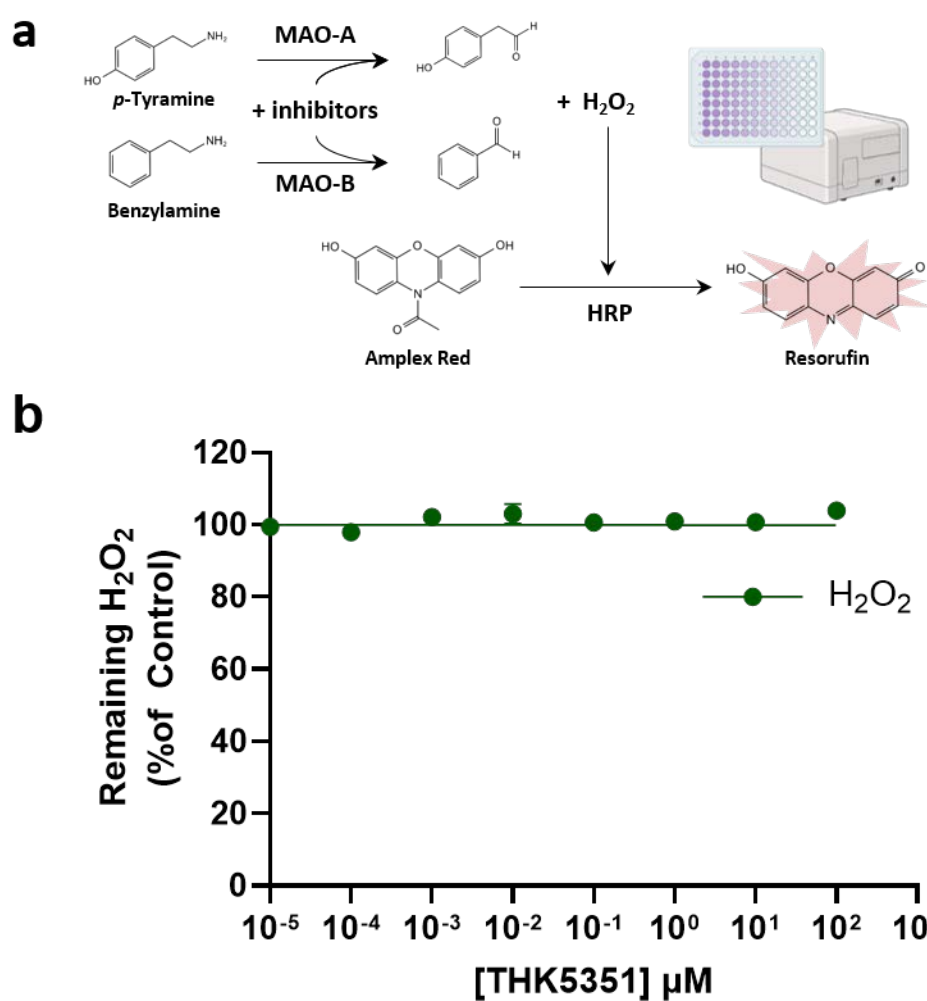


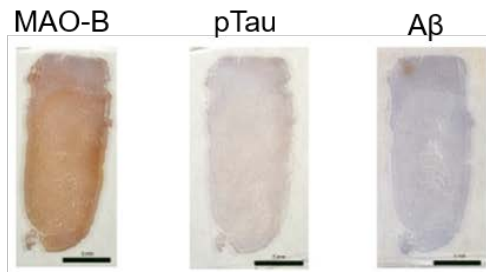
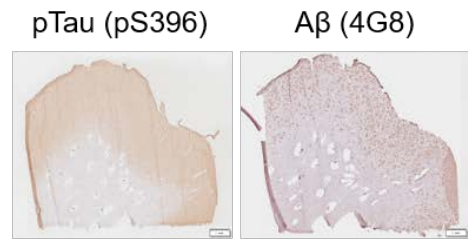
Supplementary Information

In-depth Multimodal Validation of ^{18}F -THK5351 for Imaging MAO-B-mediated Reactive Astrogliosis in Alzheimer's and Related Neurodegenerative Diseases

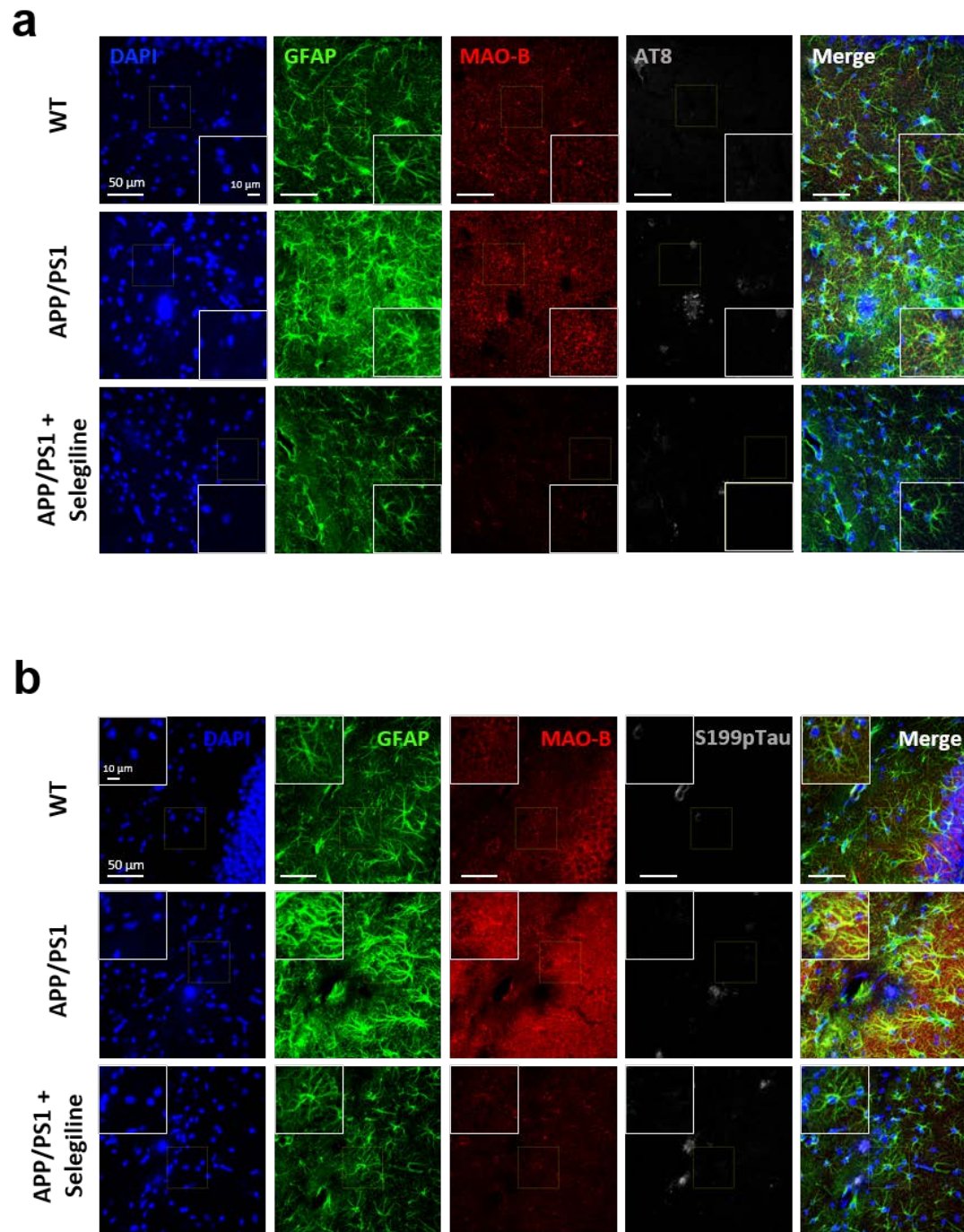
Heejung Chun^{1,2,9*}, Wongu Youn^{3,9}, Hyunkeong Lim^{4,9}, Ryuichi Harada^{5,6}, Hae Young Ko⁴, Jiyoung Yoon^{1,2}, Soo-Jin Oh⁷, Philipp Seemann⁸, Joseph G. Pepe⁸, Brian K. Shoichet⁸, Shozo Furumoto⁶, Nobuyuki Okamura^{5,6}, Mijin Yun^{4*}, C Justin Lee^{3*}



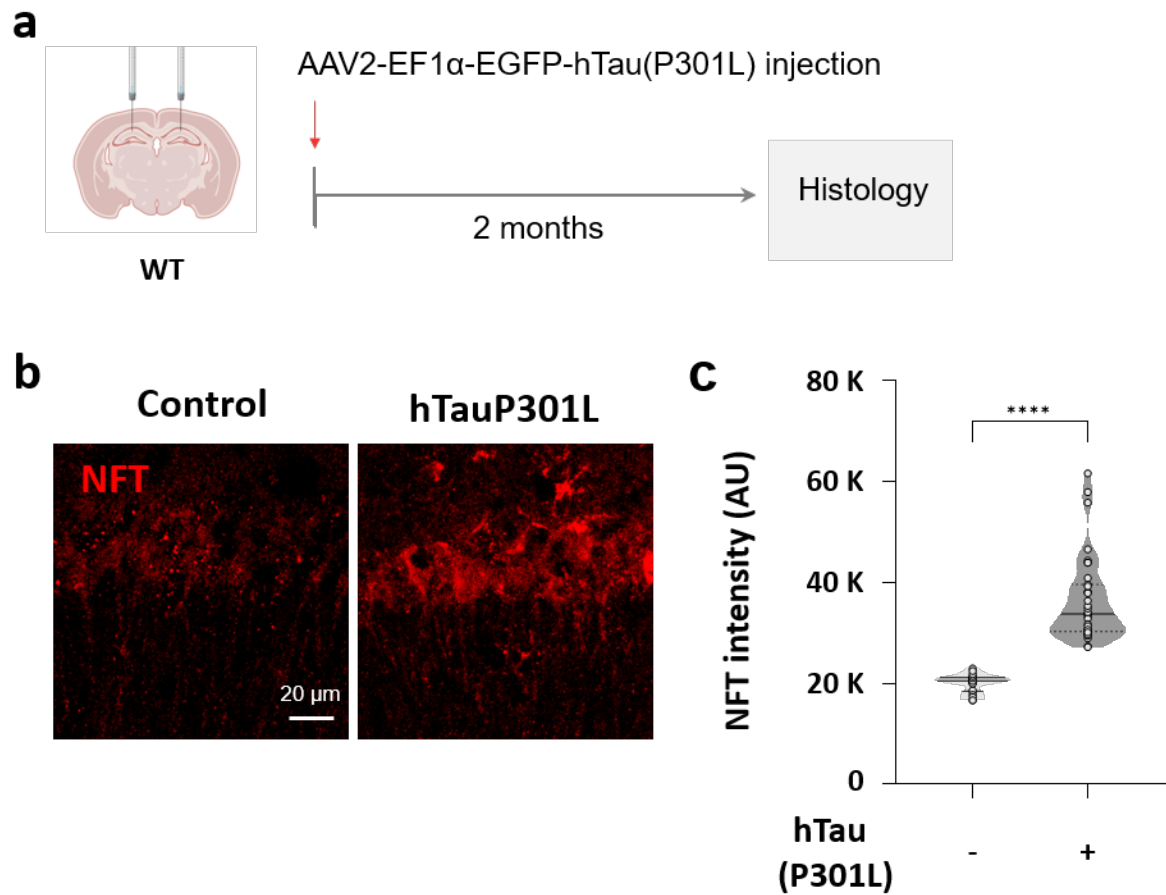
Supplementary Figure 1. Enzyme activity assay of THK5351. **a.** Schematic description of MAO-A and MAO-B enzyme assay. **b.** Amplex Red H_2O_2 assay with THK5351. Data are presented as the mean $\pm$ s.e.m.

aImmunohistochemistry**b**Immunohistochemistry

Supplementary Figure 2. IHC images of human brain section. a. IHC images of MAO-B, pTau and Aβ in human basal ganglia tissue. **b.** IHC images of pTau(S396) and Aβ in human AD brain tissue.



Supplementary Figure 3. Long-term treatment of selegiline efficiently inhibits the reactivity of astrocytes and the expression of MAO-B in APP/PS1 mice. a, b. Representative IHC images of GFAP, MAO-B and pTau(AT8 (a), S199 (b)) in selegiline-treated APP/PS1 mice brain.



Supplementary Figure 4. Overexpression of mutated tau protein, hTauP301L, in hippocampal CA1 region induces neurofibrillary tangle formation. **a.** Schematic description of hTauP301L virus injection. **b.** IHC image of NFT in hTauP301L virus injected mice or control mice hippocampus. **c.** Quantification of NFT intensity. Data are shown as violin plots with individual points, median, and interquartile range (25th-75th) indicated. **** $P < 0.0001$.