## $\alpha$ -synuclein overexpression in the retina leads to vision impairment and degeneration of dopaminergic amacrine cells

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**Supplementary Figure 1, related to Figure 1.** (a) Representative ERG traces (in dark conditions) showing the maximum amplitude obtained with flashes of 10 cd.s/m2 in animals treated with rAAV2/6-hu- $\alpha$ -syn and rAAV2/6-GFP (control). (b) Representative ERG traces (light conditions) in animals treated with rAAV2/6-hu- $\alpha$ -syn and rAAV2/6-GFP (control) after treatment with vehicle or L-DOPA. The waves refer to retinal responses after 8 min of light adaptation (last point Figure 1e).



Supplementary Figure 2, related to Figure 4. Dopaminergic amacrine cells, but not GABAergic or cholinergic, are lost in  $\alpha$ -syn injected retina. (a-b) Representative immunofluorescence on vertical retinal sections of rAAV-hu- $\alpha$ -syn injected mice (left) and rAAV-GFP mice (right) stained with antibody anti-tyrosine hydroxylase (TH) (red). White arrows indicate TH+ amacrine cells 1 month post-injection (10x Magnification, Scale bar 50µm), [one-way ANOVA group  $F_{(1,3)}$ =1161.600; p<0.0001]. (c-d) RBPMS staining (red) performed at 3 months post injection in rAAV-hu- $\alpha$ -syn injected mice and rAAV-GFP mice (right) (20x

Magnification, Scale bar 50µm) [one-way ANOVA, group  $F_{(1,27)}$ =5.204; p=0.0306]. (e-f) GAD65 and ChAT staining on retina of rAAV-hu- $\alpha$ -syn and rAAV-GFP- injected at 3 month post injection (20x Magnification, Scale bar 50µm) [GAD65: one-way ANOVA, group  $F_{(1,41)}$ = 0.733; p=0.3970; ChAT+: one-way ANOVA, group  $F_{(1,28)}$ =2.128; p=0.155]. Data represent mean ± SEM. # p < 0.05 vs rAAV-GFP.

**Supplementary Videos.** Videos showing visual task performance of C57BL/6J mice preinjection (Video 1), after intravitreal injection of (rAAV-hu- $\alpha$ -syn) combined with saline (Video 2) and L-DOPA replacement therapy (Video 3). (**Video 1**) Representative sample trial of a C57BL/6J mouse during the visual task where it is required to discriminate between a 3 cm black and white striped card and a 4 cm black and white striped card. The platform is located under the 3 cm striped card. (**Video 2**) Representative trial of the visual task showing that bilateral rAAVhu- $\alpha$ -syn intravitreal injection affects performance at the first stage of the procedure when animals are required to discriminate between a 1 cm black and white striped card and a 10 cm black and white striped card. The platform is located under the 1 cm striped card. (**Video 3**) Representative trial of the visual task showing that L-DOPA treatment rescues performance in rAAV-hu- $\alpha$ -syn injected mice on the same trial reported in Video 2.