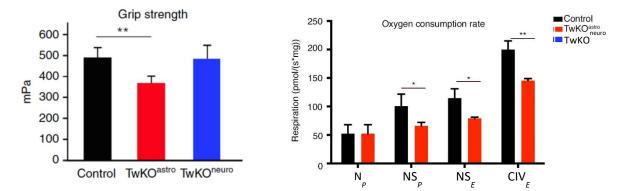


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## Loss of mtDNA activates astrocytes and leads to spongiotic encephalopathy

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The inactivation of Twinkle (TwKO), the replicative  $_{mt}$ DNA helicase, in mice neurons or astrocytes promote the early-onset spongiotic degeneration of brain parenchyma, microgliosis and secondary neurodegeneration



Oxygen flux measured by Oroboros-O2k. The data are presented as mean and error bars indicate standard deviation. \**P*<0.05., \*\**P*<0.01; cerebellum was homogenized with 6-8 strokes in a 2 mL potter-Elvehjem. N<sub>p</sub>: NADH-linked, OXPHOS; NS<sub>p</sub>: NADH&Succinate-linked, OXPHOS; NS<sub>E</sub>: NADH&Succinate-linked, ETS;  $CIV_E$ : Complex IV, ETS.

Reference: Ignatenko O, Chilov D, Paetau I, de Miguel E, Jackson CB, Capin G, Paetau A, Terzioglu M, Euro L, Suomalainen A (2018) Loss of mtDNA activates astrocytes and leads to spongiotic encephalopathy. Nat Commun 9:70.