Fibrin conduit supplemented with human mesenchymal stem cells

and immunosuppressive treatment enhances regeneration after

peripheral nerve injury.

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Abstract:

To address the need for the development of bioengineered replacement of a nerve graft, a novel

two component fibrin glue conduit was combined with human mesenchymal stem cells (MSC) and

immunosupressive treatment with cyclosporine A. The effects of MSC on axonal regeneration in the

conduit and reaction of activated macrophages were investigated using sciatic nerve injury model. A

10mm gap in the sciatic nerve of a rat was created and repaired either with fibrin glue conduit

containing diluted fibrin matrix or fibrin glue conduit containing fibrin matrix with MSC at

concentration of 80x10(6) cells/ml. Cells were labeled with PKH26 prior to transplantation. The

animals received daily injections of cyclosporine A. After 3 weeks the distance of regeneration and

area occupied by regenerating axons and ED1 positives macrophages was measured. MSC

survived in the conduit and enhanced axonal regeneration only when transplantation was combined

with cyclosporine A treatment. Moreover, addition of cyclosporine A to the conduits with transplanted

MSC significantly reduced the ED1 macrophage reaction.

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