



Supplemental Figure 1. The binucleated cell number count in regenerating livers (Post 2/3PHx surgery day 4) from WT vs. $ER\alpha$ -KO vs. $ER\beta$ -KO.

Supplemental Table 1. Genes and pathway enrichment analysis of ERa with KEGG platform

ERaKO vs Wildtype

KEGG pathway of PHx mice microarray

Term	Link	Count	%	P Value	Genes
mmu01040:Biosynth esis of unsaturated fatty acids	Biosynthesis of unsaturated fatty acids	5	0.75	0.003	SCD1, 1600014K23RIK, SCD3, SCD2, ACOT3
mmu04114:Oocyte meiosis	Oocyte meiosis	8	1.20	0.016	CCNB1, ADCY3, CDK1, PLK1, RPS6KA2, BUB1, CDC20, AURKA
mmu04914:Progeste rone-mediated oocyte maturation	Progesterone-mediated oocyte maturation	6	0.90	0.045	CCNB1, ADCY3, CDK1, PLK1, RPS6KA2, BUB1
mmu04540:Gap junction	Gap junction	6	0.90	0.047	ADCY3, CDK1, TUBB2A-PS2, TUBB2A, TUBB6, TUBB3
mmu00830:Retinol metabolism	Retinol metabolism	5	0.75	0.071	CYP2C37, CYP26B1, CYP26A1, CYP2B10, CYP2A4
mmu04115:p53 signaling pathway	p53 signaling pathway	5	0.75	0.074	CCNB1, CDK1, CDKN1A, BAI1, GADD45A
mmu04110:Cell cycle	Cell cycle	7	1.05	0.074	CCNB1, CDK1, CDKN1A, PLK1, BUB1, CDC20, GADD45A

Supplemental Table 2. Genes and pathway enrichment analysis of ERbeta with KEGG platform

ERβKO vs Wildtype

KEGG pathway of PHx mice microarray

Term	Link	Count	%	<i>P</i> Value	Genes
mmu00590:Arachi donic acid metabolism	Arachidonic acid metabolism	11	1.95	1.61E- 05	CYP2C37, CYP2C54, CYP2B13, GGT1, CYP2C50, CYP4A10, ALOX15, CYP4A32, CYP4A31, CYP2C38, CYP4A14, PLA2G2E, CYP2C39
mmu03320:PPAR signaling pathway	PPAR signaling pathway	9	1.59	3.98E- 04	CYP4A10, SCD3, SCD2, CYP4A32, CYP4A31, FABP3, UCP1, CYP4A14, CYP8B1, ADIPOQ, ACSL6
mmu00830:Retinol metabolism	Retinol metabolism	8	1.42	8.38E- 04	CYP4A10, CYP2C37, CYP2C54, CYP4A32, CYP4A31, CYP2B13, CYP4A14, CYP2C38, CYP2C39, CYP2C50
mmu00591:Linolei c acid metabolism	Linoleic acid metabolism	6	1.06	3.72E- 03	CYP2C37, CYP2C54, PLA2G2E, CYP2C38, CYP2C39, CYP
mmu00980:Metab olism of xenobiotics by cytochrome P450	Metabolism of xenobiotics by cytochrome P450	7	1.24	3.74E- 03	CYP2C37, CYP2C54, ALDH1A3, CYP2B13, CYP2C38, CYP2C39, CYP2C50
mmu00982:Drug metabolism	Drug metabolism	7	1.24	7.01E- 03	CYP2C37, CYP2C54, ALDH1A3, CYP2B13, CYP2C38, CYP2C39, CYP2C50
mmu04740:Olfact ory transduction	Olfactory transduction	34	6.02	8.22E- 03	OLFR1370, OLFR315, OLFR611, OLFR805, OLFR823, OLFR690, OLFR1157, OLFR362, OLFR1506, OLFR945, OLFR1032, OLFR575, OLFR384, OLFR533, OLFR676, OLFR843, CAMK2A, OLFR899, OLFR874, OLFR282, OLFR954, OLFR441, OLFR716, GUCY2D, OLFR620, OLFR1145, OLFR665, OLFR608, OLFR584, OLFR644, OLFR1101, OLFR12, OLFR978, OLFR830
mmu05320:Autoim mune thyroid disease	Autoimmune thyroid disease	5	0.88	8.15E- 02	IFNAB, CGA, IFNA5, H2-M1, H2-DMB2

Suppl. Fig. 2, Kao et al

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Sequence logo	Consensus binding sequence derived from Positional Weight Matrix	Matrix type	Matrix category
TGACC	V\$ER_Q6	factor-specific	matrix compiled from individual genomic sites
	V\$ER_Q6_02	family	matrix compiled from individual genomic sites
AGGTCATGACCT 0	V\$ERALPHA_01	factor-specific	ChIP-on-chip
ТGACc 0	V\$ERALPHA_Q6_01	factor-specific	matrix compiled from individual genomic sites
 T G ACC _ ⊕	V\$ERALPHA_Q6_02	factor-specific	matrix compiled from individual genomic sites
────TGACc _ @	V\$ERALPHA_Q4	factor-specific	matrix compiled from individual genomic sites

ERα (6 mouse)	
V\$ER_Q6	CAGGTCACTGTGACCTGA
V\$ER_Q6_02	CAGGTCACGGT
V\$ERALPHA_01	AGGTCACAGTGACCT
V\$ERALPHA_Q6_01	TGACCTG
V\$ERALPHA_Q6_02	GGTCACGTGACCTG
V\$ERALPHA_Q4	GTCACCGTGACCT

ERβ (3 mouse)	
V\$ER_Q6_02	CAGGTCACGGT
V\$ERBETA_Q5	GTCAGAGTGACCCAG
V\$ERBETA_Q5_01	ACAGTGACCCGGAA

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	Sequence logo	Consensus binding sequence derived from Positional Weight Matrix	Matrix type	Matrix category
	▲GGTCA •	V\$ER_Q6_02	family	matrix compiled from individual genomic sites
	<u>TGACC</u> ⊙	V\$ERBETA_Q5	factor-specific	matrix compiled from individual genomic sites
	Τ<mark>GACC</mark>Θ	V\$ERBETA_Q5_01	factor-specific	matrix compiled from individual genomic sites

Supplemental Figure 2. The sequences and scores of ER α -EREs and ER β -EREs.

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Supplemental Figure 3. The ER α - and ER β -interactome enrichment analysis with the GSEA platform.