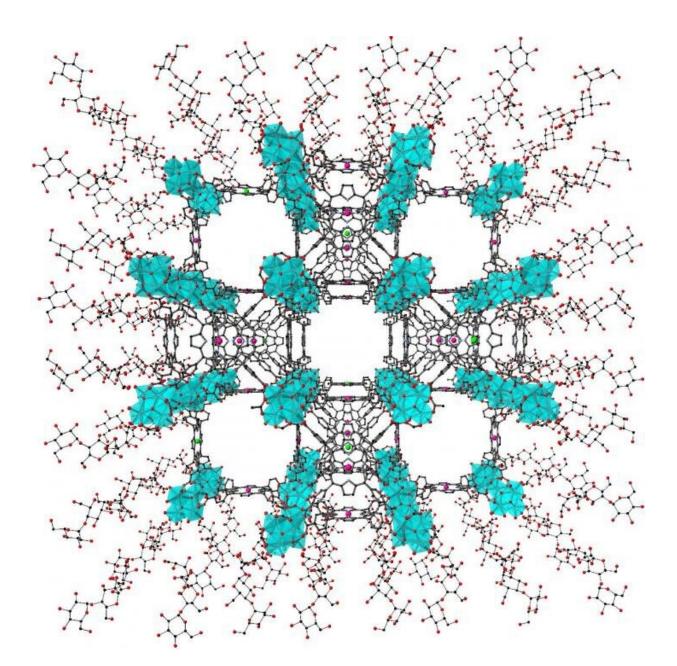


New nano drug candidate kills aggressive breast cancer cells

July 20 2020, by Matt McGowan





Chemical structure of multi-functional, anticancer drug candidate. Credit: Hassan Beyzavi, University of Arkansas.

Researchers at the University of Arkansas have developed a new nano drug candidate that kills triple negative breast cancer cells.

Triple negative <u>breast</u> cancer is one of the most aggressive and fatal types of breast cancer. The research will help clinicians target breast cancer <u>cells</u> directly, while avoiding the adverse, toxic side effects of chemotherapy.

Their study was published in June issue of Advanced Therapeutics.

Researchers led by Hassan Beyzavi, assistant professor in the Department of Chemistry and Biochemistry, linked a new class of nanomaterials, called <u>metal-organic frameworks</u>, with the ligands of an already-developed photodynamic therapy drug to create a nano-porous material that targets and kills tumor cells without creating toxicity for <u>normal cells</u>.

Metal-organic frameworks are an emerging class of nanomaterials designed for targeted drug delivery. Ligands are molecules that bind to other molecules.

"With the exception of skin cancers, breast cancer is the most common form of cancer in American women," said Beyzavi. "As we know, thousands of women die from breast cancer each year. Patients with triple negative cells are especially vulnerable, because of the toxic side effects of the only approved treatment for this type of cancer. We've addressed this problem by developing a co-formulation that targets cancer cells and has no effect on healthy cells."



Researchers in Beyzavi's laboratory focus on developing new, targeted photodynamic therapy drugs. As an alternative to chemotherapy—and with significantly fewer side effects—targeted photodynamic therapy, or PDT, is a noninvasive approach that relies on a photosensitizer that, upon irradiation by light, generates so-called toxic reactive oxygen species, which kill cancer cells. In recent years, PDT has garnered attention because of its ability to treat tumors without surgery, chemotherapy or radiation.

Beyzavi's laboratory has specialized in integrating nanomaterials, such as metal-organic frameworks, with PDT and other and therapies. Metal-organic frameworks significantly enhance the effectiveness of PDT.

Doctoral student Yoshie Sakamaki from Beyzavi's laboratrory prepared the nanomaterials and then bio-conjugated them with ligands of the PDT drug to create nanoporous materials that specifically targeted and killed tumor cells with no toxicity in normal cells.

In addition to <u>cancer treatment</u>, this novel drug delivery system could also be used with magnetic resonance imaging (MRI) or fluorescence imaging, which can track the drug in the body and monitor the progress of cancer treatment.

This collaborative project also included contributions from U of A research groups through Julie Stenken, professor of analytical chemistry; Yuchun Du, associate professor of biological sciences; and Jin-Woo Kim, professor of biological and agricultural engineering.

The American Cancer Society estimated 268,600 new cases of invasive breast cancer in 2019 and 41,760 deaths. Currently there are more than 3.1 million breast cancer survivors in the United States. Since 2007, breast cancer death rates have been steady in women younger than 50 but have continued to decrease in older women. This decrease is



believed to be the result of earlier detection and better treatments.

Triple negative breast cancer is aggressive and lacks estrogen receptors, progesterone receptors and human epidermal growth factor receptor 2, which means it cannot be treated with receptor-targeted therapy. It is difficult to treat with existing chemotherapy and often requires surgery because it quickly metastasizes throughout the body.

Cytotoxic chemotherapy is the only approved treatment for this type of breast cancer. More than 80% of women with triple negative breast <u>cancer</u> are treated with chemotherapy regimens that include anthracyclines, such as doxorubicin, which can cause cardiotoxicity as a serious side effect. Furthermore, chemotherapy treatment of <u>breast</u> <u>cancer</u> cell lines using either 5-FU, cisplatin, paclitaxel, doxorubicin or etoposide have shown multi-drug resistance.

More information: Yoshie Sakamaki et al, Maltotriose Conjugated Metal–Organic Frameworks for Selective Targeting and Photodynamic Therapy of Triple Negative Breast Cancer Cells and Tumor Associated Macrophages, *Advanced Therapeutics* (2020). DOI: 10.1002/adtp.20200029

Provided by University of Arkansas

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