

Figure # (IMC data were located at the bottom)	Group	Sample size (n)	Shapiro-Wilk normality test P value	Shapiro-Wilk normality test Passed	Applied statistical test(s) and results	P (or Adjusted)Value
1B: p-p90RSK 10 min	WT	3 (time course)	0.9999	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.991
	ERKS496A.KI	3 (time course)	0.3631	Yes		
1B: p-p90RSK 30 min	WT	3 (time course)	0.9977	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.938
	ERKS496A.KI	3 (time course)	0.5368	Yes		
1B: p-ERKS496 10 min	WT	3 (time course)	0.2103	No	One-way ANOVA non parametric Kruskal-Wallis test followed by Dunn post hoc multiple comparison tests	0.7503
	ERKS496A.KI	3 (time course)	0.1436	Yes		
1B: p-ERKS496 30 min	WT	3 (time course)	0.0365	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0004
	ERKS496A.KI	3 (time course)	0.0561	Yes		
1B: p-ERKSTEY 10 min	WT	3 (time course)	0.842	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.9977
	ERKS496A.KI	3 (time course)	0.1025	Yes		
1B: p-ERKSTEY 30 min	WT	3 (time course)	0.0048	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.9991
	ERKS496A.KI	3 (time course)	0.2476	Yes		
1D	WT	15	0.12	Yes	Unpaired Student's t-test, two-tailed	0.0362
	ERKS496A.KI	21	0.745	Yes		
1E	WT	8	0.7681	Yes	Unpaired Student's t-test, two-tailed	0.0444
	ERKS496A.KI	10	0.3932	Yes		
1F	WT	8	0.6041	Yes	Unpaired Student's t-test, two-tailed	0.0079
	ERKS496A.KI	9	0.171	Yes		
1G: Vehicle	CA.MEKSa	5	0.6909	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0001
	CA.MEKSa+oxLDL	5	0.814	Yes		
	CA.MEKSa+Vehicle	5	0.814	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0001
1H: WT	CA.MEKSa	5	0.1633	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	CA.MEKSa+MEA	5	0.1538	Yes		
1H: WT	CA.MEKSa+oxLDL	5	0.461	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	CA.MEKSa+WT	5	0.474	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00026
1J: oxLDL	CA.MEKSa+oxLDL+ERKS496A.KI	5	0.3254	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000003
	Vehicle	5	0.3254	Yes		
1J: TRX1, WT	oxLDL	5	0.8075	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0003
	WT	5	0.035	Yes		
1J: TRX1, oxLDL	ERKS496A.KI	5	0.6972	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00032
	Vehicle	5	0.3254	Yes		
1J: HO1, WT	oxLDL	5	0.2005	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000178
	WT	5	0.0265	Yes		
1J: HO1, oxLDL	ERKS496A.KI	5	0.4597	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000954
	Vehicle	5	0.3254	Yes		
1J: CD36, WT	oxLDL	5	0.6773	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000008
	WT	5	0.039	Yes		
1J: CD36, oxLDL	ERKS496A.KI	5	0.3857	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.014
	Vehicle	5	0.3254	Yes		
1J: p53, WT	oxLDL	5	0.0995	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00138
	WT	5	0.0295	Yes		
1J: p53, oxLDL	ERKS496A.KI	5	0.2527	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.01803
	Vehicle	5	0.3254	Yes		
1J: p21, WT	oxLDL	5	0.0942	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT	5	0.0642	Yes		
1J: p21, oxLDL	ERKS496A.KI	5	0.5084	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	Vehicle	5	0.3254	Yes		
1J: p16, WT	oxLDL	5	0.0562	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000003
	WT	5	0.6642	Yes		
1J: p16, oxLDL	ERKS496A.KI	5	0.5084	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000032
	Vehicle	5	0.3254	Yes		
1J: GAS6, WT	oxLDL	5	0.0205	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00008
	WT	5	0.3025	Yes		
1J: GAS6, oxLDL	ERKS496A.KI	5	0.1909	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00089
	Vehicle	5	0.3254	Yes		
1J: TNFa, WT	oxLDL	5	0.0255	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00046
	WT	5	0.0605	Yes		
1J: TNFa, oxLDL	ERKS496A.KI	5	0.9983	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00095
	Vehicle	5	0.3254	Yes		
1J: DNMT3a, WT	oxLDL	5	0.0224	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00142
	WT	5	0.0589	Yes		
1J: DNMT3a, oxLDL	ERKS496A.KI	5	0.315	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00215
	Vehicle	5	0.2957	Yes		
1K: WT	oxLDL	8	0.5409	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT	8	0.5409	Yes		
1K: oxLDL	ERKS496A.KI	8	0.926	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	Vehicle	8	0.6729	Yes		
1L: WT	oxLDL	8	0.6878	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT	8	0.6878	Yes		
1L: oxLDL	ERKS496A.KI	8	0.5478	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	Vehicle	8	0.3117	Yes		
1M: WT	oxLDL	8	0.4827	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0052
	WT	8	0.4627	Yes		
1M: oxLDL	ERKS496A.KI	8	0.0241	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.012
	Vehicle	8	0.0637	Yes		
1N: WT	oxLDL	8	0.631	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT	8	0.3258	Yes		
1N: oxLDL	ERKS496A.KI	8	0.264	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000009
	Vehicle	8	0.216	Yes		
1O: oxLDL	ERKS496A.KI	8	0.216	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT	8	0.216	Yes		
1O: oxLDL	ERKS496A.KI	8	0.0245	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT	8	0.4827	Yes		
3C	WT	1,000 (sampling by base R package)	4.98E-14	No	Wilcoxon signed-rank test	0.1966
	ERKS496A	1,000 (sampling by base R package)	4.92E-14	No		
3D	WT	1,000 (sampling by base R package)	2.16E-05	No	Wilcoxon signed-rank test	2.20E-16
	ERKS496A	1,000 (sampling by base R package)	1.76E-05	No		
3E	WT	1,000 (sampling by base R package)	2.32E-04	No	Wilcoxon signed-rank test	2.20E-16
	ERKS496A.KI	1,000 (sampling by base R package)	1.60E-03	No		
3G: WT	NCD	8	0.2914	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0017
	HFD	8	0.1052	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.024
3H: WT	NCD	8	0.0263	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00000001
	HFD	8	0.1053	Yes		
3H: HFD	ERKS496A.KI	8	0.8624	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00000001
	Vehicle	8	0.216	Yes		
3I: WT	HFD	8	0.216	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT	8	0.216	Yes		
3I: HFD	ERKS496A.KI	8	0.0522	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00056
	Vehicle	8	0.3338	Yes		
3J: HFD	ERKS496A.KI	8	0.3339	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00014
	WT	8	0.1363	Yes		
3K: WT	HFD	8	0.1052	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00000001
	WT	8	0.2452	Yes		
3K: HFD	ERKS496A.KI	8	0.2452	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	2.3420E-08
	Vehicle	8	0.6159	Yes		
3M	WT	13	0.732	Yes	Unpaired Student's t-test, two-tailed	0.0001
	ERKS496A.KI	13	0.4794	Yes		
3O	WT	14	0.0069	Yes	Non parametric unpaired Mann-Whitney test	0.000001
	ERKS496A.KI	13	0.1924	Yes		
4K	siAHR	6	0.9362	Yes	Unpaired Student's t-test, two-tailed	0.0002
	Control	8	0.0929	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
5K: oxLDL	WT	8	0.0928	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000006
	ERKS496A	8	0.3233	Yes		
5K: oxLDL	WT (control)	8	0.0928	Yes	One-way ANOVA non parametric Kruskal-Wallis test followed by Dunn post hoc multiple comparison tests	0.3938
	WT (control)	8	0.0014	No		
5K: oxLDL	AKT1S36	8	0.0828	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00005
	Vehicle	8	0.2674	Yes		
6A: WT	oxLDL	8	0.7491	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000042
	WT	8	0.8754	Yes		
6A: oxLDL	ERKS496A.KI	8	0.2709	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00003
	control	8	0.1588	Yes		
6A: WT (mid panel)	oxLDL	8	0.1433	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0001
	Control	8	0.2119	Yes		
6A: vehicle	oxLDL	8	0.4617	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0001
	vehicle	8	0.0417	Yes		
6A: ox-LDL	AKT1S36	8	0.0552	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.002
	Vehicle	8	0.3254	Yes		
6C: WT	oxLDL	8	0.9913	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000007
	WT	8	0.0303	Yes		
6C: oxLDL	ERKS496A.KI	8	0.1041	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000078
	Vehicle	8	0.3254	Yes		
6D: WT	oxLDL	5	0.519	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT	5	0.0569	Yes		
6D: oxLDL	NEFZ2K51R	5	0.3104	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000002
	Vehicle	8	0.1058	Yes		
6E: WT	oxLDL	5	0.0952	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0016
	WT	5	0.0562	Yes		
6E: oxLDL	NEFZ2K51R	5	0.1177	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.036
	Vehicle	8	0.3076	Yes		
6F: WT	oxLDL	5	0.0962	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	2.1056E-08
	WT	5	0.0517	Yes		
6F: oxLDL	NEFZ2K51R	5	0.0517	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	1.5680E-06
	Vehicle	8	0.9544	Yes		
6G: WT	oxLDL	5	0.0459	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT	5	0.2236	Yes		
6G: oxLDL	NEFZ2K51R	5	0.2234	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00012
	Vehicle	8	0.6485	Yes		
6H: WT	oxLDL	5	0.0749	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0001
	WT	5	0.3785	Yes		
6H: oxLDL	NEFZ2K51R	5	0.3658	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0001
	Vehicle	8	0.3571	Yes		
6I: WT	oxLDL	5	0.11	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.012
	WT	8	0.1236	Yes		
6I: oxLDL	NEFZ2K51R	5	0.0751	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.041
	Vehicle	8	0.2435	Yes		
7B: Maximum respiration	WT+oxLDL	5	0.2345	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT+oxLDL	5	0.2345	Yes		
7B: Maximum respiration	ERKS496A+oxLDL	5	0.6263	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT+oxLDL	5	0.5678	Yes		
7B: SRC	WT+oxLDL	5	0.5678	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT	4 (Seahorse)	0.85	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0471
7D: Glycolysis	WT+oxLDL	4 (Seahorse)	0.0643	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0008
	WT	4 (Seahorse)	0.2942	Yes		
7D: Glycolysis	ERKS496A+oxLDL	4 (Seahorse)	0.3315	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.003
7D: Glycolytic capacity	WT+oxLDL	4 (Seahorse)	0.1612	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0008

7D: Glycolytic capacity	WT+oxLDL	4(Seahorse)	0.1812	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	ERK5+S496A+oxLDL	4(Seahorse)	0.1825	Yes		
7E: WT	Vehicle	8	0.0837	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0013
	oxLDL	8	0.3917	Yes		
7F: oxLDL	ERK5 S496A KI	8	0.0817	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.027
	Vehicle	8	0.5002	Yes		
7F: WT	oxLDL	8	0.3216	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000108
	Vehicle	8	0.1538	Yes		
7F: oxLDL	ERK5 S496A KI	8	0.1534	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000032
7H: Maximum respiration	NRF2WT+oxLDL	5(Seahorse)	0.3497	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	NRF2WT+oxLDL	5(Seahorse)	0.2022	Yes		
7H: Maximum respiration	NRF2K518R+oxLDL	5(Seahorse)	0.0202	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	NRF2WT+oxLDL	5(Seahorse)	0.1831	Yes		
7H: SRC	NRF2WT+oxLDL	5(Seahorse)	0.7558	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.01
	NRF2WT+oxLDL	5(Seahorse)	0.2162	Yes		
7H: SRC	NRF2K518R+oxLDL	5(Seahorse)	0.0202	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0012
	NRF2K518R+oxLDL	5(Seahorse)	0.9554	Yes		
7J: Non-glycolytic acidification	NRF2WT	4(Seahorse)	0.3408	Yes	One-way ANOVA Tukey post hoc multiple comparison test	0.0084
	NRF2WT	4(Seahorse)	0.9813	Yes		
7J: Non-glycolytic acidification	NRF2K518R	4(Seahorse)	0.0202	Yes	One-way ANOVA Tukey post hoc multiple comparison test	0.0048
	NRF2K518R	4(Seahorse)	0.2679	Yes		
7K: NRF2WT	Vehicle	8	0.11	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	oxLDL	8	0.051	Yes		
7K: oxLDL	NRF2K518R	8	0.1531	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.015
	Vehicle	8	0.259	Yes		
7L: NRF2WT	Vehicle	8	0.54	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	6.2314E-11
	oxLDL	8	0.4844	Yes		
7L: oxLDL	NRF2K518R	8	0.0662	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	2.9228E-08
S1B: Baseline	WT	13	0.4918	Yes	Unpaired Student's t-test, two-tailed	0.1848
	ERK5 S496A KI	26	0.0205	Yes		
S1B: Time of sacrifice	WT	13	0.7965	Yes	Unpaired Student's t-test, two-tailed	0.0743
	ERK5 S496A KI	26	0.2748	Yes		
S1C: HDL	ERK5 S496A KI	11	0.272	Yes	Unpaired Student's t-test, two-tailed	0.3637
	WT	11	0.1749	Yes		
S1C: LDL	ERK5 S496A KI	19	0.5809	Yes	Unpaired Student's t-test, two-tailed	0.083
	WT	7	0.2884	Yes		
S1D	ERK5 S496A KI	7	0.1458	Yes	Unpaired Student's t-test, two-tailed	0.4344
	WT	7	0.1238	Yes		
S1E	ERK5 S496A KI	7	0.6839	Yes	Unpaired Student's t-test, two-tailed	0.279
	WT	7	0.541	Yes		
S1F: WT	Vehicle	8	0.549273	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	9.3913E-09
	oxLDL	8	0.1549	Yes		
S1F: oxLDL	WT	8	0.146479	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	1.46207E-05
	ERK5 S496A KI	8	0.166912	Yes		
S1G: Vehicle	Vehicle	8	0.1659	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	3.17047E-09
	oxLDL	8	0.607	Yes		
S1G: oxLDL	Vehicle	8	0.807	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	8.15884E-08
	MitoTEMPOL	8	0.4209	Yes		
S1H: WT	WT	8	0.0205	Yes	One-way ANOVA Tukey post hoc multiple comparison test followed by Dunn post hoc multiple comparison tests	2.40571E-05
	HFD	8	0.541	Yes		
S1H: HFD	WT	8	0.541	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	3.73288E-10
	ERK5 S496A	5	0.21	Yes		
S1I: oxLDL	Vehicle	5	0.7487	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	4.71E-13
	ERK5 S496A	5	0.3633	Yes		
S1I: oxLDL	XMD8-32	5	0.1281	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	4.71E-13
	Vehicle	5	0.1533	Yes		
S1I: oxLDL	AX15836	5	0.8493	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	4.71E-13
	WT	6	0.1259	Yes		
S2A	ERK5 S496A KI	5	0.0205	Yes	Unpaired Student's t-test, two-tailed	0.03
	WT	6	0.8407	Yes		
S2B	ERK5 S496A KI	5	0.502	Yes	Unpaired Student's t-test, two-tailed	0.0194
	WT	5	0.9414	Yes		
S2C	ERK5 S496A KI	7	0.721	Yes	Unpaired Student's t-test, two-tailed	0.0399
	Vehicle	7	0.1632	Yes		
S2F: WT	Vehicle	8	0.1632	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.001
	oxLDL	8	0.0927	Yes		
S2F: oxLDL	WT	8	0.1638	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0092
	ERK5 S496A KI	5	0.3254	Yes		
S3-p:p90RSK WT	NCD	5	0.0841	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	HFD	5	0.3254	Yes		
S3-p:ERK5S496 WT	WT	5	0.2257	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	ERK5 S496A KI	5	0.0815	Yes		
S3-p:ERK5S496 HFD	WT	5	0.1937	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000002
	ERK5 S496A KI	5	0.3486	Yes		
S3-H01 WT	NCD	5	0.0205	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000706
	HFD	5	0.1763	Yes		
S3-H01 HFD	WT	5	0.1763	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.018122
	ERK5 S496A KI	5	0.2383	Yes		
S3-CD36 WT	WT	5	0.0205	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000102
	HFD	5	0.1111	Yes		
S3-CD36 HFD	WT	5	0.1111	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000378
	ERK5 S496A KI	5	0.4816	Yes		
S3-p21 WT	WT	5	0.0205	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000009
	HFD	5	0.471	Yes		
S3-p21 HFD	WT	5	0.471	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000273
	ERK5 S496A KI	5	0.0205	Yes		
S3-p16 WT	WT	5	0.9573	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000002
	HFD	5	0.9573	Yes		
S3-p16 HFD	WT	5	0.9573	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	ERK5 S496A KI	5	0.3254	Yes		
S3-p53 WT	NCD	5	0.0205	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000866
	HFD	5	0.1613	Yes		
S3-p53 HFD	WT	5	0.1613	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.018064
	ERK5 S496A KI	5	0.2059	Yes		
S3-GAS6 WT	NCD	5	0.3254	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.024662
	HFD	5	0.3226	Yes		
S3-GAS6 HFD	WT	5	0.3226	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.001363
	ERK5 S496A KI	5	0.0205	Yes		
S3-TNF α WT	NCD	5	0.3254	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000021
	HFD	5	0.091	Yes		
S3-TNF α HFD	WT	5	0.091	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000215
	ERK5 S496A KI	5	0.3254	Yes		
S3-DNMT3a WT	NCD	5	0.0558	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.005179
	HFD	5	0.1598	Yes		
S3-DNMT3a HFD	WT	5	0.1598	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000779
	ERK5 S496A KI	5	0.1185	Yes		
S3-AHR WT	HFD	5	0.8644	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.001048
	WT	5	0.5653	Yes		
S3-AHR HFD	ERK5 S496A KI	5	0.5653	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.026874
	NCD	5	0.3254	Yes		
S3:53BP1 WT	WT	5	0.9204	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	ERK5 S496A KI	5	0.3254	Yes		
S3:53BP1 HFD	WT	5	0.3254	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000005
	ERK5 S496A KI	5	0.2335	Yes		
S3:LAMINB1 WT	NCD	5	0.3254	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	HFD	5	0.4718	Yes		
S3:LAMINB1 HFD	WT	5	0.4718	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000012
	ERK5 S496A KI	5	0.431	Yes		
S3:PCNA WT	NCD	5	0.3254	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	HFD	5	0.9666	Yes		
S3:PCNA HFD	WT	5	0.8135	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000007
	ERK5 S496A KI	5	0.683	Yes		
S4A: IL-1b WT	NCD	4	0.2453	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000023
	HFD	4	0.3343	Yes		
S4A: IL-1b HFD	WT	4	0.1921	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.00014
	ERK5 S496A KI	4	0.683	Yes		
S4A: TNF α WT	NCD	4	0.2069	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0006
	HFD	4	0.9098	Yes		
S4A: TNF α HFD	WT	4	0.2929	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0021
	ERK5 S496A KI	4	0.683	Yes		
S4A: iNOS WT	NCD	4	0.2715	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000995
	HFD	4	0.2515	Yes		
S4A: iNOS HFD	WT	4	0.2152	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0015
	ERK5 S496A KI	4	0.391	Yes		
S4C: IL-1a	WT	3(Cytokine profile)	0.7307	Yes	One-way ANOVA Tukey post hoc multiple comparison test	0.000001
	ERK5 S496A KI/HFD	3(Cytokine profile)	0.3929	Yes		
S5B: Maximum respiration	WT+NCD	12(Seahorse from 5 independent experiments)	0.1685	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000015
	WT+HFD	12(Seahorse from 5 independent experiments)	0.429	Yes		
S5B: Maximum respiration	WT+HFD	12(Seahorse from 5 independent experiments)	0.429	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000075
	ERK5 S496A+HFD	12(Seahorse from 5 independent experiments)	0.1105	Yes		
S5B: SRC	WT+NCD	12(Seahorse from 5 independent experiments)	0.1816	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.007
	WT+HFD	12(Seahorse from 5 independent experiments)	0.7593	Yes		
S5B: SRC	WT+HFD	12(Seahorse from 5 independent experiments)	0.7593	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.0221
	ERK5 S496A+HFD	12(Seahorse from 5 independent experiments)	0.1517	Yes		
S5D: Glycolysis	WT+NCD	5(Seahorse)	0.8641	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT+HFD	5(Seahorse)	0.6487	Yes		
S5D: Glycolysis	WT+HFD	5(Seahorse)	0.6487	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	ERK5 S496A+HFD	5(Seahorse)	0.6292	Yes		
S5D: Glycolytic capacity	WT+HFD	5(Seahorse)	0.5927	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001
	WT+HFD	5(Seahorse)	0.5927	Yes		
S5D: Glycolytic capacity	ERK5 S496A+HFD	5(Seahorse)	0.9748	Yes	Two-way ANOVA Tukey post hoc multiple comparison test	0.000001

2M (IL-6 #1)	WT	1716	<0.0001	No	Non parametric unpaired Mann-Whitney test	<0.0000000000000001	Unpaired t test with Welch's correction <0.0000000000000001
	KI	1291	<0.0001	No			
2M (IL-6 #7)	WT	2417	<0.0001	No	Non parametric unpaired Mann-Whitney test	3.3687E-06	Unpaired t test with Welch's correction 0.0006890363
	KI	1858	<0.0001	No			
2M (IL-6 #9)	WT	2817	<0.0001	No	Non parametric unpaired Mann-Whitney test	<0.0000000000000001	Unpaired t test with Welch's correction 4.2E-14
	KI	2170	<0.0001	No			
S2D:TRX #5	WT	1775	<0.0001	No	Non parametric unpaired Mann-Whitney test	0.009122289	Unpaired t test with Welch's correction 0.00033139
	KI	1775	<0.0001	No			
S2D:TRX #6	WT	122	<0.0001	No	Non parametric unpaired Mann-Whitney test	0.001532638	Unpaired t test with Welch's correction 0.001272231
	KI	202	<0.0001	No			
S2D:DNNMT3a #5	WT	2536	<0.0001	No	Non parametric unpaired Mann-Whitney test	5.3E-13	Unpaired t test with Welch's correction 1.87804E-07
	KI	1775	<0.0001	No			
S2D:DNNMT3a #6	WT	122	<0.0001	No	Non parametric unpaired Mann-Whitney test	0.00107075	Unpaired t test with Welch's correction 0.000729619
	KI	202	<0.0001	No			
S2D:GAS6 #5	WT	2536	<0.0001	No	Non parametric unpaired Mann-Whitney test	8.99327E-07	Unpaired t test with Welch's correction 4.01757E-07
	KI	1775	<0.0001	No			
S2D:GAS6 #6	WT	122	<0.0001	No	Non parametric unpaired Mann-Whitney test	0.003054729	Unpaired t test with Welch's correction 0.001822599
	KI	202	<0.0001	No			
S2D:TYRO3 #6	WT	202	<0.0001	No	Non parametric unpaired Mann-Whitney test	2.21641E-05	Unpaired t test with Welch's correction 0.000739053
	KI	1775	<0.0001	No			
S2D:IL-6 #5	WT	2536	<0.0001	No	Non parametric unpaired Mann-Whitney test	1.71063E-10	Unpaired t test with Welch's correction 7.0827E-11
	KI	1775	<0.0001	No			
S2E:Distance #5#8	WT	122	<0.0001	No	Non parametric unpaired Mann-Whitney test	2.09186E-09	Unpaired t test with Welch's correction 0.000906126
	KI	1781	<0.0001	No			
S2H:p16 #7	WT	2417	<0.0001	No	Non parametric unpaired Mann-Whitney test	<0.0000000000000001	Unpaired t test with Welch's correction <0.0000000000000001
	KI	1858	<0.0001	No			
S2H:p16 #9	WT	2170	<0.0001	No	Non parametric unpaired Mann-Whitney test	<0.0000000000000001	Unpaired t test with Welch's correction <0.0000000000000001
	KI	2170	<0.0001	No			
S2H:p-PKC #1	WT	1716	<0.0001	No	Non parametric unpaired Mann-Whitney test	<0.0000000000000001	Unpaired t test with Welch's correction <0.0000000000000001
	KI	1291	<0.0001	No			
S2H:p-PKC #7	WT	1716	<0.0001	No	Non parametric unpaired Mann-Whitney test	6.22427E-08	Unpaired t test with Welch's correction 8.44638E-08
	KI	1858	<0.0001	No			
S2H:p-PKC #9	WT	2817	<0.0001	No	Non parametric unpaired Mann-Whitney test	<0.0000000000000001	Unpaired t test with Welch's correction <0.0000000000000001
	KI	2170	<0.0001	No			
S2K(Top2B)	ERK5 S496A KI	575	<0.0001	No	Non parametric unpaired Mann-Whitney test	0.0671	Unpaired t test with Welch's correction 0.0397
	WT	701	0.0071	No			
S2K(TRX)	ERK5 S496A KI	575	0.3646	Yes	Non parametric unpaired Mann-Whitney test	0.0872	Unpaired t test with Welch's correction 0.0233
	WT	701	0.011	No			
S2K(K67)	ERK5 S496A KI	575	<0.0001	No	Non parametric unpaired Mann-Whitney test	0.4144	Unpaired t test with Welch's correction 0.0686
	WT	701	<0.0001	No			
S2K(p53)	ERK5 S496A KI	575	<0.0001	No	Non parametric unpaired Mann-Whitney test	0.5338	Unpaired t test with Welch's correction 0.2765
	WT	701	<0.0001	No			
S2K(Tyro3)	ERK5 S496A KI	575	0.0003	No	Non parametric unpaired Mann-Whitney test	0.0206	Unpaired t test with Welch's correction 0.0139
	WT	701	<0.0001	No			
S2K(DNMT3a)	ERK5 S496A KI	575	<0.0001	No	Non parametric unpaired Mann-Whitney test	0.0065	Unpaired t test with Welch's correction 0.0022
	WT	701	<0.0001	No			
S2K(Gas6)	ERK5 S496A KI	575	0.0298	No	Non parametric unpaired Mann-Whitney test	3.11022E-05	Unpaired t test with Welch's correction 1.47754E-05
	WT	701	<0.0001	No			
S2K(IL-6)	ERK5 S496A KI	575	0.0001	No	Non parametric unpaired Mann-Whitney test	3.11183E-06	Unpaired t test with Welch's correction 0.0687
	WT	701	0.0001	No			
Figure #							
Group		Cell number	D'Agostino & Pearson test P value	D'Agostino & Pearson test P value	Applied statistical test(s) and results		P (or Adjusted)Value
S2G		p16	<0.0001	No	Non parametric Kruskal-Wallis test followed by Dunn post hoc multiple comparison tests		<0.0000000000000001
		p53	<0.0001	No			
S2G		p-TERF2IP	<0.0001	No	Non parametric Kruskal-Wallis test followed by Dunn post hoc multiple comparison tests		<0.0000000000000001
		p53	<0.0001	No			