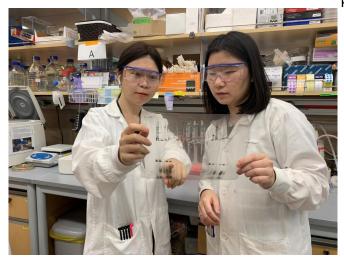


Scientists find new mechanism of cancer formation

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The novel mechanism of cancer formation discovered by NUS researchers Dr Song Yangyang (left), Dr Polly Leilei Chen (right), and their team could lead to more effective treatment. Credit: National University of Singapore

A team of scientists at the National University of Singapore (NUS), led by Dr. Polly Leilei Chen from the Cancer Science Institute of Singapore and Yong Loo Lin School of Medicine at NUS, has discovered a previously unknown mechanism of cancer formation, the understanding of which may lead to more effective treatment.

Their findings concern a process called RNA editing. The DNA code of a gene gets transcribed into an intermediate code known as RNA, before being translated into a protein molecule that plays a particular role in the cell. Sometimes, the RNA gets modified, or edited, before the final translation, yielding a different protein product. RNA editing is a phenomenon that gives the cell finer control over its proteins.

The research team examined the RNA encoding a

protein called 'coatomer subunit?' (COPA), which influences the development of cancers of the liver, esophagus, stomach and breast, and examined whether RNA transcribed from the COPA gene was edited or altered in clinical samples of cancerous liver tissues. They discovered that any given cell contains a mix of both edited and unedited versions of COPA. When the unedited or 'wild type' COPA is predominant, the cell is more likely to become cancerous. Conversely, when edited COPA is predominant, it is thought to suppress a convoluted molecular signaling network called the PI3K/AKT/mTOR signaling pathway. When this pathway gets out of control, it triggers excessive cell multiplication which can lead to cancer.

Their findings were first published in the scientific journal, the *Journal of Hepatology*, on 18 July 2020.

The researchers are now trying to find a way to boost the natural RNA editing mechanisms in the cancerous cell to tip the balance in favor of the edited version of COPA, thereby suppressing the cancer.

"With this <u>new knowledge</u>, we can now look into how A-to-I RNA editing contributes to cancer by altering their <u>protein sequences</u> and how we can restore <u>cancer</u>-suppressing processes mediated by RNA editing in the cell," said Dr. Chen.

More information: Yangyang Song et al, RNA editing mediates the functional switch of COPA in a novel mechanism of hepatocarcinogenesis, *Journal of Hepatology* (2020). DOI: 10.1016/j.jhep.2020.07.021

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