, W., Li, H., Jiang, R., Yu, S., & Cui, C. (2020). Functional Tissue Engineering of Articular Cartilage Through Dynamic Loading of Chondrocyte-Seeded Agarose Gels. *Journal of Biomedical Engineering*, 12(2), 122–205. <https://doi.org/10.1089/1548-3659.2019.00001>
⁴ISO 10885-2:2019. *Biological Evaluation of Medical Devices, Part 5: Tests for In-Vitro Cytotoxicity*. International Organization for Standardization: Geneva, Switzerland, 2019.