#### Check for updates

#### **OPEN ACCESS**

EDITED AND REVIEWED BY Jaewon Ko, Daegu Gyeongbuk Institute of Science and Technology (DGIST), Republic of Korea

\*CORRESPONDENCE Robert Hindges Import robert.hindges@kcl.ac.uk Zsolt Lele Import Lele Import Lele

#### SPECIALTY SECTION

This article was submitted to Neurodevelopment, a section of the journal Frontiers in Neuroscience

RECEIVED 30 November 2022 ACCEPTED 20 December 2022 PUBLISHED 10 January 2023

#### CITATION

Hindges R and Lele Z (2023) Editorial: Cell adhesion molecules in neural development and disease. *Front. Neurosci.* 16:1112300. doi: 10.3389/fnins.2022.1112300

#### COPYRIGHT

© 2023 Hindges and Lele. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner(s) are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.

# Editorial: Cell adhesion molecules in neural development and disease

# Robert Hindges<sup>1,2\*</sup> and Zsolt Lele<sup>3\*</sup>

<sup>1</sup>Centre for Developmental Neurobiology, King's College London, London, United Kingdom, <sup>2</sup>MRC Centre for Neurodevelopmental Disorders, King's College London, London, United Kingdom, <sup>3</sup>Laboratory of Molecular Neurobiology, Institute of Experimental Medicine, Budapest, Hungary

#### KEYWORDS

cell adhesion molecules, protocadherin, teneurin, N-cadherin, circuit assembly

### Editorial on the Research Topic Cell adhesion molecules in neural development and disease

Cell-to cell adhesion is a defining, hence essential condition of being a multicellular organism. It has been more than 60 years that Weiss published the first in a series of pioneering papers detailing various aspects of cellular adhesion (Weiss, 1959). Since then, a large number of papers has been published on this fascinating Research Topic describing all the studies that contributed to the state-of-the-art knowledge of today. In this Research Topic of Frontiers in Neuroscience, we collected a series of papers, both original research articles and reviews to emphasize the importance of cell adhesion molecules in neural development and disease. Two of the original research papers presents novel data involving protocadherins. Members belonging to this family have previously been demonstrated to be responsible for dendritic selfavoidance (Kostadinov and Sanes, 2015; Lefebvre et al., 2015; Ing-Esteves, 2018), axon sorting of olfactory sensory (Mountoufaris, 2017), and serotonergic neurons (Chen, 2017; Katori, 2017). In this Research Topic, Pancho et al. demonstrates the importance of PCDH19 in interneuron migration while Luo et al. propose the involvement of PCDH11x in target specification of hippocampal mossy fibers. An excellent overview provided by Moreland and Poulain outlining the role various cell adhesion molecules play in neural circuit assembly. As a perfect continuation of this Research Topic, another review by Meltzer and Schuldiner discusses the involvement of CAMs in neuronal remodeling. As a sharp contrast to these broad reviews, and as a reflection of recent surprising developments, László and Lele tell everything you wanted to know about N-cadherin in neural development and disease. An important general issue is the fine balancing of activities controlled by adhesion molecules. This includes not only the positive regulation of cell-cell contacts, but can also involve negative activities. Here, Baeriswyl et al. characterize such balance between positive and negative action in the context of Purkinje cell migration. Two reports focus on the teneurin family of cell adhesion molecules. A review by Dodsworth and Lovejoy focuses on the teneurin C-terminal associated peptides (TCAP), which are encoded by the last exon of teneurins. Interestingly, despite a general transsynaptic interaction of full-length teneurins with latrophilins, evidence suggests that released TCAP molecules have an additional binding capacity to these partners and might elicit distinct cellular process. The presence of teneurins at synapses and their ability of heterocomplexes in cis is described in an article by Cheung et al. The results suggest that the diversity of molecular complexes at synaptic localizations is bigger than previously thought, which thus would increase the combinatoric power to control synaptic specificity. Finally, the process of how synapse formation is controlled through structural domains of different proteins across species is presented in an article by González-Calvo et al. therefore enabling us to recognize the evolutionary conservation of these fundamental processes.

It is evident that many open questions about the structure and roles of cell adhesion molecules still exist. However, the recent progresses made are encouraging and point toward a better understanding not only in biochemical and cell biological terms, but importantly also in the context of disorders, where there is a clear need for the development of novel therapeutic strategies.

# Author contributions

RH and ZL wrote the summary. Both authors contributed to the article and approved the submitted version.

### Funding

Support to RH has been provided by the Leverhulme Trust (RPG-2021-385) and the Medical Research Council (MR/W006251/1). Support to ZL has been provided by the National Research, Development and Innovation Fund of Hungary under the 'Frontline' - Research Excellence Program KKP\_19 (KKP 129961) and the National Program in Brain Sciences (2017-1.2.1-NKP-2017-00002) funding scheme.

## **Conflict of interest**

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

### Publisher's note

All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

### References

Chen, W. V, Nwakeze, C. L, Denny, C. A, O'Keeffe, S, Rieger, M. A, Mountoufaris, G, et al. (2017). Pcdhalphac2 is required for axonal tiling and assembly of serotonergic circuitries in mice. *Science* 356, 406–411. doi: 10.1126/science.aal3231

Ing-Esteves, S, Kostadinov, D, Marocha, J, Sing, A. D, Joseph, K. S, Laboulaye, M. A, et al. (2018). Combinatorial effects of alpha- and gamma-protocadherins on neuronal survival and dendritic self-avoidance. *J. Neurosci.* 38, 2713–2729. doi: 10.1523/JNEUROSCI.3035-17.2018

Katori, S, Noguchi-Katori, Y, Okayama, A, Kawamura, Y, Luo, W, Sakimura, K, et al. (2017). Protocadherin-alphaC2 is required for diffuse projections of serotonergic axons. *Sci. Rep.* 7, 15908. doi: 10.1038/s41598-017-16 120-y

Kostadinov, D., and Sanes, J. R. (2015). Protocadherin-dependent dendritic self-avoidance regulates neural connectivity and circuit function. *Elife* 4, 22. doi: 10.7554/eLife.08964.022

Lefebvre, J. L., Sanes, J. R., and Kay, J. N. (2015). Development of dendritic form and function. *Annu. Rev. Cell Dev. Biol.* 31, 741–777. doi: 10.1146/annurev-cellbio-100913-013020

Mountoufaris, G, Chen, W. V, Hirabayashi, Y, O'Keeffe, S, Chevee, M, Nwakeze, C. L, et al. (2017). Multicluster Pcdh diversity is required for mouse olfactory neural circuit assembly. *Science* 356, 411–414. doi: 10.1126/science.aai8801

Weiss, L. (1959). Studies on cellular adhesion in tissue culture. I. The effect of serum. *Exp. Cell Res.* 17, 499–507. doi: 10.1016/0014-4827(59)90070-9