



## 3D Ultrasound to Diagnose Cardiovascular Disease



Researchers have developed a new type of imaging technology to diagnose cardiovascular disease and other disorders by measuring ultrasound signals from molecules exposed to a fast-pulsing laser.

The new method could be used to take precise three-dimensional images of plaques lining arteries, said Ji-Xin Cheng, an associate professor of biomedical engineering and chemistry at Purdue University.

Other imaging methods that provide molecular information are unable to penetrate tissue deep enough to reveal the three-dimensional structure of the plaques, but being able to do so would make better diagnoses possible, he said.

"You would have to cut a cross section of an artery to really see the three-dimensional structure of the plaque," Cheng said. "Obviously, that can't be used for living patients."

The imaging reveals the presence of carbon-hydrogen bonds making up lipid molecules in arterial plaques that cause heart disease. The method also might be used to detect fat molecules in muscles to diagnose diabetes and for other lipid-related disorders, including neurological conditions and brain trauma. The technique also reveals nitrogen-hydrogen bonds making up proteins, meaning the imaging tool also might be useful for diagnosing other diseases and to study collagen's role in scar formation.

"Being able to key on specific chemical bonds is expected to open a completely new direction for the field," Cheng said

Findings are detailed in a paper to be published in *Physical Review Letters* and expected to appear in the June 17 issue. The findings represent the culmination of four years of research led by Cheng and doctoral student Han-Wei Wang.

The new technique uses nanosecond laser pulses in the near-infrared range of the spectrum. The laser generates molecular "overtone" vibrations, or wavelengths that are not absorbed by the blood. The pulsed laser causes tissue to heat and expand locally, generating pressure waves at the ultrasound frequency that can be picked up with a device called a transducer.

"We are working to miniaturize the system so that we can build an endoscope to put into blood vessels using a catheter," Cheng said. "This would enable us to see the exact nature of plaque formation in the walls of arteries to better quantify and diagnose cardiovascular disease."

Lihong Wang, Gene K. Beare Distinguished Professor of Biomedical Engineering at Washington University in

St. Louis, is a pioneer of using the "photoacoustic" imaging of blood vessels based on the absorption of light by the electrons in hemoglobin.

The Purdue researchers are the first to show that a strong photoacoustic signal can arise from the absorption of light by the chemical bonds in molecules. The near-infrared laser causes enough heating to generate ultrasound but not enough to damage tissue.

"You can measure the time delay between the laser and the ultrasound waves, and this gives you a precise distance, which enables you to image layers of the tissues for three-dimensional pictures," Cheng said. "You do one scan and get all the cross sections. Our initial target is cardiovascular disease, but there are other potential applications, including diabetes and neurological conditions."

The approach represents a major improvement over another imaging technique, called coherent anti-Stokes Raman scattering, or CARS, which has been used by the Purdue-based lab to study three-dimensional plaque formation in arteries.

Also leading the research are Michael Sturek, chair of the Department of Cellular and Integrative Physiology at the Indiana University School of Medicine; Robert P. Lucht, Purdue's Ralph and Bettye Bailey Professor of Combustion in Mechanical Engineering; and David Umulis, a Purdue assistant professor of agricultural and biological engineering. Other authors of the paper include Purdue graduate students Ning Chai, Pu Wang and Wei Dou and Washington University postdoctoral researcher Song Hu.

Findings are based on research with pig tissues in laboratory samples and also with live fruit flies.

"You can see fat inside fly larvae, representing the potential to study how obesity affects physiology in humans," Cheng said.

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