

Fig S2**3 months
NWD1 v AIN****12 months
NWD1 v AIN****12 month
cross over back
to AIN**

<u>Pathway</u>	NES	P	FDR	NES	P	FDR	NES	P	FDR
RIBOSOME	3.19	≤0.001	0.000	3.10	≤0.001	0.000	-3.13	≤0.001	0.000
DNA REPLICATION	2.41	≤0.001	0.000	2.26	≤0.001	0.000	-2.47	≤0.001	0.000
SPLICEOSOME	2.34	≤0.001	0.000	2.08	≤0.001	0.000	-2.34	≤0.001	0.000
RNA_POLYMERASE	2.19	≤0.001	0.000	1.88	≤0.001	0.001	-2.00	≤0.001	0.001
MISMATCH_REPAIR	2.13	≤0.001	0.000	2.14	≤0.001	0.000	-2.21	≤0.001	0.000
CELL_CYCLE	1.97	≤0.001	0.001	1.91	≤0.001	0.001	-2.08	≤0.001	0.000
NUCLEOTIDE EXCISION REPAIR	1.83	≤0.001	0.006	1.90	≤0.001	0.001	-1.78	0.003	0.009
PURINE METABOLISM	1.74	≤0.001	0.015	1.60	≤0.001	0.038	-1.54	0.002	0.079
OOCYTE MEIOSIS	1.73	0.002	0.016	1.42	0.030	0.160	-1.26	0.091	0.333
PYRIMIDINE METABOLISM	1.72	≤0.001	0.015	1.80	≤0.001	0.003	-1.67	≤0.001	0.026
BASE EXCISION REPAIR	1.68	0.004	0.020	2.09	≤0.001	0.000	-1.99	≤0.001	0.001
RNA DEGRADATION	1.68	≤0.001	0.019	1.63	0.006	0.031	-1.36	0.078	0.213
LINOLEIC ACID METABOLISM	1.66	0.004	0.020	1.46	0.062	0.153			
ARACHIDONIC ACID METABOLISM	1.59	0.015	0.038						
WNT SIGNALING PATHWAY	1.52	≤0.001	0.063						

<u>Pathway</u>	NES	P	FDR	NES	P	FDR	NES	P	FDR
ABC TRANSPORTERS	-2.34	≤0.001	≤0.001	-1.98	≤0.001	0.006	1.85	≤0.001	0.010
CITRATE CYCLE -TCA CYCLE	-2.28	≤0.001	≤0.001	-2.02	≤0.001	0.006	2.01	≤0.001	0.002
PEROXISOME	-2.20	≤0.001	≤0.001		≤0.001			≤0.001	≤0.001
OXIDATIVE PHOSPHORYLATION	-2.19	≤0.001	≤0.001	-2.37	≤0.001	≤0.001	2.11	≤0.001	0.001
PARKINSONS DISEASE	-2.14	≤0.001	≤0.001	-2.12	≤0.001	0.002	1.93	≤0.001	0.004
PPAR SIGNALING PATHWAY	-2.10	≤0.001	≤0.001		≤0.001				
FATTY ACID METABOLISM	-2.08	≤0.001	≤0.001		≤0.001				
GLYCOLYSIS GLUCONEOGENESIS	-2.07	≤0.001	≤0.001	-2.03	≤0.001	0.005			
RETINOL METABOLISM	-2.07	≤0.001	≤0.001		≤0.001				
ALZHEIMERS DISEASE	-2.02	≤0.001	≤0.001	-2.17	≤0.001	≤0.001	1.95	≤0.001	0.004
FRUCTOSE/MANNOSE METABOLISM	-1.98	≤0.001	≤0.001	-2.32	≤0.001	≤0.001	2.28	≤0.001	≤0.001
AMINO SUGAR/NUCLEOTIDE SUGAR METAB.	-1.92	0.002	0.002				1.72	0.010	0.023
GLYOXYLATE/DICARBOXYLATE METAB	-1.85	0.006	0.006						
SPHINGOLIPID METABOLISM	-1.84	≤0.001	0.006				1.89	0.005	0.008
BUTANOATE METABOLISM	-1.82	0.002	0.006						
OTHER GLYCAN DEGRADATION	-1.80	0.008	0.008	-1.53	0.058	0.133	2.12	≤0.001	≤0.001
PENTOSE PHOSPHATE PATHWAY	-1.78	0.004	0.010	-1.48	0.033	0.138			