

Gene mutation combo linked to common cancer in women

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Ronald Chandler is an assistant professor of obstetrics, gynecology and reproductive biology in the College of Human Medicine. Credit: MSU

Michigan State University researchers, in collaboration with the Van Andel Institute, have identified a combination of two gene mutations



that is linked to endometrial cancer.

"More than 63,000 women are likely to be diagnosed with endometrial cancer this year, making it the most commonly identified type of gynecologic cancer," said Ronald Chandler, an assistant professor of obstetrics, gynecology and <u>reproductive biology</u> in the College of Human Medicine, who led the study.

The research, published in *Nature Communications*, found that <u>mutations</u> of the ARID1A and PIK3CA genes are frequently found together in the development of endometrial cancer, as well as in endometriosis-associated <u>ovarian cancer</u>. The findings could mean that better, more targeted treatments can be developed to help fight the disease.

ARID1A is a tumor suppressor. When it mutates, chromatin, cellular material that keeps DNA compacted in cells, loses its structure, allowing cancer to spread. PIK3CA is an instructional gene that tells the body to produce certain proteins and leads to uncontrolled growth of cells when it mutates.

Similarly, these same mutations often are found in women who have endometriosis—a painful condition in which uterine tissue grows outside the uterus—but many of those affected never develop <u>endometrial</u> <u>cancer</u>.

"We're trying to understand why some women with the same set of mutations get cancer and some don't," Chandler said. "There is something else involved."

Endometrial cancer commonly occurs in women after menopause, suggesting that other factors, such as hormonal changes, environmental exposures or obesity, could combine with the mutations to trigger cancer.



"The hardest part of our job is figuring out what's causing it," Chandler said. "Is it something in the environment or it is something else? It's a complex process. These are big questions that we're trying to address."

Through a <u>collaborative effort</u>, Chandler's team provided genetic material to Van Andel Institute. VAI researchers then analyzed the samples with a next-generation sequencer, a machine that rapidly sequences the human genome. This helped MSU researchers identify which gene mutations were associated with the <u>cancer</u>.

"Ten years ago, this research would not have been possible," said Marie Adams, VAI's genomics core manager and a study co-author. "Next-generation sequencers are powerful new tools we can use to process large-scale projects and provide actionable results in weeks as opposed to years."

More information: Mike R. Wilson et al. ARID1A and PI3-kinase pathway mutations in the endometrium drive epithelial transdifferentiation and collective invasion, *Nature Communications* (2019). DOI: 10.1038/s41467-019-11403-6

Provided by Michigan State University

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