

## Supporting Information

### 1. Power calculations

For power calculations we used the `pwr.t2n.test` package in R. Cohen's effect size  $d$  was calculated as follows:

$$d = \frac{m_1 - m_2}{S_p}, \text{ where } S_p = \sqrt{\frac{(n_1 - 1)S_1^2 + (n_2 - 1)S_2^2}{n_1 + n_2 - 2}}; \mu_1 \text{ and } \mu_2 \text{ are the mean lifespans}$$

associated with each genotype;  $\sigma_p$  is the pooled standard deviation;  $S_1^2$  and  $S_2^2$  are variances associated with the mean lifespans within each genotype. These mean lifespans were taken from the most associated SNP ( $2R\_1632386$ ;  $p$ -value= $5.9 \times 10^{-08}$ ;  $\mu_1=56.57$  and  $\mu_2=45.97$ ;  $S_1^2=89.86$  and  $S_2^2=63.57$ ).

### 2. Broad sense heritability

To estimate broad sense heritability ( $H^2$ ) we partitioned the phenotypic variance between lines and the error variance for each line using ANOVA. Broad sense

heritability was estimated by  $H^2 = \frac{S_L^2}{S_L^2 + S_E^2}$ , where  $S_L^2$  is the among-line variance and

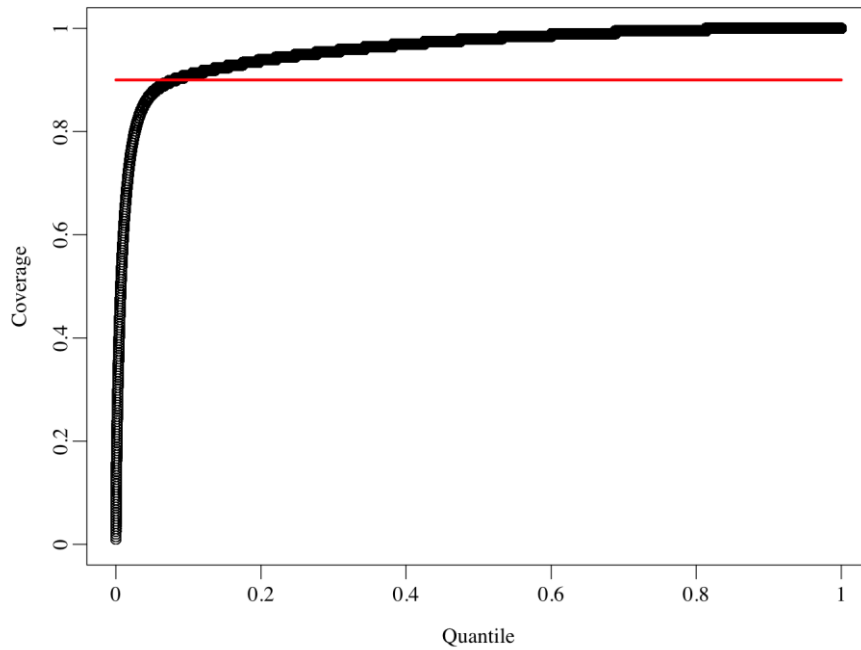
$S_E^2$  is the within-line variance.

### 3. *Drosophila melanogaster* lines

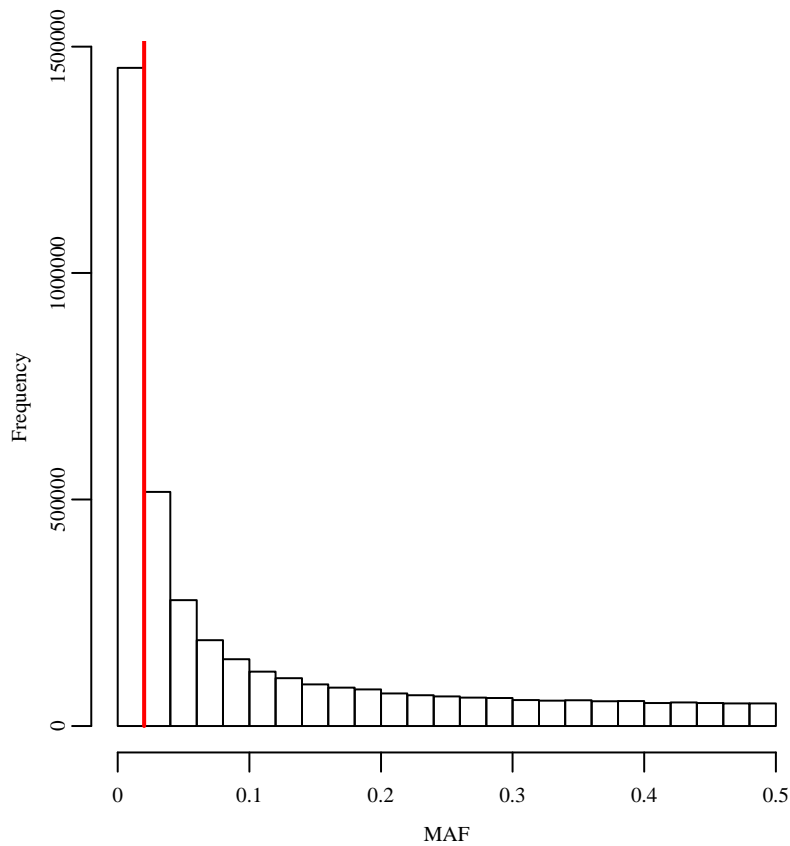
The *Drosophila* Genetic Reference Panel, Freeze 2.0 (1, 2), comprises 205 *D. melanogaster* lines derived by 20 generations of full-sib mating from wild-type caught females from Raleigh, North Carolina. Longevity was assayed as previously described (3, 4). All flies were reared from egg to adult on 10 ml standard cornmealagar-molasses medium at 25° in shell vials. The density of the stocks was controlled for three generations prior to the start of the longevity assays by restricting egg laying to 3 days and initiating the cultures with 10 pairs of flies. A total of 25 virgin males and females per line were collected in a 24-hr period and 2-day-old flies were housed in five replicate vials with five same-sex individuals per vial. Flies were transferred to fresh medium every 2 days and the number of live flies was recorded until all were dead. The assays were performed in three temporally overlapping blocks; no block effects were observed.

## Supplementary Graphs and Tables

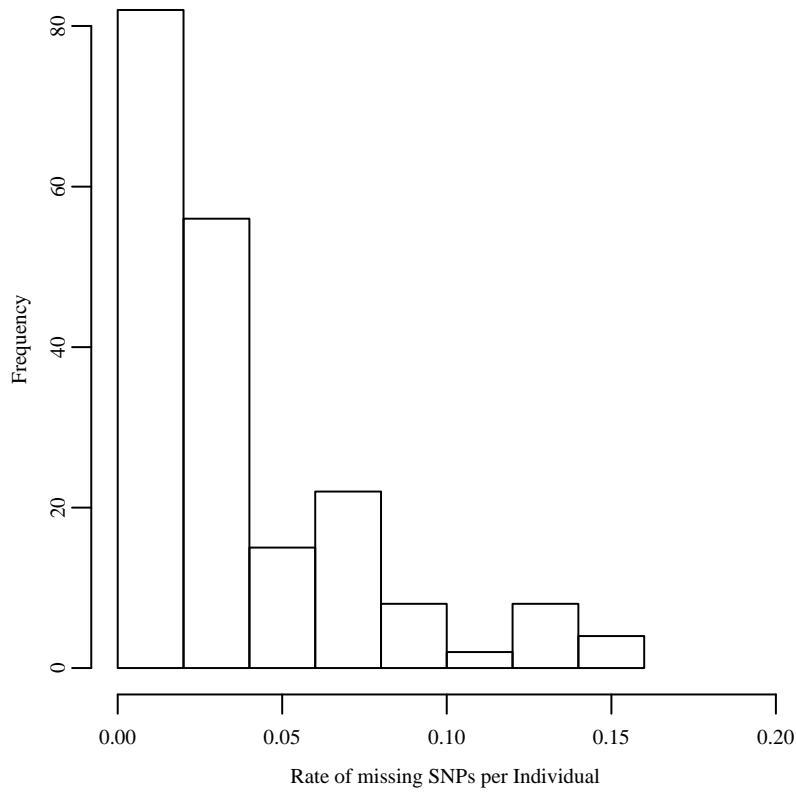
**Supplementary Figure 1. SNP call rate.** The red horizontal line represents the 0.9 SNP call rate. Coverage represents the proportion of genotypes present per SNP



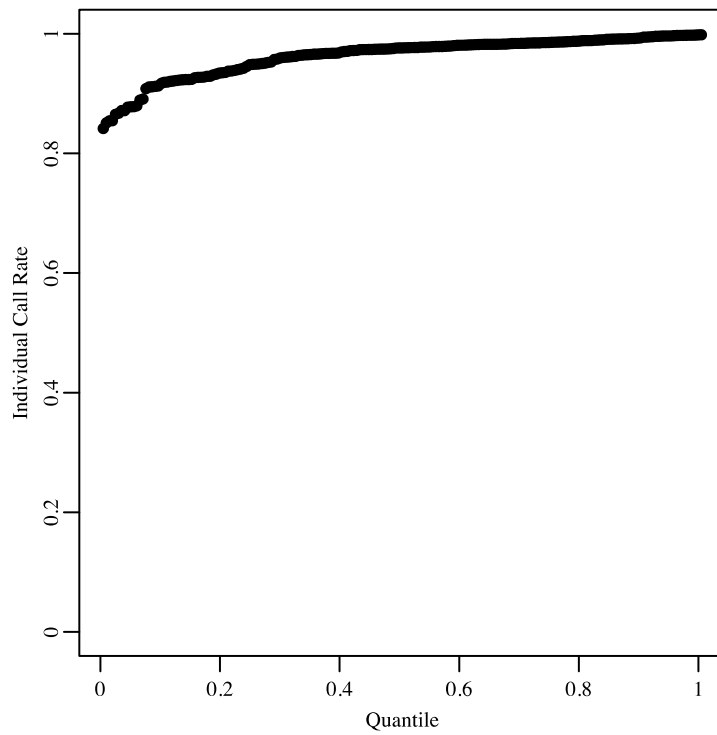
**Supplementary Figure 2. Minor allele frequency (MAF).** The red vertical line represents MAF=0.02



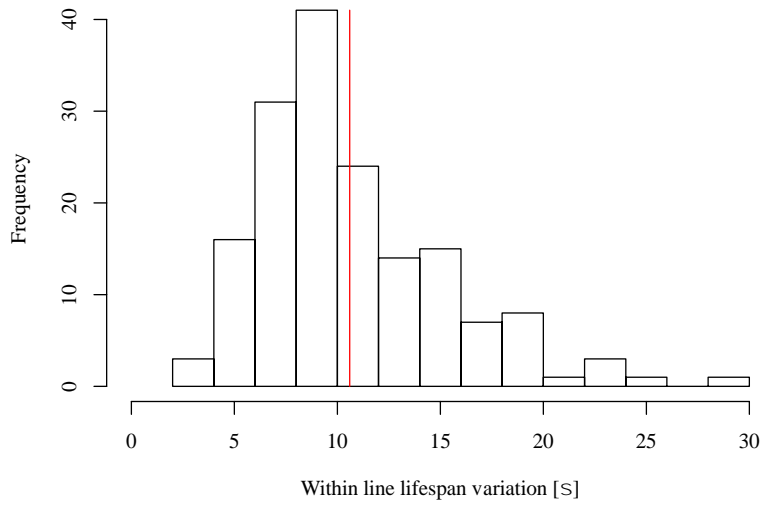
**Supplementary Figure 3. Histogram of individual call rate**



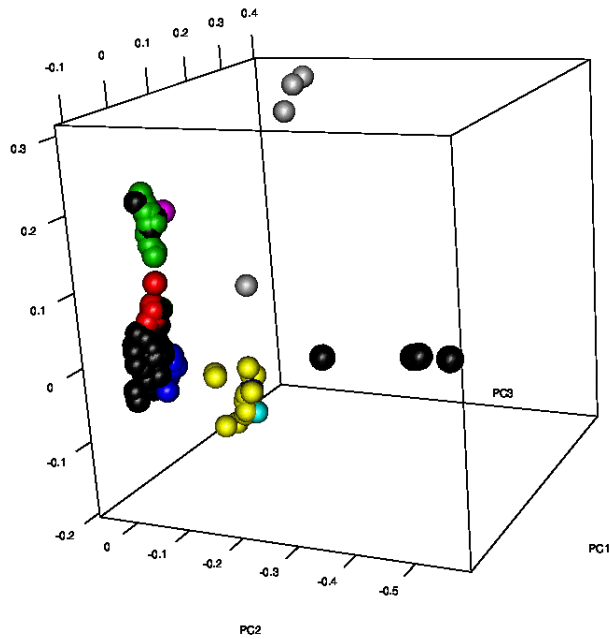
**Supplementary Figure 4. Individual call rate.** Individual call rate represents the proportion of genotypes present per fly



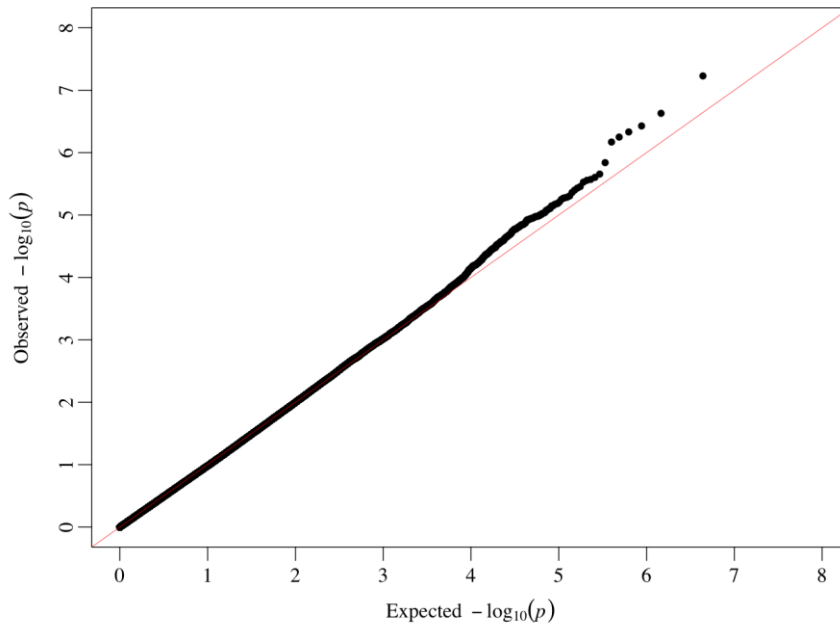
**Supplementary Figure 5. Lifespan variation within individual fly lines (165 lines).** The red vertical line represents the mean of the standard deviation between the fly lines (10.6)



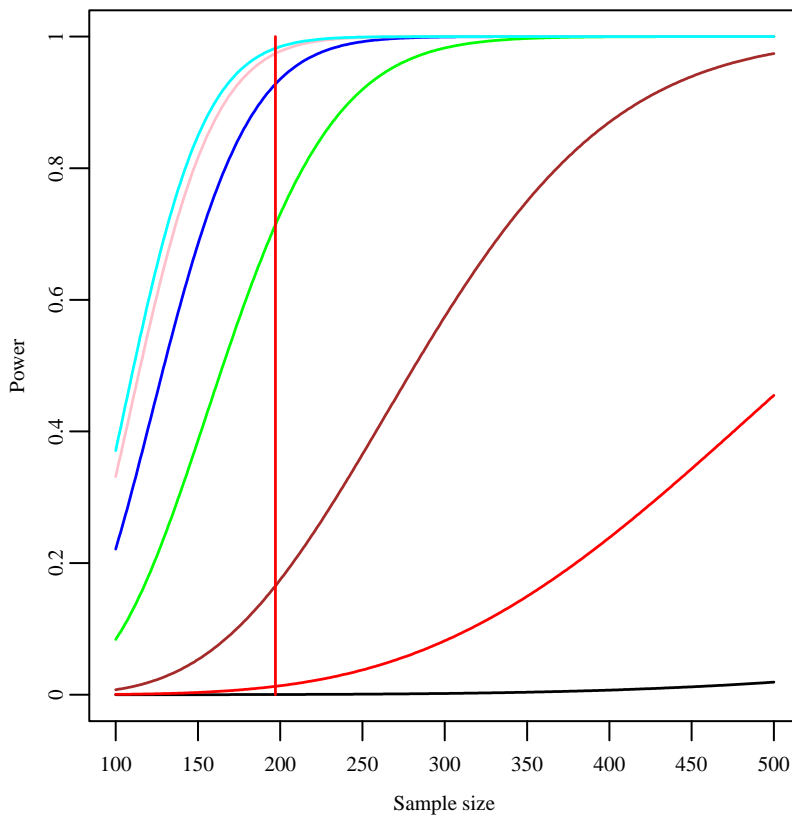
**Supplementary Figure 6. Principal component analysis (PCA), 197 lines.** PC1- principal component 1; PC2- principal component 2; PC3- principal component 3; Colours represent the inversion haplotype for *In(3R)Mo* and *In(2L)t*. 00- black, 01- red, 11- light blue, 12- pink, 22- grey, 10- blue, 20- yellow, 02- green



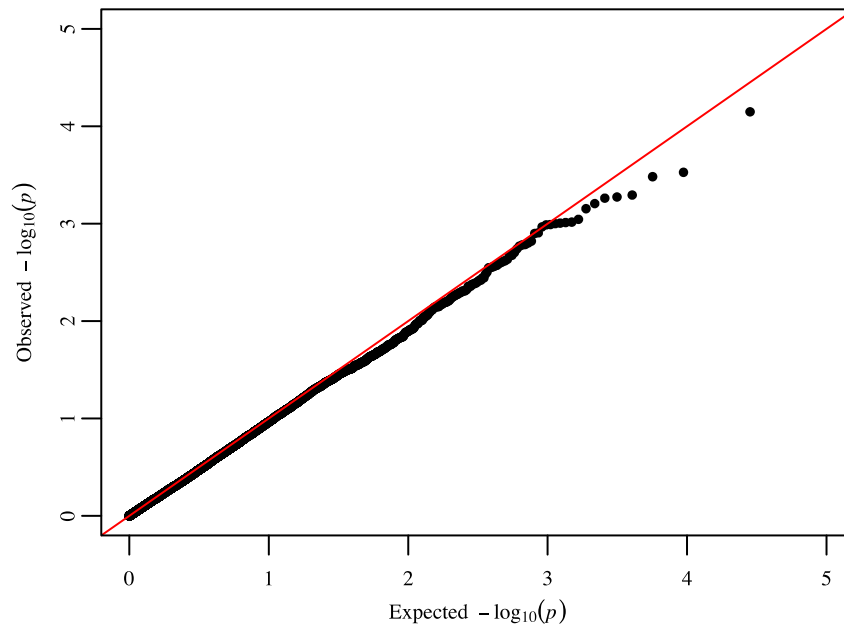
**Supplementary Figure 7. SNP-based QQ-plot**



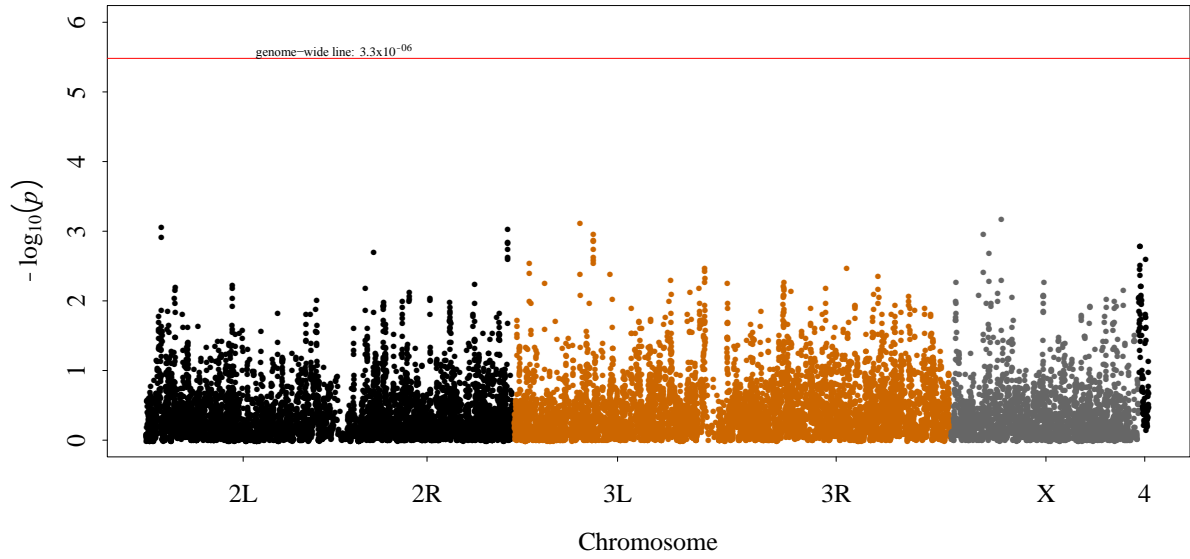
**Supplementary Figure 8. Power to detect single-SNP association.** Effect size = 10days (Cohen's  $d=1.15$ ), pooled standard deviation = 9.25,  $\mu_1=56.57$ ,  $\mu_2=45.97$ ; red line represent MAF 0.1, brown line 0.2, green line 0.3, blue line 0.4 and pink line 0.5. The red vertical line represents the DGRP sample size (197 lines)



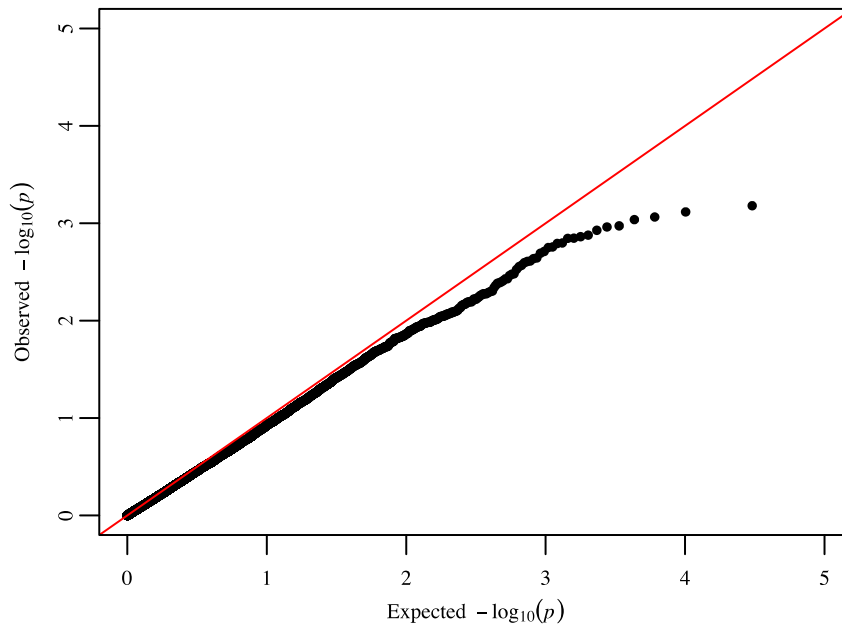
**Supplementary Figure 9. Gene-based QQ-plot**



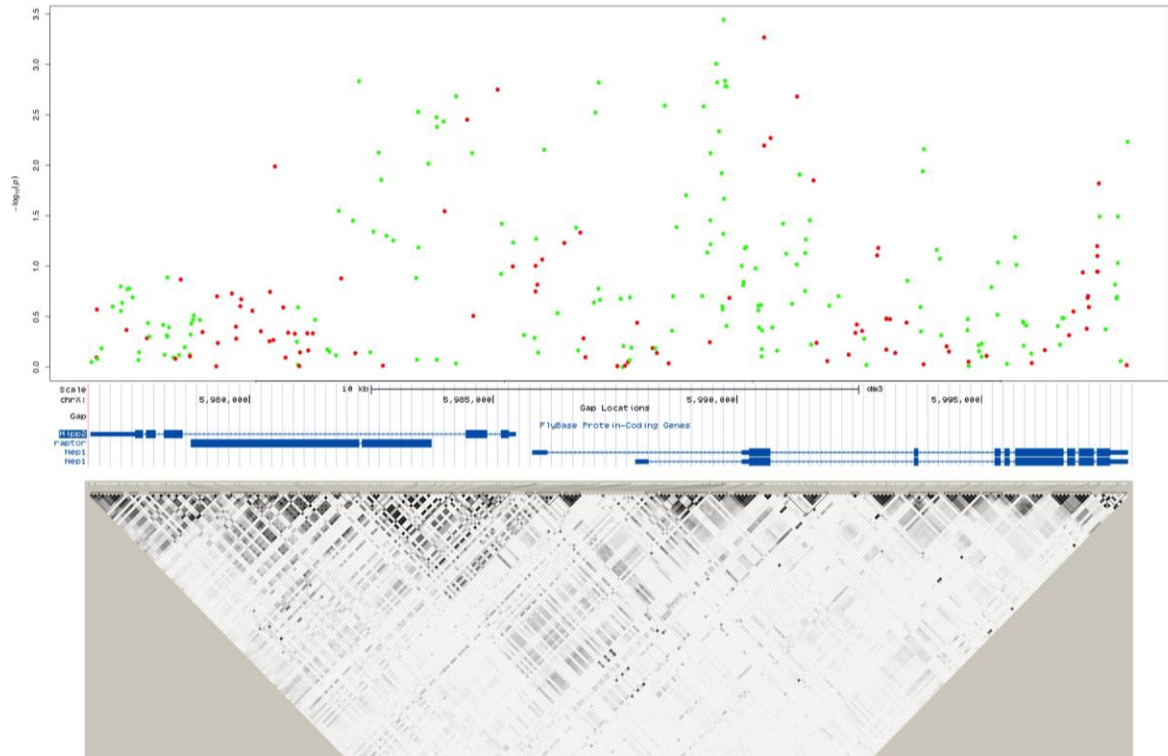
**Supplementary Figure 10. Manhattan plot for gene-based analysis (genes  $\pm 5\text{kb}$ ).** Each point represents a gene. The height of the points represents the strength of association with lifespan, expressed as  $-\log_{10}(p\text{-value})$ . The red horizontal line represents genome-wide Bonferroni significance threshold  $p=3.30 \times 10^{-6}$



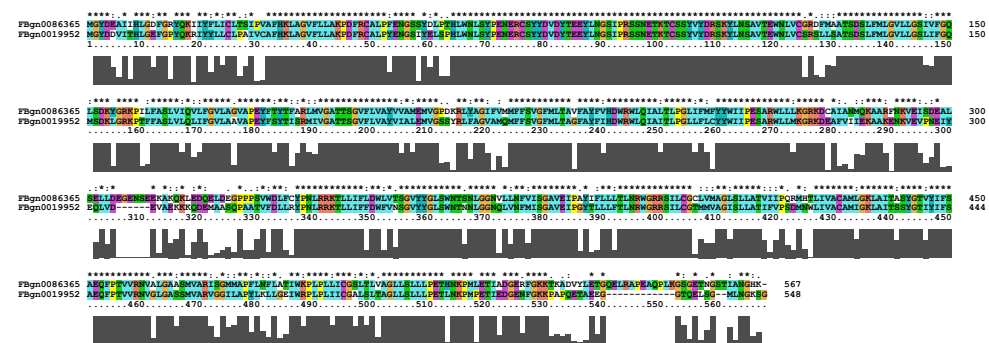
**Supplementary Figure 11. Gene-based QQ-plot (genes  $\pm 5\text{kb}$ )**



**Supplementary Figure 12. LD structure of *Mipp2* and *Nep1*.** Each point in the top half of the graph represents a SNP. The y-axis represents the strength of association expressed in  $-\log_{10}(p\text{-value})$ . Red dots represent SNPs with negative  $\beta$  coefficients and green dots SNPs with positive  $\beta$  coefficients. The middle panel shows the positions of *Mipp2* and *Nep1* in the *Drosophila* genome using the UCSC genome browser <http://genome-euro.ucsc.edu/cgi-bin/hgGateway>; The bottom panel depicts the LD structure within the two genes, expressed in terms of  $R^2$ . Black squares represent  $R^2=1$ . The LD structure was produced using Haploview (5)

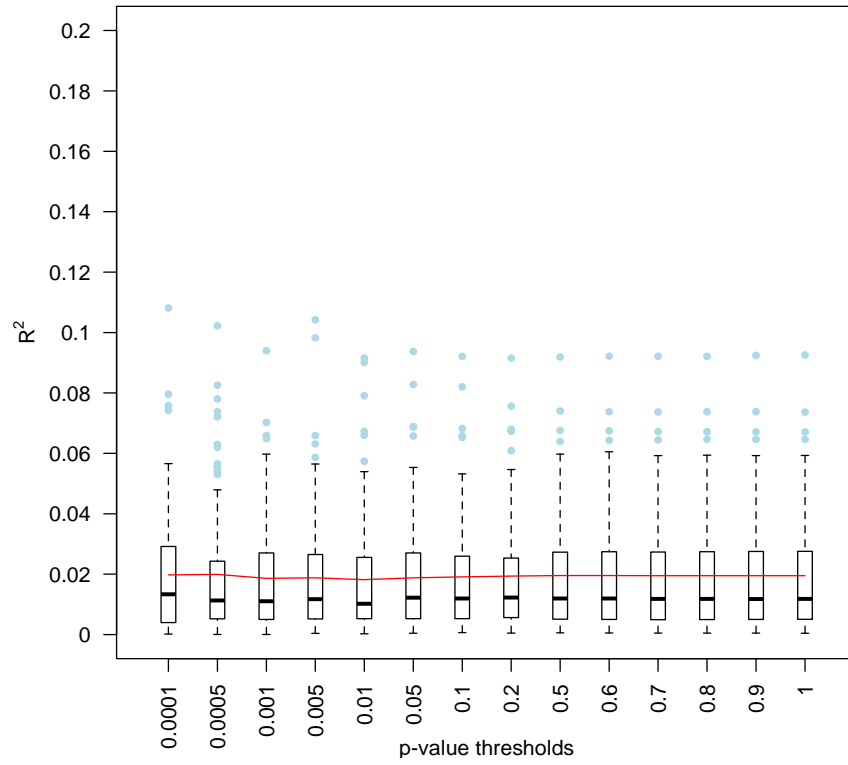


**Supplementary Figure 13. Protein alignment of *Orct* (*FBgn0019952*) and *Orct2* (*FBgn0086365*).** The protein alignment was produced using ClustalX (6)





**Supplementary Figure 14. Polygenic score (permuted lifespan phenotype).** The lifespan data were permuted 100 times. Each box represents the interquartile range (IQR) with the median as a black horizontal line; the whiskers represent values  $1.5 \times \text{IQR}$ ; outliers are represented as light blue points. The orange line connects the means within each  $p$ -value threshold



**Supplementary Table 1. DGRP lines, genotypes and phenotypes.** <sup>1</sup>*Wolbachia* status: 1- absent, 2- present; <sup>2</sup>Number of flies used for determining the mean and median lifespan; The lifespan data for several lines were derived from (3), where only the mean lifespan was given, hence for standard deviation, number of flies used and median lifespan is given as NA or not assigned.

| Line     | <i>Wolbachia</i> status <sup>1</sup> | Mean lifespan | Standard deviation | Number flies used <sup>2</sup> | Median lifespan | Number missing SNPs | Frequency missing SNPs |
|----------|--------------------------------------|---------------|--------------------|--------------------------------|-----------------|---------------------|------------------------|
| line_21  | 2                                    | 49.76         | 10.04              | 25                             | 51              | 28607               | 0.0130                 |
| line_26  | 1                                    | 46.39         | 9.13               | 23                             | 46              | 37755               | 0.0172                 |
| line_28  | 1                                    | 54.00         | 13.13              | 24                             | 56              | 23626               | 0.0108                 |
| line_31  | 1                                    | 71.88         | 7.98               | 25                             | 71              | 264022              | 0.1204                 |
| line_32  | 1                                    | 52.56         | 6.60               | 25                             | 54              | 42210               | 0.0192                 |
| line_38  | 1                                    | 48.27         | 19.75              | 22                             | 51              | 131573              | 0.0600                 |
| line_40  | 2                                    | 56.84         | 9.86               | 25                             | 58              | 6940                | 0.0032                 |
| line_41  | 1                                    | 59.24         | 23.78              | 25                             | 68              | 54027               | 0.0246                 |
| line_42  | 1                                    | 45.80         | 4.26               | 25                             | 46              | 35373               | 0.0161                 |
| line_45  | 1                                    | 48.27         | 6.76               | 22                             | 47              | 55160               | 0.0251                 |
| line_48  | 2                                    | 57.83         | 13.74              | 24                             | 61              | 167964              | 0.0766                 |
| line_49  | 2                                    | 53.84         | 6.11               | 25                             | 53              | 177048              | 0.0807                 |
| line_57  | 1                                    | 59.67         | 11.81              | 24                             | 60              | 34166               | 0.0156                 |
| line_59  | 1                                    | 58.52         | 22.65              | 25                             | 67              | 88646               | 0.0404                 |
| line_69  | 2                                    | 45.13         | 11.41              | 23                             | 45              | 82957               | 0.0378                 |
| line_73  | 2                                    | 52.33         | 6.04               | 24                             | 51              | 86777               | 0.0396                 |
| line_75  | 2                                    | 44.56         | 15.63              | 25                             | 48              | 38276               | 0.0175                 |
| line_83  | 1                                    | 68.40         | 14.24              | 20                             | 70              | 33671               | 0.0154                 |
| line_85  | 1                                    | 41.95         | 12.38              | 19                             | 44              | 201093              | 0.0917                 |
| line_88  | 1                                    | 58.79         | 11.56              | 24                             | 61              | 182007              | 0.0830                 |
| line_91  | 1                                    | 53.64         | 9.08               | 25                             | 53              | 71934               | 0.0328                 |
| line_93  | 1                                    | 46.58         | 9.61               | 24                             | 50              | 30236               | 0.0138                 |
| line_100 | 2                                    | 63.08         | 24.52              | 24                             | 75              | 136767              | 0.0623                 |
| line_101 | 1                                    | 75.83         | 7.97               | 24                             | 76              | 191396              | 0.0873                 |
| line_105 | 1                                    | 63.89         | 11.87              | 18                             | 68.5            | 45362               | 0.0207                 |
| line_109 | 1                                    | 60.60         | 15.39              | 25                             | 60              | 136599              | 0.0623                 |
| line_129 | 1                                    | 58.13         | 6.06               | 24                             | 58              | 27485               | 0.0125                 |
| line_136 | 2                                    | 76.56         | 11.83              | 25                             | 78              | 291551              | 0.1329                 |
| line_138 | 1                                    | 40.64         | 5.44               | 25                             | 41              | 50682               | 0.0231                 |
| line_142 | 2                                    | 63.48         | 10.79              | 25                             | 68              | 70638               | 0.0322                 |
| line_149 | 2                                    | 53.48         | 8.10               | 23                             | 56              | 48985               | 0.0223                 |
| line_153 | 2                                    | 54.96         | 9.68               | 25                             | 56              | 119735              | 0.0546                 |
| line_158 | 1                                    | 54.85         | 8.04               | 20                             | 52.5            | 85662               | 0.0391                 |
| line_161 | 1                                    | 67.72         | 8.27               | 25                             | 66              | 76017               | 0.0347                 |
| line_176 | 2                                    | 44.42         | 16.19              | 24                             | 44.5            | 25060               | 0.0114                 |
| line_177 | 1                                    | 30.16         | 7.94               | 19                             | 30              | 12027               | 0.0055                 |
| line_181 | 2                                    | 54.20         | 14.25              | 20                             | 58              | 19962               | 0.0091                 |
| line_189 | 2                                    | 62.32         | 14.20              | 22                             | 64              | 9351                | 0.0043                 |
| line_195 | 1                                    | 61.45         | 5.39               | 20                             | 62.5            | 61031               | 0.0278                 |
| line_208 | 1                                    | 53.76         | 2.60               | 25                             | 54              | 38622               | 0.0176                 |
| line_217 | 1                                    | 67.52         | 8.12               | 23                             | 68              | 74816               | 0.0341                 |
| line_223 | 2                                    | 48.73         | 5.33               | 22                             | 48              | 8088                | 0.0037                 |
| line_227 | 2                                    | 47.95         | 11.44              | 20                             | 49              | 35679               | 0.0163                 |
| line_228 | 1                                    | 65.81         | 4.17               | 21                             | 66              | 58009               | 0.0264                 |
| line_229 | 1                                    | 45.27         | 9.97               | 22                             | 45              | 58098               | 0.0265                 |
| line_233 | 1                                    | 54.24         | 9.02               | 25                             | 54              | 56039               | 0.0255                 |
| line_235 | 1                                    | 45.04         | 22.05              | 25                             | 50              | 28884               | 0.0132                 |
| line_237 | 2                                    | 60.65         | 6.19               | 17                             | 61              | 295495              | 0.1347                 |
| line_239 | 1                                    | 68.48         | 16.22              | 25                             | 71              | 41365               | 0.0189                 |
| line_256 | 2                                    | 67.67         | 12.10              | 24                             | 70              | 46500               | 0.0212                 |
| line_280 | 2                                    | 57.59         | 10.64              | 17                             | 62              | 37967               | 0.0173                 |
| line_287 | 2                                    | 41.09         | 18.80              | 23                             | 44              | 107899              | 0.0492                 |
| line_303 | 1                                    | 57.36         | NA                 | NA                             | NA              | 347361              | 0.1583                 |
| line_306 | 2                                    | 58.78         | NA                 | NA                             | NA              | 71315               | 0.0325                 |
| line_307 | 1                                    | 44.64         | NA                 | NA                             | NA              | 11333               | 0.0052                 |
| line_309 | 1                                    | 52.57         | 4.64               | 23                             | 54              | 167005              | 0.0761                 |
| line_310 | 2                                    | 22.13         | 7.82               | 24                             | 21              | 51190               | 0.0233                 |
| line_313 | 1                                    | 80.29         | NA                 | NA                             | NA              | 57064               | 0.0260                 |
| line_315 | 1                                    | 66.92         | NA                 | NA                             | NA              | 39840               | 0.0182                 |
| line_317 | 2                                    | 53.76         | 13.29              | 17                             | 56              | 161118              | 0.0734                 |
| line_318 | 2                                    | 49.36         | NA                 | NA                             | NA              | 58536               | 0.0267                 |

| Line     | <i>Wolbachia</i> status <sup>1</sup> | Mean lifespan | Standard deviation | Number flies used <sup>2</sup> | Median lifespan | Number missing SNPs | Frequency missing SNPs |
|----------|--------------------------------------|---------------|--------------------|--------------------------------|-----------------|---------------------|------------------------|
| line_320 | 2                                    | 65.70         | 13.07              | 23                             | 69              | 9102                | 0.0041                 |
| line_321 | 2                                    | 49.46         | 21.64              | 24                             | 47.5            | 50223               | 0.0229                 |
| line_324 | 1                                    | 48.00         | NA                 | NA                             | NA              | 3943                | 0.0018                 |
| line_325 | 1                                    | 42.61         | 4.68               | 23                             | 42              | 172915              | 0.0788                 |
| line_332 | 1                                    | 55.92         | NA                 | NA                             | NA              | 83533               | 0.0381                 |
| line_335 | 2                                    | 67.75         | NA                 | NA                             | NA              | 84844               | 0.0387                 |
| line_338 | 2                                    | 60.89         | 12.90              | 18                             | 66              | 281675              | 0.1284                 |
| line_340 | 2                                    | 59.17         | 9.01               | 24                             | 60              | 33414               | 0.0152                 |
| line_348 | 1                                    | 54.00         | 14.75              | 21                             | 57              | 8446                | 0.0039                 |
| line_350 | 1                                    | 59.25         | NA                 | NA                             | NA              | 171217              | 0.0781                 |
| line_352 | 2                                    | 46.78         | 11.52              | 23                             | 48              | 194440              | 0.0886                 |
| line_354 | 1                                    | 59.83         | 7.32               | 23                             | 58              | 5064                | 0.0023                 |
| line_355 | 2                                    | 52.45         | 8.35               | 22                             | 54              | 5391                | 0.0025                 |
| line_356 | 2                                    | 37.96         | 11.63              | 24                             | 41              | 78752               | 0.0359                 |
| line_357 | 1                                    | 58.36         | NA                 | NA                             | NA              | 111899              | 0.0510                 |
| line_358 | 1                                    | 62.67         | NA                 | NA                             | NA              | 71305               | 0.0325                 |
| line_359 | 1                                    | 56.28         | NA                 | NA                             | NA              | 51370               | 0.0234                 |
| line_360 | 2                                    | 39.91         | 5.90               | 23                             | 40              | 103452              | 0.0472                 |
| line_362 | 2                                    | 60.33         | NA                 | NA                             | NA              | 76686               | 0.0350                 |
| line_365 | 2                                    | 48.24         | 8.40               | 25                             | 47              | 38529               | 0.0176                 |
| line_367 | 1                                    | 64.15         | 8.53               | 20                             | 66              | 57968               | 0.0264                 |
| line_370 | 2                                    | 56.82         | 11.05              | 22                             | 57              | 16173               | 0.0074                 |
| line_371 | 1                                    | 39.12         | 3.11               | 25                             | 40              | 8108                | 0.0037                 |
| line_373 | 1                                    | 68.24         | 8.05               | 25                             | 68              | 129299              | 0.0589                 |
| line_374 | 2                                    | 66.04         | 8.24               | 23                             | 66              | 12381               | 0.0056                 |
| line_375 | 1                                    | 62.60         | NA                 | NA                             | NA              | 48611               | 0.0222                 |
| line_377 | 1                                    | 66.89         | 6.66               | 18                             | 68              | 243461              | 0.1110                 |
| line_379 | 1                                    | 68.08         | 16.81              | 25                             | 71              | 50561               | 0.0231                 |
| line_380 | 2                                    | 57.96         | 5.60               | 24                             | 58              | 105240              | 0.0480                 |
| line_381 | 1                                    | 41.08         | 8.83               | 25                             | 40              | 169648              | 0.0773                 |
| line_382 | 2                                    | 76.56         | 8.43               | 25                             | 78              | 15422               | 0.0070                 |
| line_383 | 2                                    | 56.84         | 10.78              | 25                             | 58              | 22528               | 0.0103                 |
| line_385 | 1                                    | 50.36         | 9.94               | 25                             | 54              | 110417              | 0.0503                 |
| line_386 | 1                                    | 54.40         | 5.06               | 25                             | 55              | 111546              | 0.0509                 |
| line_391 | 1                                    | 49.55         | 5.75               | 22                             | 50              | 30426               | 0.0139                 |
| line_392 | 1                                    | 66.29         | 7.29               | 21                             | 65              | 74115               | 0.0338                 |
| line_395 | 1                                    | 55.79         | 9.85               | 24                             | 58.5            | 5285                | 0.0024                 |
| line_399 | 1                                    | 59.26         | NA                 | NA                             | NA              | 41828               | 0.0191                 |
| line_405 | 2                                    | 50.56         | 9.35               | 25                             | 50              | 174491              | 0.0795                 |
| line_406 | 1                                    | 58.00         | 18.10              | 24                             | 64              | 20083               | 0.0092                 |
| line_409 | 2                                    | 23.92         | 9.52               | 24                             | 21              | 267506              | 0.1219                 |
| line_426 | 1                                    | 69.84         | 8.58               | 25                             | 70              | 267796              | 0.1221                 |
| line_427 | 1                                    | 52.08         | 7.40               | 24                             | 50              | 40502               | 0.0185                 |
| line_437 | 1                                    | 57.76         | 29.40              | 21                             | 72              | 10576               | 0.0048                 |
| line_439 | 1                                    | 57.80         | 12.72              | 25                             | 62              | 52056               | 0.0237                 |
| line_440 | 2                                    | 65.00         | 8.25               | 24                             | 68              | 155776              | 0.0710                 |
| line_441 | 2                                    | 65.32         | 17.65              | 25                             | 70              | 19207               | 0.0088                 |
| line_443 | 1                                    | 60.64         | 7.87               | 25                             | 60              | 156133              | 0.0712                 |
| line_461 | 2                                    | 36.96         | 8.34               | 24                             | 34.5            | 27413               | 0.0125                 |
| line_486 | 2                                    | 65.33         | 8.42               | 24                             | 70              | 61682               | 0.0281                 |
| line_491 | 1                                    | 61.13         | 8.06               | 23                             | 62              | 32192               | 0.0147                 |
| line_492 | 1                                    | 62.79         | 5.42               | 24                             | 64              | 17547               | 0.0080                 |
| line_502 | 1                                    | 69.91         | 14.05              | 23                             | 72              | 144007              | 0.0656                 |
| line_505 | 2                                    | 56.63         | 13.01              | 19                             | 58              | 6193                | 0.0028                 |
| line_508 | 1                                    | 53.60         | 8.08               | 25                             | 56              | 17207               | 0.0078                 |
| line_509 | 1                                    | 62.54         | 6.23               | 24                             | 63              | 31366               | 0.0143                 |
| line_513 | 2                                    | 51.38         | 18.29              | 24                             | 52              | 33730               | 0.0154                 |
| line_517 | 1                                    | 63.20         | 5.64               | 25                             | 64              | 39249               | 0.0179                 |
| line_528 | 2                                    | 51.08         | 14.87              | 24                             | 54              | 269113              | 0.1227                 |
| line_530 | 2                                    | 46.41         | 7.87               | 22                             | 49.5            | 26680               | 0.0122                 |
| line_531 | 2                                    | 53.61         | 19.39              | 23                             | 60              | 61208               | 0.0279                 |
| line_535 | 2                                    | 49.32         | 6.00               | 19                             | 50              | 30966               | 0.0141                 |
| line_551 | 2                                    | 59.68         | 8.98               | 25                             | 59              | 158784              | 0.0724                 |
| line_555 | 2                                    | 51.44         | 9.70               | 25                             | 53              | 46876               | 0.0214                 |
| line_559 | 1                                    | 59.76         | 6.67               | 25                             | 60              | 159498              | 0.0727                 |
| line_563 | 1                                    | 50.25         | 11.23              | 24                             | 56              | 281634              | 0.1284                 |
| line_566 | 1                                    | 65.32         | 10.60              | 25                             | 64              | 109330              | 0.0498                 |
| line_584 | 2                                    | 62.48         | 9.08               | 25                             | 65              | 57830               | 0.0264                 |
| line_589 | 2                                    | 65.96         | 15.57              | 24                             | 69.5            | 5253                | 0.0024                 |

| Line     | <i>Wolbachia</i> status <sup>1</sup> | Mean lifespan | Standard deviation | Number flies used <sup>2</sup> | Median lifespan | Number missing SNPs | Frequency missing SNPs |
|----------|--------------------------------------|---------------|--------------------|--------------------------------|-----------------|---------------------|------------------------|
| line_595 | 2                                    | 60.00         | 12.18              | 21                             | 60              | 56380               | 0.0257                 |
| line_596 | 1                                    | 37.94         | 11.18              | 16                             | 39.5            | 5999                | 0.0027                 |
| line_627 | 1                                    | 65.17         | 8.42               | 23                             | 66              | 239238              | 0.1091                 |
| line_630 | 1                                    | 66.08         | 13.17              | 25                             | 67              | 318694              | 0.1453                 |
| line_634 | 2                                    | 52.96         | 14.44              | 25                             | 53              | 141968              | 0.0647                 |
| line_639 | 2                                    | 75.82         | 15.23              | 22                             | 79              | 52102               | 0.0238                 |
| line_642 | 1                                    | 62.09         | 12.63              | 23                             | 64              | 35075               | 0.0160                 |
| line_646 | 2                                    | 58.00         | 17.57              | 23                             | 64              | 56570               | 0.0258                 |
| line_703 | 1                                    | 32.60         | 4.41               | 25                             | 33              | 38586               | 0.0176                 |
| line_705 | 2                                    | 63.67         | NA                 | NA                             | NA              | 51922               | 0.0237                 |
| line_707 | 2                                    | 60.64         | NA                 | NA                             | NA              | 47184               | 0.0215                 |
| line_712 | 2                                    | 45.52         | 7.84               | 25                             | 47              | 42014               | 0.0192                 |
| line_714 | 1                                    | 69.28         | 10.04              | 25                             | 72              | 77196               | 0.0352                 |
| line_716 | 2                                    | 60.00         | 14.50              | 25                             | 64              | 48417               | 0.0221                 |
| line_721 | 2                                    | 43.36         | 9.37               | 25                             | 42              | 44840               | 0.0204                 |
| line_727 | 2                                    | 39.38         | 10.82              | 21                             | 41              | 18656               | 0.0085                 |
| line_730 | 2                                    | 53.28         | 14.39              | 25                             | 55              | 33330               | 0.0152                 |
| line_732 | 1                                    | 66.76         | NA                 | NA                             | NA              | 168565              | 0.0768                 |
| line_737 | 2                                    | 53.16         | NA                 | NA                             | NA              | 113123              | 0.0516                 |
| line_738 | 2                                    | 59.88         | 7.98               | 24                             | 60              | 177872              | 0.0811                 |
| line_748 | 2                                    | 58.17         | 6.62               | 24                             | 58              | 8035                | 0.0037                 |
| line_757 | 1                                    | 29.65         | 18.25              | 20                             | 29              | 19606               | 0.0089                 |
| line_761 | 2                                    | 46.80         | 5.80               | 25                             | 46              | 35037               | 0.0160                 |
| line_765 | 1                                    | 33.96         | NA                 | NA                             | NA              | 40839               | 0.0186                 |
| line_774 | 1                                    | 64.36         | NA                 | NA                             | NA              | 195136              | 0.0890                 |
| line_776 | 2                                    | 59.75         | 6.22               | 24                             | 59              | 66813               | 0.0305                 |
| line_783 | 2                                    | 59.46         | 8.96               | 24                             | 60              | 36733               | 0.0167                 |
| line_786 | 2                                    | 53.83         | NA                 | NA                             | NA              | 45760               | 0.0209                 |
| line_787 | 2                                    | 44.92         | NA                 | NA                             | NA              | 55841               | 0.0255                 |
| line_790 | 2                                    | 49.48         | 8.66               | 23                             | 52              | 29843               | 0.0136                 |
| line_796 | 2                                    | 42.32         | 2.78               | 25                             | 43              | 74641               | 0.0340                 |
| line_799 | 1                                    | 60.00         | NA                 | NA                             | NA              | 32325               | 0.0147                 |
| line_801 | 2                                    | 69.44         | 6.34               | 25                             | 72              | 134359              | 0.0613                 |
| line_802 | 2                                    | 49.84         | 18.12              | 25                             | 54              | 326828              | 0.1490                 |
| line_804 | 2                                    | 67.24         | 13.26              | 25                             | 71              | 72753               | 0.0332                 |
| line_805 | 2                                    | 51.17         | 7.08               | 24                             | 51              | 25221               | 0.0115                 |
| line_808 | 1                                    | 52.48         | NA                 | NA                             | NA              | 43379               | 0.0198                 |
| line_810 | 1                                    | 49.12         | 15.80              | 25                             | 55              | 18254               | 0.0083                 |
| line_812 | 1                                    | 57.04         | 7.27               | 24                             | 59              | 142941              | 0.0652                 |
| line_818 | 2                                    | 60.36         | 8.65               | 25                             | 62              | 71898               | 0.0328                 |
| line_819 | 2                                    | 40.40         | 10.23              | 25                             | 40              | 3750                | 0.0017                 |
| line_820 | 2                                    | 38.92         | NA                 | NA                             | NA              | 64348               | 0.0293                 |
| line_821 | 2                                    | 72.70         | 10.42              | 23                             | 74              | 126181              | 0.0575                 |
| line_822 | 2                                    | 53.58         | NA                 | NA                             | NA              | 64906               | 0.0296                 |
| line_837 | 2                                    | 60.09         | 13.78              | 23                             | 60              | 18713               | 0.0085                 |
| line_843 | 1                                    | 53.38         | 6.96               | 24                             | 56              | 23051               | 0.0105                 |
| line_849 | 1                                    | 58.05         | 16.62              | 20                             | 59.5            | 148060              | 0.0675                 |
| line_850 | 2                                    | 44.74         | 5.43               | 23                             | 44              | 5883                | 0.0027                 |
| line_852 | 2                                    | 49.88         | NA                 | NA                             | NA              | 22057               | 0.0101                 |
| line_853 | 2                                    | 47.32         | 10.02              | 25                             | 49              | 193021              | 0.0880                 |
| line_855 | 2                                    | 54.58         | 8.32               | 24                             | 56.5            | 94566               | 0.0431                 |
| line_857 | 1                                    | 54.83         | 8.17               | 23                             | 55              | 159737              | 0.0728                 |
| line_859 | 2                                    | 60.84         | NA                 | NA                             | NA              | 38944               | 0.0178                 |
| line_861 | 2                                    | 55.67         | 14.72              | 24                             | 58.5            | 79452               | 0.0362                 |
| line_879 | 2                                    | 59.68         | 7.82               | 19                             | 56              | 35422               | 0.0162                 |
| line_882 | 2                                    | 53.54         | 19.83              | 24                             | 58.5            | 25058               | 0.0114                 |
| line_884 | 2                                    | 62.17         | 7.62               | 24                             | 62              | 150768              | 0.0687                 |
| line_887 | 2                                    | 48.04         | 10.11              | 23                             | 51              | 24927               | 0.0114                 |
| line_890 | 2                                    | 50.79         | 7.41               | 24                             | 51              | 38702               | 0.0176                 |
| line_892 | 2                                    | 51.42         | 16.77              | 24                             | 58.5            | 32480               | 0.0148                 |
| line_894 | 1                                    | 61.54         | 8.79               | 24                             | 62              | 38462               | 0.0175                 |
| line_897 | 2                                    | 49.60         | 7.80               | 20                             | 52              | 18548               | 0.0085                 |
| line_900 | 1                                    | 46.71         | 10.34              | 21                             | 48              | 49074               | 0.0224                 |
| line_907 | 1                                    | 60.44         | NA                 | NA                             | NA              | 168053              | 0.0766                 |
| line_908 | 1                                    | 47.29         | 12.74              | 21                             | 45              | 46539               | 0.0212                 |
| line_911 | 1                                    | 37.68         | NA                 | NA                             | NA              | 92701               | 0.0423                 |
| line_913 | 2                                    | 56.61         | 11.17              | 18                             | 58              | 319509              | 0.1456                 |

**Supplementary Table 2. Genetic variation within 165 DGRP lines.** <sup>a</sup>Within replicate line data was available for 165 fly lines. Mean lifespan was calculated from 165 fly lines as well as the rest of the calculations; <sup>b</sup> Total genetic variance; <sup>c</sup> Variance within replicates or lines; <sup>d</sup> Total phenotypic variance ( $S_G^2 + S_E^2$ ); <sup>e</sup> Broad sense heritability ( $S_G^2/S_P^2$ ); <sup>f</sup> Coefficient of genetic variation ( $100S_G/\text{Mean}$ ); <sup>g</sup> Coefficient of environmental variation ( $100S_E/\text{Mean}$ );

| Mean lifespan <sup>a</sup> | $S_G^2$ <sup>b</sup> | $S_E^2$ <sup>c</sup> | $S_P^2$ <sup>d</sup> | $H^2$ <sup>e</sup> | $CV_G^f$ | $CV_E^g$ |
|----------------------------|----------------------|----------------------|----------------------|--------------------|----------|----------|
| 55.149                     | 93.748               | 133.41               | 227.158              | 0.413              | 17.557   | 20.944   |

**Supplementary Table 3. Single-SNP GWAS, genes near the top 50 SNPs; NA - not within a gene**

| SNP             | CHR | P-value                | $\beta$<br>coefficient | Within<br>Gene | 5'      | 5' distance<br>[bp] | 3'      | 3' distance<br>[bp] |
|-----------------|-----|------------------------|------------------------|----------------|---------|---------------------|---------|---------------------|
| 2L_10068812_SNP | 2L  | 9.41x10 <sup>-06</sup> | -7.41                  | CG31714        |         |                     |         |                     |
| 2L_10070707_SNP | 2L  | 6.77x10 <sup>-06</sup> | -6.24                  | CG31714        |         |                     |         |                     |
| 2L_1632386_SNP  | 2L  | 5.90x10 <sup>-08</sup> | -5.85                  | NA             | chinmo  | 18872               | RFeSP   | 18204               |
| 2L_1632388_SNP  | 2L  | 3.74x10 <sup>-07</sup> | -5.66                  | NA             | chinmo  | 18870               | RFeSP   | 18206               |
| 2L_1696065_SNP  | 2L  | 2.49x10 <sup>-06</sup> | -7.09                  | chinmo         |         |                     |         |                     |
| 2L_1835028_SNP  | 2L  | 1.11x10 <sup>-05</sup> | 4.44                   | NA             | c-cup   | 2472                | wry     | 1603                |
| 2L_2279849_SNP  | 2L  | 2.21x10 <sup>-06</sup> | -11.63                 | NA             | CG17242 | 10731               | CG4271  | 5315                |
| 2L_3480710_SNP  | 2L  | 6.77x10 <sup>-07</sup> | -9.84                  | NA             | CG15414 | 45                  | Thor    | 1098                |
| 2L_3746990_SNP  | 2L  | 1.14x10 <sup>-05</sup> | -9.63                  | CG10019        |         |                     |         |                     |
| 2L_3752571_SNP  | 2L  | 2.35x10 <sup>-07</sup> | -12.55                 | CG10019        |         |                     |         |                     |
| 2R_19786647_SNP | 2R  | 4.66x10 <sup>-07</sup> | -12.29                 | Lpt            |         |                     |         |                     |
| 2R_4308343_SNP  | 2R  | 8.41x10 <sup>-06</sup> | -7.84                  | NA             | CSN7    | 150                 | CG43296 | 3132                |
| 2R_4308355_SNP  | 2R  | 7.86x10 <sup>-06</sup> | -7.89                  | NA             | CSN7    | 138                 | CG43296 | 3144                |
| 3L_11792808_SNP | 3L  | 5.37x10 <sup>-06</sup> | -3.74                  | CG10361        |         |                     |         |                     |
| 3L_14778027_SNP | 3L  | 3.71x10 <sup>-06</sup> | -10.20                 | bmm            |         |                     |         |                     |
| 3L_14778725_SNP | 3L  | 3.50x10 <sup>-06</sup> | -10.17                 | bmm            |         |                     |         |                     |
| 3L_14780164_SNP | 3L  | 4.00x10 <sup>-06</sup> | -10.14                 | CG13472        |         |                     |         |                     |
| 3L_14781414_SNP | 3L  | 1.45x10 <sup>-06</sup> | -11.72                 | CG13472        |         |                     |         |                     |
| 3L_17762728_SNP | 3L  | 1.13x10 <sup>-05</sup> | -9.58                  | NA             | Adgf-A  | 5471                | CG42815 | 21452               |
| 3L_18140585_SNP | 3L  | 6.51x10 <sup>-06</sup> | -4.60                  | NA             | CG7330  | 1438                | gk      | 2067                |
| 3L_18810814_SNP | 3L  | 4.36x10 <sup>-06</sup> | 4.29                   | CG14073        |         |                     |         |                     |
| 3L_18934159_SNP | 3L  | 1.06x10 <sup>-05</sup> | -5.94                  | CG32204        |         |                     |         |                     |
| 3L_1966180_SNP  | 3L  | 7.14x10 <sup>-06</sup> | -5.81                  | CG1140         |         |                     |         |                     |
| 3L_4628971_SNP  | 3L  | 5.16x10 <sup>-06</sup> | -7.43                  | Rpd3           |         |                     |         |                     |
| 3L_5319539_SNP  | 3L  | 1.12x10 <sup>-05</sup> | -5.30                  | NA             | shep    | 48496               | lama    | 17181               |
| 3L_5373941_SNP  | 3L  | 7.12x10 <sup>-06</sup> | -11.04                 | Ir64a          |         |                     |         |                     |
| 3L_5636181_SNP  | 3L  | 2.69x10 <sup>-06</sup> | -9.57                  | Blimp-1        |         |                     |         |                     |
| 3L_8650506_SNP  | 3L  | 6.13x10 <sup>-06</sup> | -4.02                  | NA             | h       | 18353               | Pex7    | 6395                |
| 3L_9507749_SNP  | 3L  | 2.97x10 <sup>-06</sup> | -3.42                  | CG33700        |         |                     |         |                     |
| 3R_14921157_SNP | 3R  | 1.15x10 <sup>-05</sup> | -7.33                  | ATPsyn-d       |         |                     |         |                     |
| 3R_15338010_SNP | 3R  | 1.08x10 <sup>-05</sup> | -4.16                  | det            |         |                     |         |                     |
| 3R_15338014_SNP | 3R  | 1.02x10 <sup>-05</sup> | -4.11                  | det            |         |                     |         |                     |
| 3R_15340424_SNP | 3R  | 6.62x10 <sup>-06</sup> | -4.38                  | Dys            |         |                     |         |                     |
| 3R_15950064_SNP | 3R  | 1.17x10 <sup>-05</sup> | -8.84                  | NA             | Gr92a   | 2066                | CG5023  | 45095               |
| 3R_18577501_SNP | 3R  | 5.63x10 <sup>-06</sup> | -7.22                  | CG7023         |         |                     |         |                     |
| 3R_19071977_SNP | 3R  | 5.64x10 <sup>-07</sup> | -5.62                  | CG4467         |         |                     |         |                     |
| 3R_20944700_SNP | 3R  | 9.10x10 <sup>-06</sup> | -5.99                  | CG31510        |         |                     |         |                     |
| 3R_21259405_SNP | 3R  | 7.92x10 <sup>-06</sup> | -9.58                  | Fur1           |         |                     |         |                     |
| 3R_21913681_SNP | 3R  | 1.04x10 <sup>-05</sup> | -5.28                  | dys            |         |                     |         |                     |
| 3R_23482833_SNP | 3R  | 9.26x10 <sup>-06</sup> | -3.51                  | NA             | Mlc1    | 452                 | tau     | 115                 |
| 3R_24748071_SNP | 3R  | 1.19x10 <sup>-05</sup> | -3.92                  | Doa            |         |                     |         |                     |
| 3R_25189263_SNP | 3R  | 1.05x10 <sup>-05</sup> | -6.41                  | NA             | Cnx99A  | 43898               | Ptp99A  | 13725               |
| 3R_25562159_SNP | 3R  | 5.27x10 <sup>-06</sup> | -4.18                  | CG7601         |         |                     |         |                     |
| 3R_25921654_SNP | 3R  | 1.04x10 <sup>-05</sup> | -8.93                  | sima           |         |                     |         |                     |
| 3R_25921693_SNP | 3R  | 9.79x10 <sup>-06</sup> | -8.94                  | sima           |         |                     |         |                     |
| 3R_25921696_SNP | 3R  | 9.79x10 <sup>-06</sup> | -8.94                  | sima           |         |                     |         |                     |
| 3R_8922024_SNP  | 3R  | 1.19x10 <sup>-05</sup> | -7.43                  | timeout        |         |                     |         |                     |
| X_20940365_SNP  | X   | 4.95x10 <sup>-06</sup> | -14.60                 | bves           |         |                     |         |                     |
| X_604933_SNP    | X   | 8.31x10 <sup>-06</sup> | -3.53                  | sdk            |         |                     |         |                     |
| X_9282626_SNP   | X   | 2.78x10 <sup>-06</sup> | 7.06                   | mgl            |         |                     |         |                     |

**Supplementary Table 4. Top 30 genes, gene-based analysis.** \*Empirical *p*-values, based on 1,000,000 permutations

| FlyBaseGene ID | Gene Symbol     | Chr | N SNPs | Gene-based <i>p</i> -value* |
|----------------|-----------------|-----|--------|-----------------------------|
| FBgn0036603    | <i>CG13062</i>  | 3L  | 22     | 7.10x10 <sup>-05</sup>      |
| FBgn0036870    | <i>CG14095</i>  | 3L  | 5      | 2.97x10 <sup>-04</sup>      |
| FBgn0037985    | <i>ssp5</i>     | 3R  | 18     | 3.29x10 <sup>-04</sup>      |
| FBgn0051956    | <i>pgant4</i>   | 2L  | 74     | 5.08x10 <sup>-04</sup>      |
| FBgn0039462    | <i>CG14252</i>  | 3R  | 62     | 5.32x10 <sup>-04</sup>      |
| FBgn0087005    | <i>rtp</i>      | 3R  | 2      | 5.47x10 <sup>-04</sup>      |
| FBgn0029843    | <i>Nep1</i>     | X   | 161    | 6.22x10 <sup>-04</sup>      |
| FBgn0039075    | <i>CG4393</i>   | 3R  | 194    | 7.01x10 <sup>-04</sup>      |
| FBgn0037156    | <i>CG11523</i>  | 3L  | 5      | 9.02x10 <sup>-04</sup>      |
| FBgn0016120    | <i>ATPsyn-d</i> | 3R  | 10     | 9.63x10 <sup>-04</sup>      |
| FBgn0051928    | <i>CG31928</i>  | 2L  | 33     | 9.73x10 <sup>-04</sup>      |
| FBgn0036008    | <i>CG3408</i>   | 3L  | 68     | 9.87x10 <sup>-04</sup>      |
| FBgn0050154    | <i>CG30154</i>  | 2R  | 21     | 9.99x10 <sup>-04</sup>      |
| FBgn0036208    | <i>CG10361</i>  | 3L  | 70     | 1.02x10 <sup>-03</sup>      |
| FBgn0263004    | <i>CG43312</i>  | 3L  | 2      | 1.03x10 <sup>-03</sup>      |
| FBgn0044324    | <i>Chro</i>     | 3L  | 26     | 1.08x10 <sup>-03</sup>      |
| FBgn0031596    | <i>CG15429</i>  | 2L  | 46     | 1.24x10 <sup>-03</sup>      |
| FBgn0262818    | <i>CG43189</i>  | 2R  | 3      | 1.26x10 <sup>-03</sup>      |
| FBgn0025638    | <i>Roc1a</i>    | X   | 5      | 1.51x10 <sup>-03</sup>      |
| FBgn0036165    | <i>chrb</i>     | 3L  | 137    | 1.58x10 <sup>-03</sup>      |
| FBgn0035011    | <i>CG13589</i>  | 2R  | 17     | 1.64x10 <sup>-03</sup>      |
| FBgn0037307    | <i>Tim17a2</i>  | 3R  | 9      | 1.65x10 <sup>-03</sup>      |
| FBgn0039385    | <i>CG5913</i>   | 3R  | 19     | 1.70x10 <sup>-03</sup>      |
| FBgn0033769    | <i>CG8768</i>   | 2R  | 34     | 1.82x10 <sup>-03</sup>      |
| FBgn0039890    | <i>CG2316</i>   | 4   | 25     | 1.98x10 <sup>-03</sup>      |
| FBgn0037960    | <i>mthl5</i>    | 3R  | 48     | 2.11x10 <sup>-03</sup>      |
| FBgn0051998    | <i>CG31998</i>  | 4   | 16     | 2.11x10 <sup>-03</sup>      |
| FBgn0032217    | <i>CG4972</i>   | 2L  | 52     | 2.32x10 <sup>-03</sup>      |
| FBgn0260003    | <i>Dys</i>      | 3R  | 3104   | 2.40x10 <sup>-03</sup>      |
| FBgn0031601    | <i>Dim1</i>     | 2L  | 24     | 2.44x10 <sup>-03</sup>      |

**Supplementary Table 5. Top 30 genes, gene-based analysis (genes±5kb).** \*Empirical *p*-values, based on 1,000,000 permutations

| FlyBaseGene ID | Gene Symbol     | Chr | N SNPs | Gene-based <i>p</i> -value* |
|----------------|-----------------|-----|--------|-----------------------------|
| FBgn0029843    | <i>Nep1</i>     | X   | 245    | 6.61x10 <sup>-04</sup>      |
| FBgn0035827    | <i>CG14252</i>  | 3L  | 337    | 7.57x10 <sup>-04</sup>      |
| FBgn0262275    | <i>mir-2280</i> | 2L  | 312    | 8.62x10 <sup>-04</sup>      |
| FBgn0015300    | <i>Ssl</i>      | 2R  | 224    | 9.17x10 <sup>-04</sup>      |
| FBgn0086075    | <i>CR34704</i>  | 3L  | 316    | 1.06x10 <sup>-03</sup>      |
| FBgn0052783    | <i>CG32783</i>  | X   | 20     | 1.09x10 <sup>-03</sup>      |
| FBgn0031367    | <i>c-cup</i>    | 2L  | 349    | 1.18x10 <sup>-03</sup>      |
| FBgn0053703    | <i>CG33703</i>  | 3L  | 369    | 1.32x10 <sup>-03</sup>      |
| FBgn0010408    | <i>RpS9</i>     | 3L  | 375    | 1.37x10 <sup>-03</sup>      |
| FBgn0017556    | <i>Prosa4T2</i> | 2R  | 240    | 1.42x10 <sup>-03</sup>      |
| FBgn0029501    | <i>Crtp</i>     | 2R  | 253    | 1.43x10 <sup>-03</sup>      |
| FBgn0051998    | <i>CG31998</i>  | 4   | 60     | 1.59x10 <sup>-03</sup>      |
| FBgn0039890    | <i>CG2316</i>   | 4   | 66     | 1.61x10 <sup>-03</sup>      |
| FBgn0053702    | <i>CG33702</i>  | 3L  | 371    | 1.75x10 <sup>-03</sup>      |
| FBgn0061188    | <i>Yu</i>       | 2R  | 213    | 1.78x10 <sup>-03</sup>      |
| FBgn0262988    | <i>CG43296</i>  | 2R  | 195    | 1.95x10 <sup>-03</sup>      |
| FBgn0025387    | <i>CG12184</i>  | X   | 93     | 2.03x10 <sup>-03</sup>      |
| FBgn0053701    | <i>CR33701</i>  | 3L  | 390    | 2.27x10 <sup>-03</sup>      |
| FBgn0035012    | <i>CG13590</i>  | 2R  | 199    | 2.31x10 <sup>-03</sup>      |
| FBgn0035011    | <i>CG13589</i>  | 2R  | 188    | 2.44x10 <sup>-03</sup>      |
| FBgn0263344    | <i>CR43425</i>  | 4   | 28     | 2.47x10 <sup>-03</sup>      |
| FBgn0036008    | <i>CG3408</i>   | 3L  | 409    | 2.53x10 <sup>-03</sup>      |
| FBgn0053700    | <i>CG33700</i>  | 3L  | 701    | 2.80x10 <sup>-03</sup>      |
| FBgn0035281    | <i>Cpr62Bc</i>  | 3L  | 377    | 2.82x10 <sup>-03</sup>      |
| FBgn0039889    | <i>Arl4</i>     | 4   | 49     | 2.98x10 <sup>-03</sup>      |
| FBgn0260003    | <i>Dys</i>      | 3R  | 3271   | 3.30x10 <sup>-03</sup>      |
| FBgn0044324    | <i>Chro</i>     | 3L  | 80     | 3.34x10 <sup>-03</sup>      |
| FBgn0053978    | <i>CG33978</i>  | 4   | 91     | 3.45x10 <sup>-03</sup>      |
| FBgn0037202    | <i>Ssl1</i>     | 3L  | 69     | 3.72x10 <sup>-03</sup>      |
| FBgn0052786    | <i>CG32786</i>  | X   | 9      | 3.73x10 <sup>-03</sup>      |

**Supplementary Table 6. Genes belonging to the IIS and TOR pathways**

|                   |   |
|-------------------|---|
| IIS pathway genes | <i>I4-3-epsilon</i> (FBgn0020238), <i>Akt1</i> (FBgn0010379), <i>B4</i> (FBgn0023407), <i>chico</i> (FBgn0024248), <i>dock</i> (FBgn0010583), <i>foxo</i> (FBgn0038197), <i>hpo</i> (FBgn0261456), <i>Ilp1</i> (FBgn0044051), <i>Ilp2</i> (FBgn0036046), <i>Ilp3</i> (FBgn0044050), <i>Ilp4</i> (FBgn0044049), <i>Ilp5</i> (FBgn0044048), <i>Ilp6</i> (FBgn0044047), <i>Ilp7</i> (FBgn0044046), <i>Ilp8</i> (FBgn0036690), <i>Impl2</i> (FBgn0001257), <i>InR</i> (FBgn0013984), <i>Lnk</i> (FBgn0028717), <i>melt</i> (FBgn0023001), <i>Phlpp</i> (FBgn0032749), <i>Pi3K21B</i> (FBgn0020622), <i>Pi3K92E</i> (FBgn0015279), <i>Pten</i> (FBgn0026379), <i>S6KII</i> (FBgn0262866), <i>sgg</i> (FBgn0003371), <i>step</i> (FBgn0086779), <i>wdb</i> (FBgn0027492), <i>Pdk1</i> (FBgn0020386), <i>B4</i> (FBgn0023407)  |
| TOR pathway genes | <i>Atg1</i> (FBgn0260945), <i>chrp</i> (FBgn0036165), <i>dm</i> (FBgn0262656), <i>Dredd</i> (FBgn0020381), <i>eIF-4B</i> (FBgn0020660), <i>eIF-4E</i> (FBgn0015218), <i>eIF4G</i> (FBgn0023213), <i>gig</i> (FBgn0005198), <i>HLH106</i> (FBgn0261283), <i>L</i> (FBgn0001332), <i>lkb1</i> (FBgn0038167), <i>Lst8</i> (FBgn0264691), <i>Mipp2</i> (FBgn0026060), <i>Mo25</i> (FBgn0017572), <i>par-1</i> (FBgn0260934), <i>path</i> (FBgn0036007), <i>pgc</i> (FBgn0016053), <i>pico</i> (FBgn0261811), <i>Pka-C1</i> (FBgn0000273), <i>Raga</i> (FBgn0037647), <i>RagC</i> (FBgn0033272), <i>raptor</i> (FBgn0029840), <i>Rheb</i> (FBgn0041191), <i>ric1</i> (FBgn0031006), <i>RpS6</i> (FBgn0261592), <i>S6k</i> (FBgn0015806), <i>scyl</i> (FBgn0041094), <i>Sesn</i> (FBgn0034897), <i>Sik2</i> (FBgn0025625), <i>Sin1</i> (FBgn0033935), <i>slif</i> (FBgn0037203), <i>SNF1A</i> (FBgn0023169), <i>Thor</i> (FBgn0261560), <i>Tor</i> (FBgn0021796), <i>tor</i> (FBgn0003733), <i>Crtc</i> (FBgn0036746), <i>Tsc1</i> (FBgn0026317), <i>Tif-IA</i> (FBgn0032988) |

**Supplementary Table 7. Ranking and *p*-values of the top 20 GO categories**

| GO ID      | Go category  | <i>p</i> -value        | Number of genes |
|------------|--|------------------------|-----------------|
| GO:0046692 | sperm competition  | 2.21x10 <sup>-04</sup> | 22              |
| GO:0022626 | cytosolic ribosome   | 2.68x10 <sup>-04</sup> | 4               |
| GO:0046527 | glucosyltransferase activity                                 | 3.04x10 <sup>-04</sup> | 11              |
| GO:0019318 | hexose metabolic process                                     | 4.44x10 <sup>-04</sup> | 63              |
| GO:0006006 | glucose metabolic process                                    | 4.87x10 <sup>-04</sup> | 43              |
| GO:0035074 | pupation   | 5.51x10 <sup>-04</sup> | 5               |
| GO:0006007 | glucose catabolic process                                    | 5.65x10 <sup>-04</sup> | 33              |
| GO:0006433 | prolyl-tRNA aminoacylation                                   | 5.76x10 <sup>-04</sup> | 2               |
| GO:0004827 | proline-tRNA ligase activity                                 | 5.76x10 <sup>-04</sup> | 2               |
| GO:0006096 | glycolysis   | 8.47x10 <sup>-04</sup> | 25              |
| GO:0043564 | Ku70:Ku80 complex  | 9.67x10 <sup>-04</sup> | 3               |
| GO:0003684 | damaged DNA binding  | 1.10x10 <sup>-03</sup> | 23              |
| GO:0004197 | cysteine-type endopeptidase activity                         | 1.10x10 <sup>-03</sup> | 28              |
| GO:0000090 | mitotic anaphase   | 1.25x10 <sup>-03</sup> | 17              |
| GO:0046365 | monosaccharide catabolic process                             | 1.29x10 <sup>-03</sup> | 34              |
| GO:0019320 | hexose catabolic process                                     | 1.29x10 <sup>-03</sup> | 34              |
| GO:0045254 | pyruvate dehydrogenase complex                               | 1.42x10 <sup>-03</sup> | 4               |
| GO:0006085 | acetyl-CoA biosynthetic process                              | 2.11x10 <sup>-03</sup> | 4               |
| GO:0005996 | monosaccharide metabolic process                             | 2.27x10 <sup>-03</sup> | 67              |
| GO:0008534 | oxidized purine nucleobase lesion DNA N-glycosylase activity | 2.81x10 <sup>-03</sup> | 2               |

**Supplementary Table 8. Ranking and *p*-values of the top 20 GO categories (genes±5kb)**

| GO ID      | Go category  | <i>p</i> -value        | Number of genes |
|------------|--|------------------------|-----------------|
| GO:0007442 | Hindgut morphogenesis                              | 7.27x10 <sup>-04</sup> | 19              |
| GO:0016485 | Protein processing                                 | 8.38x10 <sup>-04</sup> | 696             |
| GO:0051604 | Protein maturation                                 | 9.21x10 <sup>-04</sup> | 699             |
| GO:0004165 | dodecenoyl-CoA delta-isomerase activity            | 9.47x10 <sup>-04</sup> | 5               |
| GO:0035079 | Polytene chromosome puffing                        | 1.12x10 <sup>-03</sup> | 8               |
| GO:0007350 | Blastoderm segmentation                            | 1.21x10 <sup>-03</sup> | 21              |
| GO:0000090 | Mitotic anaphase                                   | 1.24x10 <sup>-03</sup> | 17              |
| GO:0006508 | proteolysis  | 1.44x10 <sup>-03</sup> | 671             |
| GO:0043564 | Ku70:Ku80 complex                                  | 1.57x10 <sup>-03</sup> | 3               |
| GO:0070011 | Peptidase activity acting on L-amino acid peptides | 1.61x10 <sup>-03</sup> | 596             |
| GO:0004177 | Aminopeptidase activity                            | 1.81x10 <sup>-03</sup> | 39              |
| GO:0042600 | chorion  | 2.14x10 <sup>-03</sup> | 38              |
| GO:0006426 | glycyl-tRNA aminoacylation                         | 2.39x10 <sup>-03</sup> | 2               |
| GO:0004820 | glycine-tRNA ligase activity                       | 2.39x10 <sup>-03</sup> | 2               |
| GO:0008233 | Peptidase activity                                 | 2.41x10 <sup>-03</sup> | 602             |
| GO:0004822 | isoleucine-tRNA ligase activity                    | 2.42x10 <sup>-03</sup> | 2               |
| GO:0006428 | isoleucyl-tRNA aminoacylation                      | 2.42x10 <sup>-03</sup> | 2               |
| GO:0008527 | Taste receptor activity                            | 2.44x10 <sup>-03</sup> | 62              |
| GO:0045254 | Pyruvate dehydrogenase complex                     | 2.48x10 <sup>-03</sup> | 4               |
| GO:0004175 | endopeptidase activity                             | 2.85x10 <sup>-03</sup> | 445             |

**Supplementary Table 9. Polygenic score (permuted vs. original lifespan).** SEM- standard error of the mean

| p threshold | Mean $R^2$<br>(Lifespan) | SEM<br>(Lifespan) | Mean $R^2$<br>(permuted lifespan) | SEM<br>(permuted lifespan) | One-sided<br>t-test p-value |
|-------------|--------------------------|-------------------|-----------------------------------|----------------------------|-----------------------------|
| ≤1.0        | 0.042                    | 0.0030            | 0.019                             | 0.0019                     | 9.83x10 <sup>-10</sup>      |
| ≤0.9        | 0.042                    | 0.0031            | 0.019                             | 0.0019                     | 9.18x10 <sup>-10</sup>      |
| ≤0.8        | 0.042                    | 0.0031            | 0.019                             | 0.0019                     | 8.39x10 <sup>-10</sup>      |
| ≤0.7        | 0.043                    | 0.0031            | 0.019                             | 0.0019                     | 7.05x10 <sup>-10</sup>      |
| ≤0.6        | 0.043                    | 0.0031            | 0.020                             | 0.0019                     | 6.56x10 <sup>-10</sup>      |
| ≤0.5        | 0.043                    | 0.0031            | 0.020                             | 0.0019                     | 5.88x10 <sup>-10</sup>      |
| ≤0.2        | 0.043                    | 0.0031            | 0.019                             | 0.0019                     | 2.19x10 <sup>-10</sup>      |
| ≤0.1        | 0.044                    | 0.0031            | 0.019                             | 0.0019                     | 5.26x10 <sup>-11</sup>      |
| ≤0.05       | 0.044                    | 0.0031            | 0.019                             | 0.0019                     | 1.76x10 <sup>-11</sup>      |
| ≤0.01       | 0.047                    | 0.0032            | 0.018                             | 0.0019                     | 4.57x10 <sup>-13</sup>      |
| ≤0.005      | 0.047                    | 0.0031            | 0.019                             | 0.0002                     | 7.23x10 <sup>-13</sup>      |
| ≤0.001      | 0.047                    | 0.0032            | 0.019                             | 0.0019                     | 1.57x10 <sup>-12</sup>      |
| ≤0.0005     | 0.046                    | 0.0031            | 0.020                             | 0.0021                     | 4.79x10 <sup>-11</sup>      |
| ≤0.0001     | 0.042                    | 0.0030            | 0.020                             | 0.0020                     | 4.02x10 <sup>-09</sup>      |

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