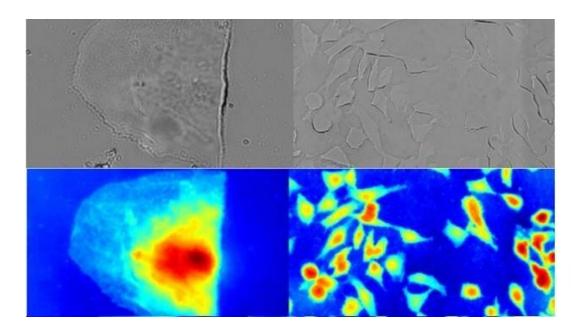


Designer lens helps see the big picture

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Quantitative phase images reveal more details than classical microscopy images. The KAUST technique captures both bright-field images (top) and phase images (bottom) in a single measurement. Credit: KAUST

Microscopes have been at the center of many of the most important advances in biology for many centuries. Now, KAUST researchers have shown how a standard microscope can be adapted to provide even more information.

In its simplest form, microscopy creates an image of an object by measuring the <u>intensity of light</u> passing through it. This requires a



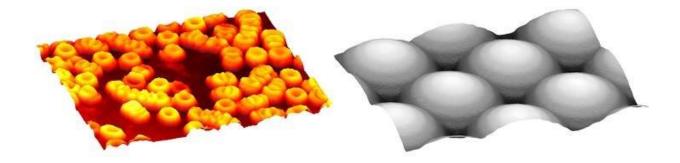
sample that scatters and absorbs light in different ways. Many <u>living cells</u>, however, absorb very little visible light, meaning that there is only a small difference between light and dark regions, known as the contrast. This makes it difficult to see the finer detail.

But the light passing through the sample changes not only its intensity, but also its phase: the relative timing of the peaks in the optical wave. "Phase-contrast microscopy converts phase into larger amplitude variations and hence allows the viewing of fine, detailed transparent structures," explains KAUST Ph.D. student Congli Wang.

Measuring the phase of light is trickier than measuring its intensity. Most phase-contrast microscopes must include a component that converts the phase change to a measurable intensity change. But this conversion is not precise; it only approximates the phase information.

Wang and his colleagues from the KAUSTVisual Computing Center, under the supervision of Wolfgang Heidrich, a professor of computer science, have now developed a new method for quantitative phase and intensity imaging. Crucial to the performance of their <u>microscope</u> was an element known as a wavefront sensor. Wavefront sensors are customdesigned optical sensors that can encode the wavefront, or phase, information into intensity images.





Quantitative phase imaging enables researchers to detect geometry of small transparent samples with fine structures; for example, three-dimensional red blood cells (left) and a microlens array (right). Credit: KAUST

The team designed an innovative high-resolution wavefront sensor, and the team members are now incorporating it into a commercial microscope to improve the performance of microscopy imaging. They then reconstructed the phase-contrast image using a computer algorithm they developed to numerically retrieved quantitative phase from an image pair: a calibration image obtained without the sample and a measurement image obtained with the sample in place.

This approach streamlines several aspects of microscopy. While other methods have achieved quantitative phase imaging in the past, they have required expensive or complicated setups, specialized light sources or a long time to generate the image. "Our method allows snapshot acquisition of high-resolution amplitude bright-field and accurate quantitative phase images via affordable simple optics, common



white-<u>light</u> source and fast computations at video rates in real time," says Heidrich. "It is the first time, to our knowledge, that all these advantages are combined into one technique."

More information: Congli Wang et al. Quantitative Phase and Intensity Microscopy Using Snapshot White Light Wavefront Sensing, *Scientific Reports* (2019). DOI: 10.1038/s41598-019-50264-3

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