



# Evaluating Abdominal Wall Health at the Mesh-Tissue Interface in Ventral Hernia Repair

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# Disclosures

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- Abdominal Core Health Quality Collaborative – salary support
- Bard-Davol, Advanced Medical Solutions – research support

# You Would Think...

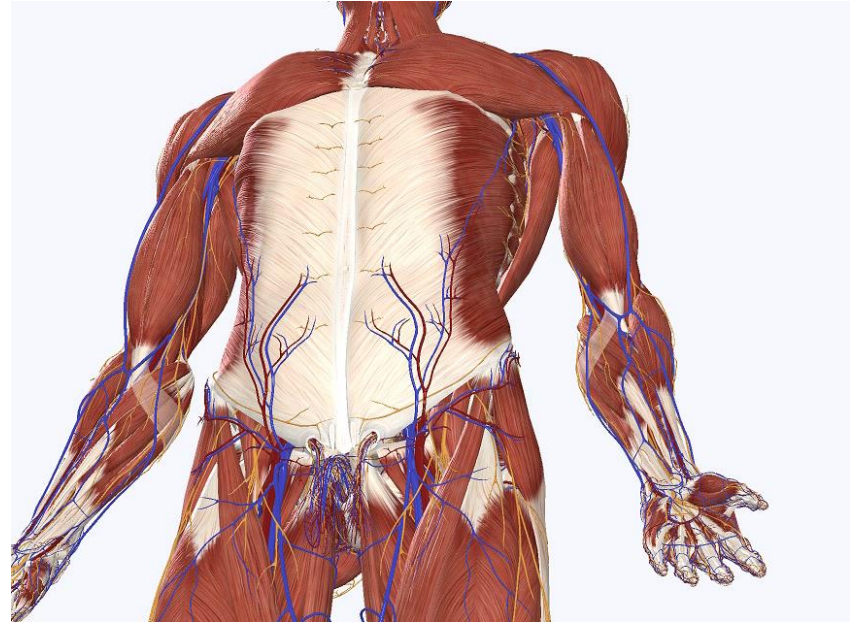
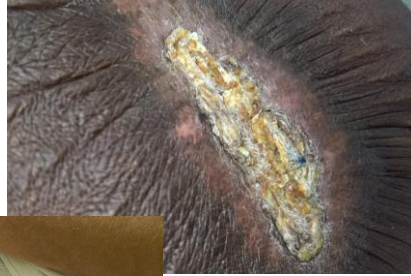
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# You Would Think...



# It Get's Complicated

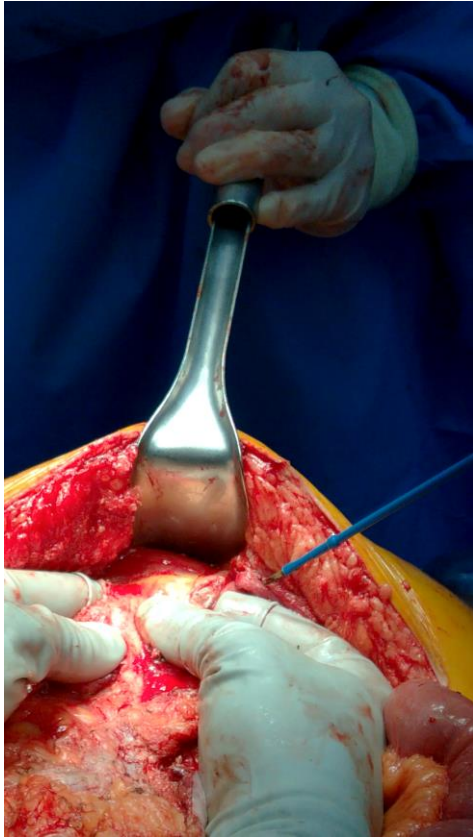




# Ventral Hernia Repair – Unintended Consequences

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- The vast majority of patients do well
- Some do not –why?



# How Often Does This Happen?

- 611,000 ventral hernia repairs done per year in U.S. (Schlosser et al, *Hernia* 2022)
- 5.6% mesh related complications **requiring excision** at 5 years in healthy Danish people (Kokotovic et al, *JAMA* 2017)
- Over 20 years (assume we are like healthy Danes, and 80% repairs use mesh):
  - At least **550,000** people with mesh related complications needing surgical intervention

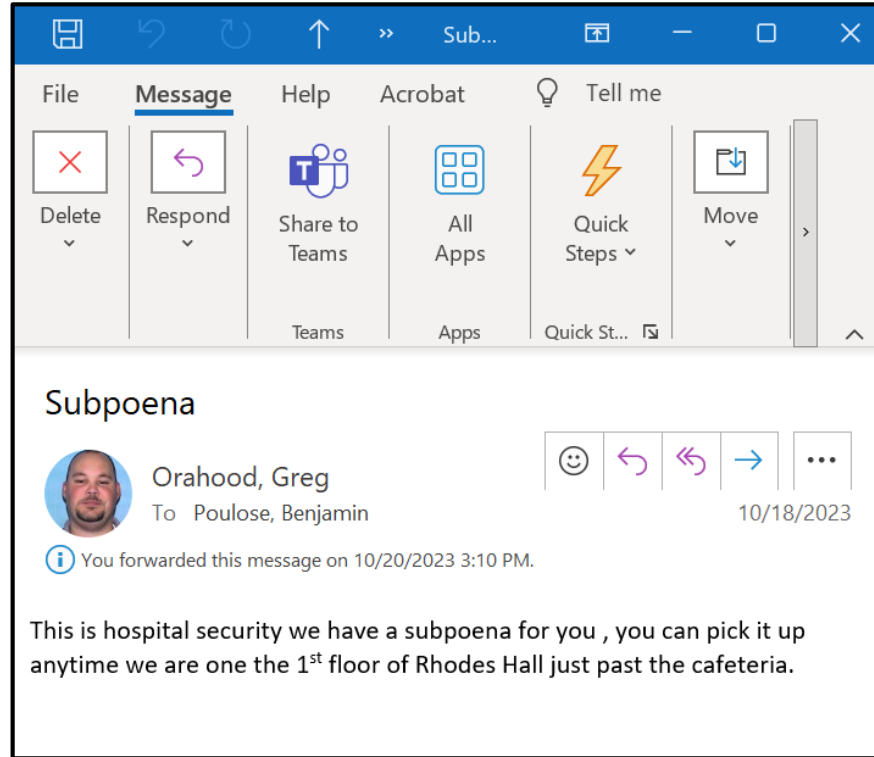


# The Mess about Mesh

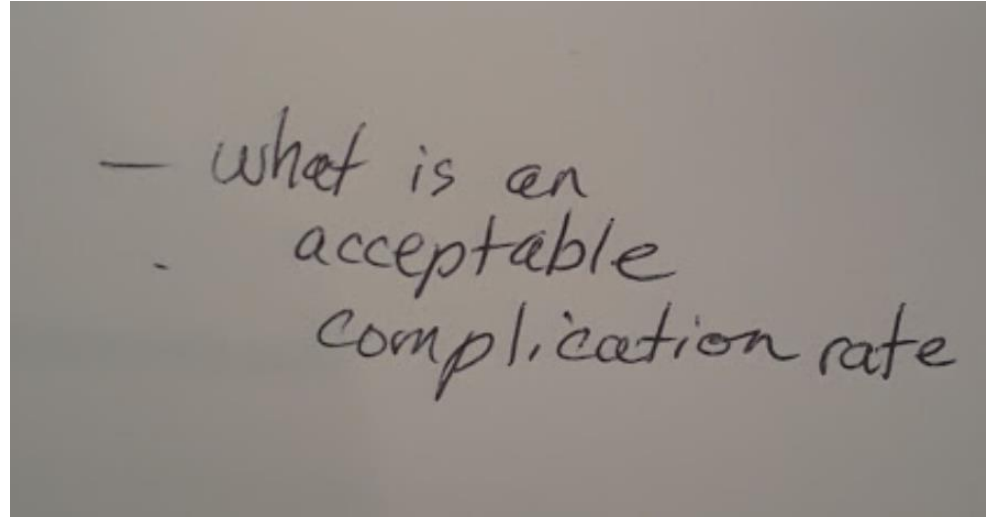
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- Thousands of individual lawsuits across the country
- Multiple MDLs (Multi-District Litigation) actions

# Take your PPIs!!



# The Real Problem



...This is a basic science problem

# Overall Question

— What Happens Right Here Over Time?



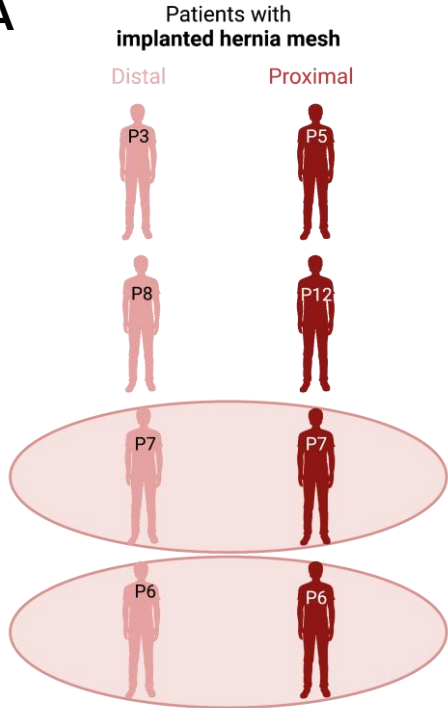
**Study aim:** Evaluate the changes in gene expression of tissue samples proximal or distal from the mesh/tissue interface of patients undergoing recurrent **Ventral Hernia Repair (VHR)**

## **Experimental design**

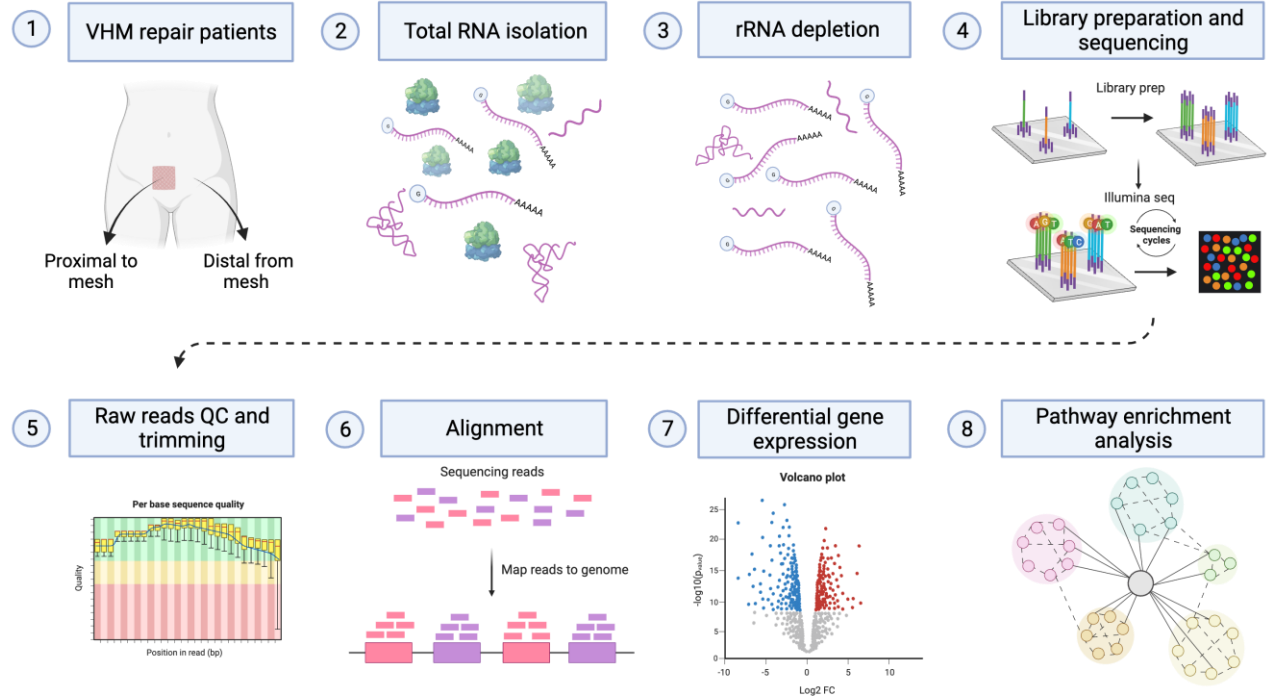
- **Samples of 1.0 cm<sup>2</sup> from the abdominal wall at the mesh/tissue interface of the previous repair site:**
  - 4 tissue samples at mesh-tissue interface (Proximal)
  - 4 tissue samples distal to the mesh (Distal)

# Methodology – Changes at the whole transcriptome level (RNA-seq)

A



B



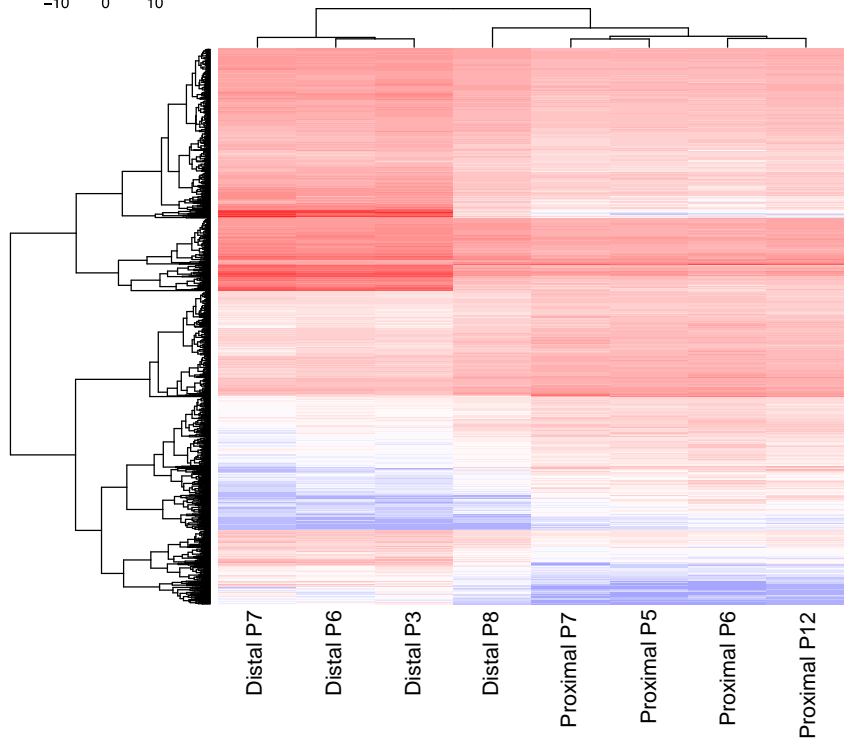
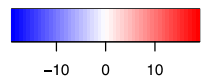
# Results

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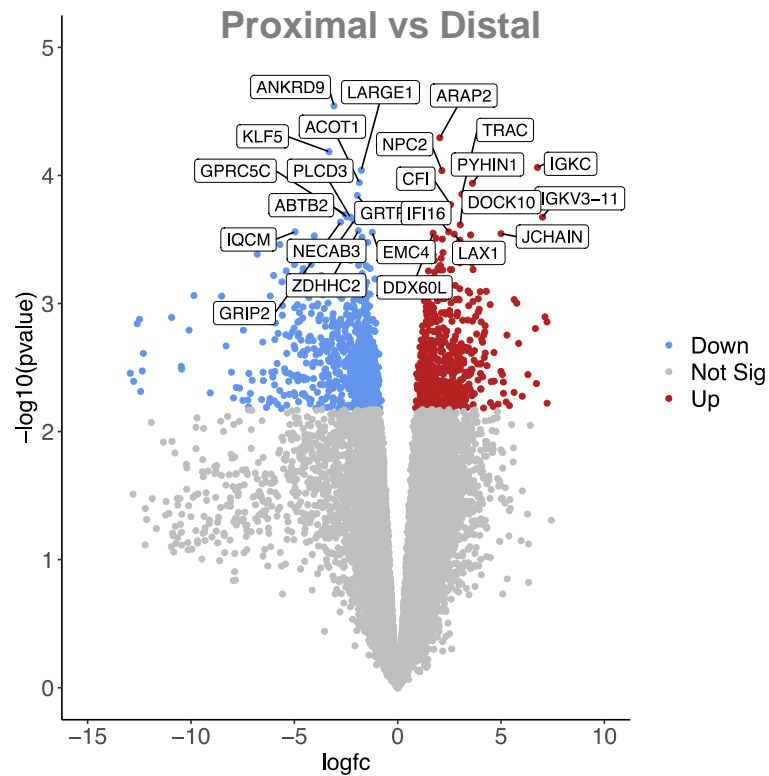
- Samples collected prospectively from 8 patients undergoing recurrent VHR
- Average age 63 years old
- Average time from previous repair 10 years
- All intraperitoneal mesh placement (polyester, polypropylene)

# Results

# Differential Gene Expression (DGE)

**A**

Fold-change > 2 and FDR < 0.1

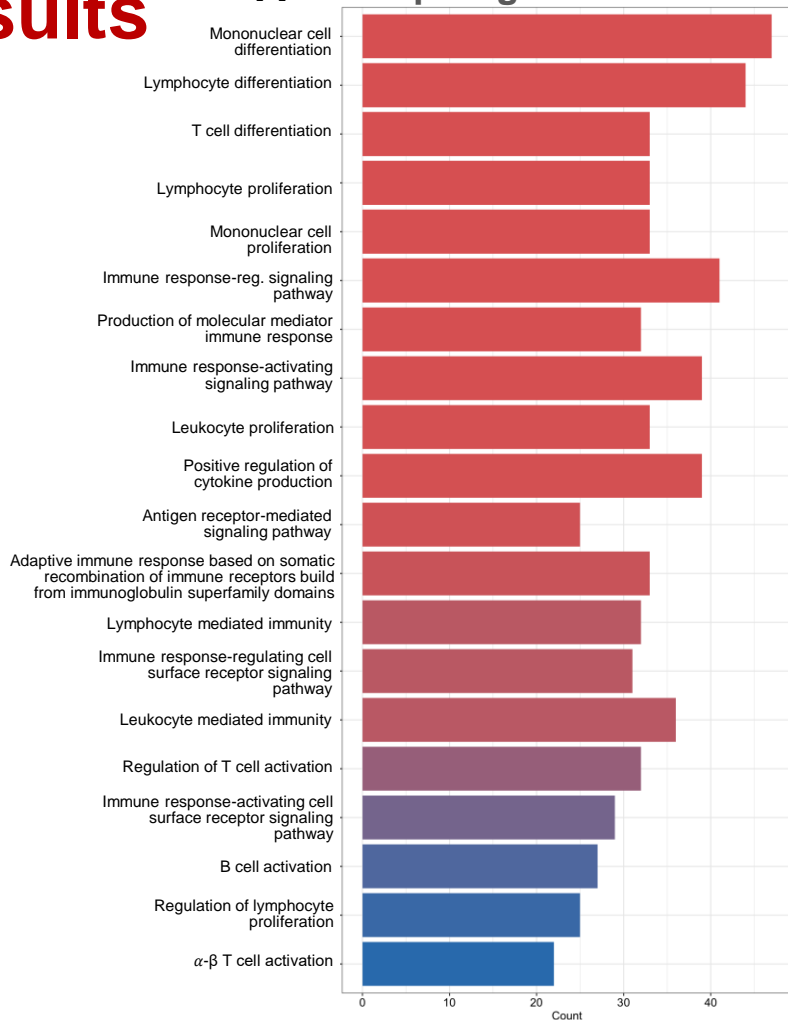
**B**

# Results

GO Enrichment: Proximal vs Distal

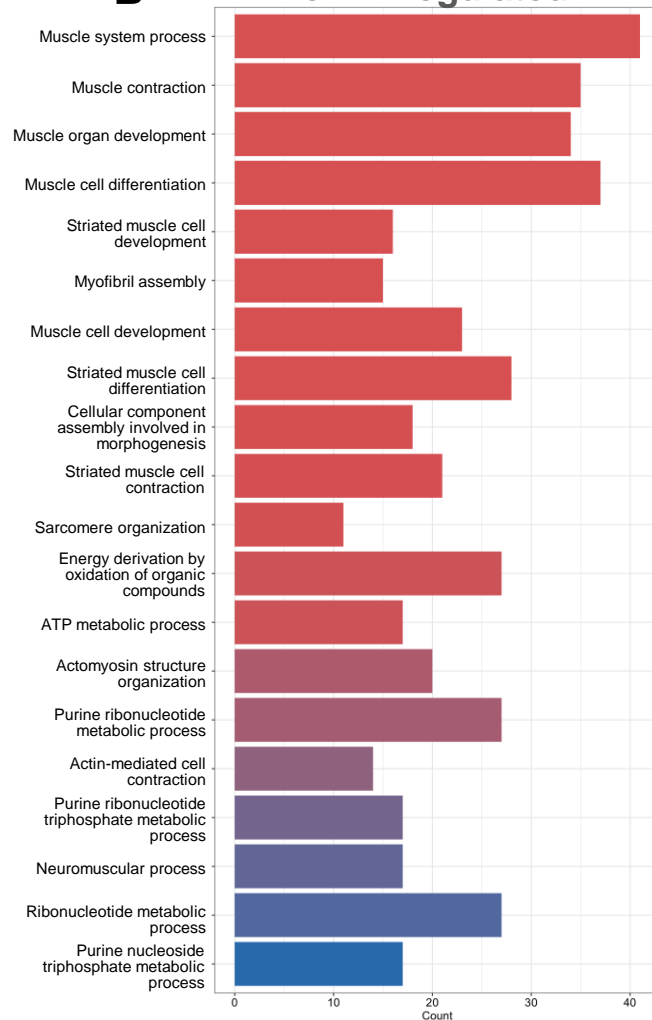
**A**

**Up-Regulated**



**B**

**Down-Regulated**





# Summary

- Marked and chronic **difference in gene expression** at the mesh/tissue interface
- Increased translation of genes related **cell proliferation** and **immune reaction** associated with **inflammation**
- **Oncogenes** upregulated

**Does this Happen?**



**Does this Happen?**



# Conclusion

- Preliminary data indicate that permanent prosthetic meshes create local changes in gene expression that can affect multiple regulatory pathways potentially impacting **cancer** and **immune regulation**, among other systems

# Acknowledgements

## Gallego-Perez Nanomedicine Lab:

Tatiana Cuellar-Gaviria, Hamza Jafri, Ludmila Diaz, Silvia Duarte-Sanmiguel, Ana Salazar-Puerta, Diego Alzate-Correa, Jon Staranan, Ana Panic, Kavya Dathathreya, Natalia Areiza, Junyan Yu

## Nanomedicine Lab

