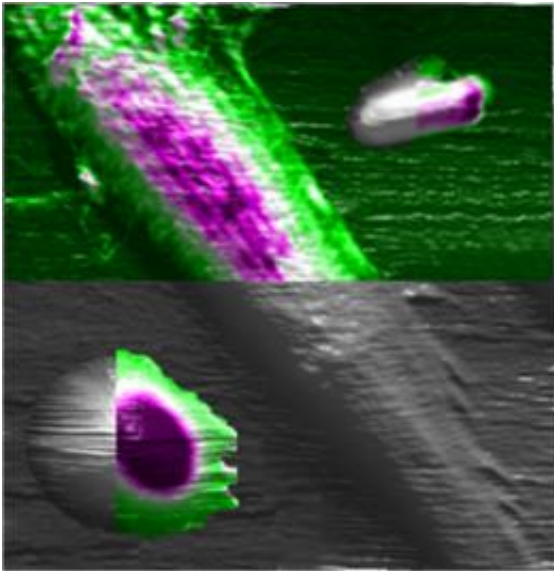


New medical, research tool possible by probing cell mechanics

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This artist's conception depicts the use of an atomic force microscope to study the mechanical properties of cells, an innovation that might result in a new way to diagnose disease and study biological processes. Here, three types of cells are studied using the instrument: a rat fibroblast is the long slender cell in the center, an E coli bacterium is at the top right and a human red blood cell is at the lower left. The colored portions show the benefit of the new technique, representing the mechanical properties of the cells, whereas the gray portions represent what was possible using a conventional approach. Credit: Purdue University image/Alexander Cartagena

(PhysOrg.com) -- Researchers are making progress in developing a system that measures the mechanical properties of living cells, a

technology that could be used to diagnose human disease and better understand biological processes.

The team used an instrument called an atomic force microscope to study three distinctly different types of [cells](#) to demonstrate the method's potentially broad applications, said Arvind Raman, a Purdue University professor of mechanical engineering.

For example, the technique could be used to study how cells adhere to tissues, which is critical for many disease and [biological processes](#); how cells move and change shape; how [cancer cells](#) evolve during metastasis; and how cells react to mechanical stimuli needed to stimulate production of vital proteins. The technique could be used to study the mechanical properties of cells under the influence of antibiotics and drugs that suppress cancer to learn more about the mechanisms involved.

Findings have been posted online in the journal *Nature Nanotechnology* and will appear in the December print issue. The work involves researchers from Purdue and the University of Oxford.

"There's been a growing realization of the role of mechanics in [cell biology](#) and indeed a lot of effort in building models to explain how cells feel, respond and communicate mechanically both in health and disease," said Sonia Contera, a paper co-author and director of the Oxford Martin Programme on Nanotechnology and an academic fellow at Oxford physics. "With this paper, we provide a tool to start addressing some of these questions quantitatively: This is a big step."

An atomic force microscope uses a tiny vibrating probe to yield information about materials and surfaces on the scale of [nanometers](#), or billionths of a meter. Because the instrument enables scientists to "see" objects far smaller than possible using light microscopes, it could be ideal for "mapping" the mechanical properties of the tiniest [cellular](#)

[structures](#).

"The maps identify the mechanical properties of different parts of a cell, whether they are soft or rigid or squishy," said Raman, who is working with doctoral student Alexander Cartagena and other researchers. "The key point is that now we can do it at high resolution and higher speed than conventional techniques."

The high-speed capability makes it possible to watch living cells and observe biological processes in real time. Such a technique offers the hope of developing a "mechanobiology-based" assay to complement standard biochemical assays.

"The atomic force microscope is the only tool that allows you to map the mechanical properties - take a photograph, if you will - of the [mechanical properties](#) of a live cell," Raman said.

However, existing techniques for mapping these properties using the atomic force microscope are either too slow or don't have high enough resolution.

"This innovation overcomes those limitations, mostly through improvements in signal processing," Raman said. "You don't need new equipment, so it's an economical way to bump up pixels per minute and get quantitative information. Most importantly, we applied the technique to three very different kinds of cells: bacteria, human red blood cells and rat fibroblasts. This demonstrates its potential broad utility in medicine and research."

The technique is nearly five times faster than standard [atomic force microscope](#) techniques.

More information: Mapping Nanomechanical Properties of Live Cells

Using Multi-harmonic Atomic Force Microscopy, *Nature Nanotechnology*.

Abstract

The nanomechanical properties of living cells, such as their surface elastic response and adhesion, have important roles in cellular processes such as morphogenesis¹, mechano-transduction², focal adhesion³, motility^{4,5}, metastasis⁶ and drug delivery^{7–10}. Techniques based on quasi-static atomic force microscopy techniques^{11–17} can map these properties, but they lack the spatial and temporal resolution that is needed to observe many of the relevant details. Here, we present a dynamic atomic force microscopy^{18–28} method to map quantitatively the nanomechanical properties of live cells with a throughput (measured in pixels/minute) that is 10–1,000 times higher than that achieved with quasi-static atomic force microscopy techniques. The local properties of a cell are derived from the 0th, 1st and 2nd harmonic components of the Fourier spectrum of the AFM cantilevers interacting with the cell surface. Local stiffness, stiffness gradient and the viscoelastic dissipation of live *Escherichia coli* bacteria, rat fibroblasts and human red blood cells were all mapped in buffer solutions. Our method is compatible with commercial atomic force microscopes and could be used to analyze mechanical changes in tumors, cells and biofilm formation with sub-10 nm detail.

Provided by Purdue University

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