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Prevention and Excision of Spinal Cord Scar Following Spinal Cord Injury Results in Motor Function and Spinal Cord Regeneration

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Abstract

Background: This paper shows originality for the treatment of acute spinal cord injuries by the absorption of fibrinogen by the omentum. This paper also shows that the statement by Ramon y Cajal that the reason spinal cords don't heal is that the scar that routinely occurs at the point of spinal cord injury must be removed in order for axons to progress into the distal spinal cord.

Case Presentation: The patient had a spinal cord injury that was reported by an MRI to have resulted in complete anatomical transection. Years after her injury, her spinal cord scar was removed and the gap in the spinal cord, which measured 4 cm (1.6 inches), was totally removed and the spinal cord defect was filled with collagen. Not only did the patient eventually walk after physiotherapy, confirmed by a video, but of extreme importance, serial MRIs presented in the paper showed that the spinal cord can spontaneously regenerate. I believe this is the first time that the spinal cord has been shown to have the ability to regenerate.

Conclusions: Removal of a spinal cord scar following an SCI can result in motor function in both animals and in a human, as demonstrated in this paper. In addition, it was learned that the spinal cord has the ability to regenerate.

Introduction

Damage to the Spinal Cord (SC) is one of the most daunting injuries a human can endure. Although improvement has occurred in the post-injury care of Spinal-Cord-Injured (SCI) patients, surgical techniques to improve acute and chronic SCI patients have been slow in advancing the remedy for this problem. Prevention of spinal cord scarring in acute SCI patients and the total removal of a spinal cord scar in chronic SCI patients appear to be required if clinical improvement is to occur.

Acute Spinal Cord Injury: Laboratory studies in cats showed that simple placement of an intact vascularized omental pedicle on an acute SCI within eight hours of injury resulted in the complete absence of a spinal cord scar (Figure 1). In contrast, when no omental pedicle was placed on an acutely injured SC, spinal cord scar routinely occurred [1,2] (Figure 2). Early omental placement on an acute spinal cord injury in animals clearly showed the prevention of a SC scar from developing. This was likely the result of the omentum's ability to absorb the edema fluid that routinely occurs following an SCI. Edema fluid contains large amounts of fibrinogen, and it is this fibrinogen that is the initiating agent in the rapid development of a spinal cord scar following an SCI. The omentum's absorption of fibrinogen in the edema fluid is likely the reason for the limitation of scarring [3–5]. If in the future the

omentum could be routinely placed on an SCI in humans shortly after injury, a decrease in spinal cord scarring might occur. This clinical suggestion merits further study.



Figure 1: Note absence of fibrosis and edema 30 days after SCI.



Figure 2: Note surface and circumferential fibrosis 30 days after SCI. Edema is still present.

Chronic Spinal Cord Injury: Ramon y Cajal, the acknowledged father of neuroscience, stated that the SC scar that develops after injury prevents axons from progressing through the scar and down into the distal spinal cord. He was the first to recognize that the goal in the treatment of a chronic spinal cord-injured patient must be the total removal of a spinal cord scar [6]. Accomplishing this treatment described by Cajal is now possible. A series of experiments was begun years ago to develop a method to remove a piece of spinal cord (comparable to removing an SC scar) to allow for restoration of distal motor function [7-12]. A 5 mm segment of the spinal cord in cats was excised at the T-9 level. In order to join the severed stumps of the divided SC, a lumbar vertebrectomy was performed. Proximal and distal vertebrae adjacent to the excised vertebral body were solidly fixed by the insertion of stainless-steel plates and screws. The firm fusion of the proximal and distal SC stumps was accomplished using a CO₂

laser set at 1–2 watts of energy. The laser beam was carefully directed under microscopic control to the cut edges of the proximal and distal SC stumps. The heat energy of the laser beam was known to penetrate only 1.2 mm into the spinal cord, thus avoiding injury to the deeply placed neuroanatomical tracts. After completing the ${\rm CO}_2$ fusion-welding process, a pedicled omental flap was wrapped around the transection site of the fused spinal cord in order to absorb axoplasmic fluid that had accumulated at the laser-treated site and also to increase omental vascularity to the cut edges of the divided SC. The final step in this surgical procedure was to immobilize the vertebral column by affixing two long stainless-steel plates on both sides of the dorsal spine processes with the plates extending above and below the SC transection site.

The experiment proved a limited success since several cats had return of Somatosensory-Evoked Potential (SEPs), but no cat had any return of motor function. The experiment was therefore discontinued since the procedure was difficult to perform (Figure 3) but, most important, it had no clinical significance.



Figure 3: A difficult operation that was unsuccessful and was discontinued.

A simple operation was then developed that involved the removal of a 3 mm-6 mm piece of spinal cord in cats followed by the insertion of semi-liquid collagen directly into the gap that separated the proximal and distal spinal cord stumps. An intact pedicled omental graft was then placed directly upon the collagen bridge. The reason a vascularized omental pedicle was placed directly on the collagen bridge was that axons traveling through (Figure 4) and exiting the collagen bridge (Figure 5) required an adequate blood supply. This vascularization of axons by the omentum was expected since it was known that placing the omentum on intact spinal cords resulted in a 58% increase in spinal cord blood flow and a 3-1 increase in blood vessel density [8,9]. This experimental operation resulted in seven of fifteen cats developing coordinated hind-forelimb walking ability (Figure 6). When the cats were autopsied, most of the collagen in the SC gap had disappeared, as the material is biodegradable, but the collagen bridge that remained was shorter in length than when it was originally inserted between the divided SC stumps at the time of surgery. This suggested that the spinal cord was in the process

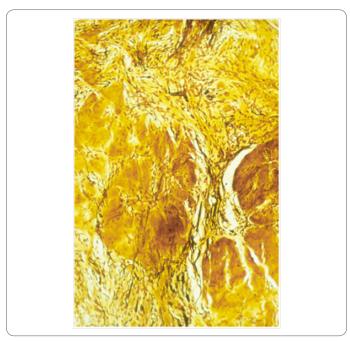


Figure 4: Axons seen in center of collagen bridge growing at 1 mm/day.

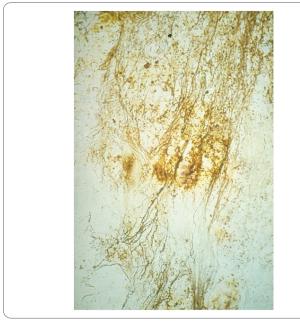


Figure 5: Axons exiting distal end of collagen bridge. Collagen is undergoing biodegradation.



Figure 6: Cats exhibiting hind-forelimb walking ability.

of longitudinal regeneration. This observation was later shown to have actually occurred by MRI studies in a patient later described in this paper.

Operative Procedures: Laboratories around the world have tried over the years to restore motor function following an SCI. The SC scar that routinely follows an SCI is the basis for the problem. It seems reasonable to believe that if this problem could be solved by complete removal of an SC scar, clinical improvement in spinal cord injuries might result.

Freeman Operation: L. Walter Freeman in 1954 was the first to report the removal of an SC segment that resulted in the return of motor function in the lower extremities of animals [10,11]. His work was of historic importance but was little appreciated. He showed that it is possible to remove a segment of the spinal cord followed by subsequent motor function in animals. This finding has now been shown to be not only possible in animals [2-5] but also in a human. Freeman removed a segment of the spinal cord to create a gap between the cut edges of the SC. This gap in the SC is closed by removing a vertebrae that shortens the spinal column. Removing a spinal cord scar in the Freeman operation requires a vertebrectomy which shortens the spinal column so that both cut ends of the divided spinal cord can be joined. If the gap in the divided SC is extensive, two vertebrae may be required. Canavero [11,12] and his associates have shown that the cut ends of a spinal cord following the complete transection of the spinal cord, could lead to motor function and electrical conductivity in animals when the cut ends of the divided cord were fused by Polyethylene Glycol (PEG) [13,14].

Omental-Collagen Operation: Both the Freeman operation and the omental-collagen procedure have been shown to restore motor function in animals following the removal of a spinal cord segment, but both operations have drawbacks. The Freeman operation requires the shortening of the spinal column by a vertebrectomy in order to close the gap in the SC after the excision of a spinal cord section. A shortcoming in the omental-collagen operation is that vascularizing the proximal and distal cut segments of the transected spinal cord by the omentum requires 1-11/2 hours of additional operative time. Although increased operating time is always undesirable, the vascularized omental pedicle application is critical as axons must have a significant blood supply as they transit through the collagen bridge into the distal SC. Whereas the Freeman and the omental-collagen operations have both shown restorative motor functions in animals following the removal of a segment of the spinal cord, the omental-collagen operation has also shown that the spinal cord can regenerate, as was confirmed by serial MRIs [10]. (See MRI study section.) As a matter of interest, the many patients who underwent the omental-collagen operation for a chronic spinal cord injury never developed an infection postoperatively. This was considered to be due to the antibacterial and anti-inflammatory material known to be present in omental tissue [15,16].

Case Presentation

The patient was a 24-year-old woman who had sustained a high-speed skiing accident on January 3, 1993. Magnetic Resonance Imaging (MRI) taken shortly after injury reported a "complete anatomical transection of her spinal cord at the T6-T7 level" (Figure 7).



Figure 7: MRI at time of injury reported total transection at T6-T7 level.

Over the next 314 years following her injury, she underwent no physiotherapy. However, in 1996, in the hope that she might regain an ability to walk, she underwent the omental-collagen operation. During this procedure, her spinal cord scar was totally removed, resulting in a large spinal cord gap that measured 1.6 inches (4 cm) between the proximal and distal spinal cord segments. Semi liquid collagen (5 cc) was inserted directly into the gap and an intact vascularized omental pedicle was placed on the collagen bridge. (Operative Photos in Addendum). The operation was uneventful as was her post-operative course. Eight months after surgery, the patient reported that in a swimming pool under direct command she could move her legs and ankles. The slight movement was confirmed by a video. At this time, the patient should have undergone extensive physiotherapy, but unfortunately this was not done. Five years after her surgery, she learned of the Neuro Institute in Arizona, which she entered in 2001 to begin intensive physical therapy. She spent one year in Arizona and showed great promise in functional areas as described by her physiotherapist in his letter (see addendum). She then returned to Germany for a year, but returned to the United States a year later. During this latter period in Arizona, she regained the ability to walk as filmed in the video. The patient was progressing satisfactorily until she was

informed that her visa to the United States was not to be extended and that she would have to leave the US. As reported by an unelected bureaucrat in Washington, the reason for this decision was that she came to the United States to recapture her ability to walk, and now that she was able to walk she would have to return to Germany. There was no consideration that she needed another year of physiotherapy in order to regain her ability to walk at a normal pace without a walker. Although great effort was made to reverse this decision, she was nevertheless forced to leave the country. On her return to Germany, she resumed her former employment, which required repeated movement in her office. Because of her slow walking ability, she could not perform her job duties and it became necessary for her to resolve the problem by returning to her wheelchair for faster mobility. It was tragic that authorities in Washington did not understand how important it was for this patient to remain in the United States for another year of physiotherapy.

MRI Studies: Following her injury in 1993, the patient had her first MRI which reported complete anatomical transection at the T6–T8 level (Figure 7). Following her surgery in 1996, she underwent further MRIs. Her first post-operative MRI, obtained one year after surgery, demonstrated a mass of soft tissue at the site of her SC injury (Figure 8).



Figure 8: One year after surgery, MRI showed a mass of soft tissue at SCI site.

The exact location of her spinal cord injury was always known because of the absence of the vertebral disc at the T6-T7 level which had been extruded at the time of her injury. Three years after surgery, her MRI showed the previous soft-tissue mass in her spinal cord was forming into a longitudinal structure (Figure 9). This strongly suggested that the spinal cord was gradually regenerating. By the 5th year after her surgery, her last MRI showed that the spinal cord had regenerated and that the previously divided proximal and distal segments of the divided spinal cord were in contact (Figure 10).



Figure 9: MRI three years after surgery, soft tissue mass appears to be developing into a longitudinal structure at the SC transection site.

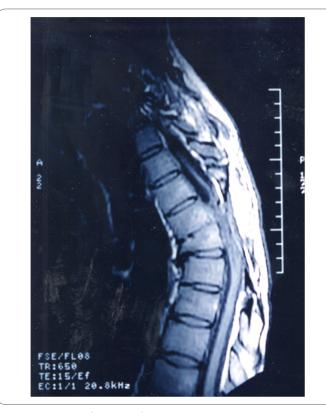
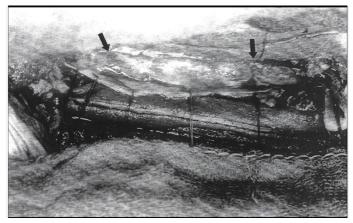


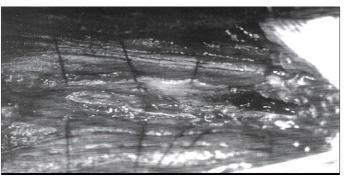
Figure 10: Final MRI five years after surgery shows regenerated SC connecting proximal and distal previously divided spinal cord segments.

Addendum - Operative Photos

Operative Photos of Patient's Omental-Collagen Procedure



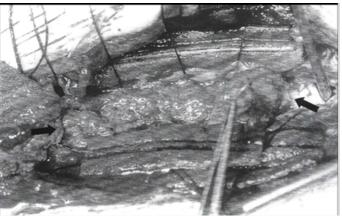
Arrows point to extensive spinal cord scar between T-6 and T-8



Photograph showing complete absence of spinal cord between T-6 and T-8



Photo showing collagen filling extensive spinal cord defect between T-6 and T-8.



Arrows showing pedicled omentum overlying collagen bridge between T-6 and T-8

Conclusion

Both the Freeman operation and the omental-collagen operation have improved motor function in animals after removing a SC segment of the spinal cord. The Freeman operation closes the gap in the divided SC by shortening the spinal column by a vertebrectomy, whereas the omental-collagen procedure fills the SC gap with collagen. Both operations have been successful in a variety of animals; however, the omental-collagen procedure has confirmed by serial MRIs that the spinal cord can regenerate in a human and lead to functional improvement. It is hoped this information will stimulate researchers and clinicians to expand these clinical findings for the future improvement of spinal cordinjured patients.

References

- 1. Goldsmith HS. Can the standard treatment of acute spinal cord injury be improved? Perhaps the time has come. Neurol Res. 2007;29(1):16–20.
- Goldsmith HS. Treatment of acute spinal cord injury by omental transposition: A new approach. J AM Coll Surg. 2009;208(2):289–292.
- 3. Goldsmith HS. Fibrinogen absorption by omentum: can it improve acute spinal cord injuries. Am J Neuroprot Neuroregen. 2016;8:54–57.
- Schachtrup C, Ryu JK, Helmrick MJ, Vagena E, Galanakis DK, Degen JL, et al. Fibrinogen triggers astrocyte scar formation by promoting the availability of active TGF-beta after vascular damage. J Neurosci. 2010;30(17):5843–5854.
- Schachtrup C, Lu P, Jones LL, Lee JK, Lu J, Sachs BD, et al. Fibrinogen inhibits neurite outgrowth via beta 3 integrinmediated phosphorylation of the EGF receptor. Proc Natl Acad Sci.USA. 2007;104(28):11814–11819.
- 6. Cajal RY. Degeneration and regeneration of the nervous

- systems (1909). New York. 1950;749-750.
- 7. Goldsmith HS. Acute spinal cord injuries: A search for functional improvement. Surg Neurol. 1999;51(2):231–233.
- 8. Goldsmith HS. Application of the omentum to the brain and spinal cord (chapter 3) In: Goldsmith HS, editor. The omentum: basic research and clinical applications. Woodbury, CT: Cine-Med Publishing; 2010;37–52.
- Goldsmith HS. Omental transposition and spinal surgery: Emphasis on revascularization and scar prevention. In: Spinal Surgery: Techniques, Complication Avoidance and Management. Ed: Edward C Benzel, Elsevier Saunders, Philadelphia; 2012;121:1183–1187.
- 10. Freeman LW. Return of spinal cord function in mammals after transecting lesions. Ann NY Acad Med Sci. 1954;58:564–569.
- 11. Freeman LW. Experimental observations upon axonal regeneration in the transected spinal cord of mammals. Clin Neurosurg. 1962;8:294–319.
- Canavero S. HEAVEN: The head anastomosis venture project outline for the first human head transplantation with spinal linkage (GEMINI). Surg Neurol Int. 2013;4(Suppl 1):S335– S342
- 13. Canavero S, Ren X, Kim CY, Rosati E. Neurologic foundations of spinal cord fusion (Gemini). Surgery. 2016;160(1):11–19.
- Ren S, Liu Z, Kim CY, Fu K, Wu Q, Hou L, et al. Reconstruction of the spinal cord of spinal transected dogs with polyethylene glycol. Surg Neurol Int. 2019;10:50.
- 15. Goldsmith HS. Naturally occurring antibacterial substance in human omentum. Surg Rounds. 2005;28:262–265.
- Chandra A, Srivastava RK, Kashyap MP, Kumar R, Srivastava RN, Pant AB. The anti-inflammatory and antibacterial basis of human omental defense: selective expression of cytokines and antimicrobial peptides. PLoS One. 2011;6(5):e20446.