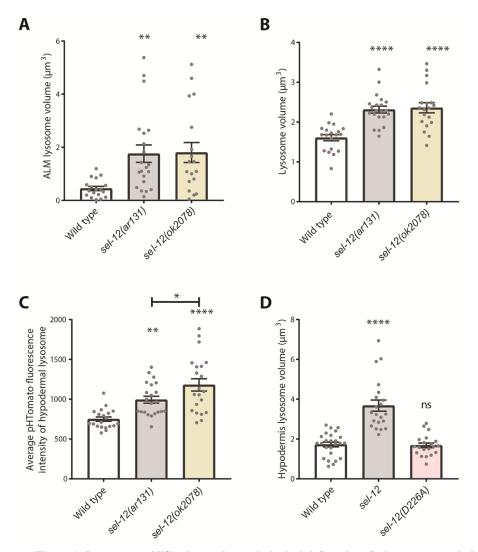
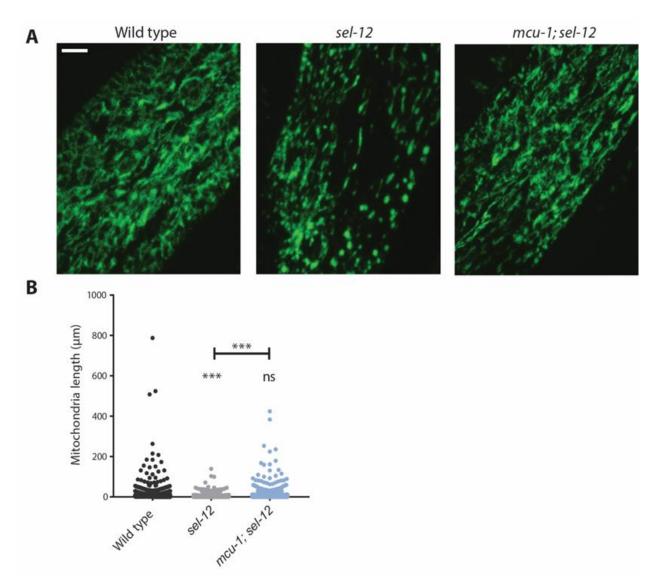
C. elegans Presenilin Mediates Inter-Organelle Contacts and Communication that Is Required for Lysosome Activity

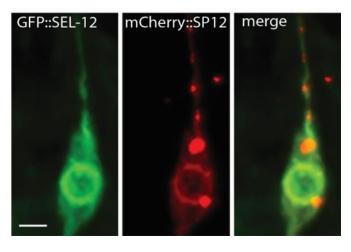
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Supplementary Figure 1. Lysosome acidification and morphological defects in *sel-12* mutants are independent of gamma-secretase activity. Related to Figures 1 and 2. (A) Quantification of average lysosome volume within the ALM TRN soma of wild type, *sel-12(ar131)*, and *sel-12(ok2078)* animals co-expressing *nuc-1*::mCherry to mark lysosomes and *mec-4p*::GFP to mark the TRNs ($n \ge 20$ animals). (B) Quantification of average hypodermal lysosome volume images in animals expressing *nuc-1*::mCherry as a marker for the lysosomal lumen ($n \ge 19$ animals). (C) Quantification of the average pHTomato fluorescence intensity per lysosome in animals expressing *nuc-1*::pHTomato controlled by the heat-shock promoter, with increased pHTomato fluorescence intensity indicating increased pH ($n \ge 20$ animals). (D) Lysosome volume (*nuc-1*::mCherry) in wild type, null *sel-12(ty11)*, and *sel-12*(D226A), which carry a point mutation in a residue necessary for gamma secretase activity ($n \ge 20$ animals). *p < 0.05, ****p < 0.0001 using Kruskal-Wallis with Dunn's multiple comparison test. Comparisons are made to wild type unless otherwise indicated. Error bars indicate mean ± SEM.



Supplementary Figure 2. *sel-12(ty11)* mutants show fragmented mitochondria in the hypodermis. Related to Figure 2. (A) Representative images of hypodermal 2xMLS::GCaMP6f expression (scale bar = 20 μ m) and (B) quantification of hypodermal mitochondrial length. ns p > 0.05, ***p < 0.001 using chi-squared test. n = 20 animals. All comparisons are made to wild type animals unless indicated. Error bars indicate mean ± SEM.



Supplementary Figure 3. SEL-12 localizes to the ER. Related to Figure 4. Confocal image of ALM soma in animal co-expressing a functional SEL-12 GFP fusion protein (*sel-12p::sel-12::*GFP) and pan-neuronal ER reporter (*rgef-1p::mCherry::SP12*) (scale bar = 5 μ m).