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# **Supplemental information**

# Neuronal loss of NCLX-dependent mitochondrial calcium efflux medi-

### ates age-associated cognitive decline

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#### Supplementary figures and legends



# Supplementary Figure 1, related to Figure 1. <sub>m</sub>Ca<sup>2+</sup> exchanger expression and cognitive assay.

**A** Western blot for NCLX expression in tissue isolated from the cortex of 2-, 9-, and 15-monthold control (Camk2a-Cre) mice. **B** Quantification of NCLX protein expression expressed as foldchange vs. 2-month-old Camk2a-Cre con. corrected to voltage-dependent anion channel (VDAC). **C** Western blot for proteins associated with  ${}_{m}Ca^{2+}$  exchange expression in tissue isolated from the cortex of 2-month-old *NCLX*<sup>fl/fl</sup> x Camk2a-Cre mice compared to age-matched controls (Camk2a-Cre). **D-K** Densitometric analysis of western blots shown in Figure S1A expressed as fold change vs. Camk2a-Cre con. corrected to VDAC. **L** Rotarod test for locomotor performance, training phase (day 1 to 3), and test phase (day 4) in 15-month-old mice. n = 3-7. All data presented as mean ± SEM; two-tailed, unpaired t-test.

Supp. Fig. 2



Supplementary Figure 2, related to Figure 2. Effect of genetic ablation of neuronal  $_mCa^{2+}$  efflux on the A $\beta$  pathway.

**A-E** Densitometry analysis of Western blots shown in Figure 2C expressed as fold change vs. Camk2a-Cre con. corrected to a loading control tubulin. n = 3. All data presented as mean  $\pm$  SEM; two-tailed, unpaired t-test.



Supplementary Figure 3, related to Figure 1, 2, 3 and 4. Full-length Western blots.