

Supplementary Data

Table S1. Details of antibodies used in Western blotting.

| Antibody | Molecular weight | Catalog number | Dilution | Company | Area |
|---------------------------|------------------|----------------|----------|-------------|---------------------|
| t-tau | 79 kDa | A0002 | 1:2000 | ABclonal | Wuhan, China |
| p-tau | 79 kDa | ab32057 | 1:2000 | Abcam | Cambridge, MA, USA |
| A β ₁₋₄₀ | 4.3 kDa | bs-0877R | 1:2000 | Bioss | Beijing, China |
| A β ₁₋₄₂ | 4.4 kDa | bs-0107R | 1:2000 | Bioss | Beijing, China |
| GAPDH | 35 kDa | E-AB-40337 | 1:2000 | Elabscience | Wuhan, China |
| Nrf2 | 100 kDa | A1244 | 1:1000 | ABclonal | Wuhan, China |
| GCLC | 73 kDa | bs-23393R | 1:1500 | Bioss | Beijing, China |
| HO-1 | 33 kDa | AF5393 | 1:1000 | Affinity | Cincinnati, OH, USA |
| NQO1 | 31 kDa | A19586 | 1:2000 | ABclonal | Wuhan, China |
| SOD1 | 16 kDa | A12537 | 1:1000 | ABclonal | Wuhan, China |
| SOD2 | 22 kDa | A1340 | 1:1000 | ABclonal | Wuhan, China |
| 4-HNE | 67 kDa | bs-6313R | 1:1000 | Bioss | Beijing, China |
| Bcl-2 | 26 kDa | A0208 | 1:1000 | ABclonal | Wuhan, China |
| Bax | 21 kDa | A19684 | 1:1000 | ABclonal | Wuhan, China |
| goat anti-mouse | | E-AB-1001 | 1:4000 | Elabscience | Wuhan, China |
| goat anti-rabbit | | E-AB-1003 | 1:4000 | Elabscience | Wuhan, China |

Table S2. Relative abundance of top 20 genera among WT, APP/PS1 and APP/PS1+HC.

| Taxa | WT | APP/PS1 | APP/PS1+HC |
|-------------------------------|-----------|----------------|-------------------|
| <i>Muribaculum</i> | 0.168931 | 0.147624 | 0.119309745 |
| <i>Lactobacillus</i> | 0.024037 | 0.090404 | 0.106769995 |
| <i>Lachnospirillum</i> | 0.069882 | 0.04285 | 0.035904689 |
| <i>Eisenbergiella</i> | 0.012297 | 0.025235 | 0.031373288 |
| <i>Akkermansia</i> | 0.006255 | 0.051867 | 0.000128735 |
| <i>Desulfovibrio</i> | 0.023127 | 0.011201 | 0.003374753 |
| <i>Bacteroides</i> | 0.004681 | 0.012425 | 0.019031598 |
| <i>Alistipes</i> | 0.008606 | 0.01562 | 0.008180541 |
| <i>Erysipelatoclostridium</i> | 0.00457 | 0.009511 | 0.016109862 |
| <i>Anaerotruncus</i> | 0.009517 | 0.005499 | 0.010060246 |
| <i>Helicobacter</i> | 0.00188 | 0.012341 | 0.009718913 |
| <i>Staphylococcus</i> | 0.002836 | 0.009946 | 0.002749092 |
| <i>Kineothrix</i> | 0.003342 | 0.002538 | 0.00727808 |
| <i>Butyrivibrio</i> | 0.004694 | 0.002369 | 0.001389474 |
| <i>Oscillibacter</i> | 0.00455 | 0.001589 | 0.002256994 |

| | | | |
|-------------------------------|----------|----------|-------------|
| <i>Ruminococcus</i> | 0.002902 | 0.001924 | 0.00329547 |
| <i>Prevotella</i> | 0.005757 | 0.000174 | 0.00004539 |
| <i>Faecalibaculum</i> | 0.001701 | 0.004107 | 0.00004997 |
| <i>Neglecta</i> | 0.003233 | 0.001031 | 0.001406 |
| <i>Candidatus Arthromitus</i> | 0.003019 | 0.000441 | 0.001529127 |

Data are presented as the mean.

Table S3. The differential metabolites of serum among WT, APP/PS1 and APP/PS1+HC.

| Metabolites | WT | APP/PS1 | APP/PS1+HC | ANOVA <i>P</i> value |
|---------------------------|-------------|-------------|-------------|----------------------|
| Phosphocholine | 9334151.018 | 26085118.25 | 13626510.64 | 0.001338 |
| Oligomycin b | 2080200.549 | 6223581.442 | 2123645.369 | 0.01381 |
| Bisphenol af | 145204716.6 | 241463850 | 164955740.4 | 0.020436 |
| 3,4-dihydroxyacetophenone | 4243171.542 | 3899588.444 | 4898134.431 | 0.015564 |
| Vitexin | 575009.3181 | 288833.2647 | 404925.8763 | 0.026925 |
| Propranolol | 1857670.674 | 821794.0851 | 2385704.623 | 0.04404 |
| D-Glutamic acid | 54131865.59 | 67947340.77 | 55742368.18 | 0.003671 |
| Oxidized glutathione | 15201929.28 | 18239389.14 | 15857783.28 | 0.015878 |

Data are presented as the mean.

Differences are considered statistically significant at ANOVA *P* value <0.05

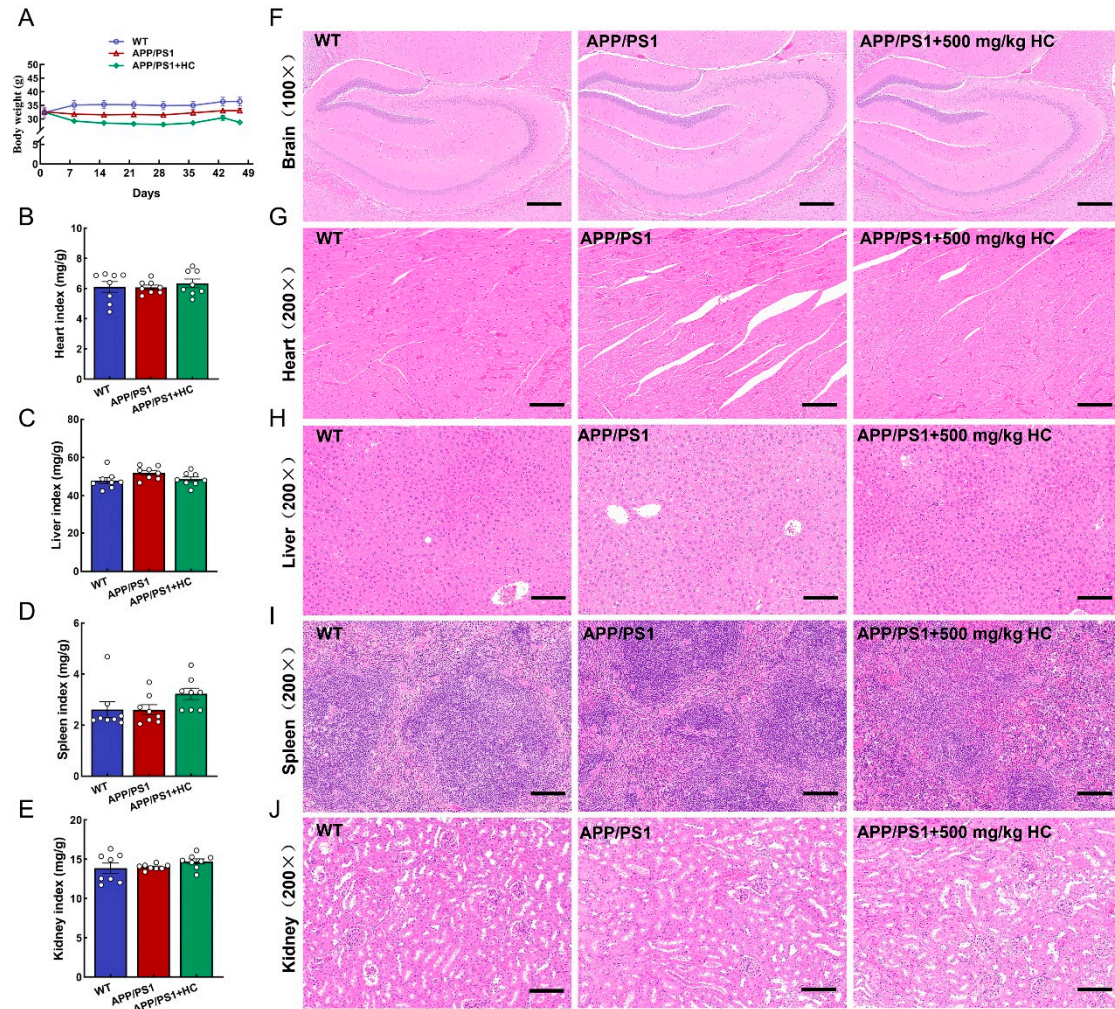


Figure S1 Safety evaluation of APP/PS1 mice treated with HC. (A) The body weight changes ($n = 8$). (B) Heart index, (C) liver index, (D) spleen index, and (E) kidney index of APP/PS1 mice ($n = 8$). Representative images of H&E staining of the (F) brain, (G) heart, (H) liver, (I) spleen, and (J) kidney (scale bar: 100 μ m) ($n = 3$). Data are expressed as mean \pm S.E.M..

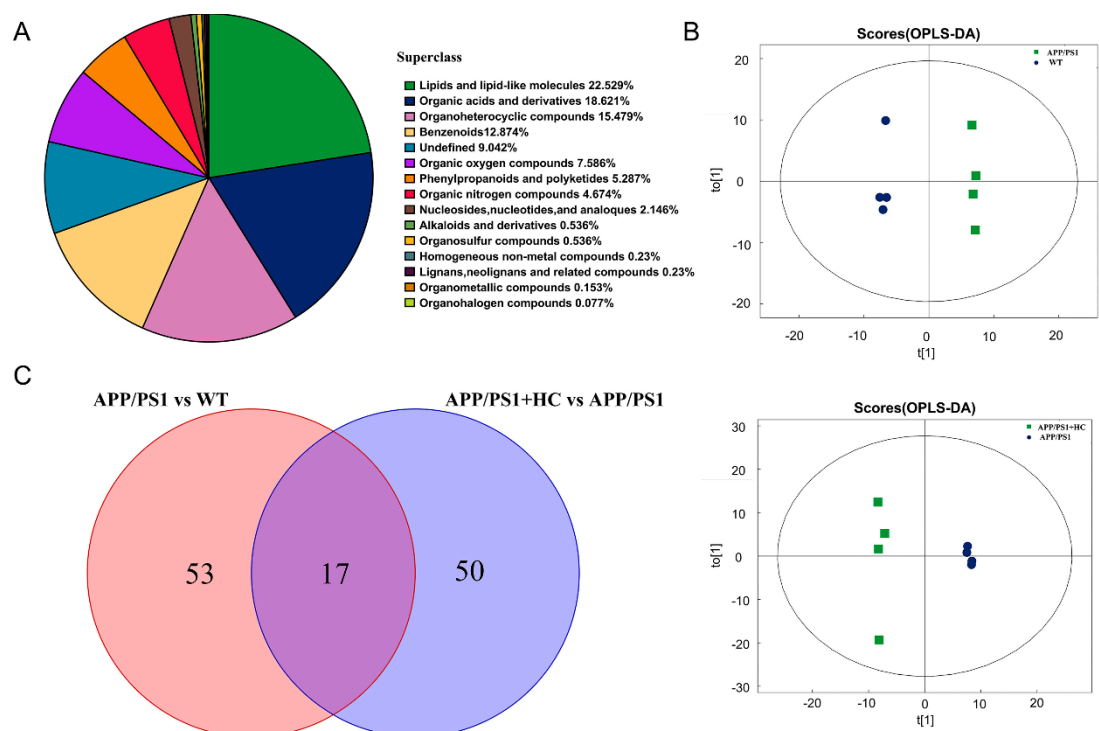


Figure S2 Classification, quantity and multidimensional statistical analysis of metabolites. (A) The proportion of identified metabolites in each chemical classification. (B) OPLS-DA score map of the WT mice and APP/PS1 mice (above), OPLS-DA score map of HC treated APP/PS1 mice and APP/PS1 mice (below) ($n = 4$). (C) Venn diagram.