

Scientists discover new way to help nerve regeneration in spinal cord injury

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There is currently no cure for spinal cord injury or treatment to help nerve regeneration so therapies offering intervention are limited. People with severe spinal cord injuries can remain paralysed for life and this is often accompanied by incontinence.

A team led by Drs Liang-Fong Wong and Nicolas Granger from Bristol's Faculty of Health Sciences has successfully transplanted genetically modified cells that secrete a treatment molecule shown to be effective at removing the scar following spinal cord damage. The scar in the damaged spinal cord typically limits recovery by blocking nerve regrowth

Previous work by the team proved olfactory ensheathing cells - which are taken from the 'smell system' where they regenerate and repair throughout life to maintain sense of smell, could be genetically modified to secrete a treatment enzyme known as chondroitinase ABC (ChABC). This treatment enzyme is key in breaking down the glial scar at the injury point of the spinal cord and helping to promote nerve regrowth.

However, while previous studies have shown

ChABC to be effective at promoting nerve regrowth when injected in experimental models of spinal cord injury as a drug treatment, it degrades rapidly at body temperature and repeated administration may be required to maintain efficacy.

In this study, researchers combined both treatments to treat rodents with spinal cord injury with genetically modified olfactory ensheathing cells to express ChABC. Following transplantation of the cells in rodent models the team were able to demonstrate the successful secretion of ChABC enzyme and removal of some of the glial scar. This led to increased nerve sprouting in the spinal cord, suggestive of successful nerve regeneration following the treatment.

The study provides an important proof-of-concept that this cell transplant strategy is a viable method to deliver this key ChABC enzyme in a rodent model of spinal cord injury and could be potentially used to allow the cells to be more efficient at repairing the spinal cord.

Dr Liang-Fong Wong from Bristol Medical School, said: "While these initial results look promising, in order to determine the longer-term survival of our genetically modified cells and assess functional recovery, such as recovery of walking or recovery of continence, we need to carry out further studies to test these cell transplants in more chronic injury models."

Dr Nicolas Granger from the Bristol Veterinary School added: "Taking this therapy further, for example by applying it to companion dogs that are naturally affected by spinal cord <u>injury</u> and remained paralysed, could help improve recovery of walking in these dogs in the longer term and pave the way for this approach to be applied to human <u>spinal cord</u> injuries."

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Research Council (BBSRC) and from the University's Elizabeth Blackwell Institute for health research.

More information: 'Transplantation of canine olfactory ensheathing cells producing chondroitinase ABC promotes chondroitin sulphate proteoglycan digestion and axonal sprouting following spinal cord injury' D Carwardine et al *PLOS ONE*.

Provided by University of Bristol

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