- 1 Supplementary Information for
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FKBP52 overexpression accelerates hippocampal-dependent memory impairments in a tau transgenic mouse model

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27 Supplementary Figure 1. FKBP52 overexpression did not affect hippocampal LTP in WT 28 mice. a The Input-Output (I-O) curve was calculated to evaluate pre- and post-synaptic excitatory 29 function. The I-O curve compares the fEPSP slope (mV/ms) versus the fiber volley amplitude (mV). 30 b LTP was measured in ex vivo hippocampal slices from WT mice infused with AAV9-GFP or AAV9-31 FKBP52. Stimulating electrode was positioned in the Schaffer collaterals (between CA3 and CA1) 32 and recordings were obtained in the CA1 pyramidal neurons. A 200Hz Theta Burst stimulation was 33 given to induce LTP (indicated by the arrow). Following this, field EPSPs were measured for 60-34 minutes. Representative traces are shown as: 1 (black) indicates baseline, 2 (teal) indicates initial 35 early LTP in the first minute following HFS, and 3 (orange) indicates late LTP in the last 60 minutes 36 of recording. Data analyzed by repeated measures two-way ANOVA. Data is shown in standard error of the mean (± SEM). n = 27 for WT-GFP and n = 23 WT-FKBP52. AAV9, adeno-associated 37 38 virus serotype 9; fEPSP = field excitatory postsynaptic potential; LTP, long-term potentiation; min 39 = minutes; mV, millivolts; ms, milliseconds; Hz, hertz.



Supplementary Figure 2. Total tau levels were increased in CA3 of rTg4510-FKBP52 mice.
Hippocampal slices were obtained from rTg4510 mice injected with AAV9-GFP and AAV9FKBP52. a Quantification of tau levels using tau (DAKO) antibody. b Quantification of oligomeric
tau levels using T22 antibody. c Quantification of pS202/T205 levels (AT8 antibody) in the
hippocampi. Protein levels were analyzed by unpaired t-test, where statistical significance is
considered by *p < 0.05 Results represented as standard error of the mean (± SEM); GFP, n = 4;
FKBP52, n = 4).



Supplementary Figure 3. Western blot membranes for total tau and phospho-tau species.
Western blot membranes of soluble fractions for total tau and tau phosphorylated species including
pT181, pS202/pT205 (AT8), and pT231 (AT180). Hippocampal tissue from WT and rTg4510 mice
expressing either AAV9-GFP or AAV9-FKBP52 was used in these blots. Membranes were run in
parallel and derived from the same experiments. Quantification of protein was performed using
Image Lab (Bio-Rad) software. Total protein was normalized to GAPDH. WT-GFP (n = 8), WTFKBP52 (n = 8), rTg4510-GFP (n = 5), rTg4510-FKBP52 (n = 5). AAV9, adeno-associated virus

58 serotype 9; GFP, green fluorescent protein; GAPDH, Glyceraldehyde 3-phosphate dehydrogenase;



WT, wild-type; Std, Protein Standards. 59

61 Supplementary Figure 4. Western blot membranes for caspase 3 and caspase 12. 62 Membranes were run in parallel and derived from the same experiments. Quantification of protein 63 was performed using Image Lab (Bio-Rad) software. Total protein was normalized to GAPDH. WT-64 GFP (n = 8), WT-FKBP52 (n = 8), rTg4510-GFP (n = 5), rTg4510-FKBP52 (n = 5). AAV9, adenoassociated virus serotype 9; GFP, green fluorescent protein; GAPDH, Glyceraldehyde 3-phosphate 65 66 dehydrogenase; WT, wild-type; Cas, caspase; kDa, kilodalton; Std, Protein Standards.

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