

Table S5: Overlaps between genes induced by tBOOH, starvation, or in *glp-1(bn18)* mutants.

Overlaps between gene sets	250 tBOOH-induced genes (RNA-seq, this study.)
329 fasting-induced genes (Uno et al. 2013)	Overlap: 44 genes p-value < 2.2e-16 Odds ratio: 13.57
1306 genes induced in <i>glp-1(bn18)</i> mutants (Steinbaugh et al. 2015)	Overlap: 61 genes p-value < 2.2e-16 Odds ratio: 4.73
221 genes induced by tBOOH, SKN-1 independent (Oliveira et al. 2009)	Overlap: 53 genes p-value < 2.2e-16 Odds ratio: 29.14

Table S6: Oxidative stress (6 mM tBOOH) response of *C. elegans nhr-49* and *fmo-2* null mutants.

Strain	Experiment no.	Median survival (hours)	Number of subjects (Dx/Nx (Cx))	p value vs. wild type
N2	1	26	67/112 (45)	NA
	2	41	41/118 (77)	NA
	3	52	48/107 (59)	NA
	4	30	76/129 (53)	NA
<i>nhr-49(nr2041)</i>	1	20	79/92 (13)	0.0016**
	2	27.5	34/44 (10)	<0.001***
	3	Not determined		
	4	Not determined		
<i>fmo-2(ok2147)</i>	1	43	76/110 (34)	0.0015**
	2	51.5	32/66 (34)	0.0849
	3	69	36/113 (77)	0.0027**
	4	30	61/129 (68)	0.0363*

Number of subjects is denoted as follows: Dx=Number of deaths that occurred during the assay; Nx=Total number of animals used in assay; Cx=Number of censored events (i.e. worms that ruptured at the vulva, underwent internal hatching of the progeny, or crawled off the plate). All p-values are derived using the log-rank (Mantel-Cox) test. *p<0.05, **p<0.01, and ***p<0.001. NA=not applicable.

Table S7: Oxidative stress (6 mM tBOOH) response of *C. elegans Pnhr-49::nhr-49::gfp* worms and non-transgenic siblings.

Strain	Experiment no.	Median survival (hours)	Number of subjects (Dx/Nx (Cx))	p value vs. non-transgenic siblings
Non-transgenic siblings	1	45	87/106(19)	NA
	2	26	101/115(4)	NA
	3	46	92/107(15)	NA
	4	45	91/102(10)	NA
<i>nhr-49p::nhr-49::gfp</i>	1	52	66/104(32)	0.0015**
	2	44	89/116(3)	0.0015**
	3	28	90/110(18)	0.0653
	4	45	95/105(9)	0.0567

Number of subjects is denoted as follows: Dx=Number of deaths that occurred during the assay; Nx=Total number of animals used in assay; Cx=Number of censored events (i.e. worms that ruptured at the vulva, underwent internal hatching of the progeny, or crawled off the plate). All p-values are derived using the log-rank (Mantel-Cox) test. **p<0.01. NA=not applicable. Cox's regression analysis of all four datasets indicated that there was a statistically significant difference between *nhr-49p::nhr-49::gfp* and non-transgenic siblings: p=0.036.

Table S8: Oxidative stress (6 mM tBOOH) response of *C. elegans nhr-49(et13)* *gof* mutants.

Strain	Experiment no.	Median survival (hours)	Number of subjects (Dx/Nx (Cx))	p value vs. wild type
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N2	1	45	81/118(37)	NA
	2	45	66/120(54)	NA
	3	47	66/116(50)	NA
	4	30	76/129 (53)	NA
<i>nhr-49(et13)</i>	1	45	54/119(65)	0.0492*
	2	52	39/136(97)	0.1352
	3	67	29/121(92)	0.0002***
	4	30	69/125 (56)	0.0007***

Number of subjects is denoted as follows: Dx=Number of deaths that occurred during the assay; Nx=Total number of animals used in assay; Cx=Number of censored events (i.e. worms that ruptured at the vulva, underwent internal hatching of the progeny, or crawled off the plate). All *p*-values are derived using the log-rank (Mantel-Cox) test. **p*<0.05, and ****p*<0.001. NA=not applicable. Cox's regression analysis of all four datasets indicated that there was a statistically significant difference between N2 and *nhr-49(et13)*: *p*<0.001.

Table S9: Oxidative stress (6 mM tBOOH) response of *C. elegans nhr-49, glp-1, and glp-1; nhr-49* mutants.

Strain	Experiment no.	Median survival (hours)	Number of subjects (Dx/Nx (Cx))	<i>p</i> value vs. wild type	<i>p</i> value vs. <i>glp-1(e2141)</i>
N2	1	23	57/116 (59)	NA	NA
	2	21	107/131 (24)	NA	NA
	3	30	65/90 (25)	NA	NA
	4	33	63/76 (13)	NA	NA
<i>glp-1(e2141)</i>	1	74	87/115 (28)	<0.0001****	NA
	2	69	84/104 (20)	<0.0001****	NA
	3	46	72/98 (26)	<0.0001****	NA
	4	46	85/113 (28)	<0.0001****	NA
<i>nhr-49(nr2041)</i>	1	Not determined			
	2	Not determined			
	3	23	60/76 (16)	<0.0001****	NA
	4	23	95/109 (14)	<0.0001****	NA
<i>glp-1(e2141); nhr-49(nr2041)</i>	1	30	62/63 (1)	0.0123*	<0.0001****
	2	27	64/66 (2)	<0.0001****	<0.0001****
	3	23	36/50 (14)	0.0101*	<0.0001****
	4	23	67/80 (13)	<0.0001****	<0.0001****

Number of subjects is denoted as follows: Dx=Number of deaths that occurred during the assay; Nx=Total number of animals used in assay; Cx=Number of censored events (i.e. worms that ruptured at the vulva, underwent internal hatching of the progeny, or crawled off the plate). All *p*-values are derived using the log-rank (Mantel-Cox) test. **p*<0.05, and ****p*<0.0001. NA=not applicable.

Table S10: Oxidative stress (6 mM tBOOH) response of *C. elegans* wild-type worms grown on RNAi clones targeting NHR-49 regulated genes.

RNAi clone	Experiment no.	Median survival (hours)	Number of subjects (Dx/Nx (Cx))	<i>p</i> value vs. control
control (L4440)	1	20	57/83(25)	NA
	2	24	54/81(32)	NA
	3	19	66/90(24)	NA
K05B2.4	1	20	67/74(7)	< 0.0001****
	2	20	70/84(7)	< 0.0001****
	3	19	62/81(19)	< 0.0001****
ZK550.6	1	28	49/71(21)	0.0036*
	2	20	60/81(30)	0.1735
	3	Undefined	17/74(55)	< 0.0001****
<i>nlp-25</i>	1	28	48/97(47)	< 0.0001****
	2	20	51/70(19)	0.049*
	3	19	67/83(13)	0.1737

<i>dhs-18</i>	1	28	44/100(55)	< 0.0001****
	2	20	54/93(35)	0.7438
	3	19	60/80(22)	0.7322
<i>icl-1</i>	1	28	37/102(63)	< 0.0001****
	2	20	60/88(24)	0.2671
	3	42	33/73(45)	< 0.0001****
<i>sodh-1</i>	1	20	59/81(19)	< 0.0001****
	2	20	61/70(8)	< 0.0001****
	3	19	57/72(15)	< 0.0001****

Number of subjects is denoted as follows: Dx=Number of deaths that occurred during the assay; Nx=Total number of animals used in assay; Cx=Number of censored events (i.e. worms that ruptured at the vulva, underwent internal hatching of the progeny, or crawled off the plate). All *p*-values are derived using the log-rank (Mantel-Cox) test. **p*<0.05, and *****p*<0.0001. NA=not applicable.

Table S11: Oxidative stress (6 mM tBOOH) response of *C. elegans glp-1* mutants grown on RNAi clones targeting NHR-49 regulated genes.

RNAi clone	Experiment no.	Median survival (hours)	Number of subjects (Dx/Nx (Cx))	<i>p</i> value vs. control
control (L4440)	1	74	61/97(34)	N/A
	2	55	39/70(29)	N/A
	3	42	94/111(22)	N/A
K05B2.4	1	28	81/104(13)	< 0.0001****
	2	47	78/102(26)	0.0125*
	3	19	75/103(27)	< 0.0001****
ZK550.6	1	47	58/74(18)	0.2017
	2	47	56/87(32)	0.9293
	3	42	50/63(13)	0.4061
<i>nlp-25</i>	1	74	66/99(30)	0.2711
	2	55	67/97(35)	0.1879
	3	42	83/95(16)	0.0669
<i>dhs-18</i>	1	47	84/103(21)	< 0.0001****
	2	74	73/88(22)	0.6203
	3	49	76/106(32)	0.0819
<i>icl-1</i>	1	74	52/99(50)	0.3797
	2	55	56/93(31)	0.9686
	3	27	68/103(34)	0.0052**
<i>sodh-1</i>	1	28	89/97(9)	< 0.0001****
	2	47	67/91(28)	< 0.0001****
	3	27	102/107(6)	< 0.0001****

Number of subjects is denoted as follows: Dx=Number of deaths that occurred during the assay; Nx=Total number of animals used in assay; Cx=Number of censored events (i.e. worms that ruptured at the vulva, underwent internal hatching of the progeny, or crawled off the plate). All *p*-values are derived using the log-rank (Mantel-Cox) test. **p*<0.05, ***p*<0.01, and *****p*<0.0001. NA=not applicable.

Table S12: Oxidative stress (6 mM tBOOH) response of *C. elegans hlh-30* mutants.

Strain	Experiment no.	Median survival (hours)	Number of subjects (Dx/Nx (Cx))	<i>p</i> value vs. wild type
N2	1	45	84/112(28)	NA
	2	46	83/94(2)	NA
	3	44	111/120(9)	NA
<i>hlh-30(tm1978)</i>	1	45	92/117(20)	0.8488
	2	46	88/94(2)	0.3601
	3	44	112/120(8)	0.8238

Number of subjects is denoted as follows: Dx=Number of deaths that occurred during the assay; Nx=Total number of animals used in assay; Cx=Number of censored events (i.e. worms that ruptured at the vulva, underwent internal hatching of the progeny, or crawled off the plate). All *p*-values are derived using the log-rank (Mantel-Cox) test. NA=not applicable.

Table S13. Worm strains used in this study.

Strain	Genotype	Reference
N2	Wild type	(Brenner 1974)
STE68	<i>nhr-49(nr2041) I</i>	(Van Gilst et al. 2005)
VE40	<i>N2 eavEx20[fmo-2p::gfp + rol-6(su1006)]</i>	This study
STE108	<i>nhr-49(et7) I</i>	(Lee et al. 2016)
STE109	<i>nhr-49(et8) I</i>	(Lee et al. 2016)
STE110	<i>nhr-49(et13) I</i>	(Lee et al. 2016)
STE117	<i>nhr-49(et13) I; eavEx20[fmo-2p::gfp + rol-6(su1006)]</i>	This study
STE71	<i>nhr-13(gk796) V</i>	(Pathare et al. 2012)
STE69	<i>nhr-66(ok940) IV</i>	(Pathare et al. 2012)
STE70	<i>nhr-80(tm1011) III</i>	(Goudeau et al. 2011)
VC1668	<i>fmo-2(ok2147) IV</i>	(Leiser et al. 2015)
JIN1375	<i>hlh-30(tm1978) IV</i>	(Lapierre et al. 2013)
VE41	<i>hlh-30(tm1978) IV; eavEx20[fmo-2p::gfp + rol-6(su1006)]</i>	This study
MAH235	<i>sqIs19[hlh-30p::hlh-30::gfp, rol-6(su1006)]</i>	(Lapierre et al. 2013)
STE119	<i>sqIs19[hlh-30p::hlh-30::gfp, rol-6(su1006)]; nhr-49(nr2041) I</i>	This study
STE118	<i>nhr-49(et13) I; hlh-30(tm1978) IV</i>	This study
KAE9	<i>seaSi39 [(pCFJ448) (eft-3p::fmo-2 + H2B::GFP) + Cbr-unc-119(+)] I; unc-119(ed3) III</i>	(Leiser et al. 2015)
CF1903	<i>glp-1(e2141ts) III</i>	(Arantes-Oliveira 2002)
AGP22	<i>nhr-49(nr2041) I; glp-1(e2141ts) III</i>	(Ratnappan et al. 2014)
VE42	<i>sek-1(km4); eavEx20[(fmo-2p::gfp + rol-6(su1006)]</i>	This study
AGP24f	<i>glmEx5 (Pnhr-49::nhr-49::GFP + Pmyo-2::mCherry)</i>	(Ratnappan et al. 2014)
KU4	<i>sek-1(km4) X</i>	(Tanaka-Hino et al. 2002)
VE43	<i>sek-1(km4) X; glmEx5[Pnhr-49::nhr-49::GFP + Pmyo-2::mCherry]</i>	This study
KU25	<i>pmk-1(km25) IV</i>	(Mizuno et al. 2004)

Table S14. List of non-Ahringer library RNAi clones used in this study.

Clone	Source
<i>nhr-28</i>	Custom made Forward primer: GCTAGCgcggtgacaatatattccatgtt Reverse primer: CCCGGGcCTCACTGCTTCGTTCCGGAC
<i>nhr-12</i>	Vidal library, Plate 70 Well C03
<i>nhr-86</i>	(Arda et al. 2010)
<i>nhr-112</i>	Custom made Forward primer: GCTAGCGAGGACAGGGATAGAgtagg Reverse primer: AAGCTTGATATGTTTGAGTAGCctgg
<i>nhr-114</i>	Custom made Forward primer: GCTAGCgttcagCTATCCGCTGTGCCTGC Reverse primer: AAGCTTGAGTCCGGCTGGCctataacg
<i>nhr-138</i>	Custom made Forward primer: GCTAGCGGCAACGAGGACAAAGgtagg

Clone	Source
	Reverse primer: CCCGGGgttacCTGTACAGAAGTAAGC
<i>nhr-273</i>	Custom made Forward primer: GCTAGCGCCGATTGATCAGAAAgtagg Reverse primer: AAGCTTCGGACGGCGCATACAAGTTG

Table S15. List of qPCR primer sequences used in this study.

Gene	Forward primer (5'-3')	Reverse primer (5'-3')
<i>fmo-2</i>	GGAACAAGCGTGTGCTGT	GCCATAGAGAAGACCATGTCCG
<i>dhs-18</i>	CATCCAAAACCTACCGGGAAC	TTTACTGCTGCCTCATCACG
<i>K05B2.4</i>	CCCTATACGAATGACAGGATTG	TGTTTGAACCTTGTGGTGAG
<i>icl-1</i>	ATTGCTTCGAGTTGATGAAGG	GATCCAAGCTGATCTTCGTAGTG
<i>ZK550.6</i>	TTCCGGAGCCAATAGAACTG	CGGGTCGAGACCATATCTTG
<i>sodh-1</i>	ATTGGTTGGAGGACACGAAG	GCTCGTGGCCTTTCTTACAG
<i>nlp-25</i>	ATCACTAATTGCGCTTCTCC	TCCTCCACCTCTGCCATAAC
<i>nhr-49</i>	TCCGAGTTCATTCTCGACG	GGATGAATTGCCAATGGAGC
<i>hlh-30</i>	CTCATCGGCCGGCGCTCATC	AGAACCGGATGCGTGGTGGG
<i>mdt-15</i>	GGAAATCCGTACAATCAGCAG	CAAGAACTGATGAAGGACCG
<i>act-1</i>	GCTGGACGTGATCTTACTGATTAC C	GTAGCAGAGCTTCTCCTTGATGTC
<i>tba-1</i>	GTACACTCCACTGATCTCTGCTGA CAAG	CTCTGTACAAGAGGCAAACAGCCAT G
<i>ubc-2</i>	AGGGAGGTGTCTTCTTCCTCAC	CGGATTTGGATCACAGAGCAGC

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