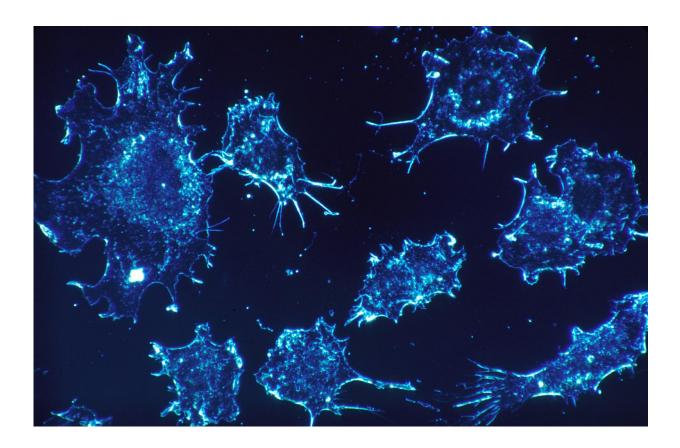


Deadly cancers show early, detectable differences from benign tumors

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Do metastatic cancer tumors "break bad" or are they "born bad"?

This question is an essential mystery in <u>cancer</u> early detection and treatment. Lacking a clear answer, patients are given the same aggressive



therapies when small, abnormal clusters of <u>cells</u> are discovered early, even though they might well be harmless.

In a study publishing the week of May 14 in the *Proceedings of the National Academy of Sciences*, a research team co-led by scientists at Duke and the University of Southern California has found that in the colorectal tumors they examined, invasive cancers are born to be bad, and this tendency can potentially be identified at early diagnosis.

"We found evidence that benign and <u>malignant tumors</u> start differently, and that cell movement—an important feature of malignancy—manifests itself very early on during tumor growth," said lead author Marc D. Ryser, Ph.D., postdoctoral fellow in Duke's departments of Surgery and Mathematics.

"By testing screen-detected, small tumors for early cell movement as a sign of malignancy, it might be possible to identify which patients are likely to benefit from aggressive treatment," Ryser said.

Ryser and colleagues built on recent research showing that in a subset of human cancers, many key traits of the final tumor are already imprinted in the genome of the founding cell. As such, they reasoned, <u>invasive</u> <u>tumors</u> would start out with the ability to spread rather than developing that trait over time. That is, they are born to be bad.

The researchers analyzed 19 human colorectal tumors with genome sequencing technology and mathematical simulation models. They found signatures of early abnormal cell movement in the majority of the invasive samples—nine of 15. This propensity is required for tumors to spread, causing them to become deadly. Early abnormal cell <u>movement</u> was not apparent in the four <u>benign tumors</u> the researchers studied.

"The early growth of the final <u>tumor</u> largely depends on the drivers



present in the founding cell," the authors wrote.

The study was small and the researchers acknowledged that verification in a larger sample is required, but the finding is a significant step toward establishing a test to distinguish between deadly and harmless growths.

"Thanks to improved screening technologies, we diagnose more and more small tumors," said senior author Darryl Shibata, M.D., professor in the Department of Pathology at Keck School of Medicine of USC. "Because treating a patient aggressively can cause them harm and sideeffects, it is important to understand which of the small screen-detected tumors are relatively benign and slowly growing, and which ones are born to be bad."

More information: Marc D. Ryser el al., "Spatial mutation patterns as markers of early colorectal tumor cell mobility," *PNAS* (2018). <u>www.pnas.org/cgi/doi/10.1073/pnas.1716552115</u>

Provided by Duke University Medical Center

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