PLATELET FUNCTION AND THROMBOGENICITY Labs



Th-Lab – Platelet function and thrombogenicity Lab

Biomechanics Research Group

Platelet Function and Thrombogenicity

Team



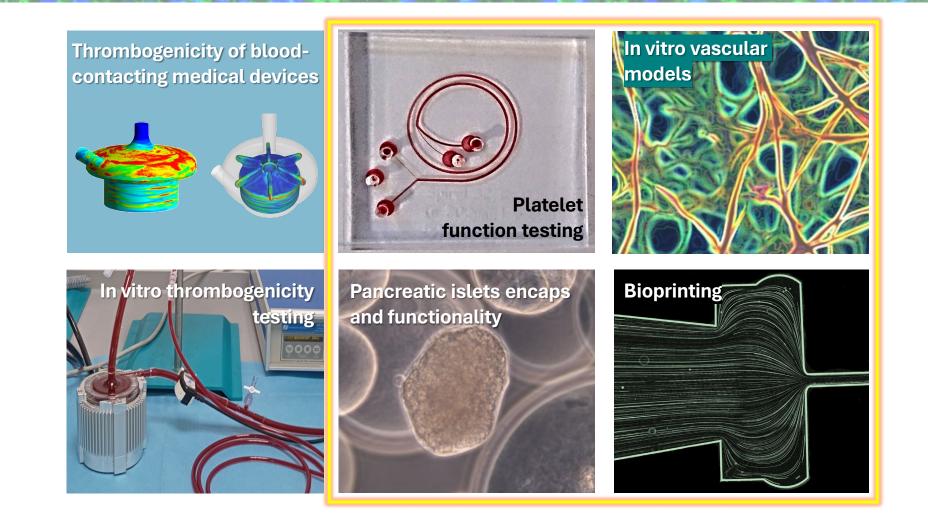
aleria Guadagno

Fabio Morelli

Platelet Function and Thrombogenicity

Tancesca Vicinan18

Research Activities



Platelet function testing THROMBOGENICITY OF BIOMATERIALS FOR BLOOD-CONTACTING MEDICAL DEVICES





Assess platelet adhesion and activation on biomaterials for blood-contacting medical devices

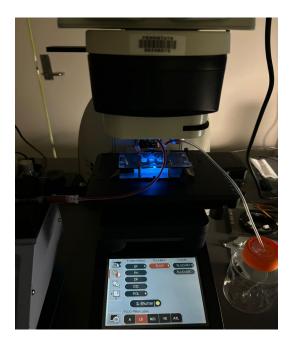
Activities

- Microfabrication
- Blood separation techniques
- Blood perfusion inside the microfluidic channels
- Investigate platelet adhesion by confocal fluorescence microscopy
- Image analysis
- Investigate platelet activation by flow cytometry

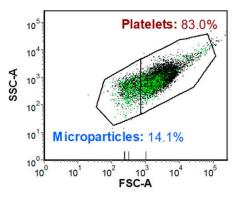
Applications

Stratification of the thrombotic risk of recipients of mechanical circulatory support devices

Background: Jingying Lin et al., 2024



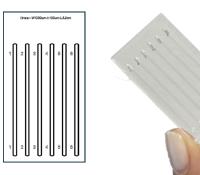
Platelet activation by flow cytometry



Imaging of platelet adhesion



Platelet Function and Thrombogenicity



Platelet function testing ATHEROGENIC FLOW DISTURBANCES AND VASCULAR INFLAMMATION IN A PERFUSABLE 3D STENOSIS MODEL



Assess the contribution of platelets and leukocytes in the development of thrombosis in a stenotic vessel model

Activities

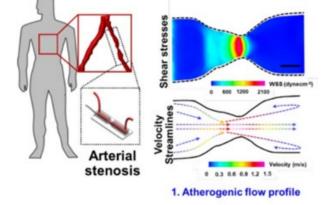
- Microfabrication
- Test different constriction geometries
- Evaluate platelets and leukocytes adhesion in stenotic channels with coating of different proteins
- Evaluate platelets and leukocytes adhesion in stenotic channels lined with endothelial cells
- Test anti-thrombotic and immunomodulatory efficacy of drugs

Applications

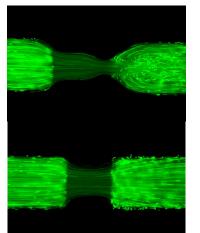
Atherogenesis

Background:

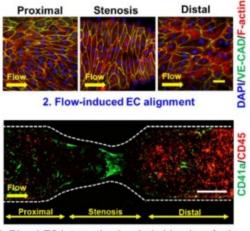
Menon et al. Biofabrication 2021



Stenosis-on-chip: blood flow disturbance



Fluorescent beads perfusion



3. Blood-EC interaction in whole blood perfusion

Platelets and leukocytes adhesion

Contacts: Alberto Redaelli, Tatiana Mencarini

Platelet Function and Thrombogenicity

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In vitro vascular models / Platelet function testing **ENDOTHELIAL CELLS - BLOOD INTERACTION**



Develop and test microfluidic platforms for studying the interaction between pathological endothelium and blood components to assess thrombogenicity

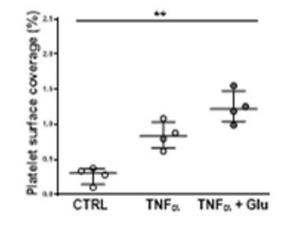
Activities

- Microfabrication
- Seed endothelial cells into silicone microchannels
- Optimise protocols for endothelial cells treatment with inflammatory stimuli
- Assess monolayer maturation by fluorescent microscopy imaging
- Perfuse whole blood within the endothelialised channels at different shear stress conditions
- Investigate thrombus formation

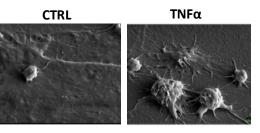
Applications

Predict cardio-vascular disease progression in patients with diabetes and chronic kidney disease

Endothelial cells PDMS Endothelia Blood PDMS (e) Nuclei







Contacts:

Silvia Bozzi, Tatiana Mencarini, Francesca Vicinanza

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Background:

Cuartas-Vèlez et al., 2023

In vitro vascular models ANGIOGENESIS-ON-A-CHIP

Fondazione IRCCS Ca' Granda Ospedale Maggiore Policlinico

Develop and test microfluidic platforms for studying angiogenesis by optimizing hydrogel matrix and channel geometries

Activities

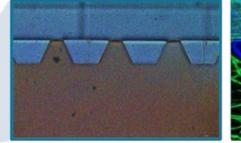
- Test different types of hydrogels (e.g., collagen, fibrin, PEG) to mimic the extracellular matrix
- Evaluate hydrogel mechanical properties
- Seed endothelial cells within the hydrogel
- Study how varying the distance between the pillars (trapezoidal structures in the image) affects vasculogenesis
- Monitor vascular network formation using microscopy and immunofluorescence staining

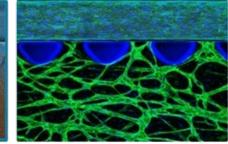
Applications

Investigate endothelial repair and regeneration after microvascular damage in patients with immune-mediated Thrombotic Thrombocytopenic Purpura (iTTP) and study how metabolic dysfunction affects angiogenesis in patients with Metabolic Associated Liver Disease (MASLD)

Background:

Zheng Y et al., 2012





Collagen gel

Vascular network

Contacts: <u>Silvia Bozzi, Tatiana Mencarini, Francesca Vicinanza</u>

In vitro vascular models 3D PERFUSABLE VESSEL-ON-CHIP



Vessel

Hydrogel

The device: High-Fidelity Vessel-on-Chip model for in vitro studies of vascular diseases and drug screening

Features: • PDMS platform

- Collagen I hydrogel constitutes the extracellular microenvironment
- Endothelial cells are seeded within the circular lumen shaped within the hydrogel

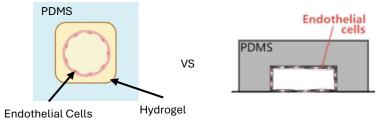
Project 1

Reconstructing the basal lamina for a realistic 3D vascular microenvironment



Project 2

Comparative analysis of endothelial cell response in a Vessel-on-Chip model with circular geometry and extracellular microenvironment vs. conventional rectangular PDMS channels

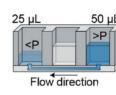


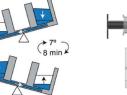
Project 3 Comparativ

Biomechanics Research Group

Comparative analysis of different perfusion methods of a Vessel-on-Chip model

Background: Cuartas-Vélez et al., Biomedical Optics Express 2023







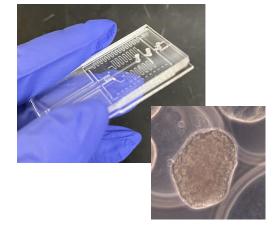
Contacts: Silvia Bozzi, Valeria Guadagno

Platelet Function and Thrombogenicity



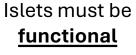
Pancreatic islets CONFORMAL ENCAPSULATION AND FUNCTIONALITY ASSESSMENT

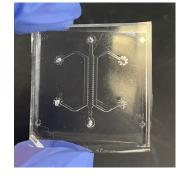


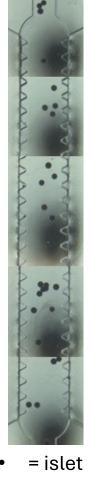


Encapsulation has the potential to allow islets <u>transplantation without immunosuppression</u>

<u>Microcapsules</u> may have failed because of **transport issues** associated with capsule **size**







Conformal Coating

encapsulation addresses all the transport issues associated with capsule size

- CAD drawing (SolidWorks, autoCAD)
- Microfabrication
- Lab work (cell culture, flow loop, microscopy)

Assessment of islets' functionality in a **physiologically relevant microenvironment** prior to the implantation

- Lab work (cell culture, gel prep)
- Microfabrication
- Lab assays (microscopy, FRAP, ELISA)

Contacts: Alberto Redaelli, Tatiana Mencarini

Activities

3D extrusion-based bioprinting HIGH CELL DENSITY CONSTRUCTS

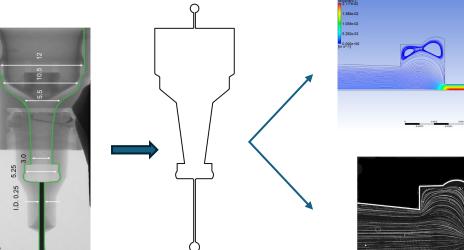


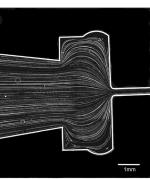
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Optimization of the bioink-nozzle coupling Task 1: Design of new gel precursors for bioink Task 2: Design a new 3D printer nozzle, coupled with the bioinks

Activities

- Rheological characterization of the newly synthesised gels with/without cells
- Ansys fluid dynamics simulation of the flow in standard and in newly designed nozzles
- Microfabrication of nozzle model devices
- Validation of the model via fluorescent beads perfusion and imaging





Contacts: Alberto Redaelli, Tatiana Mencarini

Active collaborations

