

'LAB ON A CHIP' DETECTS HUMAN AND AGRICULTURAL PATHOGENS

The UA Biosensor Lab's flexible diagnostic tool

By Susan McGinley

Detecting waterborne and foodborne contaminants usually involves collecting a water or food sample, sending it to a laboratory and waiting for the samples to be filtered, incubated, tested and identified under a microscope. If a critical infection is suspected, say for *E. coli*, the pathogen may already have multiplied and spread before the report arrives days later.

A series of "lab on a chip" (LOC) applications in development at the University of Arizona can identify pathogens in minutes rather than days, using a simple device (which may be attached to a faucet) that can deliver results locally. The degree of accuracy is three orders of magnitude greater than for conventional real-time or rapid tests, and the method can be used to test lower concentrations of pathogens.

Invented by chemical and electrical engineers in the late 1990s, the lab on a chip concept is based on the integrated circuit, where encapsulated wires and circuits are integrated into a

semiconductor chip with electricity flowing through it.

"In lab on a chip, instead of using electricity, the liquid flows," says Jeong-Yeol Yoon, an assistant professor in the Department of Agricultural and Biosystems Engineering at the University of Arizona and a member of the BIO5 Institute. When Yoon was hired at the UA in 2004 he began to test the technology on water after noticing that attempts by other scientists to test blood with the LOC failed.

"I realized lab on a chip needed to start with the simplest case. Water is easy and very dilute compared to blood," Yoon says. "When you run the test there are almost no sizeable substances except the pathogens. It may detect a single cell."

The LOC is a small glass laboratory slide filled with nanoparticles that adhere to pathogens applied in a sample drop of water. Yoon uses micro- and nanoparticles from 1000 nanometers down to 10 nanometers to scatter light and strengthen sensitivity. He combines them with antibodies to the target pathogens, leading to particle agglutination and changing the scattering signal.

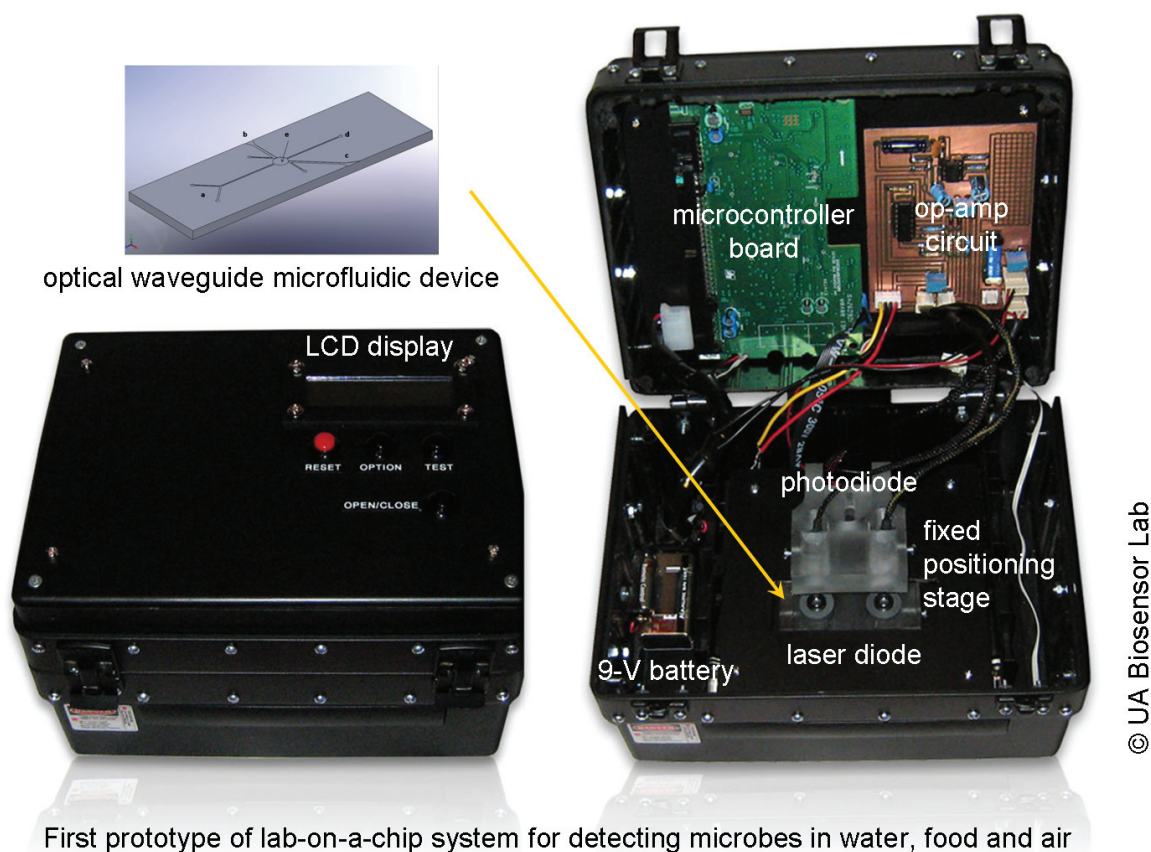
"An antibody with nanoparticles gives a stronger signal," Yoon says. "Others use fluorescent dye; we use nanoparticles as a substitute for dye."

Working with graduate and undergraduate students in his Biosensors Laboratory on campus,

Yoon has designed a test that can detect pathogens—*E. coli* and potentially *Cryptosporidium*—in drinking water networks, irrigation systems, or wastewater recycling facilities. The prototype is finished but he is currently devising different ways to attach the test to water systems, manipulate the nanoparticles and read and deliver the results by computer.

Yoon also has funding from a government agency in his native South Korea to develop tests for livestock diseases in that country. These include LOC biosensors for bovine viral diarrhea, a bacterial disease that can reduce the quality of the milk and meat, and porcine reproductive and respiratory syndrome, a common viral disease that affects both the amount and the quality of the pork produced. Yoon is also monitoring the viral pathogens for avian flu and H1N1 flu in South Korean livestock environments.

"Water supplies are affected first because the animals touch their tongues to the faucets," he says. "We're installing an LOC system on that water supply and also testing their wastewater." The goal is to meet increased consumer demand for choice and premium grades of meat, and for higher quality milk, by detecting and treating diseases earlier. Yoon is also testing air samples from livestock housing for detecting such viral pathogens. The air sample tests show about the same sensitivity and specificity as those with water samples.



To make vegetables safer, Yoon is developing a test for bacterial contamination—specifically *E. coli* and *Salmonella*—in fresh lettuce and spinach.

“We collect lettuce, grind it up, turn it into vegetable juice and then load it onto a lab on a chip and read the results,” he says. “It’s the most complicated application so far, but it’s working well. We’re detecting pathogens not from water but directly from the vegetable tissues.” The effort is funded by Desert Tech Investors.

Yoon’s ultimate step is a human biomedical application: creating an LOC test for the H1N1 virus. For this, he is using a newer “droplet” technology, where droplets on nano-structured surfaces can be manipulated using a control pad—the current prototype in the Biosensors Lab is attached to a Nintendo gaming pad.

“You can choose the reagent droplets you want (either DNA or antibody), move the sample droplet to them, and mix, rotate and/or heat them depending on the nature

of the sample,” Yoon says. “It’s more expensive because you can change the protocol. The circuit memorizes the previous movements for 100 different samples.” The rapid-response test can be done in real time and will be commercially available in about five years. 🌀

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