

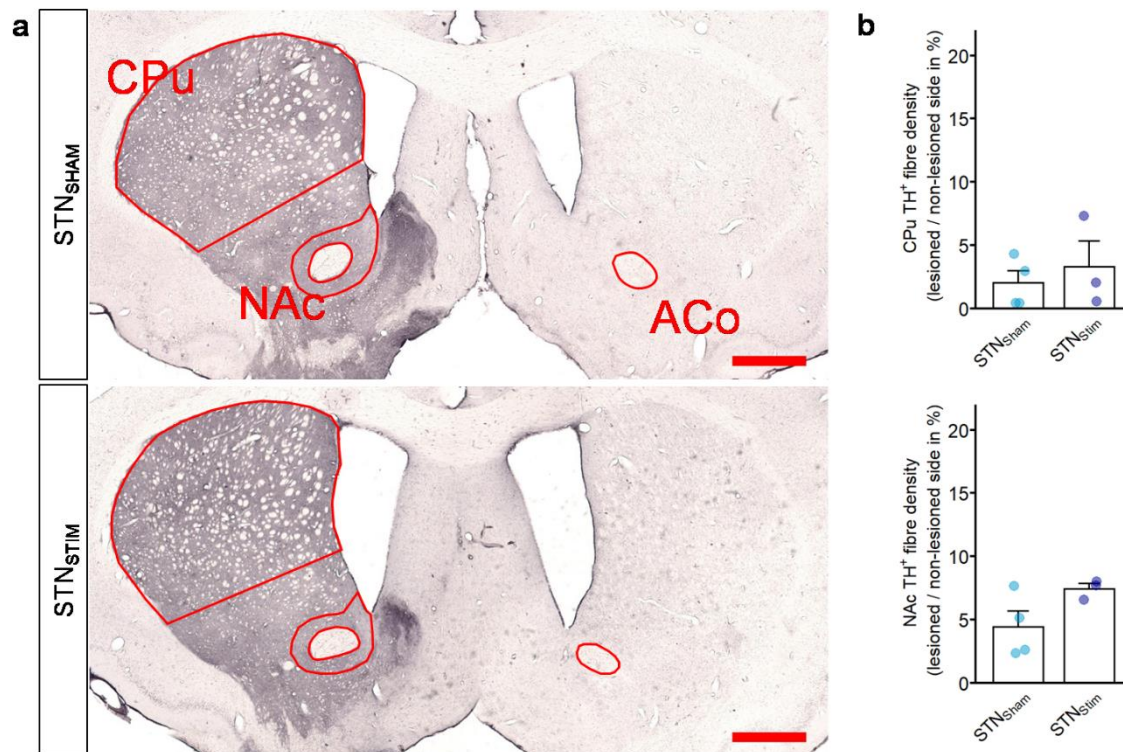
# **Subthalamic nucleus deep brain stimulation induces nigrostriatal dopaminergic plasticity in a stable rat model of Parkinson's disease**

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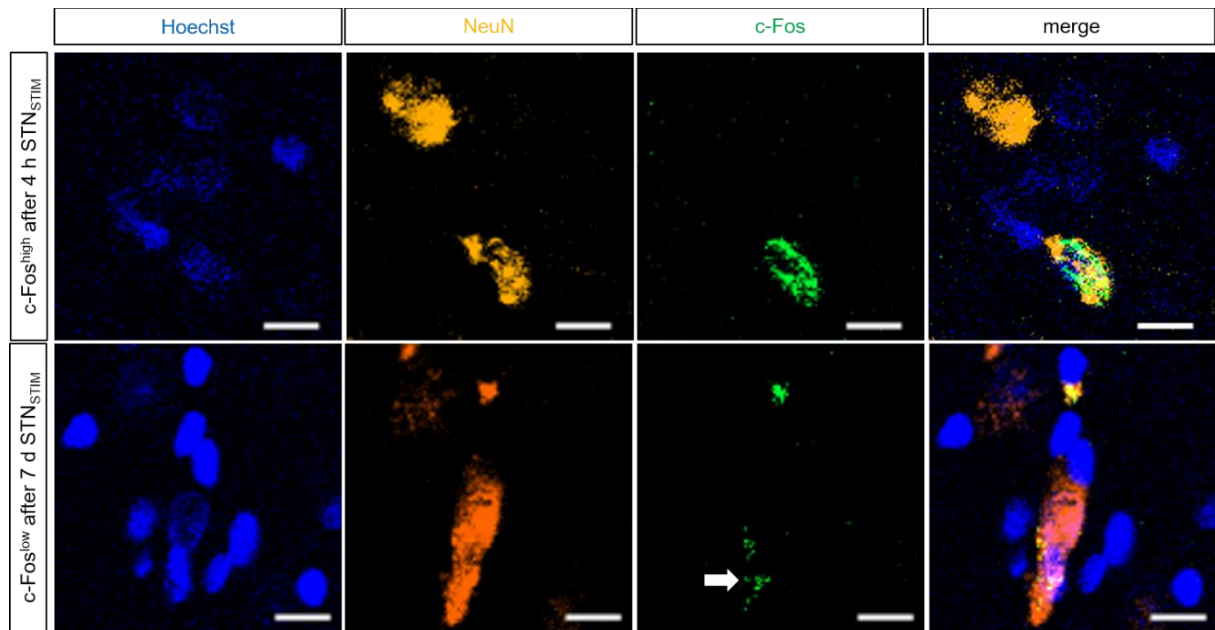
## **Supplemental Materials**

- **Figure S1.** Influence of one week of STN-DBS on striatal dopaminergic neurotransmission.
- **Figure S2.** Intensity of c-Fos immunostainings depending on STN-DBS durations.
- **Table S1.** Quantitative data and statistics of c-Fos immunohistochemistry.

Supplementary figures:



**Supplementary Figure S1:** (a) Representative immunohistological analyses of tyrosine hydroxylase (TH) stainings in the striatum of hemiparkinsonian rats after seven days of unilateral, right-sided STN-DBS. Marked regions are the dorsal striatum (= Caudate nucleus putamen; CPu) and the *Nucleus accumbens core region* (NAc) as the main target areas of the midbrain dopaminergic systems and the anterior commissure (ACo) as the reference region (white matter). Scale bars, 1000  $\mu$ m. (b) Densiometric analysis of TH<sup>+</sup> fibre densities within the CPu as the main target region of *substantia nigra* neurons and the NAc as the main target region of neurons in the ventral tegmental area showed no effects of STN-DBS in lesioned hemispheres (normalized to respective non-lesioned hemispheres). **Abbreviations:** STN - subthalamic nucleus; TH – tyrosine hydroxylase.



**Supplementary Figure S2:** Immunohistochemical c-Fos staining reveals persistent basal c-Fos expression after 7 days of continuous STN-DBS (white arrow; lower panel; c-Fos<sup>low</sup>), which differs from more widespread expression after short-term stimulation after 4h (upper panel; c-Fos<sup>high</sup>). Scale bar 10  $\mu$ m. *Abbreviation:* STN - subthalamic nucleus.

	Non-lesioned side				Lesioned side		
	STN <sub>SHAM</sub>	STN <sub>STIM</sub>	<i>P</i> -value		STN <sub>SHAM</sub>	STN <sub>STIM</sub>	<i>P</i> -value
<b>c-Fos<sup>low</sup> cells (% of cells)</b>							
<b>SNpc</b>	25.6±5.5%	19.1±4.8%	0.404		23.4±7.9%	16.8±3.1%	0.467
<b>VTA</b>	18.5±6.4%	15.3±5.9%	0.484		16.5±8.7%	18.5±8.6%	0.755
<b>c-Fos<sup>low</sup>/TH<sup>+</sup> neurons (% of all TH<sup>+</sup> neurons)</b>							
<b>SNpc</b>	21.8±5.5%	15.4±4.0%	0.382		37.5±31.5%	26.4±2.7%	0.692
<b>VTA</b>	14.8±3.52%	12.5±2.6%	0.606		11.6±2.4%	15.4±3.0%	0.355

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40 **Supplementary Table S1.** Quantitative data and statistics of c-Fos immunohistochemistry.

41 Data are presented as mean values ± S.E.M. (n=3 or 4). *P*-values are from unpaired two-sided

42 t-tests or Welch tests as appropriate to determine mean differences between groups.

43 **Abbreviations:** STN - subthalamic nucleus; TH - tyrosine hydroxylase; SNpc - Substantia

44 nigra pars compacta; VTA - ventral tegmental area.