

THE SHAPING A
TECHMED HEALTHY
EVENT FUTURE

Organ-on-a-chip: Better Designs, Better Health | Anne van der Does

Technical Innovation - Clinical biomedical research as alternative for animal testing

“The Dark Side of the Lung”

dr. Anne van der Does – PulmoScience Lab, Leiden University Medical Center



DISCLOSURE SLIDE

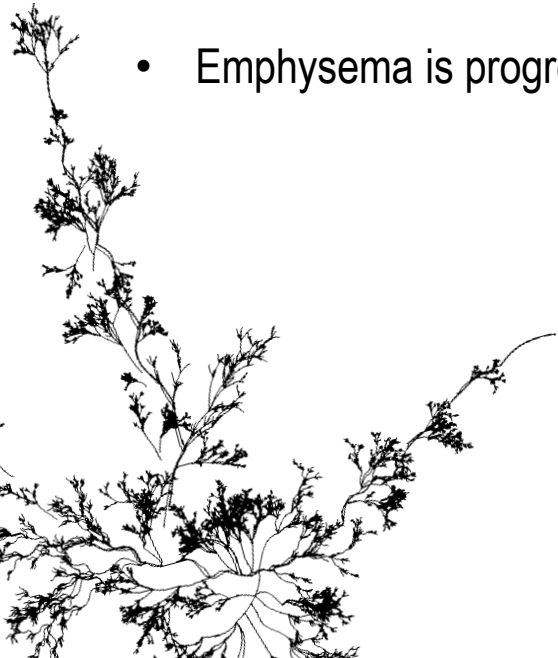
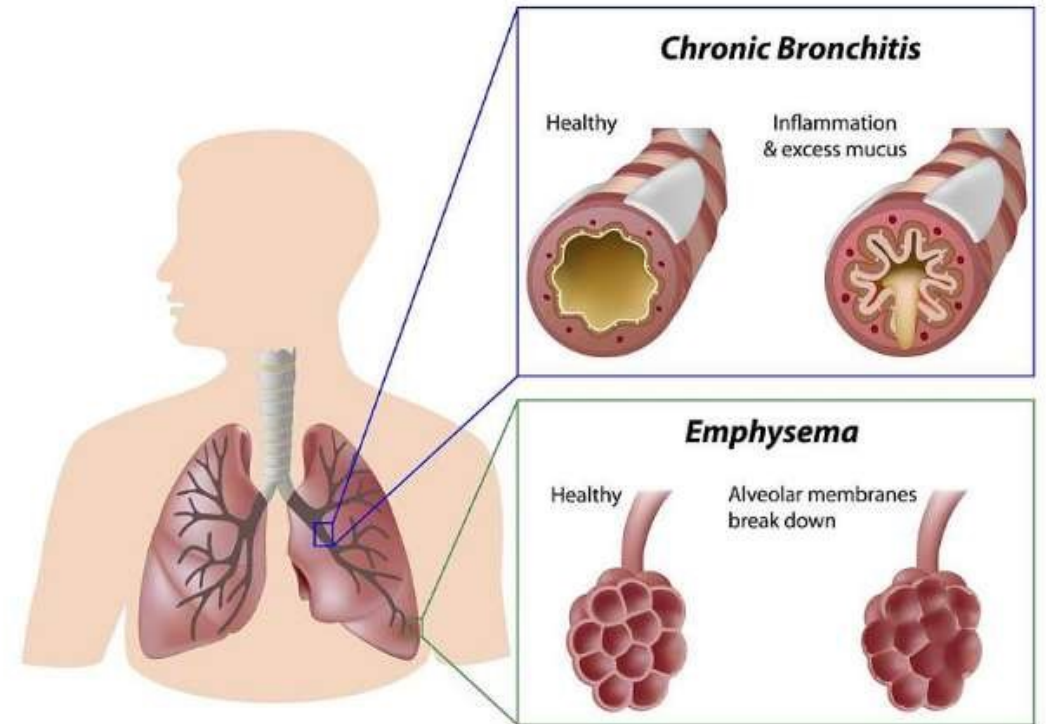
No disclosures



Chronic Obstructive Pulmonary Disease

- Third cause of death worldwide
- >500.000 patients in The Netherlands
- Characterized by two conditions: emphysema and chronic bronchitis
- Emphysema is progressive: loss of gas exchange capacity

Chronic Obstructive Pulmonary Disease (COPD)



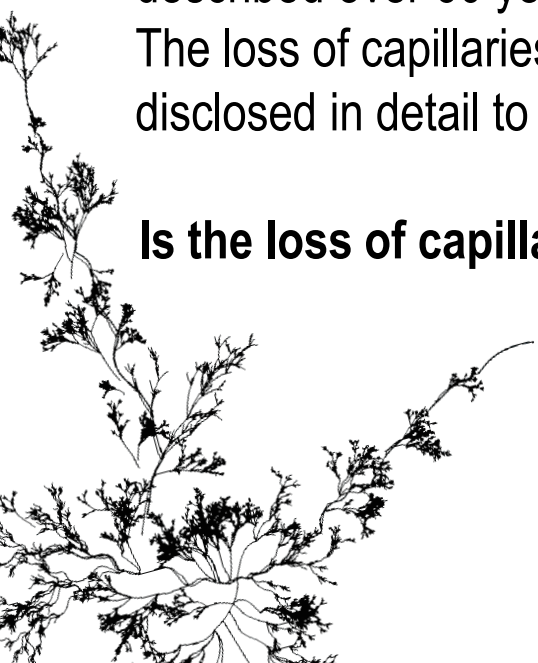
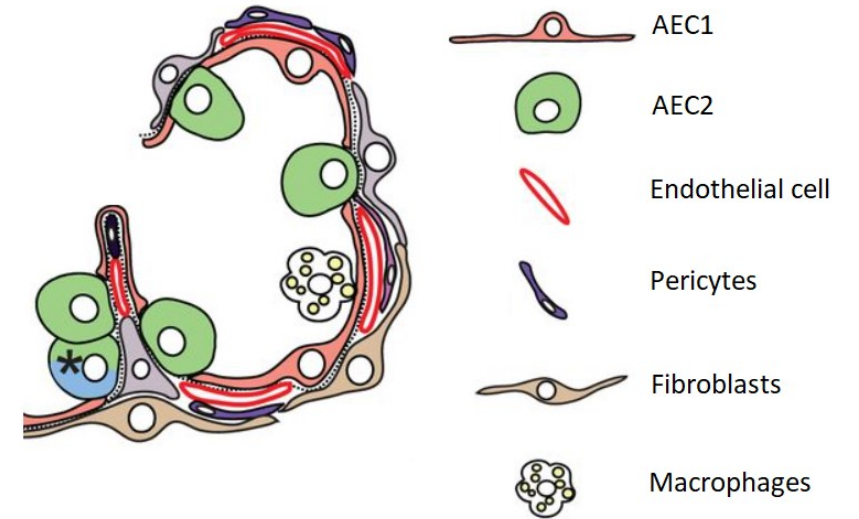
Defect in (micro)vascular compartment associated with emphysema

Cellular make up of the alveoli includes a close interaction of the type-1 alveolar epithelial cells and the (micro)vascular endothelial cells

Dysfunction of the alveolar capillary network in emphysema was first described over 60 years ago.

The loss of capillaries were not studied at that time and remain to be disclosed in detail to date.

Is the loss of capillaries a consequence of emphysema or is it driving the disease?



The Dark Side of the Lung consortium



Prof. Reinoud Gosens
RuG



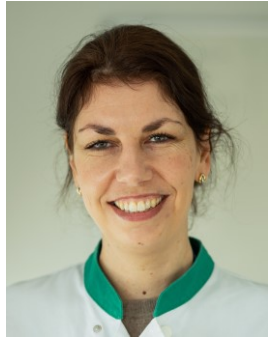
Prof. Andries v/d Meer
TU Twente



Dr. Jill Johnson
Aston University



Prof. Pieter Hiemstra
LUMC



Dr. Anne van der Does
LUMC



Prof. Marco Harmsen
UMCG



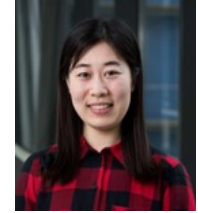
Dr. Abilash Ravi
LUMC



Dr. Xinhui Wu
RuG



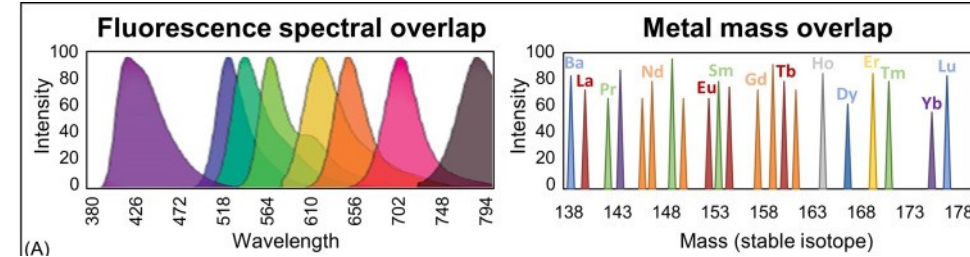
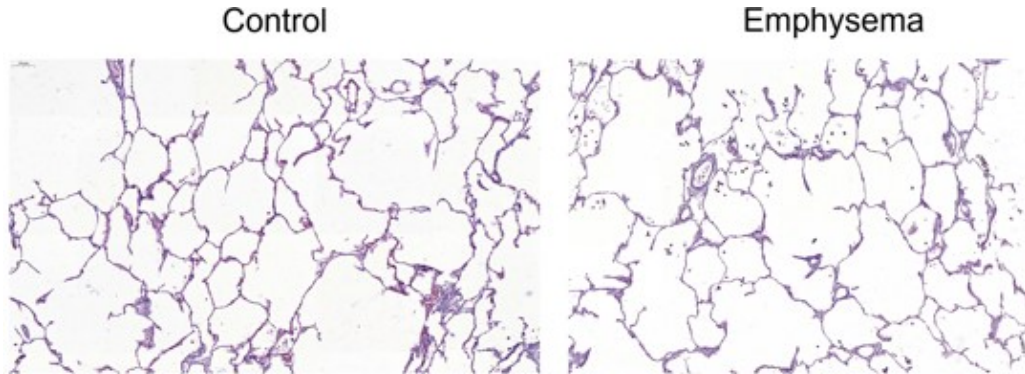
Severe emphysema: from Patient to Chip



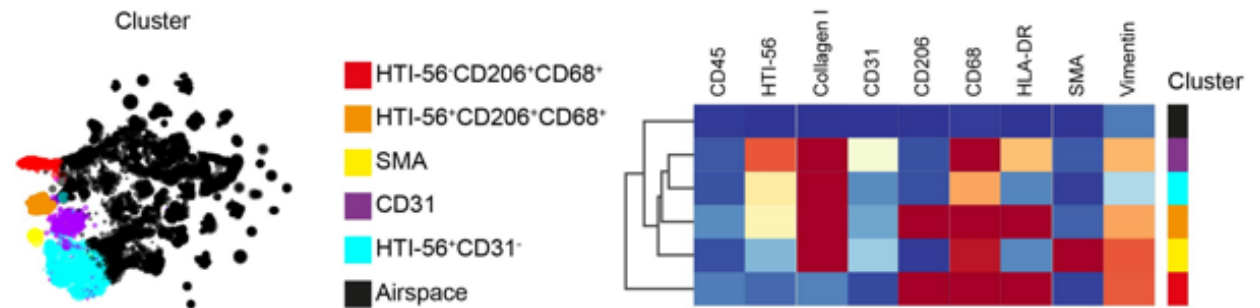
Jia Li



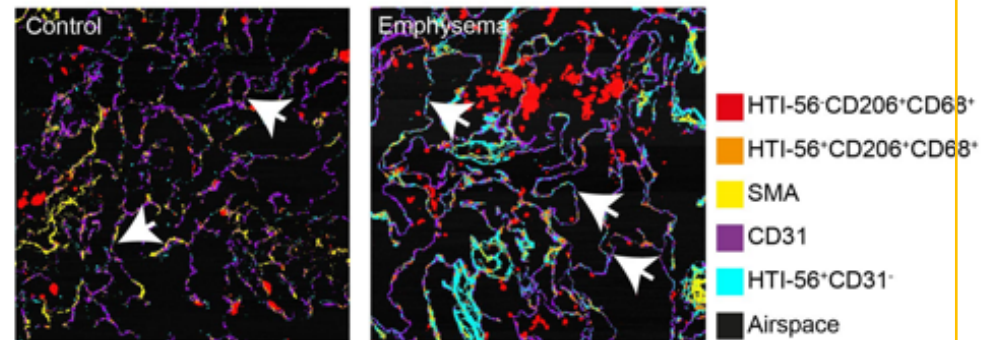
Padmini Khedoe



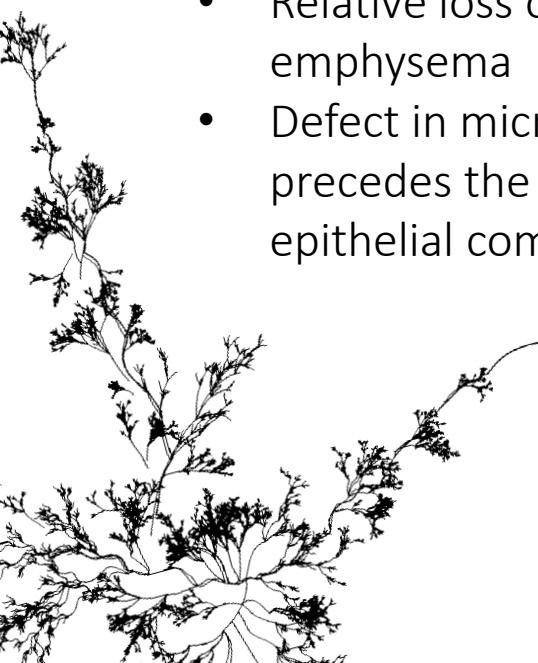
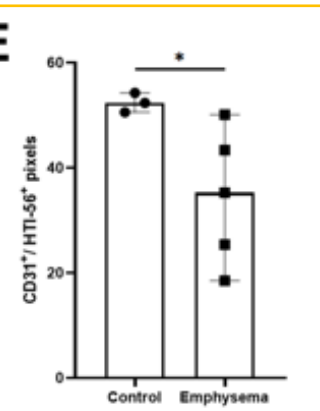
- Relative loss of microvasculature in emphysema
- Defect in microvascular compartment precedes the defect in the alveolar epithelial compartment



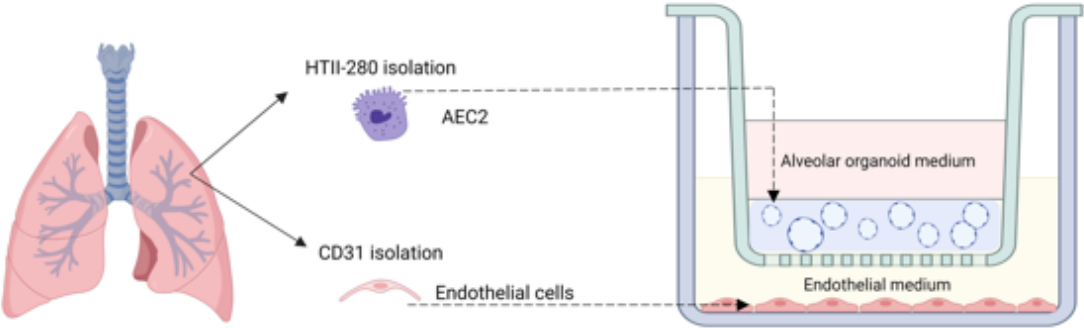
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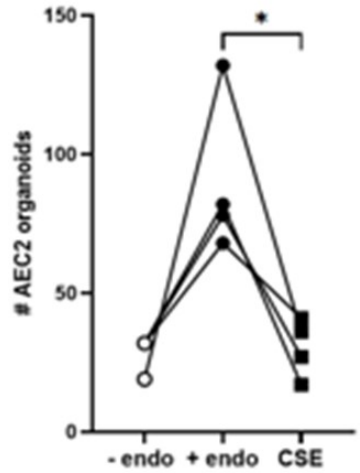
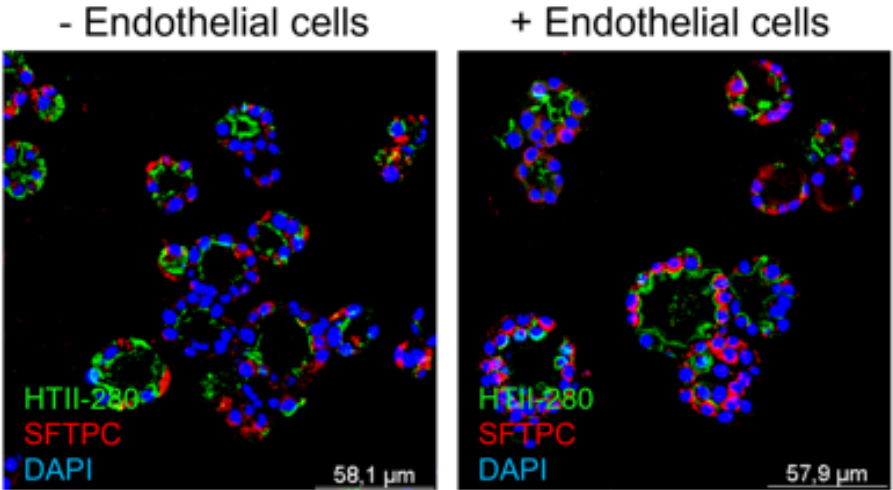
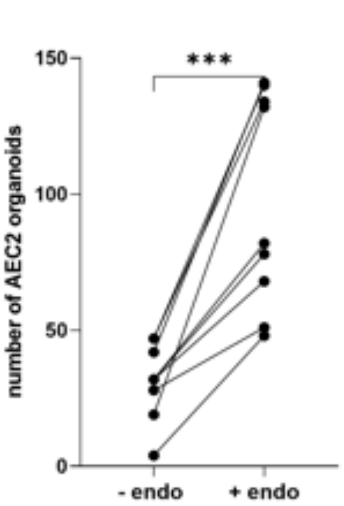
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Severe emphysema: from patient to chip

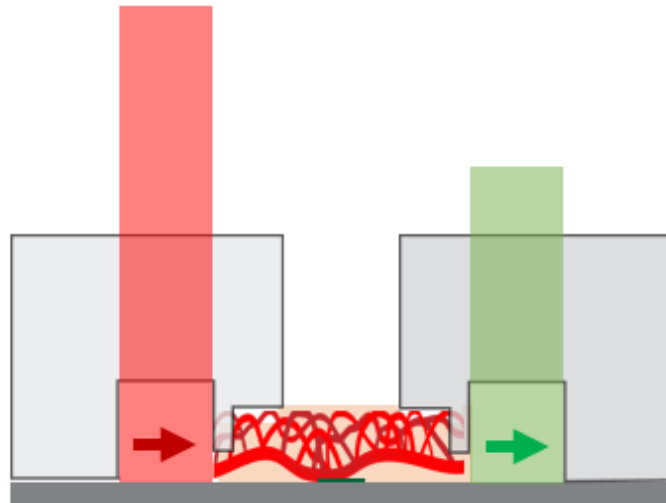
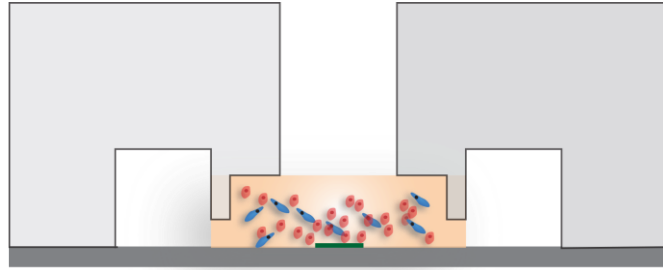
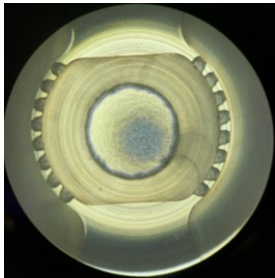
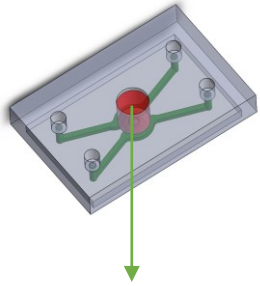


Evidence for a supporting role of endothelial cells for epithelial cells







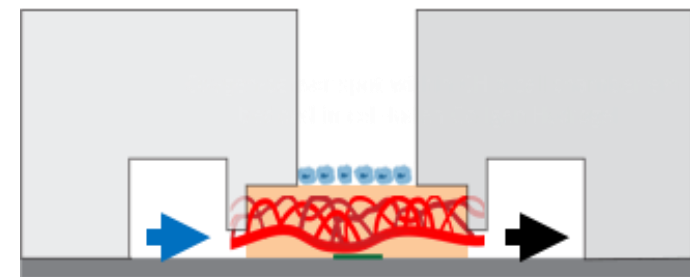
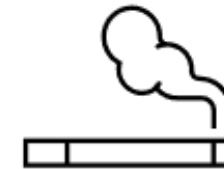
Development of the chip

Open-top; Membrane free

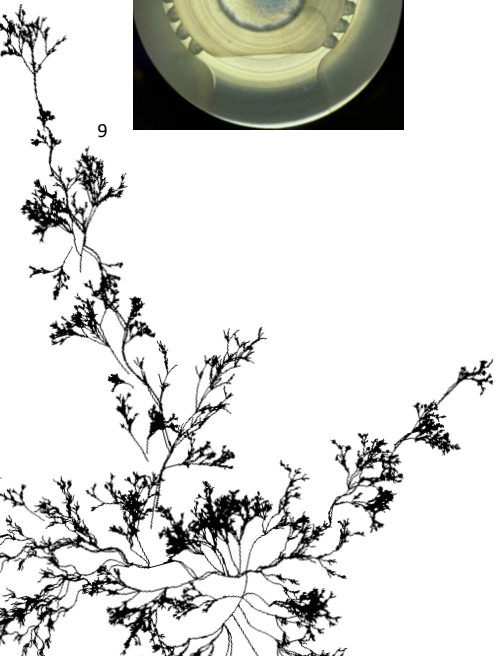


Prof. Andries v/d Meer
Tarek Gensheimer
TU Twente

-  Alveolar type II epithelial cells (AEC2)
-  Lung microvascular endothelial cells
-  Pericytes
-  Hydrogel

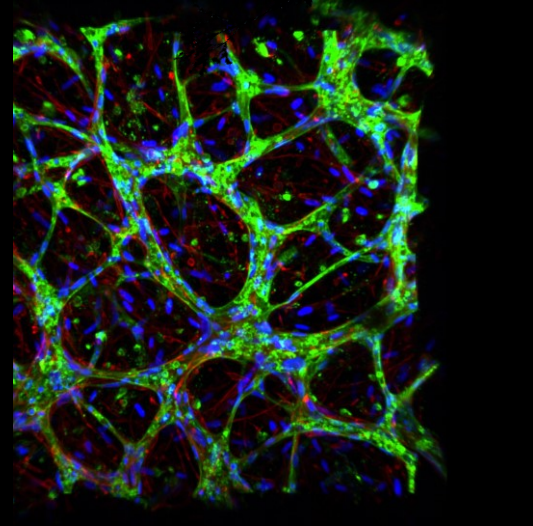
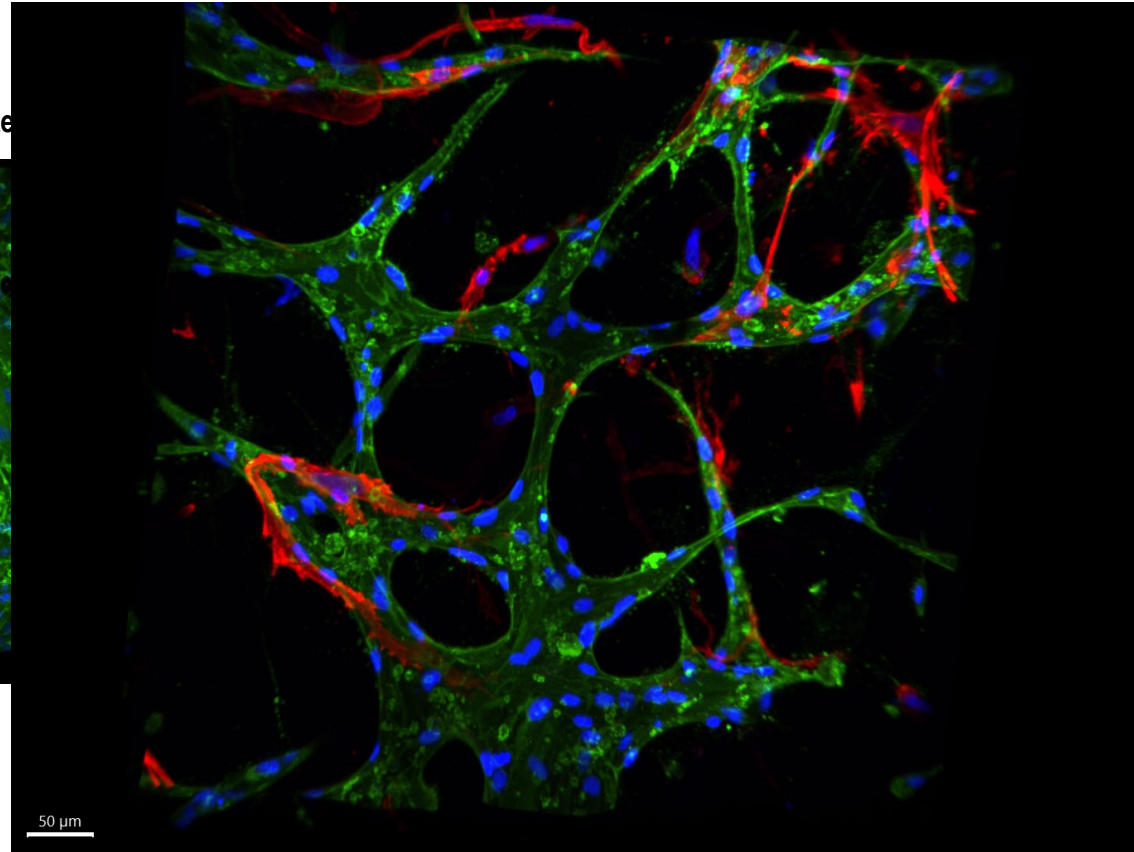
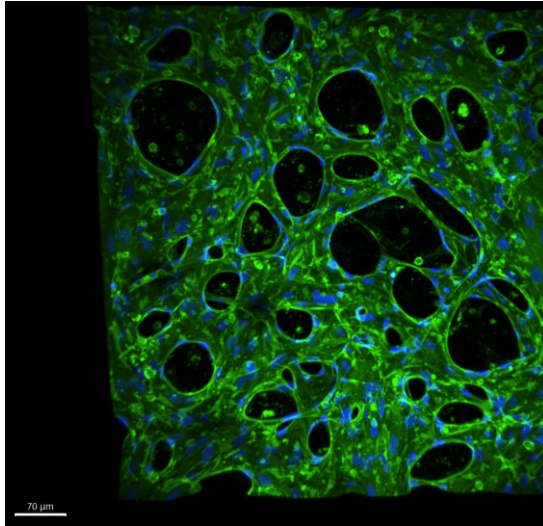


Dr. Abilash Ravi
LUMC



Control of the vessels size by combination with pericytes

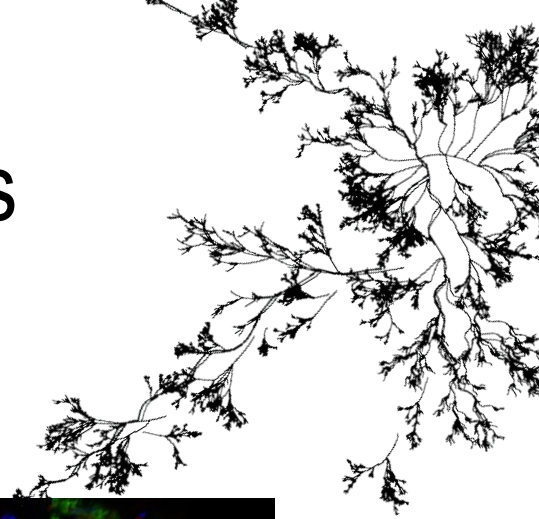
Endothelial cells without pericytes



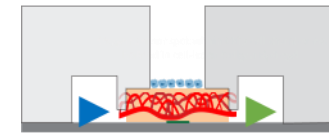
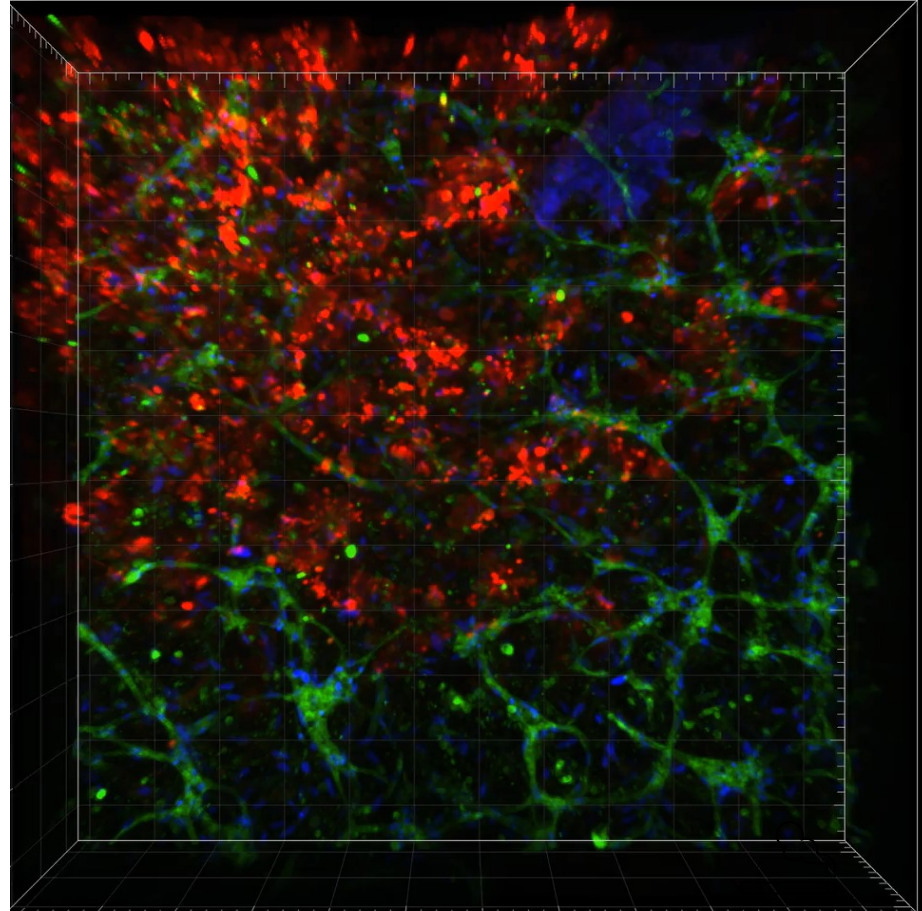
Endothelial cells – CD31

Pericytes – NG2

Day 12



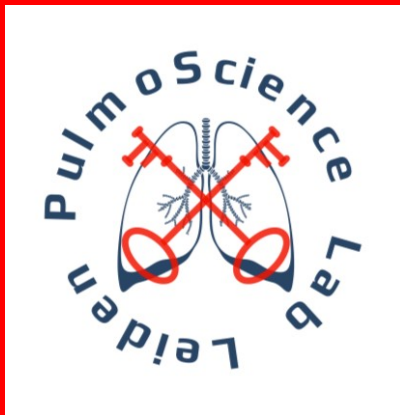
Vascularized endothelial cells co-cultured with primary AEC2



- Delineate the supportive function of endothelial cells
- Immune cell recruitment



E-mail: a.van_der_does@lumc.nl
X: LUMC PulmoScience Laboratory - @LungLabLeiden
www.pulmosciencelableiden.com





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