

Supplementary Material

Manuscript title:

Role of the CLOCK protein in liver detoxification

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Table S1. Primer sequences for quantitative real-time PCR

Gene	Forward sequence	Reverse sequence
<i>Cyp2a4</i>	AACGGTGCTTTCATTGACCC	CCTCATAGTCAAAGCGGTCCC
<i>Cyp2a5</i>	GGACAAAGAGTTCCTGTCACTGCTTC	GTGTTCCACTTTCTTGTTATGAAGTCC
<i>Cyp2j9</i>	ATGCGCCTTCCTTTCGTGG	CCAGGCTTAGAACATTCCCCTA
<i>Cyp39a1</i>	ATCCAGCCACACTCAATACCC	GGAGCCATACTCAAAGCCCTC
<i>Cyp4a10</i>	TTCCCTGATGGACGCTCTTA	GCAAACCTGGAAGGGTCAAAC
<i>Cyp2c37</i>	CACGAGGCGTTTCTCACTCA	AGGGCTGCTCAGAATCTTTGT
<i>Cyp2b10</i>	TGCTGTCGTTGAGCCAACC	CCACTAAACATTGGGCTTCCT
<i>Cyp2c55</i>	AATGATCTGGGGGTGATTTTCAG	GCGATCCTCGATGCTCCTC
<i>Cyp3a25</i>	CTTCACTGTCCAGCCTTGTGAA	AATTGGTTCCTGCTGATCTTC
<i>Cyp3a11</i>	CGCCTCTCCTTGCTGTCACA	CTTTGCCTTCTGCCTCAAGT
<i>Cyp2c70</i>	AGTATGGCCCTGTGTTTACTGT	GCCTTGGCTGGTTCTACTGAG
<i>Cyp4a12b</i>	GGGGAGATCAGACCCAAAAGC	ATTCGTCGGTGCTGAAACCAT
<i>Cyp2c40</i>	TCCGGTTTTTGACAAGGTTTCTAC	TGCCCAAGTTCCTCAAGGTATTC
<i>Cyp2j6</i>	ATGCTCGCTGCTACCGGCTC	GTGCTTCTTTGATTAAGGGC
<i>Cyp4f14</i>	ACTGGCTTATGGGTCACGTG	ACCCACCAAACGAGTCAATTC
<i>Ugt1a1</i>	GCTTCTTCCGTACCTTCTGTTG	GCTGCTGAATAACTCCAAGCAT
<i>Ugt1a5</i>	TGAGAAGGTGCTAGTGTTTCCT	GGGAACGGCATAGACTTTGAA
<i>Ugt1a9</i>	TTTCGATGTGTGCGGCTAAC	GGTCCGAGTTCTTTCCTTGAA
<i>Sult1a1</i>	CACAAGGGTCCTCTCCTTAGC	TGACAGCGGAACGTGAAGTC
<i>Sult2a7</i>	TTGTTATGAGAGAAGACACAGTCGT	GCAGACAATCTCAACAAGCCAG
<i>Clock</i>	CCAGAGGGAGAACATTCA	TGGCTCCTTTGGGTCTAT
<i>Dbp</i>	ACATCTAGGGACACACCCAGTC	AAGTCTCATGGCCTGGAATG
<i>Car</i>	CCCTGACAGACCCGGAGTTA	GCCGAGACTGTTGTTCCATAAT
<i>Bmal1</i>	CTCCAGGAGGCAAGAAGATTC	ATAGTCCAGTGGAAGGAATG
<i>Rev-erba</i>	TTTTTCGCCGGAGCATCCAA	ATCTCGGCAAGCATCCGTTG
<i>Rev-erbβ</i>	GGAGTTCATGCTTGTGAAGGCTGT	CAGACACTTCTTAAAGCGGCACTG
<i>Npas2</i>	GAACATTCCGAAGTTTAT	AATCGTTGTCAGATTTTAG
<i>Hmbs</i>	CCGAGCCAAGGACCAGGATA	CTCCTTCCAGGTGCCTCAGA

Table S2. Oligonucleotide sequences for EMSA assays

Oligonucleotide	Sequence
Cyp2a5-E-box (-1729/-1696 bp)	CTTCATTCAGGCAATTCCAGTGACCACCTGCTT
Cyp2b10-RevRE (-2346/-2319 bp)	GACTTAGGAGGAAGGTCAGAAAAACAT

Table S3. Primer sequences for ChIP assays.

Primer	Forward sequence	Reverse sequence
Cyp2a5-CLOCK	GTCTCAGAGTCCAACAGCCTAAAAC	TTTGGTTCCAGCAGGTTC
Cyp2a5-Distal	AAAGGCAGATTGAAGTTTAG	TTCCTCCTGATAGTAATGGT
Cyp2b10-REV-ERB α	CACATCTGTGGTCCCAGTA	TCCCAGGTGTCAGGATTCAG
Cyp2b10-Distal	GAAGTTCTGCTGTGGGTC	GTAACATTACTATCTAGCCTCT

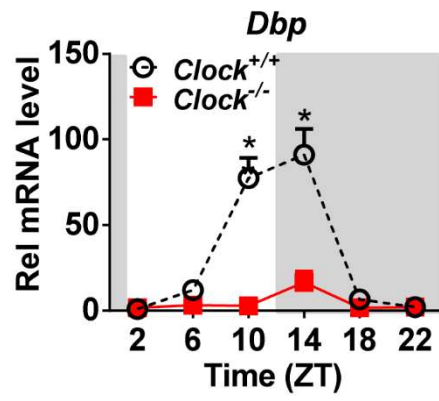


Fig S1 Expression of *Dbp* in *Clock*^{+/+} and *Clock*^{-/-} mice. Data are mean \pm SD ($n = 5$).

* $p < 0.05$ (two-way ANOVA with Bonferroni post hoc test).

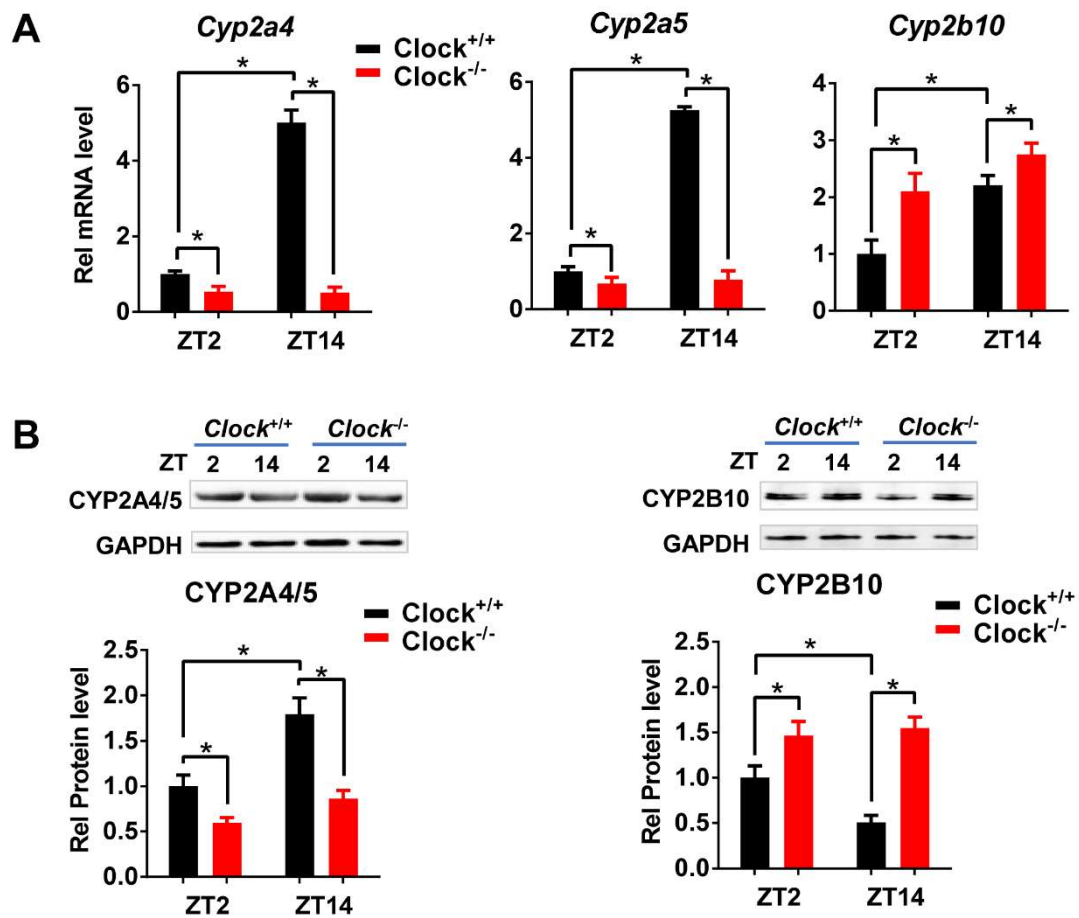


Fig S2 Hepatic mRNA and protein expression of CYP2A4/5 and CYP2B10 in female *Clock*^{+/+} and *Clock*^{-/-} mice. Data are mean \pm SD ($n = 5$). * $p < 0.05$ (two-way ANOVA with Bonferroni post hoc test).

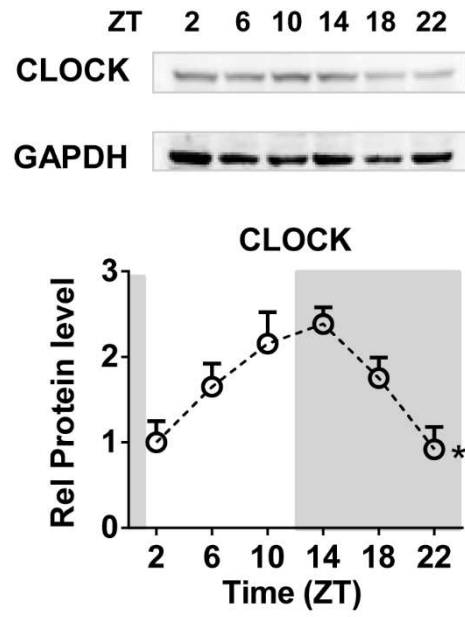


Fig S3 Circadian expression of CLOCK protein in mouse liver. Data are mean \pm SD ($n = 5$). * $p < 0.05$ (one-way ANOVA with Bonferroni post hoc test).

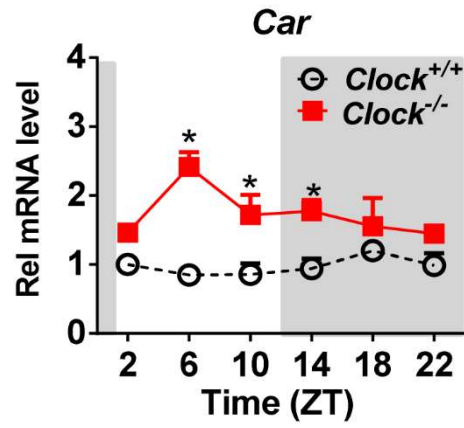


Fig S4 Expression of *Car* in *Clock*^{+/+} and *Clock*^{-/-} mice. Data are mean \pm SD ($n = 5$). * $p < 0.05$ (two-way ANOVA with Bonferroni post hoc test).

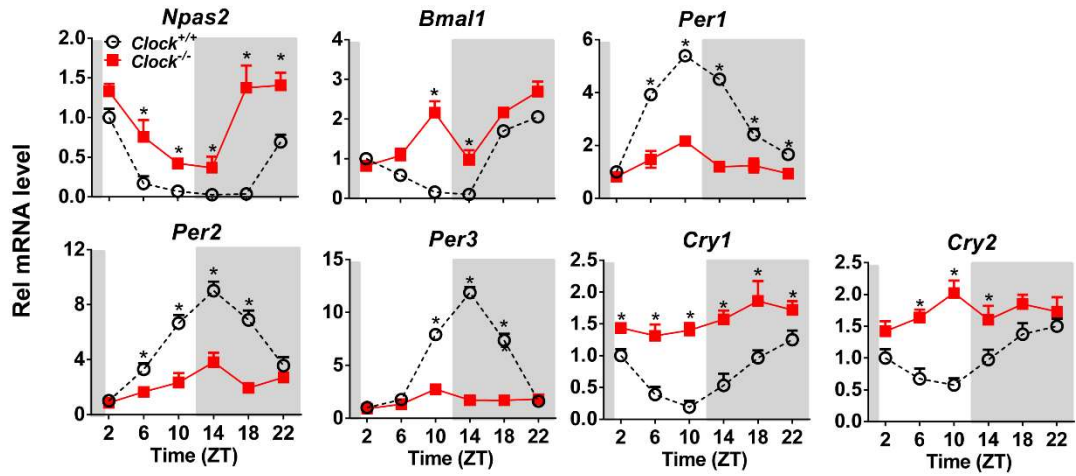


Fig S5 Core clock genes expressions in *Clock*^{+/+} and *Clock*^{-/-} mice. Data are mean ± SD ($n = 5$). * $p < 0.05$ (two-way ANOVA with Bonferroni post hoc test).

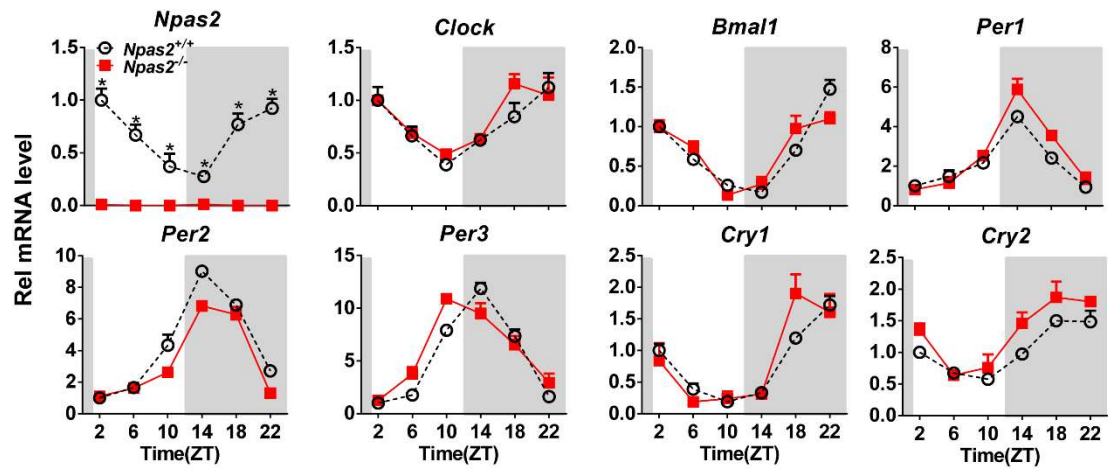


Fig S6 Core clock genes expressions in *Npas2*^{+/+} and *Npas2*^{-/-} mice. Data are mean \pm SD ($n = 5$). * $p < 0.05$ (two-way ANOVA with Bonferroni post hoc test).