How to Turn an Embryo Inside Out

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Images of an algal embryo transitioning into an adult give physicists a rare opportunity to test a mechanical model of morphogenesis.

Subject Areas: Biological Physics

A Viewpoint on:
Dynamics of a Volvox Embryo Turning Itself Inside Out
Stephanie Höhn, Aurelia R. Honerkamp-Smith, Pierre A. Haas, Philipp Khuc Trong, and Raymond E. Goldstein
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We all start life as a small ball of cells. These cells proliferate and organize into tissues, which in turn grow and fold into the complex form that defines a human being. The process that leads from a formless ball to a functional organism—known as morphogenesis—is an active area of research in biology. For physicists, it presents an exciting challenge because a growing organism can be thought of as a type of “active” matter that is capable of reshaping itself \[1, 2\]. Physics models are therefore increasingly being used to investigate morphogenetic events in plants \[3\] and animals \[4\].

Now, researchers at the University of Cambridge in the UK have compared images of a growing algal embryo with a physics-based morphogenesis model, providing one of the few experimental tests of such models \[5\]. Stephanie Höhn and her colleagues imaged embryos of the algal species *Volvox globator* as they underwent invagination—a fundamental event in morphogenesis in which a sheet of cells folds inside, triggering the formation of an internal cavity. The researchers modeled the organism as a spherical elastic shell that actively bends and contracts at certain segments, and they found the necessary balance between these motions for inversion to occur.

*Volvox* are freshwater algae generally found in ponds \[6\]. An adult *Volvox* consists of a single layer of a few thousand cells embedded within a gelatinous extracellular matrix that encloses a spherical cavity. Each cell has two flagella that face outwards and beat in the water, allowing the spheroid to swim towards light and nutrients. Embryo *Volvox* develop within a parent spheroid (Fig. 1). And, unlike adults, their flagella are oriented inwards. To form an adult, the embryonic sheet of cells must therefore turn itself inside out.

This stunning process of inversion varies from species to species \[7\]. In *V. globator*, one half of the spherical sheet of cells shrinks in radius and invaginates, initiating inversion (Fig. 2, top). In 2011, Höhn and Armin Hallmann \[8\] obtained, at different stages of inversion, the cross-sectional shape of the *V. globator* embryos. They also imaged the shape of the cells in the spherical sheet, which allowed them to relate the macroscopic shape of the embryo to the microscopic behavior of the cells. In their new work \[5\], Höhn et al. observed the live inverting embryos in three dimensions and quantified their changing shapes in real time. They also constructed and validated a continuous mechanical model that provides a tissue-based view of how inversion occurs.

Embryo *Volvox* are small (70–90 micrometers) and their growth is highly sensitive to external probes, such
as intense light or an electron beam. Both factors make imaging the organism a challenge. To reduce the organism’s exposure to intense light, Höhn and her collaborators used a high-resolution optical technique (selective plane illumination microscopy [9]) that illuminates and collects light from just a slice of the Volvox embryo. The images were then pieced together to form a 3D image.

Höhn et al. imaged the embryos at regular time intervals throughout inversion, at each time step tracing the contour of the cell sheet. From the contours, the team could measure the depth of invagination, the area of the cell sheet, and the curvature along the sheet.

With this quantitative data in hand, the researchers tried to find the simplest mechanical model that reproduced the shape changes in the embryo as it began the invagination process. As a starting point, they modeled the embryo as a thin elastic shell that was initially spherical. The researchers then derived an equation that predicts the shape of the shell when a segment bends or contracts. They then solved their equation numerically to capture later stages of inversion, when the cell sheet reconnects.

By varying the weight of these two ingredients in Höhn et al.’s model—bending and contraction—are generic. By varying the weight of these two ingredients, the model should, in principle, be applicable to other organisms with invaginating thin tissues.

**References**


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About the Author

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Arezki Boudaoud obtained a Ph.D. in 2001 from Université Pierre et Marie Curie, Paris. After a postdoctoral stay at MIT, he was hired as a CNRS Research scientist at Ecole Normale Supérieure, Paris. He initially worked in nonlinear and statistical physics, notably addressing pattern formation in thin solid and liquid sheets, and gradually switched to morphogenesis in living systems. He joined the Department of Biology at Ecole Normale Supérieure de Lyon in 2009. His team currently investigates the biomechanics of morphogenesis in plants and in animals, as well as the mechanisms that make the shape of organisms reproducible.