

DNA fragments in blood promise cheap, easy test for cancer

January 12 2023, by Amy Norton



Researchers are reporting progress on a blood test that can detect



multiple cancers in a relatively simpler, and potentially less pricey way than other tests under development.

The test picks up certain cancer signals in the blood using a fairly straightforward method: counting bits of DNA that appear to be "broken" in unexpected places.

In the new study, scientists found that the approach can distinguish people with any of 11 different types of cancer from people who are cancer-free. Not only that, it can do so using a relatively small blood sample, according to senior researcher Dr. Muhammed Murtaza, of the University of Wisconsin's School of Medicine and Public Health, in Madison.

That's relative to other blood tests under development for cancer screening, which typically require multiple tubes of blood.

"The biggest advantage of our approach would probably be in costeffectiveness," Murtaza said, although he added that it's hard to make predictions on real-world costs.

The study, published Jan. 11 in the journal *Science Translational Medicine*, is among the latest in the broader push to develop blood tests that can serve as a one-stop screening for multiple cancers.

A number of companies are working on such "multi-cancer" early detection tests. Essentially, they are all based on the fact that tumors shed bits of genetic material in the blood. If a test can detect those tumor signals, that could provide a simple, noninvasive way to screen for various cancers—including ones that now lack any screening method.

Where the tests vary is in what, exactly, they are measuring, Murtaza explained.



He and his colleagues took the approach of analyzing DNA "fragmentation patterns." Both <u>tumor cells</u> and <u>healthy cells</u> regularly release pieces of DNA into the blood. But because tumors express different genes than normal body cells do, they also differ in how those DNA bits break off.

The new test looks at the "end position" of DNA fragments in the blood, to see if the breaks happened in "unexpected places," Murtaza said.

He described it as a "simple method" that allows for a smaller blood sample containing a limited supply of DNA (which, in the world of DNA, amounts to a minimum of 1 million fragments).

For the study, the researchers analyzed blood samples from 286 healthy people, 103 who had non-cancer medical conditions, and 994 people with one of 11 types of cancer. They ranged from breast and <u>ovarian</u> <u>cancers</u> to melanoma, to rarer cancers like bile duct tumors and the deadly brain cancer glioblastoma.

Overall, the researchers found, the test did a good job of pinpointing which <u>blood samples</u> came from <u>cancer patients</u>, including those who had early-stage cancer, which is important in a screening test.

The test had an "area-under-curve" (AUC) of 0.91 for cancer overall, and 0.87 for stage 1 cancer, according to the researchers. The AUC is a measure of test accuracy that ranges from 0 to 1: the closer it is to 1, the better.

More research is still needed, according to Murtaza. One of the next steps, he said, is to see how well the test performs in detecting a single cancer, rather than multiple ones—including pancreatic cancer, a deadly disease for which there is no screening test.



A researcher who was not involved in the study agreed that the approach "potentially has some advantages" over other blood tests under development.

"It's another approach that may be a little simpler, and maybe less expensive," said Dr. David Polsky, of NYU Langone's Perlmutter Cancer Center, in New York City.

Like Murtaza, he said the test is "not ready for prime time," and more research is needed.

With any test aimed at screening healthy people for cancer, Polsky said, there are important measures of its performance: its sensitivity, which refers to the test's ability to correctly identify people who have cancer; and its specificity, or its ability to correctly identify people who do not have cancer.

Specificity is critical, Polsky noted, because if it's too low, the test will generate a lot of false-positive results that lead to unnecessary follow-up testing—and anxiety.

Some other blood tests for <u>cancer screening</u> are further along in development, including one called <u>Galleri</u> that screens for over 50 cancers with a single blood draw. It's currently being put to the test in the <u>Pathfinder 2 study</u>, being run at multiple U.S. medical centers.

There are currently no multi-cancer early detection tests approved by the U.S. Food and Drug Administration.

More information: Karan K. Budhraja et al, Genome-wide analysis of aberrant position and sequence of plasma DNA fragment ends in patients with cancer, *Science Translational Medicine* (2023). DOI: 10.1126/scitranslmed.abm6863



The American Cancer Society has more on <u>blood tests</u> under study for cancer screening.

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Citation: DNA fragments in blood promise cheap, easy test for cancer (2023, January 12) retrieved 14 February 2023 from <u>https://medicalxpress.com/news/2023-01-dna-fragments-blood-cheap-easy.html</u>

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