

Table EV 1 Exact n and p values and statistical test used

| Figure | N | P value | Statistical analysis used |
|--------------------------|--------------------------------------|----------|---------------------------|
| 1A | 8 | 0.0042 | Two tailed t-test |
| 1D | 3 | 0.0102 | Two tailed t-test |
| 1E (ATP5a) | 3 | 0.0212 | Two tailed t-test |
| 1E (HSP60) | 3 | 0.0363 | Two tailed t-test |
| 1H | 3 | 0.043 | Two tailed t-test |
| 1I | 3 | 0.0226 | Two tailed t-test |
| 1K | 3 | 0.0481 | Two tailed t-test |
| 1L | 3 | 0.0062 | Two tailed t-test |
| 1N (Acidic pH) | Control= 6; IU1 =10 | < 0.0001 | Two tailed t-test |
| 1N (Acidic / neutral pH) | Control= 6; IU1 =10 | 0.0154 | Two tailed t-test |
| 2B (TOM20) | 3 | 0.0244 | one way ANOVA/Dunnett's |
| 2B (DRP1) | 4 | 0.0101 | one way ANOVA/Dunnett's |
| 2B (Fis1) | 3 | 0.0427 | one way ANOVA/Dunnett's |
| 2B (OPA1) | 3 | 0.212 | one way ANOVA/Dunnett's |
| 2B (Mfn1) | 4 | P<0.0001 | one way ANOVA/Dunnett's |
| 2B (Mfn2) | 4 | 0.0139 | one way ANOVA/Dunnett's |
| 2D | WT=4; WT+IU1=5 | 0.0427 | Two tailed t-test |
| 2D | DRP1KO=4; DRP1 KO+IU1=4 | 0.6734 | Two tailed t-test |
| 2D | Mfn1 KO=3; Mfn1 KO+IU1=4 | 0.0236 | Two tailed t-test |
| 2D | Mfn2 KO=4; Mfn2 KO+IU1=4 | 0.5473 | Two tailed t-test |
| 2F (HSP60) | WT=6; WT+IU1=6 | 0.0007 | Two tailed t-test |
| 2F (HSP60) | Mfn2 KO=3; Mfn2 KO+IU1=3 | 0.3484 | Two tailed t-test |
| 2F (HSP60) | WT+DRP1k38A=3; DRP1k38A+IU1=3 | 0.1229 | Two tailed t-test |
| 2F (HSP60) | DRP1 KO=3; DRP1 KO+IU1=3 | 0.1984 | Two tailed t-test |
| 2F (ATP5a) | WT=3; WT+IU1=3 | 0.0331 | Two tailed t-test |
| 2F (ATP5a) | Mfn2 KO=3; Mfn2 KO+IU1=3 | 0.8911 | Two tailed t-test |
| 2F (ATP5a) | WT+DRP1k38A=3; DRP1k38A+IU1=3 | 0.1764 | Two tailed t-test |
| 2F (ATP5a) | DRP1 KO=3; DRP1 KO+IU1=3 | 0.8137 | Two tailed t-test |
| 3D | WT MEF (+/- IU1)=3 | 0.0354 | Two tailed t-test |
| 3D | Hela (+/- IU1)=3 | 0.004 | Two tailed t-test |
| 3D | PINK1 KO (+/- IU1)=3 | 0.0182 | Two tailed t-test |
| 3E HSP60 | Control MEF (+/-)=3 | 0.0315 | Two tailed t-test |
| 3E HSP60 | Hela (+/- IU1)=4 | 0.0076 | Two tailed t-test |
| 3E HSP60 | PINK1 KO (+/- IU1)=3 | 0.0478 | Two tailed t-test |
| 3E HSP60 | Control human fibroblast (+/- IU1)=3 | 0.0013 | Two tailed t-test |
| 3E HSP60 | PD patient fibroblast (+/- IU1)=3 | 0.016 | Two tailed t-test |
| 3E ATP5a | Control MEF (+/-)=5 | 0.0435 | Two tailed t-test |
| 3E ATP5a | Hela (+/- IU1)=4 | 0.0057 | Two tailed t-test |
| 3E ATP5a | PINK1 KO (+/- IU1)=3 | 0.0092 | Two tailed t-test |
| 3E ATP5a | Control human fibroblast (+/- IU1)=3 | 0.0142 | Two tailed t-test |
| 3E ATP5a | PD patient fibroblast (+/- IU1)=6 | 0.0003 | Two tailed t-test |
| 4B | 3 | 0.0002 | one way ANOVA/Dunnett's |
| 4E | 3 | 0.0012 | Two tailed t-test |
| 5A | 3 | 0.0219 | Two tailed t-test |
| 5B HSP60 | 4 | 0.034 | Two tailed t-test |
| 5B ATP5a | 4 | 0.0038 | Two tailed t-test |
| 5C | 3 | 0.0257 | Two tailed t-test |

| Figure | N | P value | Statistical analysis used |
|----------|--------------------------------------|----------|----------------------------|
| 5D HSP60 | 3 | 0.0111 | Two tailed t-test |
| 5D ATP5a | 4 | 0.046 | Two tailed t-test |
| 5E | 3 | 0.0318 | Two tailed t-test |
| 5F HSP60 | 4 | 0.041 | Two tailed t-test |
| 5F ATP5a | 3 | 0.0073 | Two tailed t-test |
| 5G | 3 | 0.0483 | Two tailed t-test |
| 5H HSP60 | 4 | 0.036 | Two tailed t-test |
| 5H ATP5a | 3 | 0.0261 | Two tailed t-test |
| 5I | 5 | 0.0488 | Two tailed t-test |
| 5J HSP60 | 4 | 0.0436 | Two tailed t-test |
| 5J ATP5a | 3 | 0.0418 | Two tailed t-test |
| 6A | WT=50; Pink1=120; Pink1 USP14 KD=57 | < 0.0001 | Log-rank test |
| 6B | WT=7; PINK1=5; PINK1 USP14KD= 4 | <0.0001 | One way ANOVA/Newman-Keuls |
| 6C | WT=5; PINK1=5; PINK1 USP14 KD=4 | 0.0001 | One way ANOVA/Newman-Keuls |
| 6D | WT=14; PINK1=11; PINK1 USP14 KD=11 | <0.0001 | One way ANOVA/Newman-Keuls |
| 7A | WT=60; Parkin=60; Parkin USP14 KD=50 | 0.0001 | Log-rank test |
| 7B | WT=3; Parkin=3; Parkin USP14 KD=3 | 0.0095 | One way ANOVA/Newman-Keuls |
| 7C | WT=3; Parkin=3; Parkin USP14 KD=3 | 0.0005 | One way ANOVA/Newman-Keuls |

| Figure | N | P value / P value summary | Statistical analysis used |
|-------------------------------------|---|---------------------------|---------------------------|
| S1A | 3 | P<0.0001 | one way ANOVA/Dunnett's |
| S1A | 3 | ctrl vs 200uM p=0.0025 | Two tailed t-test |
| S1A | 3 | ctrl vs 300uM p=0.0003 | Two tailed t-test |
| S1A | 3 | ctrl vs 400uM p=0.0002 | Two tailed t-test |
| S1A | 3 | ctrl vs 500uM p=0.0002 | Two tailed t-test |
| S1B | 3 | 0.0003 | one way ANOVA/Dunnett's |
| S2A SH-SY5U | 4 | 0.0302 | Two tailed t-test |
| S2A MEF | 3 | 0.0002 | Two tailed t-test |
| S3A | 6 | 0.0018 | Two tailed t-test |
| S3C | 3 | 0.0285 | Two tailed t-test |
| S3D ATP5a | 3 | 0.0285 | Two tailed t-test |
| S3D HSP60 | 3 | 0.0038 | Two tailed t-test |
| S3F | 3 | 0.0413 | Two tailed t-test |
| S3G | 3 | 0.0423 | Two tailed t-test |
| S3J | 3 | 0.0031 | Two tailed t-test |
| S3K | 3 | 0.0427 | Two tailed t-test |
| S4A | 3 | 0.0022 | Two tailed t-test |
| S4B | 3 | 0.029 | Two tailed t-test |
| S5A ATP5a (WT vs IU1 only) | 4 | 0.0002 | Two tailed t-test |
| S5A ATP5a (WT vs IU1 + NH4Cl) | 4 | 0.9892 | Two tailed t-test |
| S5A HSP60 (WT vs IU1 only) | 4 | 0.0143 | Two tailed t-test |
| S5A HSP60 (WT vs IU1 + NH4Cl) | 4 | 0.9354 | Two tailed t-test |
| S5B HSP60 (WT vs WT+IU1) | 3 | 0.0414 | Two tailed t-test |
| S5B HSP60 (KO vs KO+IU1) | 3 | 0.2059 | Two tailed t-test |
| S5B ATP5a (WT vs WT+IU1) | 3 | 0.044 | Two tailed t-test |
| S5B ATP5a (KO vs KO+IU1) | 3 | 0.2229 | Two tailed t-test |
| S6B USP14 | 3 | 0.0445 | Two tailed t-test |
| S6B DRP1 | 3 | 0.0413 | Two tailed t-test |
| S6B Mfn2 | 4 | 0.0007 | Two tailed t-test |
| S6B OPA1 | 3 | 0.0033 | Two tailed t-test |
| S6B Fis1 | 4 | 0.0006 | Two tailed t-test |
| S6B TOM20 | 4 | 0.0002 | Two tailed t-test |
| S6D USP14 | 3 | 0.0005 | Two tailed t-test |
| S6D DRP1 | 3 | 0.0012 | Two tailed t-test |
| S6D Mfn2 | 4 | 0.0092 | Two tailed t-test |
| S6D OPA1 | 4 | 0.0275 | Two tailed t-test |
| S6D Fis1 | 3 | 0.9079 | Two tailed t-test |
| S6D TOM20 | 5 | 0.0397 | Two tailed t-test |
| S7B | 3 | 0.0028 | Two tailed t-test |
| S8B (PHB2 F/F vs F/F + IU1) | 4 | 0.0135 | Two tailed t-test |
| S8B (PHB2 F/F Cre vs F/F Cre + IU1) | 4 | 0.4843 | Two tailed t-test |
| S8D (PHB2 F/F vs F/F + IU1) | 4 | 0.0264 | Two tailed t-test |
| S8D (PHB2 F/F Cre vs F/F Cre + IU1) | 4 | 0.8414 | Two tailed t-test |
| S8E (PHB2 F/F vs F/F + IU1) | 4 | 0.0479 | Two tailed t-test |

| Figure | N | P value | Statistical analysis used |
|-------------------------------------|----|-----------------------------|----------------------------|
| S8E (PHB2 F/F Cre vs F/F Cre + IU1) | 4 | 0.1528 | Two tailed t-test |
| S10A | 60 | 0.058 | Log-rank test |
| S10B | 3 | 0.465 | Two tailed t-test |
| S10C | 3 | 0.6967 | Two tailed t-test |
| S11B | 3 | 0.0243 | Two tailed t-test |
| S11C | 3 | 0.0051 | Two tailed t-test |
| S12A | 4 | 0.8274 | one way ANOVA/Dunnett's |
| S12B | 3 | 0.036 | one way ANOVA/Dunnett's |
| S12B | 3 | ctrl vs IU1 1uM; p=0.1996 | Two tailed t-test |
| S12B | 3 | ctrl vs IU1 10uM; p=0.0102 | Two tailed t-test |
| S12B | 3 | ctrl vs IU1 100uM; p=0.0038 | Two tailed t-test |
| S12C | 3 | 0.0148 | one way ANOVA/Dunnett's |
| S12D | 5 | 0.0292 | one way ANOVA/Tukey |
| S13A | 60 | 0.33 | Log-rank test |
| S14A | 4 | P<0.0001 | One way ANOVA/Newman-Keuls |
| S15 | 2 | | |