

**Table S3. Downregulated genes by equol treatment in both wild-type and *nfe2l2a<sup>fh318</sup>* larvae.**

| Genes             | Gene names in ZFIN | Human homologs | Equol-treated vs untreated in wild-type larvae | Equol-treated vs untreated in <i>nfe2l2a<sup>fh318</sup></i> larvae |
|-------------------|--------------------|----------------|--|---|
| ptch1.1           | ptch1              | PTCH1          | 0.210526316                                    | 0.644444444   |
| si:ch211-250g4.3  | si:ch211-250g4.3   |                | 0.25   | 0.625   |
| si:dkey-22i16.7   | si:dkey-22i16.7    |                | 0.333333333                                    | 0.432432432   |
| si:ch73-31d8.2    | si:ch73-31d8.2     |                | 0.338461538                                    | 0.450704225   |
| cabp2a            | cabp2a             | CABP2          | 0.340425532                                    | 0.642857143   |
| lrata             | lrata              | LRAT           | 0.340425532                                    | 0.416666667   |
| zanl              | zanl               | ZAN            | 0.367816092                                    | 0.655172414   |
| si:ch211-173m16.1 | si:ch211-173m16.1  |                | 0.378378378                                    | 0.239130435   |
| si:ch211-286k11.4 | cdhr5b             | CDHR5          | 0.382352941                                    | 0.407407407   |
| CR933839.1        |                    |                | 0.396551724                                    | 0.514285714   |
| lss               | lss                | LSS            | 0.409090909                                    | 0.647058824   |
| CABZ01072043.1    |                    |                | 0.409090909                                    | 0.485714286   |
| matn3a            | matn3a             | MATN3          | 0.411504425                                    | 0.552083333   |
| cers3a            | cers3a             | CERS3          | 0.413793103                                    | 0.580645161   |
| lrcc38a           | lrcc38a            | LRRC38         | 0.416666667                                    | 0.414634146   |
| CABZ01079764.1    |                    |                | 0.418867925                                    | 0.336065574   |
| CR855996.2        |                    |                | 0.423312883                                    | 0.447368421   |
| tm6sf2            | tm6sf2             | TM6SF2         | 0.425531915                                    | 0.325   |
| mf12              | meltf              | MELTF          | 0.427672956                                    | 0.654761905   |
| si:dkey-76i15.1   | vimr2              |                | 0.428571429                                    | 0.615384615   |
| nkx2.4b           | nkx2.4b            | NKX2-4         | 0.454545455                                    | 0.5625  |
| bfb               | bfb                | CFB            | 0.454545455                                    | 0.486111111   |
| asb14b            | asb14b             | ASB14          | 0.469387755                                    | 0.641025641   |
| cngb3.2           | cngb3.2            | CNGB3          | 0.471428571                                    | 0.277777778   |
| clul1             | clul1              | CLUL1          | 0.473684211                                    | 0.416666667   |
| lox15b            | lox15b             | LOXL4          | 0.475  | 0.666666667   |
| si:dkeyp-69c1.9   | si:dkeyp-69c1.9    |                | 0.476190476                                    | 0.590909091   |
| S1PR3             |                    | S1PR3          | 0.47826087                                     | 0.571428571   |
| cnga3b            | cnga3b             | CNGA3          | 0.478873239                                    | 0.612903226   |
| abcb11a           | abcb11a            | ABCB11         | 0.487804878                                    | 0.523809524   |
| rpe65a            | rpe65a             | RPE65          | 0.492662474                                    | 0.377300613   |
| abcg5             | abcg5              | ABCG5          | 0.5  | 0.555555556   |
| laynb             | laynb              | LAYN           | 0.5  | 0.606060606   |
| slc6a19a.1        | slc6a19a.1         | SLC6A19        | 0.505376344                                    | 0.328125  |
| CT990561.1        |                    |                | 0.511111111                                    | 0.625   |
| CABZ01068356.1    |                    |                | 0.515625                                       | 0.49122807  |
| fgf11a            | fgf11a             | FGF11          | 0.516129032                                    | 0.625   |
| si:dkey-35m8.1    | si:dkey-35m8.1     |                | 0.518518519                                    | 0.466666667   |
| vav3a             | vav3a              | VAV3           | 0.520547945                                    | 0.65625   |
| slc52a3           | slc52a3            | SLC52A3        | 0.526315789                                    | 0.44  |
| itih6             | itih6              | ITIH6          | 0.527777778                                    | 0.636363636   |
| slc16a7           | slc16a7            |                | 0.529411765                                    | 0.5   |
| cdkn2d            | cdkn2d             | CDKN2D         | 0.530612245                                    | 0.387096774   |
| si:dkey-103j14.5  | si:dkey-103j14.5   | ASRGL1         | 0.53125  | 0.52  |
| rpe65c            | rpe65c             | RPE65          | 0.533333333                                    | 0.371428571   |
| CABZ01079986.1    |                    |                | 0.533333333                                    | 0.625   |
| RBP1              |                    | RBP1           | 0.534883721                                    | 0.565217391   |
| si:ch211-198d23.1 | si:ch211-198d23.1  |                | 0.535714286                                    | 0.607142857   |
| lxb1b             | lxb1b              | LBX1           | 0.541666667                                    | 0.666666667   |
| zgc:65851         | zgc:65851          |                | 0.544927536                                    | 0.595092025   |
| ccl19a.1          | ccl19a.1           | CCL19          | 0.545454545                                    | 0.470588235   |
| MYO7B (1 of many) |                    | MYO7B          | 0.545454545                                    | 0.447368421   |
| cacnb4a           | cacnb4a            | CACNB4         | 0.547619048                                    | 0.666666667   |
| st6galnac4        | st6galnac4         | ST6GALNAC4     | 0.551724138                                    | 0.636363636   |
| c3a.2             | c3a.2              | C3             | 0.554744526                                    | 0.659259259   |
| emp3b             | emp3b              | EMP3           | 0.555555556                                    | 0.451612903   |
| itln3             | itln3              | ITLN1          | 0.555555556                                    | 0.666666667   |
| rdh5              | rdh5               | RDH5           | 0.557894737                                    | 0.351351351   |
| si:ch211-170d12.2 | si:ch211-170d12.2  |                | 0.559139785                                    | 0.612244898   |
| si:rp71-68n21.9   | si:rp71-68n21.9    | KLHL9          | 0.56   | 0.558823529   |
| alas2             | alas2              | ALAS2          | 0.560283688                                    | 0.59  |
| chia.6            | chia.6             | CHIA           | 0.561403509                                    | 0.511627907   |

|                   |                   |          |             |             |
|-------------------|-------------------|----------|-------------|-------------|
| si:dkey-196j8.2   | si:dkey-196j8.2   |          | 0.5625      | 0.625       |
| cyp2r1            | cyp2r1            | CYP2R1   | 0.5625      | 0.64556962  |
| myo15b            | myo15b            | MYO15B   | 0.567164179 | 0.590909091 |
| nr5a2             | nr5a2             | NR5A2    | 0.567567568 | 0.583333333 |
| igflr1            | igflr1            | IGFLR1   | 0.568181818 | 0.575757576 |
| si:ch211-224m9.4  | si:ch211-224m9.4  |          | 0.568965517 | 0.666666667 |
| rca2.2            | rca2.2            | C4BPA    | 0.571428571 | 0.652173913 |
| tfcp2l1           | tfcp2l1           | TFCP2L1  | 0.571428571 | 0.666666667 |
| grik1b            | grik1b            | GRIK1    | 0.576271186 | 0.636363636 |
| pacrg             | pacrg             | PACRG    | 0.578947368 | 0.62962963  |
| PLEKHG3           |                   | PLEKHG3  | 0.578947368 | 0.666666667 |
| si:dkeyp-77c8.3   | si:dkeyp-77c8.3   |          | 0.578947368 | 0.515151515 |
| si:dkey-1j5.4     | si:dkey-1j5.4     |          | 0.580645161 | 0.52        |
| CR848841.1        |                   |          | 0.581818182 | 0.620689655 |
| mmachc            | mmachc            | MMACHC   | 0.583333333 | 0.558823529 |
| rdh8b             | rdh8b             | RDH8     | 0.583333333 | 0.432432432 |
| abca4a            | abca4a            | ABCA4    | 0.586206897 | 0.588235294 |
| tmie              | tmie              | TMIE     | 0.587301587 | 0.472727273 |
| crym              | crym              | CRYM     | 0.588235294 | 0.53125     |
| opn6a             | opn6a             | OPN5     | 0.588235294 | 0.379310345 |
| ngs               | ngs               |          | 0.589041096 | 0.555555556 |
| slc5a11           | slc5a11           | SLC5A11  | 0.590909091 | 0.634146341 |
| col8a1a           | col8a1a           | COL8A1   | 0.591549296 | 0.65        |
| si:ch73-147f11.1  | dipk2b            | DIPK2B   | 0.592592593 | 0.666666667 |
| si:ch73-334e23.1  | si:ch73-334e23.1  |          | 0.592592593 | 0.6         |
| si:ch211-214p13.3 | si:ch211-214p13.3 |          | 0.59375     | 0.629032258 |
| fuom              | fuom              | FUOM     | 0.594594595 | 0.666666667 |
| syt6a             | syt6a             | SYT6     | 0.596491228 | 0.575757576 |
| ponzr1            | ponzr1            |          | 0.598984772 | 0.591666667 |
| slc12a1           | slc12a1           | SLC12A1  | 0.6         | 0.6         |
| syt6b             | syt6b             | SYT6     | 0.6         | 0.433333333 |
| si:dkey-96n2.1    | si:dkey-96n2.1    |          | 0.6         | 0.555555556 |
| fdxr              | fdxr              | FDXR     | 0.602941176 | 0.6         |
| fabp11b           | fabp4b            | FABP4    | 0.603448276 | 0.527272727 |
| elmod3            | elmod3            | ELMOD3   | 0.604166667 | 0.541666667 |
| si:ch73-52p7.1    | si:ch73-52p7.1    |          | 0.605263158 | 0.536585366 |
| TMEM229A          |                   | TMEM229A | 0.607142857 | 0.588235294 |
| aqp8b             | aqp8b             | AQP8     | 0.607329843 | 0.40942029  |
| pym1              | pym1              | PYM1     | 0.608       | 0.580645161 |
| hnf4g             | hnf4g             | HNF4G    | 0.608695652 | 0.625       |
| btbd17a           | btbd17a           | BTBD17   | 0.611111111 | 0.413793103 |
| pdzd2             | pdzd2             | PDZD2    | 0.614035088 | 0.647058824 |
| syndig1l          | syndig1l          | SYNDIG1L | 0.614285714 | 0.511627907 |
| slc5a8l           | slc5a8l           | SLC5A8   | 0.615384615 | 0.416666667 |
| ccb12_2           | kyat3             | KYAT3    | 0.620689655 | 0.461538462 |
| adamts13          | adamts13          | ADAMTS13 | 0.625       | 0.512195122 |
| pex12             | pex12             | PEX12    | 0.625       | 0.666666667 |
| plch1             | plch1             | PLCH1    | 0.625       | 0.588235294 |
| cntnap1           | cntnap1           | CNTNAP1  | 0.627906977 | 0.454545455 |
| mb                | mb                | MB       | 0.633270321 | 0.623471883 |
| vil1              | vil1              | VIL1     | 0.637007078 | 0.597449909 |
| vstm4b            | vstm4b            | VSTM4    | 0.638297872 | 0.619047619 |
| cnrip1b           | cnrip1b           | CNRIP1   | 0.639344262 | 0.6         |
| sncga             | sncga             | SNCG     | 0.639344262 | 0.651515152 |
| slc13a3           | slc13a3           | SLC13A3  | 0.64        | 0.596153846 |
| zgc:162331        | zgc:162331        | SUSD3    | 0.64        | 0.41509434  |
| CU855947.1        |                   |          | 0.64        | 0.615384615 |
| slc5a9            | slc5a9            | SLC5A9   | 0.640625    | 0.452830189 |
| si:dkey-117m1.4   | si:dkey-117m1.4   |          | 0.641304348 | 0.660377358 |
| antxr1b           | antxr1b           | ANTXR1   | 0.642857143 | 0.458333333 |
| nkain2            | nkain2            | NKAIN2   | 0.64516129  | 0.303571429 |
| cnga1             | cnga1b            | CNGA1    | 0.647058824 | 0.566666667 |
| adipoqb           | adipoqb           | ADIPOQ   | 0.65        | 0.630434783 |
| mycla             | mycla             | MYCL     | 0.651376147 | 0.43220339  |
| si:ch1073-82l19.1 | zyg11l            | ZYG11B   | 0.652173913 | 0.6         |
| si:dkey-181d19.6  | si:dkey-181d19.6  |          | 0.652173913 | 0.608695652 |
| zgc:110782        | zgc:110782        |          | 0.653061224 | 0.606060606 |

|                    |                  |         |             |             |
|--------------------|------------------|---------|-------------|-------------|
| plekhd1            | plekhd1          | PLEKHD1 | 0.653846154 | 0.653846154 |
| si:ch211-102c2.4   | si:ch211-102c2.4 |         | 0.653846154 | 0.666666667 |
| brf2               | brf2             | BRF2    | 0.655172414 | 0.444444444 |
| gabra6a            | gabra6a          | GABRA6  | 0.655737705 | 0.555555556 |
| urad               | urad             | URAD    | 0.657142857 | 0.589041096 |
| enpp6              | enpp6            | ENPP6   | 0.658536585 | 0.435897436 |
| dpf1               | dpf1             | DPF1    | 0.659090909 | 0.637681159 |
| asb10              | asb10            | ASB10   | 0.662337662 | 0.632075472 |
| slc15a1b           | slc15a1b         | SLC15A1 | 0.662721893 | 0.622222222 |
| cntn4_1            | cntn4            | CNTN4   | 0.666666667 | 0.578947368 |
| faah2b             | faah2b           | FAAH2   | 0.666666667 | 0.318181818 |
| gpr88              | gpr88            | GPR88   | 0.666666667 | 0.666666667 |
| rlbp1b             | rlbp1b           | RLBP1   | 0.666666667 | 0.469565217 |
| si:dkey-156n14.3_1 | si:dkey-156n14.3 | ZXDB    | 0.666666667 | 0.566666667 |
| kcna4              | kcna4            | KCNA4   | 0.666666667 | 0.666666667 |