

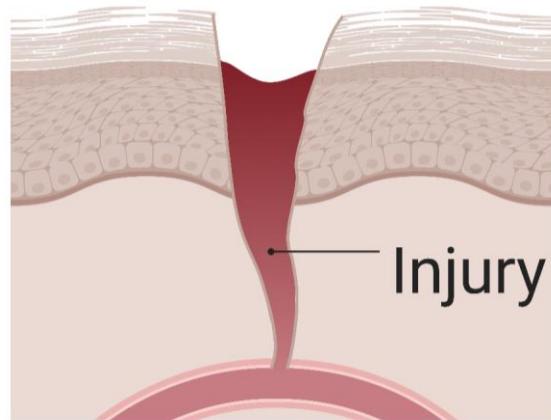
Biomaterials for Modeling Immune Mediation in Wound Healing

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The National Academies of Sciences, Engineering and Medicine
11/2/21

Why is there variability in wound healing?



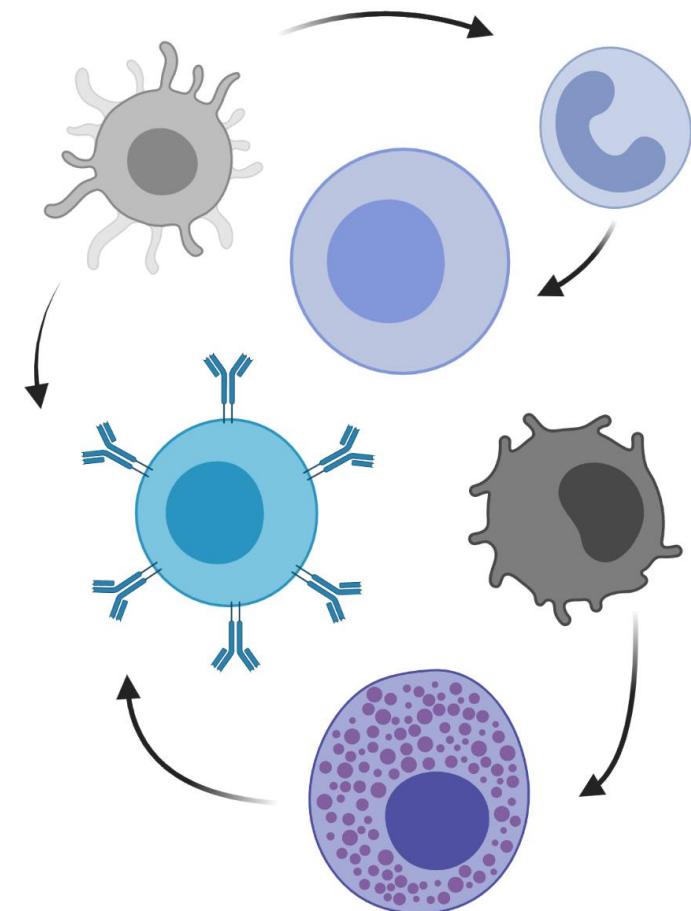
Open research questions

■ Can we understand why there is variability in wound healing?



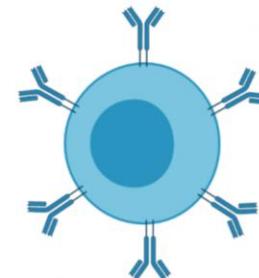
Immune cells: A coordinated team

- Multiple immune cells comprise the immune response to injury
- Orchestrate wound healing through cell-cell communications



Immune cells orchestrate healing

- B cells represent systemic response to injury
 - B cells are involved in wound healing
 - B cell dysregulation is associated with many diseases



Day 3

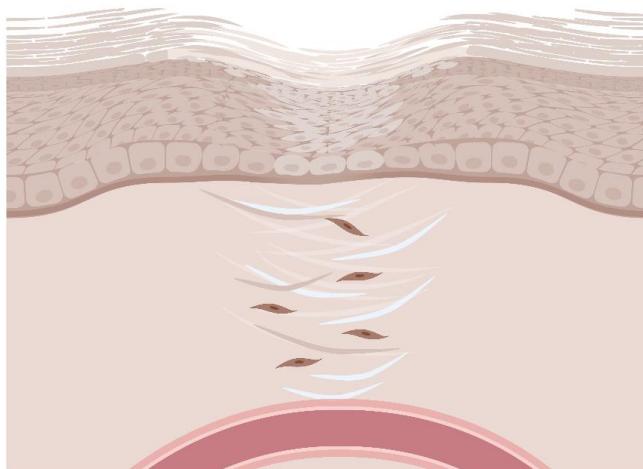
Day 7



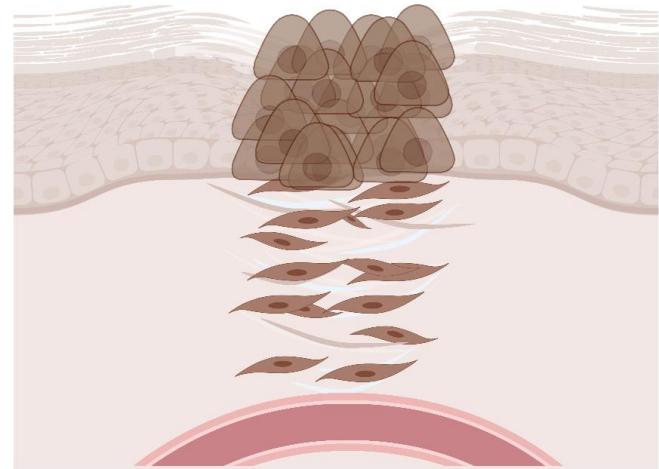
■ How do B cells respond during wound healing?

Biomaterials to mimic wound healing differences

Pro-healing

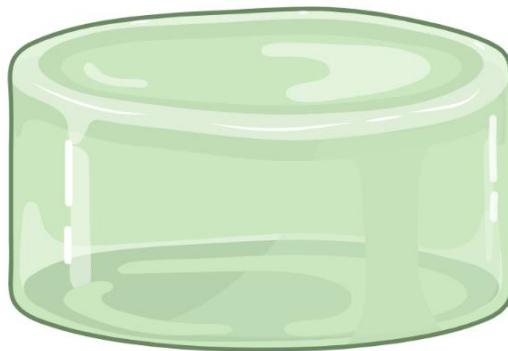


Pro-fibrotic



Biomaterials to mimic wound healing differences

Pro-healing materials

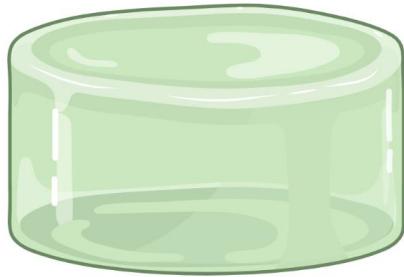


Pro-fibrotic materials



Biomaterials to manipulate wound healing

Pro-healing materials



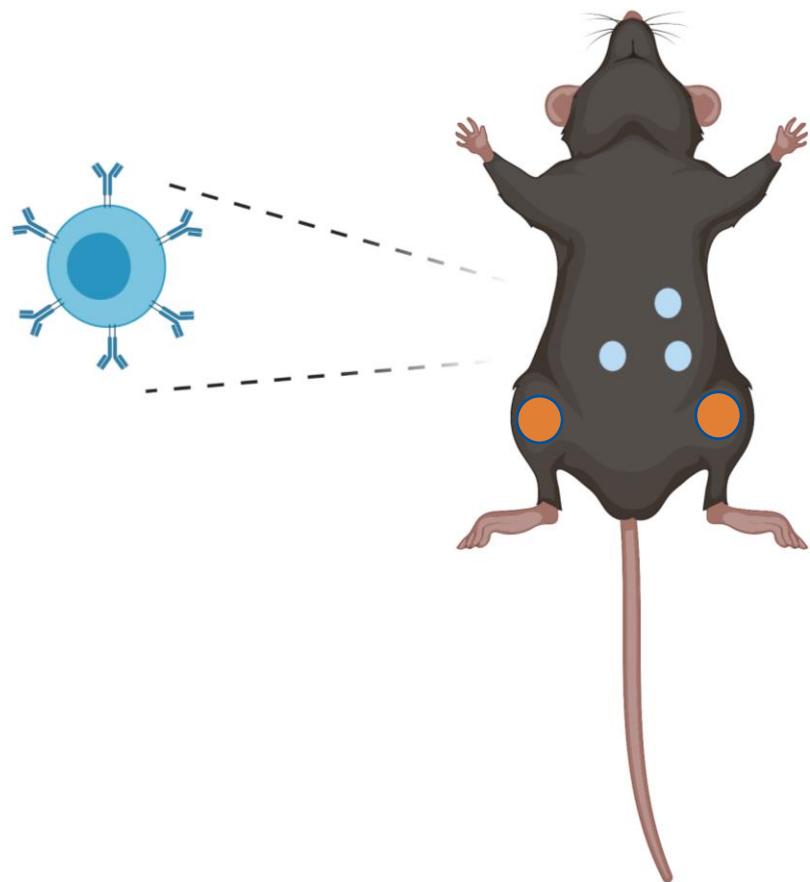
Pro-fibrotic materials



- Inform the local environment to skew wound healing
 - Pro-healing: Biomaterials that promote healing
 - Pro-fibrotic: Biomaterials that promote scarring and fibrosis

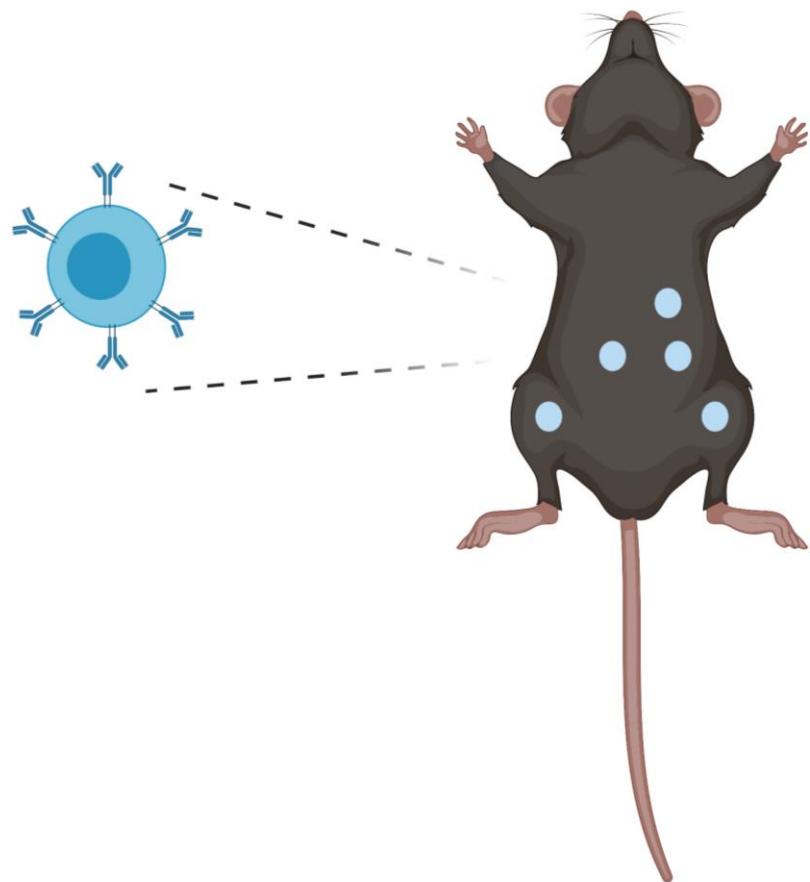
Alterations to the local environment influence systemic responses

- Create muscle injury
- Implant biomaterial in muscle
- Subsequent interrogation



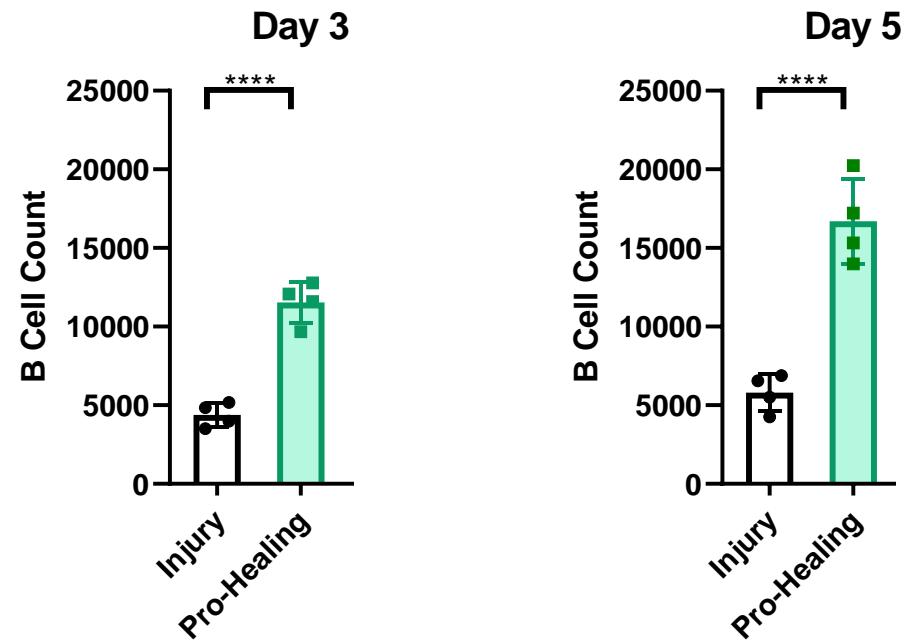
Alterations to the local environment influence systemic responses

- Create muscle injury
- Implant biomaterial in muscle
- Subsequent interrogation
 - Local environment
 - Responses in different tissues



Biomaterials to manipulate wound healing

Pro-healing materials

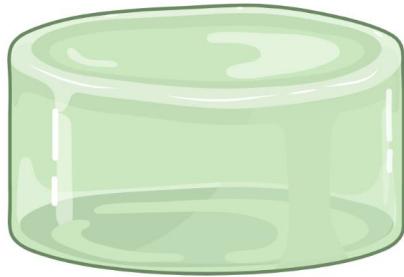


Post muscle injury with pro-healing biomaterial

- Pro-healing biomaterials recruit immature B cells early to the wound site

Biomaterials to manipulate wound healing

Pro-healing materials

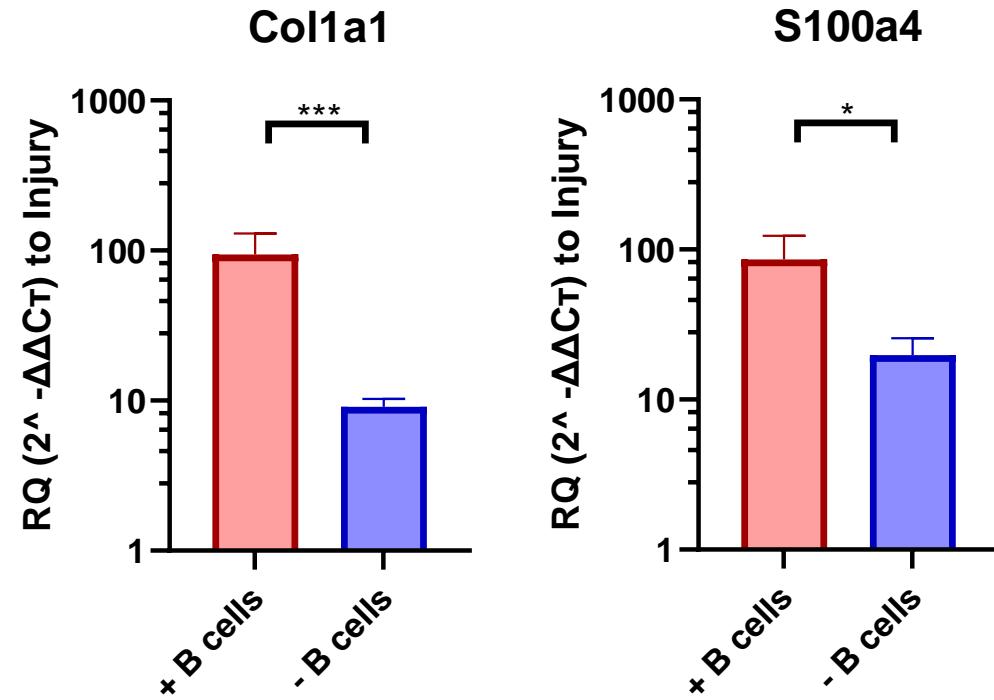


1 week post muscle injury with pro-healing biomaterial

- Pro-healing biomaterials inform tissue responses in the local lymph node

Biomaterials to manipulate wound healing

Pro-fibrotic materials



- Pro-fibrotic biomaterials recruit **Mature** B cells
- Removing mature B cells reduced fibrosis or scar formation

B cells in wound healing

Pro-healing materials



Pro-fibrotic materials



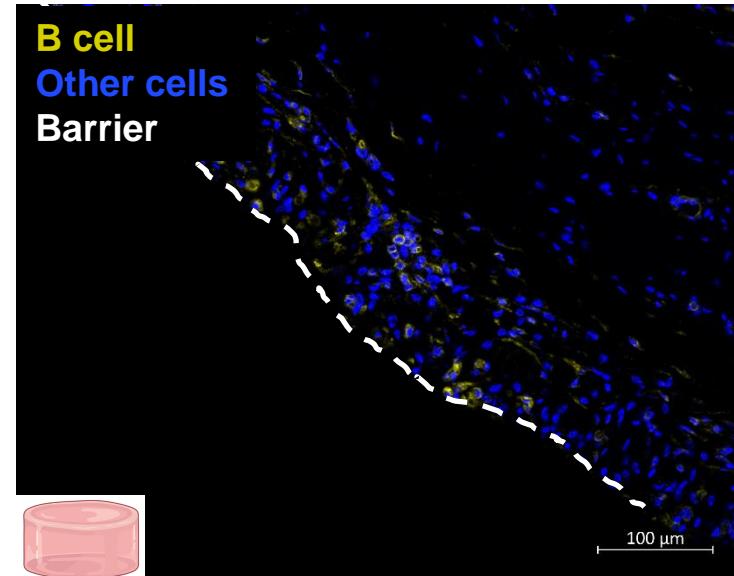
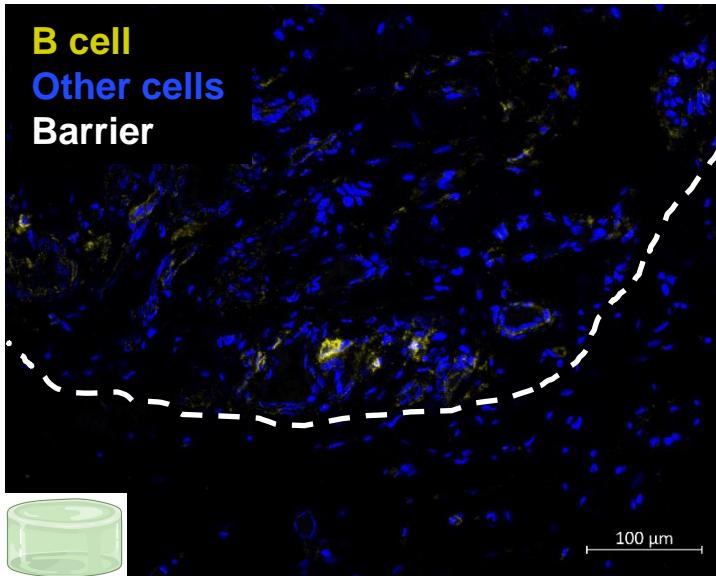
Immature B cells



Mature B cells



Biomaterials implanted in breast tissue



- Patients with naturally-derived materials

Pro-healing materials

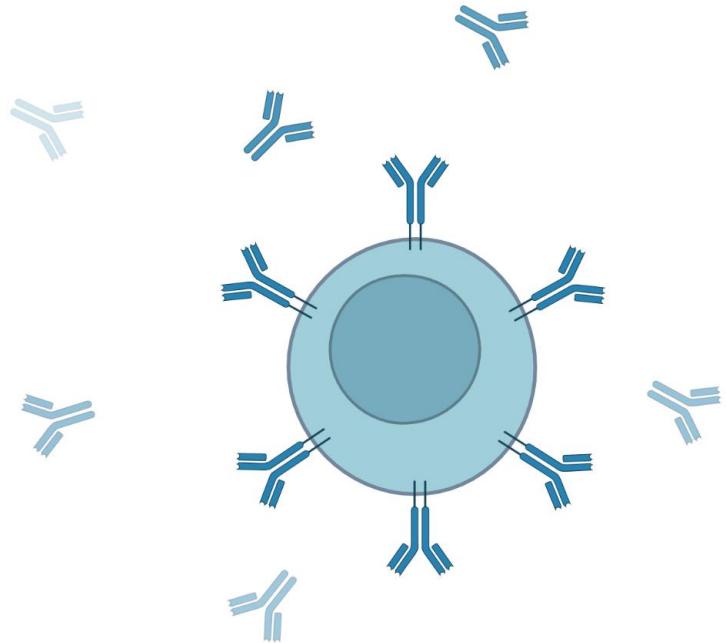


- Patients with silicone implants

Pro-fibrotic materials

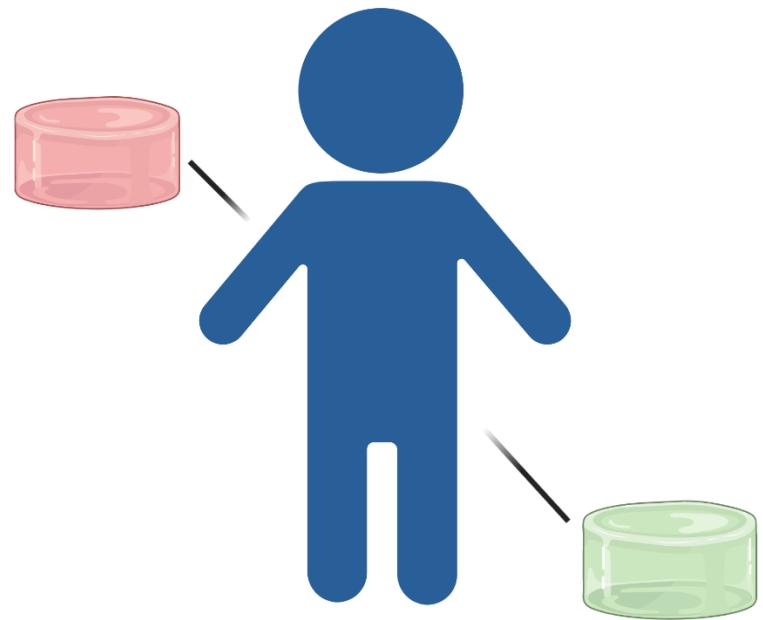
Gaps in understanding

- Adaptability of the B cell response in wound healing?
- Local tissue response can inform the systemic response



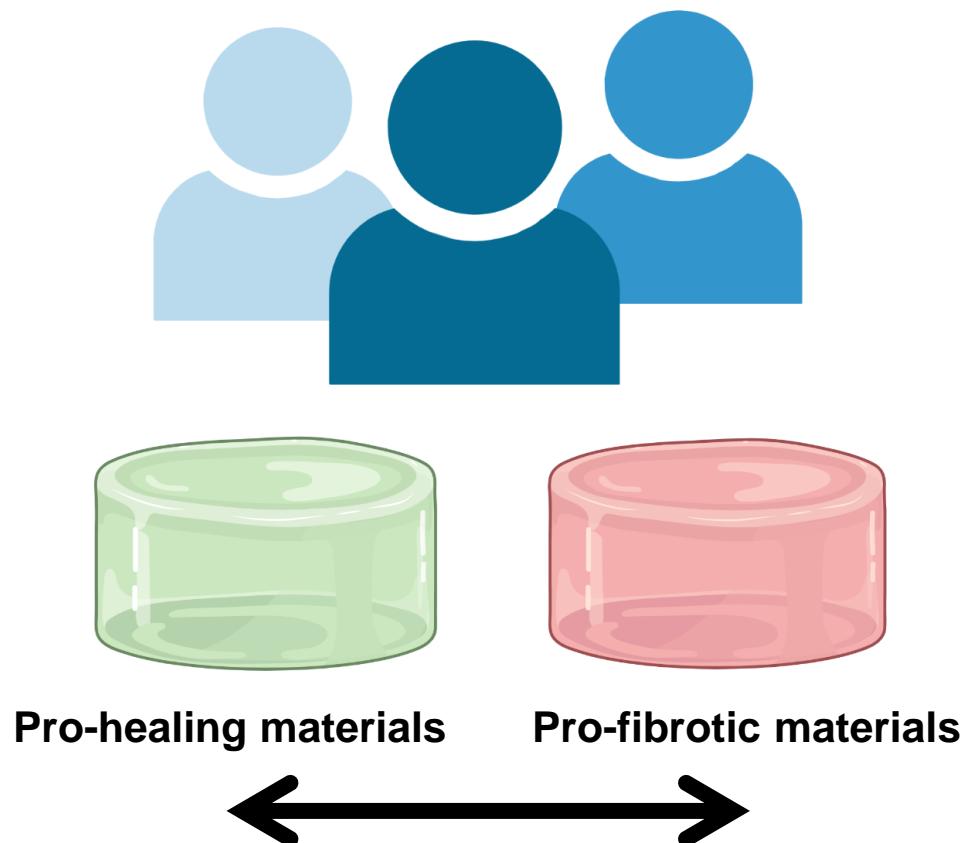
Gaps in understanding

- How do we connect the local injury response to systemic immunity?
- How do we leverage biomaterial models to understand patient-specific wound healing?



Biomaterials can be used to stratify the immune response in wound healing

- Can certain biomaterials mimic wound healing conditions of vulnerable populations?



Open research questions

- Can we interrogate the contribution of patient variability in wound healing?
 - Ancestral contributions
 - Age
 - Biological Sex
 - History of Trauma

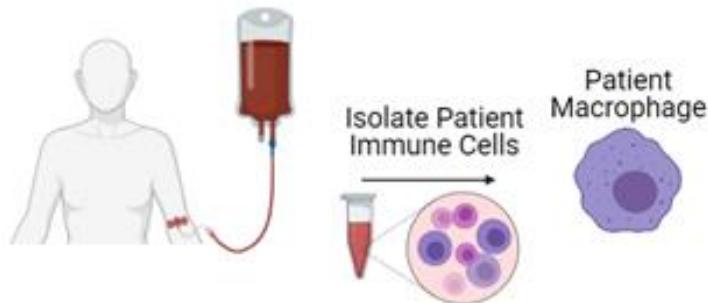


How can we use biomaterials to improve tissue regeneration?

- Biomaterial systems offer controlled systems for interrogation
- Use of organoid biomaterials to achieve patient specificity



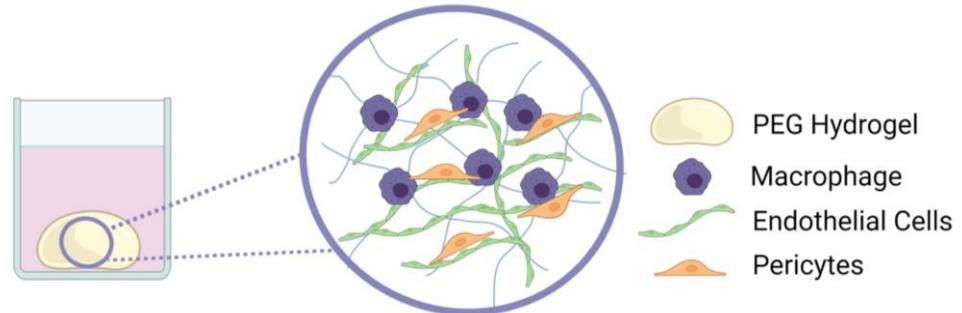
Biomaterials to interrogate age in regeneration



Immune Cells are influenced by:

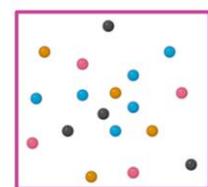
- Ancestral contributions
- **Age**
- Biological Sex
- History of Trauma

Biomaterial Model of Aging

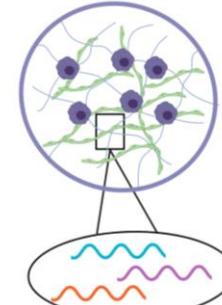


PEG Hydrogel
Macrophage
Endothelial Cells
Pericytes

Analysis in Biomaterials



Cytokine
Secretion
(ELISA)

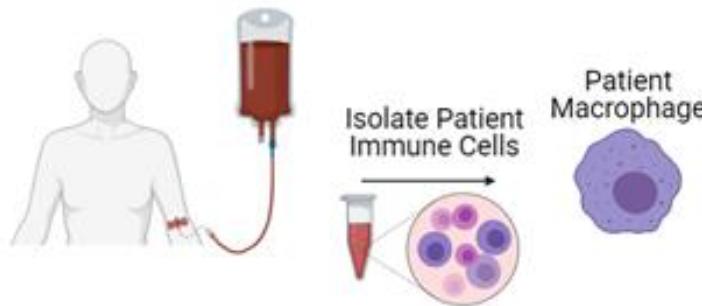


Gene Expression
(NanoString[©])



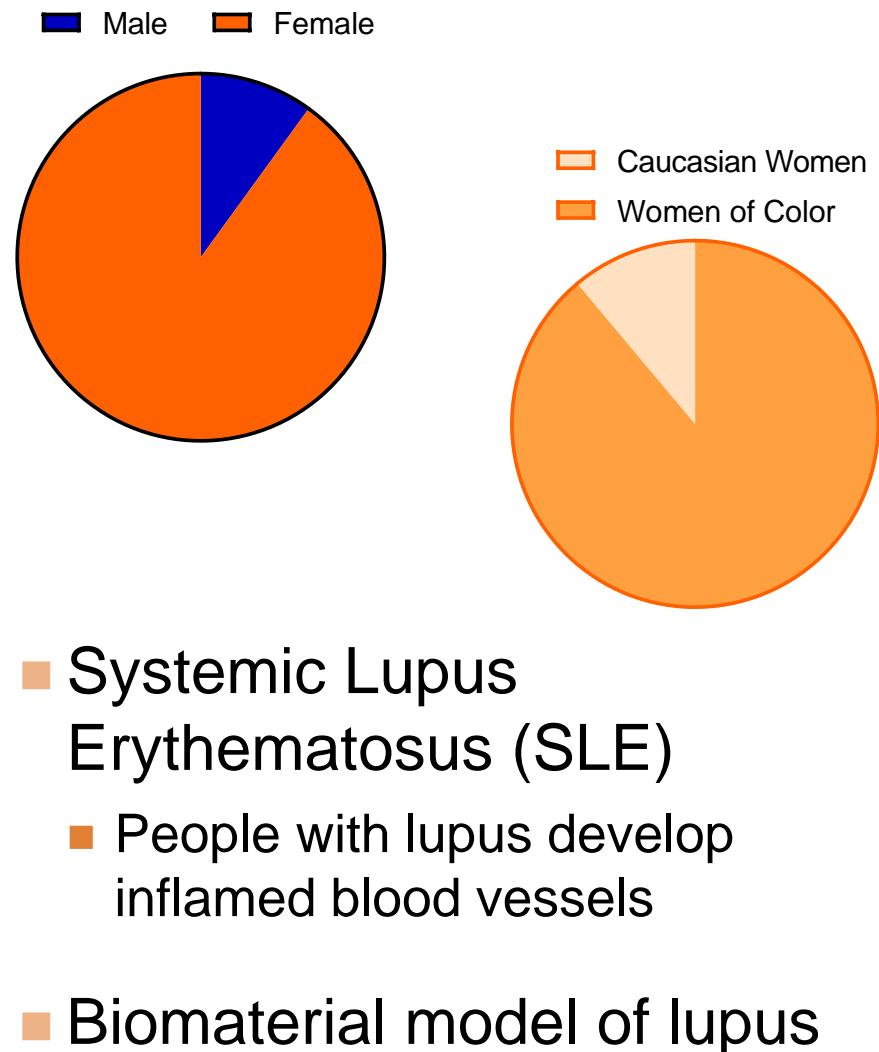
Immuno-
Fluorescence

Biomaterials to interrogate health inequities



Immune Cells are influenced by:

- Ancestral contributions
- Age
- **Biological Sex**
- History of Trauma



Thank you!

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Holly Ryan

All schematics created in BioRender



Learn more about our new lab:
www.Moore.Mse.Ufl.edu



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