

Microfluidic extrusion of cell-laden hydrogel fibers for 3D Bio-printing

Dr. Cristina Colosi

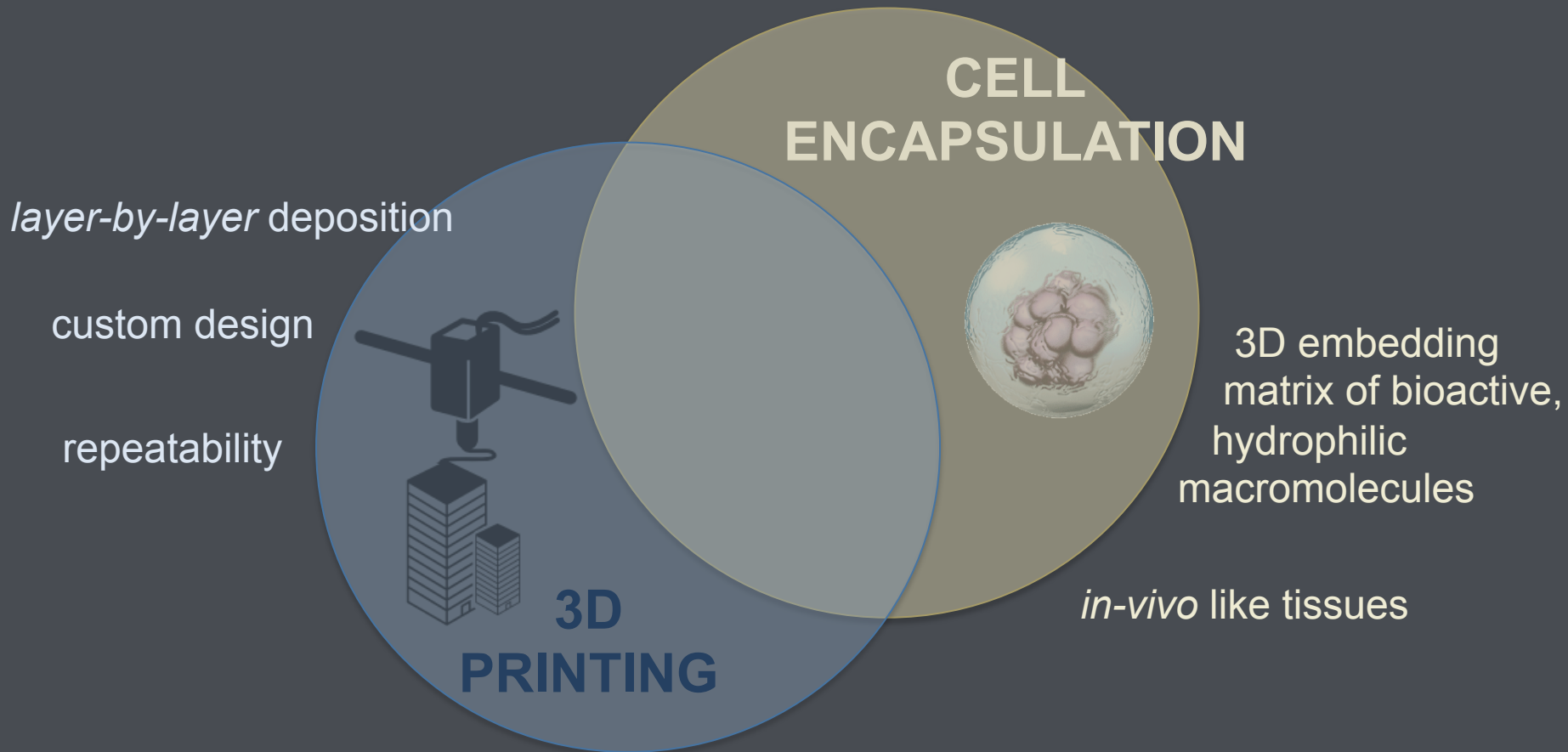
CLNS – iit@Sapienza

Bioprinting Lab.

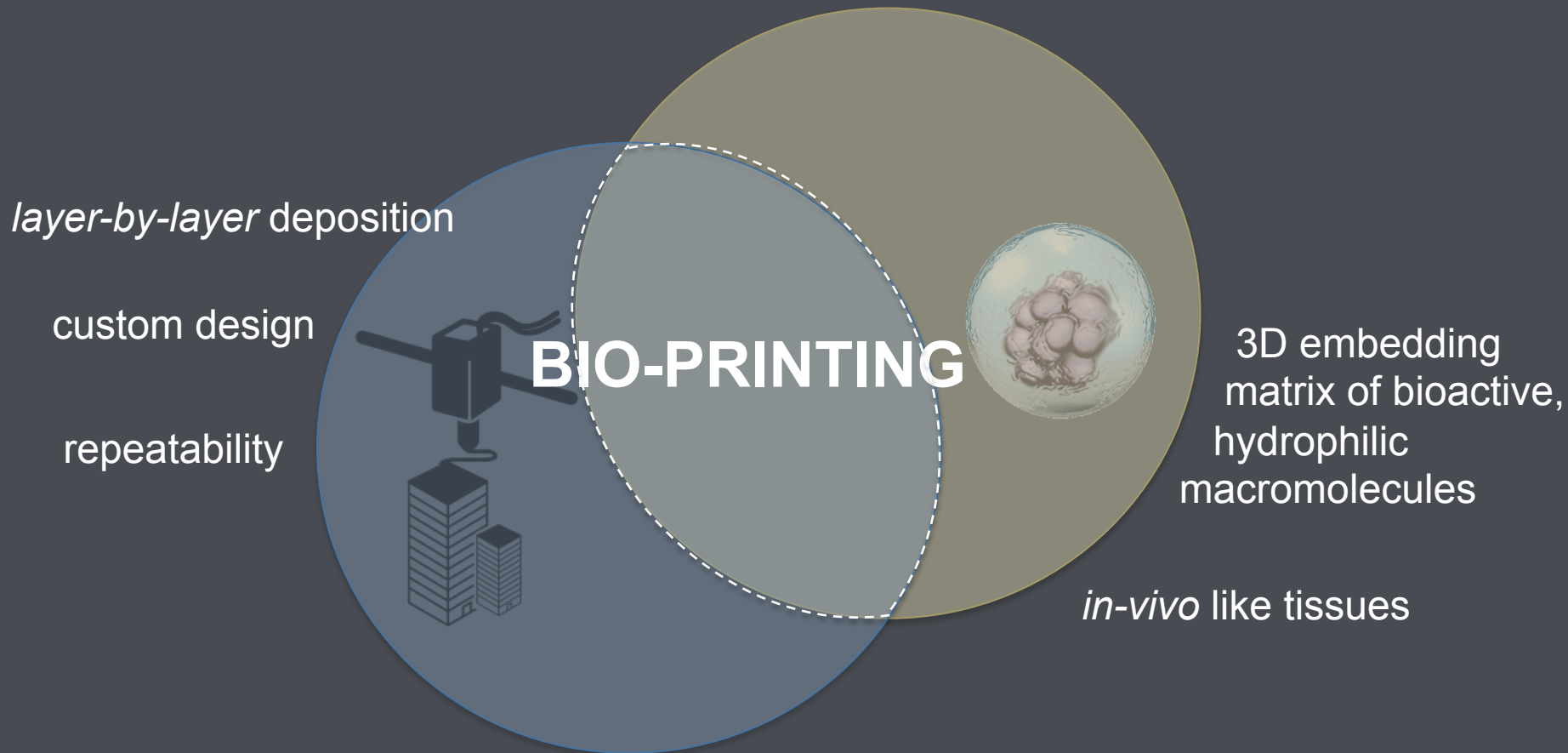
Viale Regina Elena 291 - Roma

Nano Rome, 20-23 September
2016 Innovation
Conference & Exhibition

3D Bio-Printing

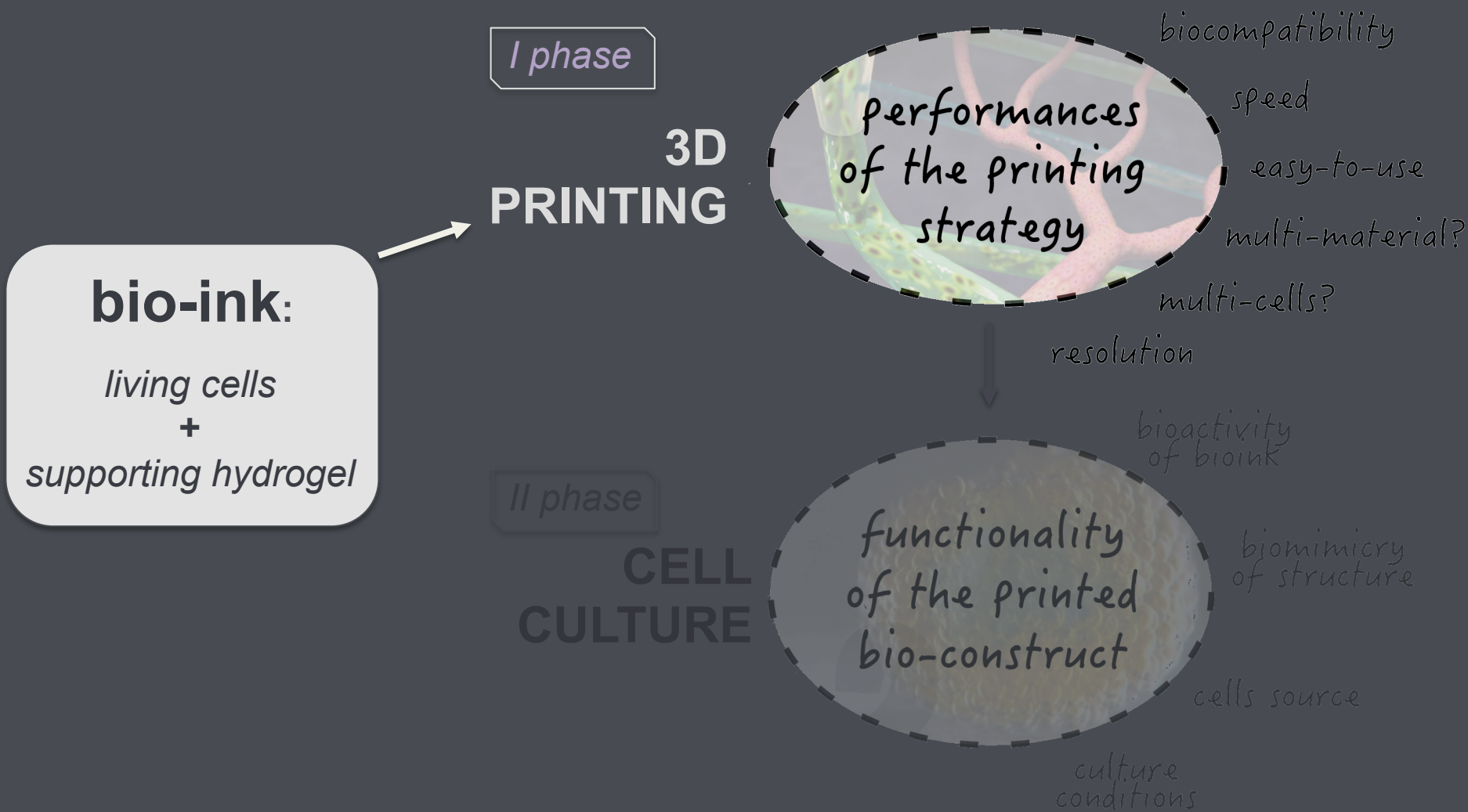


3D Bio-Printing

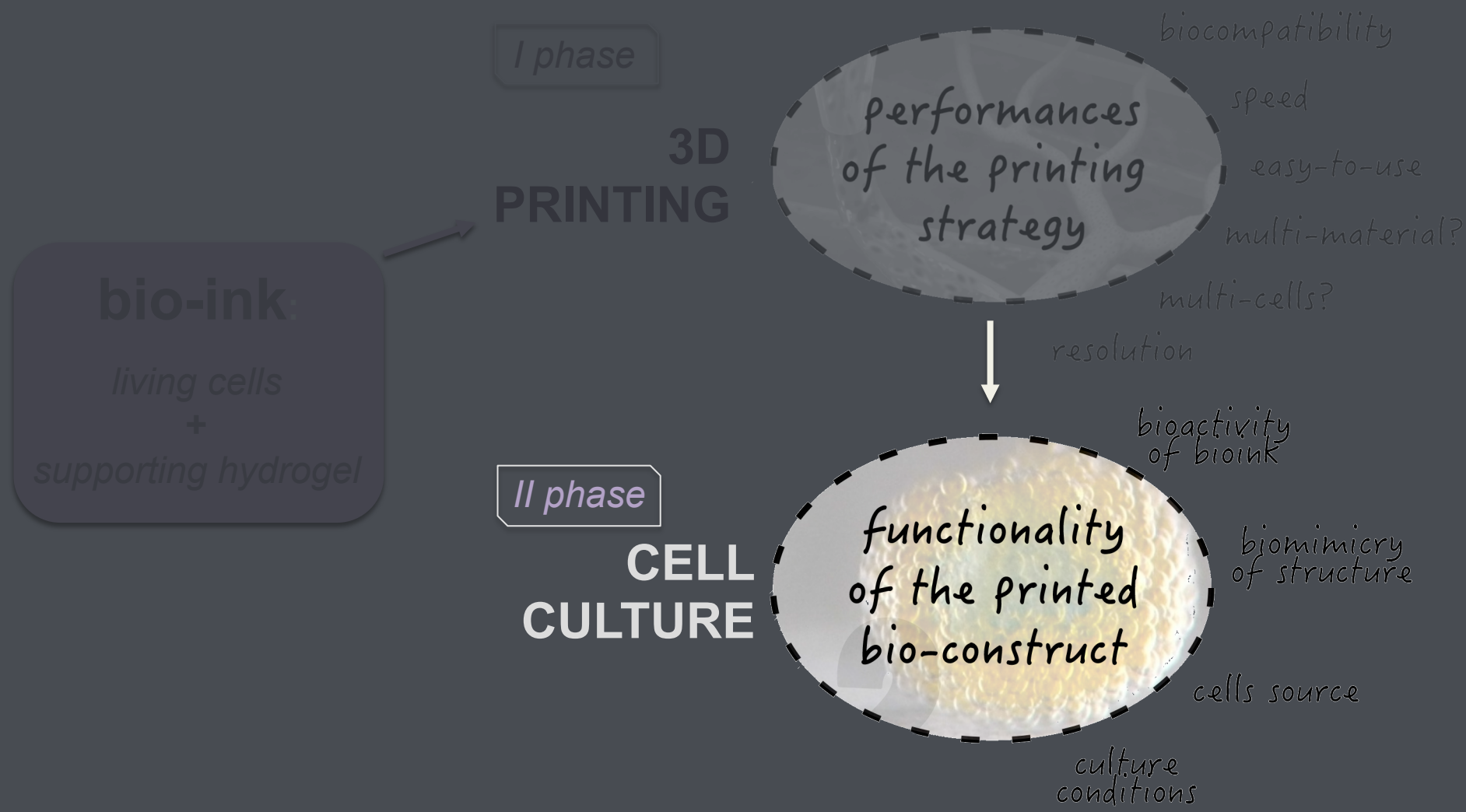


Design and fabrication of custom-designed artificial 3D living tissues

3D Bio-Printing

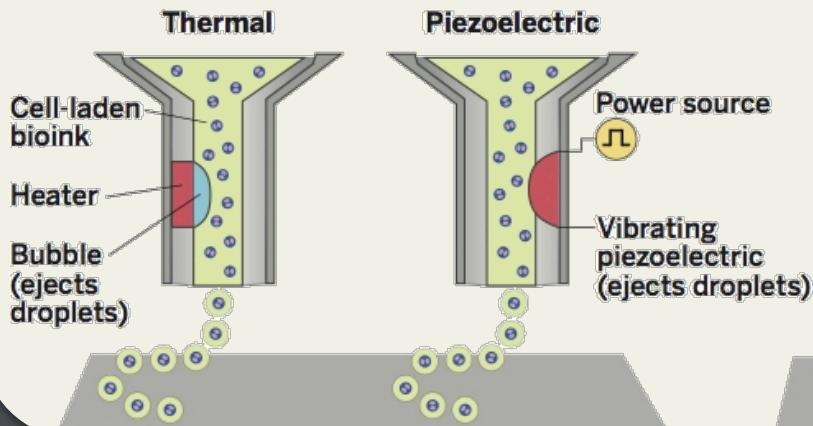


3D Bio-Printing

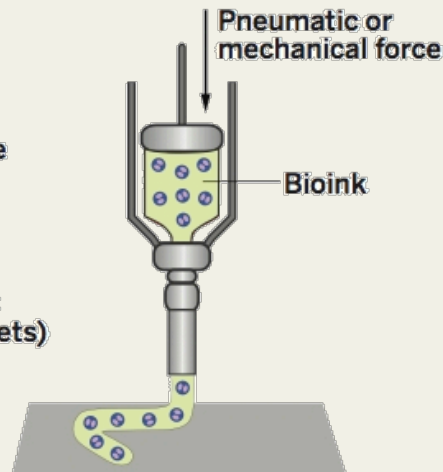


3D Bio-Printing

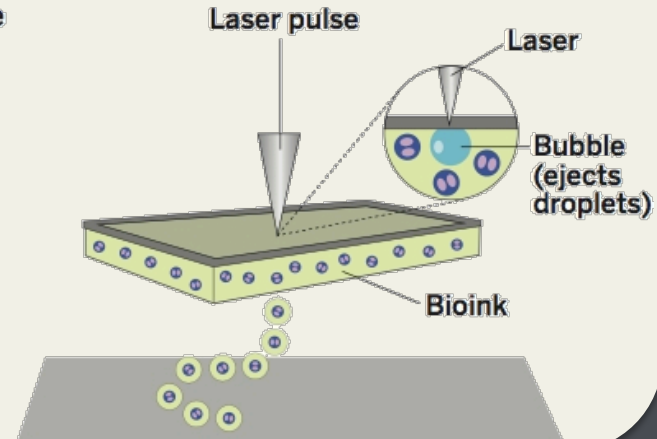
Ink-jet bioprinter



Extrusion bioprinter



Laser-assisted bioprinter



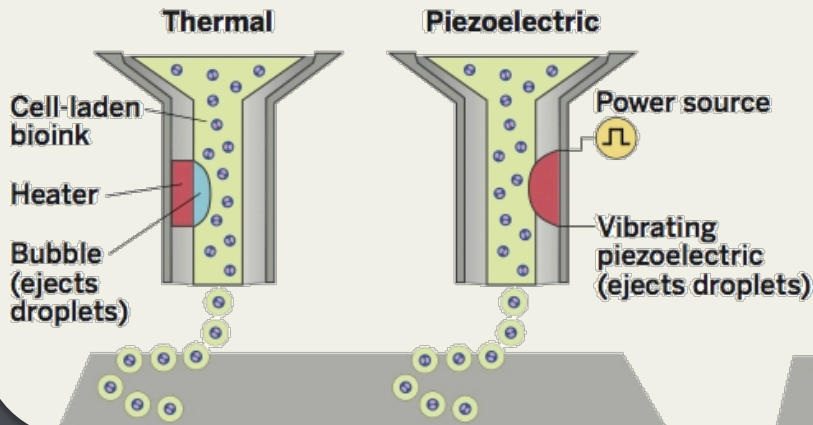
- fast
- low cost
- high precision
- suitable only for low density hydrogels
- poor mechanical properties

- fast
- low cost
- big range of suitable materials
- poor precision

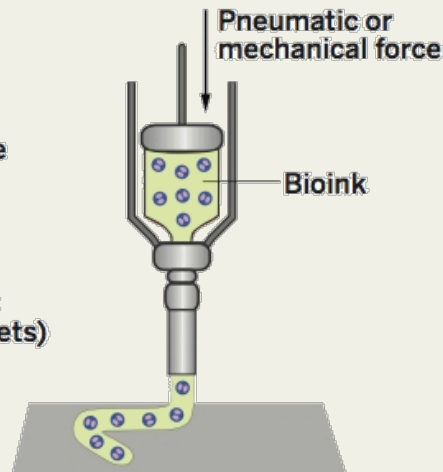
- high precision
- no material limitations
- high costs
- labor expensive

3D Bio-Printing

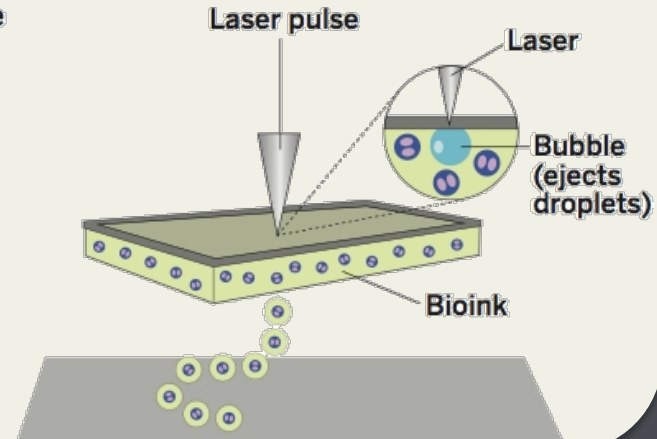
Ink-jet bioprinter



Extrusion bioprinter



Laser-assisted bioprinter



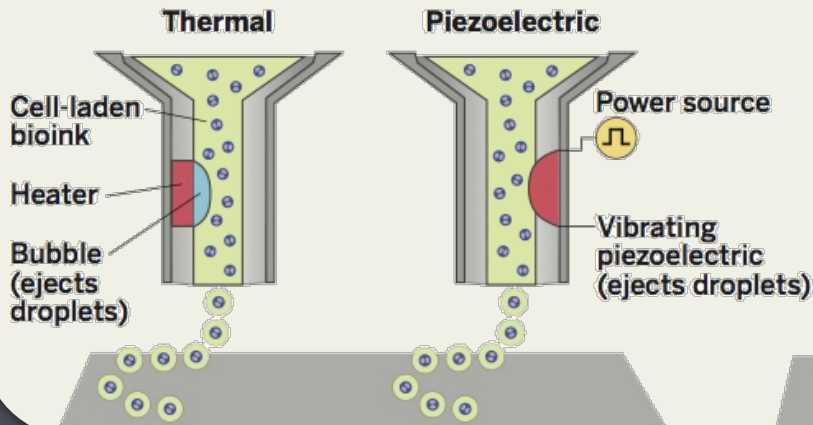
- fast
- low cost
- high precision
- suitable only for low density hydrogels
- poor mechanical properties

- fast
- low cost
- big range of suitable materials
- poor precision

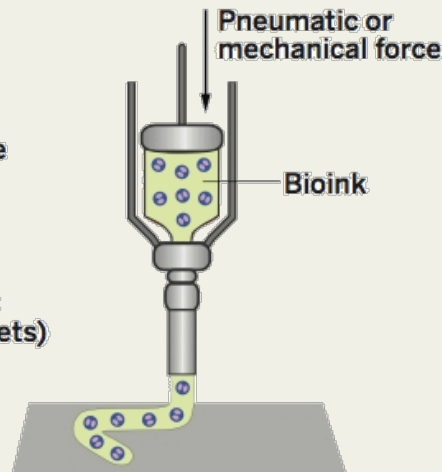
- high precision
- no material limitations
- high costs
- labor expensive

3D Bio-Printing

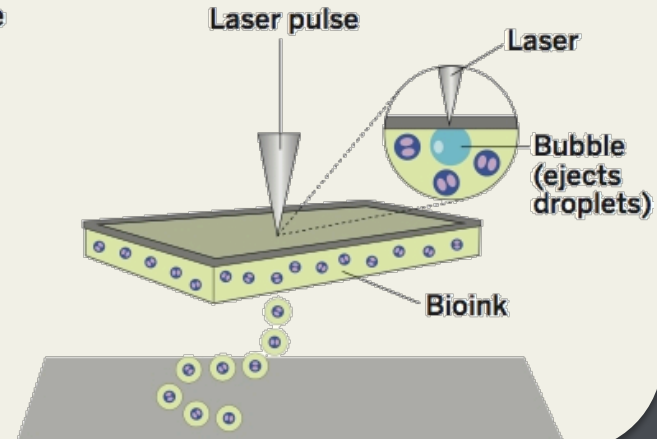
Ink-jet bioprinter



Extrusion bioprinter



Laser-assisted bioprinter



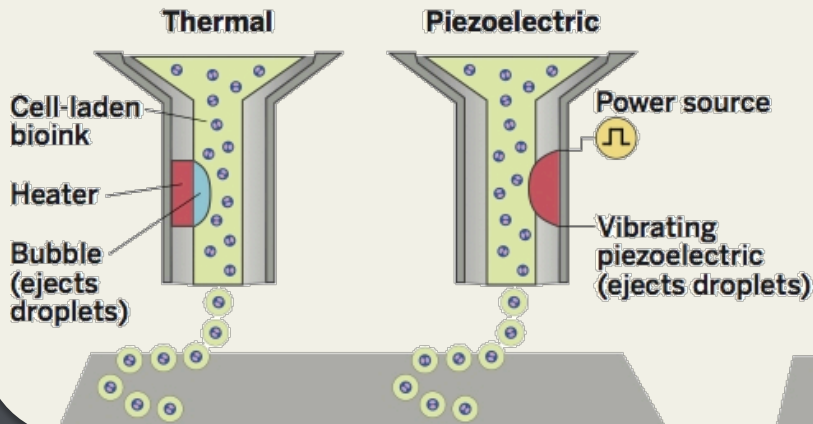
- fast
- low cost
- high precision
- suitable only for low density hydrogels
- poor mechanical properties

- fast
- low cost
- big range of suitable materials
- poor precision

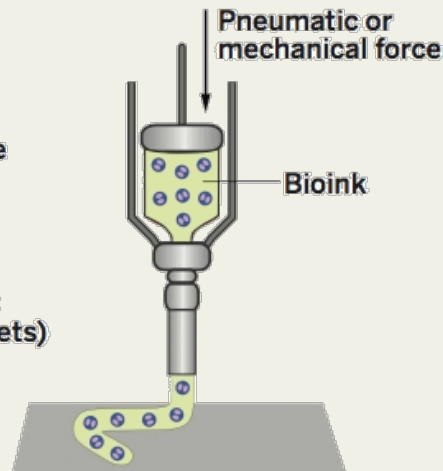
- high precision
- no material limitations
- high costs
- labor expensive

3D Bio-Printing

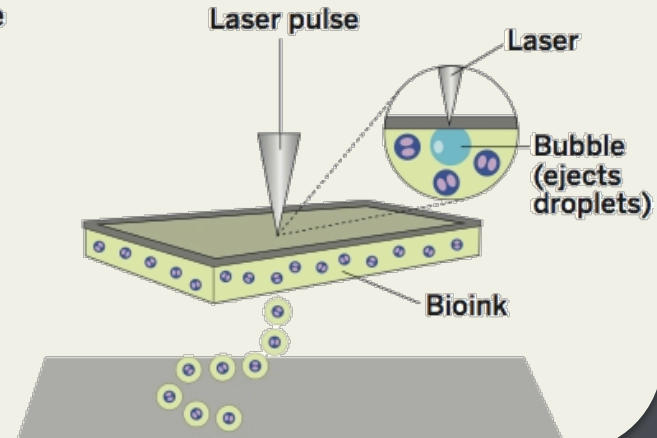
Ink-jet bioprinter



Extrusion bioprinter



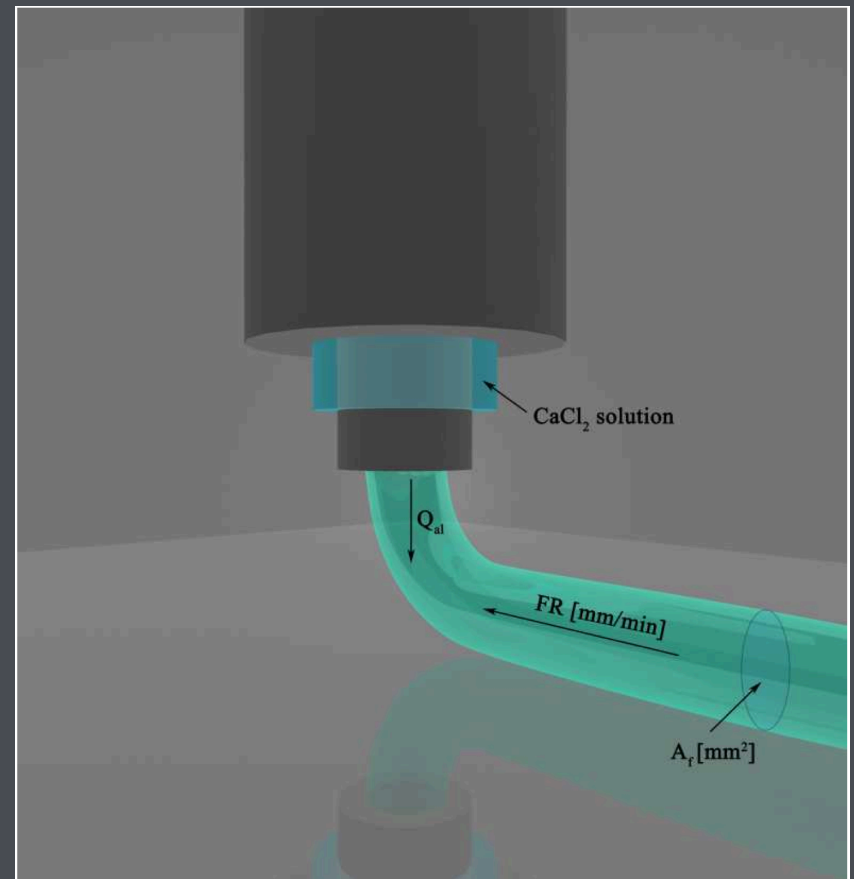
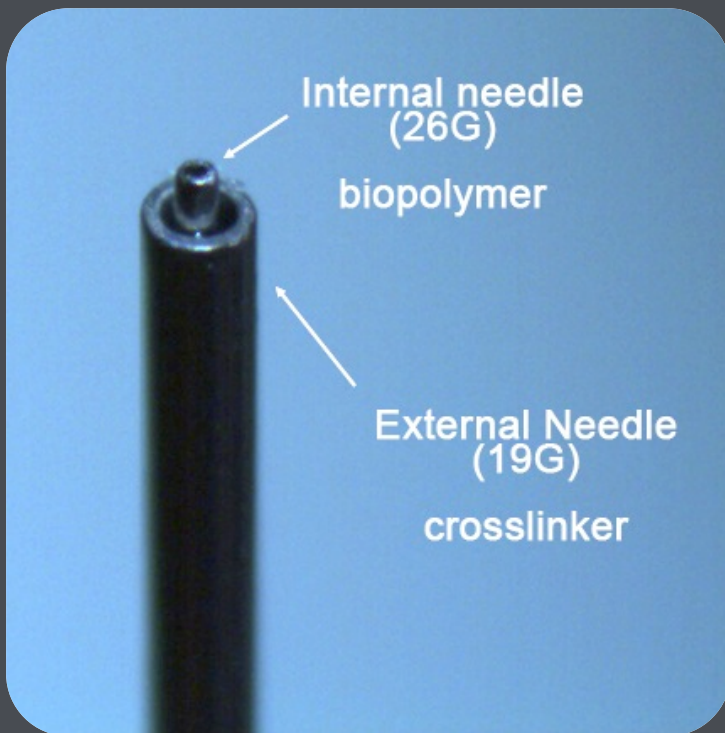
Laser-assisted bioprinter



- fast
- low cost
- big range of suitable materials
- ~~poor precision~~ → **Microfluidic extrusion**

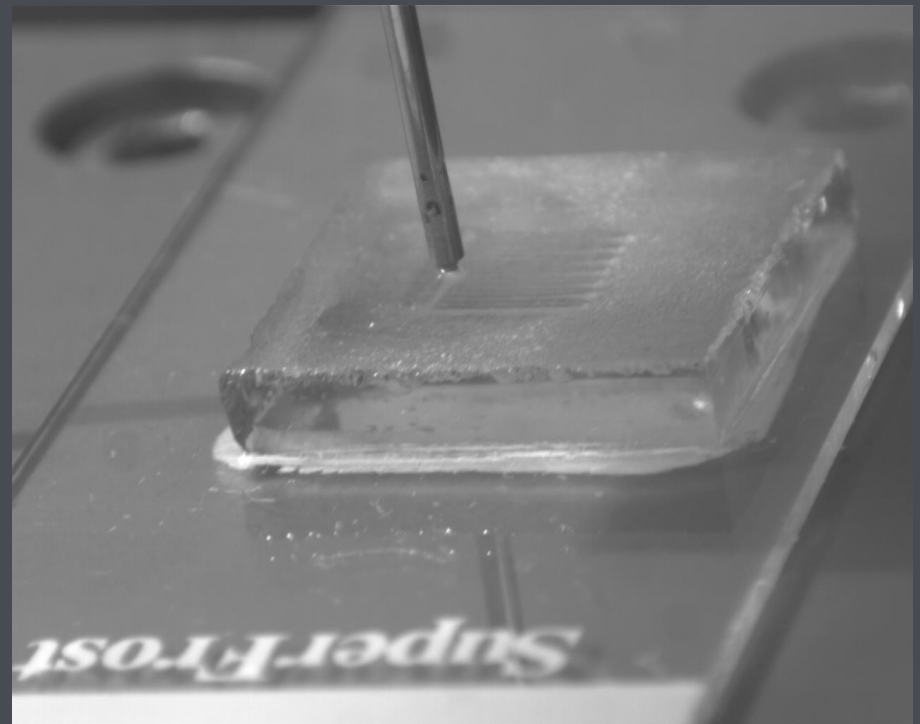
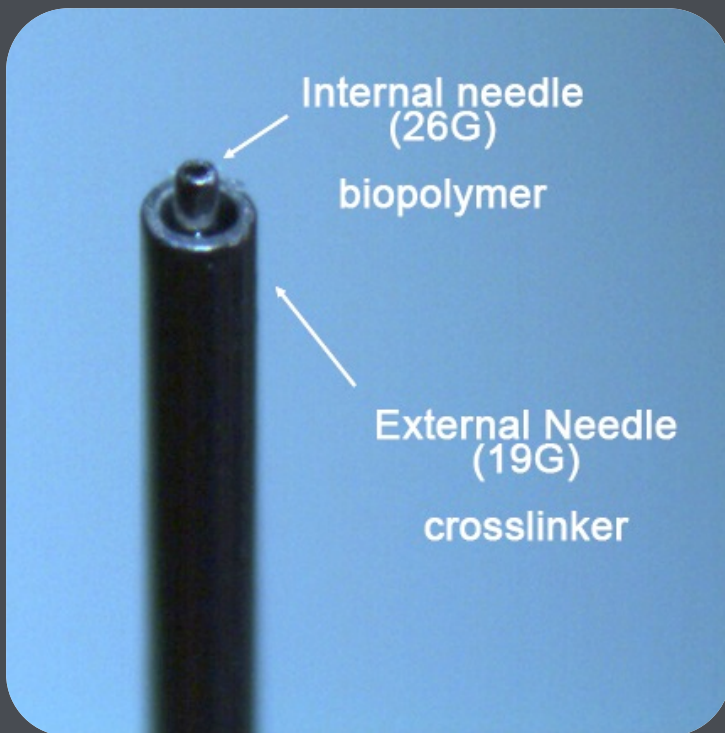
Microfluidic Coaxial Extrusion

Coaxial needle extruder for fast gelling macromolecules



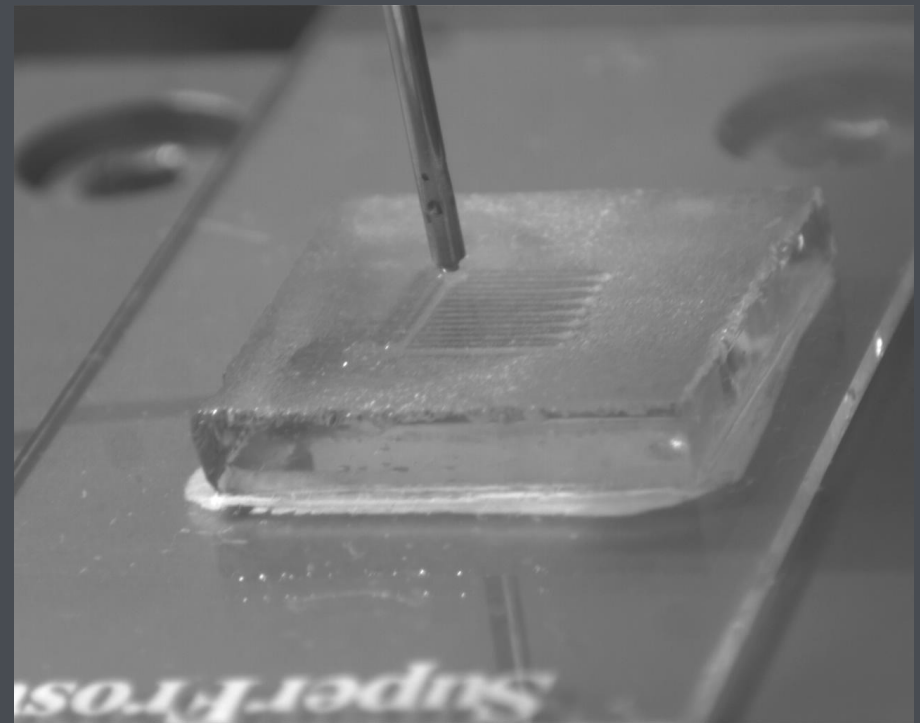
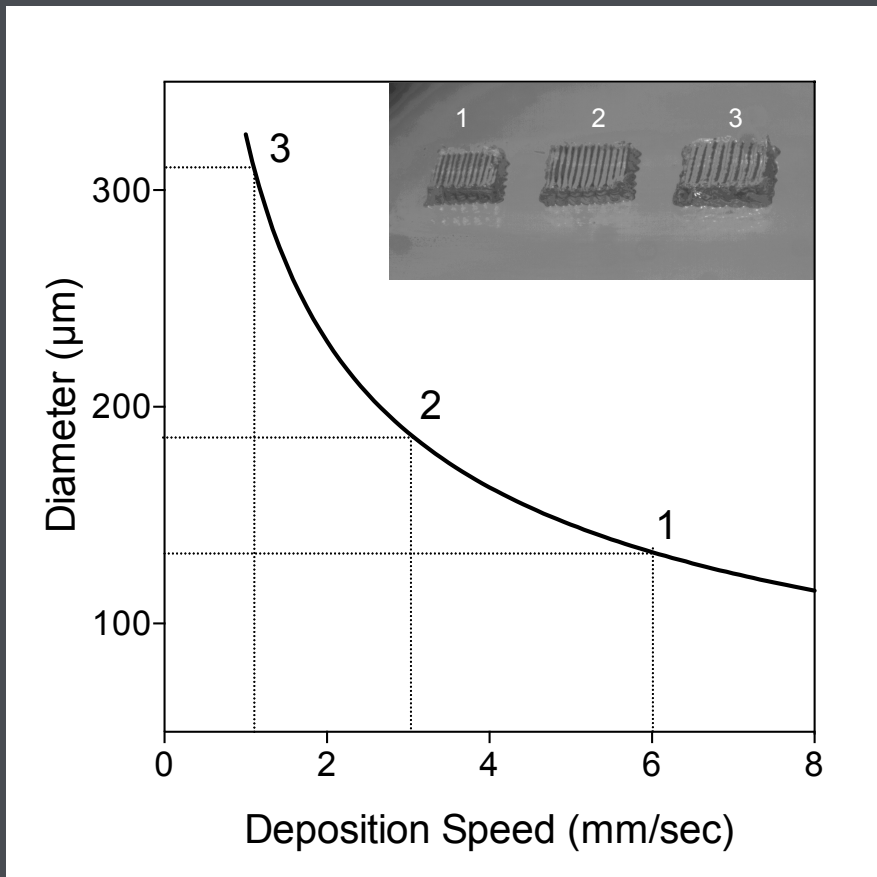
Microfluidic Coaxial Extrusion

Coaxial needle extruder for fast gelling macromolecules



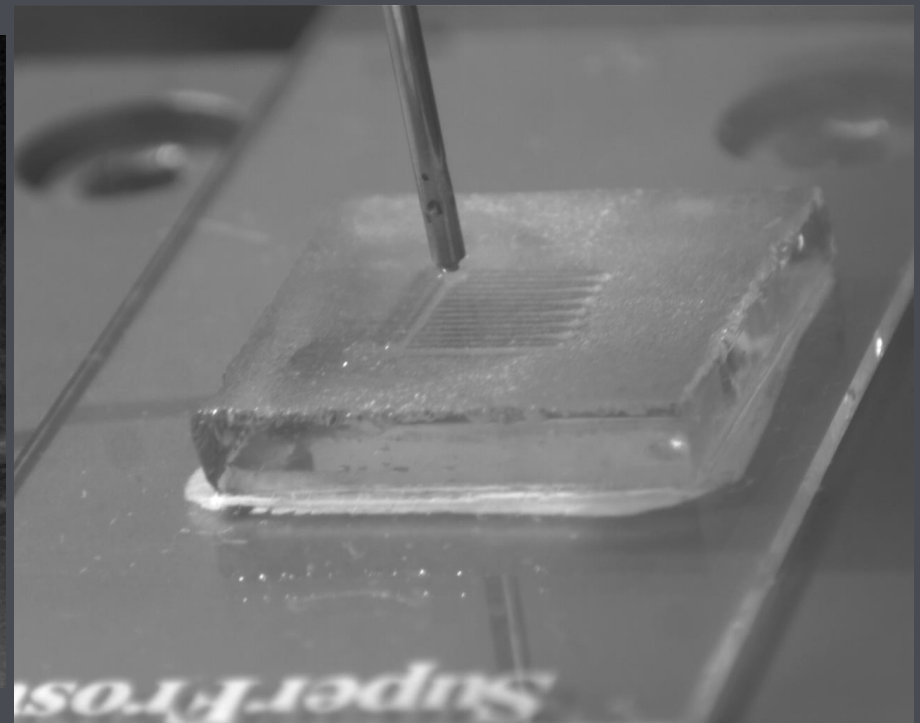
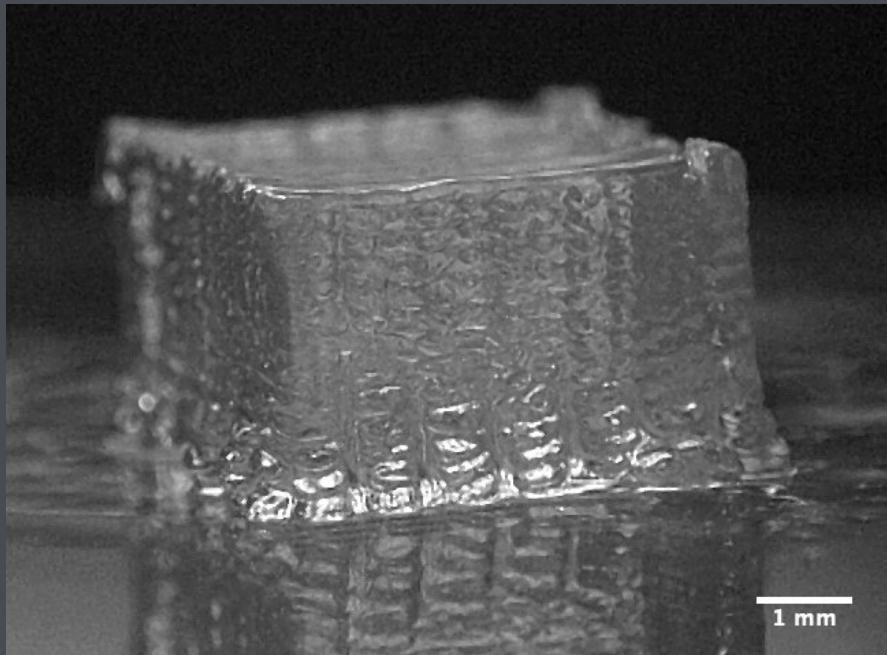
Microfluidic Coaxial Extrusion

- different fiber diameters (300 - 100 μm)



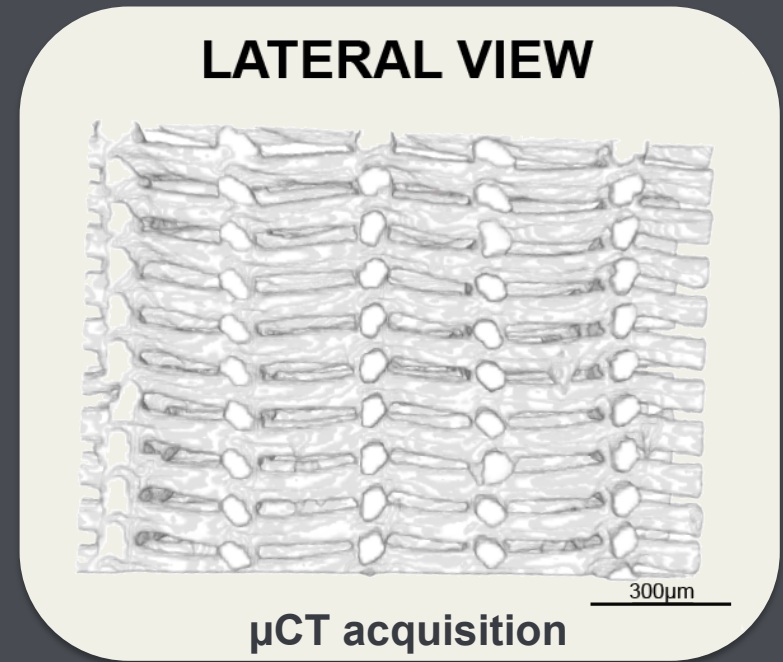
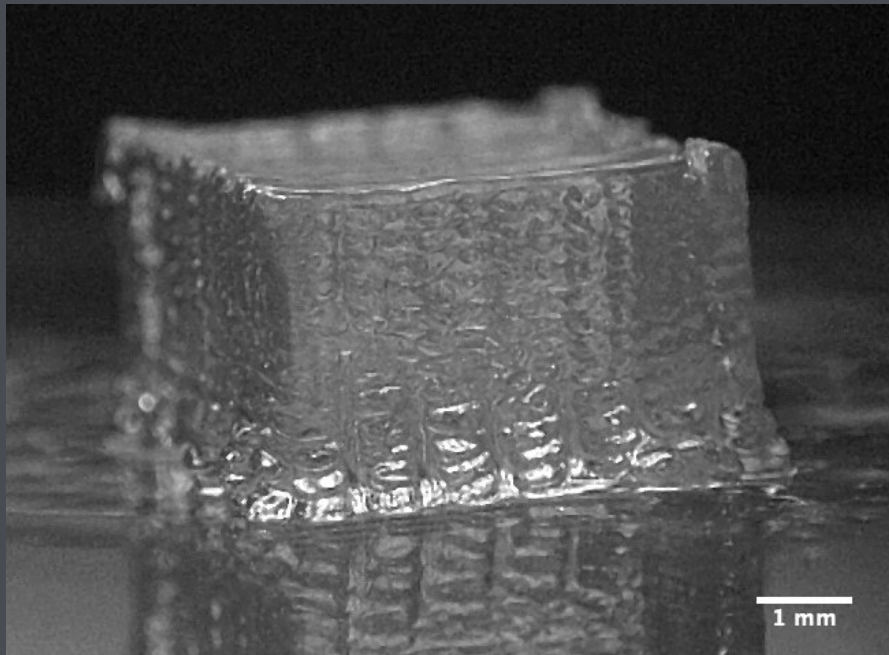
Microfluidic Coaxial Extrusion

- multilayered structures



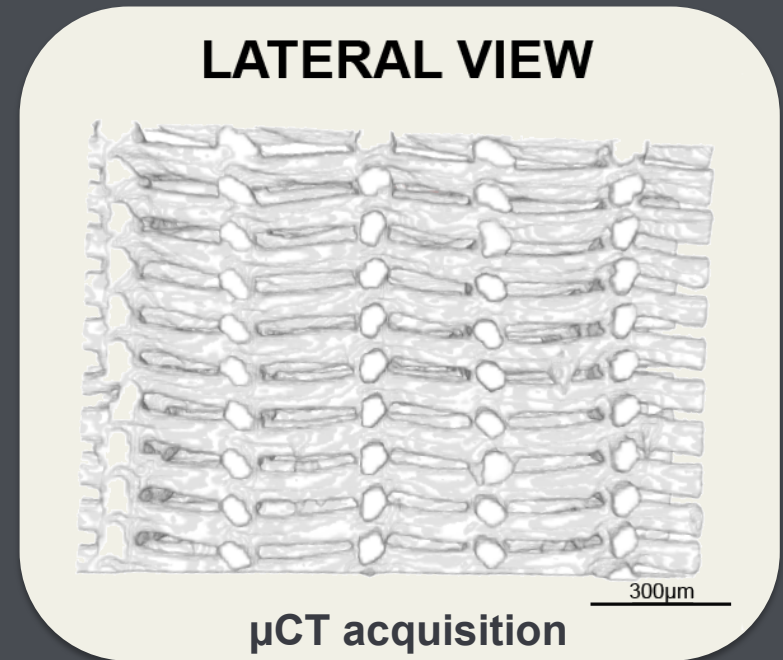
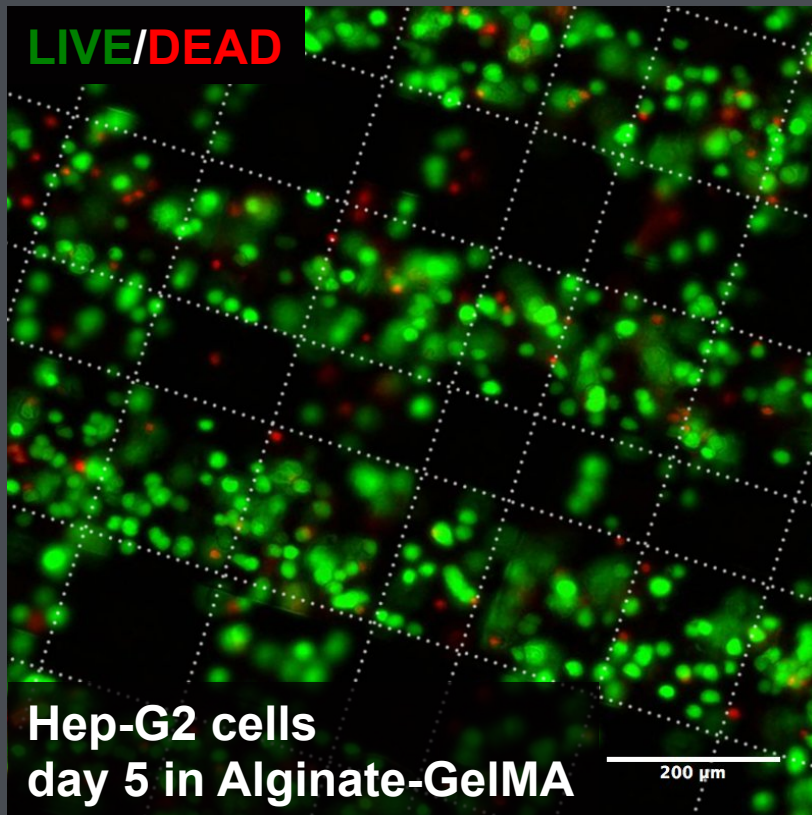
Microfluidic Coaxial Extrusion

- 3D interconnection among fibers



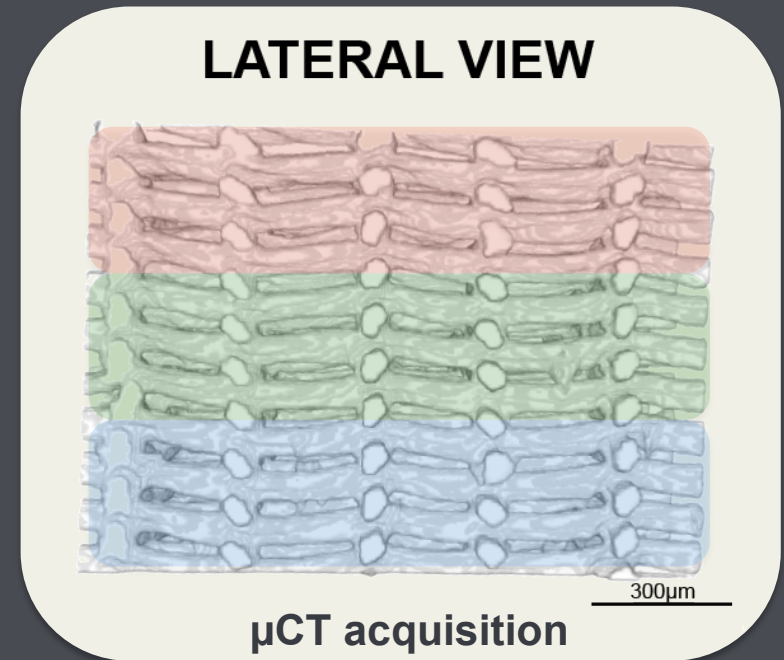
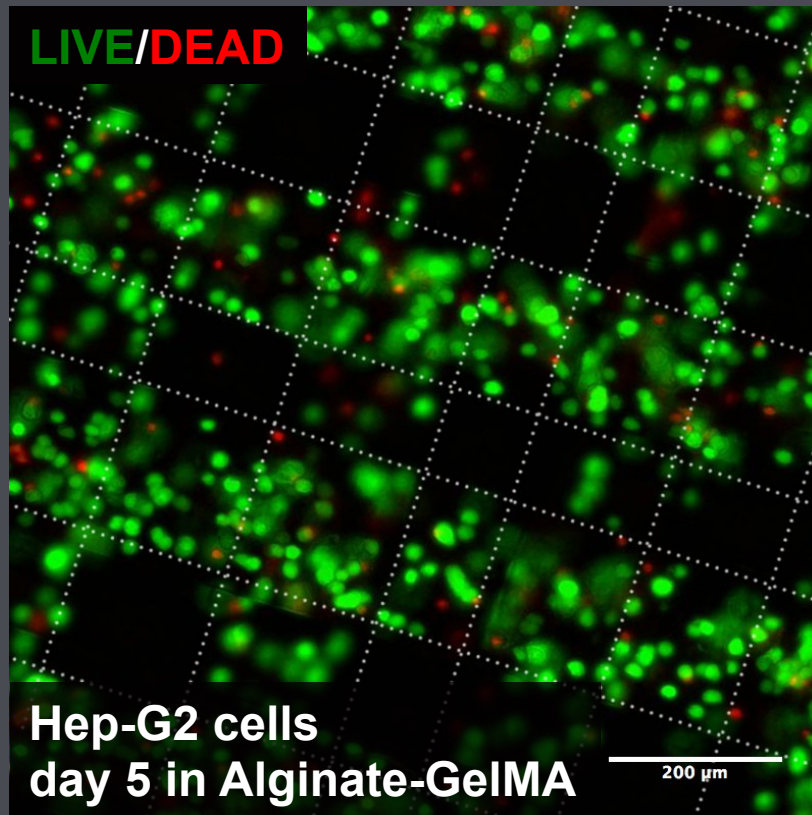
Microfluidic Coaxial Extrusion

- high cell viability



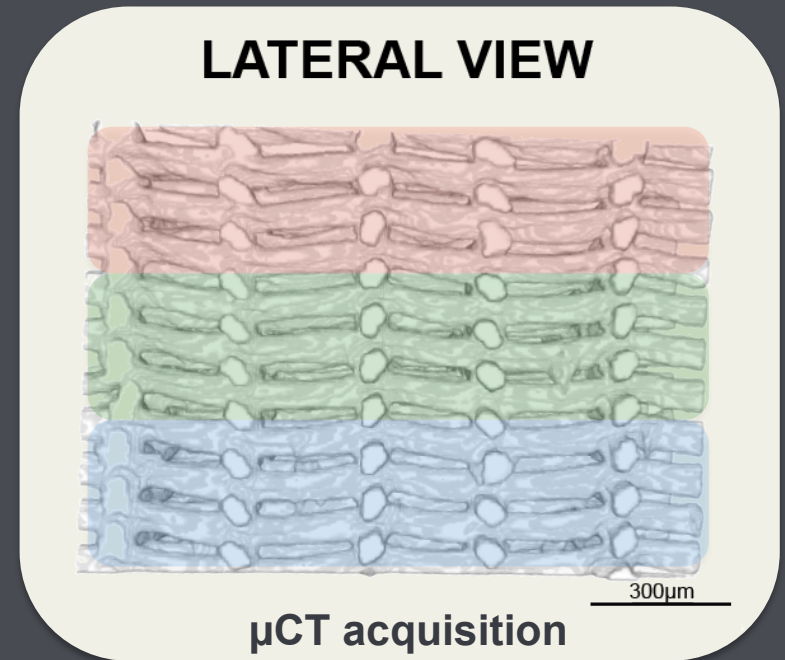
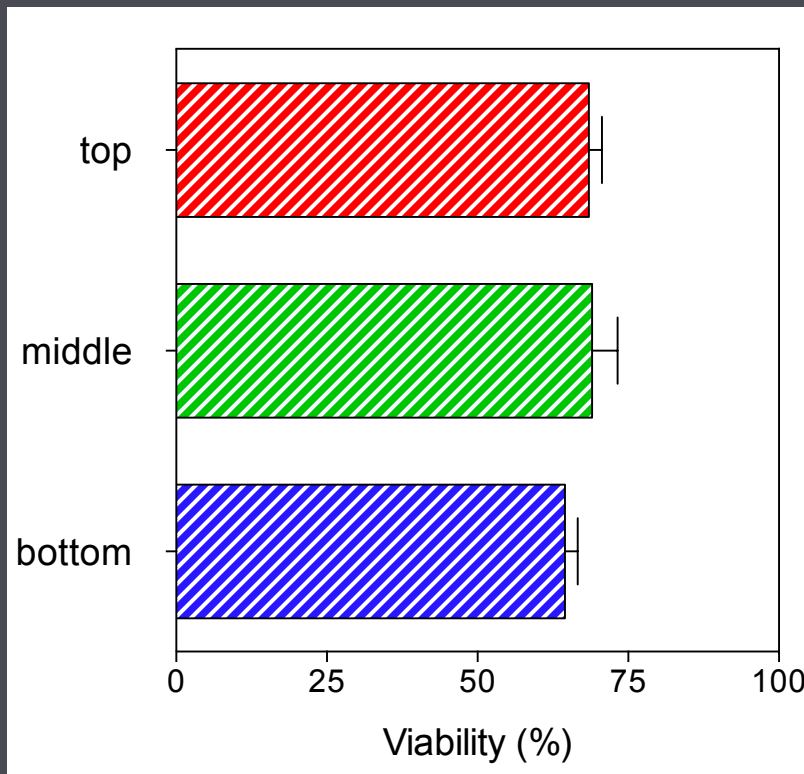
Microfluidic Coaxial Extrusion

- uniform cell viability



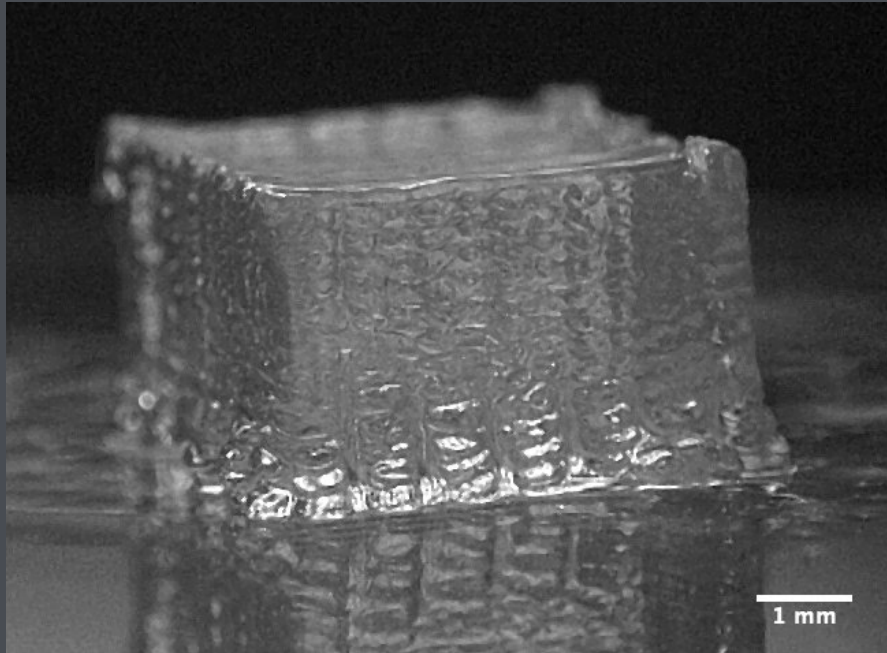
Microfluidic Coaxial Extrusion

- uniform cell viability



Microfluidic Coaxial Extrusion

- many possible extracellular matrix composition



Radical crosslinking (UV):

- gelatin (methacrylol)
- dextran (metacrylol)

Enzymatic crosslinking:

- fibrinogen

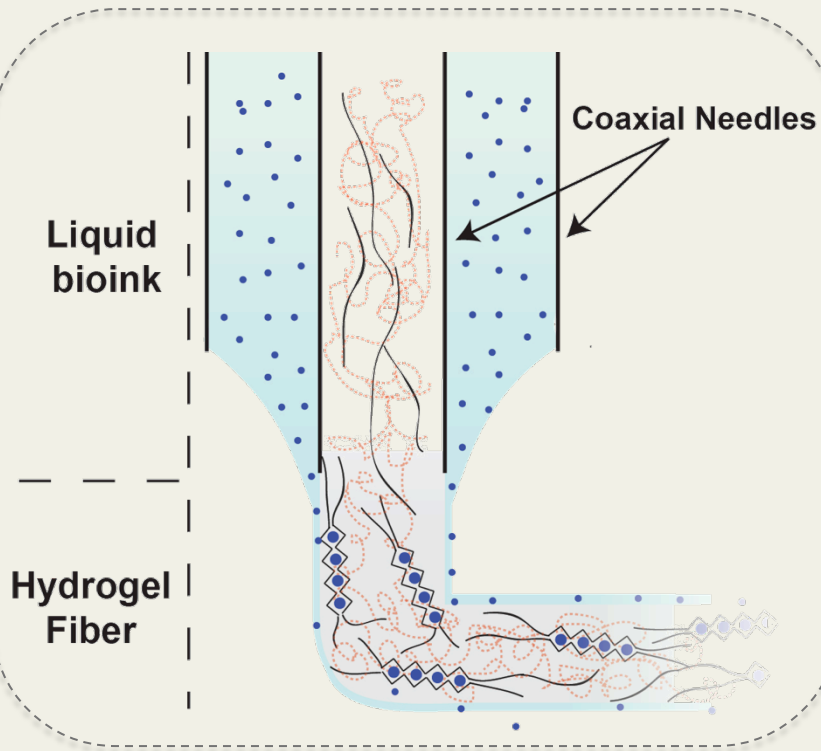
Self-assembly

- collagen
- fibroin

+
alginate

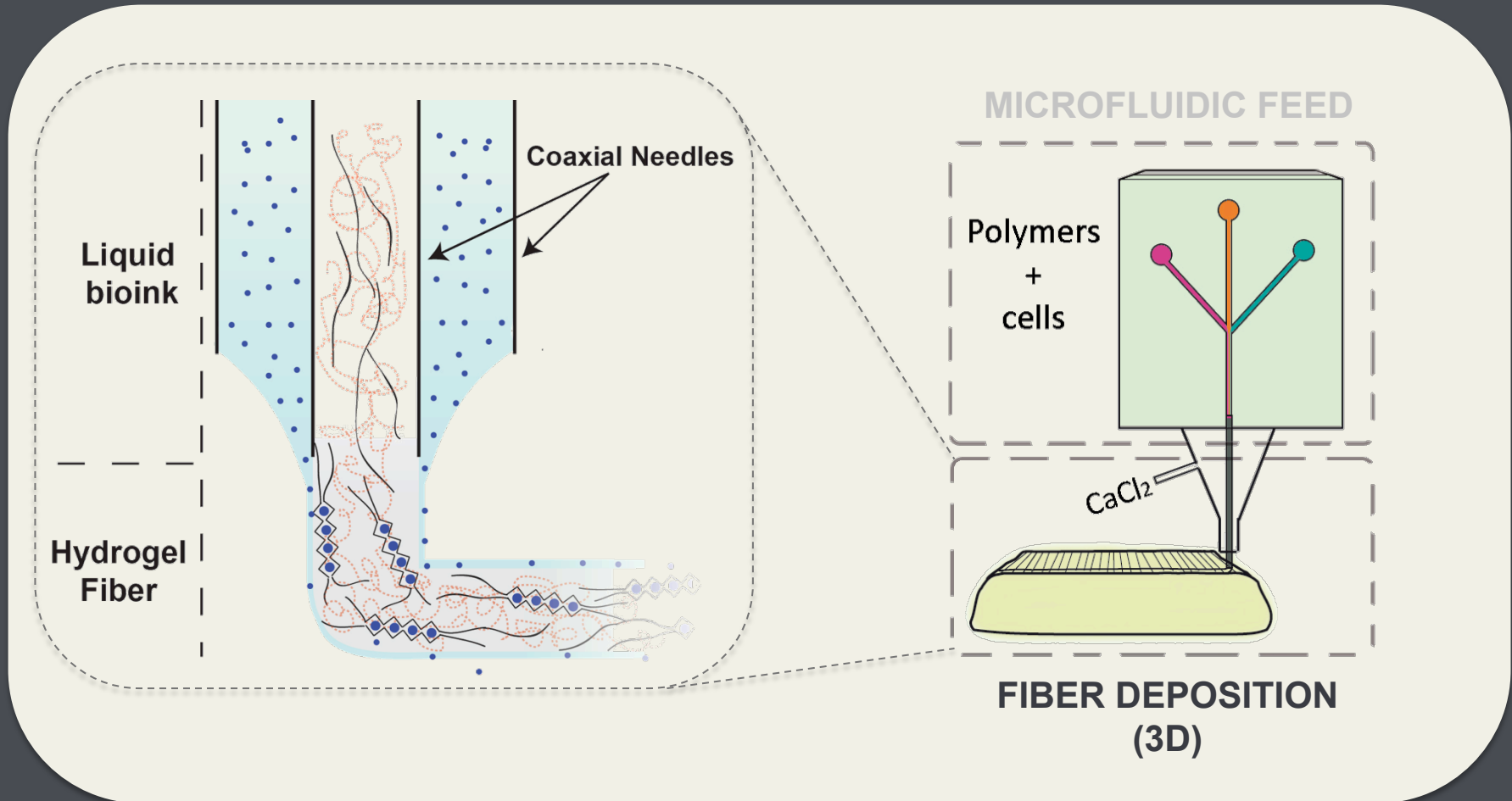
2-step crosslinking

Microfluidic Coaxial Extrusion

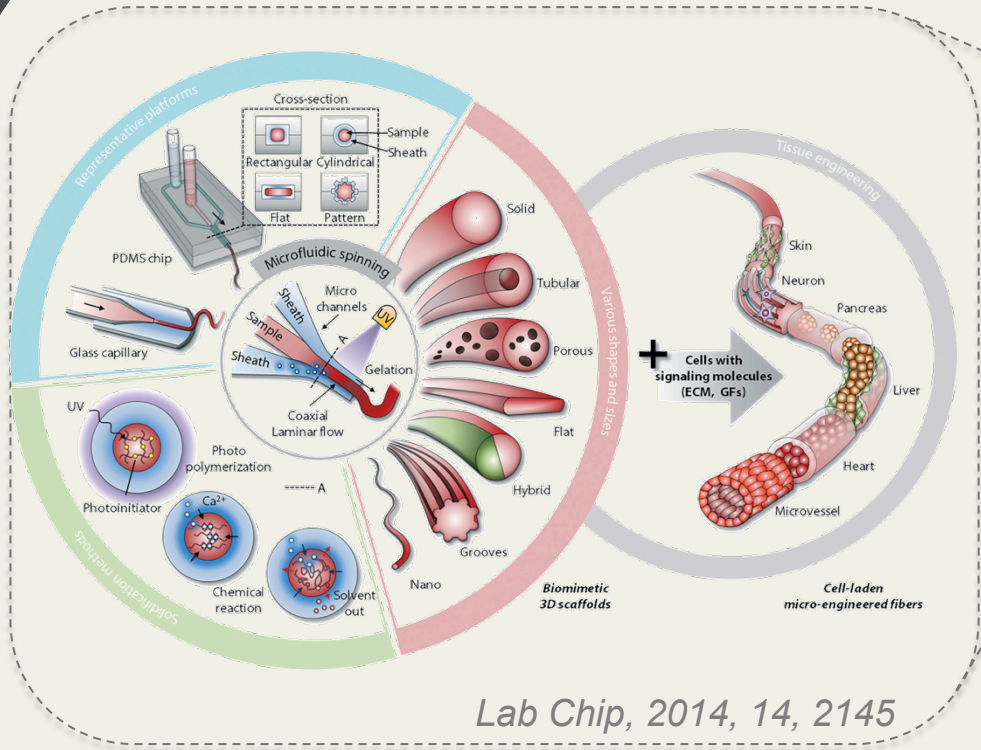


- fibers with diam = 300-100 μ m
- 3D interconnectivity
- high and uniform cell survival
- different possible compositions of embedding hydrogel

Microfluidic feeding

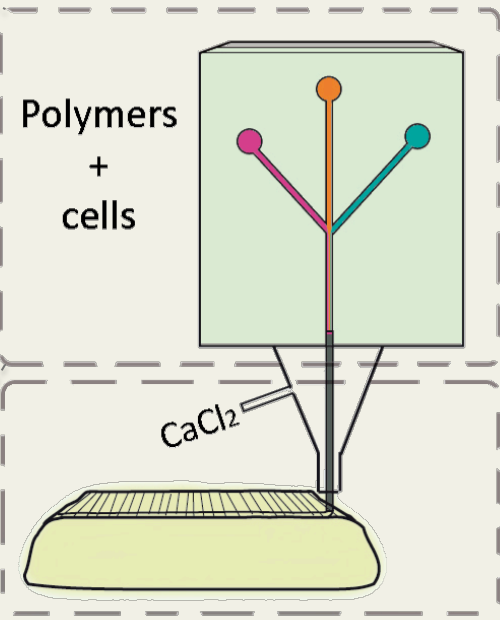


Microfluidic feeding



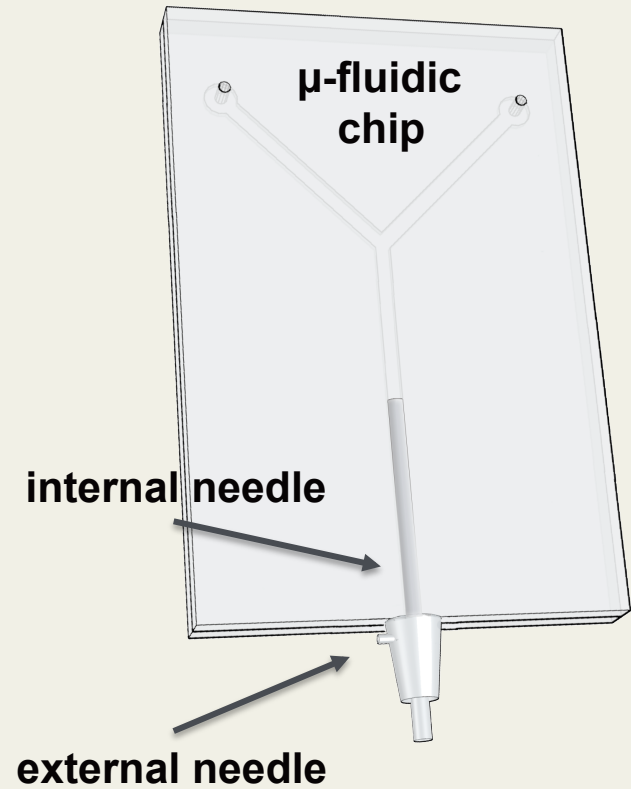
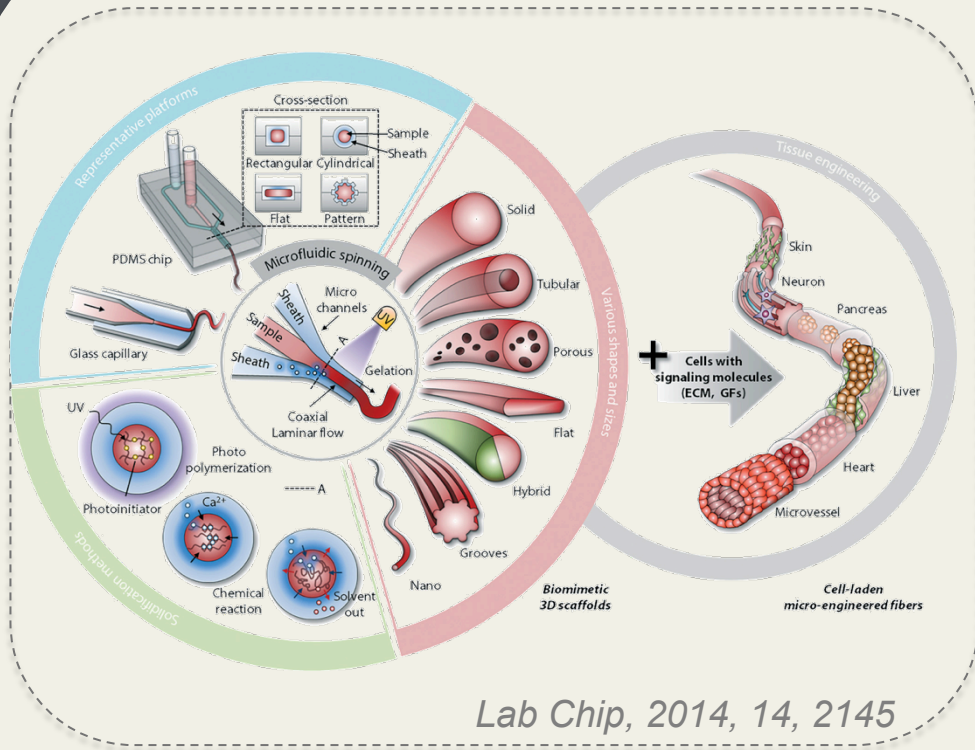
Lab Chip, 2014, 14, 2145

MICROFLUIDIC FEED



FIBER DEPOSITION (3D)

Microfluidic feeding



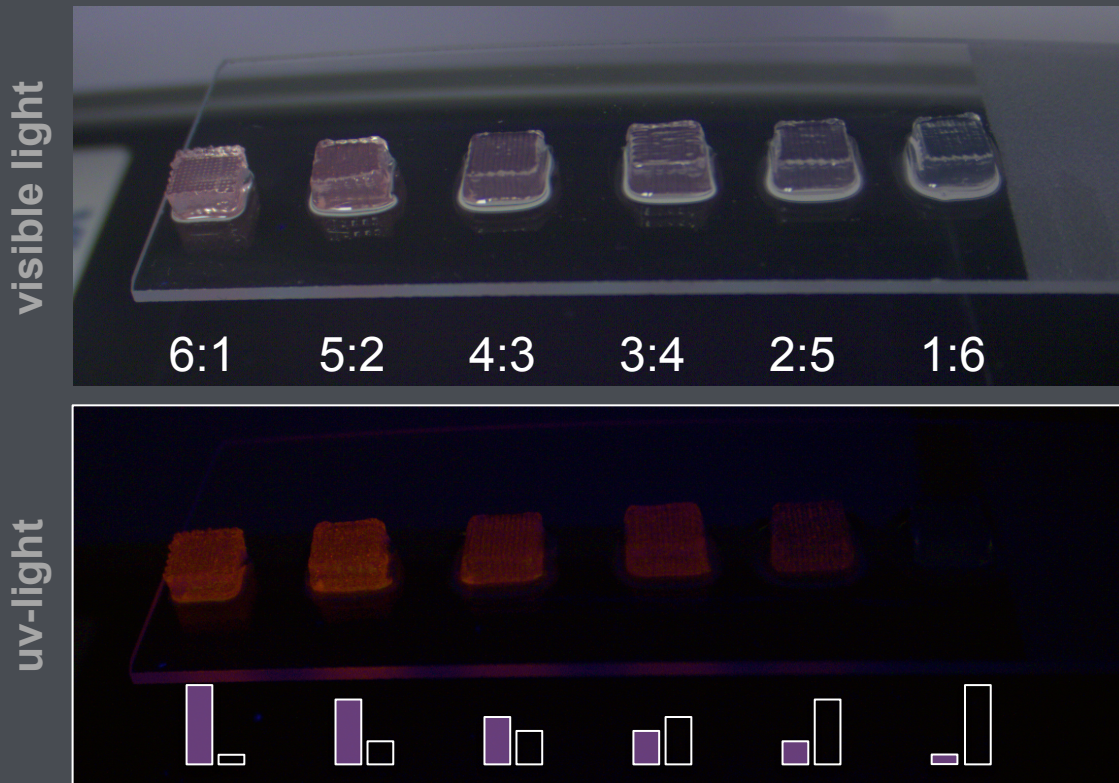
Microfluidic mixing

- Changing composition of the bioink *on-the-fly*



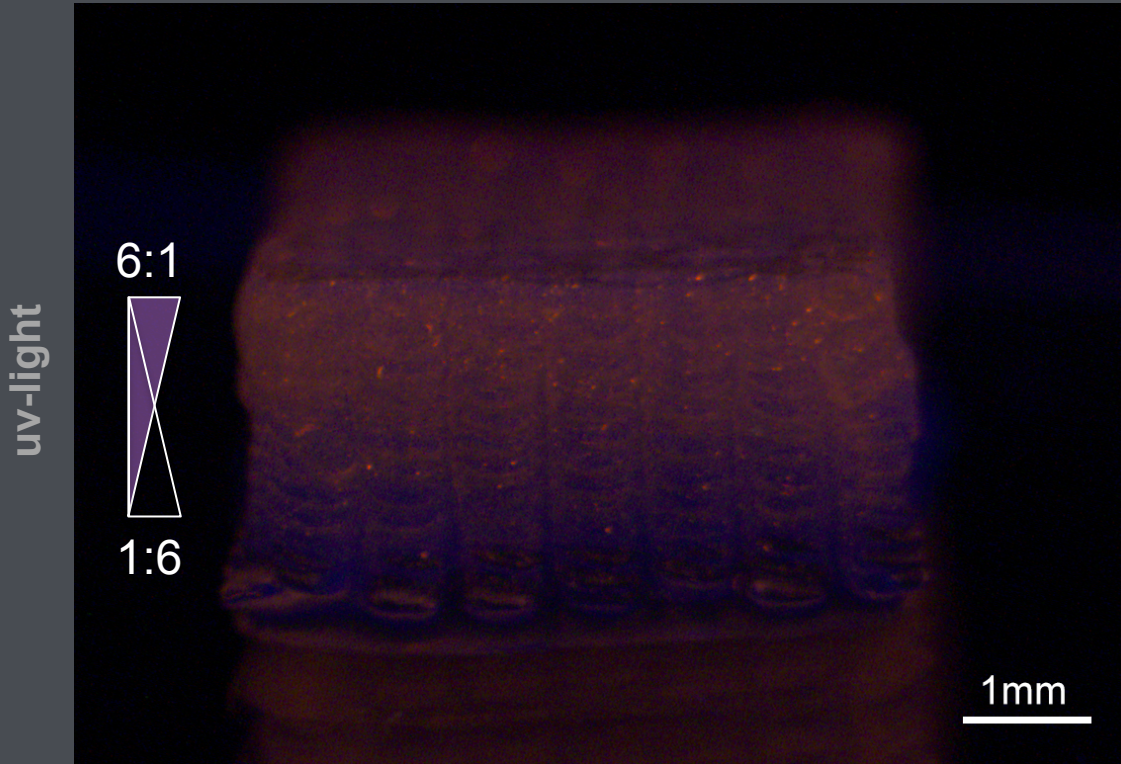
Microfluidic mixing

- Changing composition of the bioink *on-the-fly*



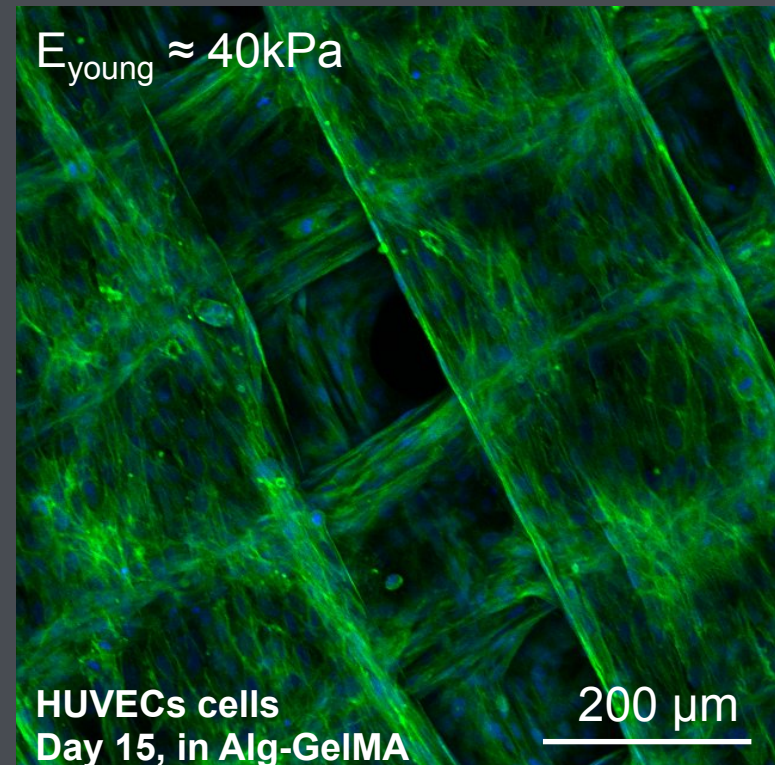
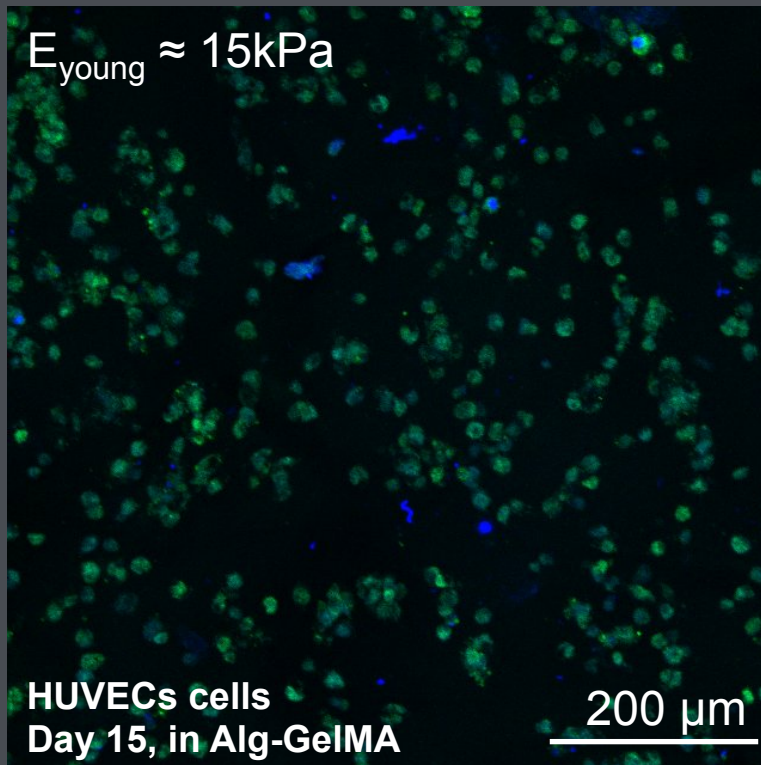
Microfluidic mixing

- Changing composition of the bioink *on-the-fly*



Microfluidic mixing

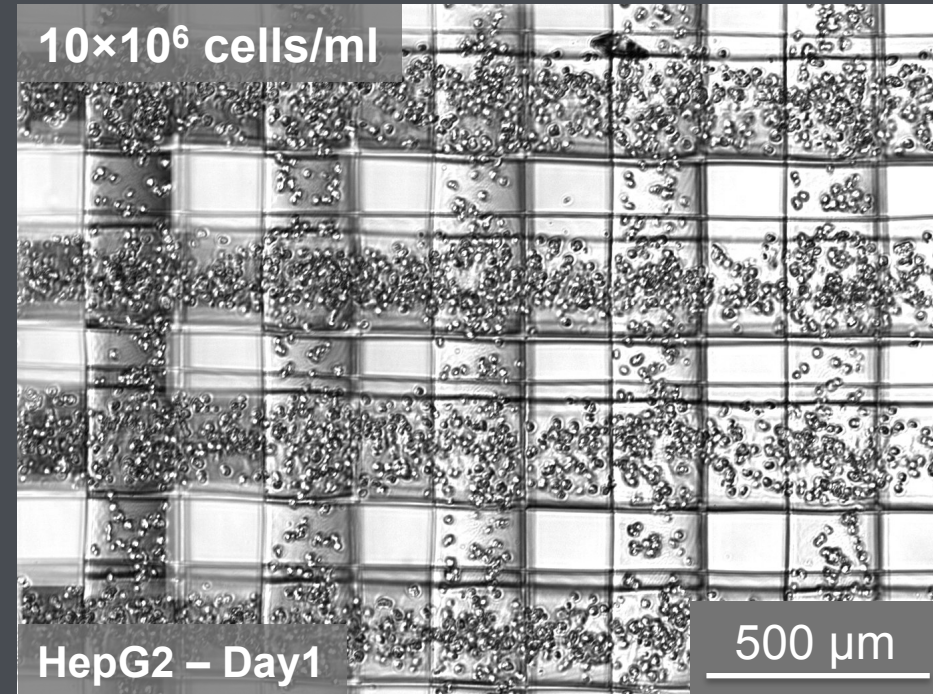
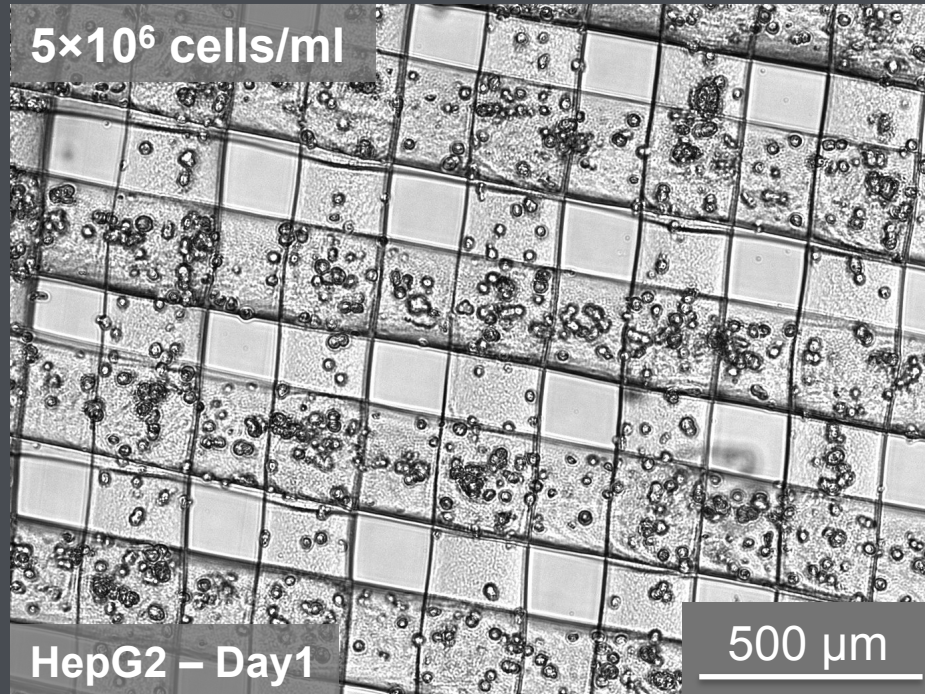
- Changing composition of the bioink:
 - components of the extracellular matrix



Adv. Mat., 2016, 28, 677

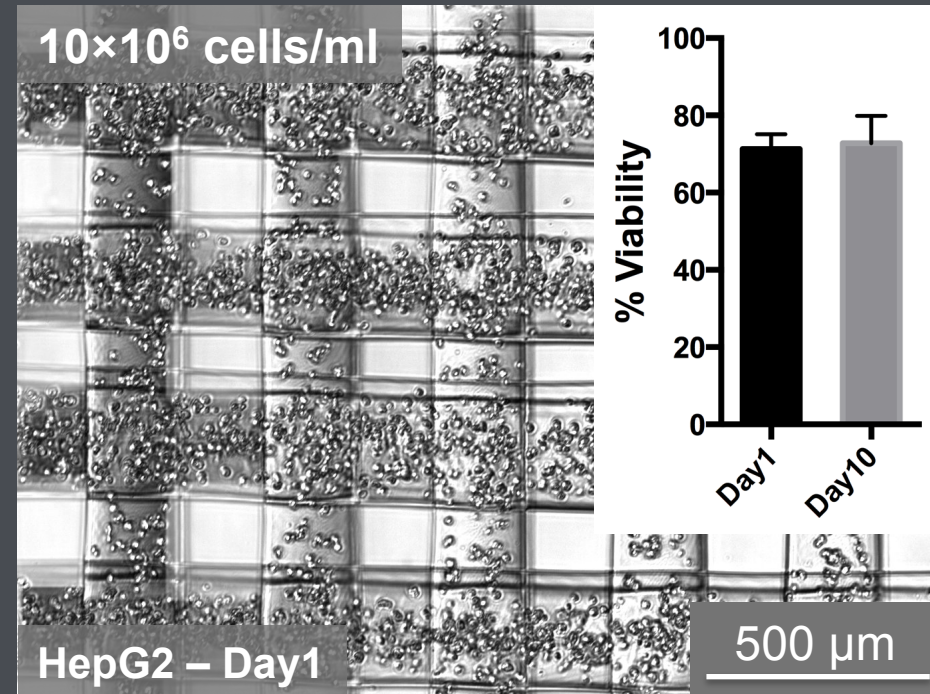
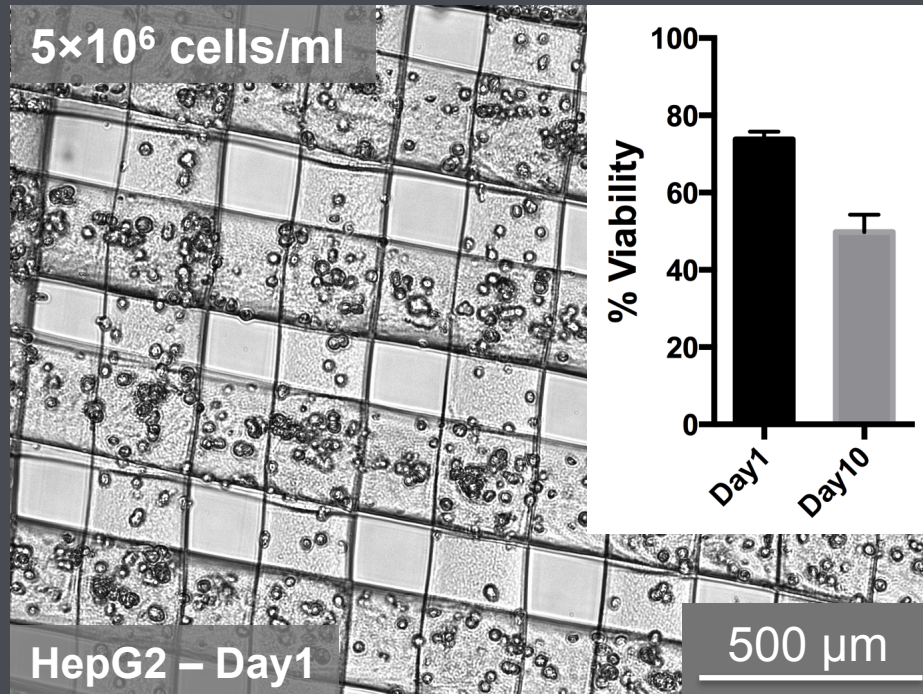
Microfluidic mixing

- Changing composition of the bioink:
 - cell density



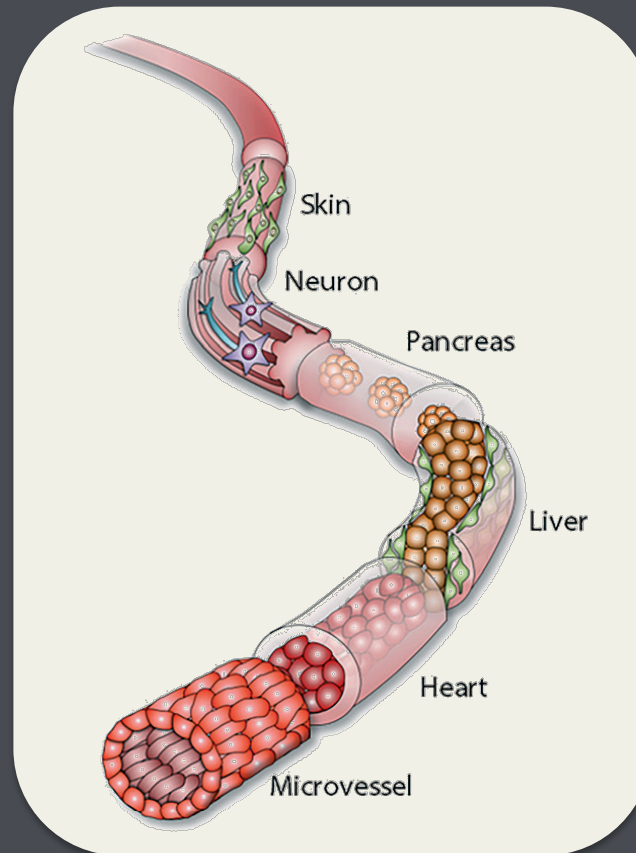
Microfluidic mixing

- Changing composition of the bioink:
 - cell density



Microfluidic patterning

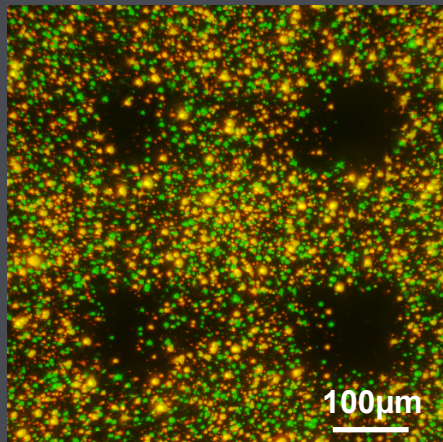
- Creating heterogeneous structures (cells ~ ECM)



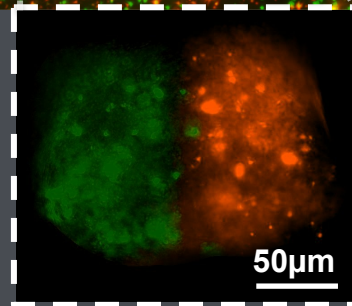
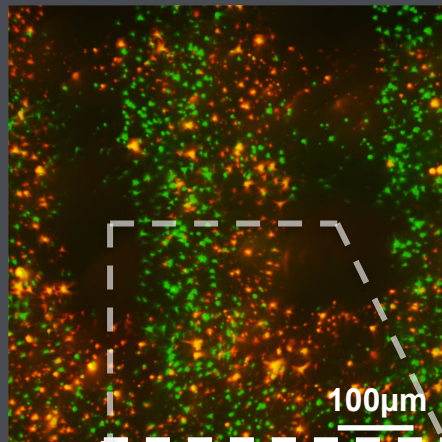
Microfluidic patterning

- Creating heterogeneous structures (cells ~ ECM) *in the same fiber*

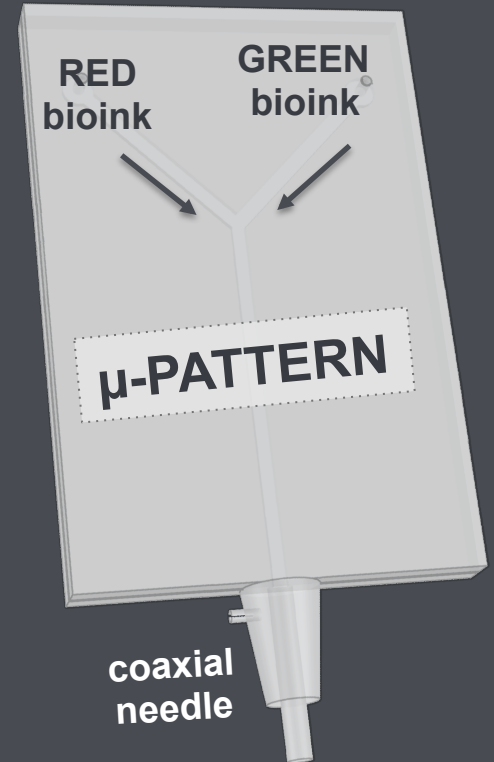
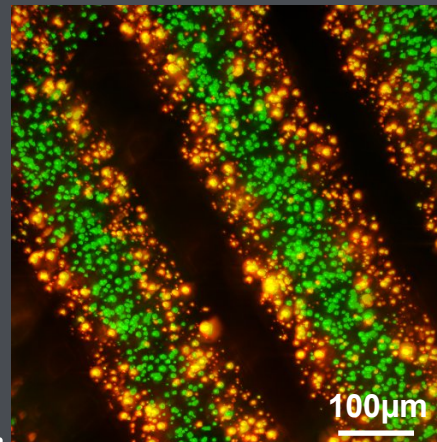
mixed



aligned in 2 parallel zones

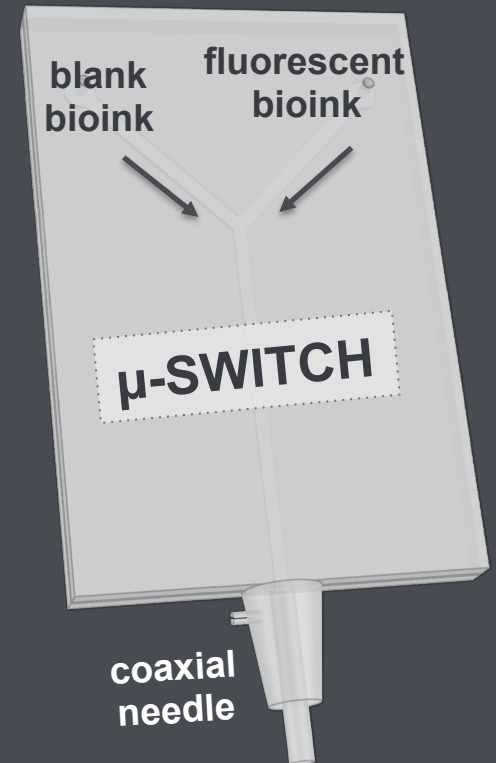
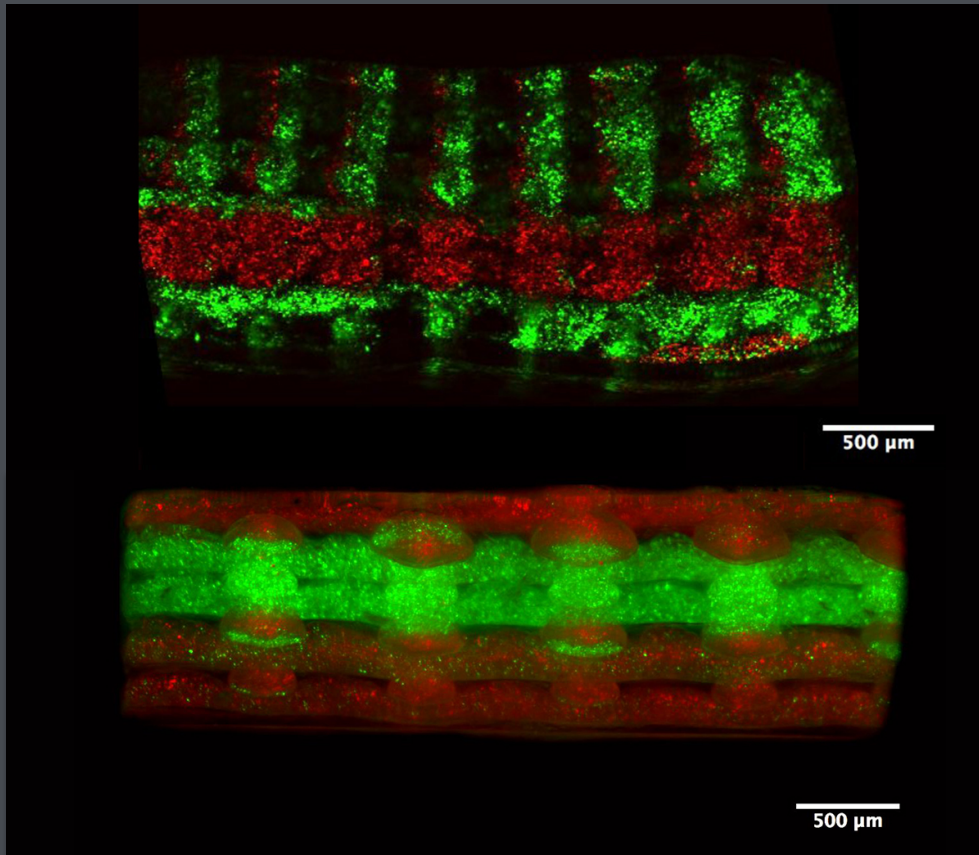


aligned in 3 parallel zones



Microfluidic switching

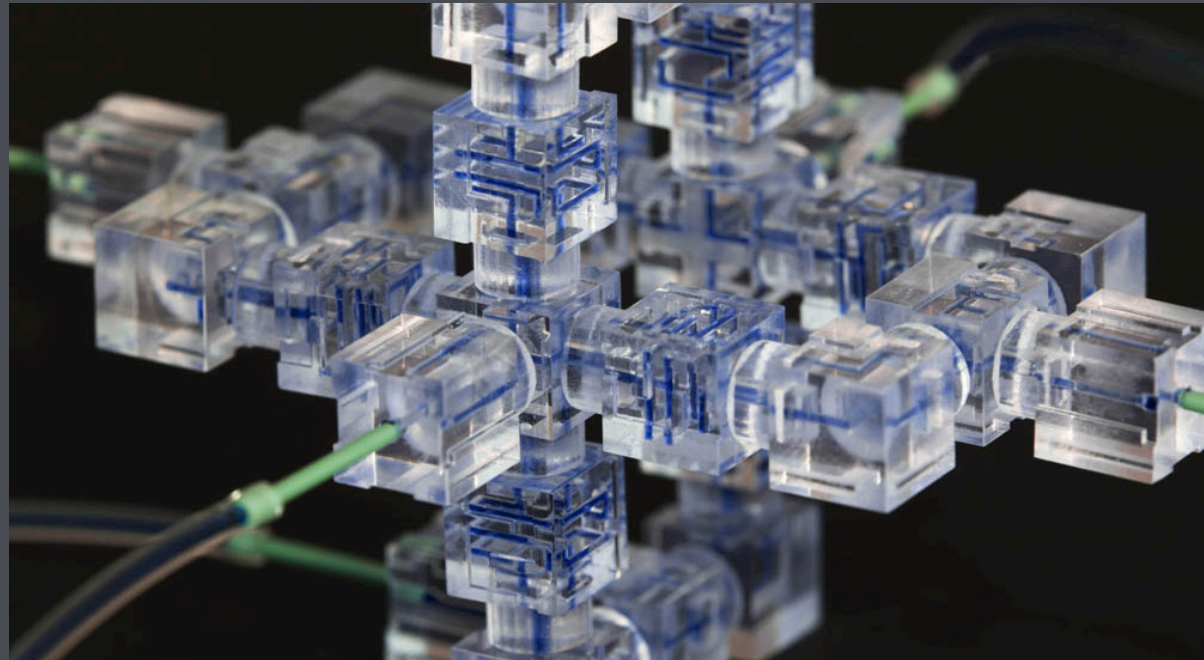
- Creating heterogeneous structures (cells ~ ECM) *in the same construct*



Adv. Mat, 2016, 28, 677

Future perspectives... modular approach

**3D-PRINTED
LEGO μ FLUIDIC
COMPONENTS
associated with
3D deposition**



PNAS, 2014, 111, 42,15013

**CUSTOMIZE THE 3D-BIOPRINTER
DEPENDING ON THE SPECIFIC APPLICATION**

thanks to:

iit

G. Ruocco
M. Levrero
L. Belloni
L. Sarra
E. Toto

Harvard**M.S.**

A. Khademhosseini
S.R. Shyn
A. Arneri
V. Manoharan

Sapienza

M. Dentini
A. Barbetta
S. Screpanti

Nano Rome, 20-23 September
2016 Innovation
Conference & Exhibition

