Supplementary information

## TDP-43 proteinopathy impairs mRNP granule mediated postsynaptic

## translation and mRNA metabolism

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Supplementary Figure S2 is associated to Figure 2. Supplementary Figure S3 is associated to Figure 7. Supplementary Figure S4 is associated to Figure 8.



**Supplementary Figure S1.** The expression profile of TDP-43 protein in the mouse CNS. (A) The representative immunofluorescent images showed the distribution of TDP-43 in the mouse CNS, including the cortical layers and the hippocampus. Scale bar: 500  $\mu$ m (B) The representative images showing the sharp-line signals in the hippocampal area indicated a predominantly somatic localization of TDP-43.by co-staining of TDP-43, neuron marker (NeuN) and nuclear marker (DAPI). Scale bar: 100  $\mu$ m (C) Co-staining of TDP-43 and glial marker GFAP showing few colocalization.



**Supplementary Figure S2.** The localization and measured distances of the respective protein pairs. The representative images and its corresponding histograms were displayed the position of COMs and distance of PSD-95, Bassoon and TDP-43 by calculated from intensity profiles along the transsynaptic axis. Scale bar: 400 nm



**Supplementary Figure S3.** The time-dependent formation of the pathological TDP-43 inclusions under proteasome inhibitor application. The HEK293T cells with TDP-43-GFP transfection were observed by confocal microscopy at 0, 4, 8 and 12 hours after MG-132 treatment. No transfection and TDP-43-GFP expression HEK 293T cells treated with vehicle (Veh) were served as control group, Scale bar: 100 µm



**Supplementary Figure S4.** The cerebral cortex images with TDP-43 and FMRP of all five FTLD-TDP patients. The immunostaining of neuronal mRNP granule marker, FMRP (green), and cytosolic TDP-43 aggregates (Red) were displayed high-degree colocalization (yellow). DAPI was served as nucleus signal. Scale bar: 50  $\mu$ m and 10  $\mu$ m (magnified images).