

2D-3D Bioprinting for medical microdevices

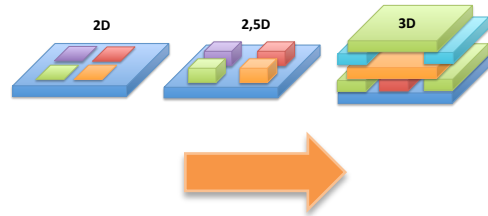
Alive Strategic Axis

Laurent Malaquin

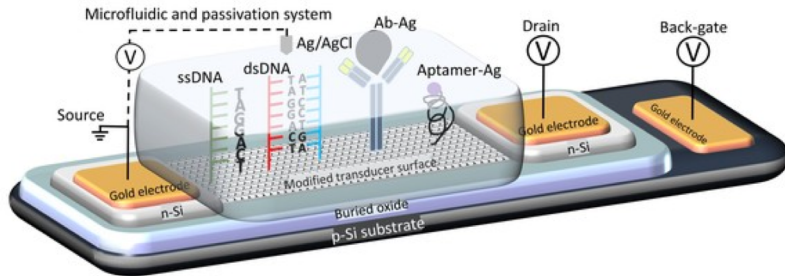
What does bioprinting means ?

> Integrating biological functions into microdevices

- Biomolecules
- Biomaterials
- Living cells
- « Vascular » structures
- Mechanics, topology

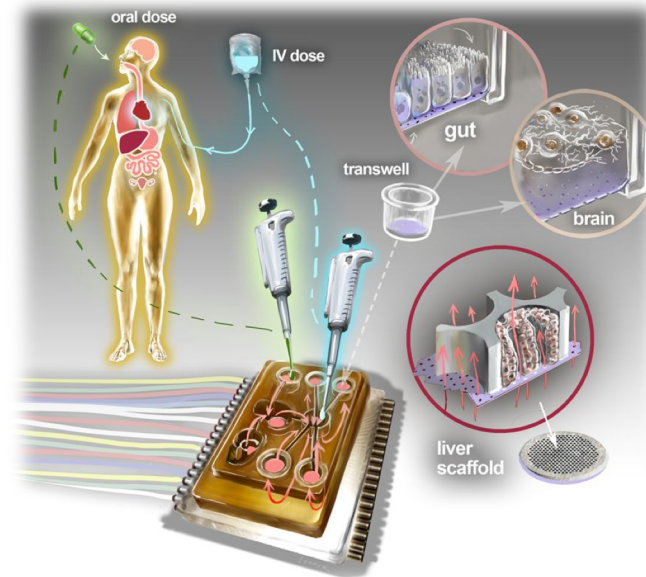


Biomarker Detection
Biomarker Analysis
Cell capture and sorting
Cell biology



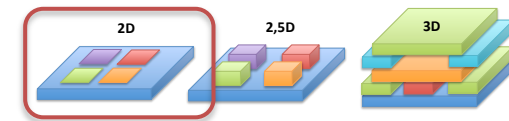
Biosensors

(Subash et al. Biosens. Bioelec. 2017)



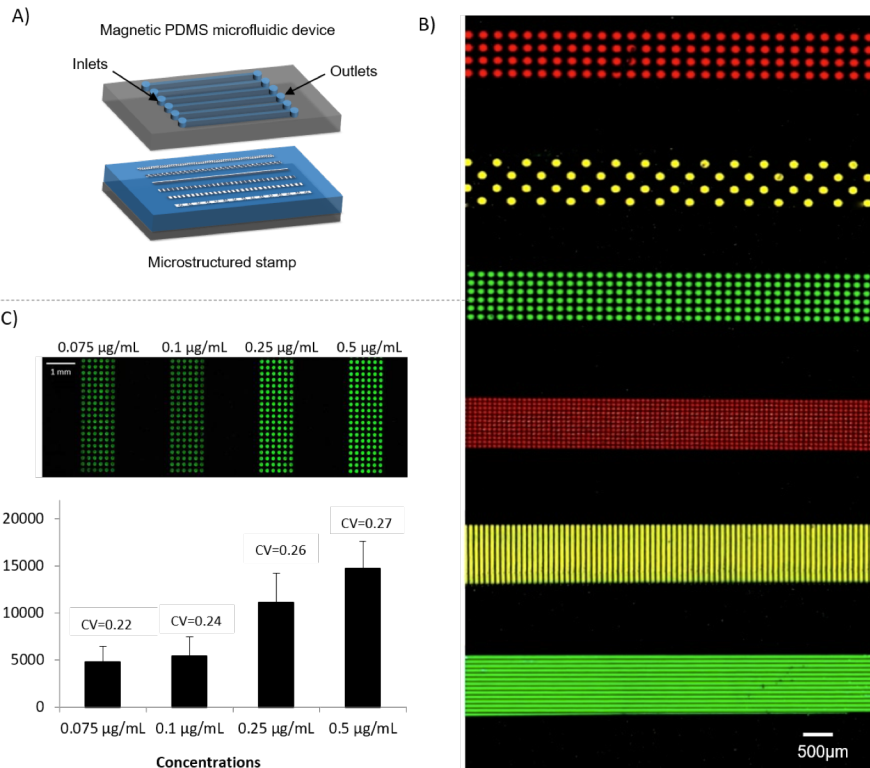
Microphysiological systems

(Edington et al., Nature 2018)



> Microcontact printing (μ CP)

Multiplexed and automated protein deposition on flat surfaces for bioassays



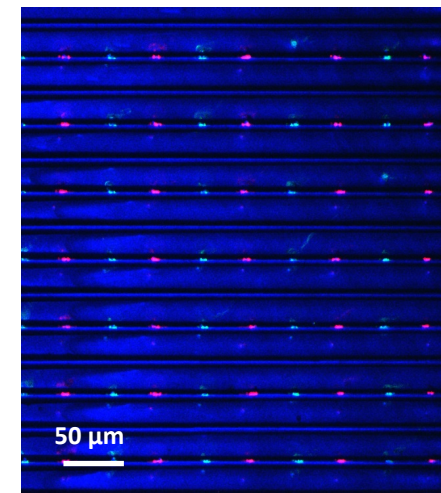
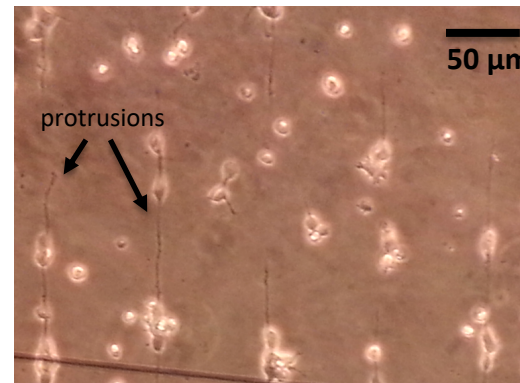
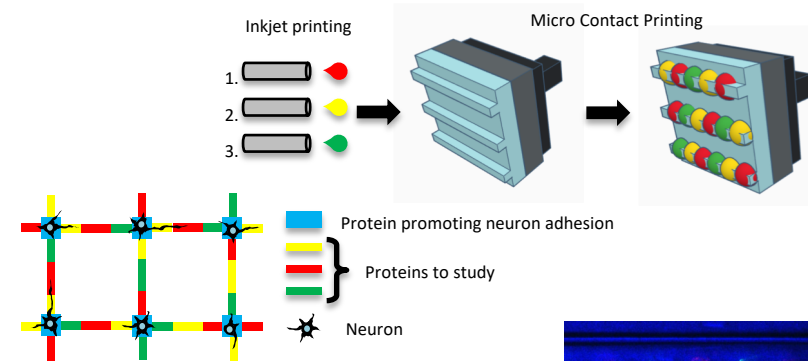
J.Foncy & A.Esteve et al, Plos One 2018

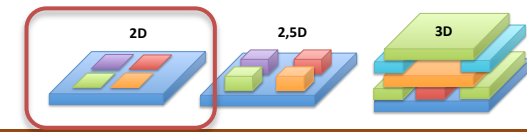
J.Fredonnet et al, Microarrays 2016

A. Esteve, MEE 2018

> Inkjet technology

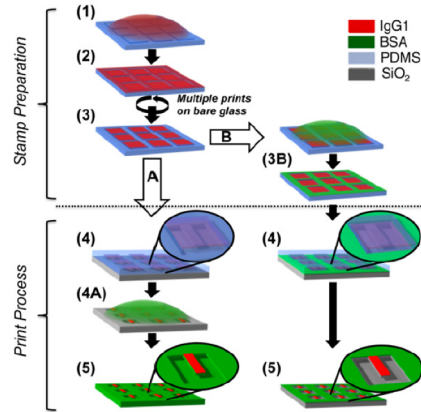
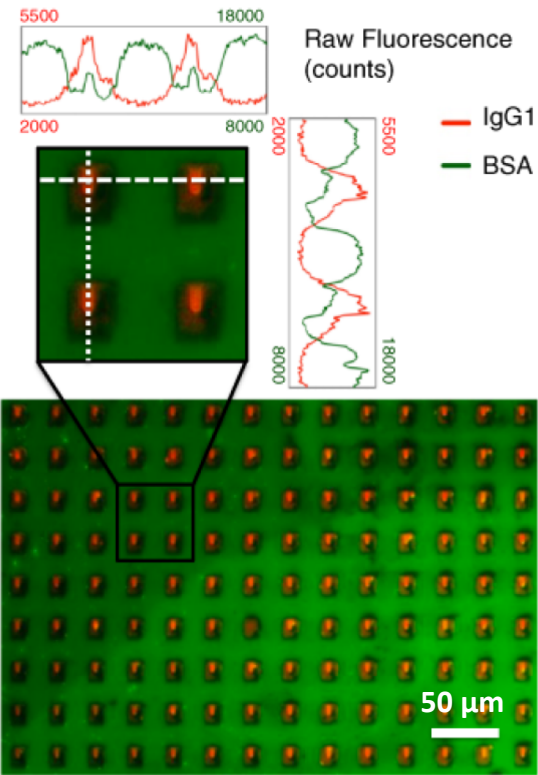
Coupling inkjet with μ CP to creating protein patterns for directed neuron growth





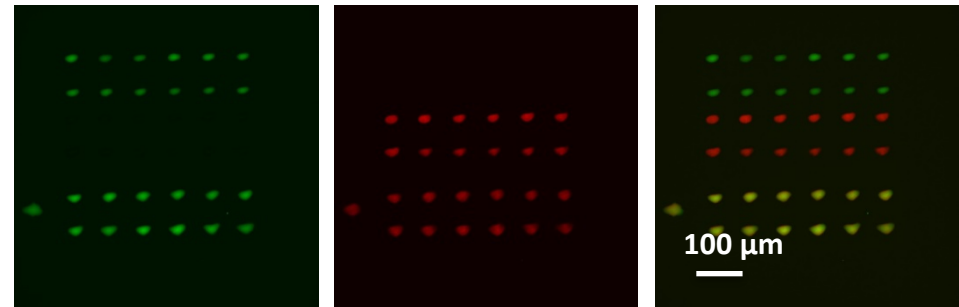
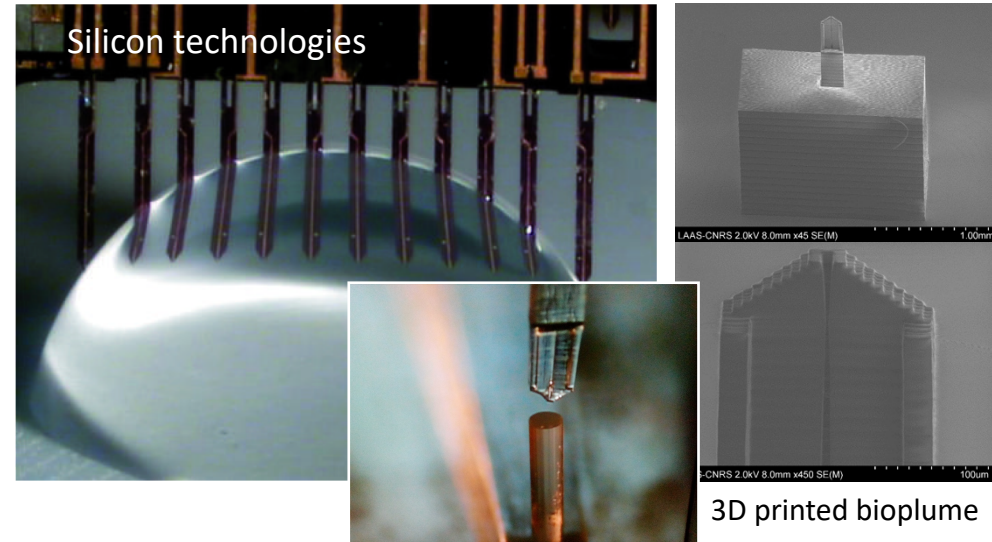
> Tailoring sensor surface

Micro contact printing of antibodies on vibrating micro/nanocantilevers



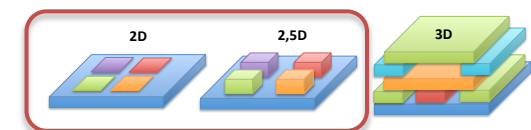
> Bioplume technology

Multiplexed DNA/protein deposition on surfaces and microdevices (optical fibers)



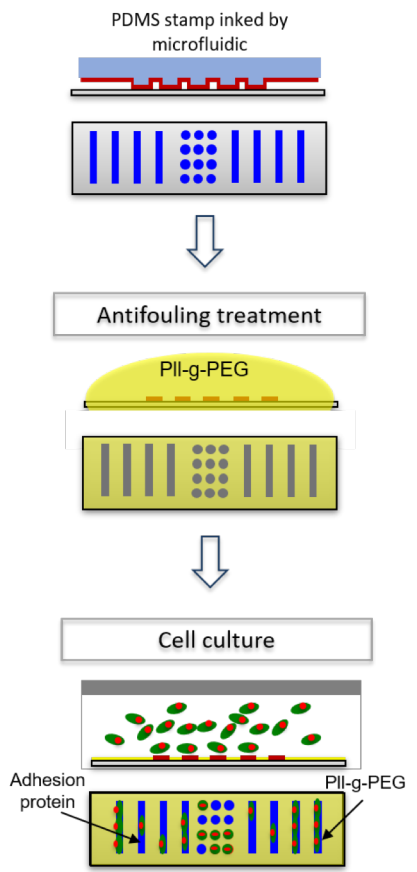
Salomon et al, *Nanotechnology*, 2012

T. Lechlé et al. *Journal of MEMs*, 2008 / *ANR Moly* (2016-2019)

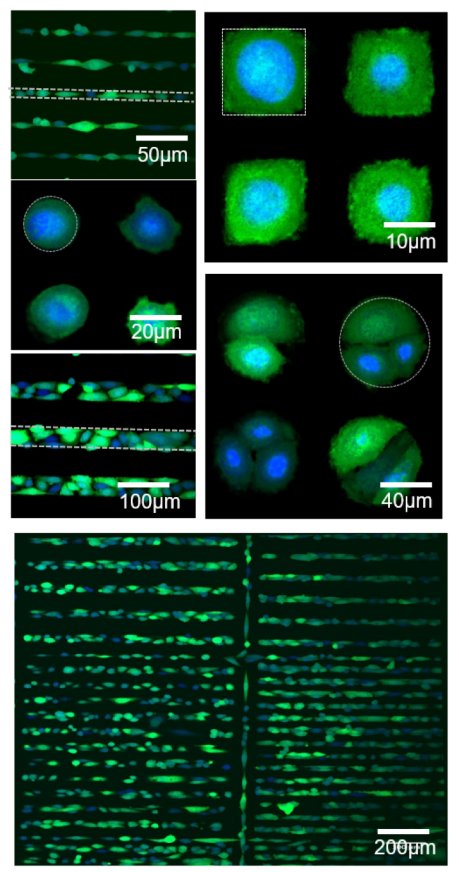


> Controlling cell shape, position, growth

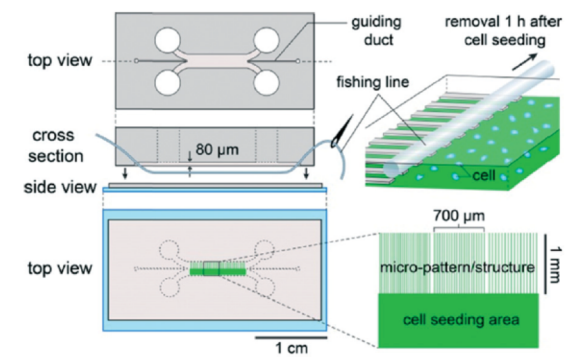
A) Surface patterning for cell culture



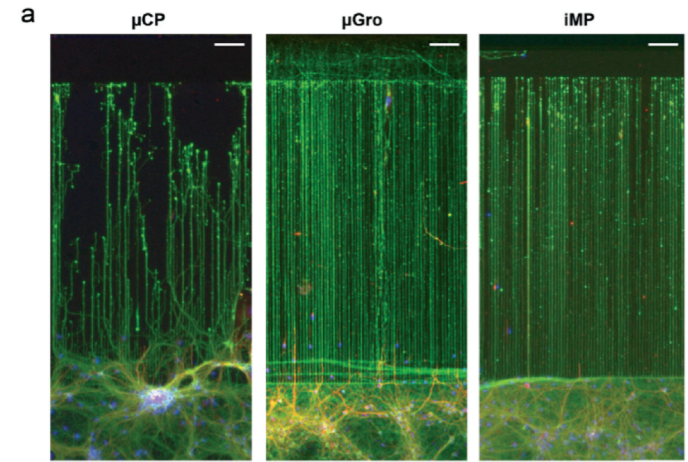
B) Confocal Images



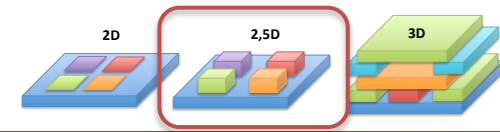
Foncy, Esteve et al, Plos One 2018



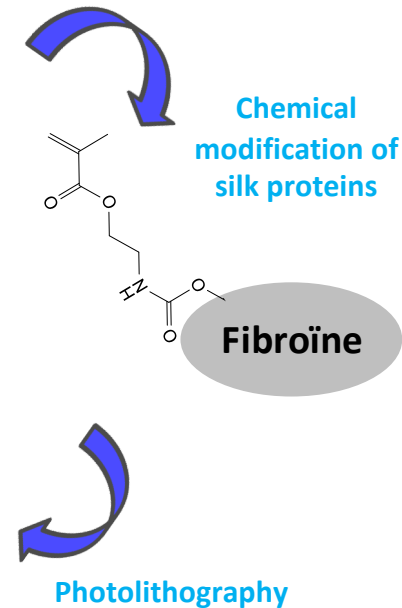
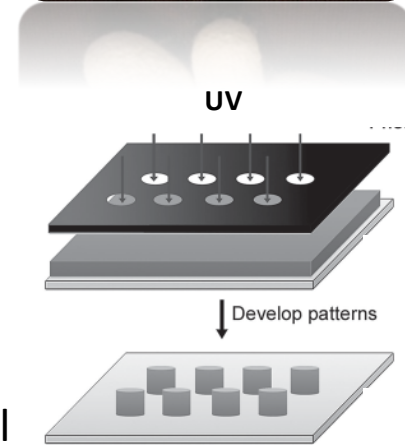
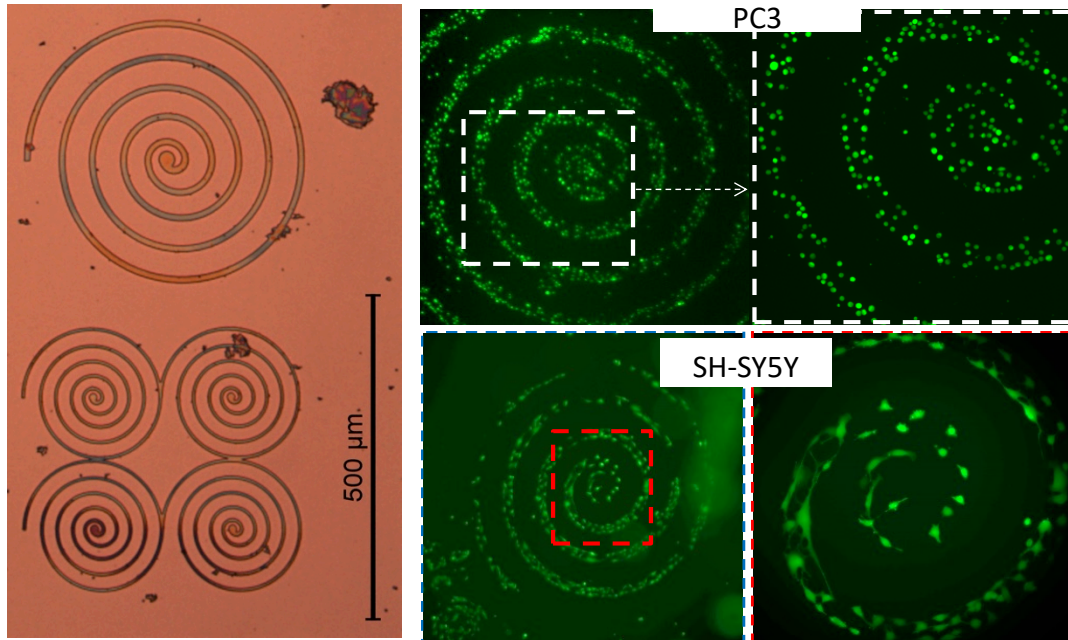
Microcontact printing and molding for cell capture and Neuron Guidance on 2D / 2.5D protein patterns



Yamada, A. et al. Lab Chip 16, 2059–2068 (2016).

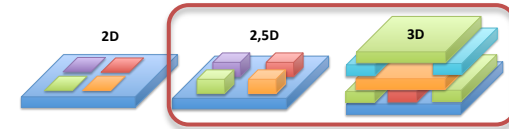


> Developing photosensitive biomaterials

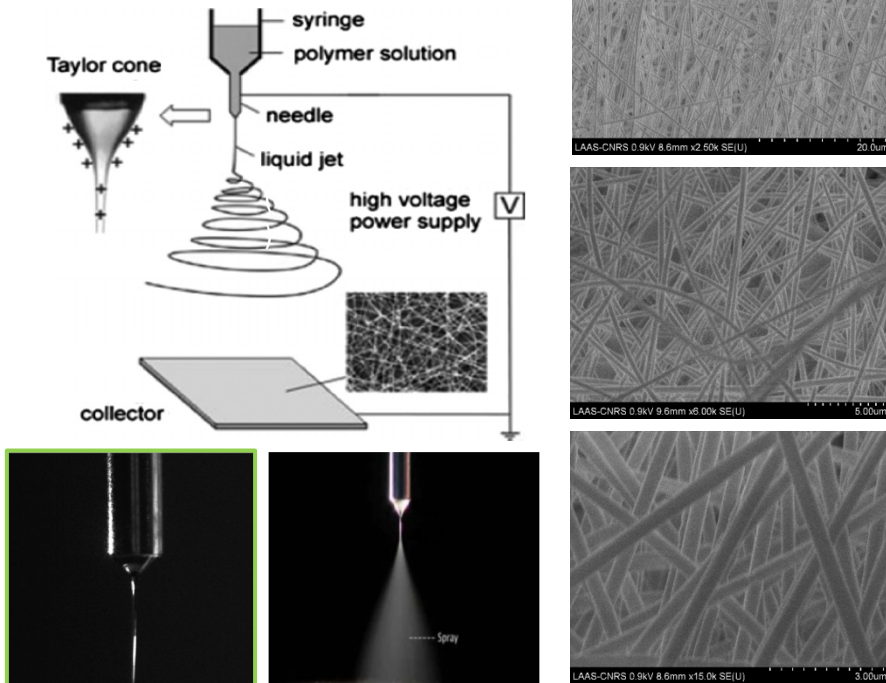


Development of photosensitive silk protein based material for cell culture in controlled stiffness conditions

Maziz et al. Submitted to Biofabrication, 2018



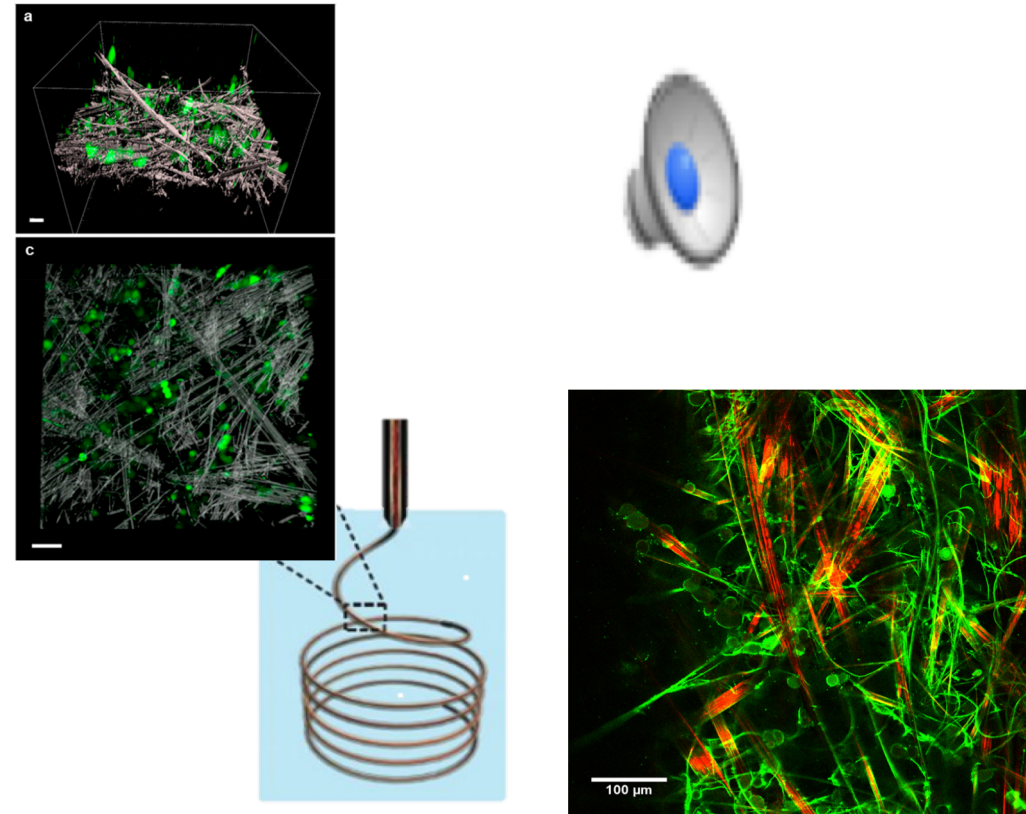
> Electrospinning



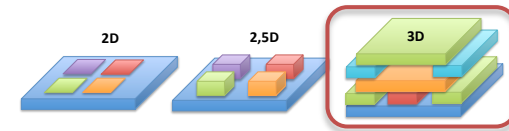
Fabrication of silk/PEG fibers or droplets with diameters ranging from 150 nm up to 4 μm

A. Maziz, C. Bergaud
A. Chalard et al, ACS Applied Materials and Interfaces, 2018

> Wet spinning (solvent exchange)

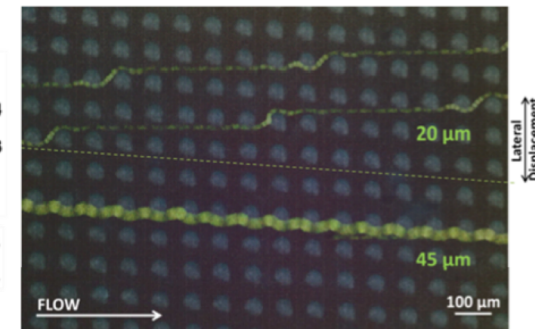
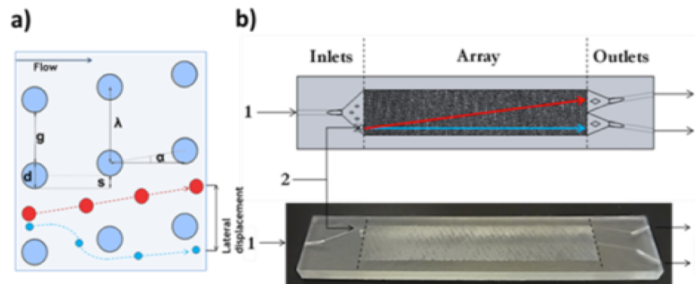
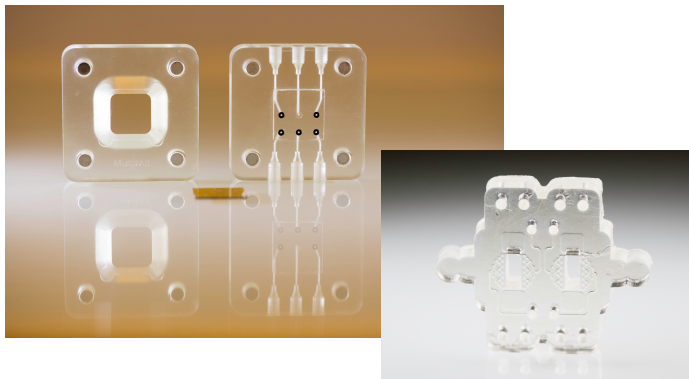


Fabrication of macromolecular Alkyl-galactonamides hydrogels fibers (100-500 μm) for neuron cell culture.

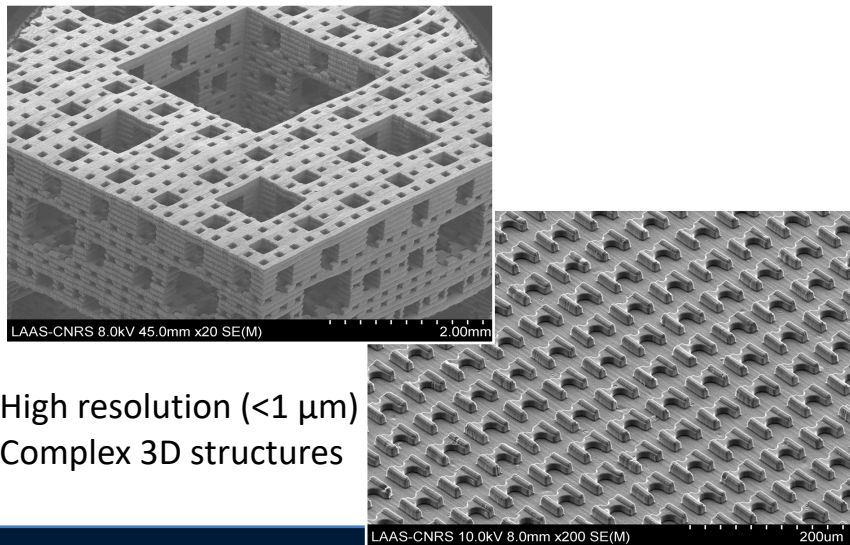


> 3D printed microfluidic devices for biomarker analysis and cell sorting

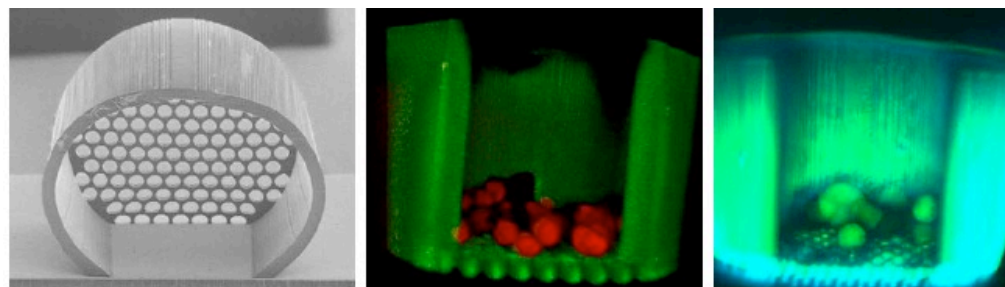
- Sorting of cells and particles (Deterministic lateral displacement)



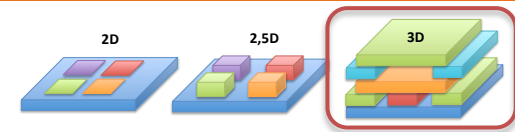
- Sorting CTC with 3D printed filters



High resolution (<1 μm)
Complex 3D structures

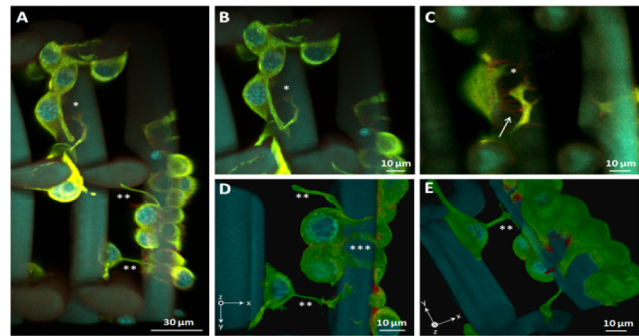
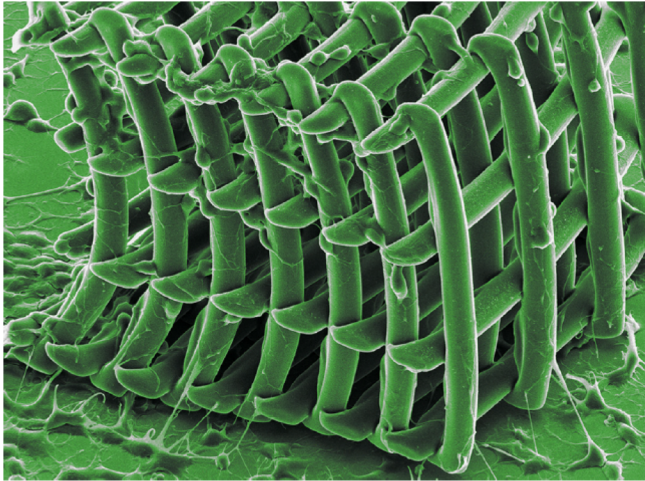


F.Mézières et al, JASA 2016
A.Accardo Add. Manufact. , 2018
A. Cerf, A. Esteve, K. Jimenez



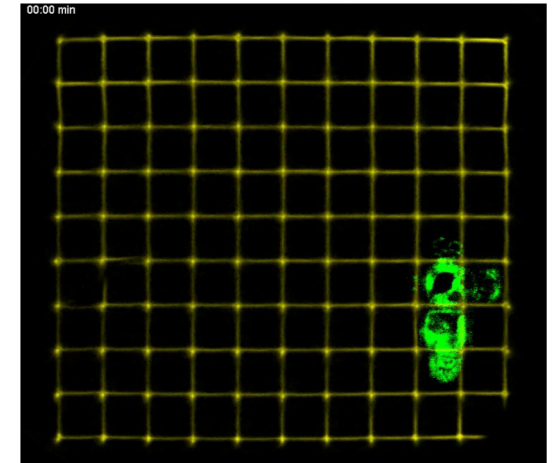
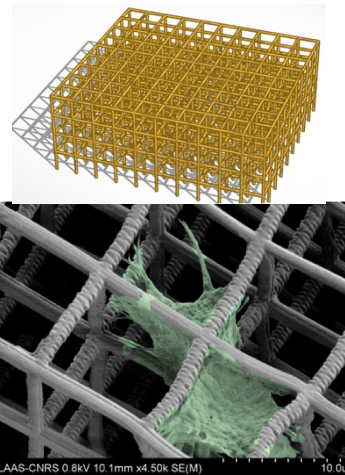
> Controlling the topology and mechanical properties of cell microenvironment

- 3D scaffolds for neuron cell growth



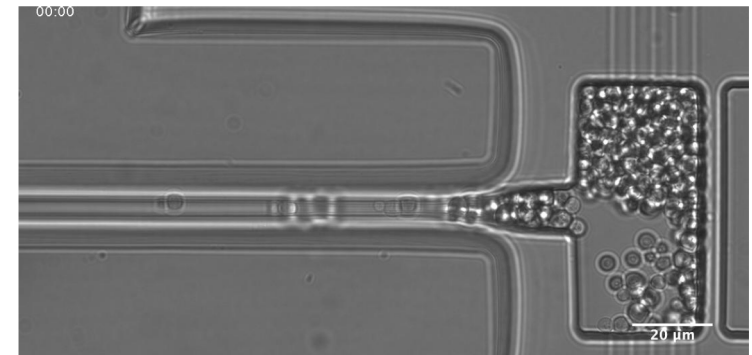
Accardo et al, *Materials Today*, 2018
 Acardo et al *Small*, 2017 / E. Desvignes
 M. Delarue et al, *Nature Physics*, 2017

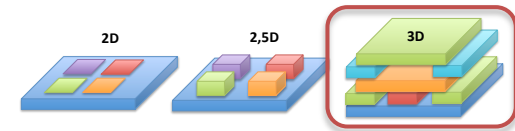
- 3D models for cell migration studies



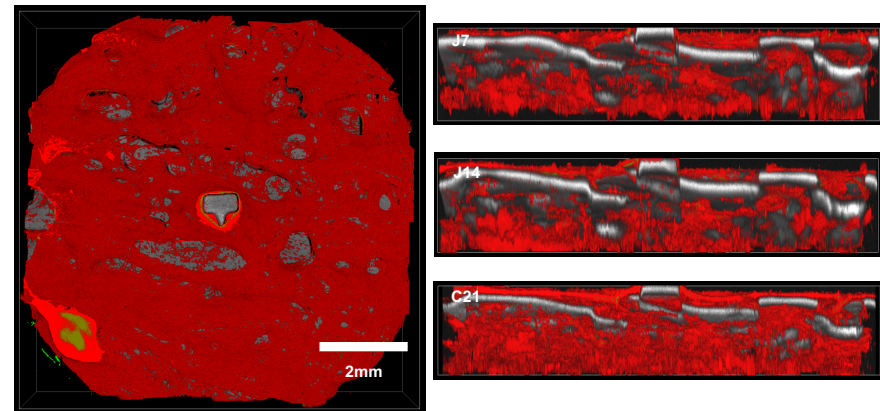
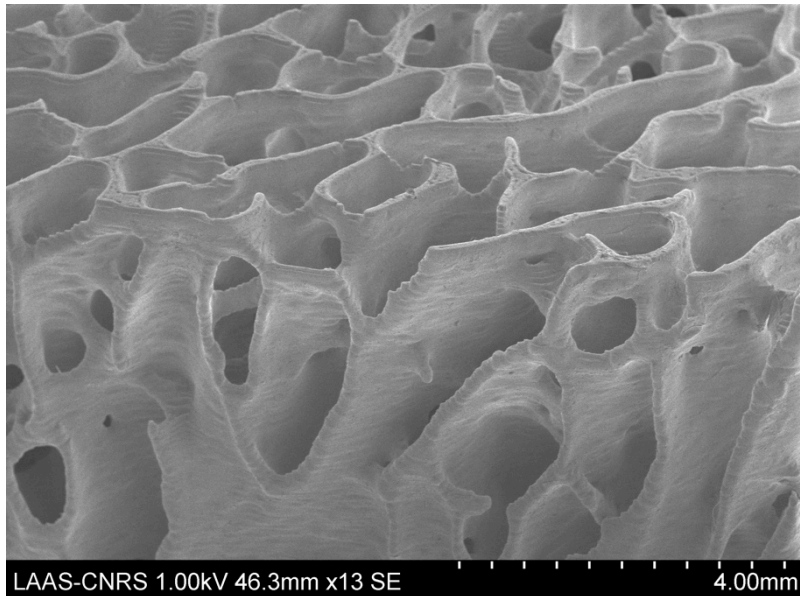
- Mechanical environment

Controlling pressure,
 supply, confinement
 (yeasts)





> Controlling topological environment for cell culture



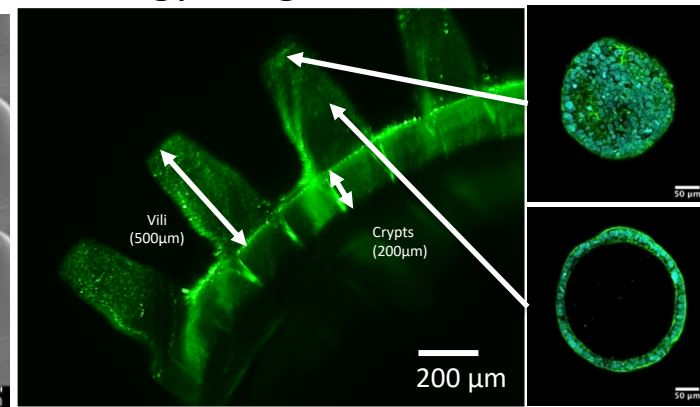
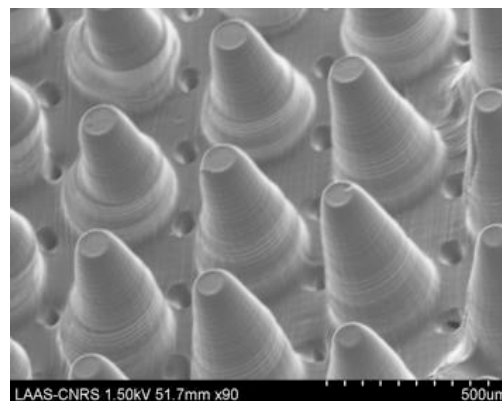
Growth and differentiation of Mesenchymal Stem Cell on trabecular bone scaffolds

- 3D models for oncology / regenerative medicine

- Controlling stiffness, topology, material chemistry of the microenvironnement
- Controlling cell position by printing ?

F. Mézières et al, JASA 2016

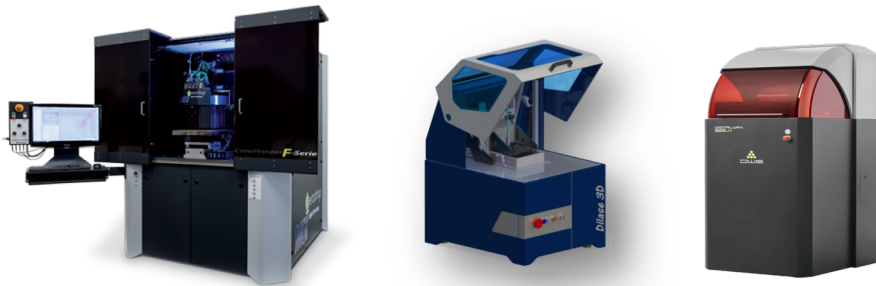
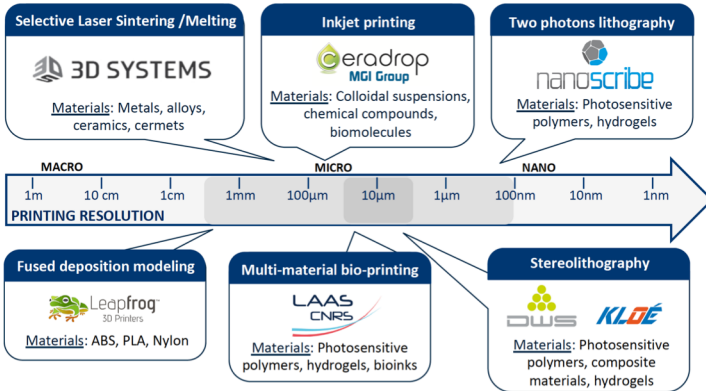
J. Creff, A. Besson



> MultiFAB platform
(LAAS-CNRS / CIRIMAT)



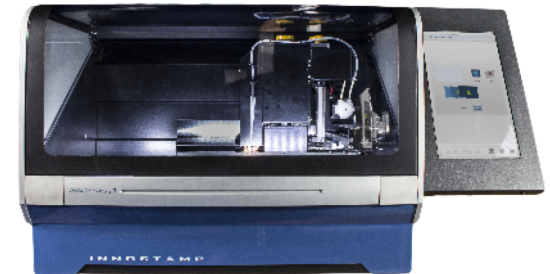
Cutting edge technologies for 3D printing



> Joined Laboratories



Multiplexed microcontact printing with reversible clamping



www.biosoftlab.com

<https://www.laas.fr/projects/MultiFAB/>



PROJET COFINANCÉ PAR LE FONDS EUROPÉEN DE DÉVELOPPEMENT RÉGIONAL

- > *Novel opportunities for microdevices fabrication and integration*

- > *An open field of research for cell biology*
 - *Fundamental studies in cell biology*
 - *Models systems for drug screening and diagnosis*
 - *Regenerative medicine*

- > **Challenges :**
 - technological development
 - material development