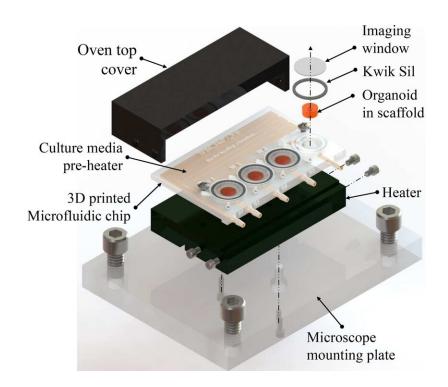


Tiny brains grown in 3D-printed bioreactor

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Scientists from MIT and the Indian Institute of Technology Madras have grown small amounts of self-organizing brain tissue, known as organoids, in a tiny 3-D-printed system that allows observation while they grow and develop. The work is reported in *Biomicrofluidics*.

Current technology for real-time observation of growing organoids



involves the use of commercial culture dishes with many wells in a glassbottomed plate placed under a microscope. The plates are costly and only compatible with specific microscopes. They do not allow for the flow or replenishment of a nutrient medium to the growing tissue.

Recent advances have used a technique known as microfluidics, where a nutrient medium is delivered through small tubes connected to a tiny platform or chip. These <u>microfluidic devices</u> are, however, expensive and challenging to manufacture.

The current advance uses 3-D printing to create a reusable and easily adjustable platform that costs only about \$5 per unit to fabricate. The design includes imaging wells for the growing organoids and microfluidic channels to provide a nutrient medium and preheating that supports tissue growth.

A biocompatible type of resin used in dental surgery was used for the 3-D-printed <u>device</u>. The printed chip was cured by exposing it to UV light, then sterilized before <u>live cells</u> were placed in the wells. After sealing the top of the wells with a glass slide, the nutrient medium and drugs for use in the study were added through small inlet ports.

"Our design costs are significantly lower than traditional petri dish- or spin-bioreactor-based <u>organoid</u> culture products," said author Ikram Khan. "In addition, the chip can be washed with distilled water, dried, and autoclaved and is, therefore, reusable."

The investigators tested their device with organoids derived from <u>human</u> <u>cells</u>. They observed the growing brain organoids with a microscope and were able to successfully follow their growth and development for seven days. The small bit of <u>brain tissue</u> developed a cavity or ventricle surrounded by a self-organizing structure that resembles a developing neocortex.



The percentage of cells in the core of the organoid that died during this one-week period was smaller in the 3-D-printed device than in regular culture conditions. The investigators believe that their cell design protects the tiny growing brain.

Khan said, "One advantage offered by our microfluidic device is that it allows constant perfusion of the culture chamber, which more closely mimics a physiological <u>tissue</u> perfusion than conventional <u>culture</u>, and thus reduces cell death at the organoid core."

The investigators hope to increase the capacity of their device by scaling up the number of available wells. Other improvements will allow for additional instruments to be integrated into the design.

More information: "A low-cost 3D printed microfluidic bioreactor and imaging chamber for live-organoid imaging" *Biomicrofluidics* (2021). <u>aip.scitation.org/doi/10.1063/5.0041027</u>

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