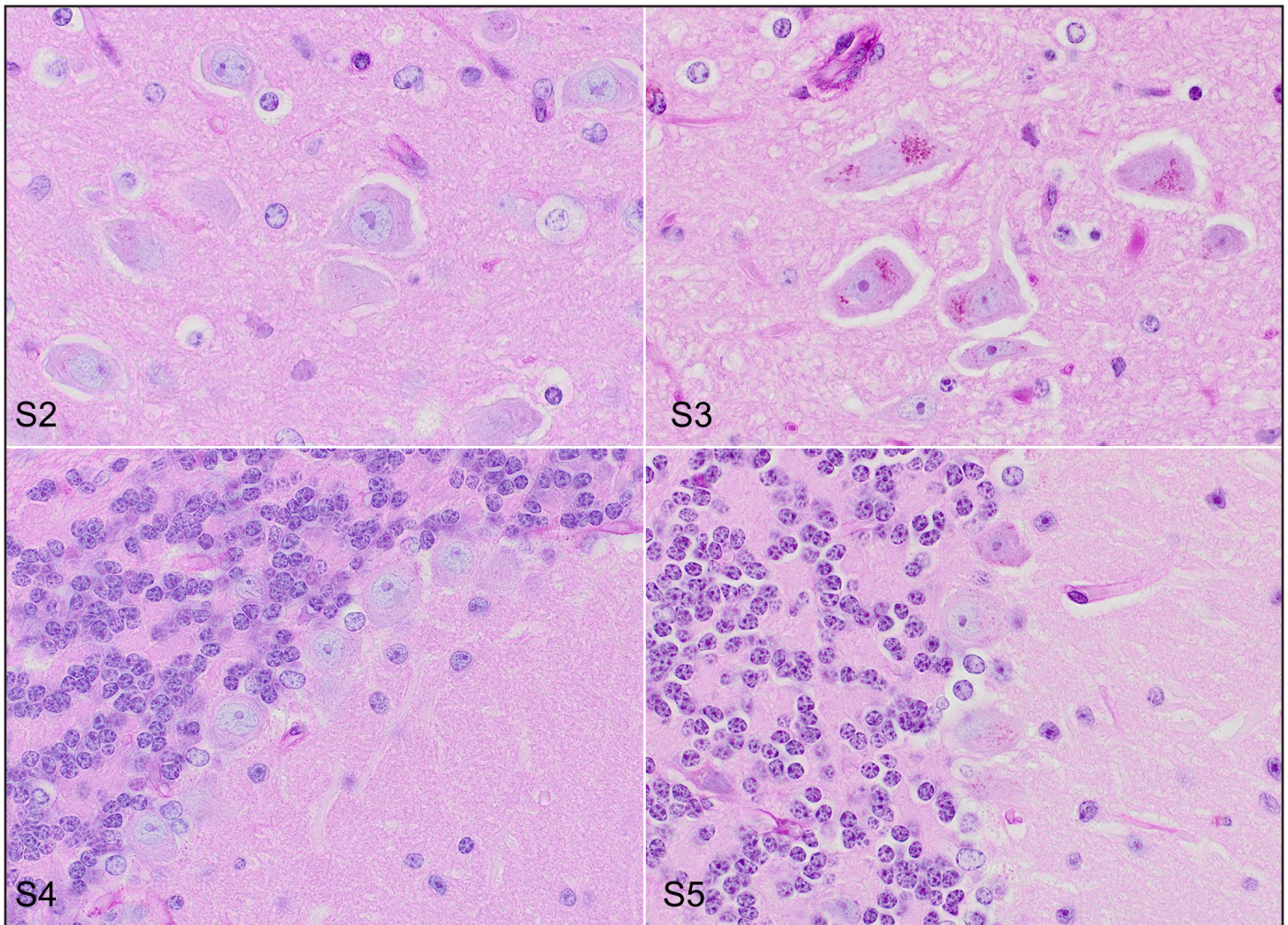


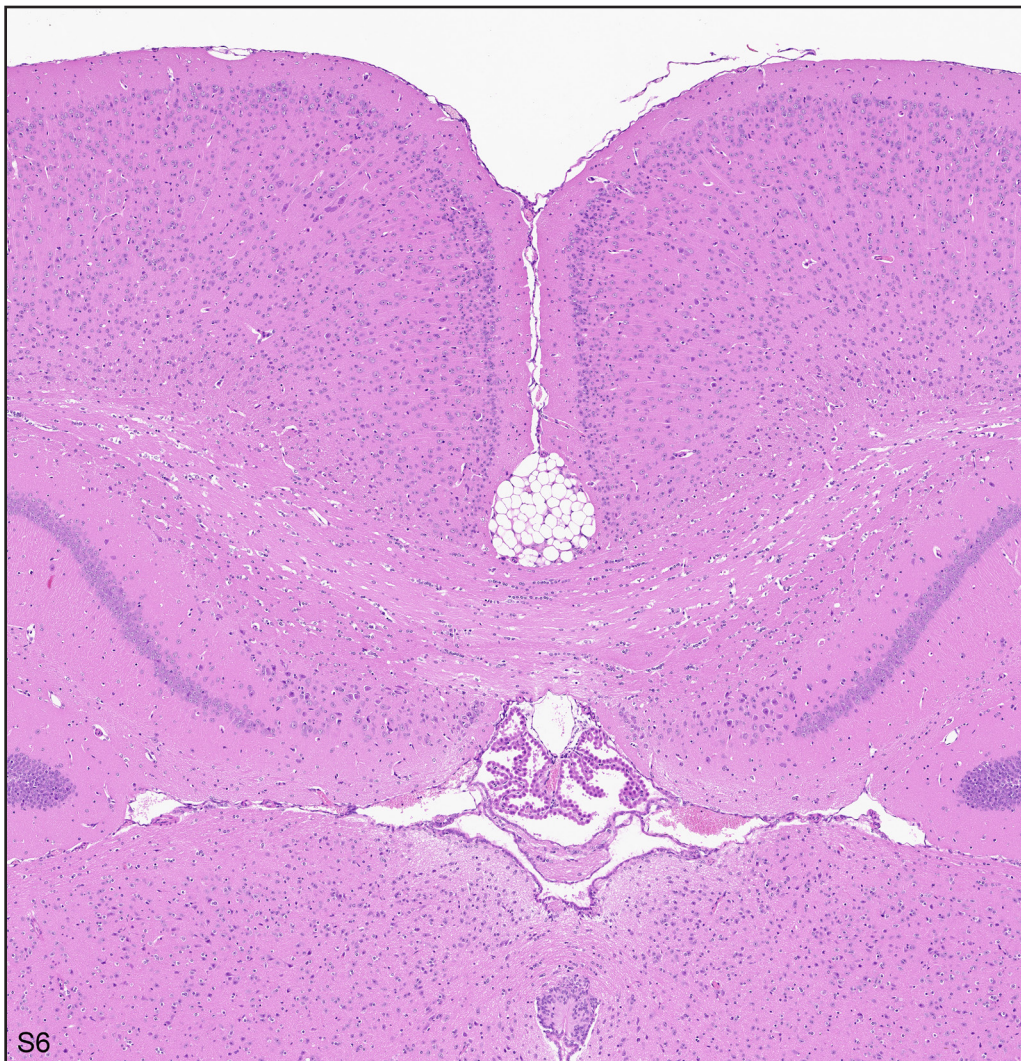
S1

Supp Fig S1 – 56-week-old female C57BL/6J mouse, coronal section of the brain just caudal to the optic chiasm. Immunohistochemistry for p75NTR highlights a group of neurons in the medio-dorsal amygdalar nuclei (dotted box, higher magnification of the region in the dotted box is in the lower right). While this animal showed multiple clusters of PAS-positive granular deposits in the hippocampus (see Fig 7), no immunoreactivity for p75NTR is evident at this level (dashed box, higher magnification of the region in the dashed box is in the upper right).

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Supp Figs S2-5. Figs S2 and S3. Deep cerebellar nuclei, Periodic acid-Schiff (PAS). **Fig. S2.** Neurons contain few punctate PAS-positive lipofuscin granules in the deep cerebellar nucleus in this 17-week-old female C57BL/6J mouse. **Fig. S3.** Neurons exhibit moderate accumulation of PAS-positive lipofuscin granules in the deep cerebellar nucleus of this 78-week-old female C57BL/6J mouse. **Figs. S4 and S5.** Cerebellar Purkinje cells, PAS stain. **Fig. S4.** Purkinje cells in this 17-week-old female C57BL/6J mouse contain no PAS-positive material. **Fig. S5.** Few PAS-positive granules are in the Purkinje cells of this 78-week old female C57BL/6J mouse.



Supp Fig S6. 10-week-old female C57BL/6J mouse. A small lipomatous hamartoma is evident in the interhemispheric sulcus. Hematoxylin and eosin (HE).

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Supplemental Table 1. Spontaneous incidental/background brain pathology in C57BL/6J Mice. Details concerning primary antibodies and procedures used for the immunohistochemical examination of selected cases/lesions.

Epitope/Target	Primary antibody	Source	Antigen retrieval	Working dilution	Incubation time	Mouse tissue/s used as control	Staining platform and detection system
GFAP	Rb pAb #Z0334	Agilent/Dako	HIER low pH	1:5000	45 min RT	FFPE brain (C+) FFPE kidney (C-)	Leica BOND RXm, Bond Polymer Refine Detection Kit (#DS9800)
IBA1	Rb pAb #019-19741	Wako	HIER low pH	1:1500	45 min RT	FFPE liver (C+ and C-)	Leica BOND RXm, Bond Polymer Refine Detection Kit (#DS9800)
Neurofilament-L	Rb mAb #2837	Cell Signaling Technology	HIER low pH	1:500	45 min RT	FFPE brain (C+) FFPE kidney (C-)	Leica BOND RXm, Bond Polymer Refine Detection Kit (#DS9800)
p75NTR	Rb mAb #8238	Cell Signaling Technology	HIER low pH	1:100	45 min RT	FFPE brain (C+) FFPE heart (C-)	Ventana discovery ultra, UltraView Universal DAB Detection Kit (#760-500)
Phospho-Tau (Thr181)	Rb mAb #12885	Cell Signaling Technology	HIER high pH	1:1200	45 min RT	FFPE brain from B6.Cg-Tg(Thy1-MAPT*P301S)2541Godt (C+) FFPE kidney (C-)	Leica BOND RXm, Bond Polymer Refine Detection Kit (#DS9800)
Synaptophysin	Rb mAb #36406	Cell Signaling Technology	HIER high pH	1:100	45 min RT	FFPE pancreas (C+ and C-)	Leica BOND RXm, Bond Polymer Refine Detection Kit (#DS9800)
Ubiquitin	Ms mAb #3936	Cell Signaling Technology	HIER low pH	1:1000	45 min RT	FFPE brain from B6.Cg-Tg(Thy1-APP ^{Sw} ,Thy1-PSEN1*L166P)21Jckr (C+) FFPE kidney (C-)	Ventana discovery ultra, UltraView Universal DAB Detection Kit (#760-500)
Vinculin	Rb mAb #13901	Cell Signaling Technology	HIER low pH	1:1200	45 min RT	FFPE liver (C+ and C-)	Leica BOND RXm, Bond Polymer Refine Detection Kit (#DS9800)

Abbreviations used in the table: Rb-Rabbit, Ms-Mouse, pAb-Polyclonal Antibody, mAb-Monoclonal Antibody, HIER-heat induced epitope retrieval, min-minutes, RT-room temperature, C+ positive control, C- negative control.