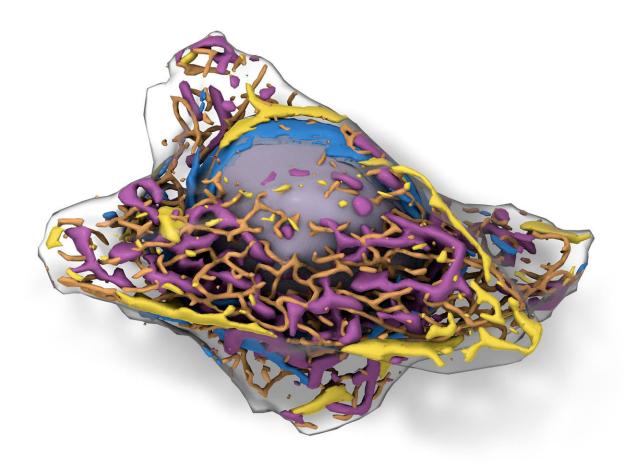


Allen Integrated Cell released online

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Multiple structures visualized together inside a human cell. Researchers took images of fluorescently labeled cells and applied artificial intelligence. AI computer vision used what it learned from cells *with fluorescent labels to find cellular structures in cells *without fluorescent labels. This label-free model can be used on relatively easy to collect brightfield microscope images. Credit: Allen Institute for Cell Science



The Allen Institute for Cell Science today launched the first predictive and comprehensive, 3D model of a live human cell, the <u>Allen Integrated</u> <u>Cell</u>. By allowing researchers around the world to see many structures inside a living cell together at the same time, the Allen Integrated Cell provides a baseline for understanding cells and studying human disease models.

"This is a new way to see inside living human <u>cells</u>," said Rick Horwitz, Ph.D., Executive Director of the Allen Institute for Cell Science. "It's like seeing the whole cell for the first time. In the future, this will impact drug discovery, disease research and how we frame basic studies involving human cells."

"Nearly every biologist has mental models of whole cells that are pieced together over their careers with information from dozens of different types of data," said Graham Johnson, Ph.D., Director of the Animated Cell. "The Allen Integrated Cell provides a new option, where 3-D visualizations of whole <u>living cells</u> link to analysis tools to allow for more direct data-driven exploration and hypothesis generation."

The Allen Integrated Cell summarizes a large collection of live human cells, gene edited by Allen Institute scientists to incorporate fluorescent protein tags. These tags illuminate specific structures inside of cells, such as the nucleus and mitochondria. Scientists took pictures of tens of thousands of these glowing cells and used artificial intelligence to study them.

First, researchers developed a computer algorithm that studied the shape of the plasma membrane, the nucleus and other fluorescently labeled cell structures to learn their spatial relationships. A powerful probabilistic model emerged from this training, that accurately predicts the most probable shape and location of structures in any cell, based solely on the shape of the plasma membrane and the nucleus.



Second, researchers took images of those same fluorescently labeled cells and applied a different machine learning algorithm. This algorithm used what it learned from cells with fluorescent labels to find cellular structures in cells without fluorescent labels. This label-free model can be used on relatively easy to collect brightfield microscope images to visualize the integration of many structures inside of cells, simultaneously and with high precision. Viewed side-by-side, the images generated by the label-free method look nearly identical to the fluorescently labeled photographs of cells.

"Fluorescence microscopy is expensive and toxic to cells; increasingly so when you tag multiple structures," said Molly Maleckar, Ph.D., Director of Modeling. "Our approach allows scientists to view cells and conduct experiments at the reduced cost of brightfield microscopy, with the <u>structure</u>-identifying power of fluorescence microscopy - and without its toxic effects. It's really the best of both worlds."

"Until now, our ability to see what is going on inside of human cells has been very limited," said Michael Elowitz, Ph.D., Professor of Biology, Bioengineering, and Applied Physics at California Institute of Technology. "Previously, we could only see the proteins that we deliberately labeled. But the Allen Integrated Cell is like the ultimate free lunch. We get to sample a 'buffet' of many different proteins and organelles, without having to label anything at all. This opens up a totally new and much more powerful way of doing cell biology. It's a total game changer."

Provided by Allen Institute for Brain Science

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