

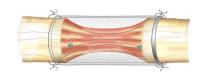
HOW NERVE CONDUITS WORK.

Peripheral nerves have the ability to effectively regenerate if given the proper environment. Autografts and allografts provide a multitubular internal structure and scaffolding that supports and physically guides axonal regeneration.¹ However, nerve conduits are hollow and provide only gross guidance.^{1,2,3,4} Regeneration through a conduit relies on the formation of a fibrin cable. This rudimentary structure, not the conduit itself, is what provides the physical support that makes regeneration possible. The process is shown below:



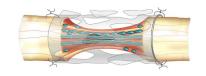
HOURS

Fluid seeps from the nerve ends into the void of the conduit.



DAYS

An hourglass-shaped fibrin cable forms. The regeneration potential is dependent upon the presence, integrity, and cross-sectional area of this fibrin cable.



MONTHS

Cell migration and axonal regeneration occurs within the cable and is restricted by the thinnest portion.



YEARS

Often the resulting tissue is visibly thinner, containing a limited number of regenerated axons.



CLINICAL EXAMPLE

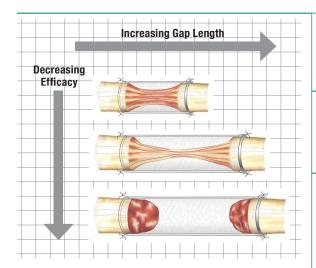
A thinning nerve cable seen in a 10mm gap previously repaired with a conduit.

Image courtesy of Jonathan Isaacs, MD Virginia Commonwealth University Health System

)

THE LENGTH LIMITATION OF CONDUITS.

Regeneration within a conduit occurs predominantly through the fibrin cable. The integrity of the fibrin cable is a function of the conduit dimensions. ^{5,6} The illustrations below demonstrate what happens as the length of the gap increases.



At short gap lengths, the fibrin cable is robust enough to allow regeneration.

As the gap length increases, the integrity of the fibrin cable diminishes and thinning restricts the regenerative space.

If the cable does not form, axons are not able to cross the gap. This results in no regeneration or possibly a neuroma.



CLINICAL EXAMPLE

Referred patient where multiple conduits failed in 18mm gaps.

- * Denotes thinned atrophic nerve tissue.

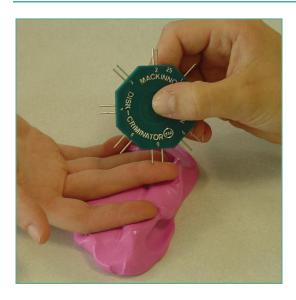
Image courtesy of Bauback Safa, MD The Buncke Clinic

CLINICAL OUTCOMES WITH CONDUITS.

Landmark clinical studies have examined the efficacy of commercially available conduits. As summarized below, results in gaps ranging from 0-5mm demonstrate that conduits can be a successful nerve repair option. Results using conduits in gaps greater than 5mm are highly variable and less reliable.

Type of Nerve	Gap Length	% Failure*	Other Findings	Clinical Publication	
Digital Nerves	0-4 mm	0%	34% failure rate	Weber et al., 2000 ⁷	
(Sensory)	5-7 mm	39%	in gaps 5mm		
	8-25 mm	29%	or greater		
Digital Nerves (Sensory)	6-18 mm	25%	100% of gaps > 16mm failed	Lohmeyer et al., 20098	
Digital Nerves (Sensory)	5-30 mm	14%	27% reported poor resolution of pain	Mackinnon and Dellon, 1990 ⁹	
Sensory, Mixed and Motor Nerves**	2.5-20 mm	57%	31% required revision***	Wangensteen and Kalliainen, 2009 ¹⁰	

^{*}No or poor sensory recovery as defined by the MRCC scale



CLINICAL EXAMPLE

Sensory nerve assessment of 2 point discrimination.



^{**}Mostly sensory nerves in the upper extremity, but also includes mixed and motor nerves elsewhere in the body
***When quantitative measurements were made

SUMMARY

A clear understanding of nerve regeneration through conduits and the inherent limitations is essential in considering treatment options. Conduits offer advantages, including off-the-shelf availability, alleviation of tension¹¹, and reduced potential for fascicular mismatch.^{12,13,14} They can be effective at short gaps (typically 5mm or less).⁷ However, there is a higher risk of conduit failure as gap length increases due to reliance upon the fibrin cable.^{7,8,9,10,15}

Clinical studies have shown that gaps greater than 5mm have:

- · High failure rates
- · Limited pain resolution
- · High revision rates

The strengths and limitations of conduits should be considered when evaluating treatment options for peripheral nerve injuries.

REFERENCES

- Lundborg G. 2000. A 25-year perspective of peripheral nerve surgery: evolving neuroscientific concepts and clinical significance. J Hand Surg Am 25(3):391-414.
- Dahlin LB, Lundborg G. 2001. Use of tubes in peripheral nerve repair. Neurosurg Clin N Am 12(2):341-352.
 Taras JS, Jacoby SM. 2008. Repair of lacerated peripheral nerves with nerve conduits. Tech Hand Up Extrem Surg 12(2):100-106.
- Lundborg G, Gelberman RH, Longo FM, Powell HC, Varon S. 1982. In vivo regeneration of cut nerves encased in silicone tubes: growth across a six-millimeter gap. J Neuropathol Exp Neurol 41(4):412-422.
- Moore AM, Kasukurthi R, Magill CK, Farhadi HF, Borschel GH, Mackinnon SE. 2009. Limitations of conduits in peripheral nerve repairs. Hand (N Y) 4(2):180-186.
- Zhao Q, Dahlin LB, Kanje M, Lundborg G. 1993. Repair of the transected rat sciatic nerve: matrix formation with implanted silicone tubes. Res Neurol and Neurosci 5: 197-204.
 Weber RA, Breidenbach WC, Brown RE, Jabaley ME, Mass DP. 2000. A randomized prospective study
- Weber HA, Breidenbach WC, Brown HE, Jabaley ME, Mass DP. 2000. A randomized prospective study
 of polyglycolic acid conduits for digital nerve reconstruction in humans. Plast Reconstr Surg 106(5):1036-1045.
 Lohmeyer JA, Siemers F, Machens HG, Mailander P. 2009. The clinical use of artificial nerve conduits for
- digital nerve repair: a prospective cohort study and literature review. J Reconstr Microsurg 25(1):55-61.
- 9. Mackinnon SE and Dellon AL. 1990. Clinical nerve reconstruction with a bioabsorbable polyglycolic acid tube. Plast Reconstr Surg 85(3):419-424.

 10. Wasserbase K. J. Killinian K. 2000. Callege Tube Conduits in Parishers Nerve Repair A Patropactive.
- Wangensteen KJ, Kalliainen LK. 2009. Collagen Tube Conduits in Peripheral Nerve Repair: A Retrospective Analysis. Hand (N Y).
- Schmidhammer R, Zandieh S, Hopf R, Mizner I, Pelinka LE, Kroepfl A, Redl H. 2004. Alleviated tension at the repair site enhances functional regeneration: the effect of full range of motion mobilization on the regeneration of peripheral nerves--histologic, electrophysiologic, and functional results in a rat model. J Trauma 56(3):571-584.
- Evans PJ, Bain JR, Mackinnon SE, Makino AP, Hunter DA. 1991. Selective reinnervation: a comparison of recovery following microsuture and conduit nerve repair. Brain Res 559(2):315-321.
- Hasegawa J, Shibata M, Takahashi H. 1996. Nerve coaptation studies with and without a gap in rabbits. J Hand Surg [Am] 21(2):259-265.
 The pictulogical analysis of higherical conduit.
- Zhang P, Zhang C, Kou Y, Yin X, Zhang H, Jiang B. 2009. The histological analysis of biological conduit sleeve bridging rhesus monkey median nerve injury with small gap. Artif Cells Blood Substit Immobil Biotechnol 37(2):101-104.
- Meek MF, Coert JH. 2002. Clinical use of nerve conduits in peripheral-nerve repair: review of the literature. J Reconstr Microsurg 18(2):97-109.

AXOGEN'S SOLUTIONS FOR REPAIRING NERVE GAPS.





Designed for more reliable results:

- Unique length for bridging gaps up to 5mm
- Semi-translucent for visualization of the nerve

0 mm	5 mm	10 mm	15 mm	20 mm	25 mm	30 mm	35 mm	40 mm	45 mm	50 mm





Provides multi-tubular structure:

• Decellularized nerve allograft v

- Decellularized nerve allograft with preserved extracellular matrix
- Clean and clear pathways allow axon growth

To Place an Order, Contact:

AxoGen Customer Service
Phone: 888-AXOGEN1 (888-296-4361)
Fax: 386-462-6801
Email: customerservice@axogeninc.com
www.axogeninc.com

AxoGuard® Manufactured By:

Cook Biotech Incorporated 1425 Innovation Place West Lafayette, Indiana 47906 U.S.A. Phone: 765-497-3355 Toll Free: 888-299-4224 Fax: 765-497-2361



13859 Progress Blvd., Suite 100, Alachua, Florida 32615

©2010 AxoGen, Inc.. Avance®, AxoGuard® Nerve Protector and AxoGuard® Nerve Connector and their logos are trademarks of AxoGen, Inc.