

**Uncovering Buffered Pleiotropy: A Genome-Scale Screen for *mel-28* Genetic Interactors in *Caenorhabditis elegans***

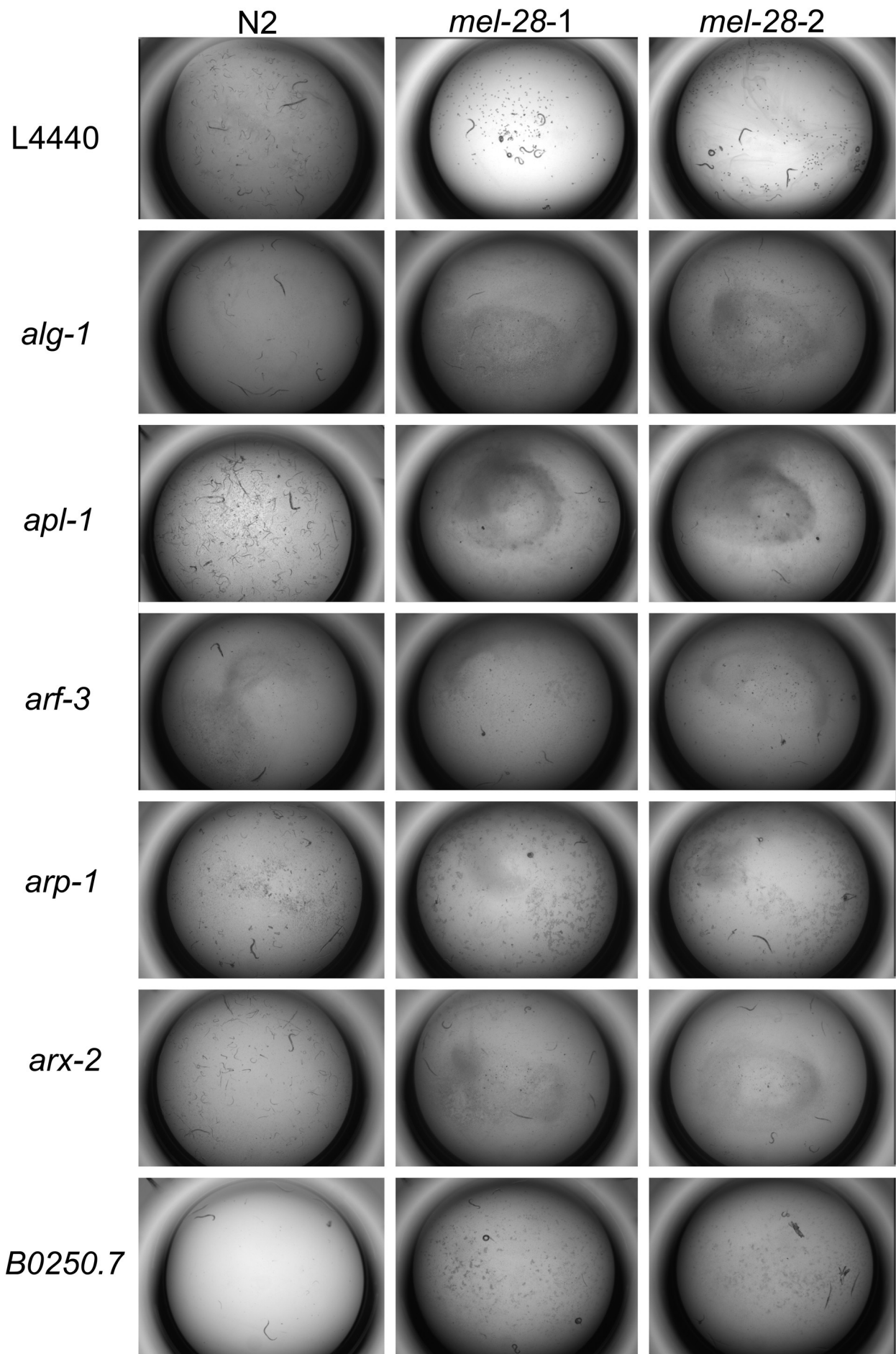
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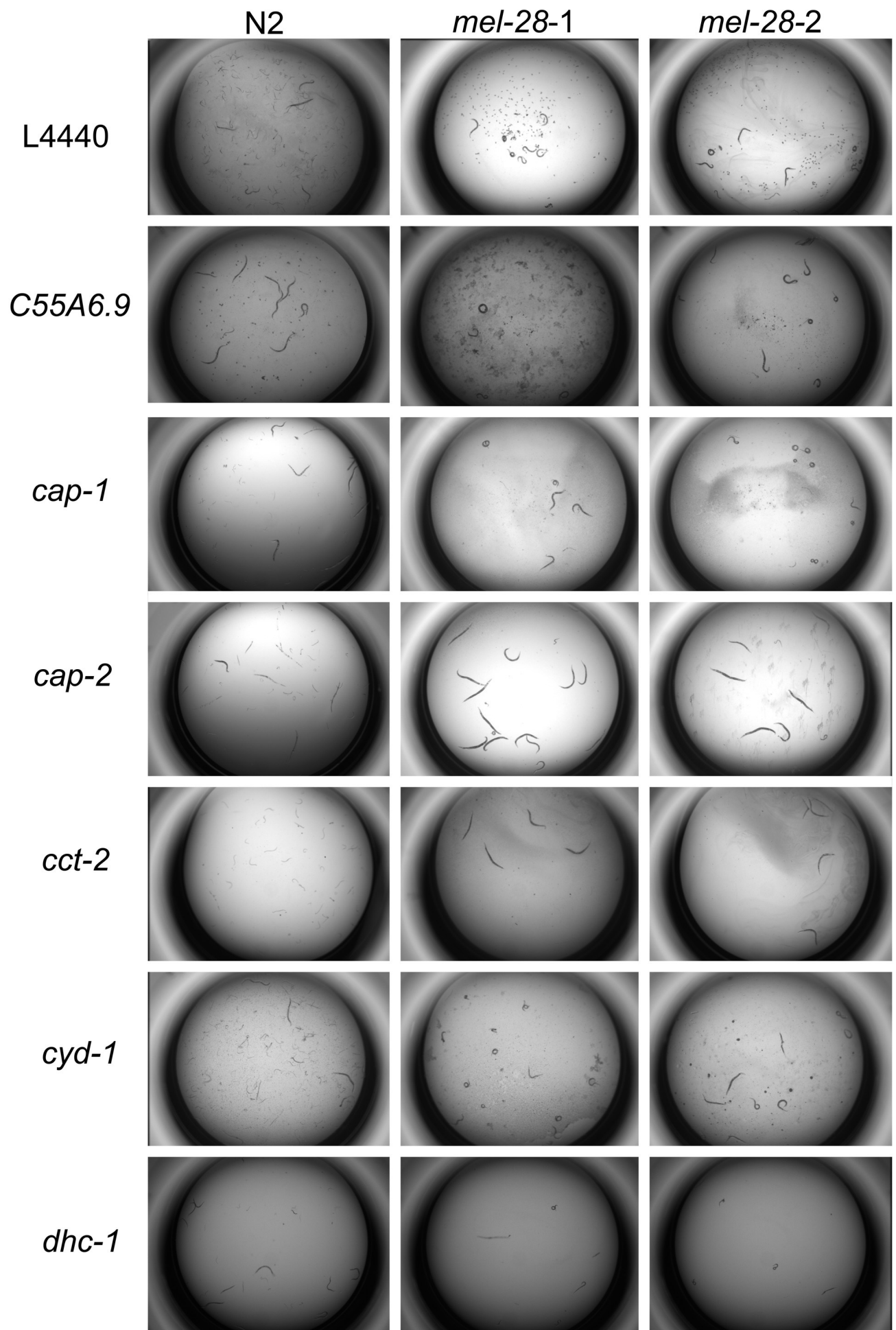
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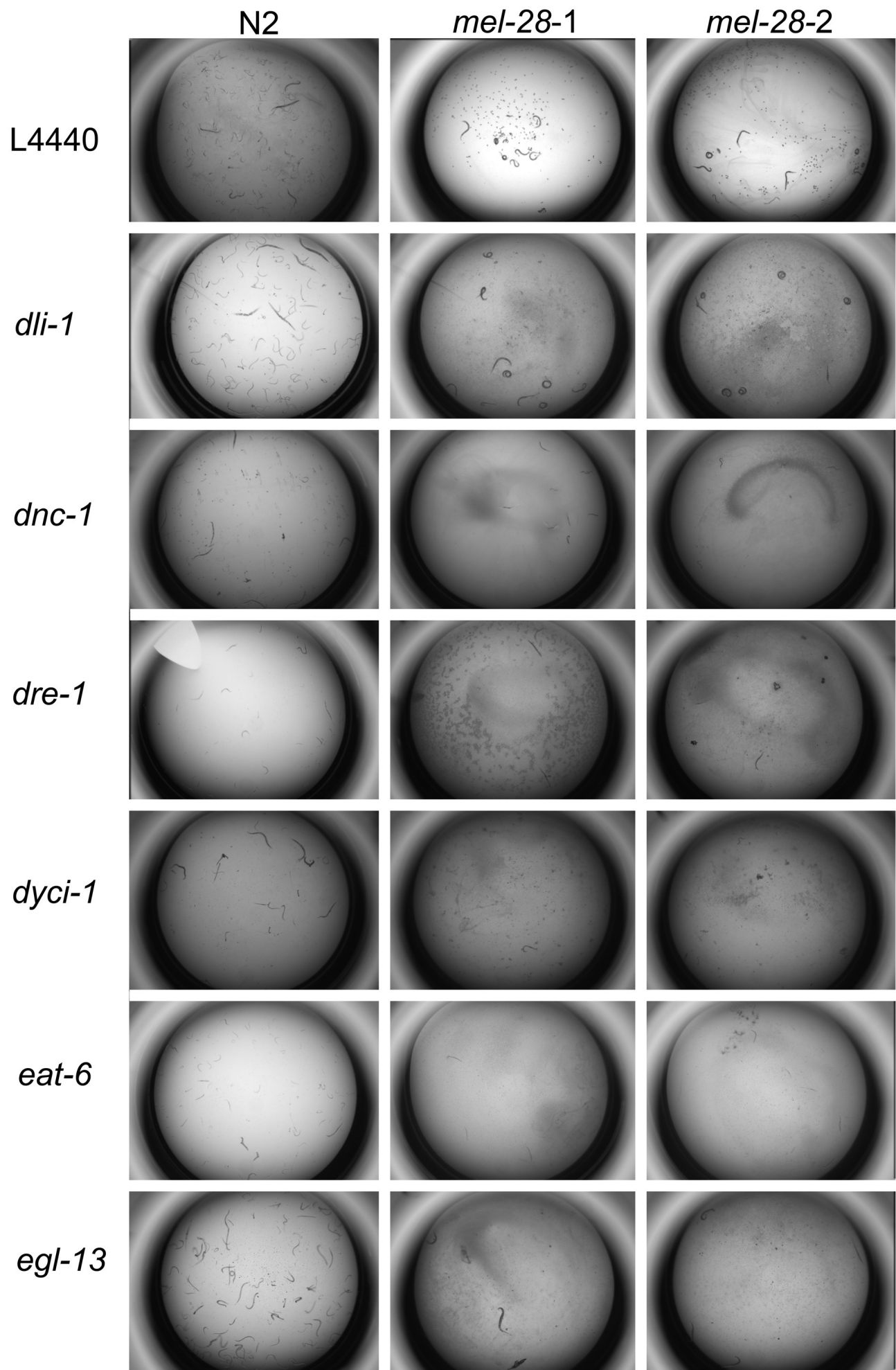
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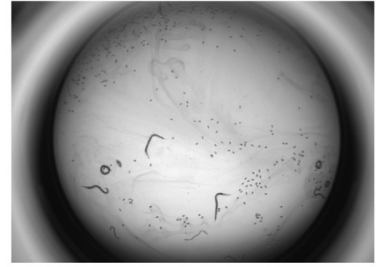
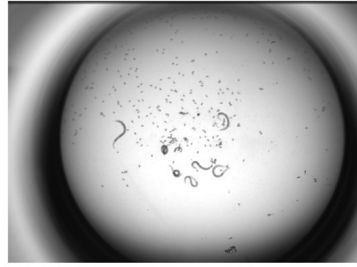
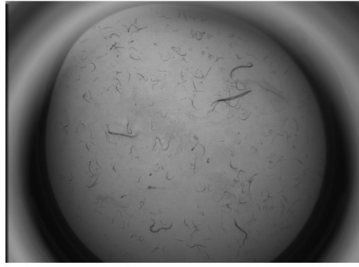


N2

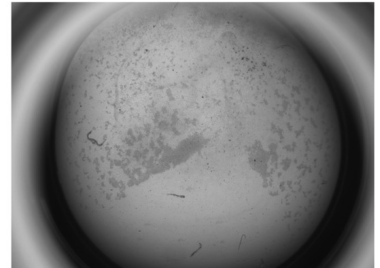
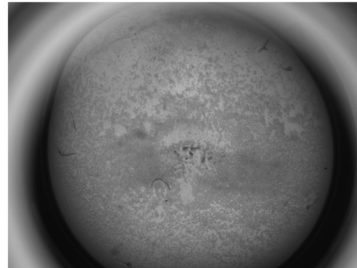
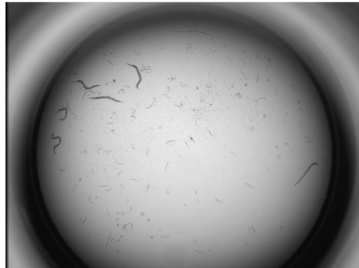
*mel-28-1*

*mel-28-2*

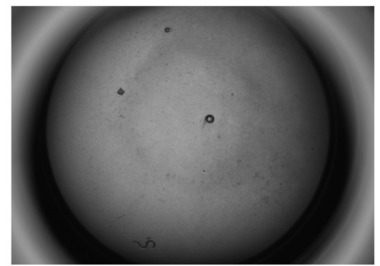
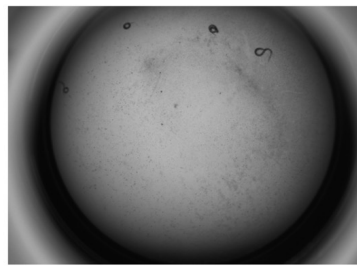
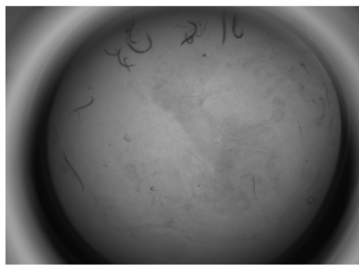
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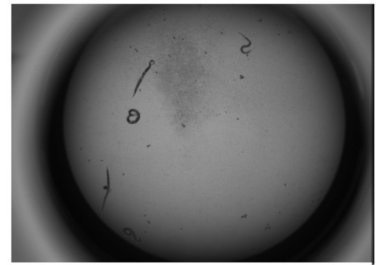
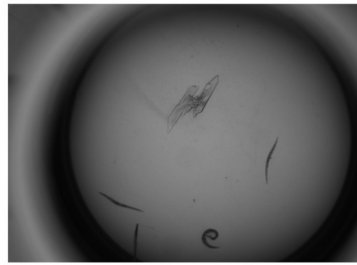
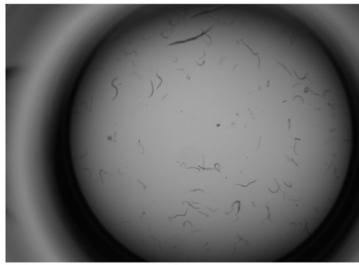
*ego-2*



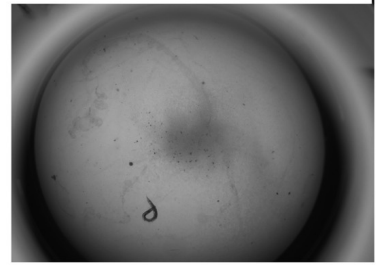
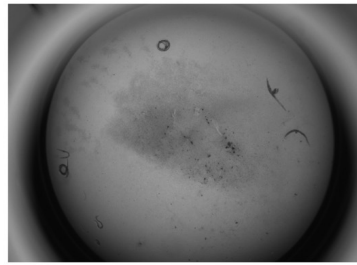
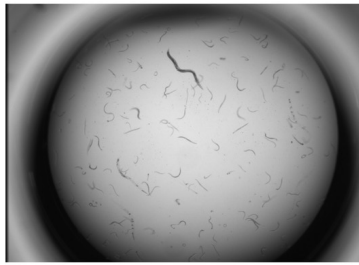
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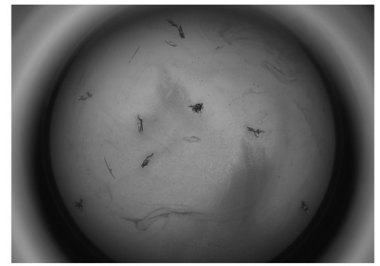
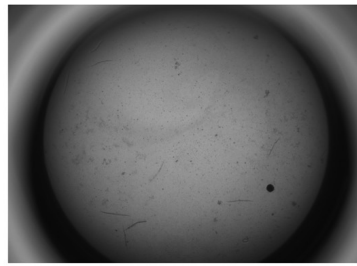
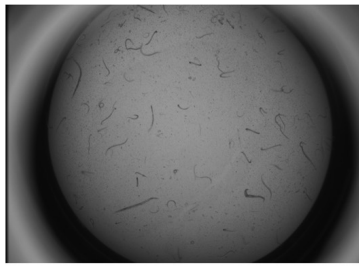
*exos-3*



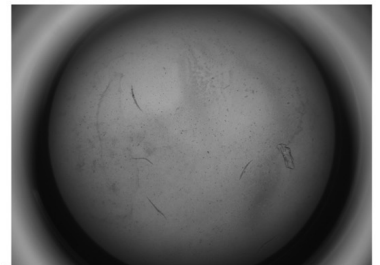
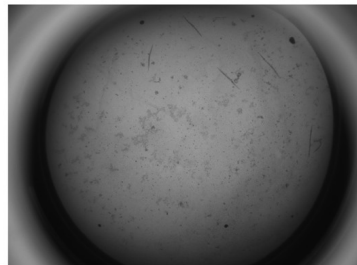
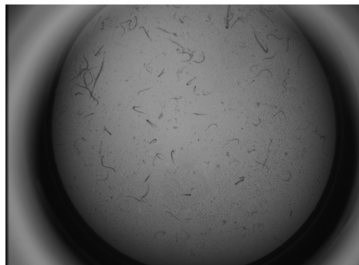
F19B6.1

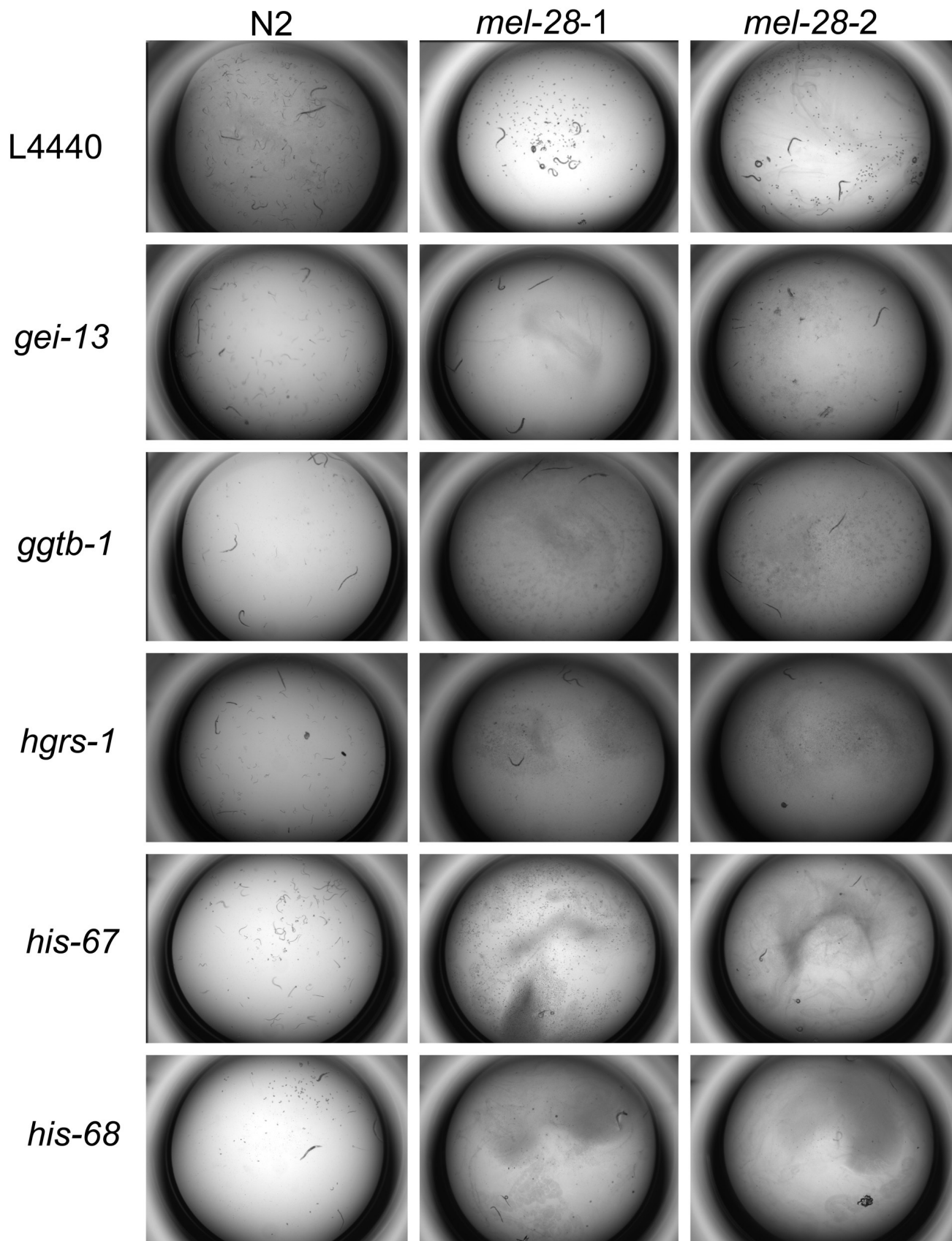


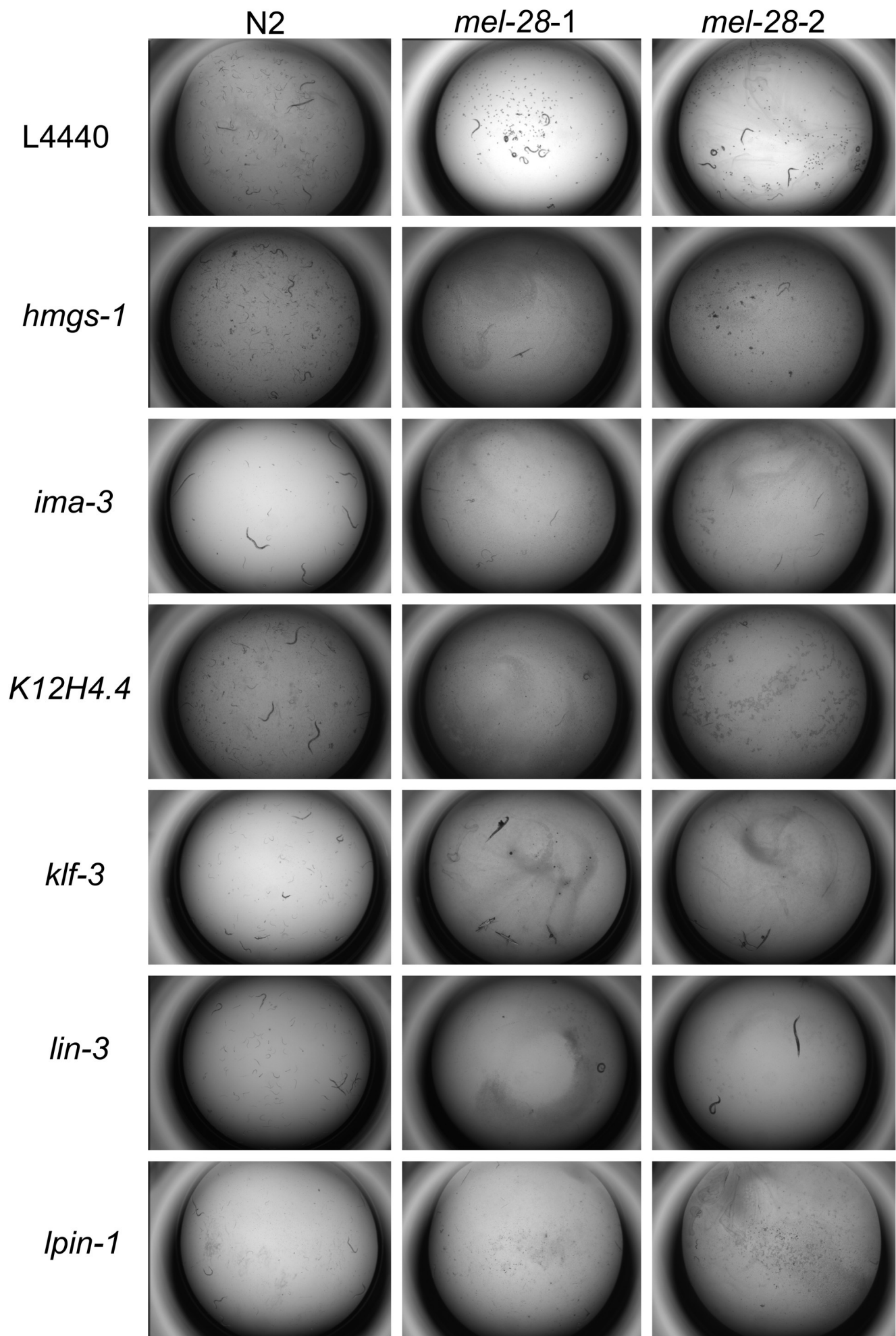
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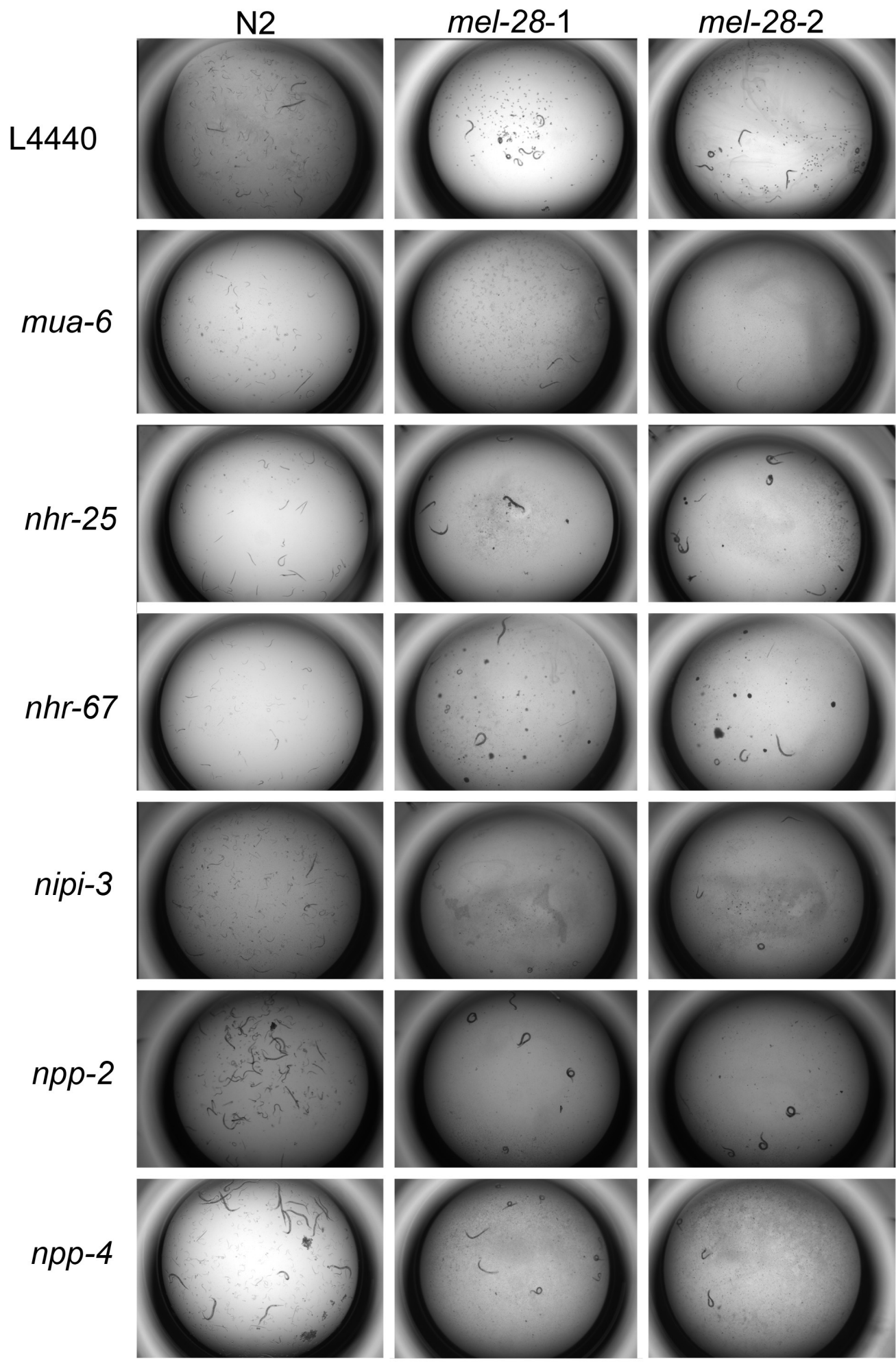


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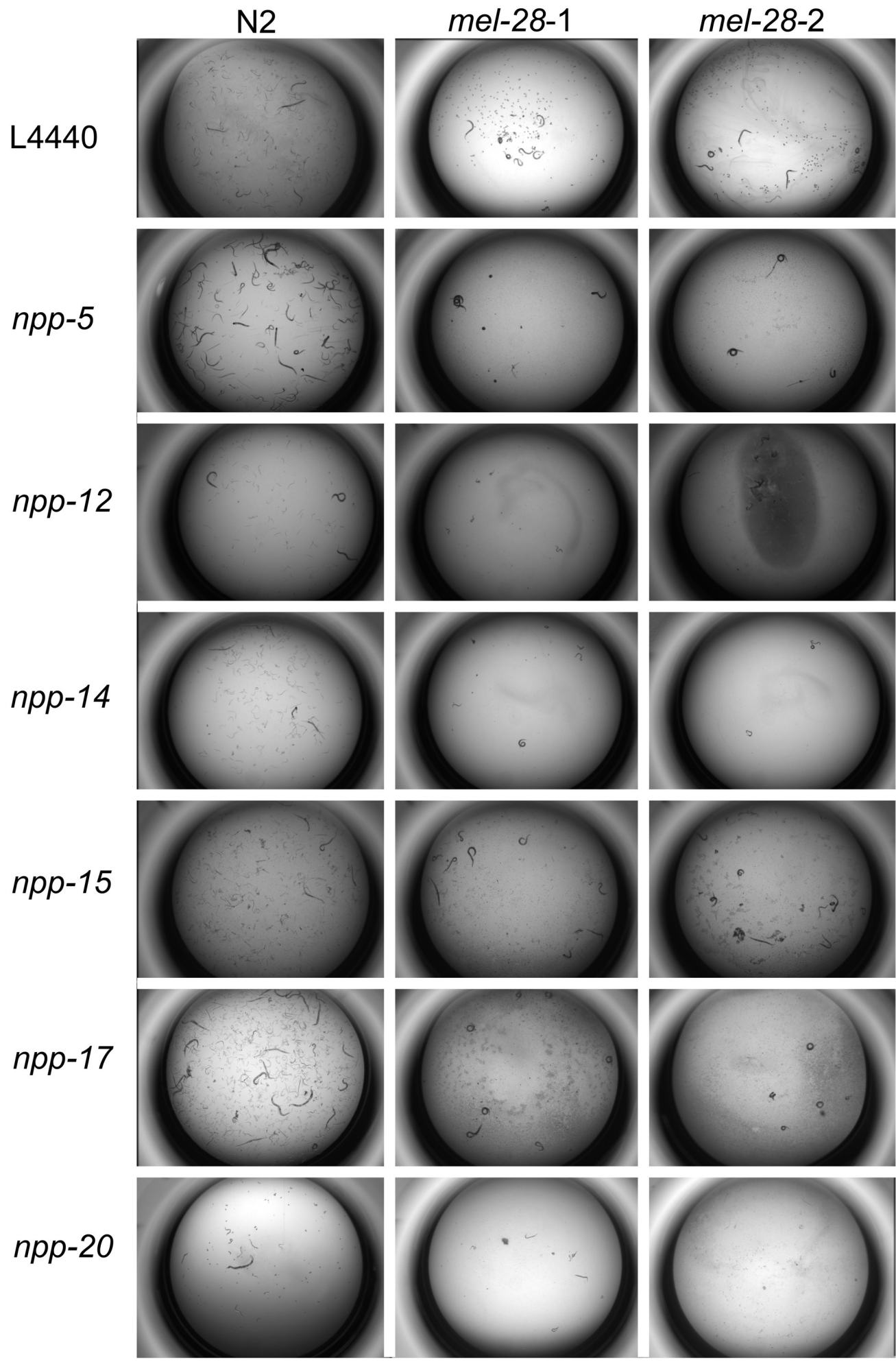


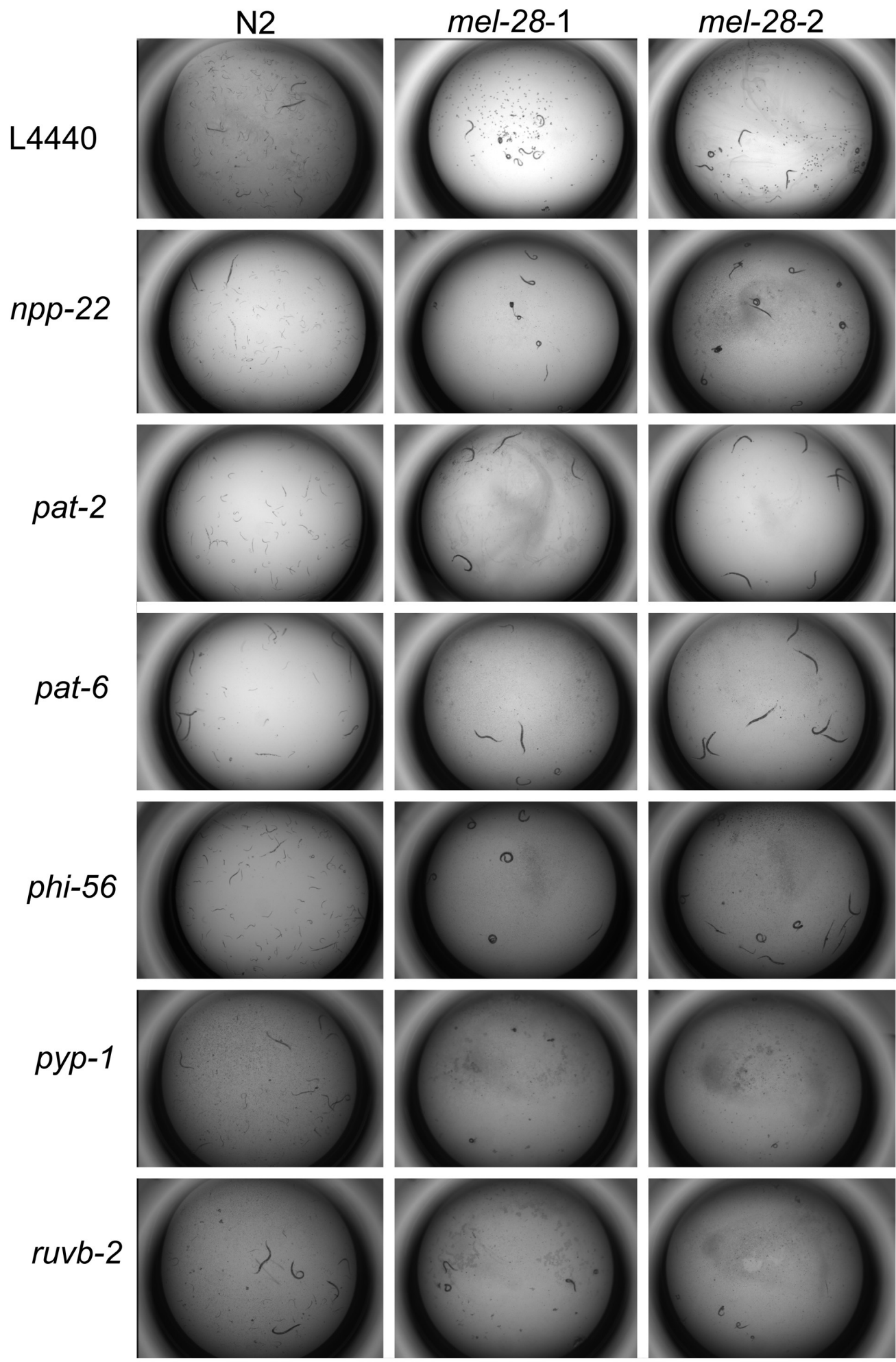


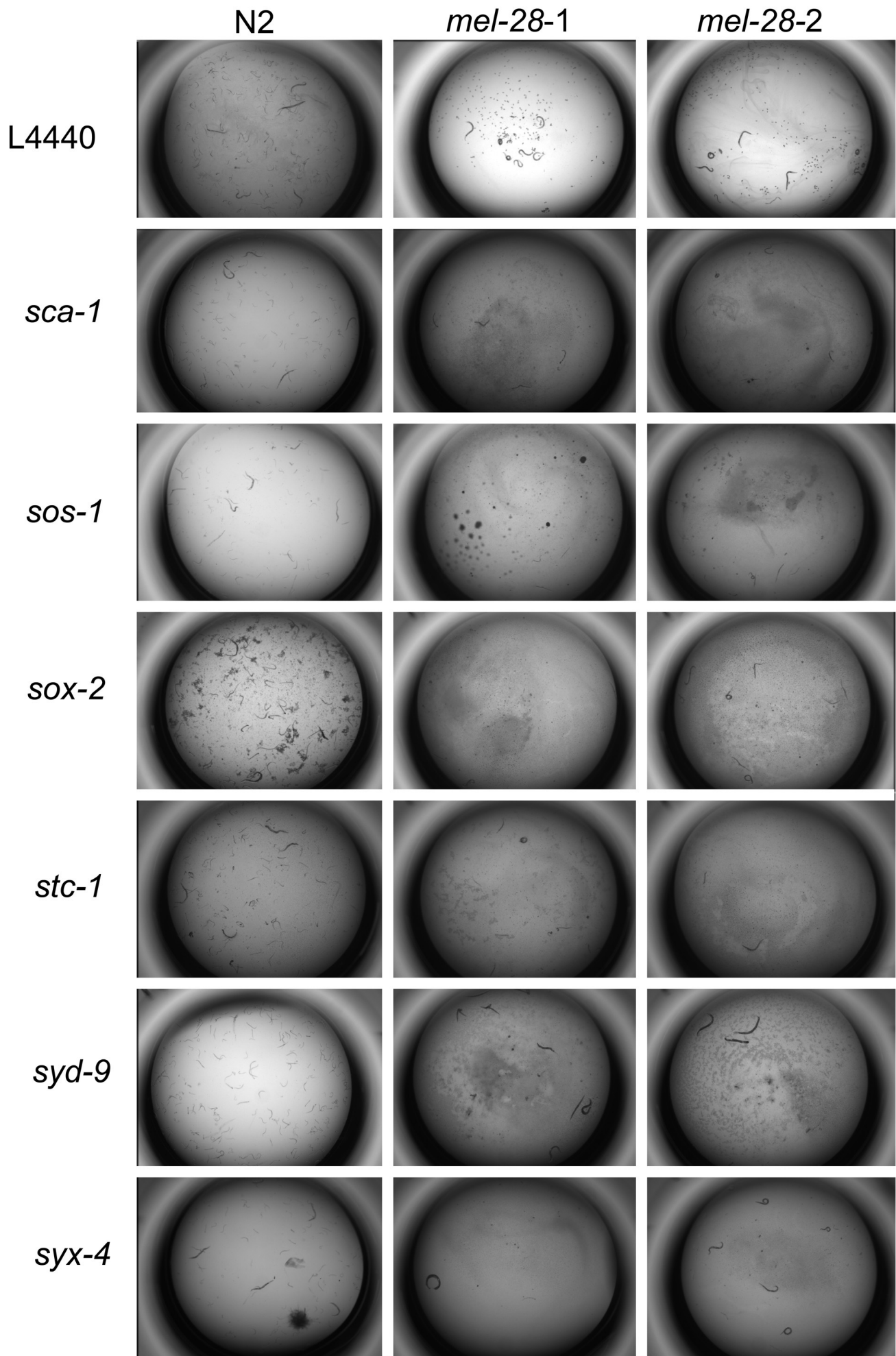












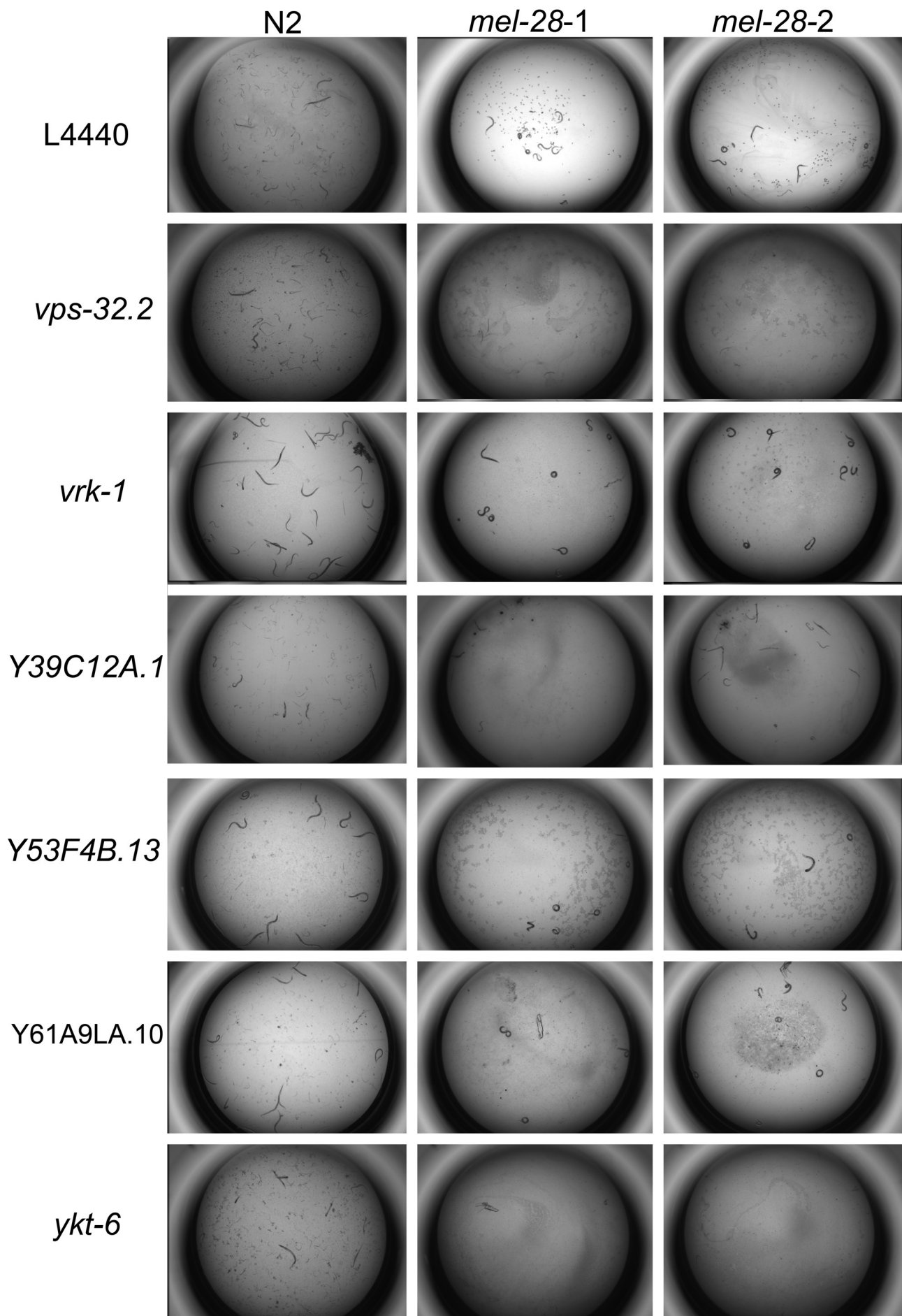


Figure S1 *mel-28* genetic interactor phenotypes

**Table S1 Brood size data for *mel-28* genetic interactors**

gene	progeny/a dult (N2)	eggs/ adult ( <i>mel-28</i> )	ratio N2 brood size: <i>mel-28</i> brood size	genetic interaction
<i>pat-2</i>	25.3	0.8	31.6	synthetic sterility
<i>pat-6</i>	7.3	0.8	9.1	enhancement
<i>klf-3</i>	13.9	0.0	>3	synthetic sterility
<i>his-67</i>	9.9	2.7	3.7	enhancement
<i>his-68</i>	10.2	0.0	>3	enhancement
<i>pyp-1</i>	2.8	0.0	>3	enhancement
<i>ruvb-2</i>	9.5	0.7	13.6	enhancement
<i>dhc-1</i>	1.7	0.0	>3	enhancement
<i>dyci-1</i>	4	0.0	>3	enhancement
<i>dli-1</i>	16	0.4	40	synthetic sterility
<i>dnc-1</i>	14.1	0.6	23.5	synthetic sterility
<i>cap-1</i>	12	0.6	20	enhancement
<i>cap-2</i>	8.4	1.4	6	enhancement
<i>arp-1</i>	16.7	1.8	9.3	synthetic sterility
<i>npp-2</i>	8.7	0.7	12.4	enhancement
<i>npp-4</i>	14.7	0.0	>3	synthetic sterility
<i>npp-5</i>	18.2	1.8	10.1	synthetic sterility
<i>npp-12</i>	21.3	0.3	71	synthetic sterility
<i>npp-14</i>	16.2	0.0	>3	synthetic sterility
<i>npp-15</i>	16	0.0	>3	synthetic sterility
<i>npp-17</i>	12.3	0.3	41	enhancement
<i>npp-20</i>	8.5	0.0	>3	enhancement
<i>npp-22</i>	25	0.0	>3	synthetic sterility
<i>ima-3</i>	6	0.0	>3	enhancement
<i>vrk-1</i>	4	1.1	3.6	enhancement
<i>lpin-1</i>	5	0.0	>3	enhancement
<i>cct-2</i>	17.3	0.2	86.5	synthetic sterility
<i>stc-1</i>	13.6	1.3	10.5	synthetic sterility
<i>sca-1</i>	13.2	0.0	>3	synthetic sterility
<i>phi-56</i>	12.3	0.0	>3	enhancement
<i>K12H4.4</i>	11.8	0.0	>3	enhancement
<i>ggtb-1</i>	2.6	0.0	>3	enhancement
<i>ykt-6</i>	11.7	1.0	11.7	enhancement
<i>syx-4</i>	24.3	0.4	60.8	synthetic sterility
<i>mua-6</i>	18.5	0.7	26.4	synthetic sterility
<i>syd-9</i>	55	0.2	275	synthetic sterility

<i>arf-3</i>	4.2	0.0	>3	enhancement
<i>hgrs-1</i>	17.8	0.0	>3	synthetic sterility
<i>vps-32.2</i>	11.3	0.0	>3	enhancement
Y61A9LA.10	9.4	0.0	>3	enhancement
<i>eif-1</i>	3.7	0.0	>3	enhancement
Y39C12A.1	20.8	0.7	29.7	synthetic sterility
F52C6.2	16	1.8	8.9	synthetic sterility
F52C6.3	14.7	0.0	>3	synthetic sterility
<i>exos-3</i>	11.2	3.0	3.7	enhancement
<i>alg-1</i>	4.9	0.0	>3	enhancement
<i>cyd-1</i>	27.8	0.0	>3	synthetic sterility
<i>ego-2</i>	15	0.6	25	synthetic sterility
<i>dre-1</i>	9.3	0.0	>3	enhancement
<i>nipi-3</i>	8.1	1.0	8.1	enhancement
<i>hmgs-1</i>	15.7	0.0	>3	synthetic sterility
<i>nhr-25</i>	17.1	0.0	>3	synthetic sterility
<i>nhr-67</i>	23.3	0.0	>3	synthetic sterility
<i>apl-1</i>	32	0.0	>3	synthetic sterility
F19B6.1	15.6	0.0	>3	synthetic sterility
C55A6.9	6.8	0.2	34	enhancement
<i>egl-13</i>	25	0.0	>3	synthetic sterility
<i>sox-2</i>	17	0.0	>3	synthetic sterility
<i>gei-13</i>	15	0.4	37.5	synthetic sterility
Y53F4B.13	3.8	0.0	>3	enhancement
<i>lin-3</i>	26.7	0.7	38.1	synthetic sterility
<i>arx-2</i>	27.2	0.0	>3	synthetic sterility
<i>eat-6</i>	10.9	0.0	>3	enhancement
B0250.7	3.8	1.0	3.8	enhancement
<i>sos-1</i>	7.8	0.0	>3	enhancement

For each strain (N2 or *mel-28*) the number of progeny per adult was tallied. We did this by adding up the total number of mothers and dividing this number by the total number of progeny from at least two RNAi experiments. For the *dhc-1* gene these tallies came from a single RNAi experiment (see methods for details).

**Table S2 GO-enriched terms amongst *mel-28* genetic interactors**

N	P value	GO attribute ID	GO attribute name
2	2.04E-05	GO:0008290	F-actin capping protein complex
2	6.11E-05	GO:0005787	signal peptidase complex
3	1.06E-05	GO:0005869	dynactin complex
7	2.39E-11	GO:0005643	nuclear pore
3	1.92E-05	GO:0051028	mRNA transport
7	6.97E-11	GO:0046930	pore complex
10	5.02E-13	GO:0005635	nuclear envelope
5	3.34E-06	GO:0006997	nucleus organization
5	2.25E-05	GO:0007338	single fertilization
5	2.73E-05	GO:0009566	fertilization
10	2.10E-08	GO:0031967	organelle envelope
10	2.20E-08	GO:0031975	envelope
12	2.05E-09	GO:0012505	endomembrane system
7	1.07E-05	GO:0051656	establishment of organelle localization
7	1.30E-05	GO:0051640	organelle localization
7	2.82E-05	GO:0032940	secretion by cell
7	3.11E-05	GO:0046903	secretion
8	2.83E-05	GO:0015031	protein transport
11	1.16E-06	GO:0044428	nuclear part
8	3.15E-05	GO:0045184	establishment of protein localization
14	6.34E-08	GO:0051649	establishment of localization in cell
14	1.62E-07	GO:0051641	cellular localization
21	7.91E-10	GO:0043234	protein complex
20	2.37E-09	GO:0006898	receptor-mediated endocytosis
11	8.38E-06	GO:0008104	protein localization
40	9.36E-12	GO:0040007	growth
20	2.91E-08	GO:0006897	endocytosis
21	3.49E-08	GO:0016192	vesicle-mediated transport
42	7.57E-11	GO:0009792	embryo development ending in birth or egg hatching
23	1.75E-08	GO:0044422	organelle part
21	6.33E-08	GO:0044446	intracellular organelle part
21	6.69E-08	GO:0032991	macromolecular complex
42	1.80E-10	GO:0009790	embryo development
33	6.12E-09	GO:0002119	nematode larval development
33	6.24E-09	GO:0002164	larval development
33	7.71E-09	GO:0009791	post-embryonic development

31	1.40E-08	GO:0051234	establishment of localization
30	2.70E-08	GO:0006810	transport
44	3.85E-08	GO:0048856	anatomical structure development
46	7.61E-08	GO:0007275	multicellular organismal development
45	1.52E-07	GO:0044767	single-organism developmental process
24	2.20E-06	GO:0040010	positive regulation of growth rate
24	2.26E-06	GO:0040009	regulation of growth rate
34	2.33E-07	GO:0051179	localization
26	1.56E-06	GO:0040011	locomotion
46	5.04E-07	GO:0032502	developmental process
46	1.46E-06	GO:0044707	single-multicellular organism process
25	6.07E-06	GO:0045927	positive regulation of growth
46	3.83E-06	GO:0032501	multicellular organismal process
19	7.29E-05	GO:0005515	protein binding
25	1.35E-05	GO:0040008	regulation of growth
27	1.21E-05	GO:0048518	positive regulation of biological process
33	5.12E-06	GO:0000003	reproduction
30	5.50E-05	GO:0043229	intracellular organelle
30	5.63E-05	GO:0043226	organelle

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We used FuncAssociate ([http://llama.mshri.on.ca/funcassociate\\_client/html/](http://llama.mshri.on.ca/funcassociate_client/html/)) to determine the GO term enrichment within the list of 65 *mel-28* genetic interactors identified. The background set was the entire *C. elegans* genome.