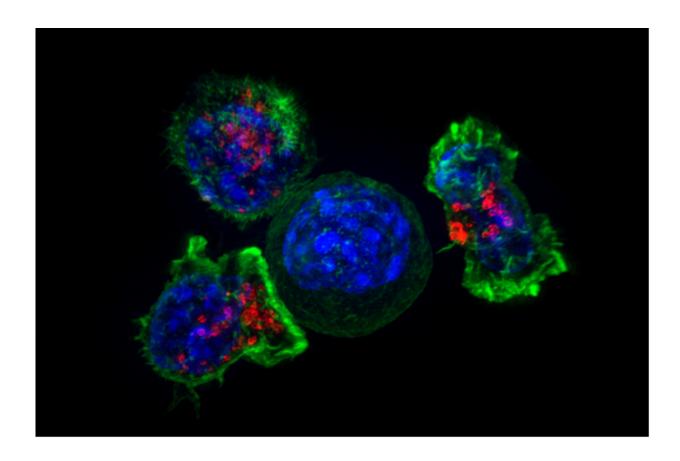


Fully reprogrammed virus offers new hope as cancer treatment

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Killer T cells surround a cancer cell. Credit: NIH

A cancer treatment that can completely destroy cancer cells without affecting healthy cells could soon be a possibility, thanks to research led by Cardiff University.



The team of researchers has successfully 'trained' a respiratory virus to recognise ovarian cancer and completely destroy it without infecting other cells.

The reprogrammed virus could also be used to treat other cancers such as breast, pancreatic, lung and oral.

Dr. Alan Parker from Cardiff University's School of Medicine said: "Reprogrammed viruses are already being used in gene therapy procedures to treat a range of diseases, demonstrating they can be trained from being life-threatening into potentially lifesaving agents.

"In cancer treatment, up until now, reprogramed viruses have not been able to selectively recognise only the cancer cells and would also infect <u>healthy cells</u>, resulting in unwanted side effects.

"We've taken a common, well-studied virus and completely redesigned it so that it can no longer attach to non-cancerous cells but instead seeks out a specific marker protein called $\alpha v \beta 6$ integrin, which is unique to certain cancer cells, allowing it to invade them.

"In this case we introduced the reprogrammed virus to ovarian cancer which it successfully identified and destroyed.

"This is an exciting advance, offering real potential for patients with a variety of cancers."

Once the virus enters the cancer cell it uses the cell's machinery to replicate, producing many thousands of copies of itself, prior to bursting the cell and thereby destroying it in the process. The newly released viral copies can then bind and infect neighbouring cancer cells and repeat the same cycle, eventually removing the tumour mass altogether. The virus also activates the body's natural immune system, helping it to recognise



and destroy the malignant cells.

The reprogrammed virus is from a group of respiratory viruses called adenoviruses. The advantage of using these viruses is that they are relatively easy to manipulate and have already been safely used in <u>cancer treatment</u>.

The technique used to reprogramme the virus to identify the protein common to ovarian, breast, pancreatic, lung and oral cancers could also be used to manipulate it so that it would recognise proteins common to other groups of cancers.

Additional refinement to the viral DNA could also allow the virus to produce anticancer drugs, such as antibodies, during the process of infecting cancer cells. This effectively turns the cancer into a factory producing drugs that will cause its own destruction.

The research was carried out in a laboratory, using mice with ovarian cancer, and has not yet reached clinical trials. The next step is to test the technique with other cancers, with a view to starting clinical trials in five years' time.

Dr. Catherine Pickworth from Cancer Research UK said: "It's encouraging to see that this <u>virus</u>, which has been modified to recognise markers on <u>cancer</u> cells, has the ability to infect and kill ovarian <u>cancer</u> cells in the lab. Viruses are nature's nanotechnology and harnessing their ability to hijack <u>cells</u> is an area of growing interest in <u>cancer research</u>. The next step will be more research to see if this could be a safe and effective strategy to use in people."

The paper "Ad5NULL-A20 – a tropism-modified, ανβ6 integrinselective oncolytic adenovirus for epithelial <u>ovarian cancer</u> therapies" is published in *Clinical Cancer Research*.



More information: Hanni Uusi-Kerttula et al. Ad5NULL-A20 - a tropism-modified, ανβ6 integrin-selective oncolytic adenovirus for epithelial ovarian cancer therapies, *Clinical Cancer Research* (2018). DOI: 10.1158/1078-0432.CCR-18-1089

Provided by Cardiff University

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