

Estrogen receptor mutation study suggests potential treatments for endometrial cancers

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Jay Gertz, Ph.D. and Zannel Blanchard, Ph.D. Credit: Huntsman Cancer Institute

Researchers at Huntsman Cancer Institute identified potential new treatment options for people with endometrial cancer. Endometrial cancer is the most common gynecological cancer and high levels of estrogen promote its development. The study, published in *Molecular Cancer Research* discovered that estrogen receptor mutations found in endometrial cancers cause large changes in endometrial cancer cells.



Estrogen is a reproductive hormone that binds and activates estrogen receptors. Cancer can cause estrogen receptors to remain in a state of constant activity. That increases shedding of the endometrial lining.

"Our goal was to characterize estrogen receptor mutations in endometrial <u>cancer</u> to see how they affected gene expression, along with how these mutations made cells more aggressive and fast-growing," says Zannel Blanchard, Ph.D., postdoctoral fellow at Huntsman Cancer Institute and lead investigator on the study. "We found that the mutations caused large changes in <u>gene expression</u> and cellular behavior."

The team used their findings to identify potential treatments for <u>endometrial cancers</u> with high levels of estrogen receptor activity. They found that inhibitors of CDK9, a protein that works with <u>estrogen</u> receptors, were effective at reducing the growth and aggressiveness of endometrial cancer cells.

"Besides surgery to treat endometrial cancer, there's only one drug approved by the Food and Drug Administration for treating primary endometrial cancer, and it was approved in the 1970s," says Jay Gertz, Ph.D., the senior author and researcher at Huntsman Cancer Institute and associate professor of oncological sciences at the University of Utah. "Our results help us to really start moving toward personalized—or precision—medicine for endometrial cancer."

The study was presented to a breast and gynecologic cancer research advocacy group made up of patients. Participants volunteer their time and meet once a month at Huntsman Cancer Institute.

"We can write support letters to help with upcoming research grants and offer a patient perspective to researchers," says Deb Jordan, an endometrial cancer patient at Huntsman Cancer Institute and advocacy group participant. "We also get to hear about the ongoing research at



Huntsman Cancer Institute. That puts my mind at ease, to learn all that is being done for endometrial cancer."

"It's exciting because the study suggests that there may be more options for endometrial cancer patients," says Blanchard. "There's more going on when you look deeper and we get to share these findings with patients who have been through treatment."

Gertz says the <u>treatment options</u> for endometrial cancer are limited and patients play an important role in inspiring researchers to find new therapies. The study suggests that molecular evaluation of tumors could lead to more personalized treatment options for <u>endometrial cancer</u> patients.

More information: Zannel Blanchard et al, Allele-specific Gene Regulation, Phenotypes, and Therapeutic Vulnerabilities in Estrogen Receptor Alpha–mutant Endometrial Cancer, *Molecular Cancer Research* (2023). DOI: 10.1158/1541-7786.MCR-22-0848

Provided by Huntsman Cancer Institute

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