## **Supplemental Material**

## Figures S1-S5

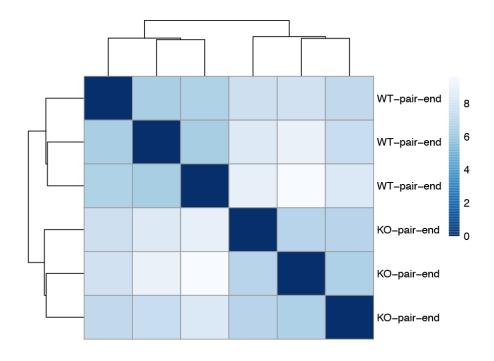


Figure S1

Clustering analysis showed consistent expression patterns in the three biological replicates of each sample (KO or WT), suggesting good repeatability.

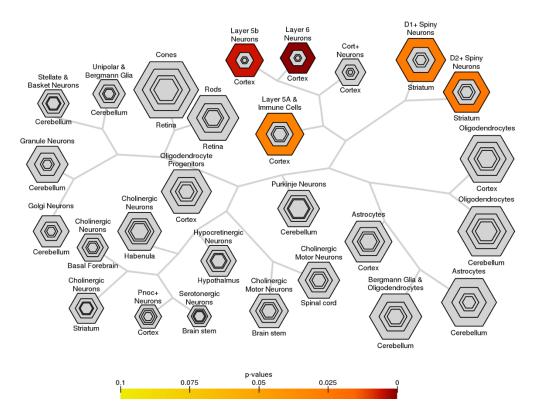


Figure S2 Cell-type specific expression analysis (CSEA) of 354 DEGs.

CSEA shows that some DEGs are expressed in spiny GABAergic neurons, cortical immune cells and neurons. Different cell types are represented by a multilayer hexagonal and color coded by p values (Fisher's exact test) as shown in the bottom scale bar. The hexagon size is scaled to the specific and enriched transcripts number at four stringency thresholds (pSI: 0.05, 0.01, 0.001, 0.0001) and the hexagon with the more stringent threshold (pSI < 0.0001) is in the center (Xu et al., 2014; Lakatos et al., 2017). For example, cortical neurons have smaller p value than spiny neurons at pSI < 0.05 level.

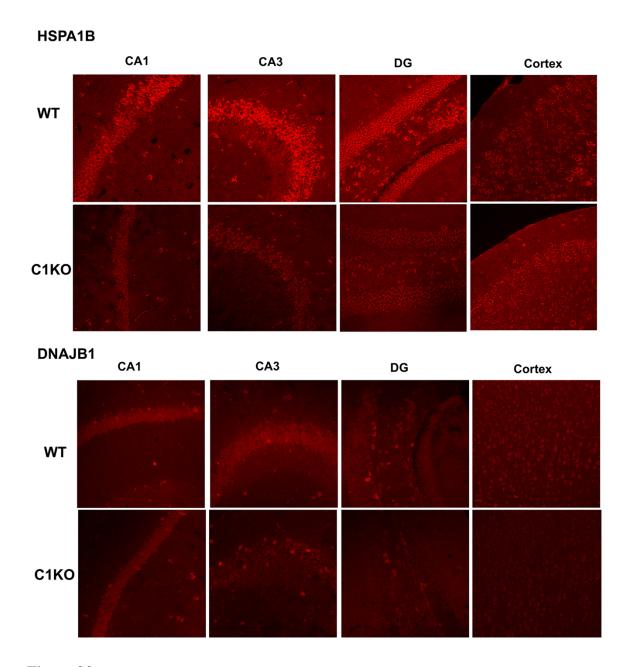


Figure S3

Representative images for HSP70 (HSPA1B) and HSP40 (DNAJB1) in hippocampus and cortex from WT and calpain-1 KO mice (C1KO).

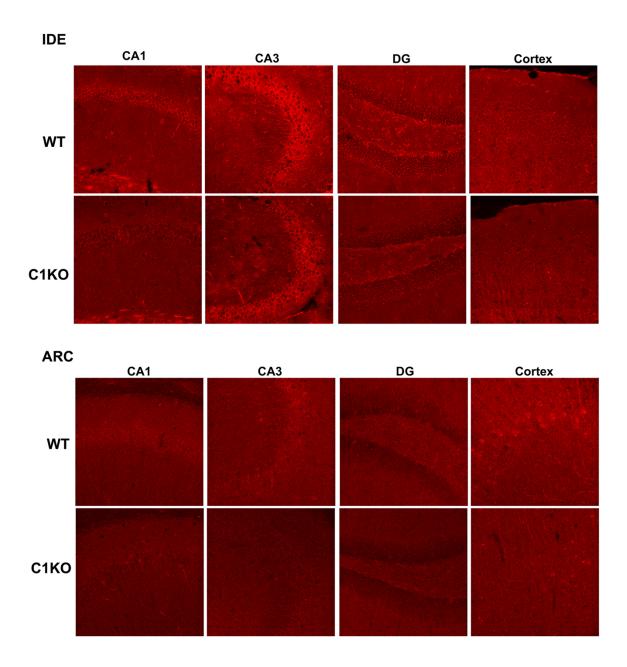


Figure S4

Representative images for Insulin Degrading Enzyme (IDE) and Activity-Regulated Cytoskeleton

Associated Protein (ARC) in hippocampus and cortex from WT and calpain-1 KO mice (C1KO).

## PER2 CA3 CA1 Cortex DG $\mathbf{WT}$ C1KO PLA2G4E CA1 CA3 DG Cortex $\mathbf{WT}$ C1KO

Figure S5

Representative images for Period Circadian Regulator 2 (PER2) and Phospholipase A2 Group IVE (PLA2G4E) in hippocampus and cortex from WT and calpain-1 KO mice (C1KO).