

Genotype	week 1	N1	week 2	N2	week 4	N4	week 5	N5
<i>dFoxo</i> ²¹ x <i>dFoxo</i> ²⁵ (1)	0.35	132					0.75	132
<i>dFoxo</i> ²¹ (1) x <i>yw</i>	0.3	15					0.92	15
<i>S6K</i> ¹⁻¹ / <i>S6K</i> ^{P1713} (2)	0.37	246					0.23	246
<i>S6K</i> ^{P1713} (2) x <i>yw</i>	0.17	233					0.68	233
<i>GMH5</i> (3) x <i>yw</i>	0.23	100	0.36	100	0.68	100	0.76	100
<i>UAS-dTOR</i> (4) x <i>GMH5</i>	0.69	26	0.72	34	0.94	39	0.99	33
<i>UAS-dTSC1&2</i> (5) x <i>GMH5</i>	0.25	100	0.34	86	0.42	41	0.45	33
<i>UAS-dnS6K</i> (6) x <i>GMH5</i>	0.25	130	0.33	167	0.4	147	0.57	107
<i>UAS-dnS6K</i> (6) x <i>yw</i>	0.21	159	0.38	134	0.45	127	0.69	103
<i>UAS-dnS6K</i> (6) x <i>Isp2-Gal4</i> (7)	0.4	129	0.37	106	0.46	47	0.58	62
<i>UAS-dnS6K</i> (6) x <i>elav-Gal4</i> (14)	0.56	100	0.27	100	0.5	100	0.6	100
<i>UAS-dnS6K</i> (6) x <i>dilp2-Gal4</i> (12)	0.54	100	0.33	100	0.23	100	0.33	100
<i>UAS-dnTOR</i> (15) x <i>Isp2-Gal4</i> (7)	0.32	100	0.44	100	0.53	100	0.68	100
<i>UAS-d4eBP</i> (8) x <i>GMH5</i> (3)	0.37	94	0.36	85	0.4	153	0.43	100
<i>UAS-dEif4e</i> (9) x <i>GMH5</i> (3)	0.7	201	0.75	112	0.77	93	0.8	64
<i>UAS-dMyc</i> (10) x <i>GMH5</i> (3)	0.22	168	0.32	160	0.64	181	0.75	100
<i>UAS-dEif4a</i> (11) x <i>GMH5</i> (3)	0.43	42	0.39	48	0.52	118	0.7	103
<i>UAS-dFoxo-RNAi</i> (VDRC) x <i>GMH5</i> (3)	0.53	100	0.58	100	0.59	100	0.56	100
<i>UAS-dFoxo-RNAi</i> (VDRC) x <i>yw</i>	0.36	100	0.48	100	0.45	100	0.63	100
<i>UAS-d4eBP-RNAi</i> (VDRC) x <i>GMH5</i> (3)	0.63	100	0.69	100	0.46	100	0.54	100
<i>UAS-d4eBP-RNAi</i> (VDRC) x <i>yw</i>	0.36	100	0.48	100	0.51	100	0.58	100
<i>UAS-d4eBP</i> (8); <i>UAS-dTOR</i> (4) x <i>GMH5</i>	0.41	100	0.38	100	0.17	100	0.29	100
<i>UAS-d4eBP</i> (8); <i>UAS-dTOR</i> (4) x <i>yw</i>	0.34	100	0.37	100	0.54	100	0.72	100
<i>UAS-dEif4e</i> (9); <i>UAS-dTSC1&2</i> (5) x <i>GMH5</i>	0.58	100	0.59	100	0.61	100	0.64	100
<i>UAS-d4eBP</i> (8); <i>UAS-dFoxo</i> (3) x <i>GMH5</i>	0.24	100	0.3	100	0.38	100	0.36	100
<i>UAS-d4eBP</i> (8); <i>UAS-dFoxo</i> (3) x <i>yw</i>	0.12	100	0.33	100	0.54	100	0.59	100
<i>UAS-dEif4e</i> (9); <i>UAS-dFoxo</i> (3) x <i>GMH5</i>	0.64	100	0.6	100	0.73	100	0.69	100
<i>UAS-dEif4e</i> (9); <i>UAS-dFoxo</i> (3) x <i>yw</i>	0.17	100	0.33	100	0.35	100	0.53	100
<i>UAS-dTOR</i> (4) x <i>yw</i>	0.39	118					0.65	92
<i>UAS-dTSC1&2</i> (5) x <i>yw</i>	0.37	173					0.72	100
<i>dilp2-Gal4</i> (12) x <i>yw</i>	0.16	48	0.38	54	0.60	49	0.66	53
<i>UAS-dnTOR</i> (15) x <i>Canton S</i>	0.31	85	0.59	132	0.71	41	0.45	38
<i>UAS-d4eBP</i> (8) x <i>yw</i>	0.26	172	0.59	132	0.71	41	0.78	14
<i>UAS-dEif4e</i> (9) x <i>yw</i>	0.42	45	0.24	160	0.60	111	0.65	100
<i>UAS-dMyc</i> (10) x <i>yw</i>	0.22	169	0.32	160	0.64	181	0.75	100
<i>GS-eif4e</i> ²³²⁶ (13) x <i>yw</i>	0.18	109					0.76	58
<i>GS-eif4e</i> ²⁷⁸³ (13) x <i>GMH5</i>	0.70	100					0.77	52
<i>GS-eif4e</i> ²²⁵¹ (13) x <i>GMH5</i>	0.63	85					0.80	46
<i>GS-eif4e</i> ²³²⁶ (13) x <i>GMH5</i>	0.75	16					0.80	31
<i>GS-eif4e</i> ²²⁵¹ (13) x <i>yw</i>	0.37	164					0.72	108
<i>UAS-dFoxo-RNAi</i> (VDRC) x <i>Isp2-Gal4</i> (7)	0.34	106	0.40	87	0.12	76	0.54	67
<i>UAS-dFoxo</i> (3x) (3) x <i>Isp2-Gal4</i> (7)	0.42	95	0.30	89	0.47	77	0.55	86

Source References: (1) Junger et al. 2003; (2) Montagne et al. 1999; (3) Wessells et al. 2004; (4) Hennig and Neufeld 2002; (5) Potter et al. 2001; (6) Barcelo and Stewart 2002; (7) Cherbas et al. 2003; (8) Mifon et al. 2001; (9) Arquier et al. 2005; (10) de la Cova and Johnston 2006; (11) Li et al. 2005; (12) Ikeya et al. 2002; (13) Krupp et al. 2003; (14) Brand and Perrimon 1993; (15) Hennig and Neufeld 2002. (VDRC) Vienna *Drosophila* Research Center.

Supplemental Table 1. Summary of failure rate vs. age of various genotypes. Genotype: flies tested are denoted by their genotype or by the cross used to generate them. week 1-5: fraction of flies tested that exhibit heart failure at that age. N1-5 number of flies tested at 1-5 weeks of age.

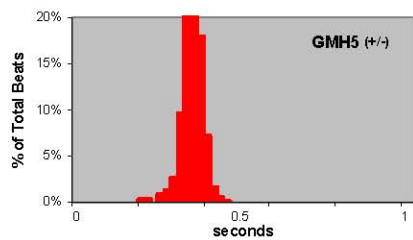
Supplemental Table 2: Cardiac Failure Rate Analysis

Genotype 1	Genotype 2	Genotype-by-Age p-value
<i>Foxo 21/25</i>	<i>Foxo21 x yw</i>	0.221
<i>S6K1-1/S6KP1713</i>	<i>S6KP1713 x yw</i>	<0.001
<i>UAS-dTOR x GMH5</i>	<i>UAS-dTOR x yw</i>	0.8476*
<i>GS-Eif4e²³²⁶ x GMH5</i>	<i>GS-Eif4e²³²⁶ x yw</i>	<0.01
<i>UAS-dnS6K x GMH5</i>	<i>UAS-dnS6K x yw</i>	0.084
<i>UAS-dnS6K x lsp2-gal4</i>	<i>UAS-dnS6K x yw</i>	<0.01
<i>UAS-dnS6K x elav-gal4</i>	<i>UAS-dnS6K x yw</i>	<0.001
<i>UAS-dnS6K x dilp2-gal4</i>	<i>UAS-dnS6K x yw</i>	<0.0001
<i>UAS-dMyc x GMH5</i>	<i>UAS-dMyc x yw</i>	0.986
<i>UAS-dTSC x GMH5</i>	<i>UAS-dTSC x yw</i>	0.2621*
<i>UAS-dnTOR x lsp2-gal4</i>	<i>UAS-dnTOR x Canton S</i>	0.214
<i>UAS-d4eBP;UASdFoxo x GMH5</i>	<i>UAS-d4eBP;UAS-dFoxo x yw</i>	<0.001
<i>UAS-Foxo(3x) x lsp2-gal4</i>	<i>UAS-Foxo-RNAi x lsp2-gal4</i>	0.387
<i>UAS-dEif4e x GMH5</i>	<i>UAS-dEif4e x yw</i>	<0.001
<i>UAS-dEif4e;UAS-dFoxo x GMH5</i>	<i>UAS-dEif4e;UAS-dFoxo x yw</i>	<0.05
<i>UAS-d4eBP x GMH5</i>	<i>UAS-d4eBP x yw</i>	<0.001
<i>UAS-d4eBPRNAi x GMH5</i>	<i>UAS-d4eBP-RNAi x yw</i>	<0.0001
<i>UAS-d4eBP;UAS-dTOR x GMH5</i>	<i>UAS-d4eBP;UAS-dTOR x yw</i>	<0.0001
<i>UAS-dFoxoRNAi x GMH5</i>	<i>UAS-dFoxoRNAi x yw</i>	<0.05

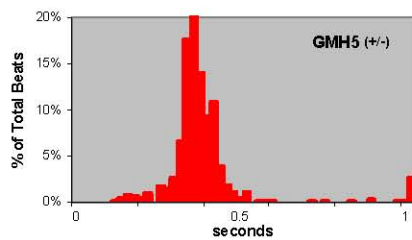
Supplemental Table 2: Statistical comparison of cardiac failure rate employing two-way ANOVA followed by a Bonferroni comparison to evaluate the effect of genotype on the effect of age on cardiac failure rate compared to out-crossed controls.

* *dTOR* and *dTSC* cardiac overexpression versus control do not show a significant difference in the slope of age-dependent changes (genotype x age). However, both *dTOR* and *dTSC* overexpression causes significantly higher failure at young ages (student's t-test at 1 week, *dTOR*: $p < 0.001$, *dTSC*: $p < 0.05$) and at older ages (student's t-test at 5 week, *dTOR*: $p < 0.0001$, *dTSC*: $p < 0.01$).

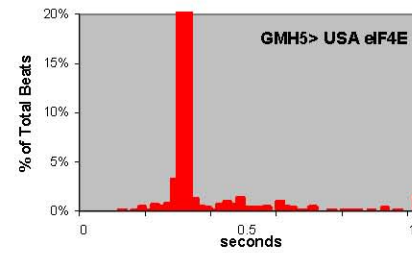
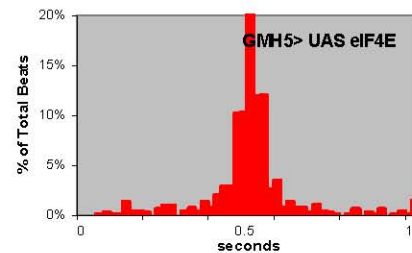
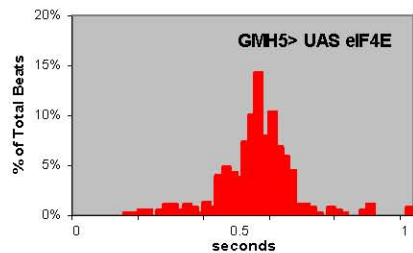
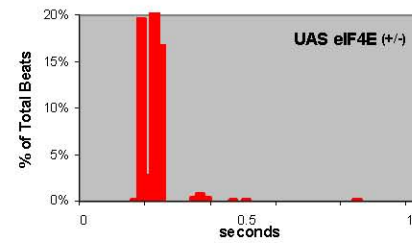
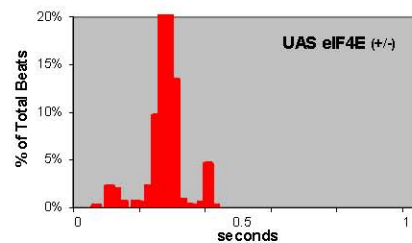
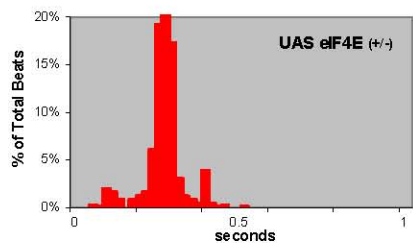
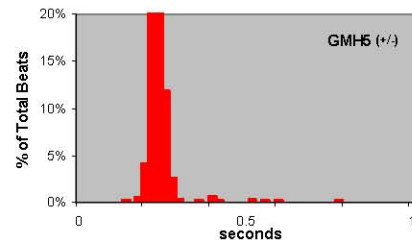
A Diastolic Intervals
(1-week old flies)



B Diastolic Intervals
(4-week old flies)



C Systolic Intervals
(4-week old flies)



Supplemental Figure 1 legend:

Distribution of diastolic and systolic Intervals in young and old flies. A) The distributions of diastolic intervals for all 1-week old flies (normalized to the median diastolic interval) are shown. The majority of diastoles from both heterozygous control groups show a relatively tight clustering between 0.3 and 0.4 seconds. The diastolic intervals for flies, where *dEif4e* has been over-expressed specifically in the heart, show a broader distribution compared to controls. B) The diastolic intervals for older (4-week old) flies where *dEif4e* has been over-expressed specifically in the heart also show a broader distribution compared to controls (especially due to 'outliers' at some distance from the main peak). C) The distribution of systolic intervals for older (4-week old) flies where *dEif4e* has been over-expressed specifically in the heart show a peak at a time point that is slightly longer than seen for controls ($p < 0.05$). Systolic intervals in *dEif4e* overexpression hearts also exhibit a broader distribution compared to controls with a significant number of very long systolic intervals corresponding to the episodes of fibrillation seen in the M-modes in panel (B), (compare GMH5 (+/-) and UAS eIF4E (+/-) with GMH5> *dEif4e*).