



**Supplementary Figure 1**

**Supplementary Figure 1 - Generation of  $\Delta$ leftPAL and  $\Delta$ IRIS mice.** Upper part (A+B): generation of  $\Delta$ leftPAL mice. A: Gene-targeting vector was constructed in order to replace the genomic fragment encompassing the hs3a+hs1,2 part of the 3'RR with a loxP/neoR gene cassette in ES cells. Mutant ES clones allowed the derivation of Neo mutant mice. Neo mutant mice allowed the derivation of  $\Delta$ leftPAL animals after cre-deletion of the neo gene. B: A specific PCR was performed to ensure correct 5' recombination (P1+P2) (amplicon of 3.7 kb) in ES cells. A specific PCR was performed to ensure correct 3' recombination (P3+P4) (amplicon of 10 kb) in ES cells. A specific PCR was performed to ensure correct Cre recombination (P5+P6) (amplicon of 2 kb vs 0.5 kb after Cre deletion) in Neo mutant mice. A specific PCR was performed to ensure homozygous  $\Delta$ leftPAL mice (P9+P10) (lack of a 0.2 kb amplicon). Lower part (C+D): Generation of  $\Delta$ IRIS mice. C: Same protocol that in the upper part except that the loxP/neoR gene cassette reintroduced inverted hs3a and hs1,2 enhancers. D: A specific PCR was performed to ensure correct 5' recombination (P1+P2) (amplicon of 3.7 kb) in ES cells. A specific PCR was performed to ensure correct 3' recombination (P3+P4) (amplicon of 10 kb) in ES cells. A specific PCR was performed to ensure correct Cre recombination (P5+P6) (amplicon of 3 kb vs 1.4 kb after Cre deletion) in Neo mutant mice. A specific PCR was performed to ensure homozygous  $\Delta$ IRIS mice (P7+P8) (lack of a 0.15 kb amplicon).

### Screening

P1	5'-GGA GGG AGA AAT ACC ACC-3'	
P2	5' AGC ACG TAC TCG GAT GGA AG 3'	
P3	5'GTG CAA TCC ATC TTG TTC AAT GGC CGA TCC 3'	
P4	5' CGG CTC TAA CAA CTG GGT CCT GTA C 3'	
P5	5'-GGT TAA TCA GTT GGG GCT GA-3'	
P6	5'-AGG TGC ATG ACA GGG TAA GG-3'	
P7	5'-GGT GAC ATT AAG CTT GGG GAT A-3'	
P8	5'-CCC TGA GAG GTT TCA CAT TTT C-3'	
P9	5'-ACC ACT GCT GGA ATC TGA CC-3'	
P10	5'-CAG GCC AAG ACT TTT CCA GT-3'	

### SHM analysis

VHJ558 SHM	5'-CAG CCT GAC ATC TGA GGA CTC TGC-3'	
SHM H4 3'	5'-CAG CAA CTA CCC TTT TGA GAC CGA-3'	

### ChIP analysis

ChIP A Fw	5'-CAG TCT CCT CAG GTG AGT CCT-3'	Maul et al, 2014
ChIP A Rev	5'-CCC AAT GAC CCT TTC TGA CT-3'	
ChIP B Fw	5'-GGA GAG CTG TCT TAG TGA TTG AGT CAA GGG-3'	
ChIP B Rev	5'-GTG TTC CTT TGA AAG CTG GAC-3'	
S $\mu$ -U Fw	5'-TCTAAATGCGCTAAACTGAGG-3'	Wang et al., 2009
S $\mu$ -U Rev	5'-AGCGTAGCATAGCTGAGCTC-3'	
I $\gamma$ 3 Fw	5'-AGATCCCAAAGCTAAGCTCCTG-3'	Wang et al., 2006
I $\gamma$ 3 Rev	5'-CAGAGAGACCCCTCCACAGT-3'	
S $\gamma$ 3-U Fw	5'-GCTGAGAGTATGCACAGCCA-3'	
S $\gamma$ 3-U Rev	5'-GGATCATGGAACTCCTCCG-3'	
S $\gamma$ 3-D Fw	5'-CAGGCTGGGAACTCTTGGG-3'	
S $\gamma$ 3-D Rev	5'-AGTTAGTCCCCATACTTGAACC-3'	
I $\gamma$ 2b Fw	5'-TATTGTTCTGGCTGCAAATGG-3'	Li et al, 2013
I $\gamma$ 2b Rev	5'-GTGCAGGCATATCTTCCATCC-3'	
S $\gamma$ 2b-U Fw	5'-AGCTCCAAAAGCTCAGCAGAC-3'	
S $\gamma$ 2b-U Rev	5'-AGCCCCAGCTTACAAAGAGCT-3'	
S $\gamma$ 2b-D Fw	5'-GGTGGGAATATGAGGGAGAAGTCCTAG-3'	
S $\gamma$ 2b-D Rev	5'-TTCCACCTGCCTCAGCTCTCCACAGC-3'	

### GLT Analysis

I $\mu$ -C $\mu$ -Fwd	5'-ACCTGGGAATGTATGGTTGTGGCTT-3'	Park et al, 2009
I $\mu$ -C $\mu$ -Rev	5'-TCTGAACCTTCAAGGATGCTCTTG-3'	
I $\gamma$ 3-C $\gamma$ 3-Fwd	5'-AACTACTGCTACCACCACCACCAG-3'	
I $\gamma$ 3-C $\gamma$ 3-Rev	5'-ACCAAGGGATAGACAGATGGGG-3'	
I $\gamma$ 1-C $\gamma$ 1-Fwd	5'-GGCCCTCCAGATCTTTGAG-3'	
I $\gamma$ 1-C $\gamma$ 1-Rev	5'-ATGGAGTTAGTTTGGCAGCA-3'	
I $\gamma$ 2b-C $\gamma$ 2b-Fwd	5'-CCAACCAGGAAGAGTCCAGAG-3'	
I $\gamma$ 2b-C $\gamma$ 2b-Rev	5'-ACAGGGATCCAGAGTTCCAAGT-3'	
I $\gamma$ 2a-C $\gamma$ 2a-Fwd	5'-GCTGATGTACCTACCGAGAGA-3'	
I $\gamma$ 2a-C $\gamma$ 2a-Rev	5'-GCTGGGCCAGGTGCTCGAGGTT-3'	
I $\alpha$ -C $\alpha$ Fw	5'-CTACCATAGGGAAGATAGCCT-3'	Park et al, 2005
I $\alpha$ -C $\alpha$ -Rev	5'-TAATCGTGAATCAGGCAG-3'	

**Supplementary table 1** - PCR primers used for the screening of  $\Delta$ leftPAL and  $\Delta$ IRIS mice, for real-time PCR analysis and sequencing

## Supplementary References

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