| 4 | |
|---|---|
| | 1 |

| Triply regulated | | Quadruply regulated | | | | |
|------------------|--------|---------------------|----------------------------|--------|----------|--------------------------------|
| Rank | Weight | PubMedID | Tag(s) | Weight | PubMedID | Tag(s) |
| 1 | 3.60% | 18778409 | Ageing | 3.20% | 18778409 | Ageing |
| 2 | 1.60% | 21589891 | Germline | 3.00% | 21931806 | Chemical stimulus |
| 3 | 1.60% | 23352664 | Lifespan | 2.40% | 23300685 | Oxidative stress, lifespan |
| 4 | 1.60% | 21931806 | Chemical stimulus | 1.90% | 20798549 | Metabolism |
| 5 | 1.50% | 20798549 | Metabolism | 1.70% | 22560298 | Gene expression |
| 6 | 1.50% | 19745112 | Gene expression | 1.70% | 20816092 | Insulin-like signaling, stress |
| 7 | 1.40% | 15772128 | Development | 1.60% | 19379696 | Germline immortality |
| 8 | 1.40% | 21765926 | Stress hormesis | 1.60% | 22083954 | Innate immunity |
| 9 | 1.30% | 21363964 | Dosage compensation | 1.50% | 21363964 | Dosage compensation |
| 10 | 1.20% | 23300685 | Oxidative stress, lifespan | 1.10% | 21723504 | Lifespan, stress response |

b

С

| | Gene Array | | Triply regulated | | Quadruply regulated | |
|---------------------|--|-----|------------------|-----|---------------------|--|
| | - | Up | Down | Up | Down | |
| | daf-2 pathway targets | 14 | 1% | 8 | % | |
| Longevity | daf-16 pathway targets | | 4% | 5% | 2% | |
| mutants and | tants and Age-regulated genes | | 3% | | 2% | |
| dauers | Shared dauer and insulin pathway targets | 1% | 0% | 1% | 0% | |
| | Most responsive daf-16 regulated genes | 16% | 6% | 14% | 2% | |
| Food | Genes in response to fasting | 0% | 11% | 0% | 11% | |
| deprivation | Starvation (dietary restriction) | 4% | | 3% | | |
| Stress | <i>skn-1</i> pathway targets | 0% | | 0% | | |
| | Hyperbaric oxygen stress (oxidative stress) | 1% | 0% | 1% | 0% | |
| | Commonly regulated in oxidative stress and aging | 4% | 0% | 4% | 0% | |
| | Commonly regulated in oxidative stress and/af-16 | 8% | 0% | 7% | 0% | |
| | Commonly regulated in stress, aging, and/af-16 | 4% | 0% | 10% | 0% | |
| Polyphenolic action | 1 mM Resveratrol regulated genes | | 1% | | 1% | |
| | 200 µM Quercetin regulated genes | | 2% | | 1% | |
| action | 300 µM Tannic acid regulated genes | | 0% | | 0% | |
| Immune | Immune effector genes | 15% | | 4% | | |
| response | daf-2 pathway and Pseudomonas aeruginosa infection | | 2% | | 2% | |
| Toxin | Genes in response to cadmium toxicity | 6% | 6% | 5% | 4% | |

| Enzyme | Total # in Genome | Abbr. | N2 WT | Punc-25::GFP | dnj-14(ok237) | dnj-14(tm3223) |
|---------------------------------------|-------------------|-------|-------|--------------|---------------|----------------|
| Cytochrome P450 | 75 | сур | 8 | 12 | 17 | 12 |
| UDP-glucuronosyl/glucosyl transferase | 72 | ugt | 7 | 8 | 15 | 12 |
| Glutathione peroxidase | 8 | gpx | 0 | 0 | 0 | 0 |
| Glutathione S-transferase | 57 | gst | 1 | 5 | 5 | 3 |
| Superoxide dismutase | 5 | sod | 0 | 0 | 1 | 0 |
| Thioredoxin | 5 | trx | 1 | 1 | 1 | 0 |
| Thioredoxin reductase | 2 | trxr | 0 | 0 | 0 | 0 |
| Catalase | 3 | ctl | 0 | 0 | 0 | 0 |
| Peroxiredoxin | 3 | prdx | 0 | 0 | 0 | 0 |
| Sestrin | 1 | sesn | 0 | 0 | 0 | 0 |
| Isocitrate dehydrogenase | 2 | idh | 0 | 0 | 0 | 0 |
| Glutaredoxin | 5 | glrx | 0 | 0 | 0 | 0 |
| Ferritin | 2 | ftn | 1 | 1 | 1 | 1 |

Figure S7. Data set associations. (A) Top ten data set associations for stringent common DEGs queried using the SPELL search engine. Gene expression data sets and their associated PubMed ID, tags and weight are indicated. (B) Manual comparisons of gene expression patterns of relevant gene clusters or transcriptional profiles in the literature to findings in this study. Only a small proportion of common DEGs overlap with expression profiles of various stress conditions. (C) Lack of enrichment of known detoxification enzymes in response to ethosuximide treatment.