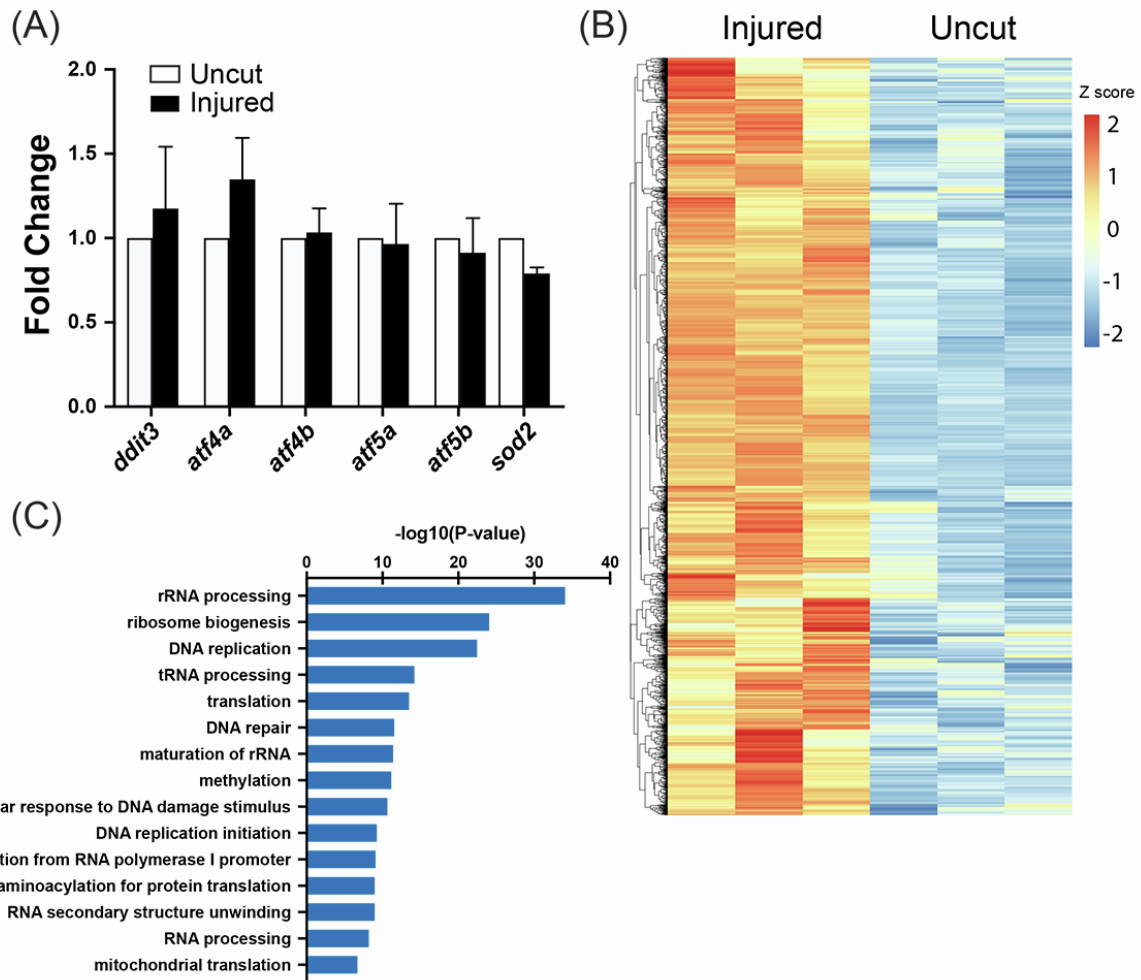


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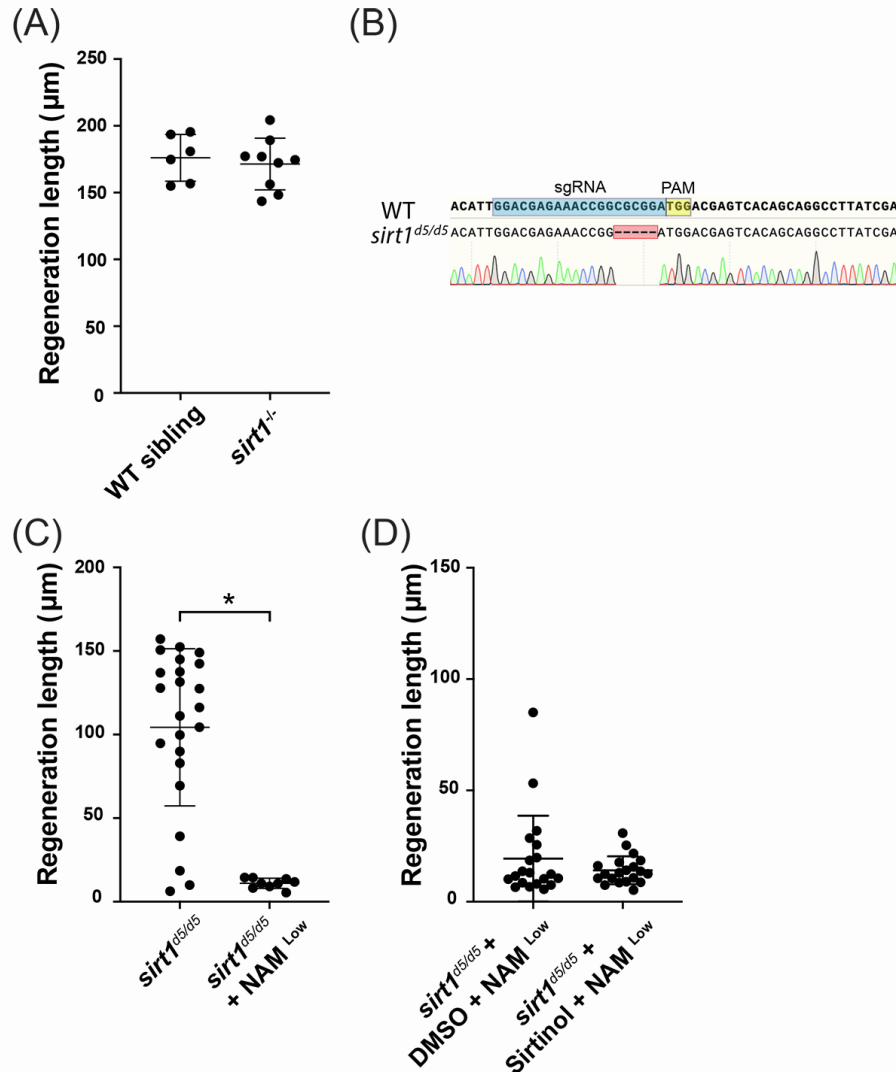
Supplemental information

**Sirt1 promotes tissue regeneration in zebrafish
through regulating the mitochondrial
unfolded protein response**

Yi-Fan Lin, Jessica Sam, and Todd Evans



Supplemental Figure 1. Transcriptional analysis of larval fin tissue during regeneration, Related to Figure 2. (A) qRT-PCR assays show that none of the known UPR^{mt} transcriptional regulators, nor *sod2* is up-regulated in regenerating larval fin tissue. (B) A heatmap shows significantly up-regulated genes (2908 genes) in regenerating larval fin tissue. (C) Top enriched processes identified from the up-regulated genes in regenerating larval fin tissue.



Supplemental Figure 2. An independent *sirt1* mutant allele also shows a consistent fin regeneration defect, Related to Figure 3. (A) The *sirt1* mutant larval fish exhibit normal regeneration growth compared to wild type siblings. (B) An independent second *sirt1* mutant allele (indicated here as d5) carries a 5 bp deletion in exon1 of the *sirt1* gene. The official name for this mutant allele is *sirt1*^{wcm19/wcm19}. (C) The *sirt1*^{d5/d5} mutant larval fish exhibit a strong regeneration defect when incubated in the low concentration of NAM. (D) The *sirt1*^{d5/d5} mutant larval fish exhibit a similar regeneration defect when treated with Sirtinol and NAM compared to NAM alone.

Table 1. List of Primers, Related to STAR Methods

cloning	
<i>hspd1</i> promoter	f: TGTTTGTGGATCCTGGACCC r: TCAGAGAGTGAGAGAGAGAG
<i>ornithine decarboxylase</i> d2 fragment	f: AGCCATGGCTTCCCGCCGGAG r: CTACACATTGATCCTAGCAGA
<i>hspd1 in situ</i>	f: TGTTGAGGACCAGAGTGCTG r: TGAGACAGATGAGGCCTGTG
<i>lonp1 in situ</i>	f: CCCATGTGCTCAGGTAAT r: GATCATCAGAGCCGGGATTA
qRT-PCR	
<i>hspd1</i>	f: ACTCCAGAGGAAATCGCTCA r: CATGCCCTCAATGATCTCAA
<i>hspa9</i>	f: GTGGCTGTGATGGATGGAAA r: ATCCTACAAGCCGCTCTCC
<i>clpp</i>	f: CAGAGAGCAACAATAAGCCGA r: CAGGTGGAGATGGGATTTAGGA
<i>lonp1</i>	f: GCTGTTGAGGAGGAAAGTGC r: CGCAGTTTATCTCCGAGGTC
<i>ddit3</i>	f: TGGGACAAAATATCGCCAAC r: AATACGACACGCTCCCCTC
<i>atf4a</i>	f: GACAGAGCAGAGCACAGCAG r: TCACACGACCCAATCAGAGA
<i>atf4b</i>	f: CGCAAACAACCTCAGTGCATC r: TAATTTTCGTGCTGTCGGGTA
<i>atf5a</i>	f: TGGTGAACGCAAACAGAAGA r: GCTGTTCCCTCCAATGAGTCC
<i>atf5b</i>	f: TCAGCACGATGGAGTTTCAG r: ATCTATGCCGCCACAATTC
<i>sod2</i>	f: TTCATCACAGCAAGCACCAT r: ATTTCAATGCAGGCTGAAGG

<i>polrmt</i>	f: TGTTGTCGCATGTGTCTCAG
	r: GCCTTCGGTAAATCGTAGCC
<i>tf2bm</i>	f: GTGTTGACTCGTGCTTTGCT
	r: CCCTCCAGTCTGCTCTCCA
<i>sirt1</i>	
mutagenesis	
<i>gRNA to target</i>	
<i>sirt1</i> (PAM site underlined)	GGACGAGAAACCGGC <u>GCGGATGG</u>
<i>sirt1 genotyping</i>	f: TGACGTCACGCCACGGCATTACCG
	r: CGGAGATCTCGGGCTCCGGGTCGGCTG