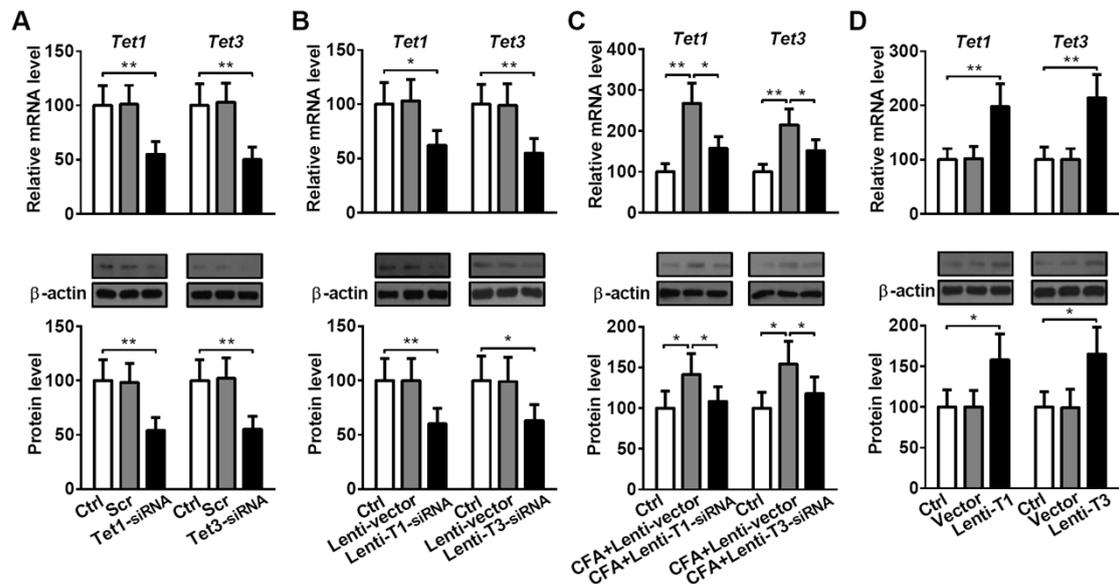
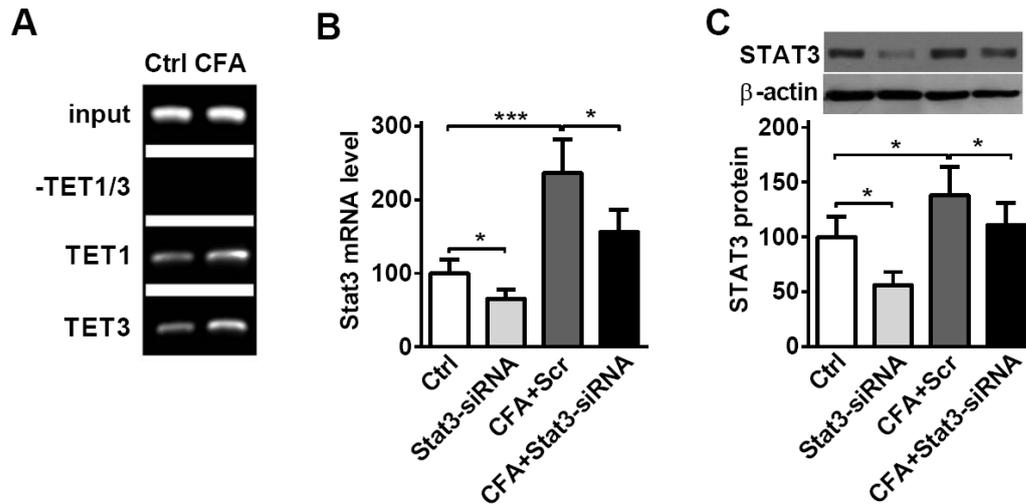


**Figure S1.** Genomic mapping and features of 5-hmC in spinal cord of control and CFA-treated mice. (A) Normalized densities of 5hmC reads across the genome-wide transcript unit of all reference genes from transcription start sites (TSS) to transcription ending sites (TES), as well as up- and downstream regions, in spinal cord from control and CFA mice. The coverage of up- and downstream interpolated 5000 bp sequence is also shown. 5-hmC densities were normalized to the total number of aligned reads from each sample (in millions). (B) Heatmap of 5-hmC levels of RefSeq genes. Two distinct clusters of genes were identified based on the dynamic changes of their 5-hmC levels in spinal cord. The heatmap represents the normalized 5-hmC density of each gene.



**Figure S2.** The validation of transfection efficiency of modulation tool *in vivo*. (A-C) The expression of *Tet1* and *Tet3* in mRNA and protein after the intrathecal injection of exogenous siRNA in naïve mice (A), endogenous siRNA mediated by lentivirus in naïve mice (B) or CFA mice (C).  $n=6/\text{group}$ . The injection of siRNA or Lenti-siRNA in naïve mice:  $*p<0.05$ ;  $**P < 0.01$  versus the corresponding control or scramble or Lenti-vector group by two-tailed unpaired Student's *t* test. The injection of Lenti-siRNA in naïve mice: One-way ANOVA (expression vs. the treated groups) followed by post *hoc* Tukey test, *Tet1* mRNA:  $F(2, 15) = 43.3$ , *Tet3* mRNA:  $F(2, 15) = 30.72$ ; *TET1* protein:  $F(2, 15) = 13.24$ ; *TET3* protein:  $F(2, 15) = 17.85$ ;  $*P < 0.05$ ,  $**P < 0.01$ . (D) The expression of *Tet1* and *Tet3* mRNA and protein after the intrathecal injection of lentivirus overexpressing *TET1* and *TET3*. The expression of mRNA and protein was measured at 1 day after 2 consecutive days injection of siRNA or at 2 d after 3 consecutive days injection of lentivirus in naïve mice or CFA mice.  $n=6/\text{group}$ ,  $*p<0.05$ ;  $**p<0.01$  versus the corresponding control or vector group by two-tailed unpaired Student's *t* test.



**Figure S3.** The binding of TET1 and TET3 with Stat3 promoter and the validation of Stat3 knockdown tool. (A) The capacity of TET1 and TET3 could bind to *Stat3* promoter in CFA group versus control by chromatin immuno-precipitation (ChIP-sqPCR) method. In input group, genome DNA was used as PCR template. -TET1/3, without TET1 and TET3 antibody in ChIP experiment. (B-C) The expression of *Stat3* in mRNA and protein after the intrathecal injection of siRNA in naïve mice (B) or CFA mice (C). n=5/group; One-way ANOVA (expression vs. the treated groups) followed by post *hoc* Tukey test, mRNA:  $F(3, 16) = 37.22$ ; protein:  $F(3, 16) = 18.48$ ; \* $p < 0.05$ ; \*\*\* $p < 0.001$ .