

Figure S1. FMRP localizes to parvalbumin (PV) and somatostatin (SOM)-expressing neurons in mouse hippocampus. Representative images showing immunostaining of FMRP and PV (top) or SOM (middle) in WT mouse hippocampus. Arrows point to individual cells co-expressing FMRP and PV or SOM. Bottom panel shows absence of FMRP labeling in *Fmr1*^{-/y} mouse hippocampus; Scale bar: 50 μ m.



Figure S2. FMRP expression and rotarod testing in *Fmr1-^{/y}*-PV mice. Quantification of FMRP fluorescence in PV-expressing cells in WT-PV and *Fmr1-^{/y}*-PV mouse (A) mPFC and (B) hippocampus. (C) Rotarod test. *Fmr1-^{/y}*-PV and WT-PV mice showed no significant differences in motor coordination and learning in the rotarod test. Values represent mean \pm SEM (n = 12-14 animals per genotype); RM two-way ANOVA followed by Bonferroni's multiple comparisons test.



Figure S3. Object recognition, MWM reversal, and visible MWM behavior of *Fmr1*-/y-PV mice. (A) Preference Index (PI) for a novel object during STM and LTM portions of object recognition test did not differ significantly between *Fmr1*-/y-PV and WT-PV mice. Values represent mean \pm SEM (n = 12-13 animals per genotype). (B) Escape latency during the reversal or flag test portions of MWM was not significantly different between *Fmr1*-/y-PV and WT-PV mice. Values represent mean \pm SEM (n = 15 animals per genotype); RM Two-way ANOVA followed by Bonferroni's multiple comparisons test, Student's *t* test.



Figure S4. FMRP expression and behavoioral testing of *Fmr1*-^{*i*}-*y*-SOM mice. Quantification of FMRP fluorescence in SOMexpressing cells in WT-SOM and *Fmr1*-^{*i*}y-SOM (A) mPFC and (B) hippocampus. Water-based Y maze, three chamber social preference and social novelty test, object recognition, MWM probe test, and visible MWM test were performed on *Fmr1*-^{*i*}y-SOM mice. (C) There was no significant difference in percentage of correct choices during the training, LTM, or reversal portions of the Y Maze task between WT-SOM and *Fmr1*-^{*i*}y-SOM mice. Values represent mean ± SEM (n = 9-12 animals per genotype. (D-E) WT-SOM and *Fmr1*-^{*i*}y-SOM mice spent more time exploring a stranger mouse (S) compared to an object (O) during social preference portion of the 3CSI Test (D). Both genotypes did not exhibit a preference for social novelty as the time spent exploring a novel stranger mouse (S2) compared to a familiar stranger mouse (S1) did not differ significantly (E). Values represent mean ± SEM (n = 9-12 animals per genotype). (F-G) PI for a novel object during STM (F) or LTM (G) portions of object recognition test was not significantly different between the genotypes. Values represent mean ± SEM (n = 9-12 animals per genotype). (H-J) Target quadrant occupancy (H), frequency of platform crossings during 60 s probe trial (I) or during flag test (J) were not significantly different between *Fmr1*-^{*i*}y-SOM and WT-SOM mice. Values represent mean ± SEM (n = 9-12 animals per genotype); ***p < 0.001, two-way ANOVA or RM two-way ANOVA followed by Bonferroni's multiple comparisons test, Student's *t* test.



Figure S5. Phosphorylation of ribosomal protein S6 (rpS6) was elevated in PV-positive neurons in the hippocampus but not mPFC of WT and *Fmr1*-^{*t*}y-PV mouse. (A) Representative images of PV, rpS6 phosphorylated at serine 235 and 236 (p-S6 235/6) and neuronal marker NeuN staining from WT-PV and *Fmr1*-^{*t*}y-PV mouse hippocampus. (B-C) Quantification of p-S6 235/6 in PV-positive neurons in the hippocampus (B) and mPFC (C) of WT-PV and *Fmr1*-^{*t*}y-PV mouse. Values represent mean ± SEM (hippocampus, n = 20-23 z-stacks, 9 sections, from 3 animals per genotype; mPFC, n = 20-27 z-stacks, 9 sections, from 3 animals per genotype); *p < 0.05 (Student's *t* test). Scale bar: 100 µm.