

Supplementary Figure Legends

Figure S1. The swim speed and locomotor activity are similar between *NCoR1*

***loxP* mice and *NCoR1* cKO mice**

(A) The swim speed ($t_{1,16} = 0.95$, $p > 0.05$) and (B) Visible platform learning performance ($F_{1,16} = 0.7$, $p > 0.05$) of *NCoR1 loxP* mice and *NCoR1* cKO mice. N = 9 each group from the same animals in Figure 2. (C) Number of crossovers ($t_{1,10} = 0.24$, $p > 0.05$) and mean movement speed ($t_{1,10} = 0.18$, $p > 0.05$) for locomotor activity measure from a different batch of *NCoR1 loxP* mice and *NCoR1* cKO mice. N = 6 each group. (D) *NCoR1* expression level in the hippocampus of *NCoR1 loxP* mice and *NCoR1* cKO mice from (C) ($t_{1,10} = 4.87$, $p < 0.001$). Data are expressed as individual values and mean $\pm$ SEM or expressed as mean $\pm$ SEM (for B). *** $p < 0.001$.

Figure S2. NMDA administration decreases the association between *NCoR1* and

DEC2 and spatial training decreases the association between *NCoR1* and

HDAC3

(A) Rats received acute PBS or NMDA (8 mM) injection to their hippocampal CA1 area and their CA1 tissue lysate was subjected to immunoprecipitation using anti-*NCoR1* antibody and immunoblotting using anti-DEC2 antibody and anti-*NCoR1* antibody at 1 h later (right panel). These tissue lysates were also

subjected to western blot determination of NCoR1 expression (right-lower panel). For the control experiment, the same tissue lysates were subjected to immunoprecipitation with IgG and immunoblotting with anti-DEC2 antibody (left panel). **(B)** The quantified result of (A) for the association between NCoR1 and DEC2 ($t_{1,6} = 9.44$, $p < 0.001$) and between NCoR1 and NCoR1 ($t_{1,6} = 10.53$, $p < 0.001$) ($N = 4$ each group). **(C)** The CA1 tissue from trained and non-trained rats was subjected to immunoprecipitation using anti-HDAC3 antibody and immunoblotting using anti-NCoR1 antibody. The same tissue lysates were immunoprecipitated with IgG and immunoblotted with anti-NCoR1 antibody to serve as the control experiment. These tissue lysates were also subjected to western blot determination of HDAC3 expression (lower panel). **(D)** The quantified result of (C) for the association between HDAC3 and NCoR1 ($t_{1,4} = 5.8$, $p < 0.01$) and for HDAC3 expression in the hippocampal lysate ($t_{1,4} = 0.07$, $p > 0.05$) ($N = 3$ each group). **(E)** DEC2 expression level was examined in trained and non-trained animals and a representative gel pattern is shown. **(F)** The quantified result of (E) ($t_{1,8} = 12.93$, $p < 0.001$). Data are expressed as individual values and mean $\pm$ SEM. ** $p < 0.01$ and *** $p < 0.001$.

Figure S3. *DEC2* siRNA transfection enhances but *DEC2*WT plasmid transfection impairs spatial learning and memory performance

(A) Mice received control siRNA or *DEC2* siRNA (15 pmol) transfection to their hippocampal CA1 area and they were subjected to water maze learning 48 h later for two consecutive days with four trials a day ($F_{1,12} = 26.09$, $p < 0.001$). $N = 7$ each group. (B) Probe trial performance (time spent in the target quadrant) ($t_{1,12} = 2.22$, $p < 0.05$) and representative swim patterns from the same animals. (C) Total distance traveled in the target quadrant from the same animals ($t_{1,12} = 4.66$, $p < 0.001$). (D) Swim speed ($t_{1,12} = 0.03$, $p > 0.05$) for the probe trial test from the same animals. (E) *DEC2* expression level in the hippocampus from the same animals after the probe trial test ($t_{1,12} = 11.26$, $p < 0.001$). (F) Mice received Flag-vector or Flag-*DEC2*WT plasmid transfection to their hippocampal CA1 area and they were subjected to water maze learning 48 h later for two consecutive days with four trials a day ($F_{1,12} = 37.23$, $p < 0.001$). $N = 7$ each group. (G) Probe trial performance (time spent in the target quadrant) ($t_{1,12} = 2.77$, $p < 0.05$) and representative swim patterns from the same animals. (H) Total distance traveled in the target quadrant from the same animals ($t_{1,12} = 2.31$, $p < 0.05$). (I) Swim speed ($t_{1,12} = 0.36$, $p > 0.05$) for the probe trial test from the same animals. (J) *DEC2* expression level in the hippocampus from the same animals after the probe trial test ($t_{1,12} = 11.83$, $p < 0.001$). The same tissue lysates were immunoprecipitated with anti-Flag antibody and immunoblotted with anti-Flag antibody to confirm the transfection and expression of the Flag-*DEC2*WT plasmid

(lower panel). Data are expressed as individual values and mean $\pm$ SEM or expressed as mean $\pm$ SEM (for A and E). * $p < 0.05$ and *** $p < 0.001$.

Figure S4. Transfection efficiency in primary culture and Neuro2A cells

(A) Immunofluorescence staining of EGFP (green) and DAPI (blue) after pcDNA3-EGFP plasmid transfection to mouse primary neurons is shown. **(B)** Number of EGFP-positive neurons over total number of neurons is calculated as the transfection efficiency. N=4. **(C)** Immunofluorescence staining of EGFP (green) and DAPI (blue) after pcDNA3-EGFP plasmid transfection to Neuro2A cells is shown. **(D)** Number of EGFP-positive neurons over total number of neurons is calculated as the transfection efficiency. N=4. Individual values are shown.