BCAS AB

# Supplemental Information

Chronic cerebral hypoperfusion shifts the equilibrium of amyloid  $\beta$  oligomers to

aggregation-prone species with higher molecular weight

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# $BCAS\,A\beta$

(a) KB staining



Bar = 100 µm

Supplemental Figure S1. BCAS decreased the area of the cingulum with refraction

(a) KB staining of the coronal section of the mice brain. The corpus callosum was surrounded by red dot-line. Bar =  $200 \mu m$ . (b) TUNEL assay of the hippocampus at week 0 or 10 after BCAS operation. Assays with DNase treated sections were used as positive controls. Bar =  $100 \mu m$ .

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#### BCAS $A\beta$



Supplemental Figure S2. BCAS had no effect on APP processing and A $\beta$  metabolism in the brain of APP/PS1 mice

(a) Quantifications of APP, sAPP $\alpha$  and sAPP $\beta$  in the Triton-X fraction by western-blot. (b) Quantification of panel (a). Actin-normalized level of expression of each protein in the Triton-X fraction was normalized to sham = 1. (c) Quantifications of APP and A $\beta$  processing enzymes in the total brain homogenates by western-blot. (d) Quantification of panel (d). Relative quantity of each protein to actin expression in Triton-X or PBS fraction was normalized to sham = 1.

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Supplemental Figure S3. Serial detergent fractionation assay of sham- or BCAS-operated APP/PS1 mice brain homogenates.

The concentration of  $A\beta_{40}$  (a, c) and  $A\beta_{42}$  (b, d) in the brain homogenates was measured by ELISA. Samples were obtained at 5 weeks (a, b) and at 15 weeks (c, d) after the BCAS operation. Statistical significance was determined by using a Student's *t*-test. n = 8 (5 weeks) and 6 (15 weeks).

# Supplemental Figure S2A



sAPPβ



sAPPα



Actin



# Supplemental Figure S2C

## BACE1



PEN2



#### Nicastrin



## PSEN1



## APH1a



## RAGE



# Supplemental Figure S2C

## LRP1



ABCB1



Actin



#### Clusterin



## Neprilysin

