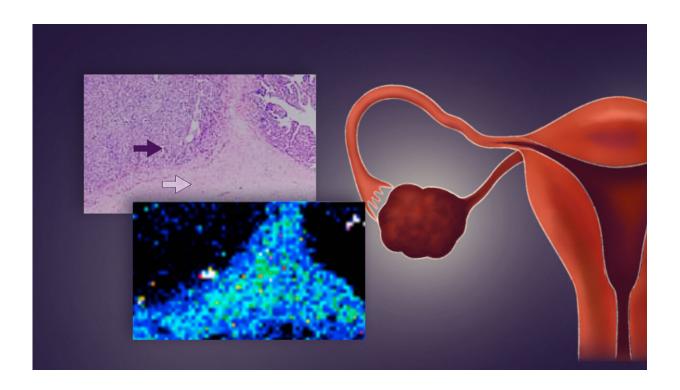


Platinum-based chemotherapy distribution within a tumor may predict ovarian cancer treatment resistance

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Tumor tissue is indicated with the filled arrow and stroma with the unfilled arrow (upper photo). A platinum-resistant patient showing platinum (blue in the bottom photo) mostly in the stroma instead of the tumor. Credit: Uno et al., and Reiko Matsushita

A team of researchers from Japan has discovered that the distribution of



platinum within a tumor following platinum-based chemotherapy treatment of ovarian cancer may predict whether the tumor will be resistant to further treatment. The research could offer ways to manage treatment for women whose tumors may be resistant to further platinumbased chemotherapy.

One of the biggest problems with treating ovarian <u>cancer</u> is completely removing the cancer surgically. To facilitate easier surgical removal, patients are given <u>platinum</u>-based chemotherapy drugs early in treatment to shrink the tumor. These drugs introduce platinum into the DNA of the <u>cancer cells</u> making up the tumor, which makes the DNA difficult to read and causes the cells to die.

However, if the tumor is not completely removed, it takes advantage of the body's weakened immunity and comes back, often growing faster than before.

Because the tumor has experienced the treatment before, it develops increased resistance to it, in the same manner that bacteria can become resistant to antibiotics. Repeated exposure leads to 80% of women developing ovarian cancer tumors that are resistant to further treatment. Therefore, a technique that would allow doctors to tell if the patient will develop resistance to these commonly used drugs would make this process a lot more effective.

A team of researchers led by Professor Hiroaki Kajiyama, Assistant Professor Nobuhisa Yoshikawa, and graduate student Kaname Uno of the Nagoya University Graduate School of Medicine reported that the answer to this problem lies in the distribution of platinum in the tumor after treatment. Their findings were published in *Scientific Reports*.

The team looked at the amount of platinum in both the tumor itself and in its stroma, which is the part of the tumor that supplies cancer cells



with nutrition and gets rid of <u>waste products</u>. They found that there were two basic types of platinum distribution: one where the platinum was evenly distributed throughout both the tumor itself and the stroma, and one where the platinum was mostly in the stroma.

The difference was staggering: only one of the 11 patients with an even distribution of platinum died in the three years after diagnosis, whereas half of 16 patients with most of the platinum in the stroma were dead within three years. In addition, the patients with tumors that showed an even distribution of platinum were also less likely to develop a tumor that was resistant to further treatment.

"We showed that there were two different patterns of platinum distribution within the tumor tissue. The difference in the distribution was shown to be significantly correlated with prognosis, suggesting that it is possible to discriminate between platinum-resistant and platinum-sensitive tumors before they recur," explains Kajiyama. "Using this method, early diagnosis of platinum resistance and selection of appropriate postoperative anticancer therapy for patients as well as a new ovarian cancer treatment strategy are possibilities."

Predicting whether a <u>tumor</u> will become resistant to platinum-based drugs would solve one of the major difficulties in successfully treating <u>ovarian cancer</u>. A future where doctors can modify chemotherapy for <u>patients</u> most at risk of resistance will improve their chances of surviving and make sure that the most effective treatment is given.

More information: Kaname Uno et al, Significance of platinum distribution to predict platinum resistance in ovarian cancer after platinum treatment in neoadjuvant chemotherapy, *Scientific Reports* (2022). DOI: 10.1038/s41598-022-08503-7



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