

New technology 'retrains' cells to repair damaged brain tissue in mice after stroke

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Graduate research associate Jordan Moore reviews brain MRI images of mice in the nanomedicine lab at The Ohio State University College of Medicine. In a new study, researchers demonstrate the potential of a new cell therapy to reverse damage from ischemic stroke by regrowing blood vessels and healthy brain tissue. Credit: The Ohio State University Wexner Medical Center

Most stroke victims don't receive treatment fast enough to prevent brain damage. Scientists at The Ohio State University Wexner Medical Center, College of Engineering and College of Medicine have developed technology to "retrain" cells to help assistant professor of biomedical engineering and repair damaged brain tissue. It's an advancement that may someday help patients regain speech, cognition and motor function, even when administered days after an ischemic stroke.

Engineering and medical researchers use a process created by Ohio State called tissue nanotransfection (TNT) to introduce genetic material into cells. This allows them to reprogram skin cells to become something different—in this case vascular cells—to help fix damaged brain tissue.

Science Advances.

In this mouse study, cells were 'pre-conditioned' with specific genes and injected into the strokeaffected brains, where they promoted the formation of new blood vessels via reprogramming and the repair of damaged brain tissue.

"We can rewrite the genetic code of skin cells so that they can become blood vessel cells," said Daniel Gallego-Perez, an assistant professor of biomedical engineering and surgery at Ohio State who is leading the research. "When they're deployed into the brain, they're able to grow new, healthy vascular tissue to restore normal blood supply and aid in the repair of damaged brain tissue."

Researchers studied the process in mice and found that those treated with this innovative cell therapy regained 90% of their motor function. MRI scans showed damaged areas of the brain were repaired within a few weeks.

"We found that the mice have a higher recovery because the cells that are being injected into the affected area also release healing signals in the form of vesicles that help in the recovery of damaged brain tissue" said Natalia Higuita Castro, surgery at Ohio State and a co-lead author on the study.

Every 40 seconds, someone in the United States suffers a stroke. It's the second-leading cause of death worldwide, and those who survive often have irreversible brain damage resulting in paralysis, speech impairment and loss of motor function. No treatments exist to address the lasting and debilitating damage to brain tissue caused by stroke.

Although medical advancements have allowed Study findings published online today in the journal doctors to clear clots in the brain faster and



improve outcomes, this is only effective if done within a few hours of the stroke, before the brain tissue dies. About 80% of ischemic stroke patients don't receive the clot-busting therapy in time to prevent permanent deficits to their speech, cognition and motor function.

Provided by The Ohio State University



Researchers Natalia Higuita Castro, Ana Salazar Puerta and Jordan Moore helped develop a new cell therapy that reprograms cells to repair brain damage caused by ischemic stroke. Credit: The Ohio State University Wexner Medical Center

"The thought was that once <u>brain tissue</u> dies, that was it," said Dr. Shahid Nimjee, a neurosurgeon at Ohio State Wexner Medical Center, a member of Ohio State's Neurological Institute and co-author of the study. "We're now learning that there could be opportunities to regenerate <u>cells</u> to restore brain function."

Researchers continue to study this approach, and they're also exploring other potential uses for this technology to treat brain disorders such as Alzheimer's and autoimmune diseases.

More information: "Nanotransfection-based vasculogenic cell reprogramming drives functional recovery in a mouse model of ischemic stroke" *Science Advances* (2021).

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