

The Adaptive Immune System Has a Critical Role in Promoting Nerve Regeneration across Acellular Nerve Allografts

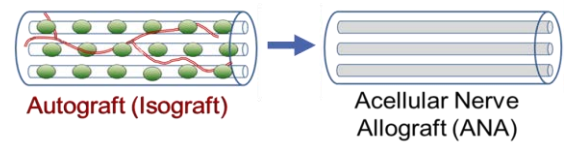
Matthew D. Wood, Deng Pan, Daniel A. Hunter, Lauren Schellhardt, Anja G. Fuchs, Alexandra E. Halevi, Alison K. Snyder-Warwick, Susan E. Mackinnon



PNSRL.WUSTL.EDU
Email: woodmd@wustl.edu

Introduction

• Due to disadvantages of nerve autografting, alternatives are being increasingly used and desired for nerve defect repair.

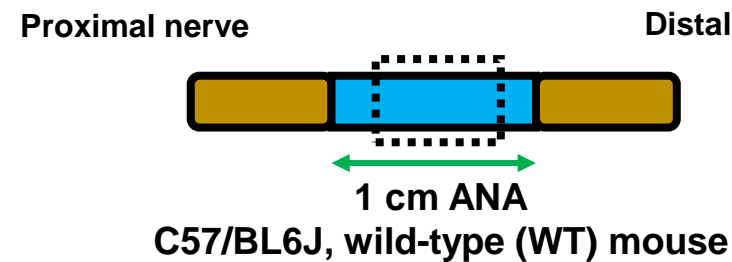


• These alternatives have limits for the repair of nerve defects. We still do not understand why these limits are present, or how to overcome these limits.

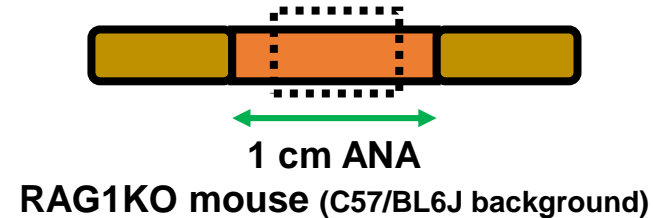
• We determined whether the adaptive immune system (T & B cells) has a role in nerve regeneration across initially acellular nerve autograft alternatives.

Methods

- Acellular nerve allografts (ANAs) were generated using a chemical detergent protocol.
- One (1) cm ANAs were used to repair sciatic nerve gaps in Rag1KO (lacking both T and B cells) or wild-type (WT; control) mice.
- Regeneration was quantified using histology, immunofluorescence, and a grid walk test.

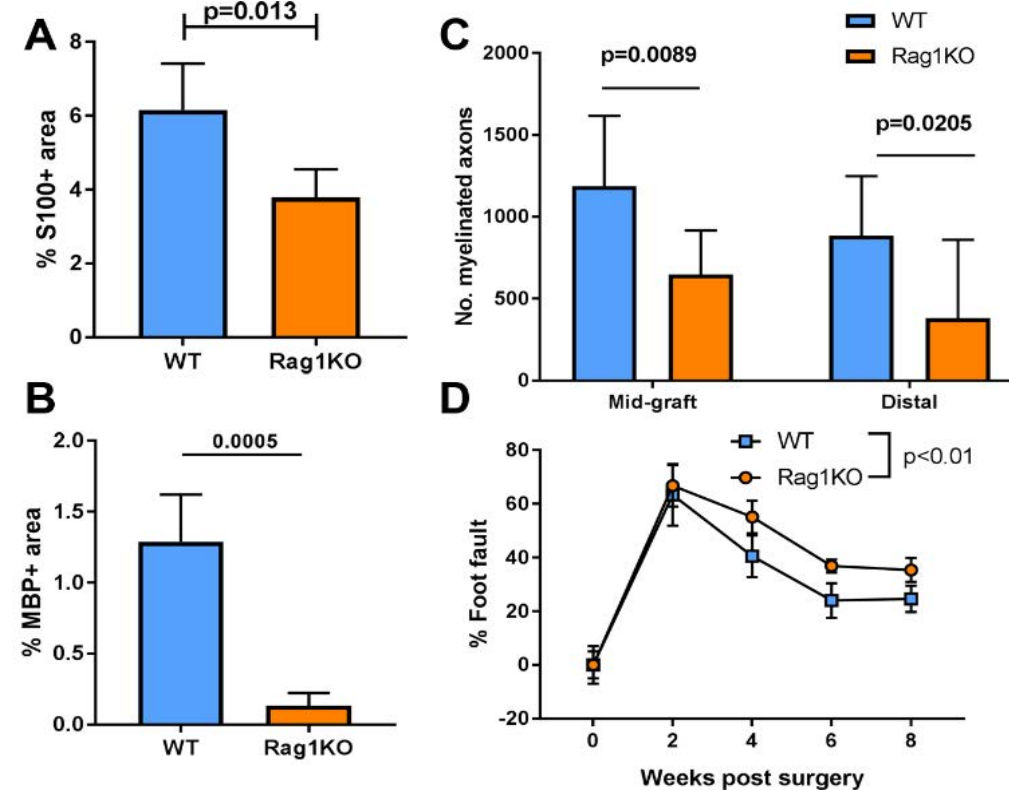


T cell sufficient

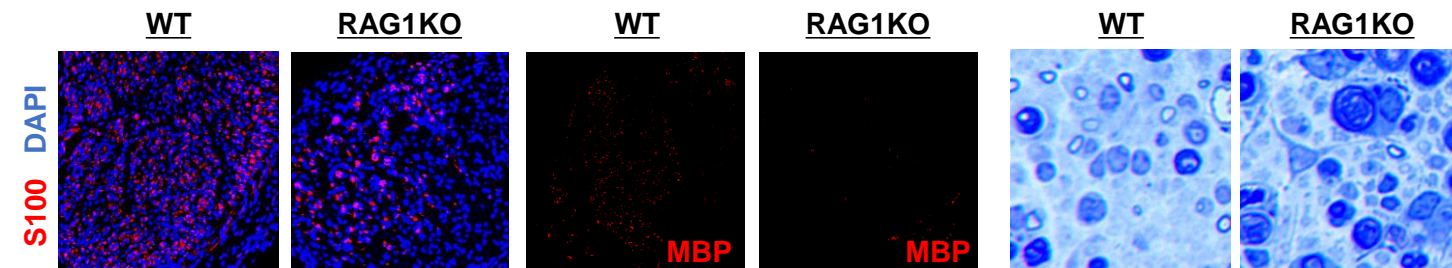


T and B cell deficient

Results



WT vs Rag1KO mice were assessed for nerve regeneration after ANA repair. After 2 weeks, Rag1KO Schwann cell (S100) quantities and myelin basic protein (MBP) within ANAs was reduced compared to WT (A, B). At 4 weeks, Rag1KO had reduced numbers of myelinated axons compared to WT (C). Starting from 4 weeks, Rag1KO had a persistent increase in foot faults compared to WT (D). Mean ± SD, n≥5/group; p values shown. Images below.



Conclusions

- This data suggests adaptative immune cells promote regeneration across ANAs.