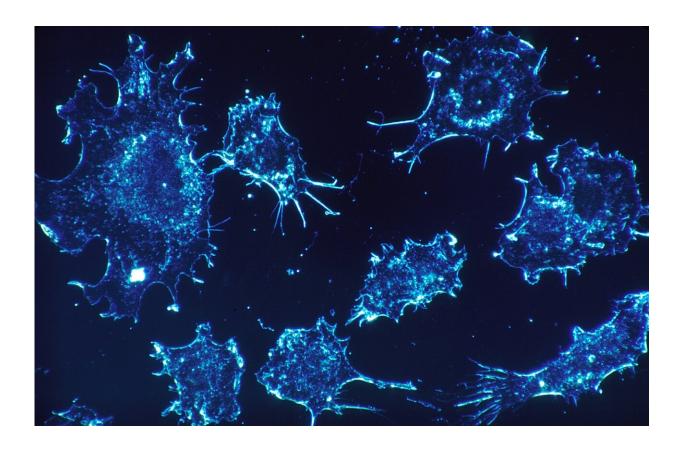


Nanorobot kills cancer cells in mice with hidden weapon

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Researchers at Karolinska Institutet in Sweden have developed nanorobots that kill cancer cells in mice. The robot's weapon is hidden in a nanostructure and is exposed only in the tumor microenvironment, sparing healthy cells. The study is <u>published</u> in the journal *Nature*



Nanotechnology.

The research group at Karolinska Institutet has previously developed structures that can organize so-called death receptors on the surface of cells, leading to cell death. The structures exhibit six peptides (amino acid chains) assembled in a hexagonal_pattern.

"This hexagonal nanopattern of peptides becomes a <u>lethal weapon</u>," explains Professor Björn Högberg at the Department of Medical Biochemistry and Biophysics, Karolinska Institutet, who led the study.

"If you were to administer it as a drug, it would indiscriminately start killing cells in the body, which would not be good. To get around this problem, we have hidden the weapon inside a nanostructure built from DNA."

Created a 'kill switch'

The art of building <u>nanoscale structures</u> using DNA as a <u>building</u> <u>material</u> is called DNA origami and is something Högberg's research team has been working on for many years. Now they have used the technique to create a 'kill switch' that is activated under the right conditions.

"We have managed to hide the weapon in such a way that it can only be exposed in the environment found in and around a solid tumor," he says. "This means that we have created a type of nanorobot that can specifically target and kill cancer cells."

The key is the low pH, or acidic microenvironment that usually surrounds cancer cells, which activates the nanorobot's weapon. In cell analyses in <u>test tubes</u>, the researchers were able to show that the peptide weapon is hidden inside the nanostructure at a normal pH of 7.4, but that



it has a drastic cell-killing effect when the pH drops to 6.5.

Reduced tumor growth

They then tested injecting the nanorobot into mice with breast cancer tumors. This resulted in a 70 percent reduction in <u>tumor growth</u> compared to mice given an inactive version of the nanorobot.

"We now need to investigate whether this works in more advanced cancer models that more closely resemble the real human disease," says the study's first author Yang Wang, a researcher at the Department of Medical Biochemistry and Biophysics, Karolinska Institutet. "We also need to find out what side effects the method has before it can be tested on humans."

The researchers also plan to investigate whether it is possible to make the nanorobot more targeted by placing proteins or peptides on its surface that specifically bind to certain types of cancer.

More information: A DNA Robotic Switch with Regulated Autonomous Display of Cytotoxic Ligand Nanopatterns, *Nature Nanotechnology* (2024). DOI: 10.1038/s41565-024-01676-4, www.nature.com/articles/s41565-024-01676-4

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