

**Computer Simulation of Flow and
Mass Transfer in a Bioprinted
In-Vitro Device of Human Cancer
Cells for Optimizing Oxygen
Supply -
a Progress Report**

Peter Farber, Simone Marasso

The Team

- National Research Council (CNR) Institute of Materials for Electronics and Magnetism (IMEM), IMEM Laboratory Torino Politecnico di Torino (PoliTO)

ChiLab - Materials and Microsystems Laboratory

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- IMH - Institute of Modelling and High-Performance Computing

Hochschule Niederrhein

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Hochschule Niederrhein
University of Applied Sciences

 **IMH**
Institut für Modellbildung
und Hochleistungsrechnen
Institute of Modelling
and High-Performance Computing

Agenda

- **A Few Words about the Labs at PoliTO**
- **A Few Words about IMH**
- **The Real Process**
- **First Results of Numerical Simulations**
- **Future Steps**

A Few Words about the Labs at PoliTO - 1

- **Project members:**

- **Nanoscience Lab**
- **PolitoBIOMed Lab**



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- **Chi-Lab**
- **Piquet Lab**



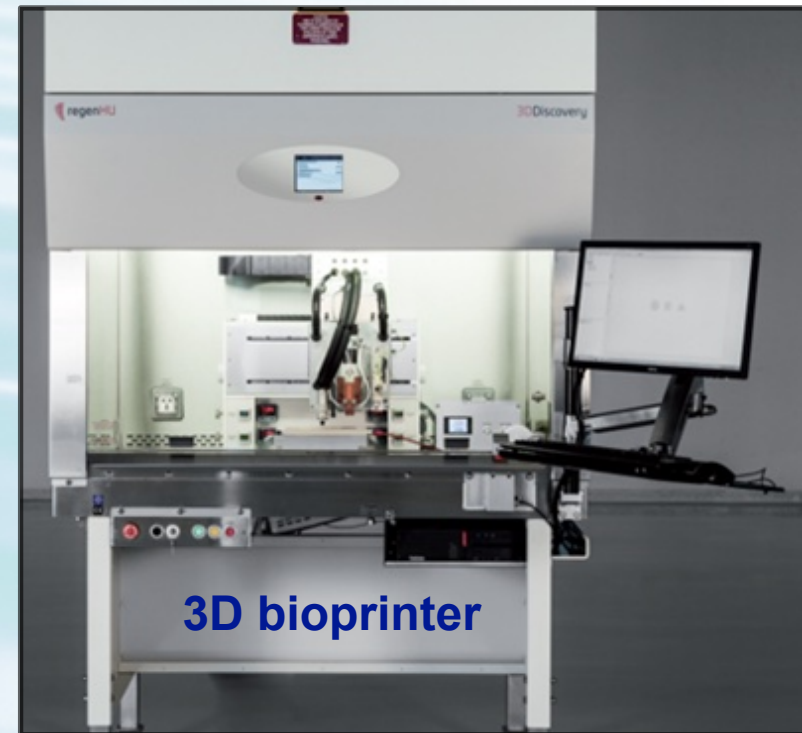
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A Few Words about the Labs at PoliTO - 2

- **Polito^{Bio}Med - biological field**
 - **Lab Cell culture & cell biology**
 - **Imaging Bioprinting**
 - **Biochemistry & molecular biology**
 - **Biomatrix preparation**



POLITECNICO
DI TORINO



Polito^{BIO}Med Lab
Biomedical Engineering Lab

A Few Words about the Labs at PoliTO - 3

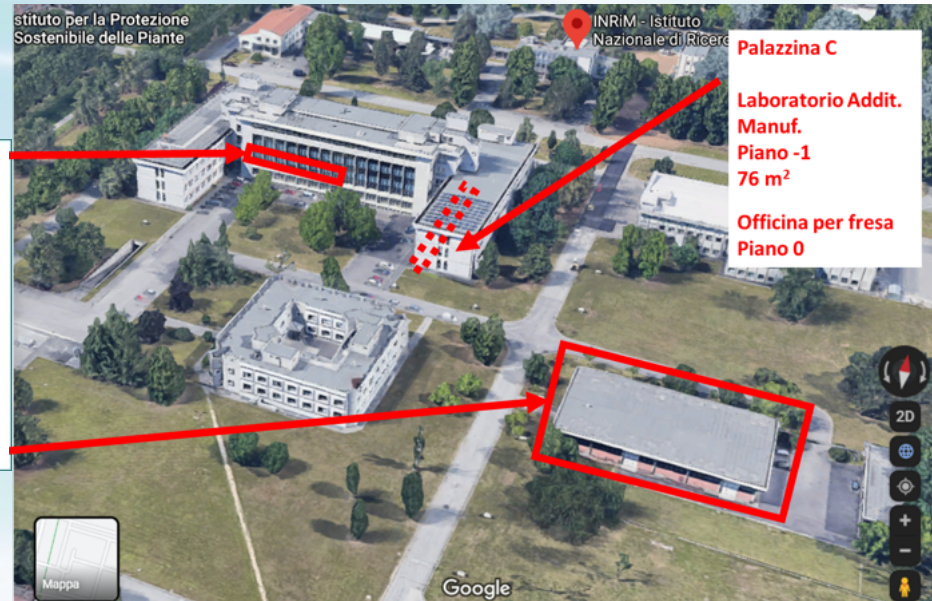
- **ChiLab - Micro&Nano Technological field**

- Lab class 100 - 15 m²
- Lab class 1000 - 45 m²
- Lab class 10 000 - 90 m²
- Biomatrix preparation



CHILAB

- **PiQuET - Micro&Nano Technological field**



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A Few Words about IMH - 1

- **Members:**

- **4 professors**

- **Computer Science, Parallel Systems**

- **Robust Design Optimization, Mathematics, Optimization**

- **Computer Simulation of Fluid Flow (Computational Fluid Dynamics, CFD)**

- **7 scientific assistants**

- **1 assistant for finance and organization**

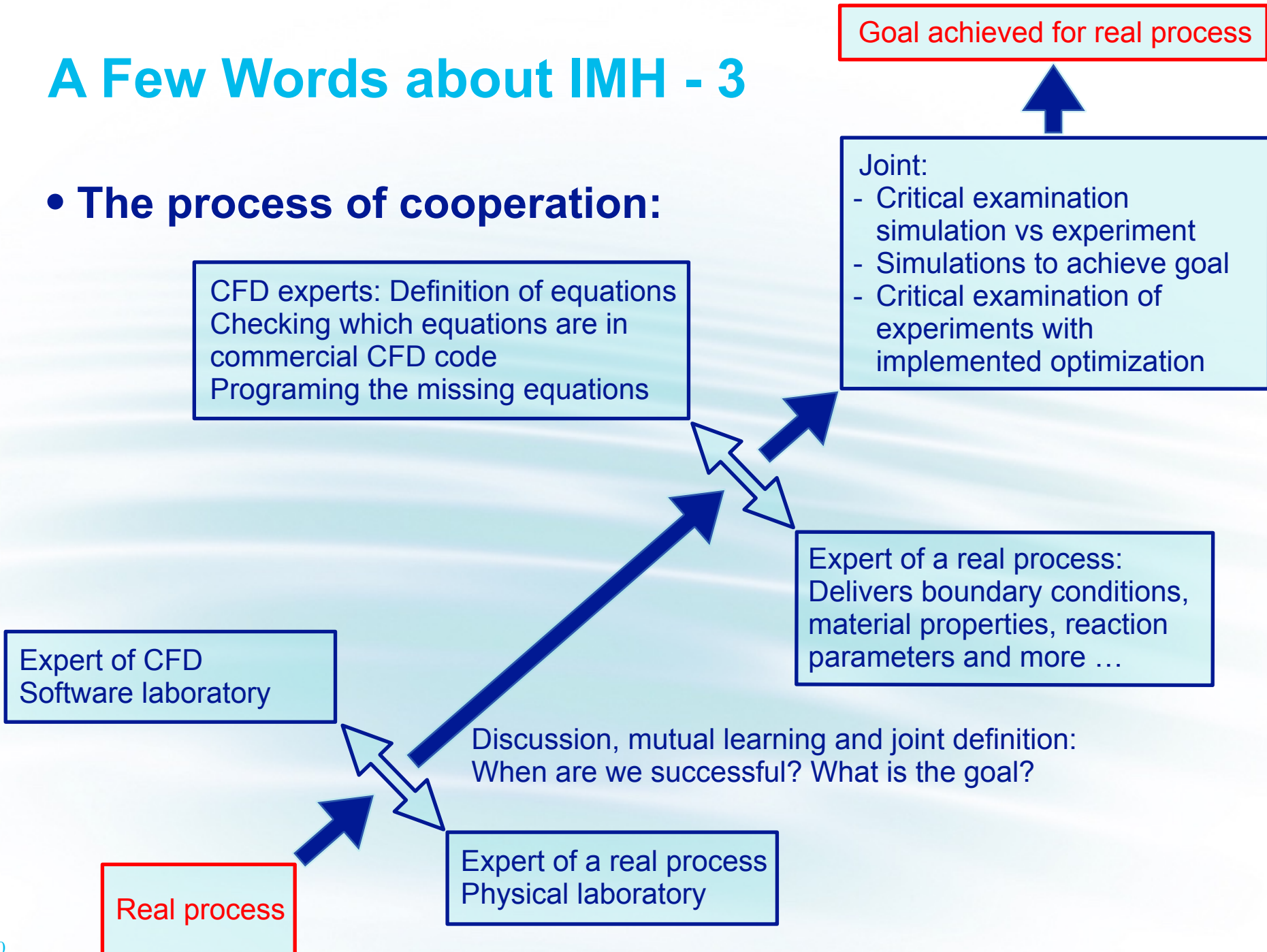


A Few Words about IMH - 2

- **CPU-cluster „Europa“ - 512 cores / 2,0 TB RAM**
- **Cluster „fornix“:**
 - **CPU-cluster - 1.216 cores / 9,7 TB RAM**
 - **GPU-cluster - 38 Nvidia A100 / 1,5 TB RAM**
- **9 workstation (together 122 cores and 544 GB RAM)**
- **75 permanent licenses Ansys Multiphysics**
- **512 permanent parallel licenses Ansys Multiphysics**
- **DSMC OpenFOAM, software for coupling to Ansys**
- **And more ...**

A Few Words about IMH - 3

• The process of cooperation:



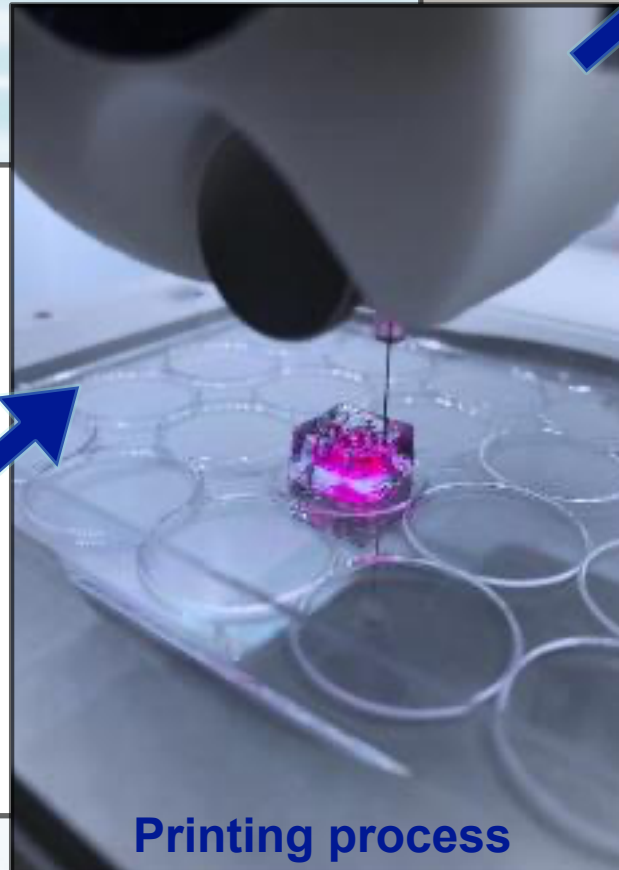
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The Real Process - 1

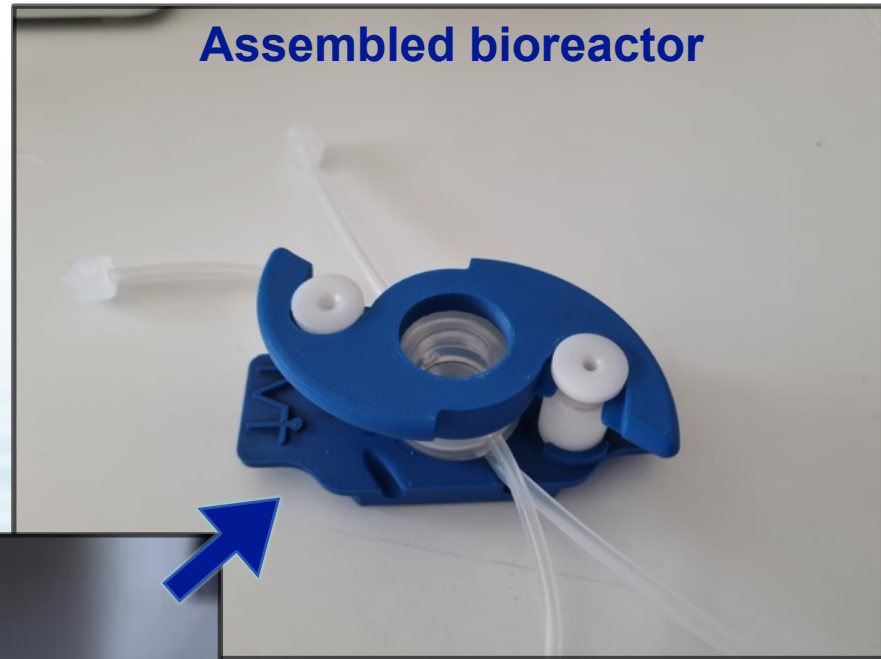
- Organ-on-a-Chip study for tumoral cells bioprinting

3D bioprinter



Printing process

Assembled bioreactor

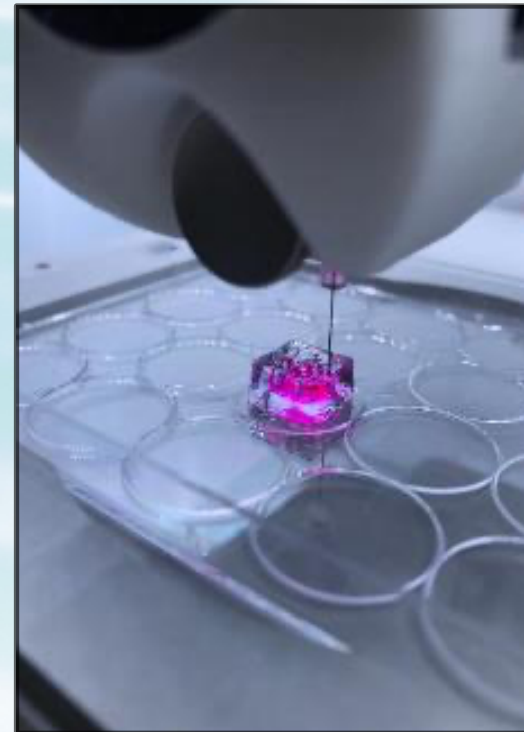
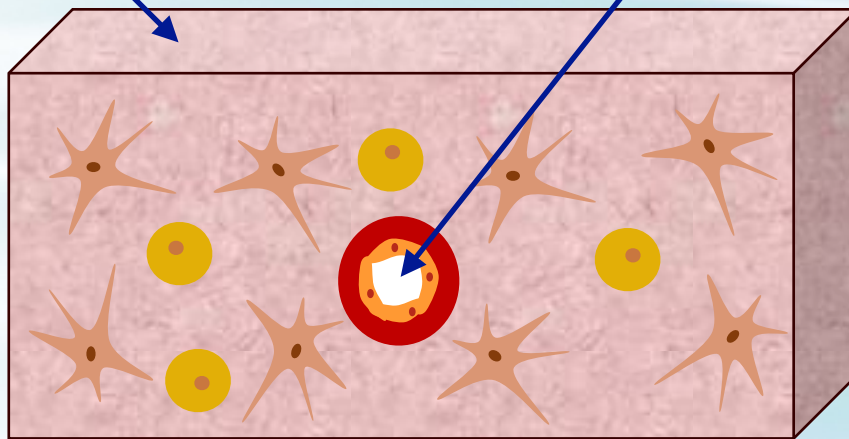


The Real Process - 2

- Sketch of the bioprinted in-vitro device of human lung-cancer

GelMA (Methacrylate Gelatin) filled with MEM

MEM with dissolved oxygen



PMRC5

A549

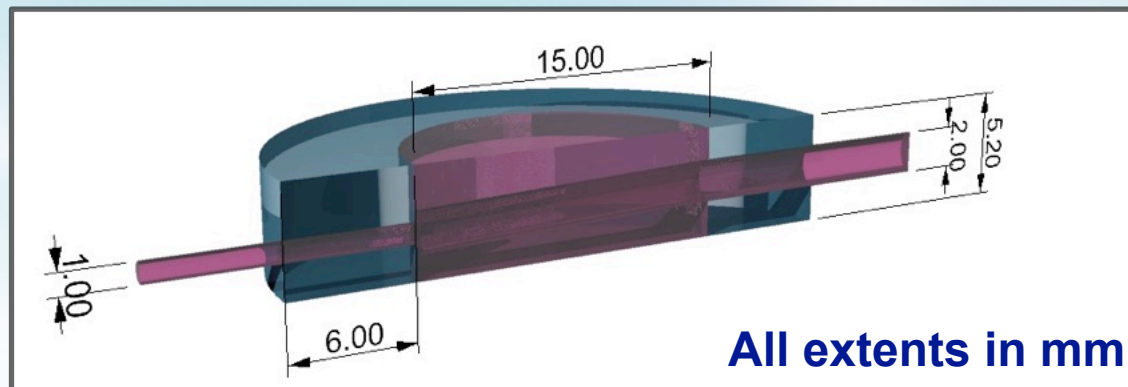
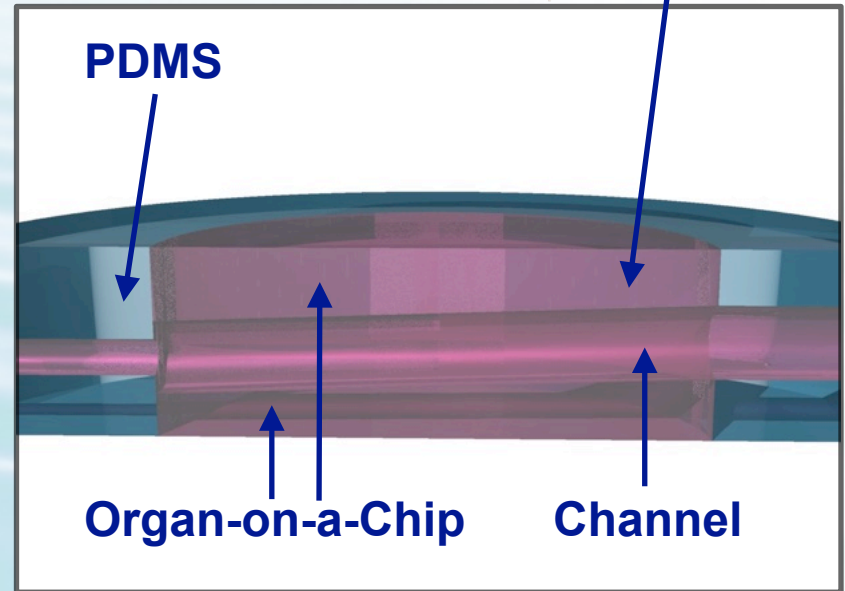
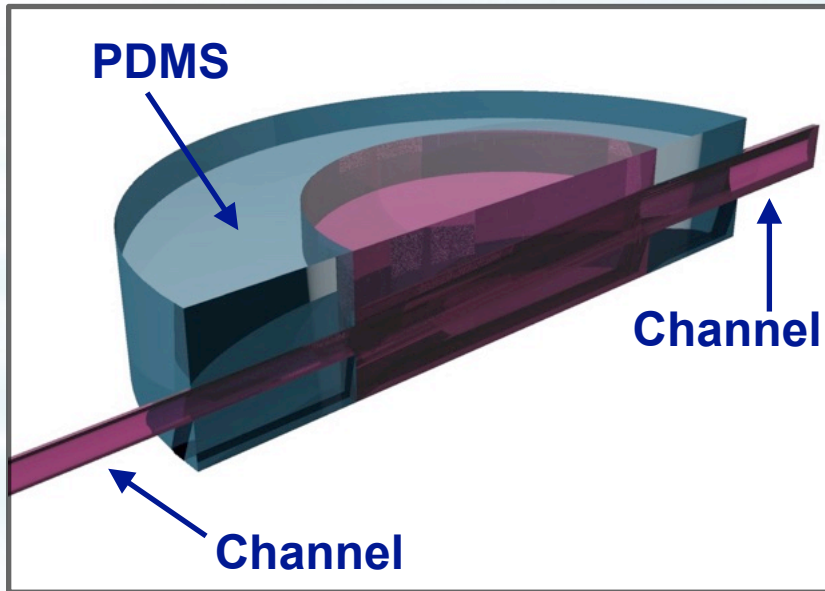
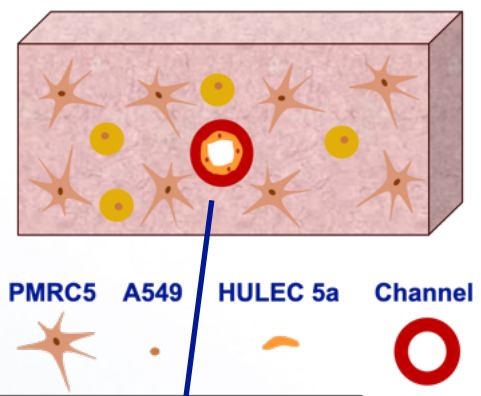
HULEC 5a

Channel



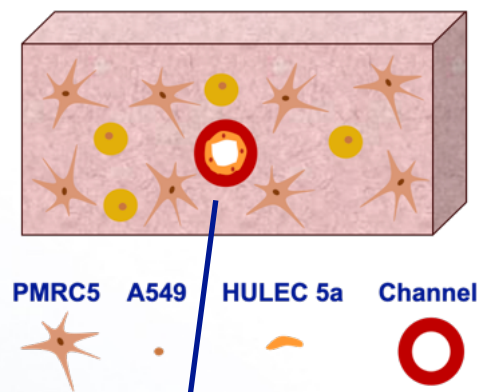
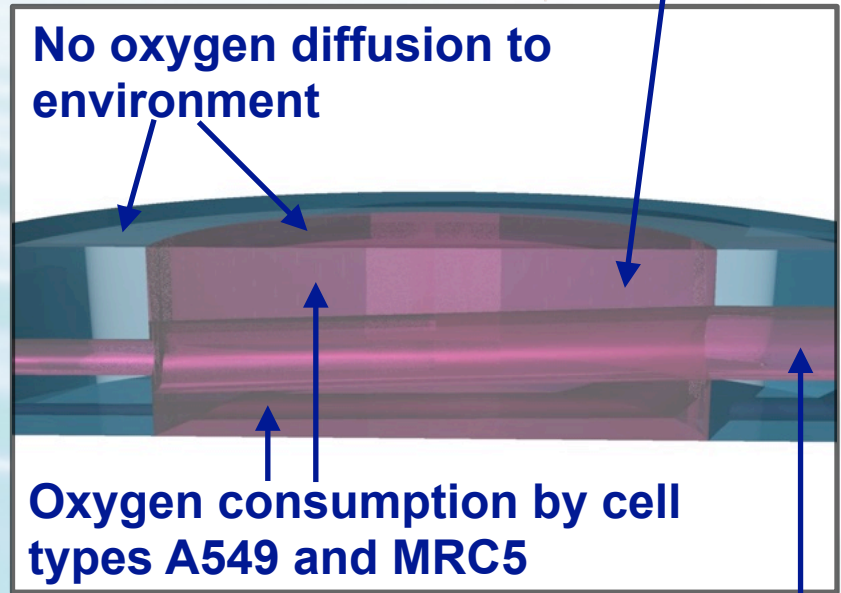
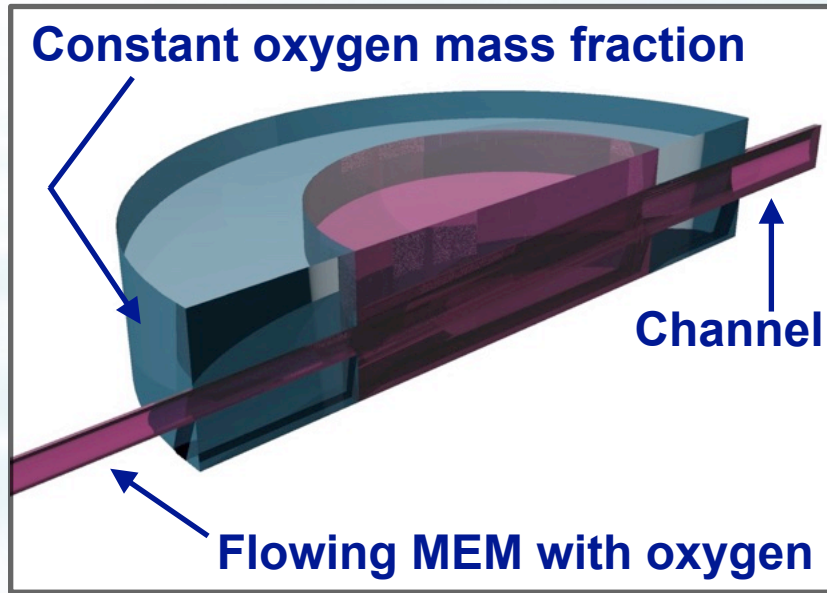
The Real Process - 3

- Solution domain: 3D CAD rendering



The Real Process - 4

- **Boundary conditions**

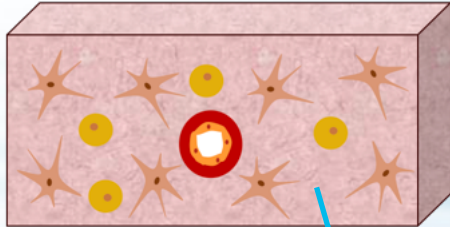


Agenda

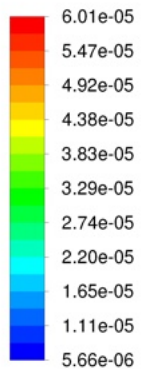
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First Results of Numerical Simulations - 1

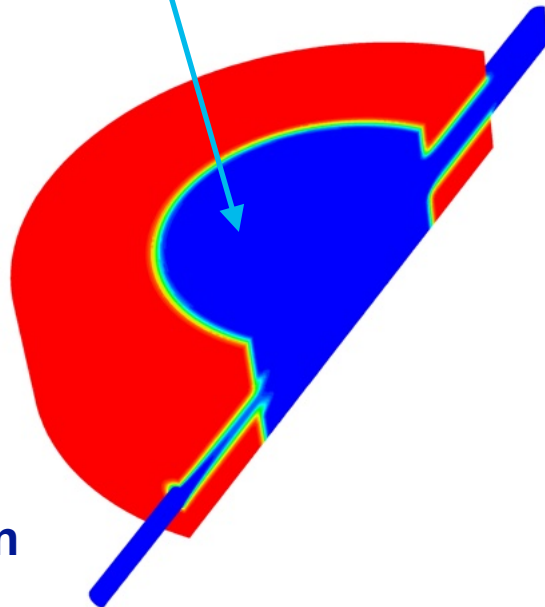
- **O₂ Mass fraction evolution**



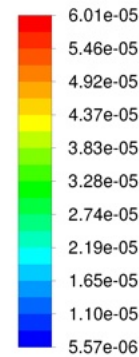
Time: after 10 s



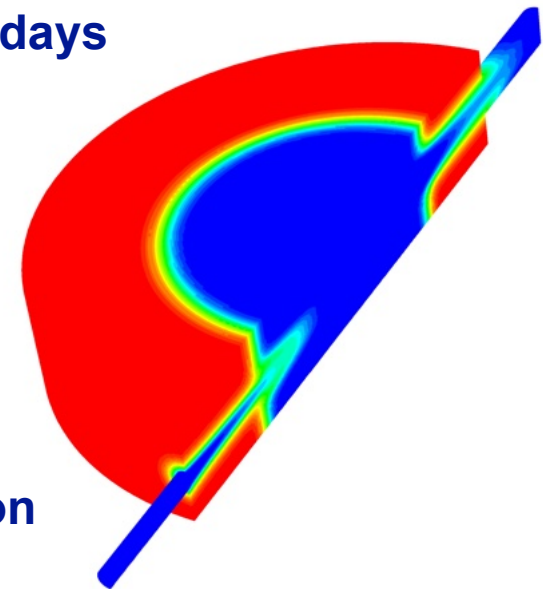
O₂ Mass fraction



Time: after 2.5 days



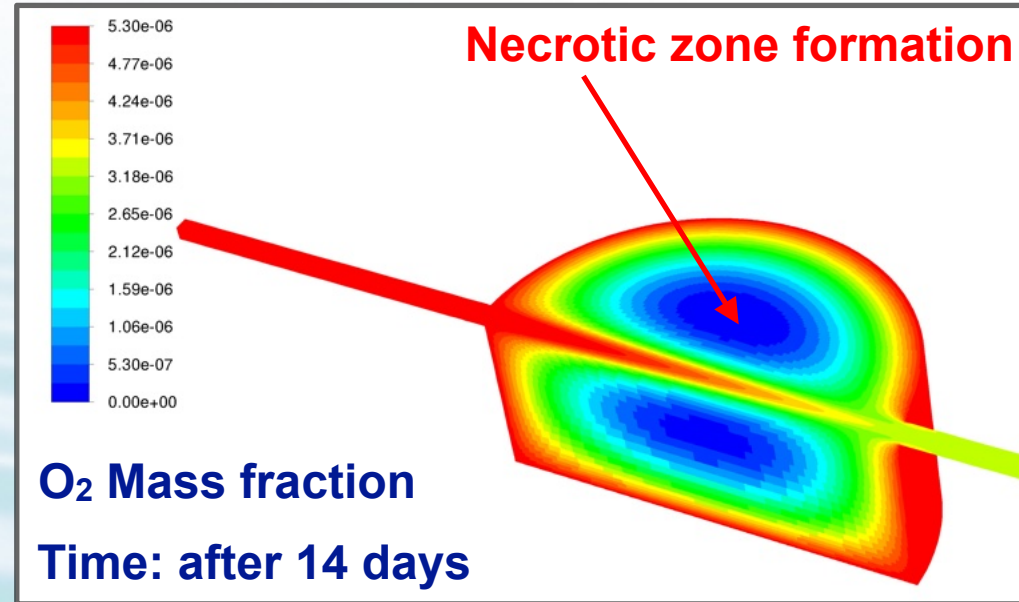
O₂ Mass fraction



First Results of Numerical Simulations - 2

- **Benefits of Computer simulation:**

- One can look inside a volume of a liquid or gas - at every point
- This generates new insights into the process
- and an increase of quality
- as well as a shortening development time and cost



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Future Steps

- **Implementation of additional program in Ansys Fluent to consider physical O₂ max fraction in MEM**
- **Check of iteration convergence and mesh convergence (i.e. check of numerical accuracy)**
- **Comparison with the O₂ evolution in the experimental model**
- **Avoid necrotic zones:**
 - **Optimization of the flow rate to obtain O₂ optimal mass consumption with respect to MEM renewal in the OoC**



References

Reference values

- MEM

	Density [kg/m ³]	Viscosity [Pa s]	Flow rate [μ l/min]
MEM	1.01E+03	9.30E-04 ^[1]	200

- O₂ mass fraction in mem

	D COEF. O ₂ [m ² /s]	Molarity in Medium	%wt of O ₂ in MEM
O₂ mass fraction in MEM	2.88E-09	1.67E-04 ^[1]	5.30E-06

- O₂ consumption per cell (Kg/s)

	Cell Type	#cell/ml	O ₂ consumption per cell [Kg/s]
O₂ consumption	A549	1.50E+06	2.66E-18 ^[2]
	MRC5	1.50E+06	1.06E-17 ^[3]

[1] <https://www.pnas.org/doi/pdf/10.1073/pnas.91.25.12248><https://www.pnas.org/doi/pdf/10.1073/pnas.91.25.12248>

[2] <https://www.pnas.org/doi/pdf/10.1073/pnas.91.25.12248>

[3] <https://www.nature.com/articles/s41598-017-00130-x>

