Human cerebral organoids and fetal brain tissue share proteomic similarities: enabling alternative models to study psychiatric disorders

Juliana Minardi Nascimento<sup>1,2</sup>, Verônica M Saia-Cereda<sup>1</sup>, Rafaela C. Sartore<sup>2,3</sup>, Rodrigo Madeiro da Costa<sup>2</sup>, Clarissa S. Schitine<sup>4,5</sup>, Hercules Rezende Freitas<sup>5,6</sup>, Michael Murgu<sup>7</sup>, Ricardo A. de Melo Reis<sup>5</sup>, Stevens K Rehen<sup>2,4\*</sup>, Daniel Martins-de-Souza<sup>1,8,9\*</sup>

## **Supplementary information**

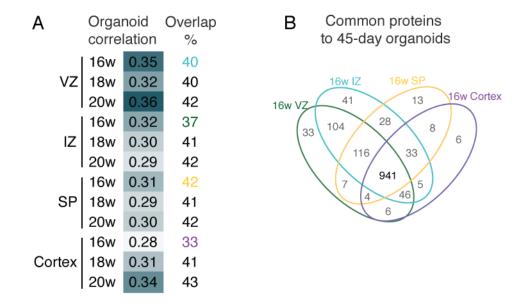
Supplementary Table 1: Protein expression of PSC-derived, 45-day cerebral organoids.

(Excel file)

Supplementary Table 2: Enriched proteins of glutamatergic and dopaminergic synapse

(Excel file)

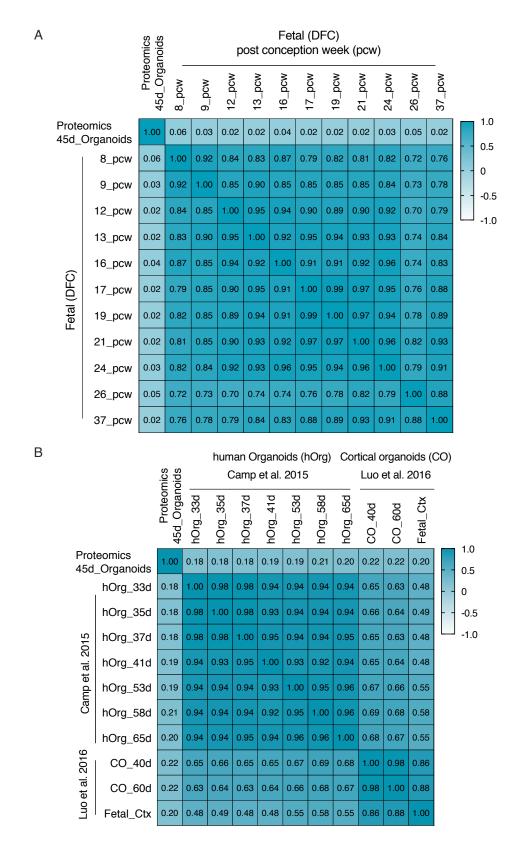
Supplementary Figure 1, 2 and 3



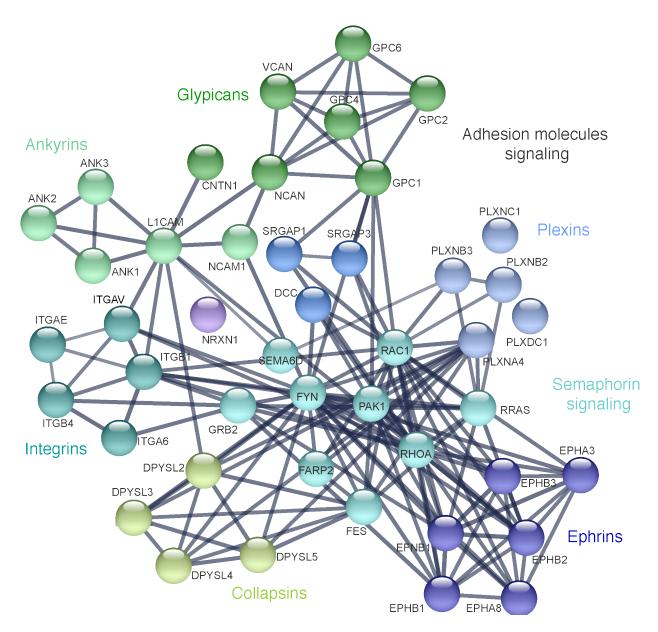
## C Common proteins of 16 week VZ, IZ, SP and cortex to organoids

	GO Terms	FDR
Biological	Cell -cell adhesion	1.22E-40
Processes	mRNA splicing	5.84E-28
	Regulation of mRNA stability	9.71E-28
	Translational initiation	2.62E-23
Cellular	Extracellular exosome	1.48E-162
Compartment	Cytosol	3.26E-110
	Myelin sheath	7.03E-68
	Membrane	1.59E-60
Molecular Function	Poly(A) RNA binding	1.69E-109
	Protein binding	1.70E-58
	Cadherin binding involved in cell-cell adhesion	1.17E-45
	RNA binding	1.02E-36

**SUPPLEMENTARY FIGURE 1:** Comparison of cerebral organoid proteomes with developing brain sub-compartments (ventricular zone, VZ; intermediate zone, IZ; subplate, SP; cortex). (A) Spearman correlation of 45-day cerebral organoid proteomes to 16-20 weeks (VZ, IZ, SP, and cortex proteomes). (B) Venn diagram of proteins commonly found in 45-day cerebral organoids and 16-week developing human brain areas. (C) Gene ontology of the proteins found in common to the 16-week (VZ, IZ, SP, and cortex) proteomes and cerebral organoids proteomes.



**SUPPLEMENTARY FIGURE 2:** Comparison of cerebral organoid proteomes with transcriptomics of the developing brain and cortical cerebral organoids. Spearman correlation of 45-day cerebral organoid proteome compared to **(A)** human fetal 8-37 post conceptional week, dorsolateral prefrontal cortex (DFC) and **(B)** human cortical organoids of 33 to 65 days *in vitro*.



**SUPPLEMENTARY FIGURE 3:** Interactive organizational network of proteins from 45-day cerebral organoids. Molecular relationships between proteins found in cerebral organoids from the dataset are based on the STRING database (<a href="https://string-db.org/">https://string-db.org/</a>). Groups of proteins are separated by different colors, as indicated.