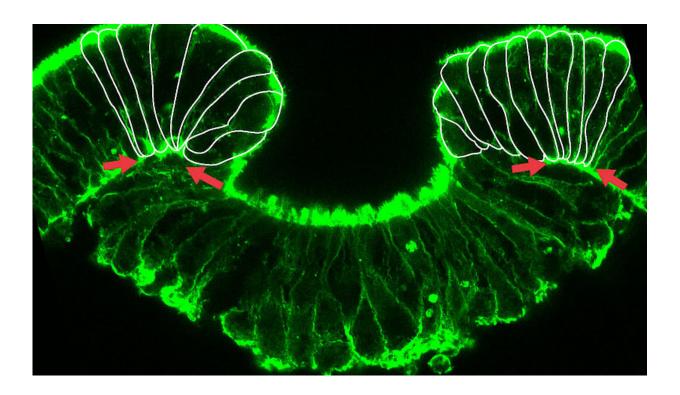


Embryonic gene regulation through mechanical forces

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Early stage of invagination of the inner cell layer, the endoderm. Cells at the margins show a strong deformation. Credit: Ulrich Technau

During embryonic development, genetic cascades control gene activity and cell differentiation. In a new publication of the journal *PNAS*, the team of Ulrich Technau of the Department of Molecular Evolution and Development at the University of Vienna reported that besides the genetic program, mechanical cues also contribute to the regulation of



gene expression during development. Comparisons with other animals suggests that this regulatory principle is ancient.

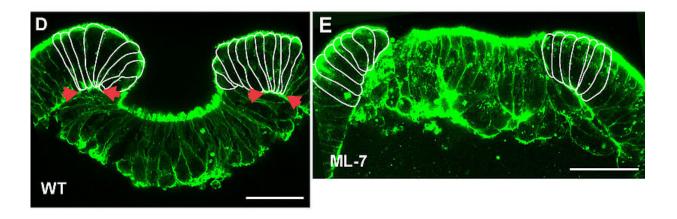
It is generally thought that <u>embryonic development</u> and cellular differentiation of animals and humans follows a precise <u>genetic program</u> of spatiotemporal <u>gene expression</u>. However, a number of recent studies suggested that mechanotransduction – the ability of cells to transform mechanical forces into biochemical signals – can also contribute to the regulation of gene <u>expression</u>, and thus may play an important role in development. While most of these studies were done in cell cultures, the team of Ulrich Technau from the University of Vienna now reports experiments with mechanosensitive gene expression during early development of the starlet sea anemone Nematostella vectensis.

The authors show in a recent publication in the *Proceedings of the National Academy of Sciences* that chemical inhibition of cellular myosin function not only blocks the morphogenetic movement of gastrulation – the process in which inner and outer cell layers are formed by invagination – but also abolishes the expression of a crucial developmental regulator gene, brachyury. This gene has a crucial ancient role in the development of virtually all animals. Surprisingly, external mechanical pressure applied to such embryos can activate or restore the expression of brachyury. Furthermore, brachyury expression in Nematostella vectensis via mechanotransduction depends on β -catenin, a key protein with a conserved dual role in cell-cell adhesion and in gene regulation following a signaling cascade.

Based on their findings, the authors propose a feedback loop whereby mechanical and genetic regulation work together to ensure robust brachyury expression. In addition, because β -catenin-dependent mechanotransduction occurs in other animals like zebrafish and the fruitfly, the findings suggest that this form of gene regulation dates back to at least 600 million years ago, the evolutionary split between



vertebrates, insects and sea anemones.



In embryos treated with the myosin inhibitor ML-7, cells at the margin do not deform and invagination is blocked. Credit: Ulrich Technau

More information: Ekaterina Pukhlyakova et al. β-Catenin–dependent mechanotransduction dates back to the common ancestor of Cnidaria and Bilateria, *Proceedings of the National Academy of Sciences* (2018). DOI: 10.1073/pnas.1713682115

Provided by University of Vienna



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