# Cytoarchitectonic Maps of five newly identified Areas in the human Dorsolateral Prefrontal Cortex



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## BACKGROUND

# The Dorsolateral Prefrontal Cortex (DLPFC)



#### Location

Middle (medial and lateral) superior frontal gyrus; Brodmann area 9 & 46

#### Functions

'higher-order' cognitive functions



SFG2

# METHODS

## **Observer-independent border detection**

## A: Cytoarchitectonical analysis

ten post-mortem brains (5 men, 5 women: 39 – 86 years old); serial, coronal sections (thickness: 20 μm; every 15<sup>th</sup> section); cell body silver staining<sup>17</sup> Box: ROI, SFG2 (pink) and SFG3 (blue)

#### **B: GLI-Profiles**



Sarkissov et al. (1995)<sup>14</sup>

attention<sup>1,2</sup>, planning, executive function, working memory<sup>-3-6</sup>, behavior<sup>7,8</sup>

#### Disorders

Schizophrenia<sup>9</sup>, depression<sup>10</sup>, obsessivecompulsive disorder<sup>11</sup>

#### Discrepancy

in maps (see figures): The locational relationship of area 46 and area 9<sup>15,16</sup>

#### Aims

- Delineation and cytoarchitectonic analysis of the human DLPFC with focus
   on the superior frontal gyrus and the middle frontal gyrus
- Generation of probabilistic cytoarchitectonic maps to assign potential functions
- High-resolution 3D reconstruction maps for interpretation and comparison of neuroimaging studies and future research projects on the human frontal lobe







microscopically defined region of interest digitized; computation of gray level index (GLI) images GLI profiles represent laminar changes between layer II and cortex/white matter borders (yellow and magenta lines), i.e., cytoarchitecture. Arrow marks the border of SFG2 and SFG3 that was computed (C, D)<sup>18,19</sup>

#### C/D: Significant border detection

Significant maxima of Mahalanobis distance (black dots) were tested for different block sizes (E; n = 20 - 30). At least three block sizes mark a significant border (Bonferroni corrected alpha level of 0.001)<sup>18,19</sup>

#### D: Mahalanobis distance

Observer-independent border detection; significant maxima of Mahalanobis distance at profile number 53 (asterisk and red line) are plotted against the profile index<sup>18,19</sup>

## RESULTS

#### Cytoarchitectonic characteristics of the DLPFC Areas



#### Location

five new cytoarchitectonic areas on the superior frontal gyrus (SFG2, SFG3 and SFG4), on the surface of frontal gyrus (MFG4 and MFG5)

#### Boundaries

<u>Rostral</u>: Frontal pole (area 10)<sup>19</sup> <u>Lateral</u>: anterior DLPFC (SFS1, SFS2, MFG1, MFG2)<sup>20</sup> <u>Ventral</u>: Broca (area 44/45)<sup>21</sup> <u>Caudal</u>: Premotor & prefrontal cortex (area 6 & 8)<sup>22</sup>

### **Delineations of ROIs**

## Location and extent of identified areas



MFG5





## **REFERENCES & ACKNOWLEDGEMENTS**

Rowe, J.B. & R.E. Passingham (2001) Neuroimage, 14: p. 77-86.
 Hoshi, E. and J. Tanji (2004) J Neurophysiol, 91(6): p. 2707-22.
 Pribram, K.H., et al.(1952) J Comp Physiol Psychol, 45(6): p. 565-75.
 Petrides, M. (2000) J Neurosci, 20(19): p. 7496-503.
 Rowe, J.B. et al. (2000) Science, 288(5471): p. 1656-60
 Barbey, A.K. et al. (2013) Cortex, 49(5): p. 1195-205.

7. Shallice, T. & P.W. Burgess (1991) Brain, 114 (Pt 2): p. 727-41.
8. Barraclough, D.J., et al. (2004) Nat Neurosci, 7(4): p. 404-10.
9. Huang, M.L. et al. (2017) Medicine (Baltimore), 96(25): e7228
10. Koenigs et al., (2009) Behav Brain Res, 201(2): 239-243
11. Russel et al., (2003) J Child Adolesc Psychopharmacol, 13 Suppl 1: S31-38
12. von Economo, C. & G.N. Koskinas (1925) Springer, Berlin

13. Brodmann, K. (1909) Verlag von J. A. Barth, Leipzig
14. Sarkissov et al., (1999) Medgiz, Moskau
15. Petrides, M. & D.N. Pandya (1999) Eur J Neurosci, 11(3): p. 1011-36
16. Rajkowska, G. & P.S. Goldman-Rakic (1995) Cereb Cortex, 5(4): p. 307-22.
17. Amunts et al. (2020) Science, 369(6506): p. 988-992

Schleicher, A. et al. (1999) Neuroimage, 9(1): p. 165-77.
 Bludau, S. et al. (2014) Neuroimage, 93 Pt 2: p. 260-75.
 Bruno et al., (2022) Front Neuroanat, 16:915877
 Amunts (1999) J Comp Neurol, 412(2): 319-341
 Sigl et al. (2016) OHBM Annual Meeting

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