Supplemental information

Histone malonylation is regulated by SIRT5 and KAT2A

Ran Zhang, Joanna Bons, Grace Scheidemantle, Xiaojing Liu, Olga Bielska, Chris Carrico, Jacob Rose, Indra Heckenbach, Morten Scheibye-Knudsen, Birgit Schilling, and Eric Verdin

Supplemental figures

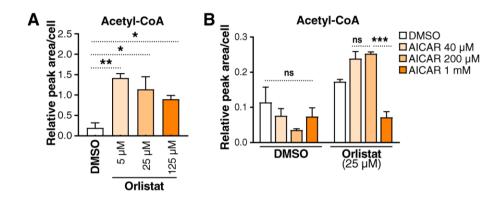


Figure S1. Acetyl-CoA measurement, Related to Figure 1. Acetyl-CoA levels in K562 cells treated with orlistat (A) or AICAR (B) at different doses as indicated for 24 h were measured with LC-MS. Quantified levels are normalized to cell count in each sample. N = 3 per treatment. Values are shown as mean \pm SEM. ns (not significant) >= 0.05, *P < 0.05, *P < 0.01, and ***P < 0.001 using unpaired student t test.

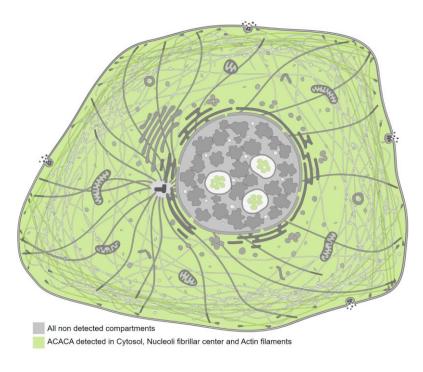


Figure S2. ACC1 subcellular localization (the HPA database, proteinatlas.org), Related to Figure 5.

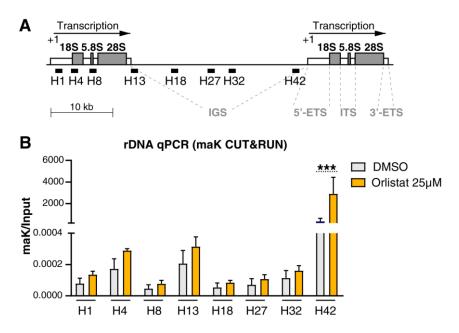


Figure S3. Lysine malonylation (maK) CUT&RUN, Related to Figure 5.

- (A) Schematic representation of a human rDNA repeat with 8 primer pairs (solid bars) and their approximate positions relative to the transcription start site are indicated. IGS, intergenic spacer. 5'and 3'-ETS, 5'and 3'-external transcribed spacer. ITS, internal transcribed spacer.
- (B) K562 cells treated with DMSO or 25 μ M or listat for 24 h were used for CUT&RUN of maK. The malonylation levels at 8 regions of rDNA were quantified using qPCR. 3 biological replicates per treatment. Values are shown as mean \pm SEM. ***P < 0.001 using two-way ANOVA with Sidak multiple comparison.

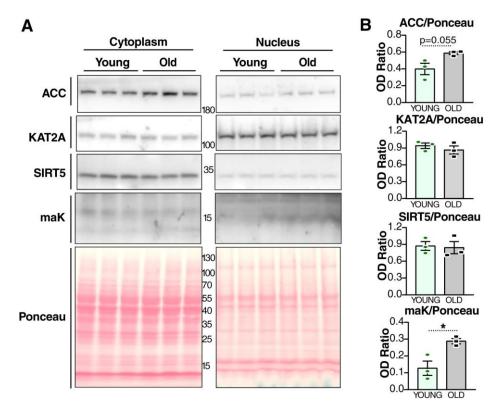


Figure S4. Western blotting of cytoplasmic and nuclear proteins, Related to Figure 6.

- (A) Western blot of cytoplasmic and nuclear protein samples prepared from young (3-6 months old) and middle-aged (12-14 months old) female mouse whole brains.
- (B) Quantification of the ODs of blots in the young and elder nuclear protein samples. n = 3 per group. Error bar: mean \pm SEM, *p<0.05 with unpaired student t test.

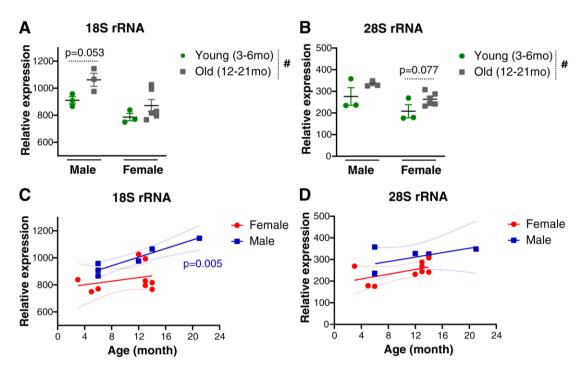


Figure S5. Ribosomal RNAs increase with age in mouse brain, Related to Figure 6.

(A and B) 18S and 28S rRNA in young (3-6 months old) and old (12-21 months old) male and female mouse brain tissues was quantified through real-time qPCR normalized to β -actin expression level. N= 3-6 per group. Error bar: mean \pm SEM. P values were indicated using unpaired student t test; #p<0.05 using two-way ANOVA.

(C and D) Correlation between rRNA expression level and age of the mouse was analyzed using linear regression, with 95% confidence interval shown between light lines. Data were gender-stratified. P value was shown as indicated.