3D Organoid-on-a-Chip in a Microphysiological System

Novel Technologies to Improve Predictivity of Non-Clinical Studies and Replace, Reduce, and Refine Reliance on Animal Testing

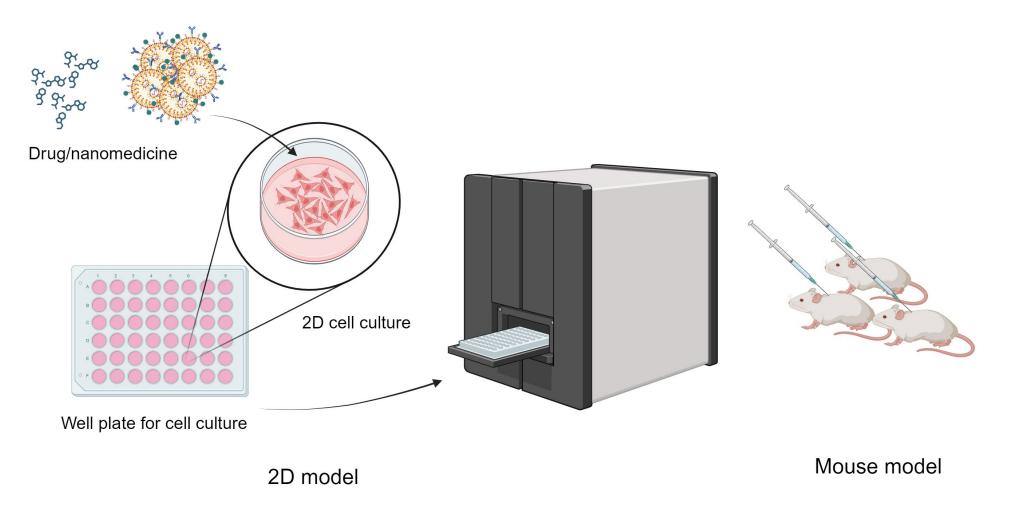
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DeLouise Lab

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Two-Dimensional (2D) Monolayer Cultures and *In Vivo* Animal Models cannot Adequately Replicate the Human Tissues

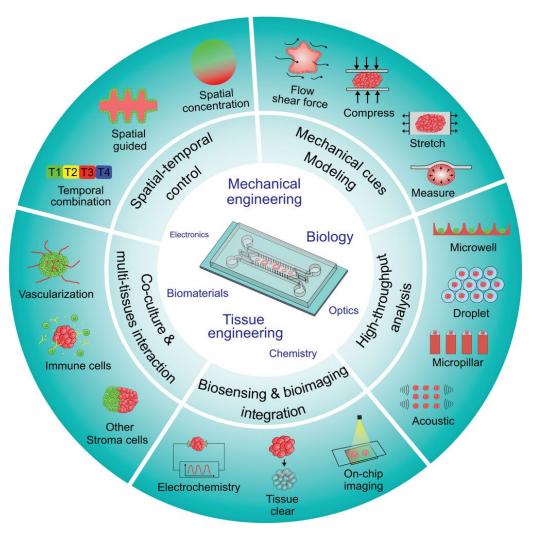




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Organoid-on-a-Chip can Model *In Vivo* Tissue Microenvironment

- Spatial-temporal control
- Ability of perfusion culture
- Mechanical cues modeling
- High-throughput analysis with more reliable drug screening results
- Multi-tissues or organs interaction
- Integration of biosensing and bioimaging
- Reduction use of animal models
- Closer features to in vivo natural systems



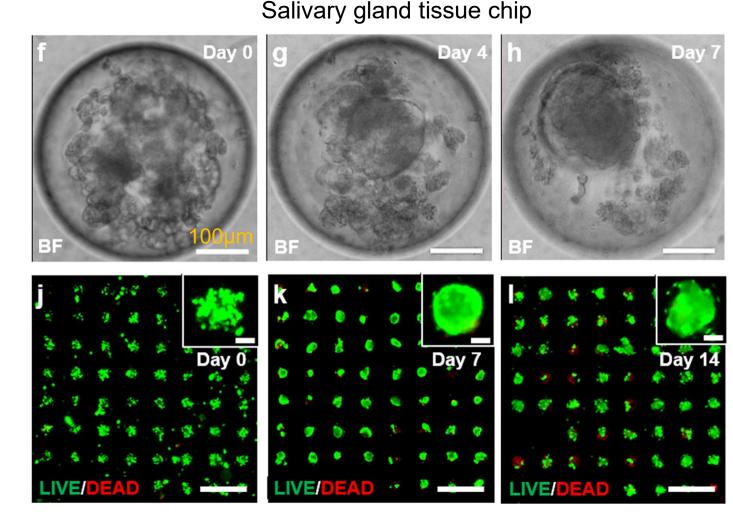
Tissue Chip Models Support Cell Viability in 3D Tissue Microenvironment and can Reduce the Use of Animal Models

~ 50 MBs/chip

40 experiments with n=50 from one mouse

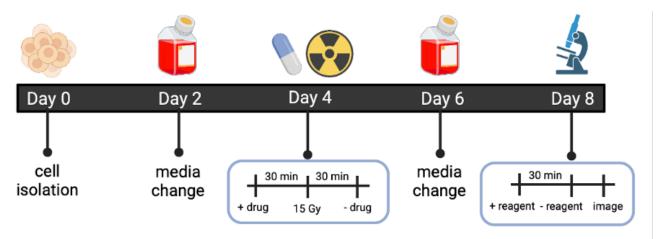
Microbubble (MB)

Salivary gland cell clusters

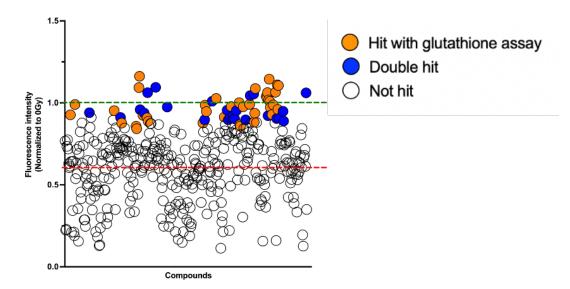


Song, Y et al. Communications Biology, 2021

Tissue Chips Support High-Throughput Screening for Drug Discovery



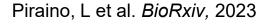
Defined Biomarkers for Radiosensitivity: Glutathione, Senescence





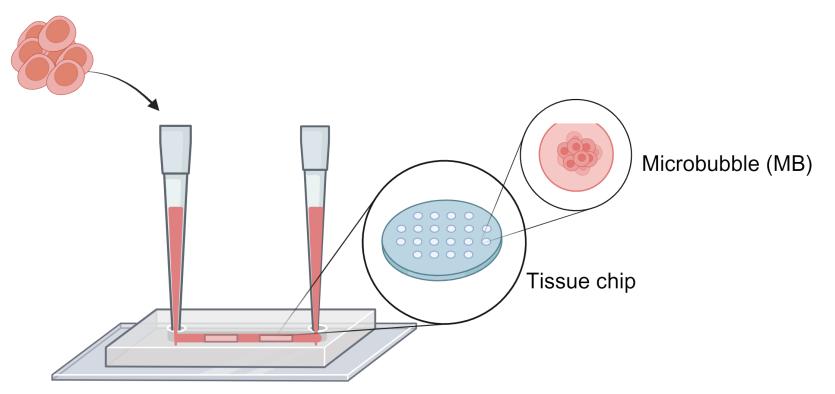
Results were confirmed by in vivo mouse model testing

SelleckChem library screening	
Drugs screened with glutathione assay	438
Hits with glutathione assay	62
Hit rate (percentage)	14.2%
Hits with Glutathione and Senescence assays	25
Double hit rate (percentage)	5.7%



Tissue Chips Under Flow in Microfluidic Devices can Provide Individualized Therapies and Precision Medicine

Induced pluripotent stem cells (iPSCs) or primary tumor cells



Microphysiological organoid-on-a-chip System

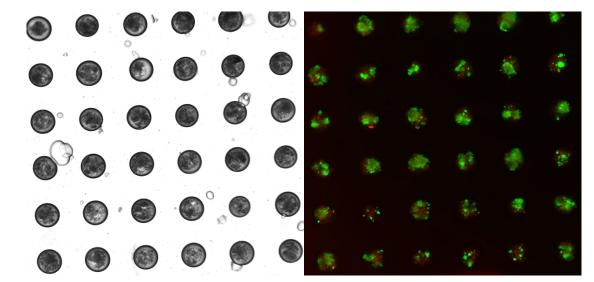


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Salivary Gland Tissue Culture in a Microfluidic Device

Salivary gland cell clusters





Live/Dead Assay on Day 7



Organoid-on-a-chip system in a microfluidic device

Summary and Conclusions

- Tissue chips support cell viability in 3D tissue microenvironment and promote cell growth in the form of 3D spheroid/organoid model
- Tissue chips enable high-throughput drug screening with defined biomarkers
- Organoid-on-a-chip microphysiological system can be used for personal medicine applications
- The models can Replace, Reduce, and Refine Reliance on Animal Testing



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