## CVR-2010-567R2

**Supplementary Figure 1**. Acute central administration of Ex-4 does not alter activity. Mice received either aCSF or Ex-4 through an ICV cannula. (A) Activity was unchanged during acute Ex-4 infusion. (B) Average activity during the dark and light period during the first 24 hours of Ex-4 infusion. Chronic ICV infusion of Ex-4 did not alter activity during either the dark (C) or light (D) periods. Arrow indicates beginning of Ex-4 infusion.

**Supplementary Figure 2**. Central administration of Ex-4 depresses temperature. Mice received either aCSF or Ex-4 through an ICV cannula. Temperature was significantly depressed. (A) Body temperature throughout the first 24 hours Ex-4 infusion. (B) Average body temperature was diminished during the dark period. Chronic ICV Ex-4 infusion significantly reduced temperature during both the dark (C) and light (D) periods. Arrow indicates beginning of Ex-4 infusion. \*p < 0.05; \*\*\* p < 0.001; significant difference between groups; two-way ANOVA with repeated measures

**Supplementary Figure 3.** Chronic ICV infusion of Ex-4 significantly reduced body weight (A) and blood glucose levels (B) over 28 days. \* p < 0.05; two way ANOVA between groups, Bonferroni post-hoc test.

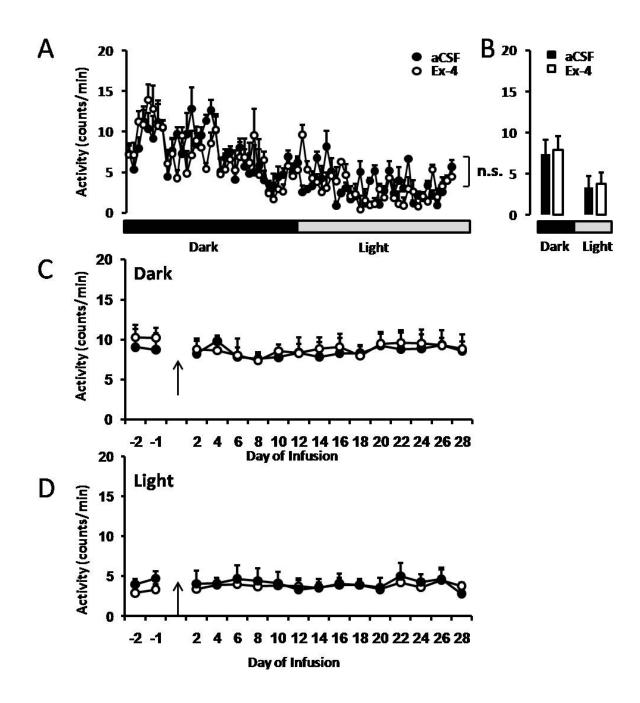


Figure S1

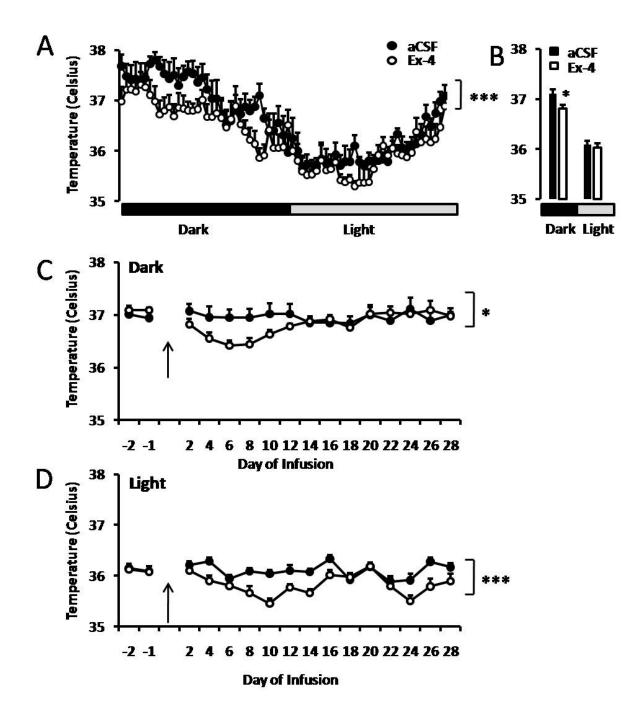


Figure S2

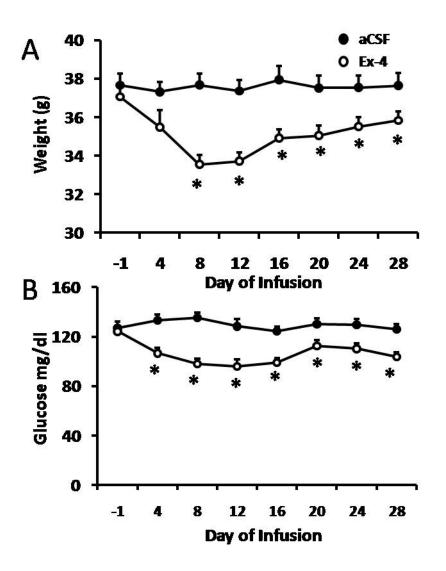


Figure S3