

Researchers target 'undercover' gene that helps cancer cells proliferate

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For decades, scientists and doctors have been searching for a cure for cancer. Now, researchers from the Cancer Science Institute of Singapore (CSI Singapore) at the National University of Singapore (NUS) have taken a significant step forward with a new concept that could ultimately result in more effective treatments.



Research has so far focused on identifying and targeting the genes that directly cause <u>cancer</u> called driver genes. However, non-driver genes can also enlist other genes to assist them in helping <u>cancer cells</u> proliferate. These little-known additional genes are named "onco-requisite factors" by the research team led by Associate Professor Takaomi Sanda from CSI Singapore. The researchers demonstrate, with a leukemia case study, why it is important to understand these genes.

In leukemia and other cancers, there is a gene that does not directly cause the cancer, but is abnormally active in cancer cells. It produces an enzyme called aldehyde dehydrogenase.

Aldehyde dehydrogenase enlists other enzymes to turbo-charge the cancer cells with energy for growth and proliferation. But crucially, it also helps reduce the wear and tear on these turbo-charged cells by removing toxic by-products of energy production. Otherwise, the cancer cells would end up killing themselves.

While it may be possible to treat cancer by depriving the cells of aldehyde dehydrogenase, more research is needed.

"Cancer is not a disease of one instrument—it is an orchestra. The performance depends not just on the soloist, but on every single instrument involved. This broadens our understanding of cancer and possible ways to disrupt its machinery," explained Assoc Prof Sanda, who is also from the NUS Department of Medicine.

The NUS researchers are now examining which stage of cancer involves <u>aldehyde dehydrogenase</u>, as that would determine whether it can be utilized to prevent cancer, or to kill the already formed cancer <u>cells</u>.

Ms Zhang Chujing, an NUS doctoral student who worked on the project with Assoc Prof Sanda, hopes other researchers will pick up on their



idea of onco-requisite factors. "This concept invites us to focus on the roles of other genes, which are not sitting in the driver seat of cancer formation, but may hold equal power in terms of cancer treatment," she said.

The paper on this work was first published in the journal Haematologica earlier this year, and was selected as one of the late-breaking abstracts at the prestigious Annual American Association for Cancer Research (AACR) meeting. In addition, the NUS team presented the findings at the Frontiers in Cancer Science conference on 4 November 2020.

Frontiers in Cancer Science is an <u>annual event</u> that brings distinguished cancer researchers with complementary knowledge and expertise from across the globe together for the exchange of ideas and information. This year's conference, which was held online from 2 to 6 November 2020, was attended by over 1,200 participants from over 40 countries.

More information: Chujing Zhang et al. Oncorequisite role of an aldehyde dehydrogenase in the pathogenesis of T-cell acute lymphoblastic leukemia, *Haematologica* (2020). DOI: 10.3324/haematol.2019.245639

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