



## LRRK2 G2019S KI/KI Mouse Model

This Parkinson's disease knock-in mouse model carries the human G2019S gain-of-function mutation within the endogenous murine LRRK2 gene.

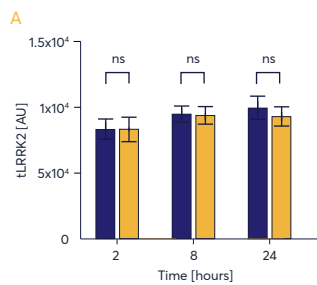
- Unchanged total LRRK2 levels
- Unchanged pS935 LRRK2 levels
- Unchanged basal motor function
- Unchanged cognitive abilities
- Increased pS1292 LRRK2 levels
- Phosphorylation status modifiable by LRRK2 inhibitor MLI-2

**iPSCs with LRRK2 can be used for *in vitro* analysis of mutations**

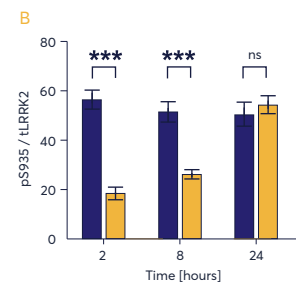
**Figure 1:**  
Time-dependent inhibition of LRRK2 kinase activity upon single oral MLI-2 treatment. LRRK2 G2019S KI/KI mice received a single dose of MLI-2 or vehicle and were sacrificed 2, 8, or 24 hours later. Brain levels of total LRRK2 (A), pS935 LRRK2 (B), and pS1292 LRRK2 (C) were quantified by immunosorbent assay. Mean ± SEM; n = 8 per group. Two-way ANOVA with Bonferroni's post hoc test; \*\*p<0.01, \*\*\*p<0.001; ns, not significant.

Figure 1

### Total LRRK2 Levels



### pS935 LRRK2 Levels



### pS1292 LRRK2 Levels

