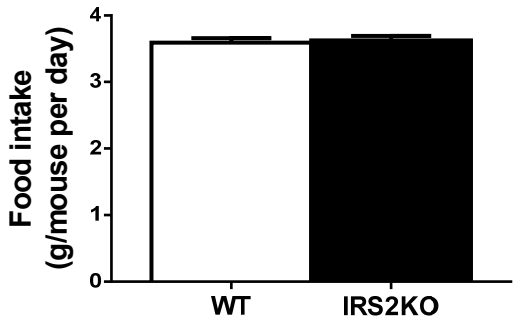

Figure S1. Mean daily food intake. Experimental groups: Wild-type (WT) and *Irs2*-deficient (IRS2KO) female mice (n=8 animals per condition).

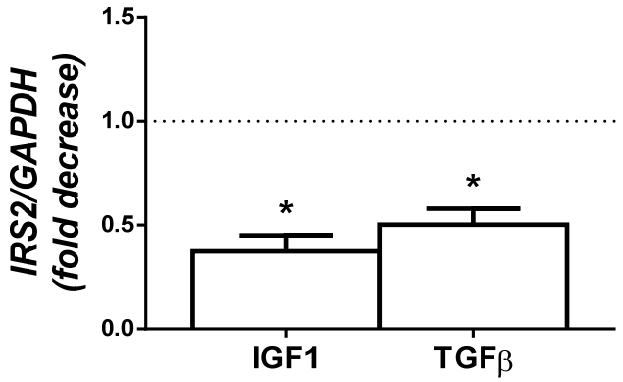

Figure S2. LX2 cells were treated with 10 nM IGF1 or 5 ng/ml TGFβ for 24 h. *IRS2* mRNA levels determined by RT-qPCR. Statistical significance was carried out by unpaired two-tailed Student's t-test: *p<0.05, IGF1 or TGFβ vs. untreated cells (n=3 independent experiments performed in duplicate).

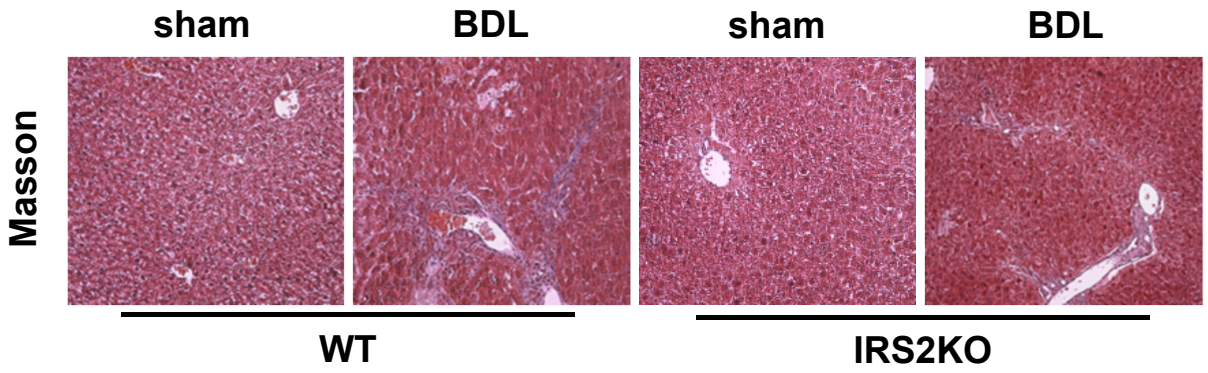

Figure S3. Representative images (20x) from Masson`s trichrome staining. Experimental groups: Wild-type (WT) and *Irs2*-deficient (IRS2KO) female mice submitted to BDL and analyzed 28 days after surgery (n=8 animals per condition).

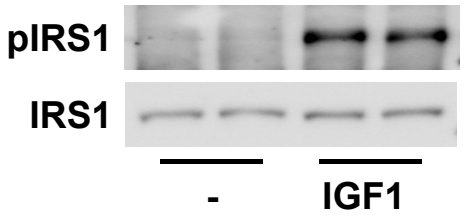

Figure S4. LX2 cells were treated with 10 nM IGF1 for 15 min and IRS1 tyrosine phosphorylation was analyzed by Western Blot. Representative images are shown (n=3 independent experiments performed in duplicate).

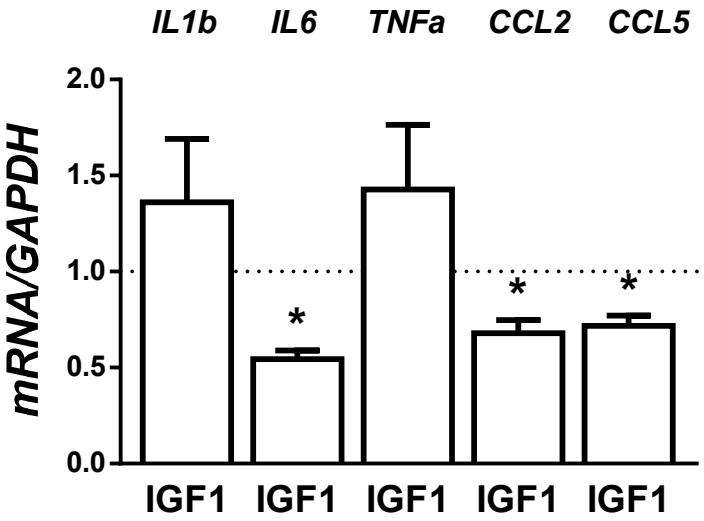

Figure S5. LX2 cells were treated with 10 nM IGF1 for 24 h. mRNA levels of pro-inflammatory cytokines determined by RT-qPCR. Statistical significance was carried out by unpaired two-tailed Student's t-test: *p<0.05, IGF1 vs. untreated cells (n=3 independent experiments performed in duplicate).

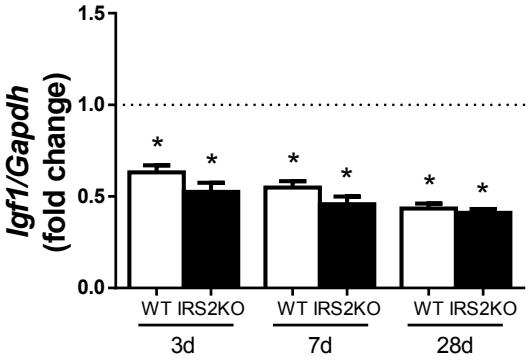

Figure S6. *Igf1* mRNA levels determined by RT-qPCR. Experimental groups: Wild-type (WT) and *Irs2*-deficient (IRS2KO) female mice submitted to BDL and analyzed at the indicated time-periods. Statistical significance was carried out by one-way ANOVA with Bonferroni post-test: *p<0.05, BDL vs. Sham (n=6-8 animals per condition).

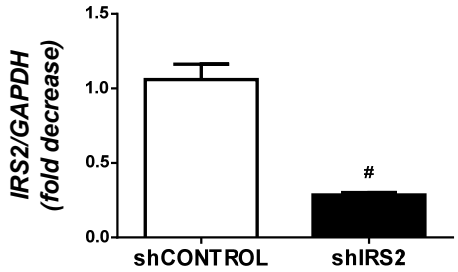

Figure S7. LX2 cells infected with scrambled (shC) or IRS2 shRNA (shIRS2) lentiviral particles. *IRS2* mRNA levels determined by RT-qPCR. Statistical significance was carried out by unpaired two-tailed Student's t-test: #p<0.05, shIRS2KO vs. shC (n=4 independent experiments performed in duplicate).

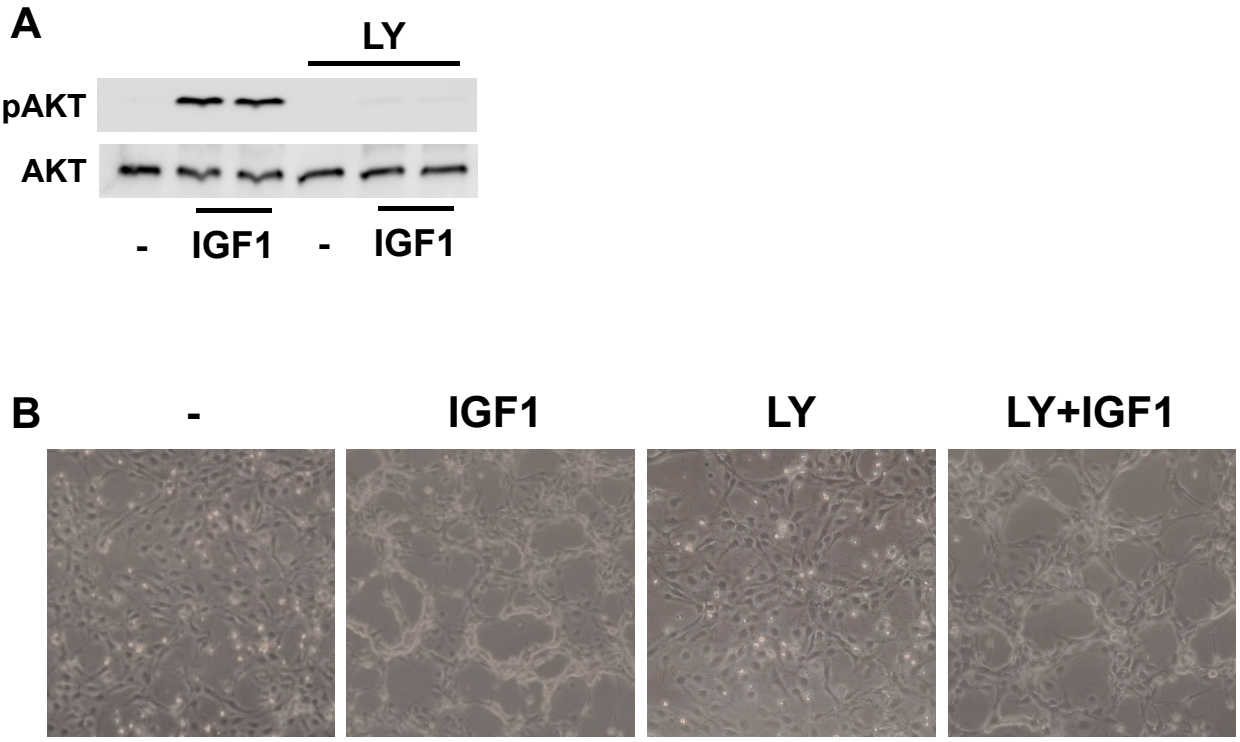

Figure S8. LX2 cells were pretreated with 25 μM LY294002 for 2 h prior to 10 nM IGF1 stimulation for 15 min (A) or 24 h (B). **A**. Representative Western blot with the indicated antibodies. **B**. Representative phase-contrast images. (n=3 independent experiments performed in duplicate).

Table S1. *Primer sequences for RT-qPCR.*

m-Gapdh	Forward	TGGAAGAGTGGGAGTTGCTGT
	Reverse	CCTGGAGAAACCTGCCAAGTA
m-Acta2	Forward	CCCAGACATCAGGGAGTAATGG
	Reverse	TCTATCGGATACTTCAGCGTCA
m-Mmp9	Forward	AAACCTTCAAAGGCCTCAAGTG
	Reverse	GTCTCGCGGCAAGTCTTCAG
m-Pai1	Forward	ATGACTGGGTGGAAAGGCATAC
	Reverse	CAGGCGTGTCAGCTCGTCTA
m-Hmox1	Forward	CACAGATGGCGTCACTTCGTC
	Reverse	GTGAGGACCCACTGGAGGAG
m-Il6	Forward	GAGGATACCACTCCCAACAGACC
	Reverse	AAGTGCATATCGTTGTTCATACA
m-Il1b	Forward	AGAAGCTGTCGCAGGTACCTG
	Reverse	GGAAAAGAAGGTGCTCATGTCC
m-Tnfa	Forward	CATCTTCTCAAAATTCGAGTGACAA
	Reverse	TGGGAGTAGACAAGGTACAACCC
m-Irs2	Forward	CACGAGCCCGTAGTTGTCAT
	Reverse	ACCGACTTGGTCAGCGAAG
m-Col1a1	Forward	TAGGCCATTGTGTATGCAGC
	Reverse	ACATGTTCAGCTTTGTGGACC
m-Igf1	Forward	TCAACAAGCCCACAGGCTATG
	Reverse	GCTCCGGAAGCAACACTCAT
h-IRS2	Forward	CAGTGCTGAGCGTCTTCTTTT
	Reverse	ACCTACGCCAGCATTGACTT
h-IL1b	Forward	TGAGCACCTTCTTTCCCTTCA
	Reverse	GTAGTGGTGGTCGGAGATTCG
h-IL6	Forward	CCTGACCCAACCACAAATGC
	Reverse	CCTTAAAGCTGCGCAGAATGA
h-TNFA	Forward	TCGAACCCCGAGTGACAAG
	Reverse	TTGGCCAGGAGGGCATT
h-CCL2	Forward	GAAGAATCACCAGCAGCAAGTG
	Reverse	CAGATCTCCTTGGCCACAATG
h-CCL5	Forward	CGTGCCCACATCAAGGAGTA
	Reverse	CGGTTCTTTCGGGTGACAAA
h-GAPDH	Forward	GAAGGTGAAGGTCGGAGT
	Reverse	CATGGGTGGAATCATATTGGAA