

Matrix Composition and Stiffness Regulate Ovarian Cancer Behavior

Introduction

- Matrix stiffness regulates ovarian cancer behavior.
- Cells interact with different extracellular matrix (ECM) components such as fibronectin and laminin.
- Fibronectin communicates between the intra and extracellular environment.
- Laminin regulates cellular activities and signaling pathways in the basement membranes of cells.



Cell membranes have many receptor-type proteins responsible for ECM interaction to transduce various signals. Both mechanical and biophysical components of the ECM contribute to the variety of intracellular signals.

Objective

- To observe how the stiffness of Dextran gels affects the matrix absorption of different proteins, fibronectin and laminin.
- To explore ensuing migration of ovarian cancer with respect to protein presence and stiffness.



 Identify key ECM components that regulate pathways in cancer that could be targeted for drug treatment.

Methods/Materials

- 2D UV Dextran Gels were crosslinked using irgacure photo initiator.
- Dextran gels were coated with fibronectin and laminin.
- Cells were seeded after ECM coating and stained using Phalloidin in a 1:1000 ratio with PBS++

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Material Synthesis and Hydrogel Formation



50mg/mL Dextran Methacrylate is mixed with irgacure and crosslinked under a UV light for either 45 seconds or 240 seconds. ECM proteins are independently incubated for one hour on the gels for absorption.



- 45 and 240 seconds showed significantly different storage moduli at ~850 Pa and ~8000 Pa • Tunable material moduli can capture different stiffnesses of
- the tissue within the body.



- Antibody staining shows the difference in absorption of ECM proteins
- Laminin formed sheet-like structures, while fibronectin was scattered, as that is how they arrange in natural conditions.







• The cell morphology images show that both the absorption of different ECM proteins and gel stiffness affect Ovarian Cancer.

- coating.







• OVCAR8 cells were grown on 45 and 240 second gels with either Fibronectin or

• Laminin appeared to be elongated, while Fibronectin appeared to be more circular Higher stiffnesses appear to have higher



• HEYA8 cells were grown on 45 and 240 second gels with either Fibronectin or

• Cells cultured on stiffer gels appear to be more elongated than on softer gels There appears to be no significant difference between the ECM proteins

Discussion

• The differing ECM proteins more significantly influenced OVCAR8's cell morphology, while the gel stiffness affected HEYA8's morphology. • Future work could be done using aspect ratios to quantify morphological

differences and ECM absorption, which determines stiffness-dependent

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