Internship Proposal

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Project Title:

Dissecting RhoGTPase regulation to understand how epithelial tissues grow and migrate **Level:**

Master Student

Project Summary:

Epithelial tissues are an essential barrier that isolate the internal organ environment from the exterior. This property relies on their ability to establish molecular and functional asymmetries along an apico-basal axis. Epithelial apico-basal organization is a conserved trait whose its disruption is a serious threat to development and human health. However, apical-basal organization must also be highly dynamic to drive cell shape changes during cell division or cell migration. Using endogenously GFP-tagged Drosophila lines, we identified two uncharacterized RhoGTPase regulators that control 1) epithelial reorganization during cell division (co-supervised by André Carvalho) or 2) cell shape change during tissue migration (co-supervised by Vítor Yang). In these projects, we will combine optogenetic tools, live imaging in Drosophila tissues and mammalian cell culture, to discover how these regulators modulate the cytoskeleton and the polarity machinery to shape animal organs.

Work to be developed by the student:

Drosophila genetics; live imaging of cell division or cell migration in Drosophila ovaries cultured ex-vivo; optogenetics; immunofluorescence; confocal microscopy of fixed tissues; mammalian cell culture; molecular biology

References:

•di Pietro, F.; Osswald et al., Systematic characterization of Drosophila RhoGEF/GAP localizations uncovers regulators of mechanosensing and junction formation during epithelial cell division. Curr Biol (in press)

•Osswald, M. et al., (2022). aPKC regulates apical constriction to prevent tissue rupture in the Drosophila follicular epithelium. Curr Biol 32, 4411-4427 e4418.

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•Zeng, J., Santos, A.F et al., (2021). Target-induced clustering activates Trim-Away of pathogens and proteins. Nat Struct Mol Biol 28, 278-289. 10.1038/s41594-021-00560-2.
•Moreira, S. et al.,(2019). PP1-Mediated Dephosphorylation of Lgl Controls Apical-basal Polarity. Cell Rep 26, 293-301 e297. 10.1016/j.celrep.2018.12.060.





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