

02/09/2024

Two-in-one breakthrough: Cutting-edge immunotherapy could hold promise for incurable brain cancer

Researchers at WEHI have identified a promising new two-in-one treatment that not only targets and destroys an aggressive form of brain cancer, but also helps the immune system develop a lasting defence against it.

This dual-action approach uses a specific immunotherapy known as CAR T cell therapy to treat gliomas, an incurable brain cancer with few treatment options.

The pre-clinical findings have revealed the therapy's ability to eliminate glioma cells and its potential to strengthen the immune system to prevent future tumour growth – two significant advances that could revolutionise the way these lethal cancers are treated in the future.

At a glance

- WEHI researchers have found a specific immunotherapy could hold promise for treating gliomas, an aggressive form of brain cancer with limited treatment options
- The new study shows that CAR T cell therapy not only kills glioma cancer cells, but can also
 provide lasting immunity and reduce the likelihood of tumour recurrence in pre-clinical models
- The findings could transform the way gliomas and other cancers are treated in the future

Overcoming barriers to save lives

In Australia, one person is diagnosed with brain cancer every 5 hours. There currently is no cure and current therapies for severe brain cancers like gliomas are largely ineffective, leaving patients with few options.

CAR T cell therapy is an innovative new approach that involves isolating a patient's immune cells, engineering them to become "super killer cells" and then re-infusing them into the patient to fight their cancer.

While CAR T cell therapy has shown promise for brain cancer treatment, finding the right proteins to target on brain cancer cells has been a major hurdle.

The WEHI-led study has revealed a protein called EphA3, found on the surface of high-grade glioma cells, could be a key target for CAR T cell therapy.

Lead researcher and laboratory head at WEHI, Professor Misty Jenkins AO, says the potential impact of this discovery is profound on many levels.

"This new way to treat gliomas could offer a potential lifeline for patients," Prof Jenkins, also head of the immunotherapy program at The Brain Cancer Centre, said.



"Our research has shown that EphA3-targeted CAR T cells can effectively seek out and eliminate glioma cells, both in lab tests and animal models.

"This therapy not only targeted and killed the cancer cells, but also triggered a long-lasting immune response, preventing future tumour growth.

"We never expected a two-in-one breakthrough that could offer such life-saving possibilities.

"This is a significant development because gliomas are incredibly challenging to treat, and current options are limited.

"Most importantly, this could mean longer survival times for patients and improved quality of life, giving hope to those affected by these aggressive cancers."

Future of cancer medicine

Prof Jenkins said CAR T cell therapy was opening up new pathways for brain cancer treatment.

"CAR T cell therapy is clever because it uses a patient's own immune cells and engineers them to recognise and kill the tumour, with no long-term side effects," she said.

"The tricky nature of brain cancer means that traditional methods don't work – chemotherapy is ineffective, and radiation kills cancerous and healthy brain tissue.

"This recent discovery is further evidence that immunotherapy is a powerful force that can help overcome many cancer challenges that were previously considered insurmountable."

Powerful new tools for cancer treatment

The project team is hopeful that future clinical trials for EphA3-targeted CAR T cell therapy could lead to the development of a treatment that not only kills and halts cancer growth, but also provides lasting immunity.

As the research advances, there is hope that partnerships with biotech companies and investors will help bring this innovative therapy to patients as quickly as possible.

"The possibility of providing patients with a long-lasting immune response against their cancer is a gamechanger, offering a significantly better outlook for patients and their families," Prof Jenkins said.

"The potential to revolutionise brain cancer treatment – like what has been achieved for other cancers – makes this a truly exciting time in the field of cancer research.

"As we stand on the brink of transforming brain cancer treatment, we are not just aiming for better outcomes – we're striving for a future where the words "brain cancer" no longer carry the same fear. The journey ahead promises breakthroughs that will redefine hope for patients around the world."

The research has been supported by many collaborators, key partners and major funders, including the National Health and Medical Research Council (NHMRC), The Brain Cancer Centre, the Robert Connor



Dawes Foundation, the Isabella and Marcus Foundation, the Medical Research Future Fund (MRFF), Cure Brain Cancer Foundation and the Zoe Stanley Research Program (Zoe's Fight Foundation).

The study, "<u>EphA3-targeted chimeric antigen receptor T cells are effective in glioma and generate curative memory T cell responses</u>", is published in the *Journal for ImmunoTherapy of Cancer* (DOI: 10.1136/jitc-2024-009486).

WEHI authors: Leesa Lertsumitkul, Melinda Iliopoulos, Stacie Wang, Alexander Davenport, Ryan Cross, Misty Jenkins.

Image Captions

Critical Mass

Chimeric Antigen Receptor (CAR) T cells, genetically engineered leukocytes (white blood cells), in their fight against glioblastoma, a deadly type of glioma. Credit: WEHI

Lead Researcher

Lead researcher on the study and laboratory head at WEHI, Professor Misty Jenkins AO. Credit: WEHI

WEHI Authors

WEHI authors of the study, including [pictured L to R]: research assistant Melinda Iliopoulos, CAR T cell researcher Dr Ryan Cross, PhD student Leesa Lertsumitkul, laboratory head Professor Misty Jenkins AO, immunotherapy researcher Dr Alex Davenport and paediatric oncologist Dr Stacie Wang. Credit: WEHI

Media enquiries

M: +61 475 751 811 E: communications@wehi.edu.au

About WEHI (Walter and Eliza Hall Institute of Medical Research)

WEHI is where the world's brightest minds collaborate and innovate to make life-changing scientific discoveries that help people live healthier for longer. Our medical researchers have been serving the community for more than 100 years, making transformative discoveries in cancers, infectious and immune diseases, developmental disorders, and healthy ageing. WEHI brings together diverse and creative people with different experience and expertise to solve some of the world's most complex health problems. With partners across science, health, government, industry, and philanthropy, we are committed to long-term discovery, collaboration, and translation. At WEHI, we are brighter together. Find out more at www.wehi.edu.au