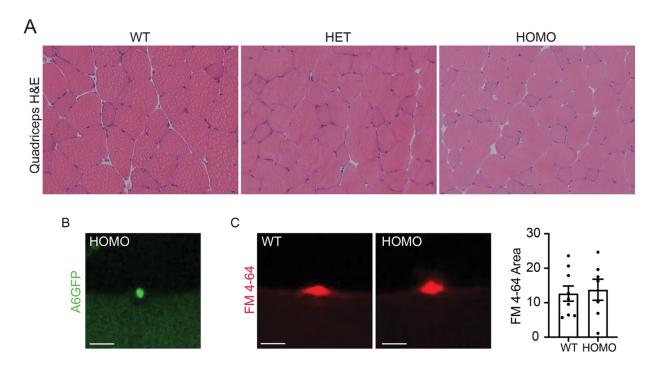
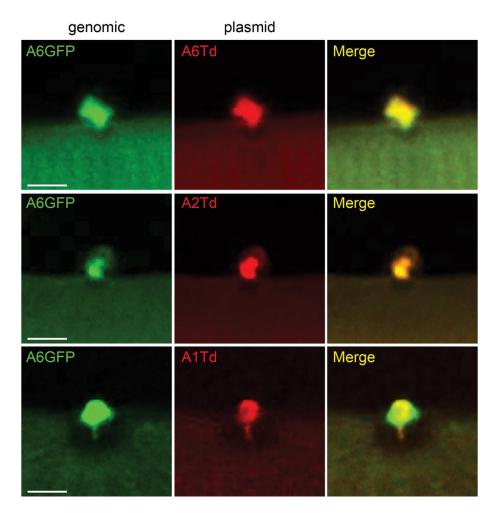
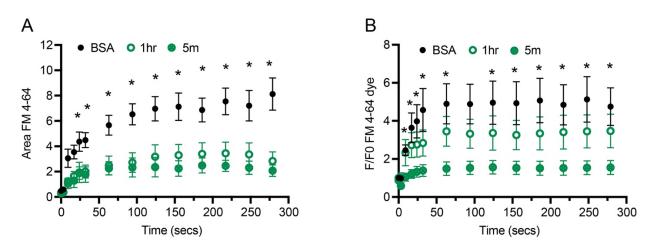
Supplemental Figures and Legends For "A conserved annexin A6-mediated membrane repair mechanism in muscle, heart, and nerve"



**Supplemental Figure 1. A)** Grossly normal muscle histopathology in heterozygous and homozygous *Anxa6gfp* tissue compared to wildtype control muscle. **B**) Upon laser-induced membrane injury, annexin A6GFP localized to the repair cap in homozygous *Anxa6gfp* myofibers. **C**) Similar FM 4-64 uptake after laser-induced membrane injury of wildtype and homozygous *Anxa6gfp* myofibers. Scale 5µm. n=3 mice per genotype, n  $\geq$  7 myofibers.



Supplemental Figure 2. Genomically-encoded annexin A6GFP colocalizes with repair complex members at the site of injury. Myofibers were isolated from heterozygous *Anxa6gfp* mice and electroporated with td-Tomato (red) tagged annexin A1, A2, or A6. Genomically-encoded annexin A6GFP (green) colocalizes with annexin A1, A2 and A6 at the site of membrane damage (merge, yellow). Z-stack images from an injured myofiber. Scale 5µm.  $n \ge 7$  myofibers per condition.



**Supplemental Figure 3.** Recombinant annexin A6 treatment reduces FM dye uptake. Quantification of FM 4-64 dye overtime showed that FM 4-64 was significantly reduced with 1hr and 5mins of recombinant annexin A6 treatment throughout the imaging series measured as (A) FM area or (B) fluorescent intensity. n = 6-8 myofibers from  $n \ge 3$  mice. \* P < 0.05 by ANVOA.

## Supplemental Movie Legends

**Supplemental Movie 1.** Z-stack imaging of genomically encoded annexin A6GFP (green) repair cap localized at the site of myofiber injury.

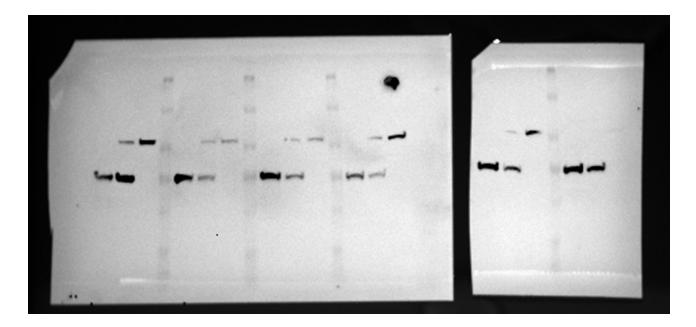
**Supplemental Movie 2.** Timelapse imaging of genomically encoded annexin A6GFP (green) localizing to the site of cardiomyocyte membrane injury over 50 seconds.

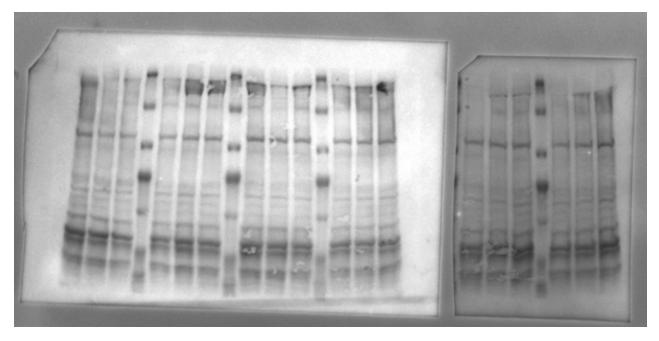
**Supplemental Movie 3.** Timelapse imaging of genomically encoded annexin A6GFP (green) localizing to the site of neuron membrane injury over 60 seconds.

**Supplemental Movie 4.** Timelapse imaging of recombinant annexin A6 tdTomato (red) colocalizing with genomically encoded annexin A6GFP (green) repair cap at the site of muscle membrane injury.

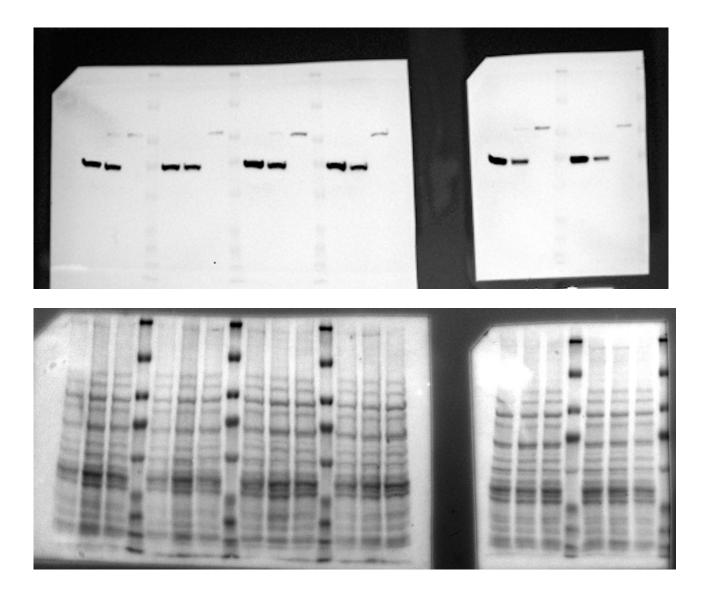
**Supplemental Movie 5.** Recombinant annexin A6 tdTomato (red) forms blebs at the site of myofiber injury.

**Supplemental Movie 6.** Timelapse imaging of recombinant annexin A6 tdTomato (red) localizing to the site of neuron transection. WGA outlines the neuron in blue.

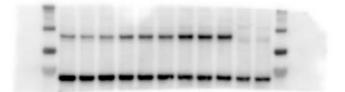




Full gels for Figure 2



Full gels for Figure 3



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Full gels for Figure 4