"Type grouping" in skeletal muscles after experimental reinnervation

G. Karpati, M.D., and W. K. Engel, M.D.

Application of histochemical techniques to the study of human muscle biopsies has helped to elucidate a number of pathologic reactions of skeletal muscle that are not evident in routinely stained paraffin-embedded tissue.[1–4] One of these abnormalities has been termed "type grouping" and defined as "groups of fibers of the same histochemical type much larger than usual . . . consisting of 50-100 fibers or more."[2] Normally, in histochemically mixed muscles, the two basic histochemical fiber types (types 1 and 2) are intermingled in a mosaic pattern when viewed in cross section. In normal human muscle the number of fibers of similar histochemical type in tight contiguity does not exceed 50 and is usually less than 15. Type grouping is commonly observed in muscle biopsies of patients with chronic peripheral neuropathies (Fig. 1) and somewhat less often in the advanced stage of motor neuron disease. It has been postulated to be a sign of reinnervation by "collateral sprouting."[2]

In this paper we present an experimental model in which typical type grouping was produced in histochemically mixed muscles of the guinea pig by experimental reinnervation. The results support the suggestion that collateral sprouting is at least one pathogenic mechanism which may be responsible for type grouping in disease in human beings.

Materials and Methods

In adult (3-month-old), male albino guinea pigs, experimental reinnervation of the gastrocnemius and plantaris muscles was produced on one side of each animal. All experimental procedures were done on animals under intraperitoneal pentobarbital sodium anesthesia (25 mg. per kilogram). The three methods used for experimental reinnervation are shown in Figure 2. In group A (12 animals) the lower one-third of one sciatic nerve was exposed from a posterior thigh approach by blunt dissection of the hamstring muscles and the exposed nerve was crushed with uniform force for ten seconds. The crushing instrument was a mosquito forceps on which the serrated jaws were enveloped in polyethylene tubing. At ten and twelve weeks after the procedure, these animals were sacrificed with an overdose of barbiturate. In group B (6 animals) one sciatic or posterior tibial nerve was exposed at the region of the sciatic nerve bifurcation as described above and with the tip of a fine jeweler's forceps the nerve was crushed vigorously at three separate places about 2 mm. apart. Ten days later the same procedure was repeated on the same nerves at a slightly more distal portion of the nerve. These animals were sacrificed six months after the second procedure. In group C (6 animals) the sciatic nerve bifurcation was exposed in the popliteal fossa and the sciatic or posterior tibial nerve was completely severed just above or below the bifurcation. The severed ends were carefully approximated by suturing of the nerve sheath with two 7-0 silk sutures. These animals were sacrificed at four to six months after the operation.

Two types of controls were used: [1] the contralateral unoperated or sham-operated side of the experimental animals and [2] completely normal adult male guinea pigs.

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