

## Seeing deep blood flow with sound and laser light

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a, Schematic of the imaging system. Light from a laser is delivered to the blood vessel region via a fibre bundle. Photoacoustic waves are generated and then detected by the ultrasonic probe. Photoacoustic signals are streamed to the computer through the DAQ module. b, Effect of blood heterogeneity on the visibility of the signals arising from the lumen region of the blood vessel. For uniform blood, the simulated photoacoustic signals from the lumen region are suppressed relative to the signals at the boundaries, resulting in little visibility in the lumen region of the image, whereas in non-uniform blood, the simulated photoacoustic signals from the lumen region case. Credit: *Nature Biomedical Engineering* (2023). DOI: 10.1038/s41551-023-01148-5



A great number of health problems, and consequently the medical treatments for them, involve how blood flows through the body. Heart attacks are caused by restricted blood flow to the heart muscle. Many symptoms of diabetes are the result of damaged blood vessels. Tumors, meanwhile, often promote the growth of new vessels that deliver blood specifically to them. And blood flow is a crucial physiological parameter for measuring brain function.

Because of this, <u>medical professionals</u> want to be able to examine <u>blood</u> vessels and assess their condition, but with many of those vessels buried quite deeply in the body, such an examination can be difficult without exploratory surgery.

New research conducted in the laboratory of Caltech's Lihong Wang, Bren Professor of Medical Engineering and Electrical Engineering, is now making it possible to image deep blood vessels in humans, and even the blood flowing through them, in a non-invasive way.

In an <u>article</u> published in the journal *Nature Biomedical Engineering*, Wang and his colleagues describe this technology, which they call photoacoustic vector tomography, or PAVT. This technology is similar in many ways to Wang's other photoacoustic imaging technologies, which make use of laser light that is well absorbed by hemoglobin, the oxygen-carrying molecule found in red blood cells.

The energy that the hemoglobin molecules absorb from the laser causes them to vibrate ultrasonically. Those vibrations travel throughout the tissue until they arrive at the skin's surface, where they are detected by sensors connected to a computer. The computer then creates an image of the features of the tissue, in this case, the blood vessels.



This is not the first time Wang's lab has shown the ability to image blood vessels using photoacoustic technology, but the new method can image <u>blood flow</u> deeper in the <u>human body</u> than previously possible and shows for the first time not just the presence of blood vessels and their oxygenation status but how the blood is flowing through the vessels.

"Before, we could only show the sizes of blood vessels, concentrations of blood, and oxygen saturations," says Wang, also the Andrew and Peggy Cherng Medical Engineering Leadership Chair. "Now, we can measure the vector flow, which indicates both flow rate and direction.

"Our field has been working on photoacoustic technology for more than 20 years, but nobody predicted anything like this. We surprised ourselves because our field didn't think this was possible."

"When I first saw our images of blood flow, I was absolutely amazed," says Yang Zhang, the lead author and a postdoctoral scholar research associate in medical engineering. "The most exciting part of this work is that we synergized engineering and physiology to overcome a hurdle previously thought to be insurmountable by the field."

The team is able to see the direction and <u>flow rate</u> because PAVT has such fine resolution that it can make out signals arising from the distribution of red blood cells deep in the body. An algorithm integrated into the system tracks the motion of these distributions and deduces the speed and direction of the flow. It's kind of like how Google determines how heavy traffic is on a freeway by looking at the speed at which mobile phones are moving in that area.

The researchers hypothesize that their images and videos of human blood flow are facilitated by the heterogenous distribution of red blood cells, which arises, in part, from the way <u>blood vessels</u> are structured throughout the body.



Wang likens the situation in veins to what happens when two rivers with different water qualities, one clear and one muddy, for example, join into one larger stream. At such a confluence, it is not uncommon to see the streams remain unmixed for a long distance even while flowing through the same channel.

A similar phenomenon is seen when two veins carrying blood with differing blood contents (oxygenated and unoxygenated) join together. Even though the blood from those two vessels has joined as a single stream, it will remain unmixed for a while. The PAVT system can distinguish these unmixed patches and track their motion.



At the confluence of the Amazon River and the Rio Negro in Brazil, the waters from each river can be seen running parallel and remaining unmixed for some time after the rivers join. Similar phenomena can be seen in blood vessels.



Credit: Portal da Copa/Wikimedia Commons

And since <u>red blood cells</u> absorb laser light from the PAVT system differently depending on whether they are oxygenated or not, PAVT can also determine how much oxygen the blood in a particular vessel is carrying. "This allows us to quantify oxygen consumption, which is an important measure of metabolism," Wang adds.

The paper describing the research, titled "Photoacoustic vector tomography for deep hemodynamic imaging," appears in the November 30 issue of *Nature Biomedical Engineering*.

**More information:** Yang Zhang et al, Photoacoustic vector tomography for deep haemodynamic imaging, *Nature Biomedical Engineering* (2023). DOI: 10.1038/s41551-023-01148-5. On arXiv: DOI: 10.48550/arxiv.2209.08706

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