A Quad-Culture Microfluidic Human Gut-on-a-Chip System

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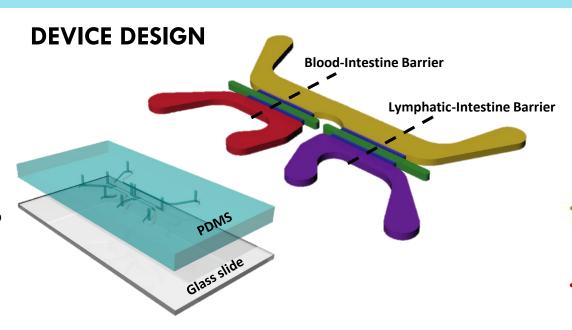


INTRODUCTION



- Successful treatments in animal models rarely translate into effective drugs for humans
- Microfluidic devices are a cheap and effective way to model the gut

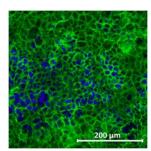
Goal: To better recapitulate the intestinal environment, we seek to mimic the anatomy of the human villi in our device.



METHODS

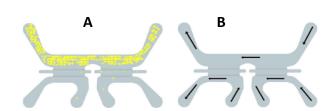
Staining and Imaging of Quad-Culture Model

- Live-dead cell assay
- Actin and nuclei visualization
- Immunofluorescence staining of mucus and cell junctions



Evaluating Barrier Function (A) and Comparing Flow vs. Static Conditions (B)

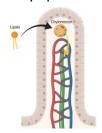
- **A.** Measure and quantify diffusion of lucifer yellow into blood and lymphatic channels.
- **B.** Evaluate villi-like structure formation, MUC2 expression, and barrier formation due to flow condition.



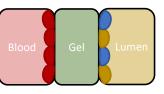
Measuring the Selective Uptake of Chylomicrons

Add fluorescently-labelled lipids to the lumen and observe chylomicron localization to lymphatic channel.

Chylomicrons: lipoproteins that transport dietary lipids in the body.



CELL BARRIERS



Lymph Gel Lumen

Blood-Intestine

Lymphatic-Intestine

(Cross-sections of device design)

- HT-29 and Caco-2: colon cancer cells and epithelial colorectal cancer cells (respectively) that form monolayer like enterocyte in the small intestine.
- Blood endothelial cells or Lymphatic endothelial cells: form interface between blood or lymph and the lumen.
- **Gel Channel**: fibrin structure provides support and acts as the extra-cellular matrix.

CONCLUSION

In progress: device fabrication and cell culture. **Future applications**: statistically reliable clinical trials with novel drugs; modelling microbiome.

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