

# Effects of dopamine D2 receptor antagonist haloperidol on movement speed in a drawing task

Sophie Sowden, Lydia Hickman, Bianca Schuster, Alicia Rybicki, Dagmar Fraser, Jennifer Cook

Centre for Human Brain Health, School of Psychology, University of Birmingham, UK

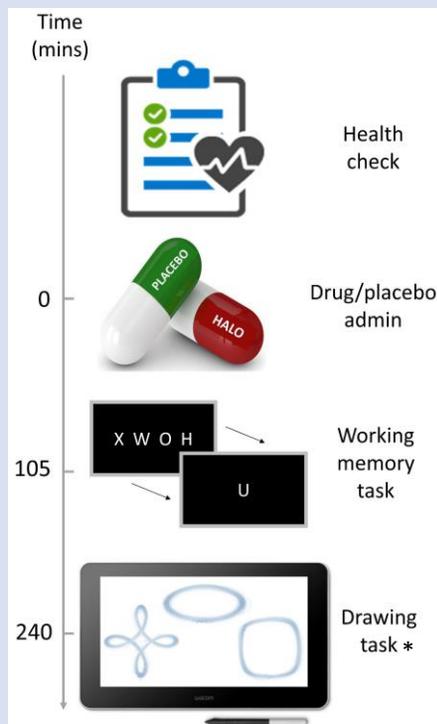


## Background

- **Decreased brain dopamine** has been related to **slowed movement** response initiation<sup>1</sup> and **reduced movement vigour** when reaching to a target<sup>2</sup>.
- **Withdrawal of dopaminergic medication** in patients with Parkinson's Disease (PD) leads to **abnormalities in handwriting kinematics**<sup>3</sup>.
- To date, few studies have investigated dopaminergic modulation of movement speed in **tasks devoid of the reward-based motivation to move**, and many use only **ellipse shapes** (considering a single angular frequency)<sup>4</sup>.
- Moreover, it is important to **control for baseline working memory (WM)**, as research has demonstrated its tight association with **baseline striatal dopamine**, with differential drug effects found in those with high vs. low WM<sup>5,6</sup>.

## Method

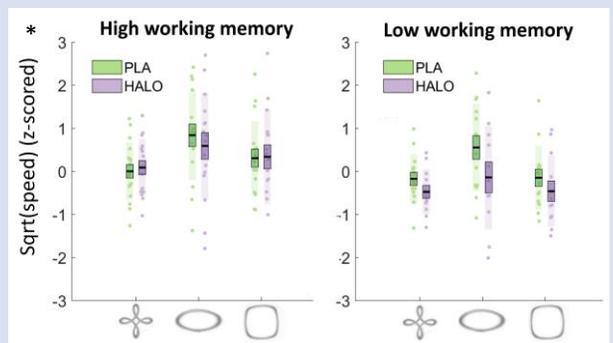
- **37 healthy adult participants** (16 female;  $M_{AGE}$  26.3;  $M_{BMI}$  23.3)
- **Double-blind, placebo-controlled** procedure to test the effect of the D2 antagonist haloperidol on movement speed during a simple drawing task.
- On two separate days, once after receiving **2.5mg haloperidol** and once after receiving **placebo**, participants completed the following procedure:



\* Stylus used to draw 3 shapes (10  $\pi$  rotations) on a Wacom touchscreen. X and y position was recorded at 133Hz. For each participant indices of overall speed were derived.

## Results

- Linear mixed effect models examined the **impact of drug on drawing speed** as a function of shape and baseline dopamine levels (indexed by placebo WM):  
$$\text{speed} \sim \text{shape} + \text{drug} + \text{shape} * \text{drug} + \text{shape} * \text{WM} + \text{drug} * \text{WM} + \text{shape} * \text{drug} * \text{WM} + (1 | \text{subject})$$
- Drawing speed differed as a function of shape [ $F(2,180) = 18.82, p < .001$ ]; with ellipse shapes of highest speed.
- Following administration of haloperidol, participants' drawing speed was significantly slowed compared to placebo [ $F(1,180) = 4.28, p < .05$ ].
- Crucially, there was an **interaction between drug and baseline WM performance** [ $F(1,180) = 6.75, p < .05$ ].
  - Those with high baseline WM showed no change in speed of drawing between placebo and haloperidol ( $p = .82$ )
  - Those with **low baseline WM** (i.e. low baseline dopamine) showed significantly **slower drawing speed following haloperidol** than placebo ( $p < .05$ )



\* Median split by placebo working memory. Drawing speed by shapes for placebo and haloperidol conditions. Thick black horizontal line represents the sample mean, surrounding box represents 1 standard error of the mean and shaded region represents 1 standard deviation.

## Discussion

- Our results support the role of dopamine in drawing speed and advocate the **use of the current task to assess movement atypicalities associated with PD**.
- Our findings are in line with those by previous studies highlighting **differential effects of haloperidol** between individuals with **low and high baseline WM**<sup>5,6</sup>.

## References

1. Quattrocchi G, Monaco J, Ho A, Irmen F et al., (2018). Pharmacological dopamine manipulation does not alter reward-based improvements in memory retention during a visuo-motor adaptation task. *eNeuro*, 5(3): 1–12.
2. Niv Y, Daw ND & Dayan P. (2006). How fast to work: Response vigor, motivation and tonic dopamine. *Advances in Neural Information Processing Systems*, 18, 1019–1026.
3. Tucha O, Mecklinger L, Thome J, Reiter A et al., (2006). Kinematic analysis of dopaminergic effects on skilled handwriting movements in Parkinson's disease *Journal of Neural Transmission*, 113(5): 609–623.
4. Huh, D., & Sejnowski, T. J. (2015). Spectrum of power laws for curved hand movements. *Proceedings of the National Academy of Sciences*, 112(29), E3950–E3958.
5. Cools, R., Gibbs, S. E., Miyakawa, A., Jagust, W., & D'Esposito, M. (2008). Working memory capacity predicts dopamine synthesis capacity in the human striatum. *Journal of Neuroscience*, 28(5): 1208–1212.
6. Frank MJ & O'Reilly RC. (2006). A mechanistic account of striatal dopamine function in human cognition: Psychopharmacological studies with cabergoline and haloperidol. *Behavioral Neuroscience*, 120(3): 497–517.



UNIVERSITY OF BIRMINGHAM



@Sophie\_Sowden



s.l.sowden@bham.ac.uk