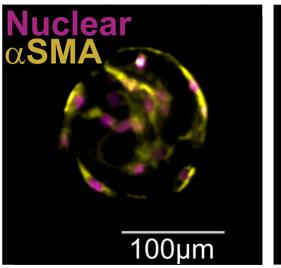
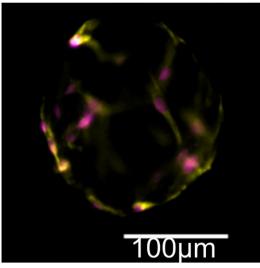


## Microtissue system allows study of deadly lung disease

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Patient Encapsulated Cells
IPF Patient Control Patient





The Wood Lab's high-throughput 3D culture workflow allows encapsulation of patient-sourced cells and the study of their fibrotic function. Credit: Katherine Cummins

Idiopathic pulmonary fibrosis (IPF) is a deadly and rapidly progressing disease with no cure.

The disease involves abnormal interactions between <u>lung cells</u>, including



fibroblasts, and their surrounding environment. Because of this, standard 2-D cell culture models used for <u>drug screening</u> tend to perform poorly when predicting response to potential therapies.

Amid the COVID-19 pandemic and rising air pollution levels, incidence of IPF is anticipated to rise, urgently increasing the need for strong model systems.

In *APL Bioengineering*, researchers from the University of Minnesota-Twin Cities and Mayo Clinic in Rochester, Minnesota, describe a 3-D cell culturing platform that allows study of lung fibroblasts and their microenvironment. The platform enables measurement of cell behaviors and microenvironment changes involved in the disease progression of IPF, and the platform's size and simplicity make it suitable for use in high-throughput drug screening protocols.

"IPF is a horrible disease that drastically impacts a patient's life and eventually causes them to die from lack of oxygen," said co-author Katherine Cummins. "It's really important to have lab tools and models that create and control the microenvironment in which the <u>cells</u> sit, because this may be key to preclinical identification of possible treatments."

Unlike rodent IPF models that do not mimic progressive disease and other <u>cell culture systems</u> that lack the surrounding microenvironment, their microtissue platform allows study of fibroblasts within an extracellular matrix (ECM). Changes in the ECM are a hallmark of IPF, so the system allows more relevant functional outputs. Moreover, its simplicity and tunability make it easy to use.

"Many organoid and lab-on-a-chip platforms can be hard to use," said coauthor David Wood. "What's exciting is that this system is very easy to use. We've already disseminated it to two other labs who are using it



completely independent of us."

Validation of the system's functioning focused primarily on ECM remodeling (i.e., cell-driven changes to the microenvironment) and cell contractility, which increases in activated, diseased fibroblasts.

Multiple tests for each of these two functions demonstrated the system robustly quantifies key aspects of fibrosis. These results were reproduced using patient-donated cells as well, indicating the system could be used for personalized medicine.

Moreover, the system's versatility allows it to be used with different cell types and other matrix components, so it could be adapted for use in the study of other diseases where cell-microenvironment interactions contribute to disease. The research team has already used the system to study liver toxicity and anticipates it could be used across multiple solid organ systems, including in the study of cancer progression and metastasis.

**More information:** "A scalable 3-D tissue culture pipeline to enable functional therapeutic screening for pulmonary fibrosis" *APL Bioengineering*, 2021, aip.scitation.org/doi/full/10.1063/5.0054967

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