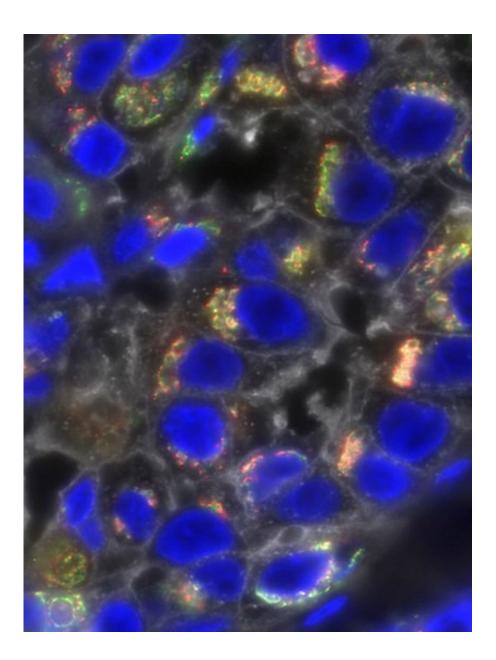


Genetic targets to chemo-resistant breast cancer identified

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Triple negative breast cancer cells, shown here, often become resistant to



chemotherapy over time. New research suggests drugs that inhibit the genes MCL1 or MYC might prevent this. Credit: NCI

Research led by Dr. Carlos Arteaga, Director of the Harold C. Simmons Comprehensive Cancer Center, has identified potential targets for treatment of triple negative breast cancer, the most aggressive form of breast cancer.

Increased activity of two genes, *MCL1* and *MYC*, is associated with the development of <u>chemotherapy resistance</u>. The increased action of these two genes boosts mitochondrial oxidative phosphorylation, which promotes the growth of chemotherapy-resistant <u>cancer stem cells</u>, the research showed.

"Alterations in these two genes are easily detectable with tumor gene tests in current use. Combinations of drugs that inhibit *MCL1* or *MYC*, or both, have the potential to reduce the development of chemotherapy resistance and should be studied in clinical trials," said Dr. Arteaga, Professor of Internal Medicine at UT Southwestern Medical Center. Dr. Arteaga holds The Lisa K. Simmons Distinguished Chair in Comprehensive Oncology.

Most breast cancers can be treated with hormone therapy, but about 15 percent of cases are triple negative breast cancer, meaning the <u>cancer</u> <u>cells</u> are not influenced by hormones like estrogen or progesterone. These triple negative breast cancers must, therefore, be treated with chemotherapy, which is toxic to healthy cells as well as cancer cells. Furthermore, most triple negative breast cancers eventually become resistant to chemotherapy and the cancer then spreads unchecked.

Drugs that inhibit activity of the MCL1 or MYC genes are in



development, Dr. Arteaga said. These drugs, given in conjunction with standard chemotherapies, could potentially slow or even prevent the development of chemotherapy resistance, improving the outlook for this aggressive form of <u>breast cancer</u>.

More information: MYC and MCL1 Cooperatively Promote Chemotherapy-Resistant Breast Cancer Stem Cells via Regulation of Mitochondrial Oxidative Phosphorylation, DOI: <u>dx.doi.org/10.1016/j.cmet.2017.09.009</u>, <u>http://www.cell.com/cellmetabolism/fulltext/S1550-4131(17)30560-0</u>

Provided by UT Southwestern Medical Center

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