

Physical modelling of ESCRT-III mediated cell division in archaea

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Studying the mechanism of cell division in evolutionary simpler cells can teach us about the mechanistic principles that have been conserved as life evolves into more complex cells. The archaeon *Sulfolobus acidocaldarius* possesses a cell division cycle similar in structure and logic to that of many eukaryotes. While the archaeal ESCRT-III homologues CdvB and CdvB1/B2 are essential for the final stage of cell division, their exact function in the timely remodelling of the cell membrane is elusive. Based on experimental findings, we suggest a mechanistic model of ESCRT-III mediated division in archaea [1]: As cells prepare to divide, they assemble a non-contractile CdvB ring of a fixed diameter at the cell midzone. The CdvB ring acts as a template for the assembly of a CdvB1/B2 polymer which, like force-generating ESCRT-III polymers in other systems, has a small preferred curvature. As a result, the loss of the CdvB scaffold drives rapid contraction of the CdvB1/B2 ring towards its preferred curvature, constricting the membrane as it does so. In order to test the proposed model, we perform coarse-grained molecular dynamics simulations of a large spherical membrane coupled to an elastic filament that can both change its curvature and disassemble. We map the cell division phase space as a function of the CdvB degradation procedure (i.e. instantaneous, sequential, randomised), as well as the intrinsic curvature and the disassembly rate of the ESCRT-III filament [2]. Our results show that cell division is not achieved by contraction of the ESCRT-III filament alone; disassembly is also required. Furthermore, we analyse the furrow constriction in time and find a good qualitative agreement between simulation and experiment. We thereby determine a molecular mechanism by which the timely degradation of one protein, CdvB, triggers the division of a single archaeal cell into two daughter cells.

- [1] G. T. Risa, F. Hurtig, S. Bray, A. E. Hafner, L. Harker-Kirschneck, P. Faull, C. Davis, D. Papatziadou, D. R. Mutavchiev, C. Fan, L. Meneguello, A. A. Pulschen, G. Dey, S. Culley, M. Kilkenny, L. Pellegrini, R. de Bruin, R. Henriques, B. Snjiders, A. Saric, A. Lindås, N. Robinson, and B. Baum, *Proteasome-mediated protein degradation resets the cell division cycle and triggers ESCRT-III-mediated cytokinesis in an archaeon*, submitted (2019)
- [2] A. E. Hafner, L. Harker-Kirschneck, D. Hrynuik, G. T. Risa, B. Baum, and A. Saric, *Physical modelling of ESCRT-III mediated cell division in archaea*, in preparation (2019)