

INTRODUCTION

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20 Nerve coaptation is important for successful nerve reconstruction. Different
21 techniques for enhancement of nerve regeneration after microsurgical nerve repair
22 have been described previously (1-9). Significant loss of nerve tissue necessitates the
23 use of a nerve graft, while nerve regeneration through a large peripheral nerve gap
24 will not be successful unless a conduit is used. Autogenous nerve grafting is the most
25 commonly used procedure for nerve repair when a significant nerve gap is present. It
26 generally provides good results, but it is hampered by donor site morbidity and
27 limited availability (10-13). Clinical implementation of nerve conduits has focused on
28 the use of autogenous tissue (veins, arteries, pseudosheaths, nerve grafts) and
29 occasionally on artificial conduits (silicone, polyglactine mesh) (10,11,14). According
30 to current literature, conduit materials do not seem to significantly improve clinical
31 outcome when compared with conventional nerve grafting (15,16). A major difficulty
32 in the use of conduits is the defect size that can be successfully bridged; in humans,
33 this is approximately 25 mm. For larger defects, nerve grafting is necessary.

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The epineurium may serve as an autologous biologically active conduit and
35 may facilitate bridging of nerve defects. The use of the epineural cuff technique was
36 first described by Snyder et al. (17,18). This technique has been validated in animal
37 models (19-23). We have previously presented the outcomes of using an epineural
38 flap in a patient with a short radial nerve defect after excision of a neuroma-in-
39 continuity, with excellent functional results after a follow-up of 17 months (24). The
40 motor recovery of the wrist, finger, and thenar extensor muscles was measured as M4
41 (excellent), whereas sensory recovery at the area of distribution the superficial radial
42 nerve over the second metacarpal was S3 (satisfactory). This article reports the
43 successful outcome of a 32-year-old patient who experienced complete transection of

44 the median nerve at the distal forearm treated with the epineural sleeve graft
45 reconstruction technique. The patient provided written informed consent for print and
46 electronic publication of this case report. The authors report no actual or potential
47 conflict of interest in relation to this article.

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CASE REPORT

50 A 32-year-old male blue-collar worker presented with a median nerve transection at
51 the distal third of his right forearm after a fall on broken glasses. Surgical exploration
52 of the wound was done 3 hours after the injury. Exploration showed complete
53 transection of the median nerve and radial artery, and the tendons of the flexor
54 digitorum superficialis (FDS) and flexor digitorum profundus (FDP) to the index,
55 middle and ring fingers, and the flexor carpi radialis (Figure 1). The nerve was cut at
56 2 close areas and contused. Microsurgical debridement was done that resulted in a
57 median nerve gap of 14 mm.

58 The tendons were repaired in a deep-to-superficial method with a 4-0 ethibond
59 suture using a modified Kessler technique; tendons' repair was reinforced with a 6-0
60 prolene continuous epitendinous suture. Next, the radial artery was repaired using 9-0
61 nylon interrupted sutures. The injury to the median nerve was then addressed. Care
62 was taken to avoid extensive iatrogenic injury to the epineurium. After microsurgical
63 debridement, the median nerve stumps were approximated with 10-0 nylon sutures,
64 leaving a 10 mm nerve defect.

65 The epineurium was then incised longitudinally proximally and distally,
66 creating two 1.5-cm-long epineural flaps, one for each end across the gap to bridge
67 (Figure 2). The epineurium was plicated serving as an additional mechanical aid to
68 bridge the nerve gap, thus reducing nerve tension. The flaps were sutured to the

69 epineurium of the other side and finally to each other with side-to-side stitches. The
70 remaining gap between the nerve stumps was filled with a blood clot derived from the
71 patient's own blood, as previously reported (19,20).

72 Postoperatively, the patient's arm was immobilized in a long arm splint in 30
73 degrees of wrist flexion and 40–60 degrees of flexion at the metacarpophalangeal
74 (MCP) joints, with the interphalangeal joints allowed to extend fully. Wrist position
75 was changed on a weekly basis, with a gradual return to a neutral position. The
76 postoperative rehabilitation protocol included active finger extension and passive
77 finger flexion during the first 4 weeks. Then, at 4 weeks, the splint was removed and
78 the patient was allowed protected early range of motion exercises. In addition,
79 tendon-gliding exercises were initiated, and resumption of light activity of daily living
80 was encouraged. The patient was informed on the necessity to protect the limb during
81 the nerve recovery period, and returned to work 8 months after initial trauma. A
82 sensory re-education program was performed in two phases, according to Dellon's
83 recommendations (25).

84 Follow-up examination was done every 2 months for the first 1 year and at the
85 last follow-up, at 18 months, for the purpose of the present study. Clinical evaluation
86 was done using the protocol described by Rosén and Lundborg (26); analysis was
87 conducted at three levels: sensory domain (sensory innervation, tactile gnosis, and
88 finger dexterity), motor domain (motor innervation and grip strength), and pain/
89 discomfort domain (hyperaesthesia and cold intolerance) (Table). The quotient in
90 each domain was calculated by dividing the obtained results with the normal outcome.
91 Every domain was assigned a mean score and a total score. Total scores ranged from
92 0 to 3, with each domain contributing 1 point. This protocol has been developed for
93 routine documentation and quantification of functional outcome following nerve

94 repair at the wrist or distal forearm level, including sensory and motor function as
95 well as evaluation of postoperative pain and discomfort in a summarized scoring
96 system (26). It is simple and flexible and includes a numerical scoring system for
97 clinical and scientific use (23). The calculated total score correlates strongly with the
98 patients' subjective opinion about how much the nerve injury influenced their
99 activities of daily living (26).

100 At the last follow-up, the patient experienced complete sensory and motor
101 function of the median nerve (Figure 3). No further operations, such as neurolysis or
102 tenolysis were necessary until the period of this study; rupture of the tendons repair
103 was not observed. Overall sensory recovery was classified as good (grade S3+)
104 according to the modified classification of the British Medical Research Council (27),
105 with a total numerical score of 2.3, based on the Rosén and Lundborg protocol (26).

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DISCUSSION

108 The idea of using the epineurium to bridge a nerve defect is not new (9,19-23).
109 Several experimental studies in animals supported the idea of using the epineurium in
110 nerve reconstructive surgery (19-23). The general idea of the epineural sleeve in nerve
111 repair is to use epineurium of the nerve stump to cover the coaptation site. Coaptation
112 site becomes separated from the surrounding tissues, thus providing optimal
113 environment for nerve regeneration. The epineural sleeve provides a favorable nerve
114 regeneration environment; it prevents protrusion of fascicles out from the suture line
115 and prevents nerve from contact with scar or fibrotic tissues around coaptation site
116 facilitating axonal regrowth (9). The interposed epineurium is not just a passive
117 conduit. It assumes a more active role in the nerve regeneration process by providing
118 neurotrophic factors. Additionally, the axoplasmic fluid is retained at the repair site

119 and this facilitates the regeneration process (22). Siemionow et al. (22) mentioned that
120 the epineural sleeve provides a biological chamber at the coaptation site where
121 axoplasmic fluid is collected, leaving a perfect milieu for regeneration of the nerve.
122 They were able to collect as much as 1 ml of fluid from the transection site in a rabbit
123 sciatic nerve model (3). The collected fluid, when administered subepineurally after
124 nerve repair, enhanced the regeneration process, as confirmed by the increased
125 number of axons and myelinated fibers (3). The authors concluded that preservation
126 of axoplasmic fluid at the repair site may have a potentially beneficial effect on
127 peripheral nerve regeneration and functional recovery. The studies by Lundborg et al.
128 (28) and Longo et al. (29) showed that the fluid collected from the divided peripheral
129 nerve stumps contained neurotrophic factor activity for several neural types, including
130 sensory, motor, and sympathetic neurons. In our practice and the present patient, we
131 used an autologous blood clot to fill the gap between the nerve stumps based on our
132 previous animal studies for lumen collapse prevention (19,20). Instead of an
133 autologous blood clot, neurotrophic factors could also be added into the epineural
134 sleeve to enhance nerve regeneration, if necessary, based on the surgeons'
135 preferences. Additionally, based on this experimental work, we have seen that the
136 epineural sleeve technique may be useful for bridging nerve gaps up to 25 mm.
137 However, since this cannot be supported from the present case report, we can only
138 recommend this technique to repair nerve gaps up to 14 mm.

139 The epineural sleeve nerve reconstruction technique has several advantages
140 compared to other techniques: a conduit of neural origin is used; no separate surgical
141 exposure for harvesting is necessary; there is no donor-recipient size mismatch; the
142 biocompatibility is perfect; there is no antigenicity or inflammatory reaction; and the
143 cost of harvesting is negligible. The disadvantages of this technique include: technical

144 difficulties in surgical manipulation of the epineurium, time-consuming harvesting
145 procedure, and limited amount of graft material. A concern is that stripping of the
146 epineurium might influence nerve electrophysiologic properties. However,
147 Karacaoglu et al. (21) noted that denuding of the sciatic nerve from the epineurium in
148 the rat did not induce significant any significant nerve function. In a study by Tetik et
149 al. (23), three methods of nerve repair involving the epineural sleeve technique were
150 compared with conventional nerve repair using the rat sciatic nerve transection model
151 in four groups. In group 1, the sciatic nerve was repaired using the conventional
152 epineural technique by placing four sutures. In groups 2, 3, and 4, the epineural sleeve
153 technique was combined with two sutures, fibrin glue, and two sutures with fibrin
154 glue, respectively. Functional recovery was evaluated using walking track analysis,
155 limb circumference, and toe contracture severity. The results showed better functional
156 recovery and a higher number of myelinated fibers in groups using the epineural
157 sleeve technique compared with conventional techniques. The addition of fibrin glue,
158 however, did not make any difference. The epineural sleeve technique was found to
159 be superior when compared with conventional nerve repair, providing faster
160 functional recovery and improved nerve regeneration. Yavuzer et al. (30) investigated
161 the effects of the turnover epineural sheath tube (TEST) when used over the primary
162 nerve repair site to improve nerve regeneration in twenty-five Wistar rats. Three
163 months postoperatively, functional analysis and nerve and muscle histomorphometric
164 studies revealed similar results in the primary repair and TEST groups, with on
165 statistically significant difference. However, during microscopic examination, a
166 decrease in both foreign material reaction and an inflammatory response with less
167 fibrosis were observed in the TEST group. The authors noted that the TEST had
168 nerve-healing properties similar to primary epineural repair, with the advantage of a

169 reduced number of sutures, thus decreasing fibrosis around the repair site. The authors
170 concluded that the TEST is a good alternative treatment modality, especially for
171 polyfascicular peripheral nerves.

172 In conclusion, the use of the epineural sleeve technique may be useful for
173 bridging short nerve defects, up to 14 mm with excellent results, as the report has
174 shown. The epineurium serves as a mechanical aid to reduce gap size, and increase
175 repair strength, effectively assisting nerve regeneration.

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LEGENDS

260

FIGURES

261 **Figure 1.** Intraoperative photograph shows complete transection of the median nerve.

262 **Figure 2.** Intraoperative photograph of the sutured nerve stumps covered with the
263 epineural sleeve.

264 **Figure 3.** Clinical photographs at the 18-month follow-up. The patient was able to
265 make **(A)** “OK” sign, **(B)** normal opposition, and **(C)** palmar abduction of the thumb.

266

267

TABLE

268 **Table.** The numerical test protocol and numerical system based on the clinical results
269 at the 18-month follow-up.