

## Scientists regenerate neurons that restore walking in mice after paralysis from spinal cord injury

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In a new study in mice, a team of researchers from UCLA, the Swiss Federal Institute of Technology, and Harvard University have uncovered a crucial component for restoring functional activity after spinal cord injury. The neuroscientists have shown that re-growing specific neurons back to their natural target regions led to recovery, while random regrowth was not effective.

In a 2018 study published in *Nature*, the team <u>identified a treatment</u> <u>approach that triggers axons</u>—the tiny fibers that link <u>nerve cells</u> and enable them to communicate—to regrow after spinal cord <u>injury</u> in rodents. But even as that approach successfully led to the <u>regeneration</u> of <u>axons</u> across severe spinal cord lesions, achieving functional recovery remained a significant challenge.

In a new study, published in *Science*, the team aimed to determine whether directing the regeneration of axons from specific neuronal subpopulations to their natural target regions could lead to meaningful functional restoration after spinal cord injury in <u>mice</u>. They first used advanced genetic analysis to identify nerve cell groups that enable walking improvement after a partial spinal cord injury.

The researchers then found that merely regenerating axons from these nerve cells across the spinal cord lesion without specific guidance had no impact on functional recovery. However, when the strategy was refined to include using chemical signals to attract and guide the regeneration of these axons to their natural target region in the lumbar spinal cord, significant improvements in walking ability were observed in a mouse model of complete spinal cord injury.



"Our study provides crucial insights into the intricacies of axon regeneration and requirements for functional recovery after <u>spinal cord</u> <u>injuries</u>," said Michael Sofroniew, MD, Ph.D., professor of neurobiology at the David Geffen School of Medicine at UCLA and a senior author of the new study. "It highlights the necessity of not only regenerating axons across lesions but also of actively guiding them to reach their natural target regions to achieve meaningful neurological restoration."

The authors say understanding that re-establishing the projections of specific neuronal subpopulations to their natural target regions holds significant promise for the development of therapies aimed at restoring neurological functions in larger animals and humans. However, the researchers also acknowledge the complexity of promoting regeneration over longer distances in non-rodents, necessitating strategies with intricate spatial and temporal features.

Still, they conclude that applying the principles laid out in their work "will unlock the framework to achieve meaningful repair of the injured spinal cord and may expedite repair after other forms of central nervous system injury and disease."

**More information:** Jordan W. Squair et al, Recovery of walking after paralysis by regenerating characterized neurons to their natural target region, *Science* (2023). <u>DOI: 10.1126/science.adi6412</u>. www.science.org/doi/10.1126/science.adi6412

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