

Ichilov  
Scientific  
Ecosystem







# Laboratory of Neuroimmunology



# ➤ Who we are and our area of interest

## Name of lab/Location

- Neuroimmunology Research Lab
- Rishonim building, 5<sup>th</sup> floor, room 62

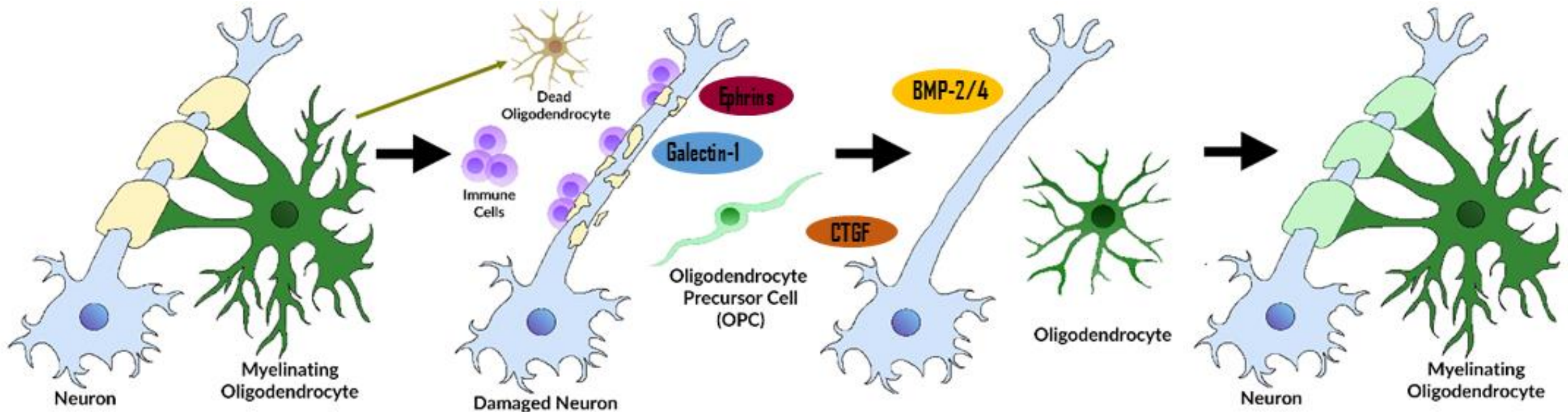
## PI/Manager

- PI: Prof. Arnon Karni
- Manager: Dr. Maya Golan



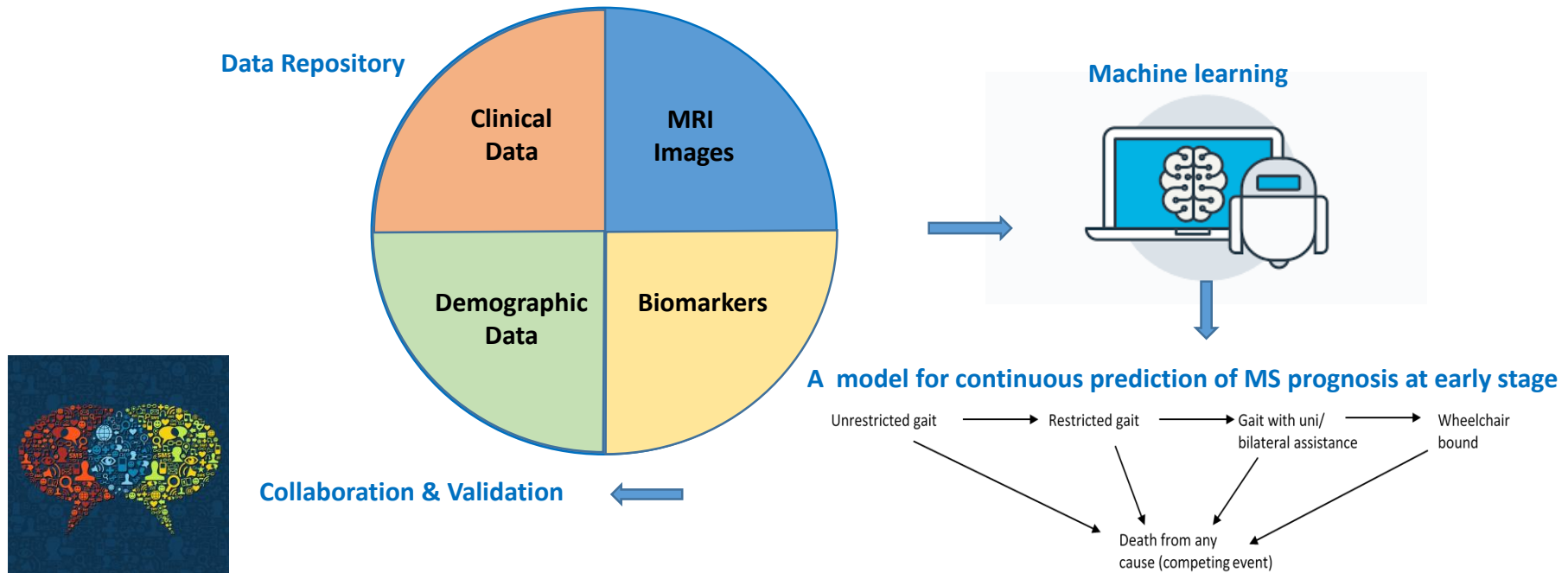
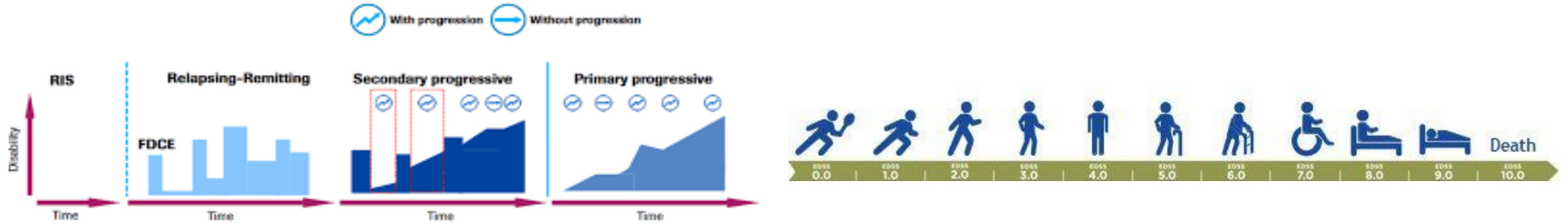
## ➤ Who we are and our area of interest

### ➤ Demyelination and Remyelination in Multiple Sclerosis (MS)





# ➤ Developing a prognostic model for disability





# Who we are and our area of interest

## Main Subjects in the lab

- Exploring and characterizing different factors that can affect remyelination failure in MS.
- Developing new agent for a novel regenerative treatment for MS and studying their mode of function.
- Developing an effective multi-parameter prognostic calculation model of disability in multiple sclerosis, using advanced statistical and machine-learning methods.
- Further evaluating the mode of action of existing disease modifying drugs.

## Keep it simple to people who are not in the field

- To discover new therapies approaches in order to cure the tissue damage in multiple sclerosis (MS).
- We are studding biological targets that can influence the optimal repair of myelin damaged in MS.
- We are developing a multi-parameter prognostic model for early stages of the disease.



# Key Capabilities

## What are we specialized in

- ELISA and Multiplex.
- WB analysis for cell signaling.
- Flow cytometry panels for T cells and macrophages sub-populations.
- Assay development of protein-protein interaction for HTS (HTRF).
- OPCs and Macrophages differentiation in-vitro.
- Mice models for MS: EAE and Cuprizone.
- IF histochemistry: for cells and neuronal histopathological specimens.
- Confocal analysis and ImageJ image analysis
- EM analysis for remyelination (G-ratio).

## What specialized equipment we use to answer Q

- ELISA reader.
- EVE cell counter.
- Inter departmental equipment:
  - Canto II- Flow cytometer.
  - Confocal Microscope.
- Tel Aviv University: EM.
- Weismann institute: HTS – high thruput screenings.

## How can we aid other scientists to answer their Q

- Usage of the ELISA reader, EVE cell counter.
- Help and consultation in IF staining, confocal analysis and imageJ software usage.
- Advice with stem cell differentiation models and demyelinating mice models.





## **What questions still needs to be answered, what is needed in order to answer them?**

- Creating a database that can interact with The Chameleon, Namer, PACS for MRI scans and our Lab bio-repository and biomarkers levels results.
- Luminex reader for multiplex arrays.
- Available ECL documentation system in the interdepartmental equipment.
- Histopathological service for research.



# Laboratory for Early Markers of Neurodegeneration



**Anat Mirelman, PhD- PI**

Mission: identify early markers of neurodegeneration which allude to a pathological process and identify markers of progression





# Tel-Aviv PD Genetic Project

**Genetics in PD**  
**N=2851 participants**

**1612 patients with PD**

**412 healthy adults**

**827 relatives**







Anat Mirelman, PhD



- Genotype- phenotype
- Motor –cognitive reserve and interactions
- Gait and wearable sensors
- Sleep

Avner Thaler, MD, PhD



- Genotype- phenotype
- Inflammatory and Biological markers
- Potential compounds for disease modification

Inbal Maidan, PhD



- Electrophysiology Markers (EEG)
- Hemodynamic responses (fNIRS)
- Deep Brain Stimulation

Amgad Droby, PhD

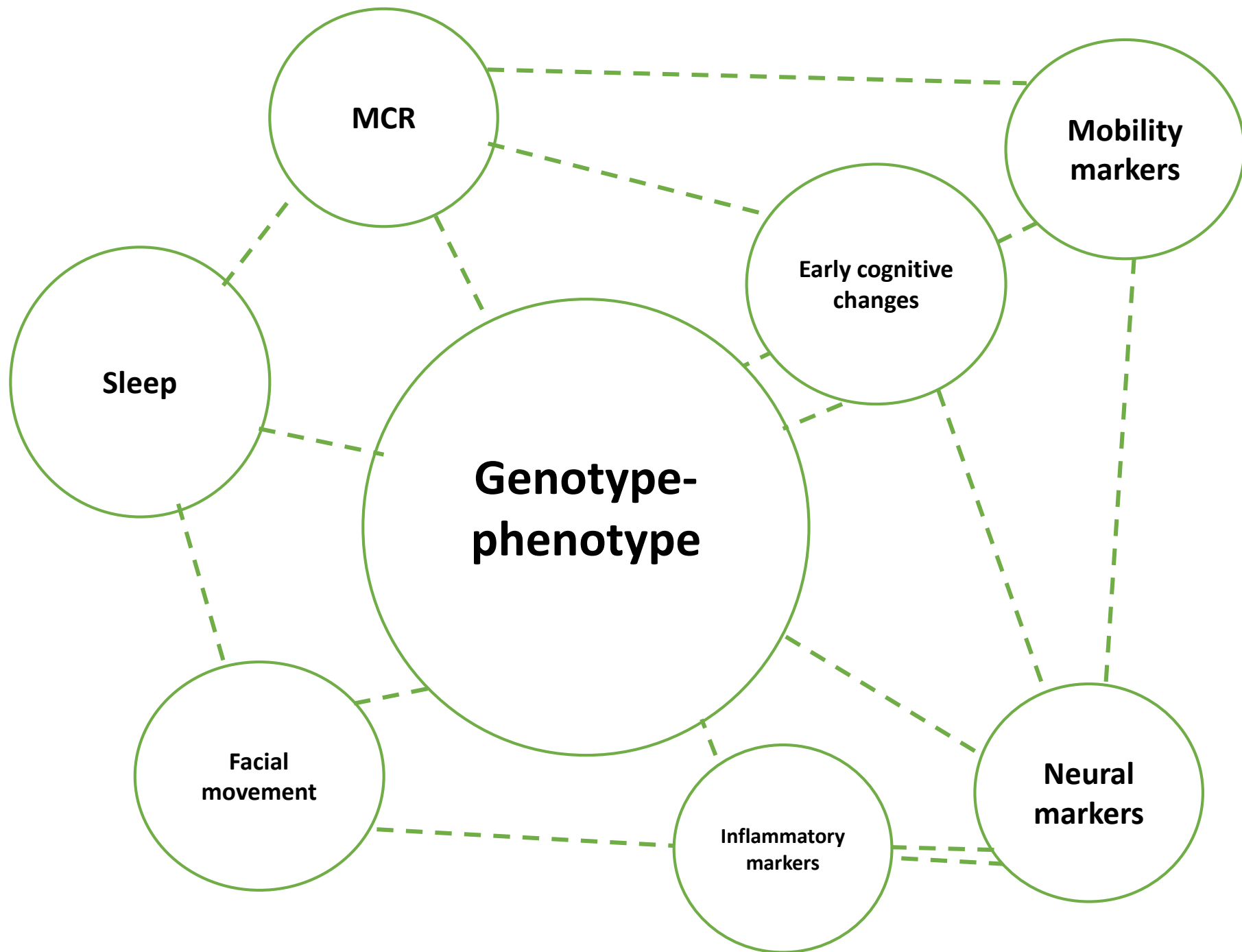


- Structural and functional markers (MRI and fMRI)
- Combining tools for neural assessment (PET, SPECT, MRI)

23 students [6 PhD, 9 MSc, 8 BSc from Sagol, Medicine, Engineering, Computer science]









LEMONS T  LEMONADE



## Key Capabilities

### What are we specialized in

- Wearable sensors for the assessment of movement
- Electrophysiology
- Multimodal integration

### What specialized equipment we use to answer Q

- Wearables
- EEG
- EMG
- fNIRS

### How can we aid other scientists to answer their Q

- We are experts in behavior and clinical phenotypes







עשיה

הצלחה

# שנה טובה

שגשוג

בריאות

אהבה

אושר



שנה טובה ומתוקה



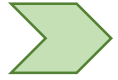


# Brain Institute



Brain Institute	Leaders	Enhancing Neuroscience research in TLVMC	Enhancing collaboration
<ul style="list-style-type: none"><li>• Neurology</li><li>• Neurosurgery</li><li>• Psychiatry</li><li>• Neuro-rehab</li><li>• Affiliated Labs</li><li>• Sagol Brain Institute</li></ul>	<ul style="list-style-type: none"><li>• Clinical directors</li><li>• Labs PIs</li></ul>	<ul style="list-style-type: none"><li>• Internal Grants - 700K (IS)</li><li>• TAU Grants<ul style="list-style-type: none"><li>• Aufzien – 1M (IS)</li><li>• Psychedelia - ?</li></ul></li><li>• Torino –TA Collab. – 800K (IS)</li></ul>	<ul style="list-style-type: none"><li>• Advertising and opening all academic activities to all Brain Institute members</li><li>• Monthly meeting</li></ul>





# Calls for Grants Timeline

- Internal grants – March 2024
- Aufzien's grants – June 2024
- Aufzien's travel grants – on going
- Torino – TA grants – March 2024
  - 1<sup>st</sup> Meeting December 10,2023 in Tel-Aviv



# TAU Synergies in Neuroscience Initiative

Nir Giladi, Tal Laviv, Bruria Ben Zeev, Nocham Wolpe



- Improving collaboration between TAU campus, its affiliated hospitals and the neuroscience industry in Israel
- Partners:
  - TAU Faculty of Medicine
  - Sagol School of Neurosciences
  - 14 affiliated hospitals
  - BrainstormIL (145 startup companies)
- 1<sup>st</sup> join meeting February 20, 2024 Smolarsh Hall
  - Neuroscience website at TAU
    - All labs will present their activities – one page



Autoimmune Neurology research laboratory

Avi Gadoth

Lab director – Valeria Briskin



# Autoimmune Neurology

- 2007
- Encephalitis, Peripheral nerves, autonomic
- Expanding
- Treatable



# Current work – 3 months

- T cell response
- Epitope mapping
- HLA correlation/association



# Future directions

- Protein production
- Synaptic transmission and mechanisms - cooperation
- Single cell RNA – mechanisms and difference from infectious
- Protein identification – Phage, proteomics...



# Cooperation

- Yifat Alcalay
- David Hagin
- Moshe Giladi
- Tamar Rubinek





# Who we are:

Name of lab/Location

## Microsurgery and Plastic Surgery laboratory

(The Plastic Surgery  
Department)

**Address:**  
**Founders building**  
6th floor  
Room 8,11

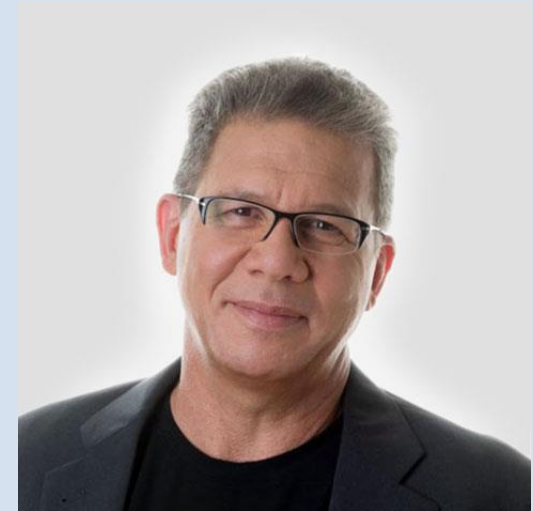
PI/Manager

Dr. Inna Solodeev, Lab Manager

Prof. Yoav Barnea



Prof. Eyal Gur





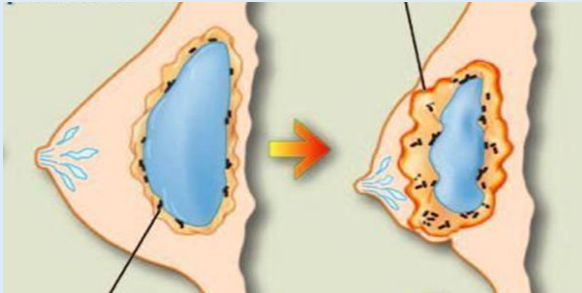
# ➤ Area of interest.

## Problems related to breast reconstruction:

### 1. Prevention and Treatment of Capsular Contracture in Implant-Based Breast Reconstruction Following Radiotherapy

Women with breast cancer who undergo postmastectomy radiation therapy has the highest risk of Capsular Contracture with 70-100% incidence rate.

Capsular Contraction

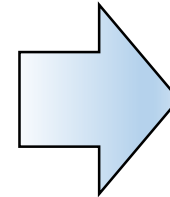


Fibroblast layer

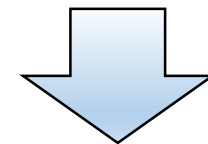
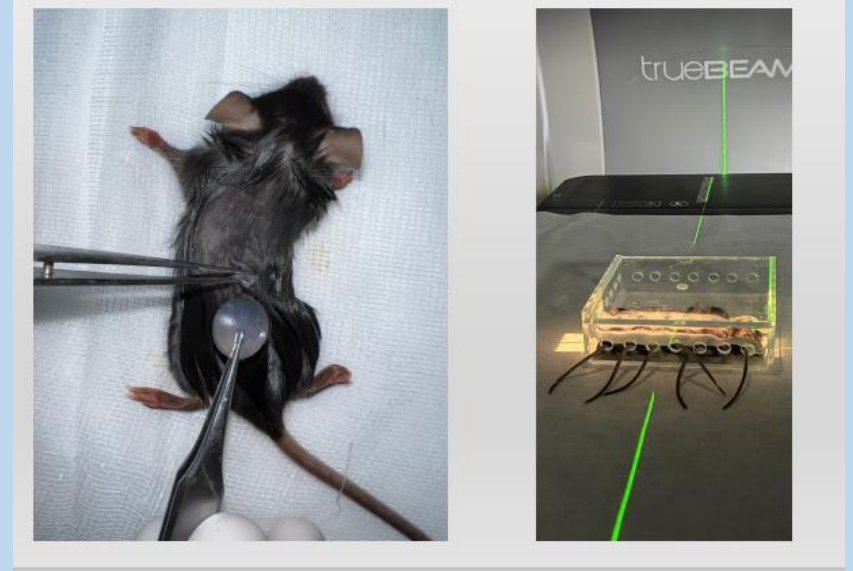
Before and after reconstruction surgery



Capsular Contracture



Mouse model of breast implant capsular contracture induced by targeted radiation

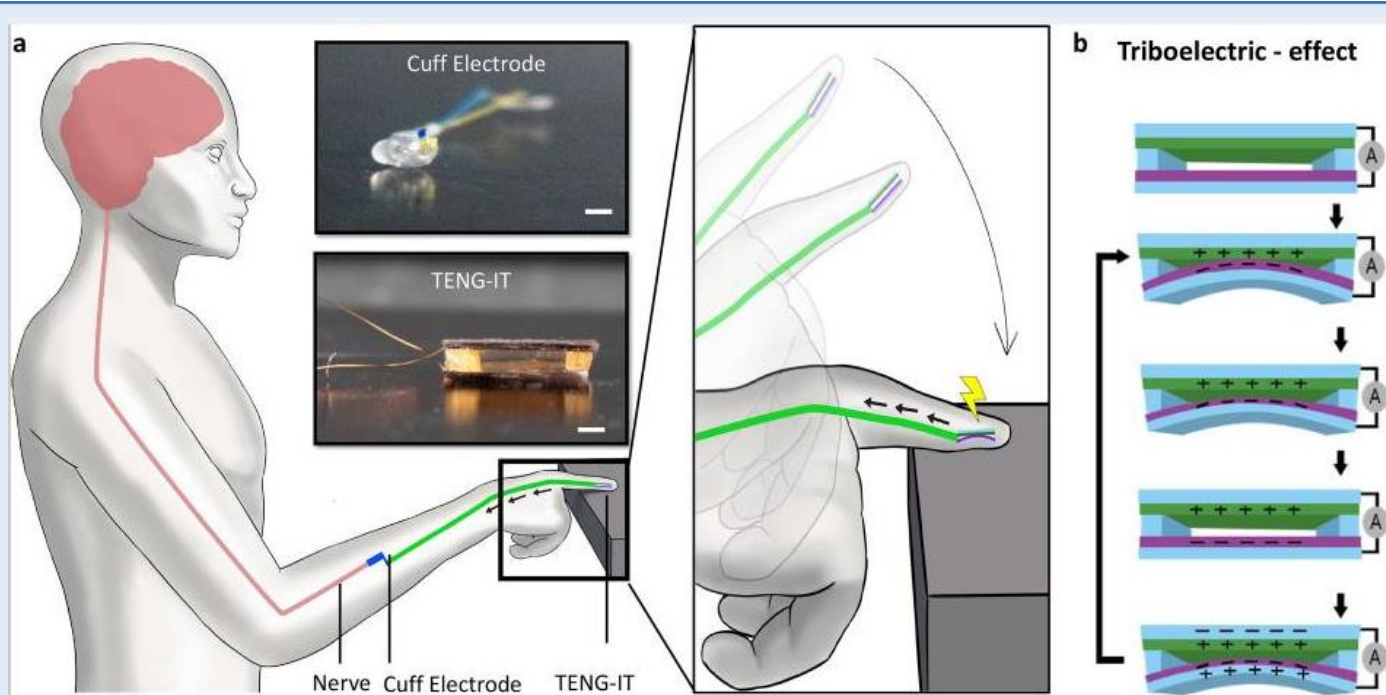


Etiology and treatment



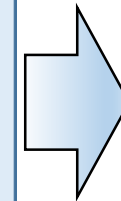
## 2. Restoring Tactile Sensing in post mastectomy breast cancer patients by a Triboelectric Nanogenerators Array

Collaboration with Tel-Aviv university



Triboelectric Nanogenerator for restoring tactile sensation.

This integrated tactile device is implanted under the skin and translates tactile pressure into electrical potential, which it relays via cuff electrodes to healthy sensory nerves, thereby stimulating them, to mimic tactile sensation.



The goal:

to restore the ability to sense touch for breast cancer post nipple-sparing mastectomies by developing

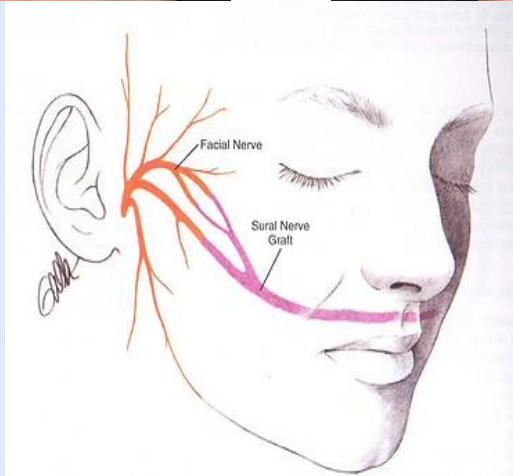
- a sensitive
- self-powered

tactile system integrated with sensory neurons.

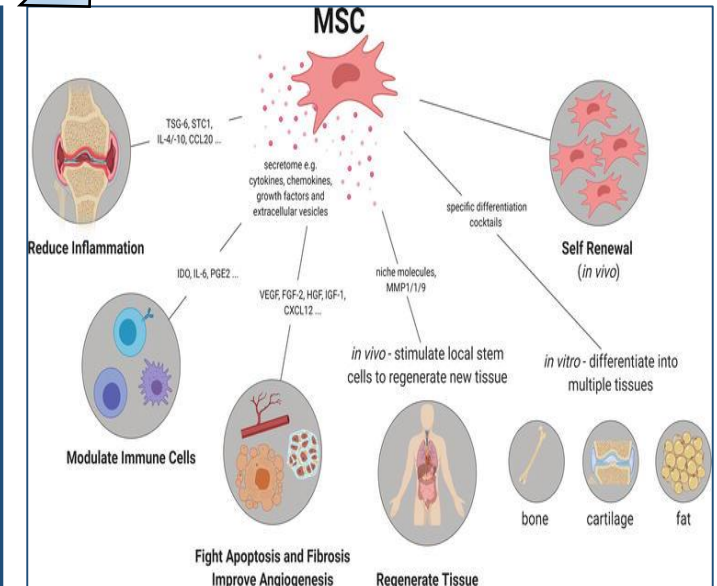
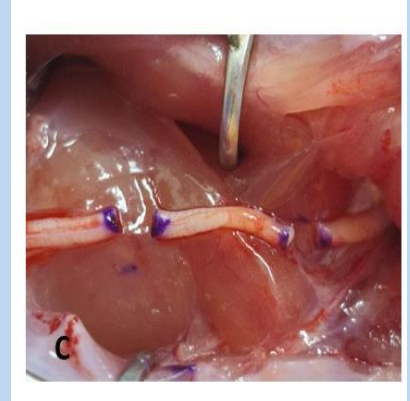
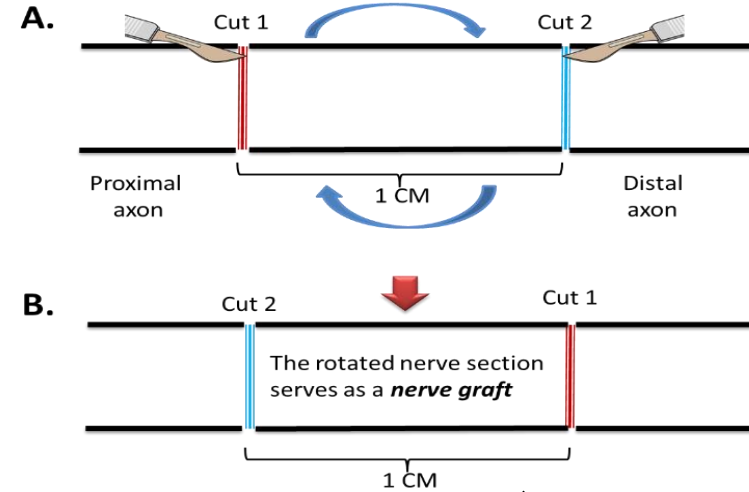


# Peripheral nerve regeneration

Facial paralysis



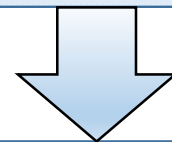
The results of these complex reconstructive procedures are dependent on the rate and quality of axonal growth through the nerve graft.





# Cryopreservation of large tissues and organs

**On day 32 days post-vitrification and initial transplantation (POD32), the rat was able to walk using the cryopreserved limb.**

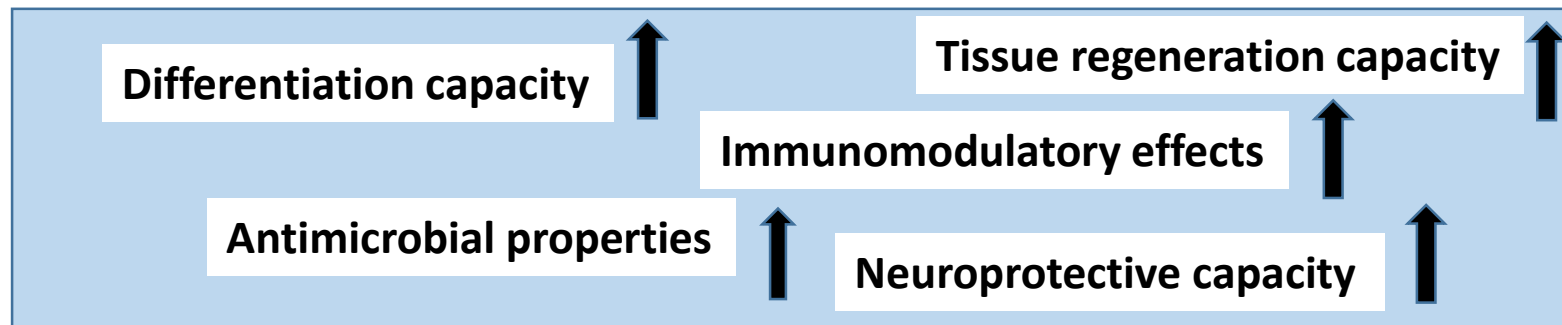
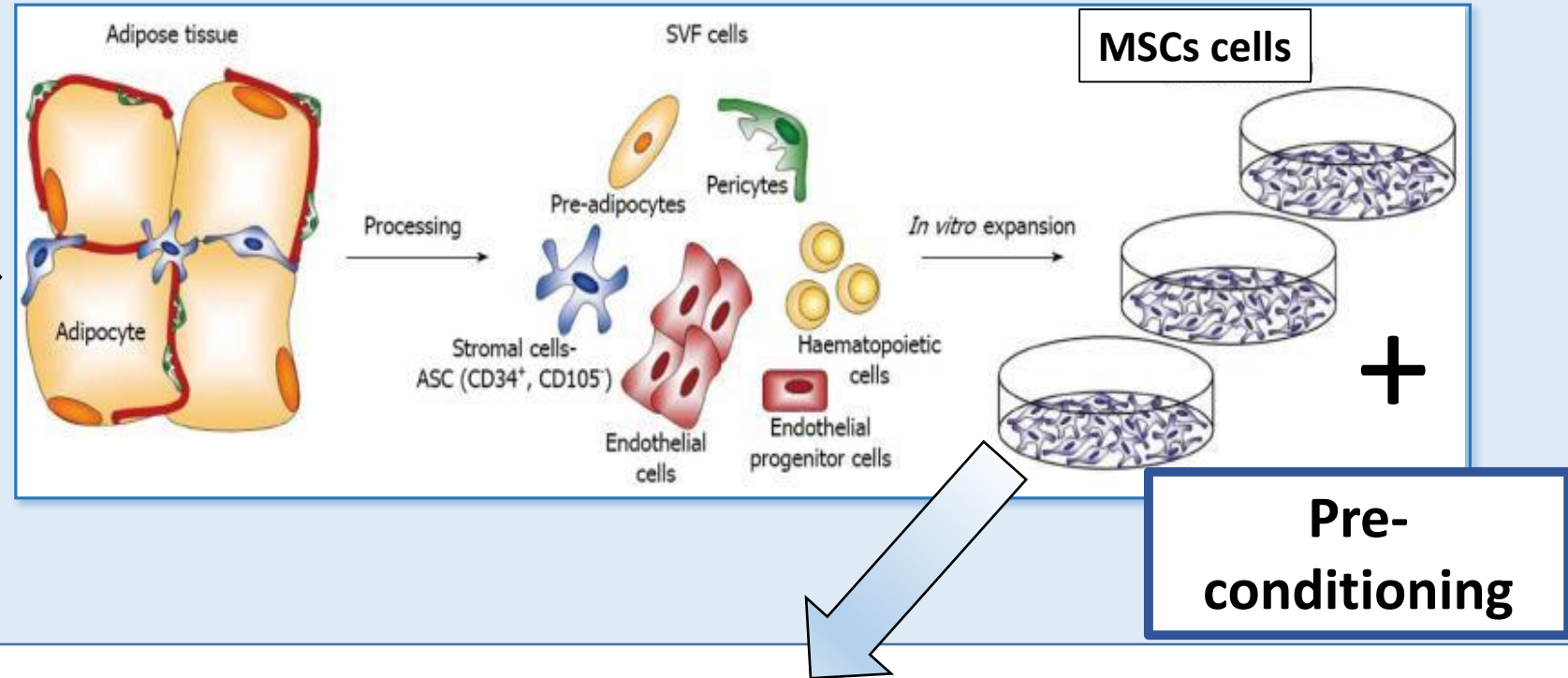
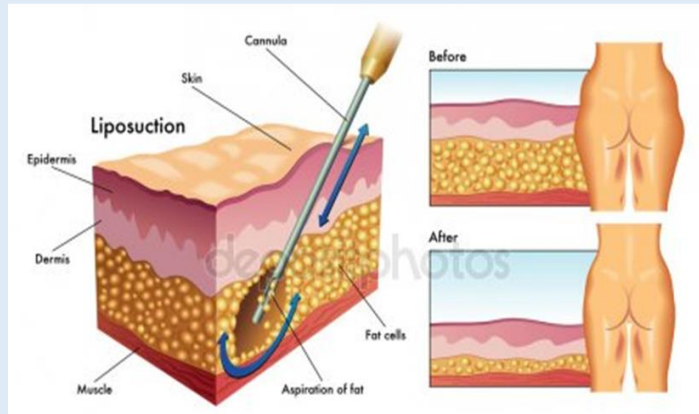


Continue to develop protocols for long-term preservation of large tissues for clinical use.



# Pre-conditioning of Human Mesenchymal Stem Cells for transplantation application

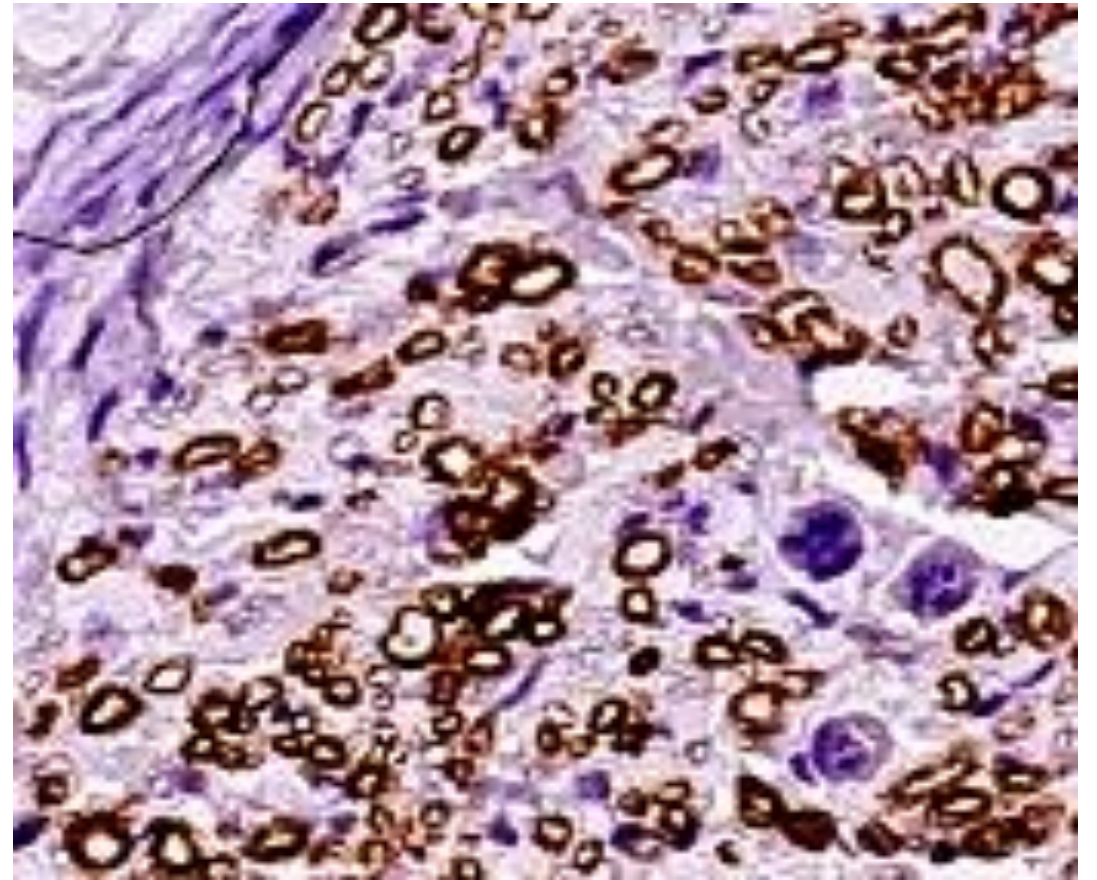
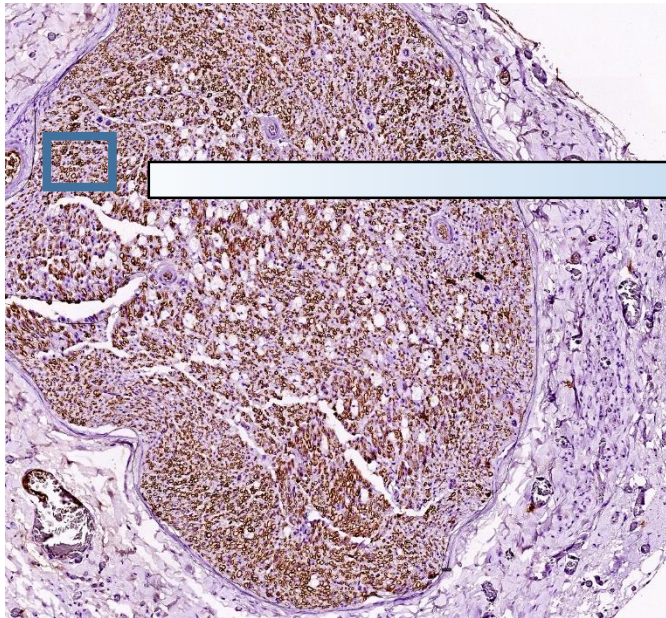
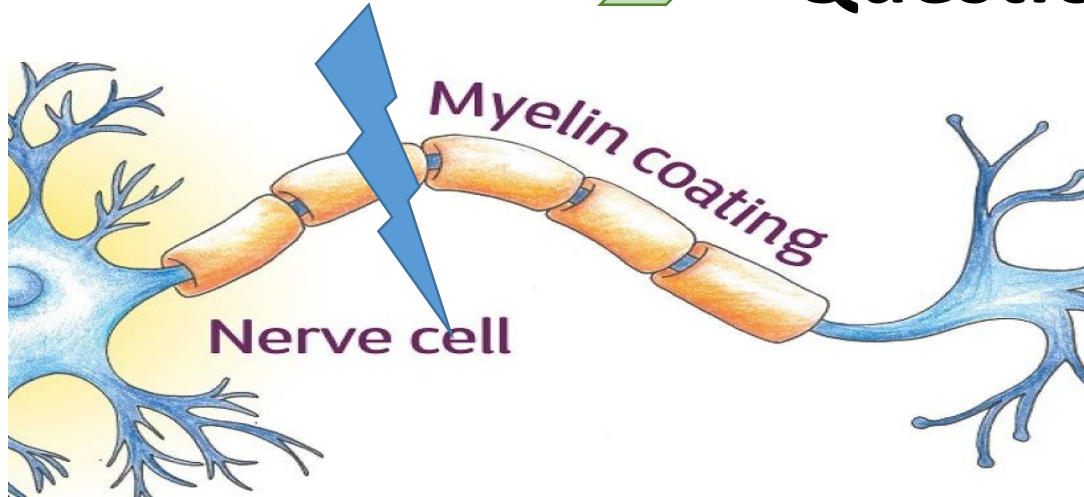
## Liposuction







Question:





## Dr. Miriam and Sheldon G. Adelson Clinic for Drug Abuse Treatment and Research



10 Dafna street



**Prof. Einat Peles** Ph.D.  
Research Director

**Anat Sason, MA**  
Research Assistant

**Yali Abramsohn**  
Clinic Director

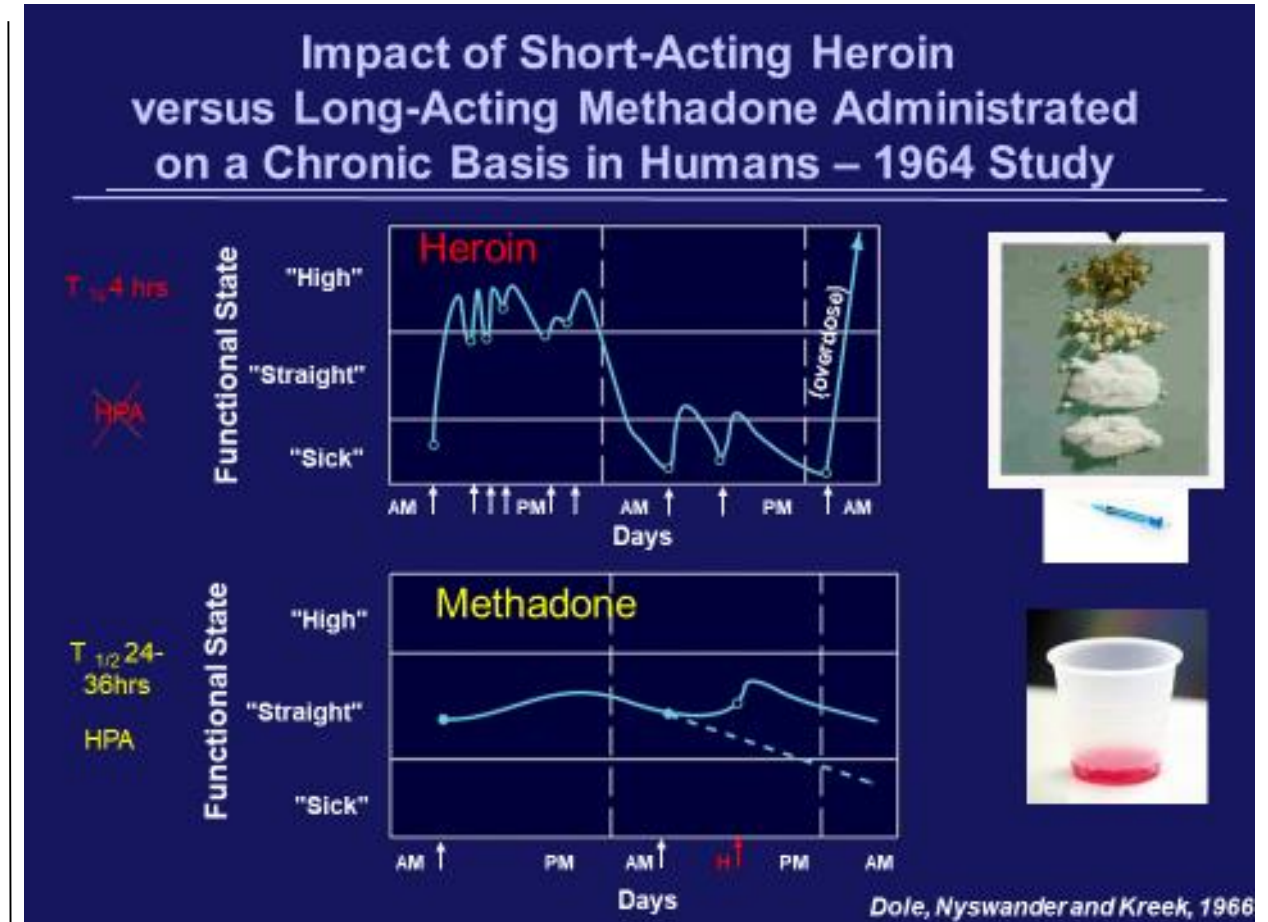
**Prof. Shaul Schreiber**  
Medical Director





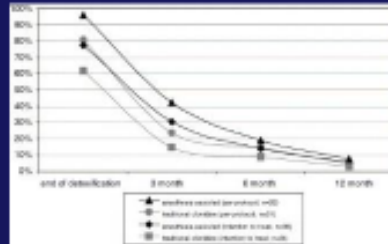
## Research Topics

- ✓ Addiction Medicine
- ✓ Opioid use disorder in Methadone Maintenance Treatment (MMT)
- ✓ Patients characteristics and outcome (predictors)
- ✓ Mental and physical comorbidity
- ✓ Pain indices
- ✓ Sleep quality
- ✓ Cognition
- ✓ Addiction risk factors
- ✓ Genetic predisposition
- ✓ Epigenetics
- ✓ Stigma, Ignorance

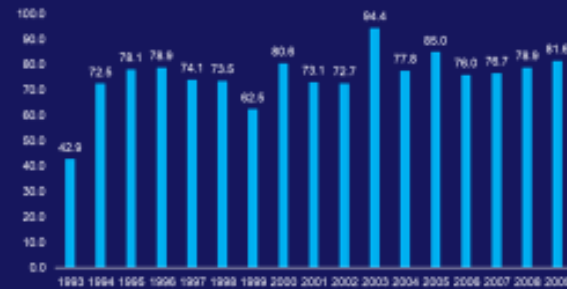




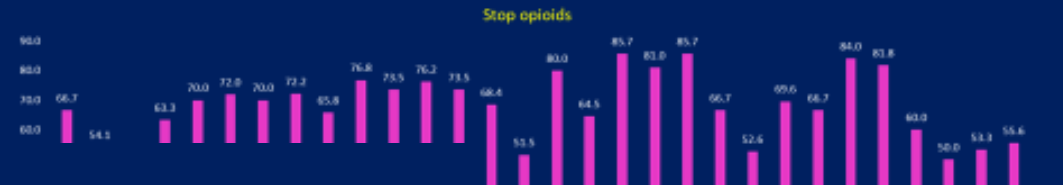
## Opioid abstinence following detoxification



Dr. Miriam & Sheldon G. Adelson Clinic  
for Drug Abuse Treatment & Research  
Tel Aviv Sourasky Medical Center

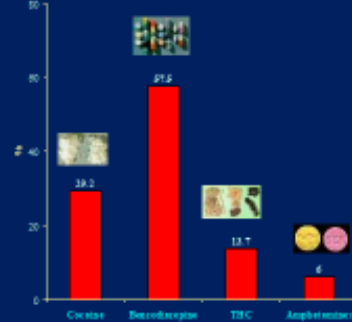


## Main goal: opioid discontinuation



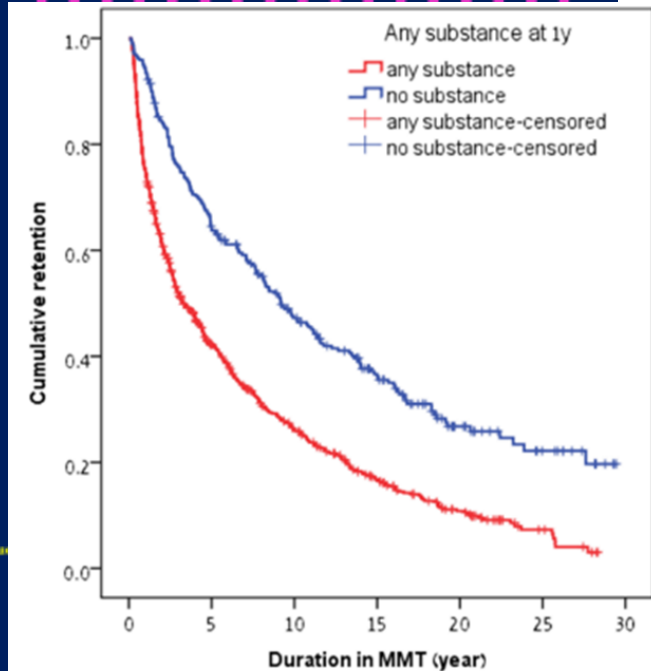
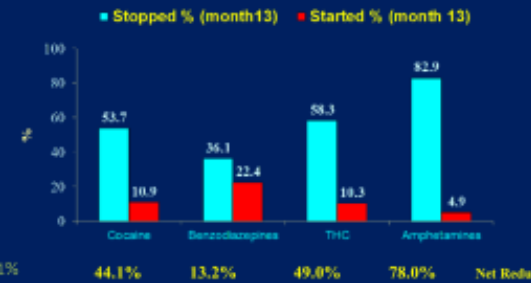
## Other substance?

### Drug Abuse on Admission (n=1120)\*



\*unknown n=4, \*\*amphetamines unknown 5.1%

### Net reduction of substance after one year

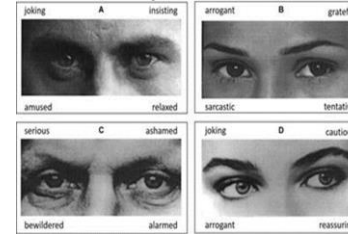




## Methodology

- Questionnaires
- computerized tasks
- Pain Algometer (pain threshold)
- Heart rate variability
- Urine test for substance
- Blood measures (methadone level)
- Genetics, Epigenetics

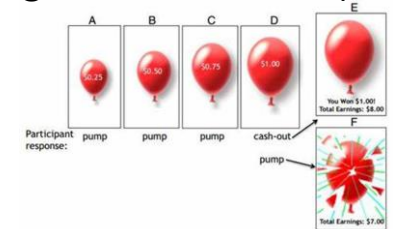
### Theory of Mind



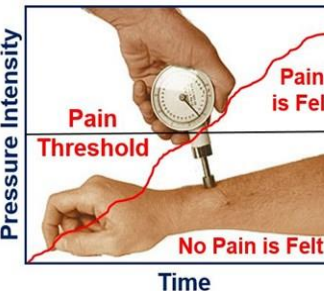
### Pain empathy



### Balloon Analogue Risk Task (BART)

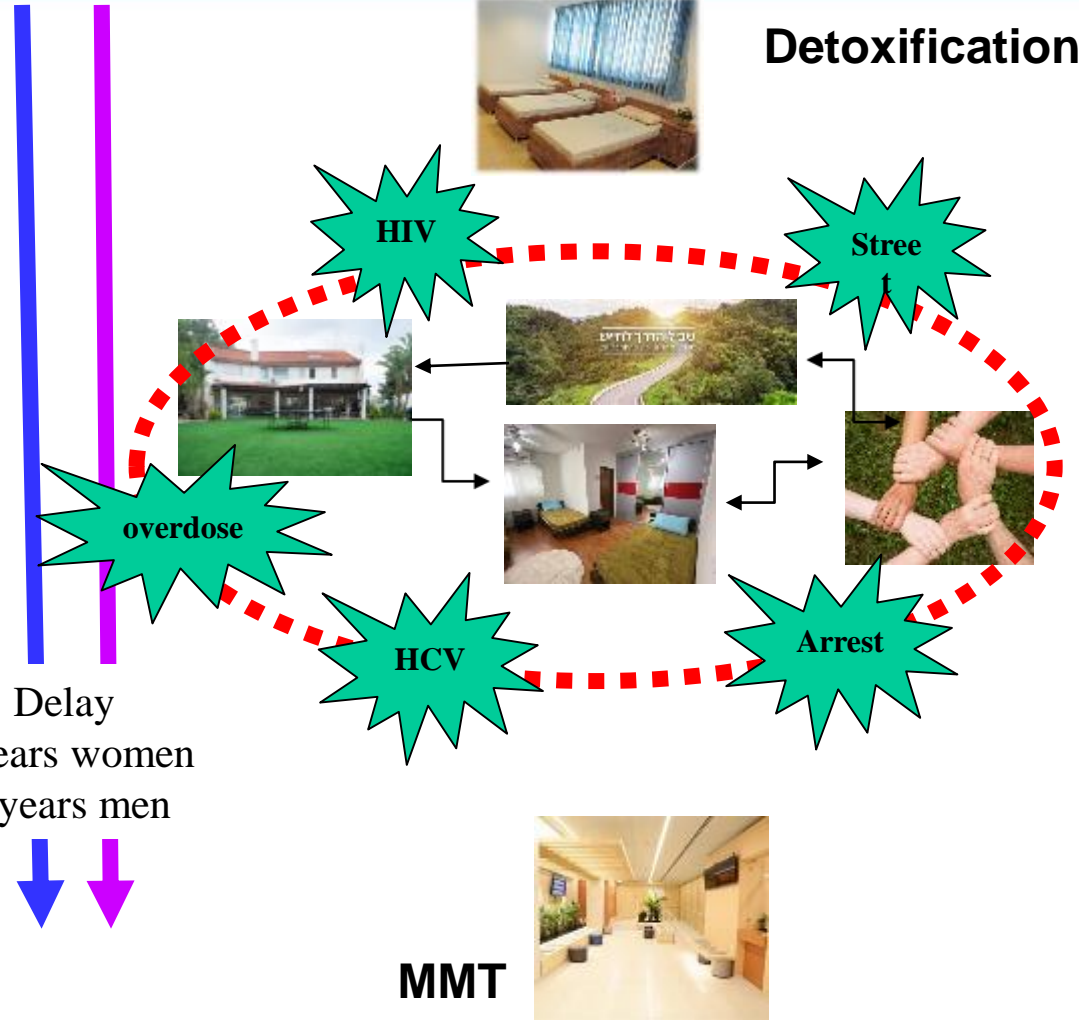
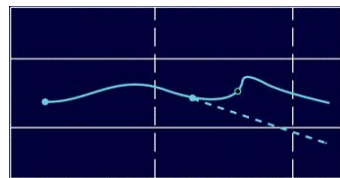
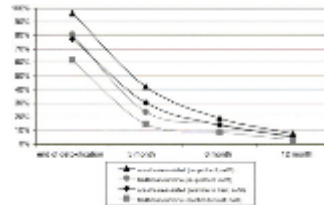
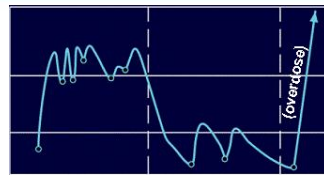


### Algometer





# Opioid Use Disorder





# ישראל מדביקה את ארה"ב

## Opioid consumption in Israel and the United States



Source: Nadav Davidovitch, Yannai Kranzler, and Oren Miron, Taub Center | Data: INCB



TEL AVIV SOURASKY  
MEDICAL CENTER  
ICHILOV



תל אביב  
אוניברסיטת תל אביב



TEL AVIV SOURASKY  
MEDICAL CENTER  
ICHILOV



תל אביב  
אוניברסיטת תל אביב



## **What questions still needs to be answered, what is needed in order to answer them?**

- Biomarker for develop addiction (grant with Prof. Segev Barak)
- Opioid epidemic- Fentanyl prevalence overdose??
- Brain imaging
- “Medical” cannabis

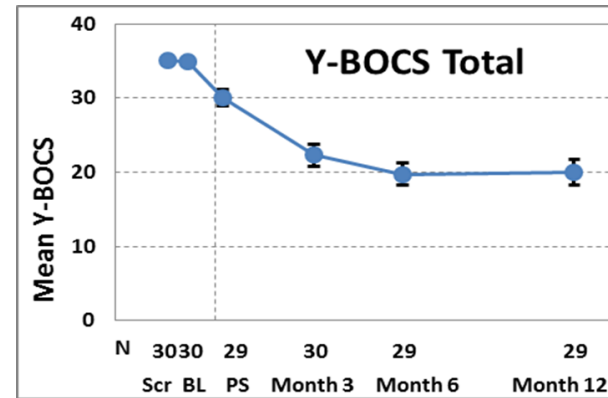




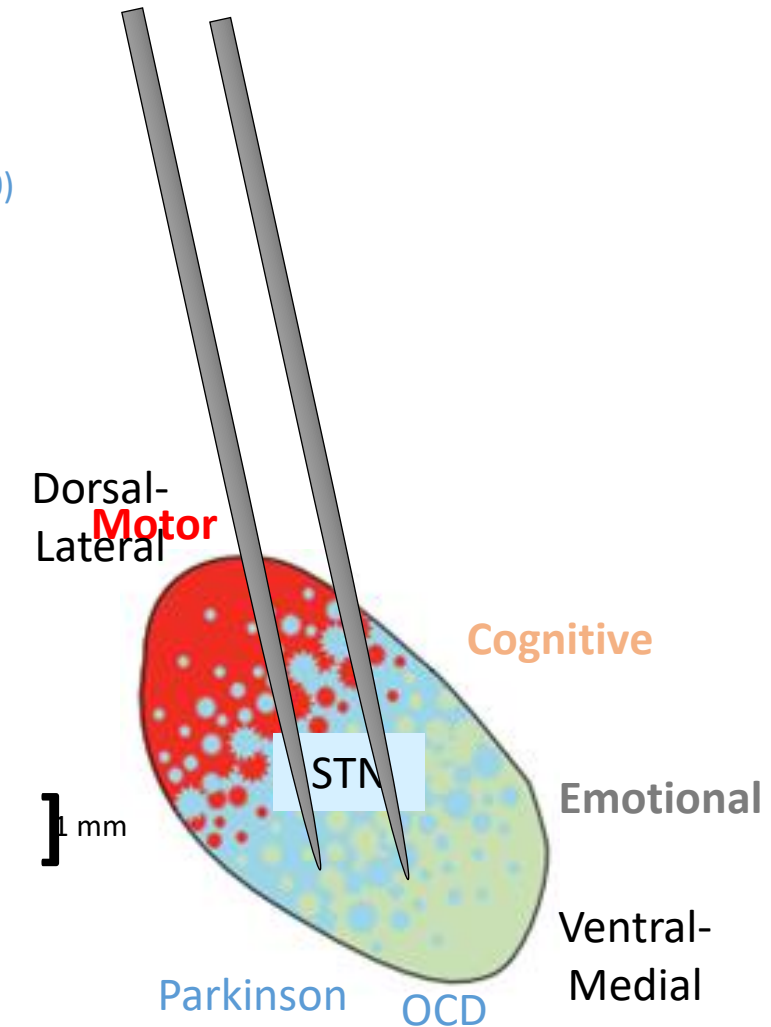
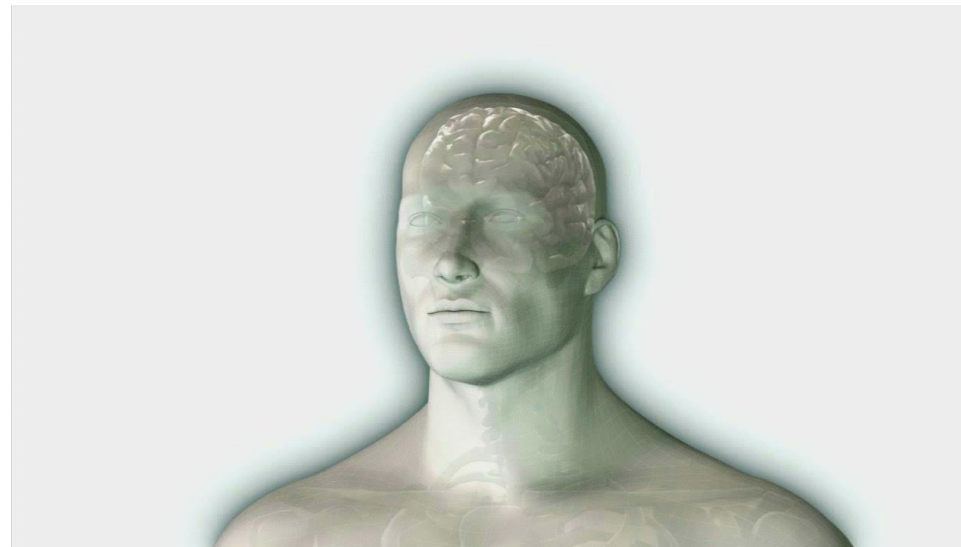


# Deep Brain Stimulation - DBS

- Movement Disorders:
  - \* Parkinson's Disease
  - \* Essential Tremor
  - \* Dystonia
- Major Depression
- **OCD**
- Tourette's Syndrome
- Addiction
- Alzheimer's Disease
- Epilepsy
- Anorexia Nervosa
- **Schizophrenia**



Menchón et al, Mol Psych, 2019 (n=30)





# Who we are and our area of interest

Name of lab/Location	PI/Manager	Main Subjects in the lab	Keep it simple to people who are not in the field
<ul style="list-style-type: none"><li>• Neuropsychiatry Lab</li><li>• Rehabilitation Building, Psychiatry Institute, Ground floor, rooms 27-30</li></ul>	<ul style="list-style-type: none"><li>• Dr. Renana Eitan (Chair of Psychiatry)</li><li>• Dr. Keren Avirame (Director of Research, Psychiatry)</li></ul>	<ul style="list-style-type: none"><li>• Electrophysiological makers of Neuropsychiatric Disorders</li><li>• Biomarkers of illness dynamics to develop personalized and adaptive neuromodulation</li></ul>	<ul style="list-style-type: none"><li>• Novel brain therapies for Neuropsychiatric disorders such as OCD, Schizophrenia and MDD</li></ul>



# ➤ Key Capabilities

## What are we specialized in

- Cortical and Sub-cortical Electrophysiology
- Disease-specific neuropsychological tasks
- Multivariate models pathological behaviors and therapy-induced effects

## What specialized equipment we use to answer Q

- Brain stimulation (DBS)
- Various electrophysiological recordings and analyses : single-unit, LFP, EEG.
- Various cognitive, emotional and social tasks

## How can we aid other scientists to answer their Q

- ISF Precise Medicine - Neuropsychiatry Group in TASMC with Dr. Inbal Meidan and Dr. Genela Morris
- Innovation Authority – Closed-loop adaptive brain stimulation in Schizophrenia with Prof. Leon Deouell
- Part of ReTune DFG Consortium – led by Prof. Kuhn (Charite Berlin)
- Collaboration with MRC Brain Network Dynamics Unit in Oxford – Prof. Denison



## **What questions still needs to be answered, what is needed in order to answer them?**

- To collect and organize a database of patients operated in Ichilov and recorded for a long period of time during different states (Sleep, Tasks, Exercise, Rest).
- To explore and develop motor, cognitive and emotional tasks.
- To group experts in electrophysiology.
- To collaborate with engineers to advance close-loop algorithm.



### Instructions:

1. Presentation in English, you can talk in Hebrew
2. **Five** minutes no longer, you will be stopped
3. use these slides, additional slides can be put during the time frame of 5 minutes
4. No need to show data the purpose is the **Ecosystem in Ichilov**
5. Keep in mind, how will I Know other labs. In order to make more collaborations



# Who we are and our area of interest

Name of lab/Location	PI/Manager	Main Subjects in the lab	Keep it simple to people who are not in the field
<ul style="list-style-type: none"><li>• Immune Diagnostic Laboratory</li><li>• The Clinical Immunology laboratory – Sourasky 1<sup>st</sup> floor, Agaf Z.</li></ul>	<ul style="list-style-type: none"><li>• David Hagin</li><li>• Freund Tal</li><li>• Yifat Alcalay</li></ul>	<ul style="list-style-type: none"><li>• Studying patients with suspected primary immunodeficiency disorders (AKA Inborn Errors of Immunity – IEIs).</li><li>• Identifying anti-viral T-cells.</li></ul>	<ul style="list-style-type: none"><li>• We are trying to better understand the effect of certain defined immune defects on the function of the immune system.</li><li>• We are trying to promote donor-derived anti-viral T-cell therapy</li></ul>



# ➤ Key Capabilities

## What are we specialized in

- Cell cultures.
- Flow-cytometry.
- Developing functional assays for evaluating immune response (mostly phospho-flow).

## What specialized equipment we use to answer Q

- Flow cytometry.
- Western.

## How can we aid other scientists to answer their Q

- Functional assays.
- Patients with defined monogenic disorders ('experiments of nature').



## **What questions still needs to be answered, what is needed in order to answer them?**

- Novel candidate genes.
- (Many of you are already helping).
- KI / KO and transfections.
- Genetics / single cell analysis.



**Dalit Ben-Yosef**

**Stem Cell Research lab, Fertility & IVF Institute**

**CORAL – Center of Regeneration and Longevity**



# The Wolfe PGD-Stem Cell Research Lab in the Institution of Reproduction and IVF

[R&D](#) > [Laboratories](#) > The Wolfe PGD-Stem Cell Research Lab - Prof. Dalit Ben Yosef



[Our vision](#)



[Research](#)



[Gallery](#)



[Our team](#)



[Current funding](#)



[Publications](#)



[From the press](#)



[Contact us](#)

## >> Our Vision

During fertilization, the human sperm and egg unite to form the developing fetus. These early phases of preimplantation development are considered one of the most fundamental questions in cell biology.

Our research lab focuses on deciphering these initial stages of embryonic development in order to understand how these processes are controlled in normal development and what happens as they stray from it, which leads to severe genetic diseases.

Our research model include human embryonic stem cells (hESC) that we derive directly from diseased embryos in order to study the mechanisms underlying the development of genetic diseases.

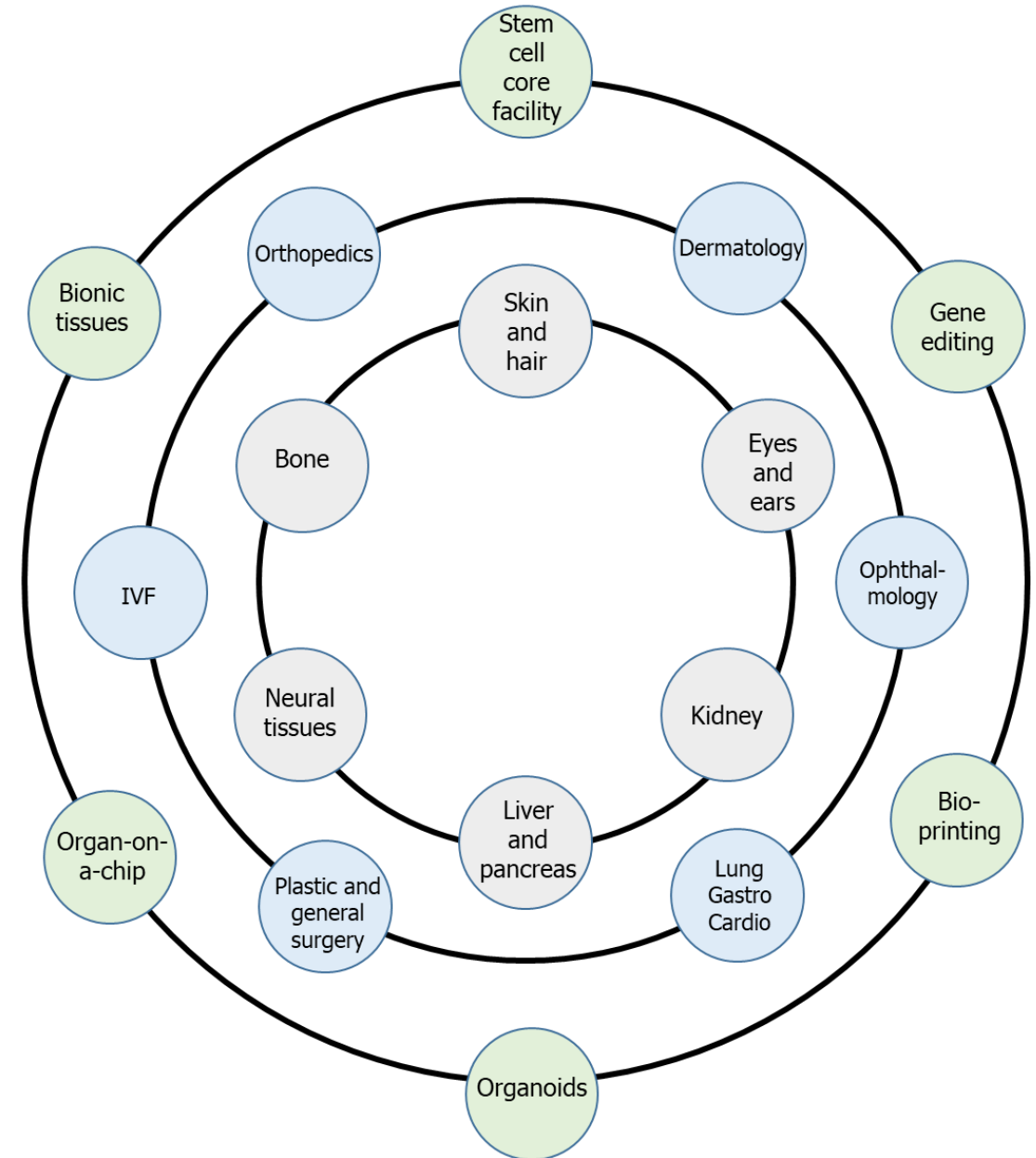




**CORAL**

Center for Regeneration And Longevity

**Implanting relevant and advanced technologies onto existing structures**





# Who we are and our area of interest

Name of lab/Location	PI/Manager	Main Subjects in the lab	Keep it simple to people who are not in the field
<ul style="list-style-type: none"><li>• Stem Cell Research lab, Fertility &amp; IVF Institute</li><li>• CORAL – Center of Regeneration and Longevity</li></ul>	<ul style="list-style-type: none"><li>• <b>Dalit Ben-Yosef</b></li><li>• Dep. Of Cell &amp; Developmental Biology &amp; Sagol School of Neuroscience, TAU</li></ul>	<ul style="list-style-type: none"><li>• Embryo Development</li><li>• Stem Cells</li><li>• Regeneration</li><li>• Genetic diseases</li></ul>	



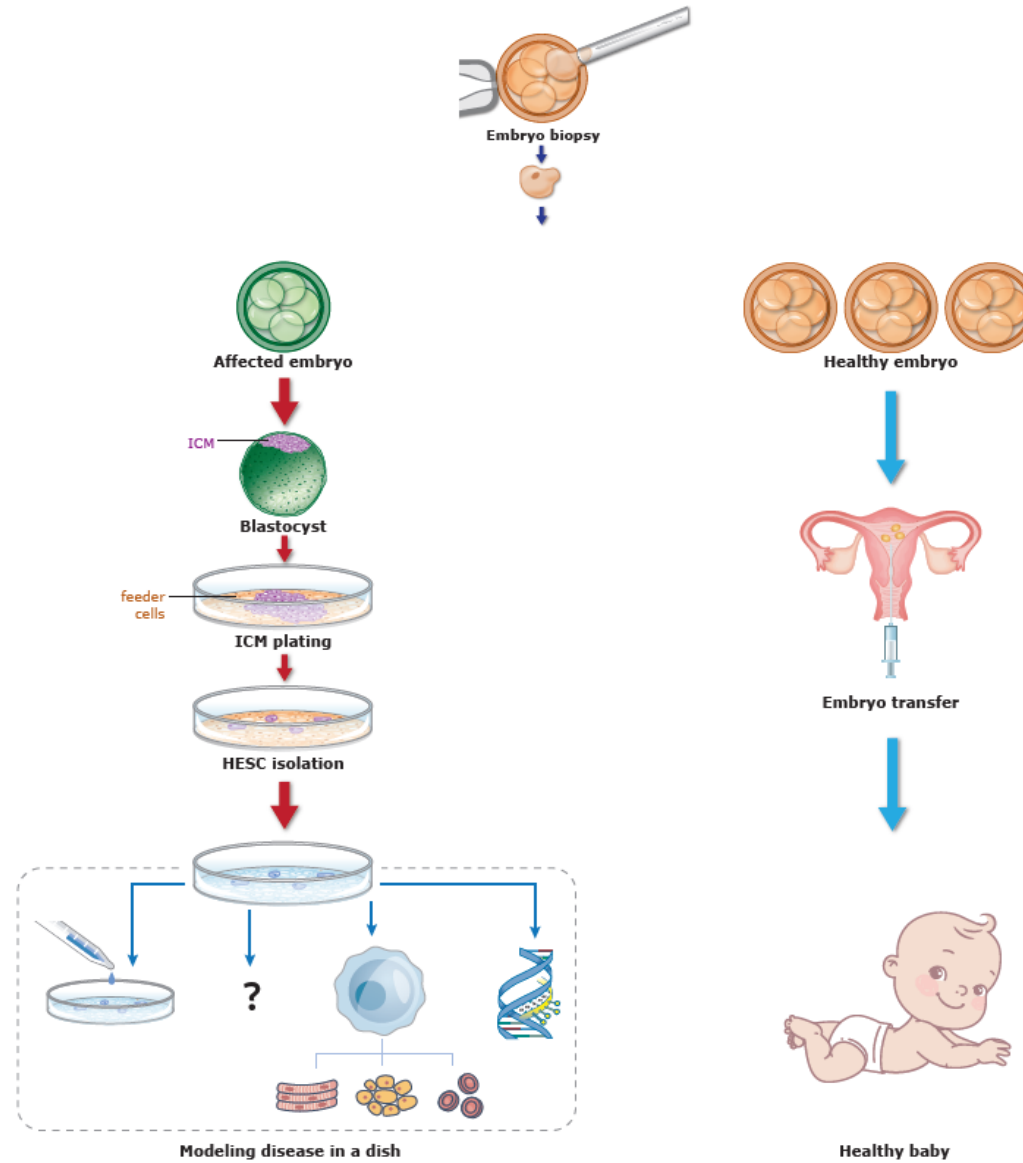
# PGD & Diseased hESC lines

iPs

Differentiation

Organoids

Genotype-phenotype





# ➤ Key Capabilities

## What are we specialized in

- Embryo micromanipulation
- ES & iP
- Stem cells & pluripotency
- Stem cell differentiation in 2D & 3D (Organoids)
- CRISPR
- Neural differentiation
- Differentiation into colon organoids

## What specialized equipment we use to answer Q

- Laser micromanipulation
- single-cell multiplex-nested PCR analysis . Simultaneous amplification of up to 30 DNA loci in a single multi-nested PCR
- Fluorescence Microscope
- Molecular analysis
- Bulk RNAseq
- Functional assays

## How can we aid other scientists to answer their Q

"שאל בני ונען"...





## **What questions still needs to be answered, what is needed in order to answer them?**

- **Bioinformatics - mainly for analyzing RNAseq data**
- **Ca<sup>2+</sup> imaging**
- **MEA – Micro Electrode Array**





# Ophthalmology stem cells lab

## Tel Aviv medical center

### Research Directors

Prof. Adiel Barak MD

Dr. Aya Barzelay MD PHD

### Group Members

Moshe Benhamou, PhD

Yahel Shechter, M.sc. student

### Physicians

Prof. Igal Leibovich MD

Dr. Benjamin Shalev MD

Dr. Ran Ben-Cnaan MD

Dr. Ilan Feldman MD

### Collaborations

Prof. Tal Dvir, laboratory of tissue engineering and regenerative medicine, Tel Aviv university

Prof. Eyal Gur MD, Dr. Nir Shani PhD, DR. Benjamin Meilik MD, Department of Plastic and reconstructive surgery



# ➤ Who we are and our area of interest

Name of lab/Location	PI/Manager	Main Subjects in the lab	Keep it simple to people who are not in the field
<ul style="list-style-type: none"><li>• Stem Cells Ophthalmology laboratory</li><li>• Tel Aviv Sourasky Medical Center</li></ul>	<ul style="list-style-type: none"><li>• Barak Adiel</li><li>• Barzelay Aya.</li></ul>	<p>ADIPOSE-DERIVED MESENCHYMAL STEM CELLS therapy for macular degeneration</p> <p>3 D printing and implanting of retinal tissue</p>	<p>Replacing dead RPE cells to prevent blindness</p>



# Key Capabilities

## What are we specialized in

- RPE cells transformation from mesenchimal cells
- 3D printing of retinal tissue
- Implanting 3D tissue into subretinal area

## What specialized equipment we use to answer Q

- 3D printing collaboration with Tal Dvir lab
- Complex vitreo retinal surgeries in Biotech farm lab.

## How can we aid other scientists to answer their Q

- Ophthalmic expertise including cornea and retinal knowledge.



## **What questions still needs to be answered, what is needed in order to answer them?**

- Refine our work on RPE transformation for ADIPOSE-DERIVED MESENCHYMAL STEM CELLS
- Collaboration with other labs to implament photoreceptor cells into 3D retinal organ
- Implantation of retinal organ in live pigs and showing functionality.
- Conversion our work to GMP standard to transfor lab woprk to first in human





# 3D bioprinting of a vascularized retina

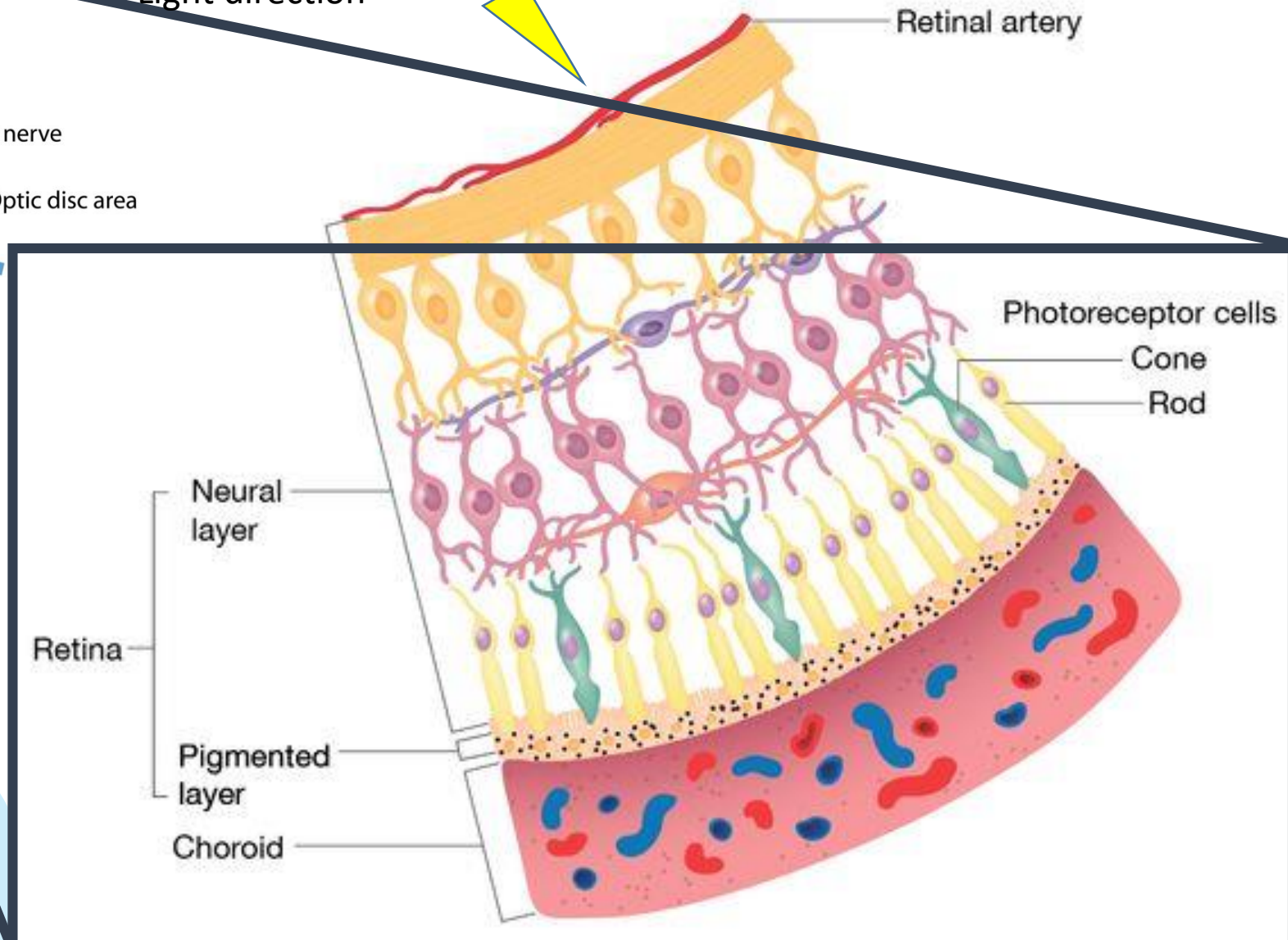
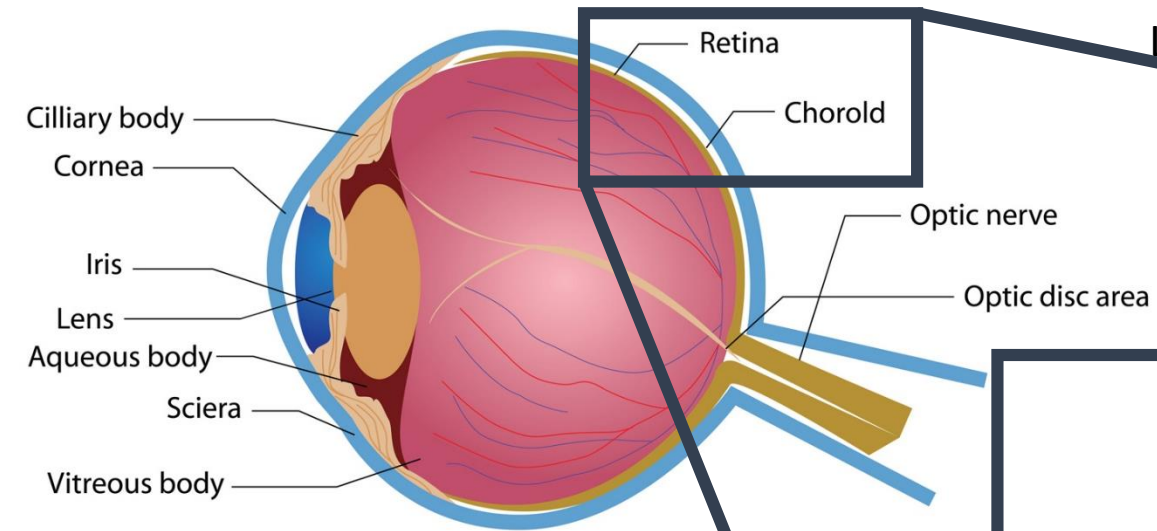
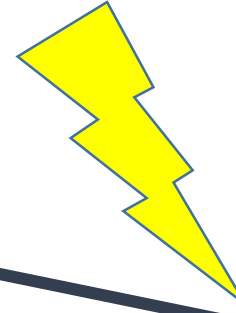


Yahel Shechter



# Retina

Light direction





# AMD

👁️ **Age-related Macular Degeneration**

👁️ Chronic disease of the central retina

👁️ The major cause of vision loss worldwide

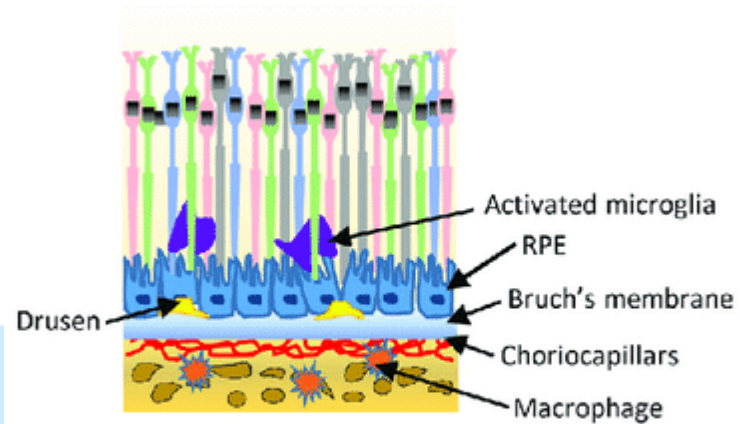
👁️ Dry (non-exudative)

👁️ 90% of cases of the disease

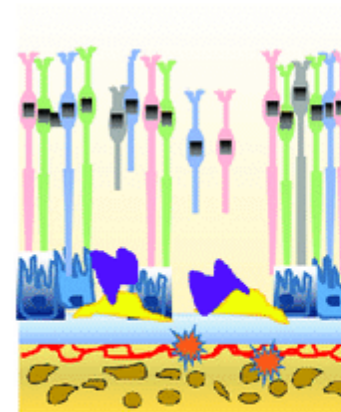
👁️ deterioration the photoreceptors and RPEs

👁️ currently has no treatment

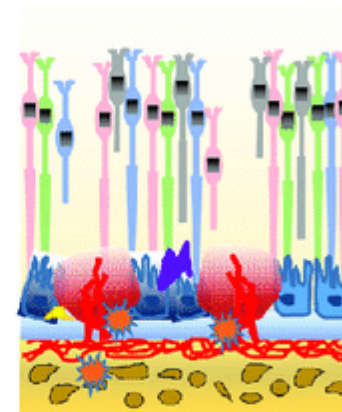
👁️ Wet (exudative)



Early AMD



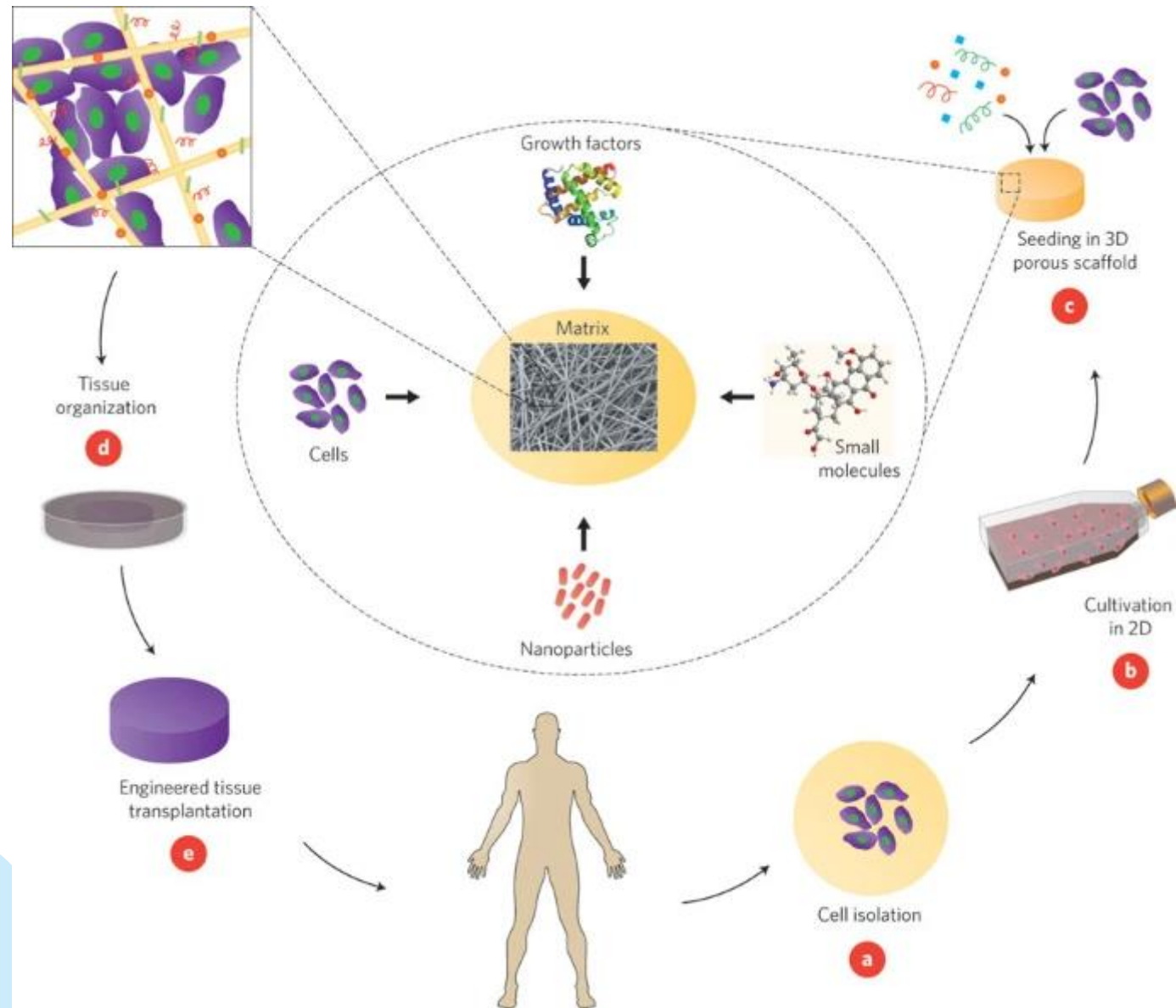
Geographic atrophy  
'Dry' AMD



Neovascular AMD  
'Wet' AMD

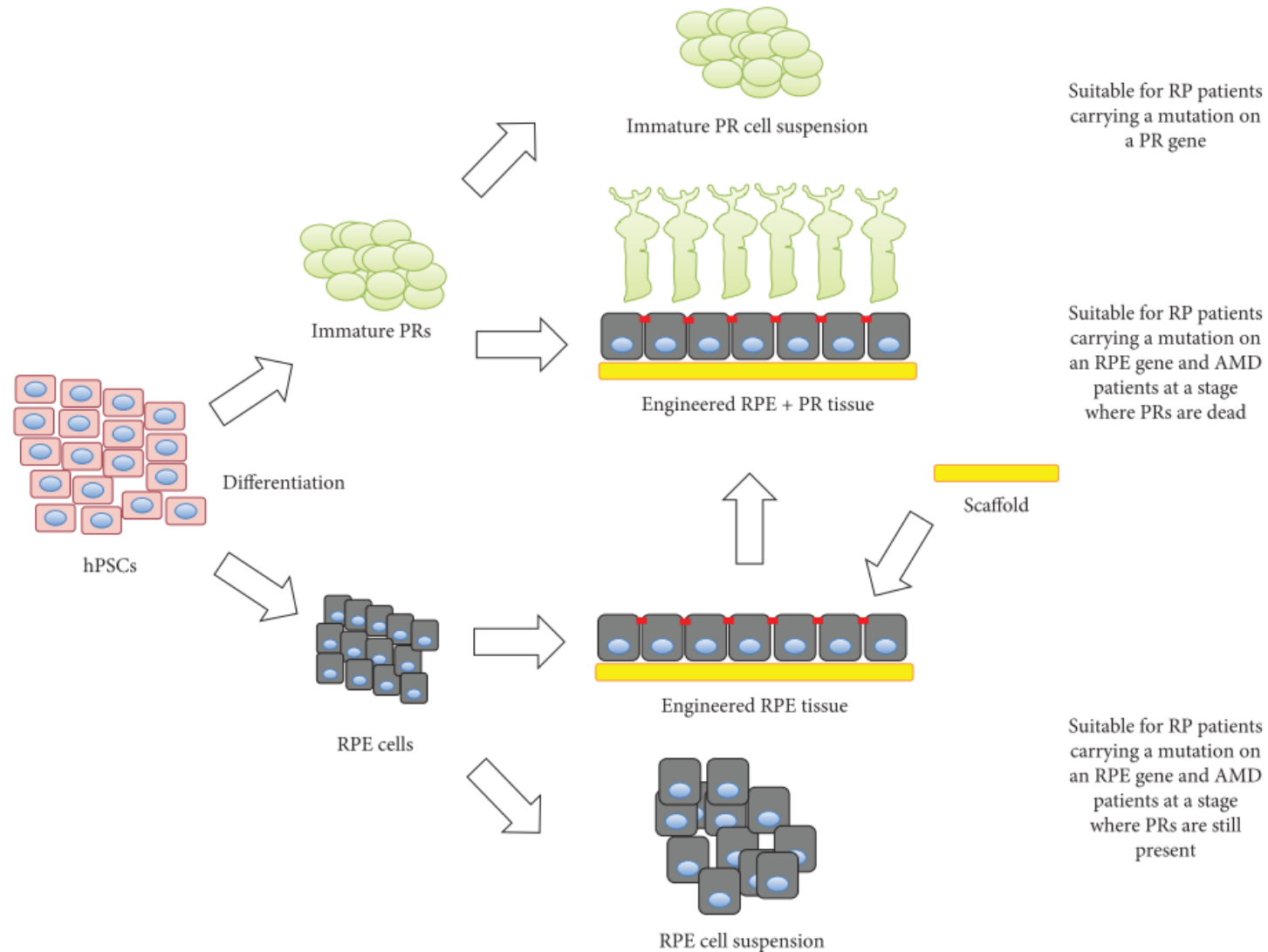


# Tissue engineering



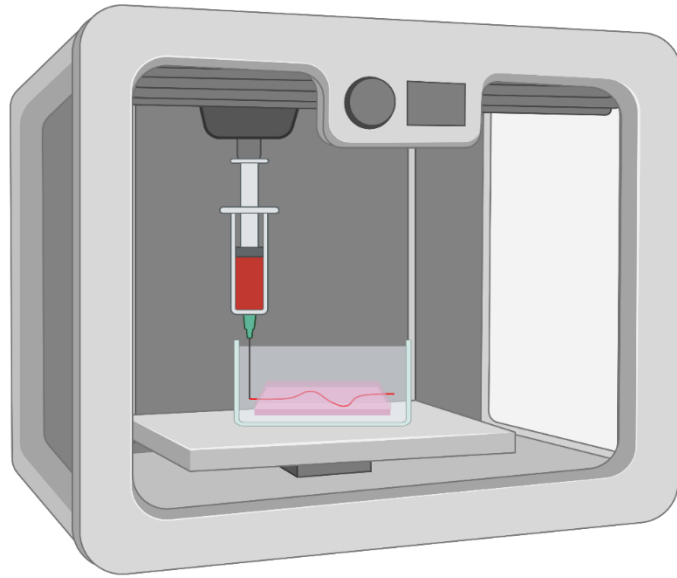


# Tissue engineering of the retina

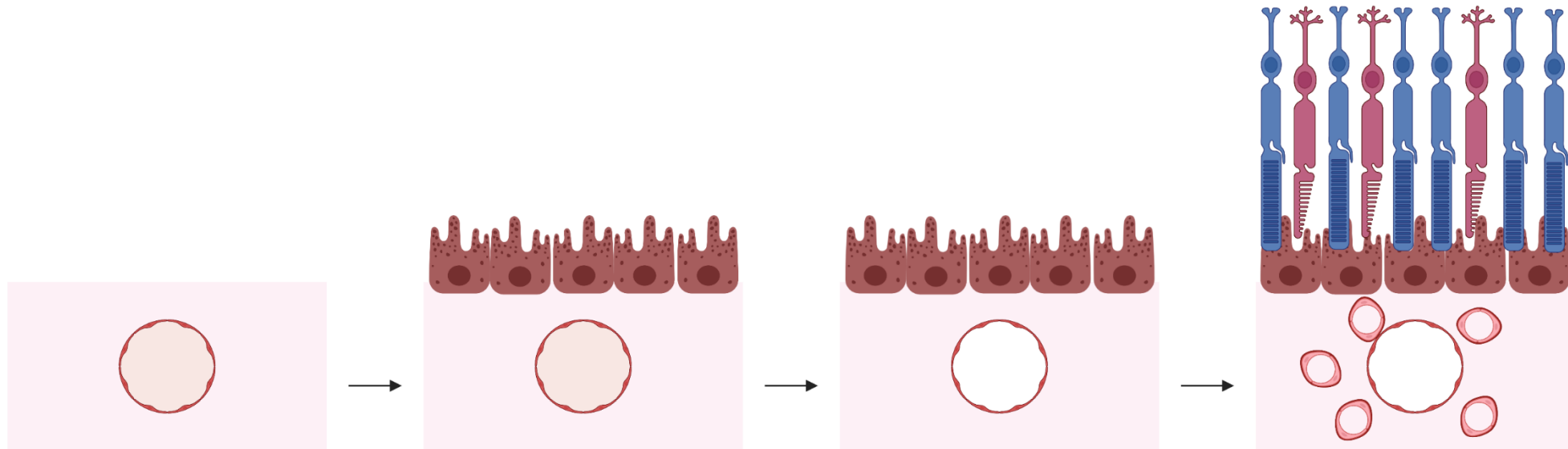
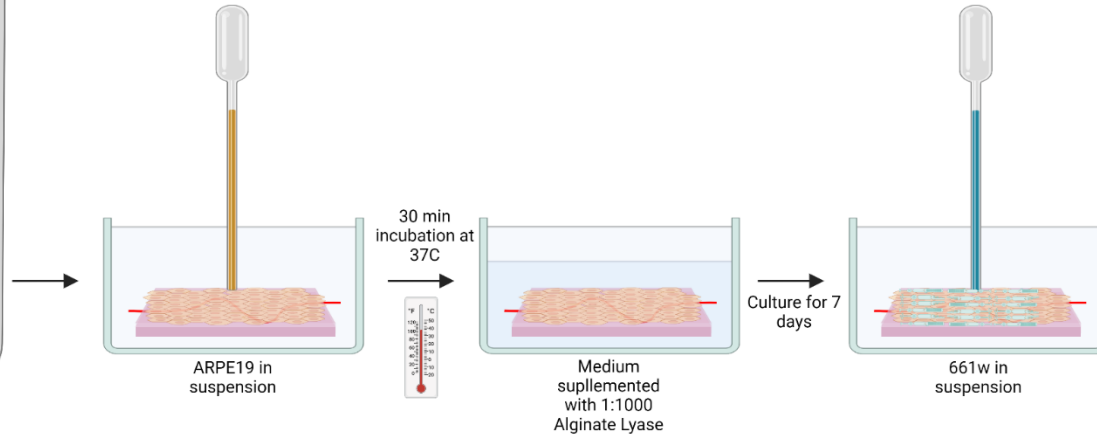




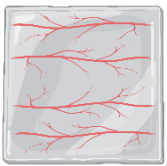
# Design



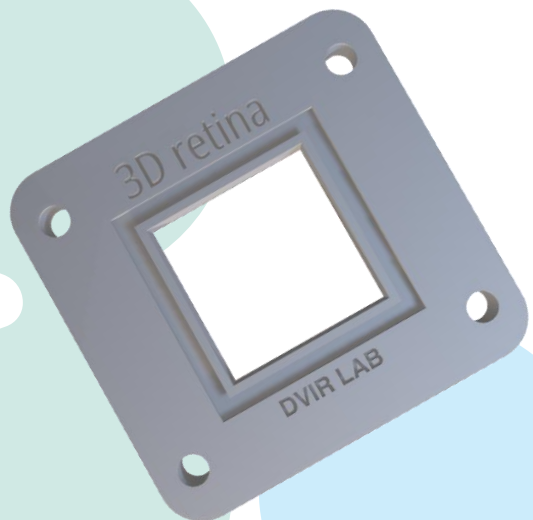
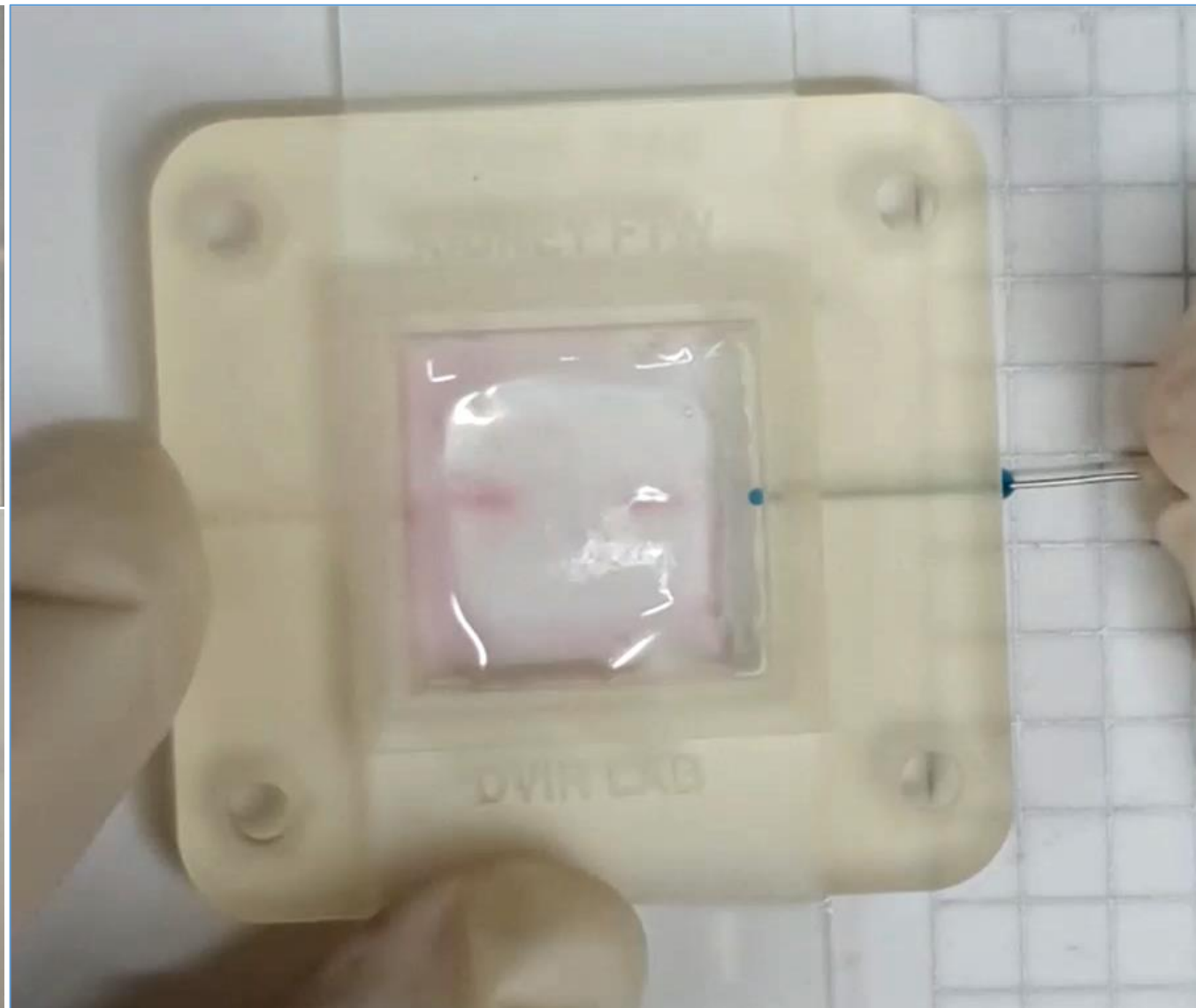
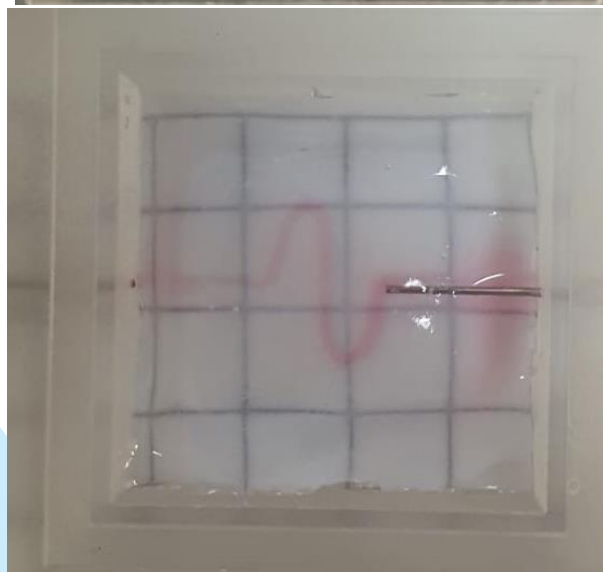
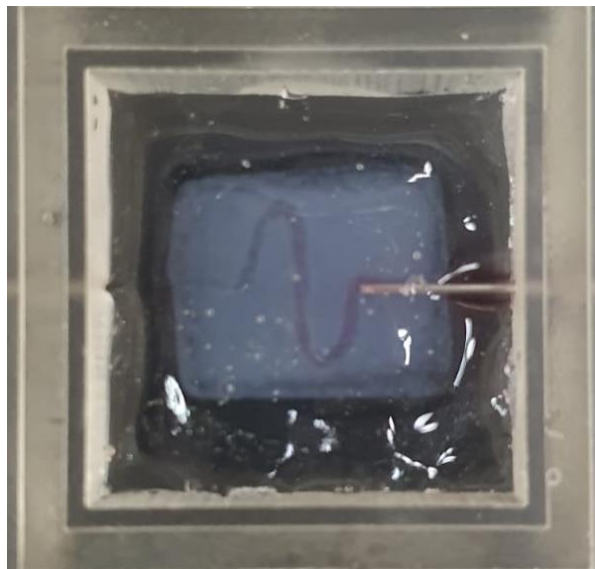
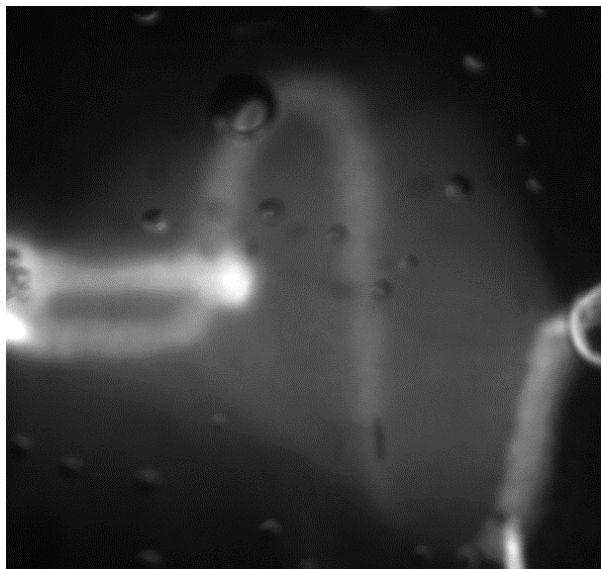
HUVEC in supporting material



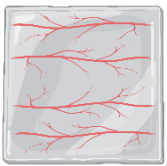




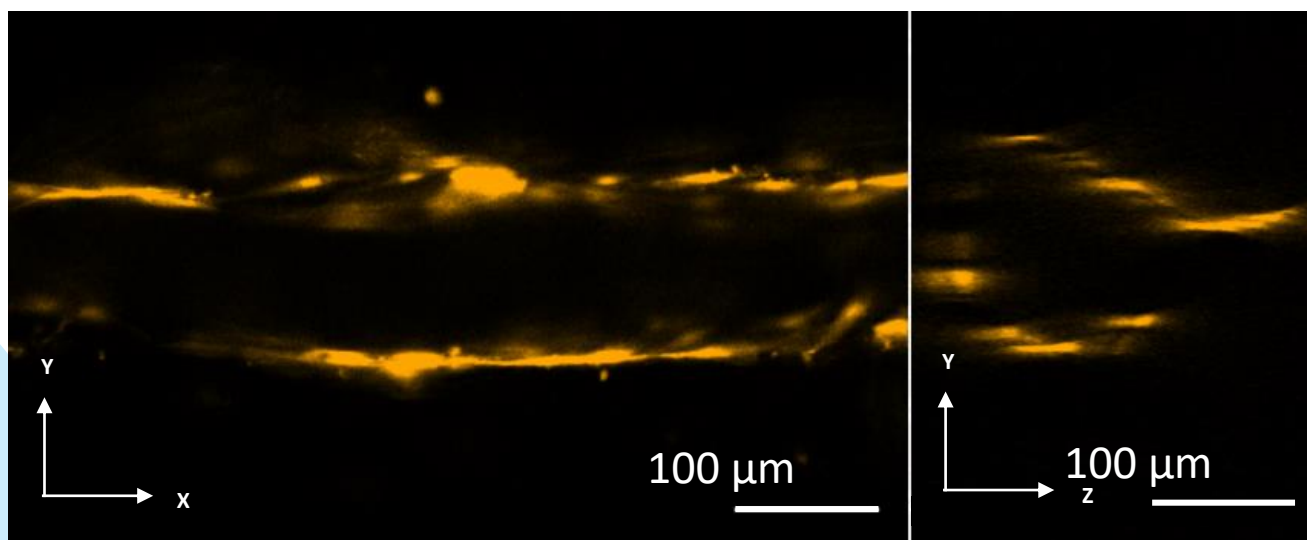
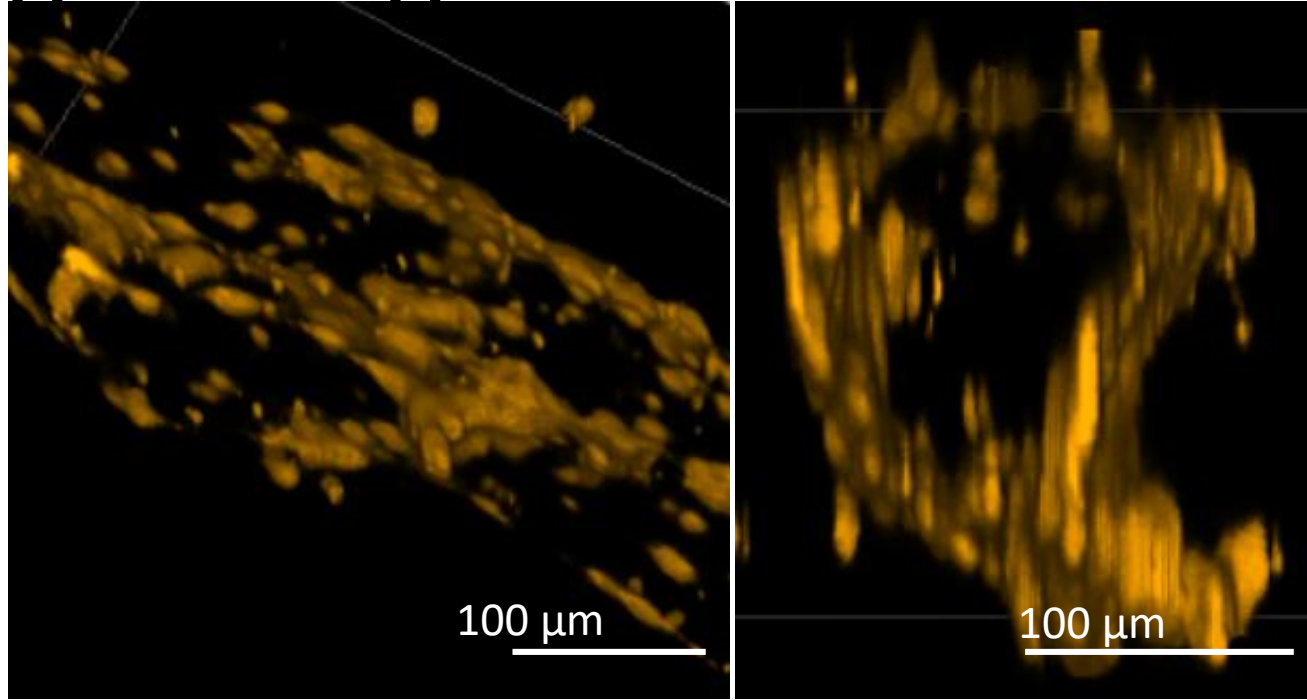
# Engineering the blood vessel layer



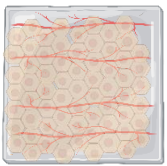




# Engineering the blood vessel layer

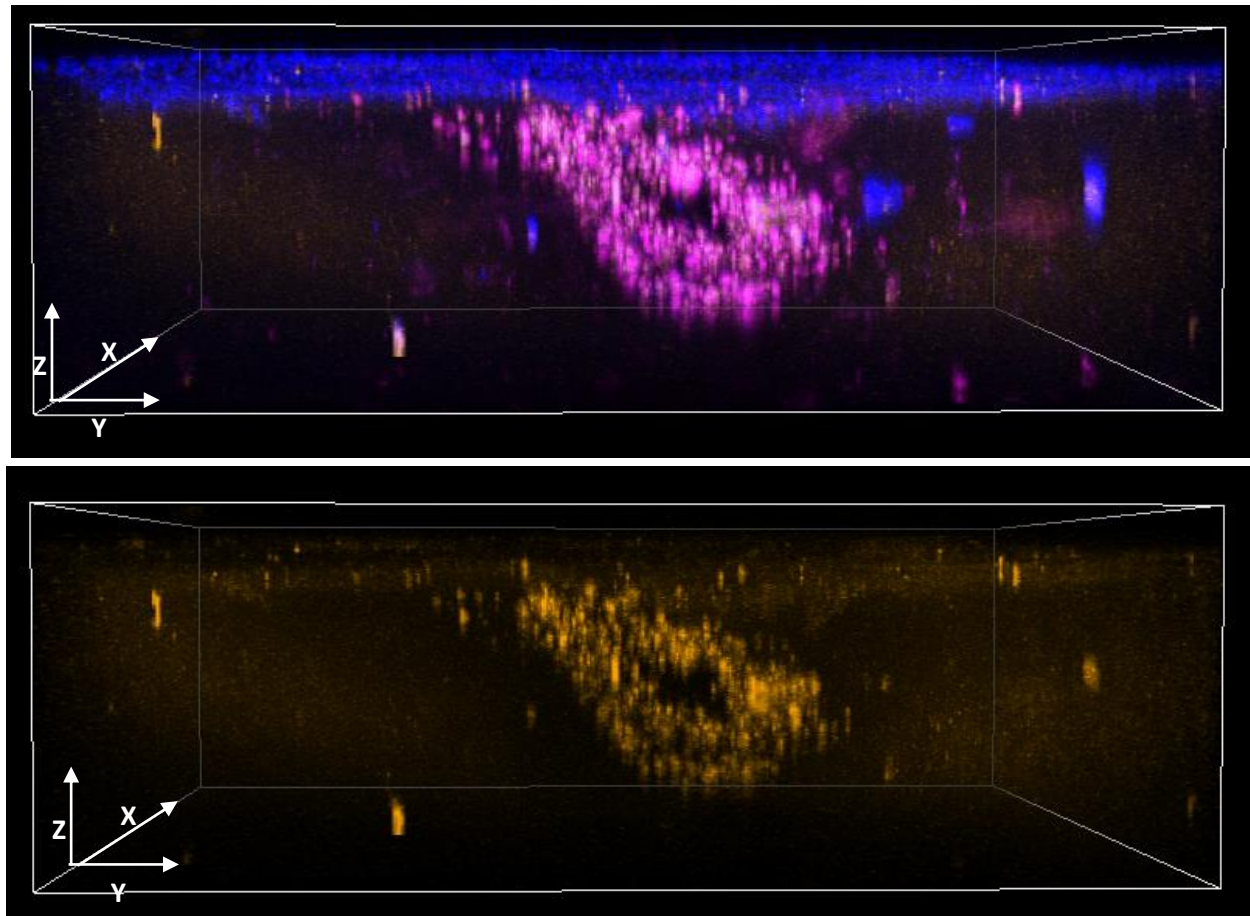




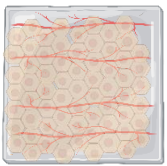


# Engineering the RPE layer

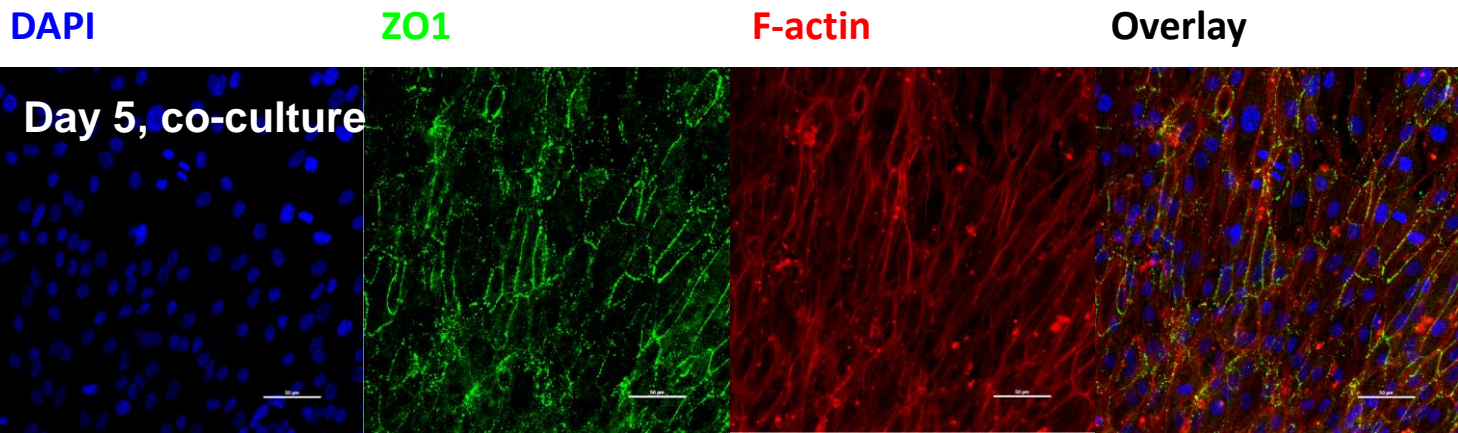
DAPI/CD31/HUVEC-CYTOPAINTER



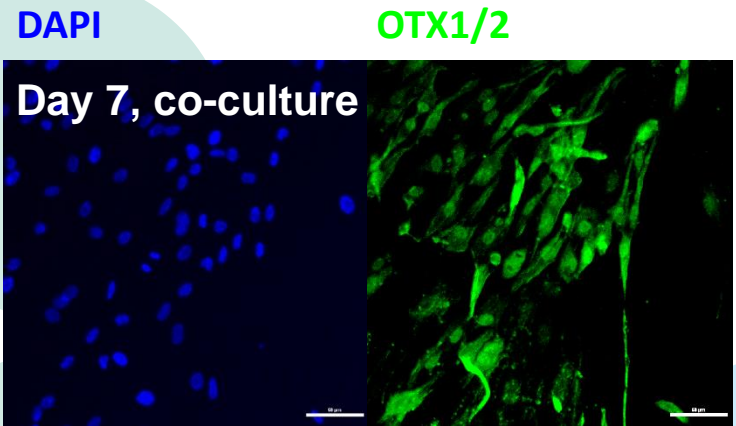




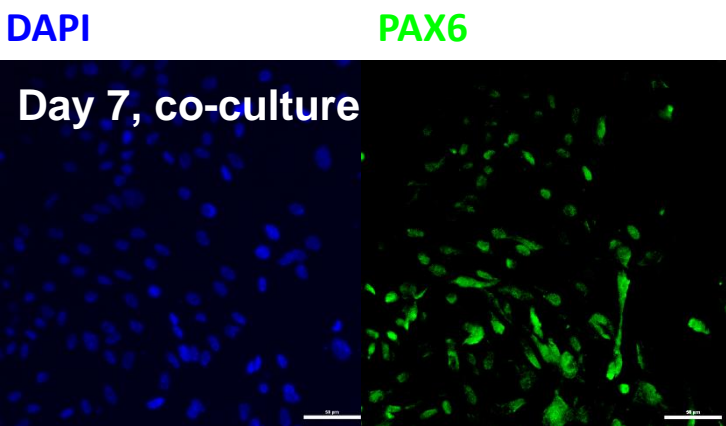
# Engineering the RPE layer



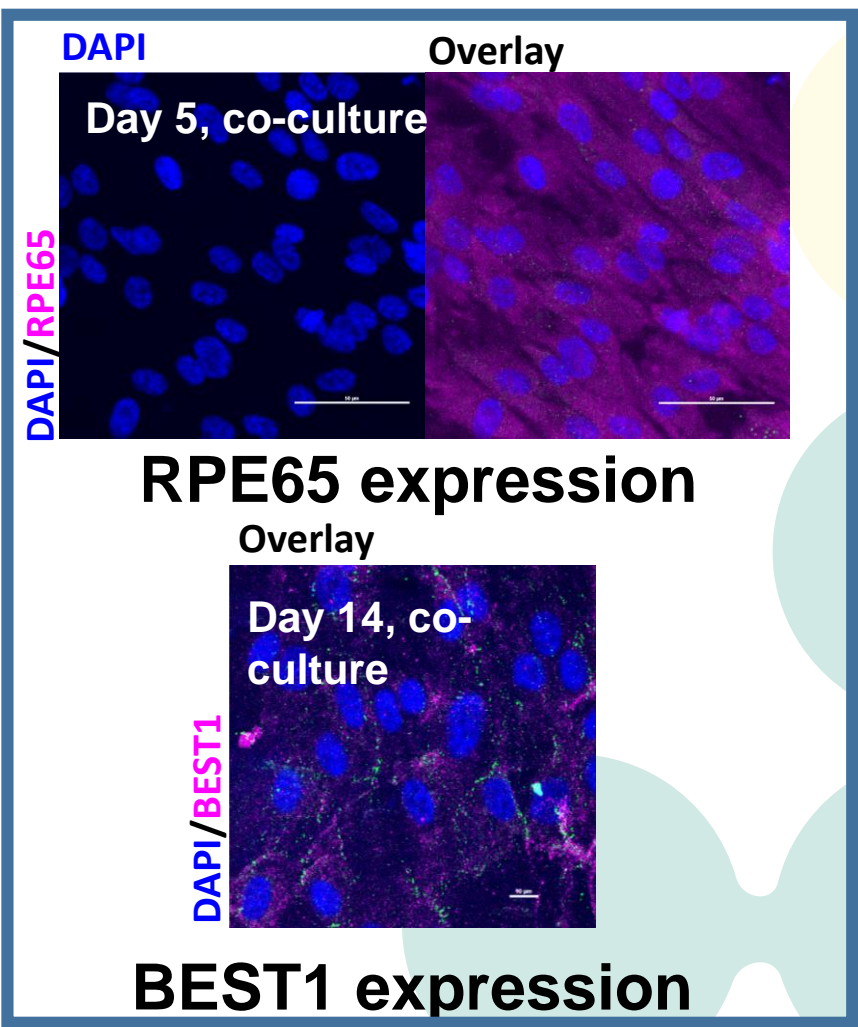
**ZO1 expression**



**OTX 1/2 expression**

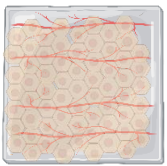


**PAX6 expression**

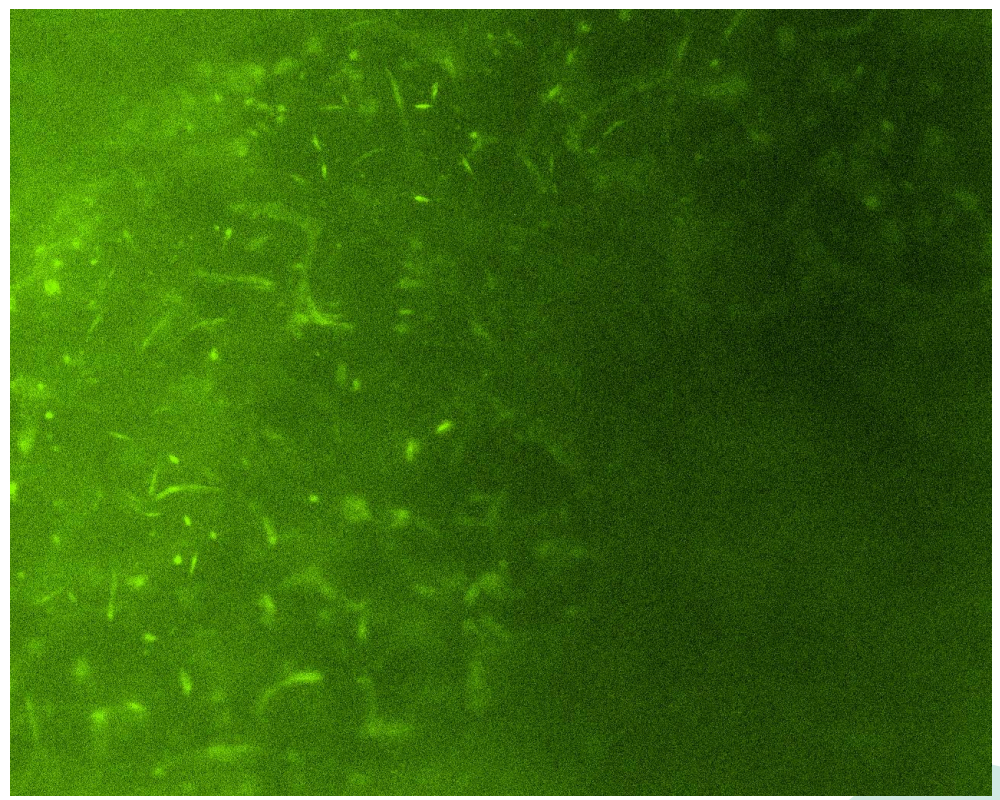
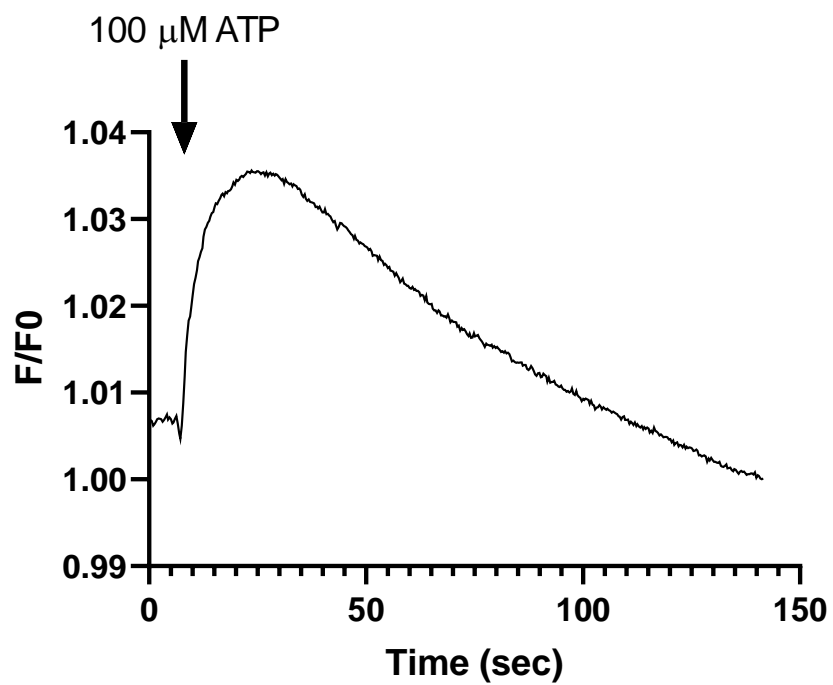


**Mature RPE markers**



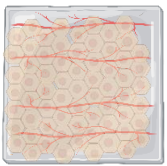


# Engineering the RPE layer

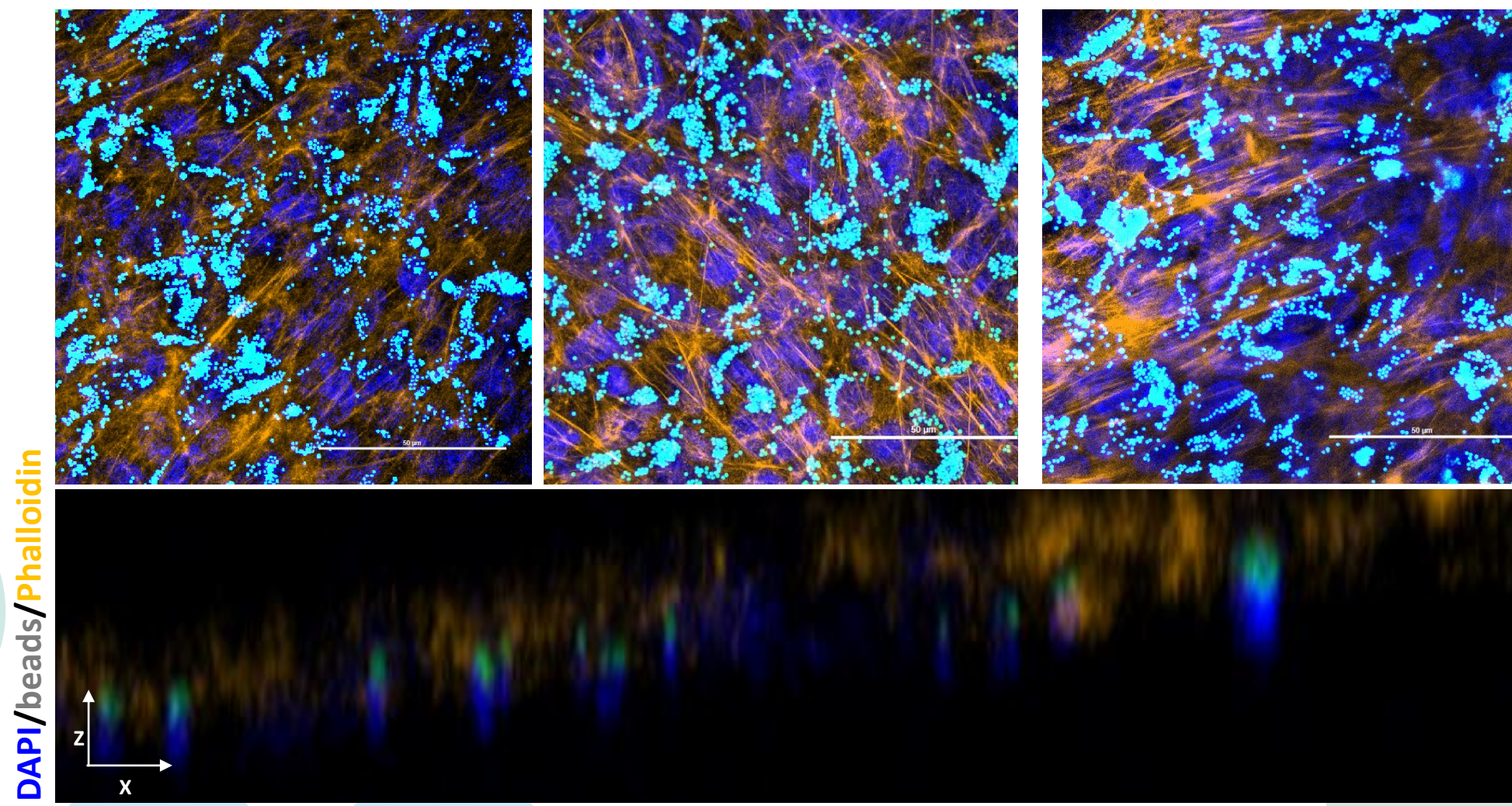


ARPE-19 transport  $\text{Ca}^{2+}$  in response to ATP after Co-culture with endothelial cells





# Engineering the RPE layer



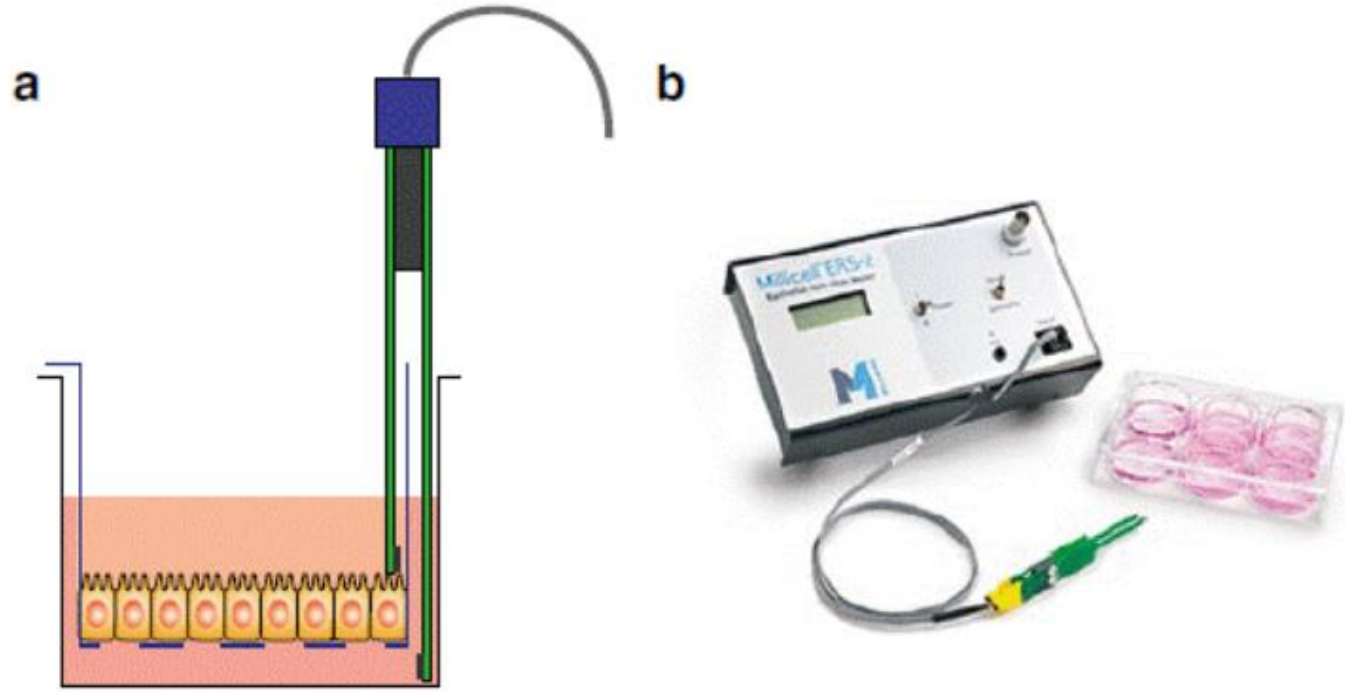
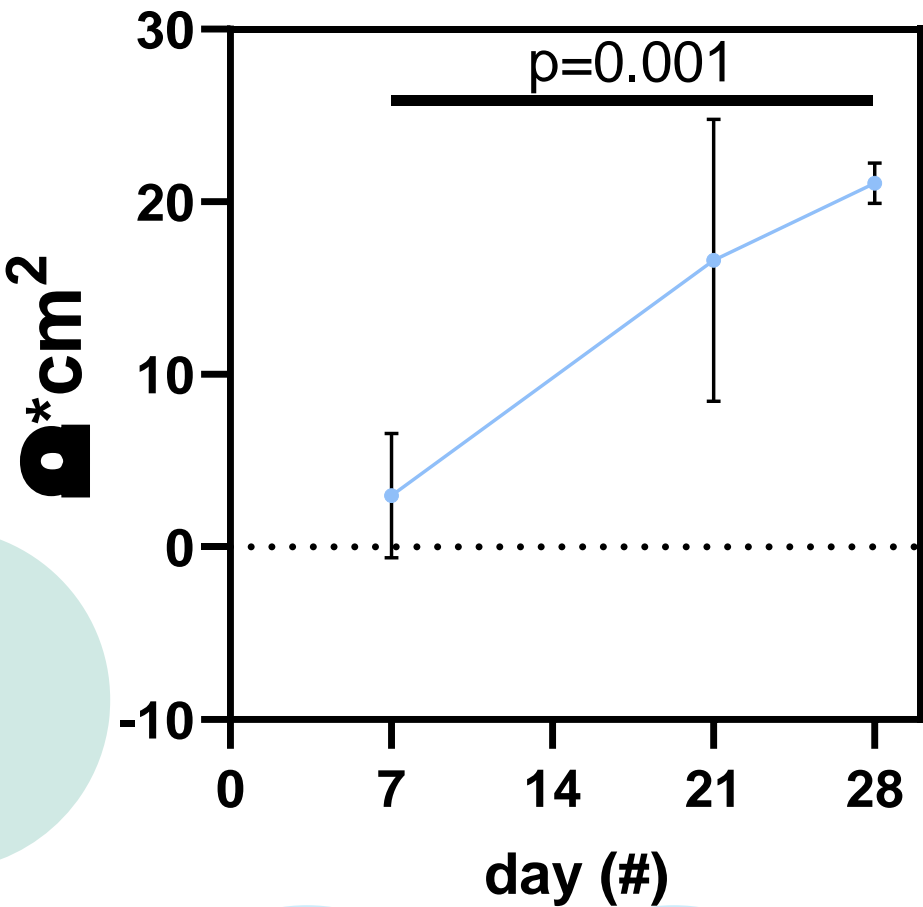
ARPE-19 preform phagocytosis of latex beads after Co-culture with endothelial cells

day 20 phagocytosis of latex beads





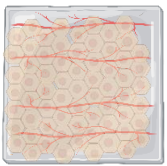
# Engineering the RPE layer



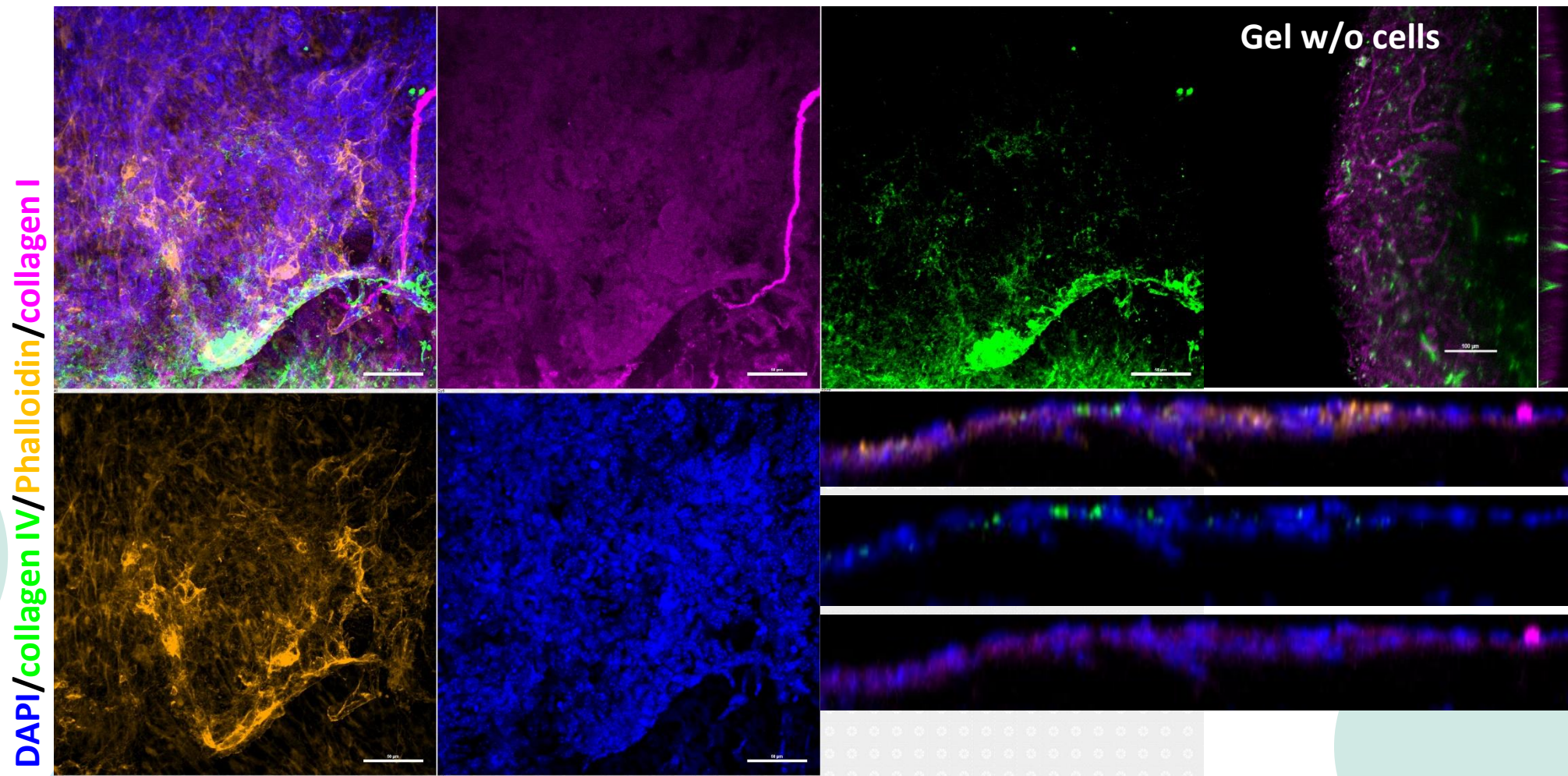
[https://ebrary.net/reet\\_ecnatsiser\\_lacirtcele\\_laillehtipesnart\\_tnemerusaem/htlaeh/24380](https://ebrary.net/reet_ecnatsiser_lacirtcele_laillehtipesnart_tnemerusaem/htlaeh/24380)

ARPE-19 form a barrier on top of the hydrogel





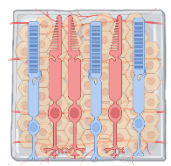
# Engineering the RPE layer



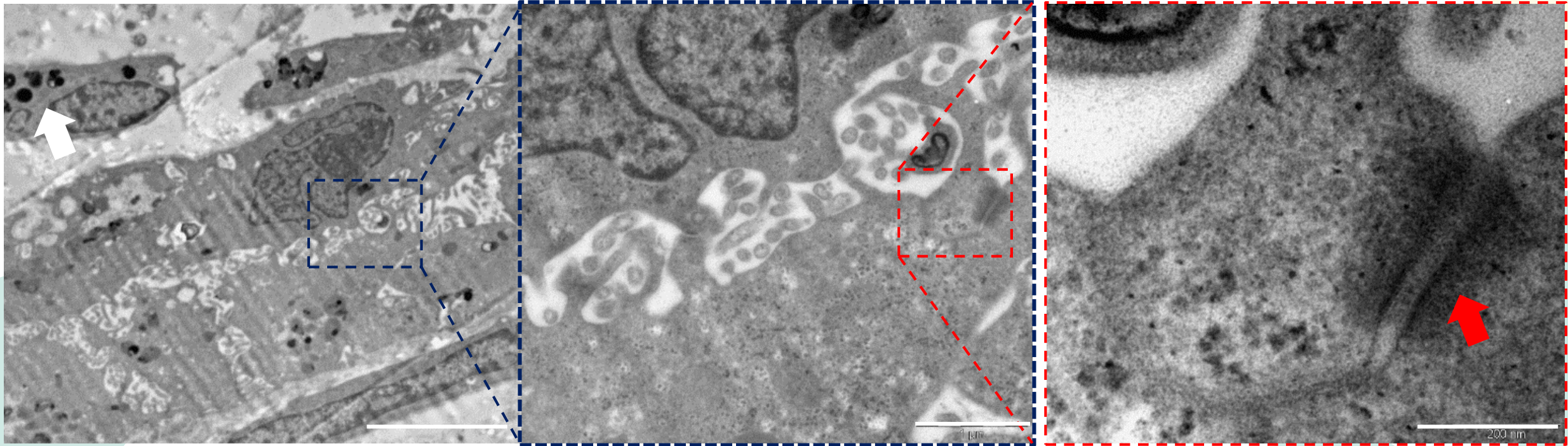
RPE cells secrete Bruch membrane proteins in the co-culture condition

Co-culture: HUVEC+ARPE19 day 94 + day 81 661w





# Engineering the RPE layer

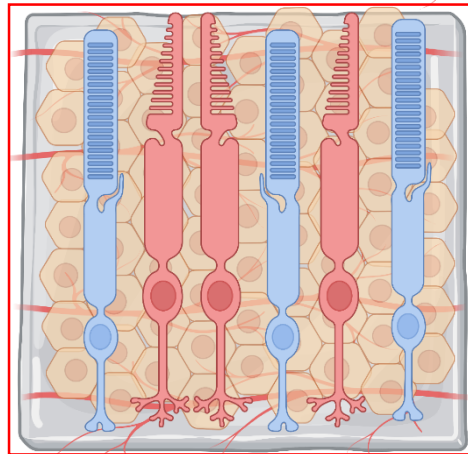


ARPE-19 cells display microvilli, pigmentation and desmosomes

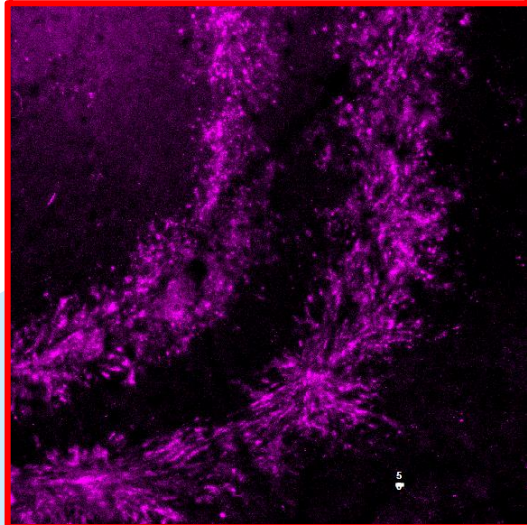
Co-culture: HUVEC+ARPE19 day 104 + day 91 661w



# Engineering the vascularized retina



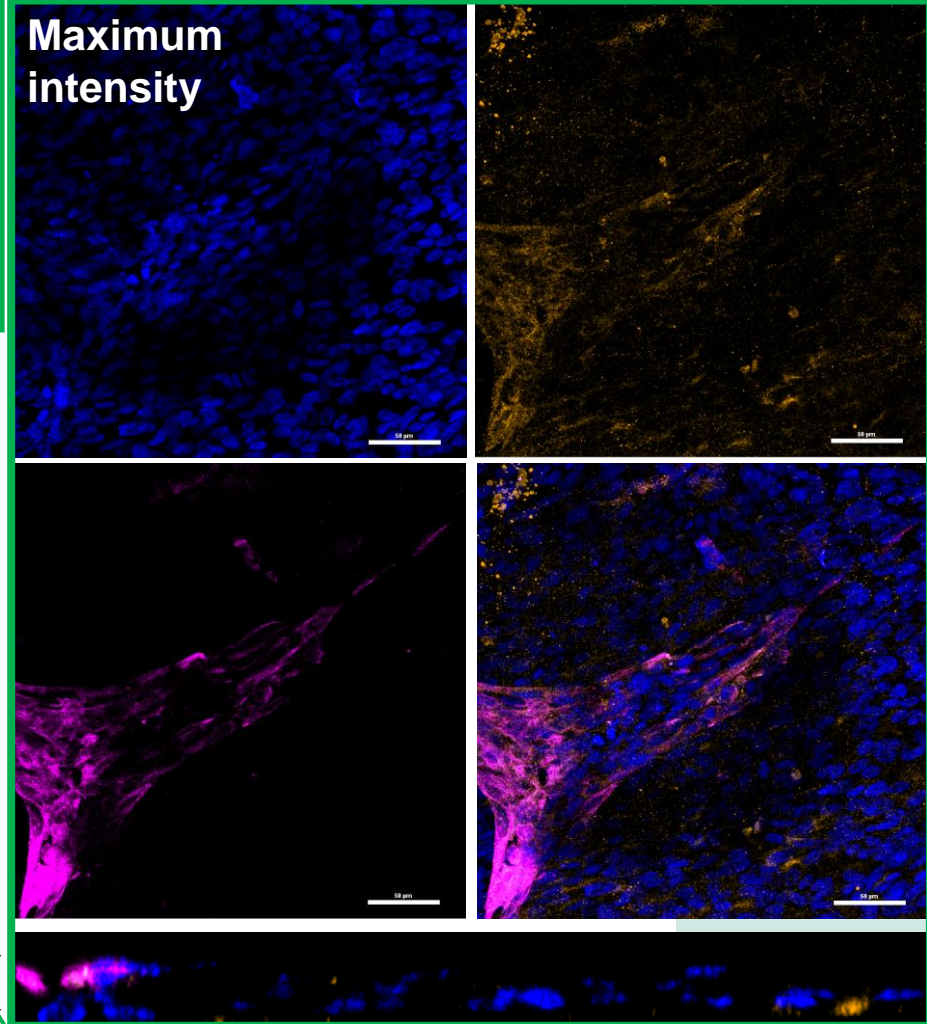
Endothelial cells



CD31

Photoreceptors →  
RPE →

Maximum  
intensity



DAPI/NESTIN/BEST







# Future work

- 👁️ *In vivo* – show incorporation into the tissue
- 👁️ Large animal model – mini pigs
- 👁️ Design of the surgical techniques



# Acknowledgments



**In collaboration with the  
Regenerative ophthalmic lab:  
Prof. Adiel Barak MD and Dr.  
Aya Barzelay MD, PhD**





# Recanati Research Center

**Giris Jacob, MD, PhD,**

Medicine F

*Medicine, Sigol, Physiology, Pharmacology & Clinical Pharmacology*

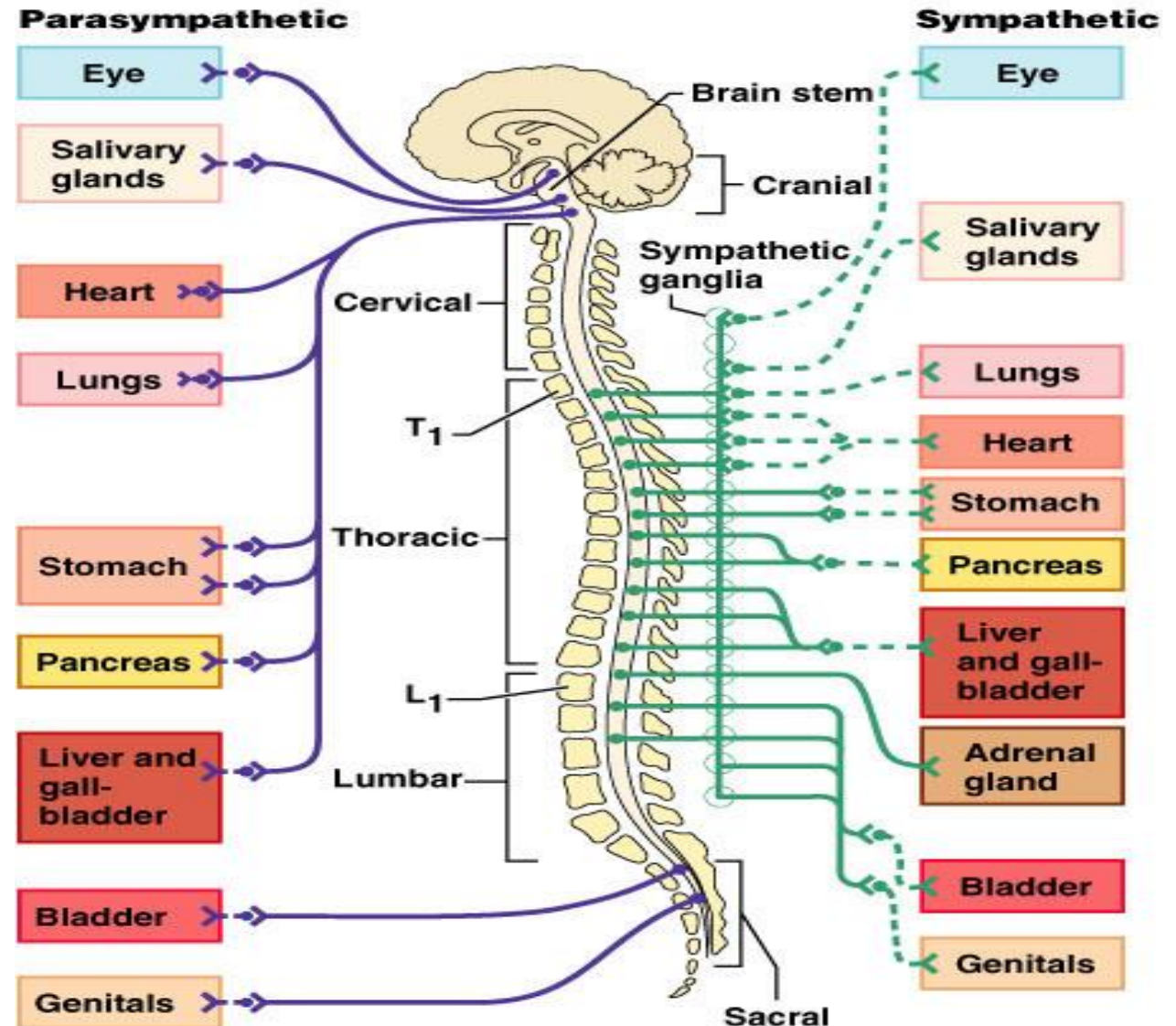


- Autonomic Nervous System
- CV Regulatory Systems
- Flow Regulations
- Hemodynamics, etc.

- Non-Invasive equipments
- Monitoring and signal processing
- Various biomarkers Blood Hemostasis
- MRI & (collaborations)

We use pharmacological tools to investigate physiologic and pathophysiologic mechanisms

## Autonomic Nervous System and Regulatory systems





# Essence of our Research

- Translational Science, cornerstone of the basic research in human: **Hypothesis-driven**.
- Understanding mechanistic processes, in physiology and pathophysiology: **Health & Disease**
- Development of new treatment and helpful medical devices: **High Tech**
- Focus, is on cardiovascular regulatory systems, with major "hint" on central and peripheral autonomic nervous system control: on Heart, Mesenteric Circulation and Cerebral regulatory mechanisms (MD, Basic Science)
- **Cannabis** effects on regulatory systems (PhD Student)

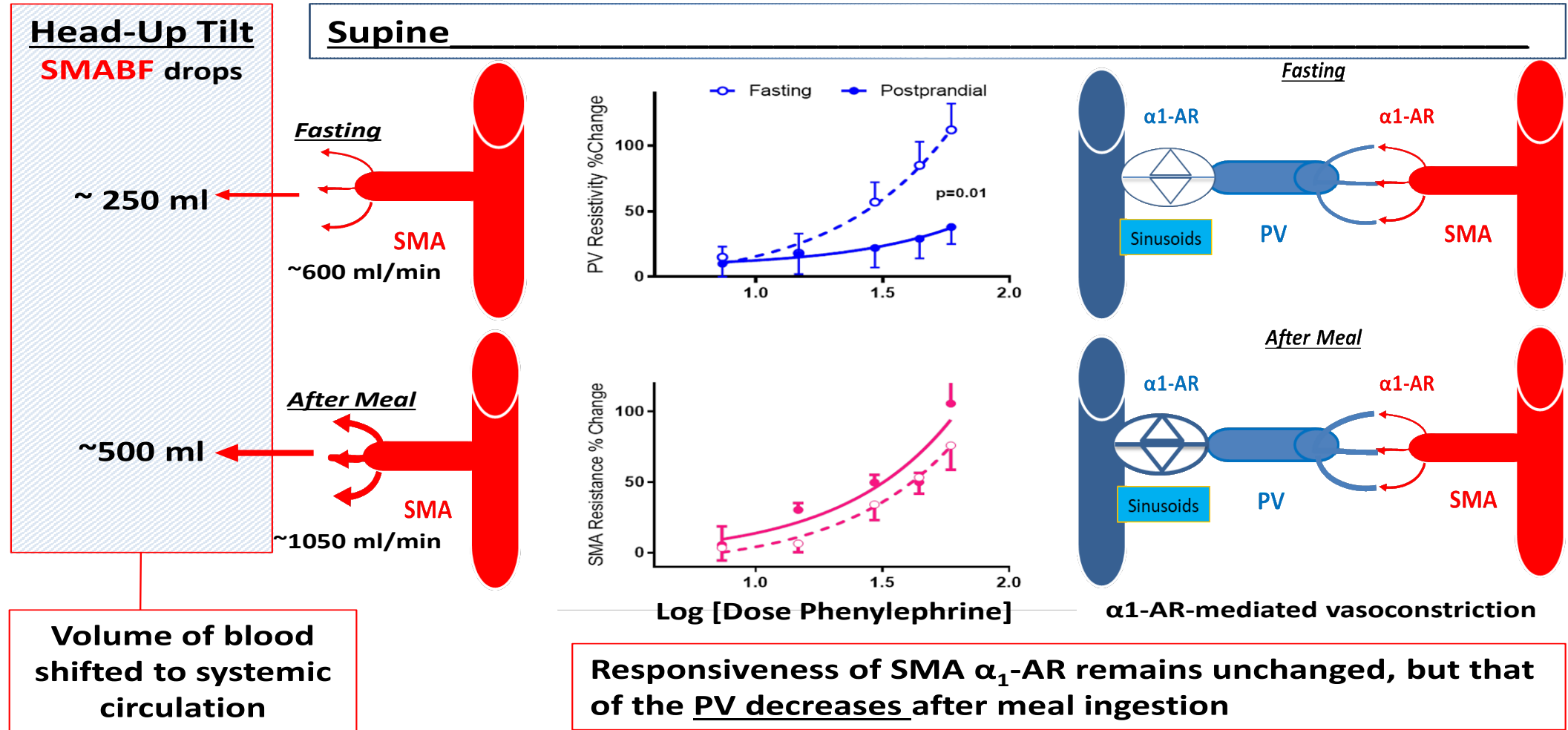


# Longitudinal View of our outcome:

- Syndormes:
  - Neuropathic Postural Tachycardia Syndrome (POTS), *NEJM*
  - Dysautonomia in Hypermobility Syndrome, *AJM*
  - Dysautonomia in Premenstrual Syndrome, *JCME*
- Mechanisms:
  - Adrenoreceptors functions in CVD, *Circulation x*
  - CBF: CO<sub>2</sub>-NO axis, *Circulation & AJP*
  - Endothelial Function, Brain, Penile, Periphery, *AJP, Urol, Circ*
  - Coagulation Physiology, effect of ANS, *Hypertension , Hemost & Throm*
  - Cannabis Effects on inhibitory pain, *Neurology*
  - Effect of meals on CV regulatory systems, *AJP*



# Effect of meal on mesenteric circulation during orthostasis, and $\alpha_1$ -AR responsiveness



AR, adrenoceptor, SMA (BF), superior mesenteric artery (blood flow), PV, portal vein, sinusoids, venous hepatic sinusoids



[Extracellular Vesicles of COVID-19 Patients Reflect Inflammation, Thrombogenicity, and Disease Severity.](#)

Aharon A, Dangot A, Kinaani F, Zavaro M, Bannon L, Bar-Lev T, Keren-Politansky A, Avivi I, **Jacob G.** *Int J Mol Sci.* 2023 Mar 21;24(6):5918. doi: 10.3390/ijms24065918.PMID: 36982991

[Diagnostic approaches to syncope in Internal Medicine Departments and their effect on mortality.](#)

Galron E, Kehat O, Weiss-Meilik A, Furlan R, **Jacob G.** *Eur J Intern Med.* 2022 Aug;102:97-103. doi: 10.1016/j.ejim.2022.05.015. Epub 2022 May 20.

[Effect of ingesting a meal and orthostasis on the regulation of splanchnic and systemic hemodynamics and the responsiveness of cardiovascular  \$\alpha\_1\$ -adrenoceptors.](#)

Zreik F, Meshulam R, Shichel I, Webb M, Shibolet O, **Jacob G.** *Am J Physiol Gastrointest Liver Physiol.* 2021 Nov 1;321(5):G513-G526. doi: 10.1152/ajpgi.00142.2021. Epub 2021 Sep 15.PMID: 34523347

[COVID-19-Associated Hyper-Fibrinolysis: Mechanism and Implementations.](#)

**Jacob G,** Aharon A, Brenner B. *Front Physiol.* 2020 Dec 16;11:596057. doi: 10.3389/fphys.2020.596057. eCollection 2020.PMID: 33391014

[Vagal and Sympathetic Function in Neuropathic Postural Tachycardia Syndrome.](#)

**Jacob G,** Diedrich L, Sato K, Brychta RJ, Raj SR, Robertson D, Biaggioni I, Diedrich A. *Hypertension.* 2019 May;73(5):1087-1096. doi: 10.1161/HYPERTENSIONAHA.118.11803.PMID: 30879357

[Cannabis analgesia in chronic neuropathic pain is associated with altered brain connectivity.](#)

Weizman L, Dayan L, Brill S, Nahman-Averbuch H, Hendler T, Sharon H, **Jacob G.** *Neurology.* 2018 Oct 2;91(14):e1285-e1294.