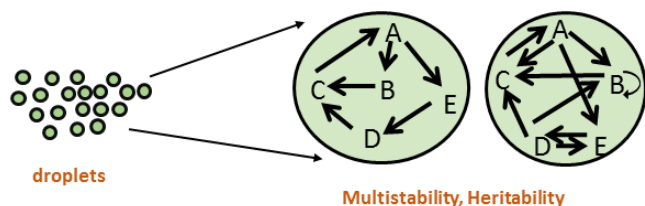


Cooperative self-assembly in origin of life

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Self-assembly and network formation in droplets

Self-assembly must have been an intriguing feature of prebiotic molecules to generate first life-like scenarios on Earth. Both small molecule-based chemical systems as well as RNA are shown to possess self-assembly properties to generate more complex prebiotic-relevant products.¹

Recent works has shown how small RNA fragments of *Azoarcus* group I intron spontaneously self-assemble to form fully-functional ribozymes in a cooperative and autocatalytic fashion,^{2,3} overcoming the hurdle of error-catastrophe in a pure replication-based origin-of-life system.^{4,5} However it is not clear whether such system can show Darwinian-like evolution, specifically heritability, a key feature for any kind of life to develop on Earth. Exploring heritability involves examining how the final state of a system depends on the initial state which has been transmitted from a previous system to later, analogous to genetic material transmission & replication.

Recent works has shown how small RNA

In this work, by exploring huge diversity of initial states and their progress towards the final state when each of them is provided with same material to self-assemble, we are addressing two main questions 1) Multistability: whether these diverse starting initial states leads to development of multiple final stable-states or rather converge to a single universal state? Having multiple states competing for common resource is more origin-of-life relevant than mere dependence of system state on environmental conditions. 2) Heritability: whether the observable effect of initial state on final state can be propagated to next generations. To address these issues we have developed a high-throughput experimental set-up by combining droplet-microfluidics^{6,7} with next-generation sequencing where we will have unprecedented resolution for millions of self-assemblies of *Azoarcus* RNA fragments in droplets. Initial results on a sub-set of conditions show that there is strong effect of initial state on the final state. Now we are exploring full-diversity of *Azoarcus* RNA fragments to identify all the possible states and their heritability.

References

1. Deamer, D. *et al. Philos. Trans. R. Soc. Lond., B, Biol. Sci.* **361**, (2006).
2. Hayden, E. J. & Lehman, N. *Chem. Biol.* **13**, (2006).
3. Vaidya, N. *et al. Nature* **491**, (2012).
4. Eigen, M. & Schuster, P. *Die Naturwissenschaften* **64**, (1977).
5. Kun, A., Santos, M. & Szathmary, E. *Nat. Genet.* **37**, (2005).
6. Pekin, D. *et al. Lab on a chip* **11**, (2011).
7. Ryckelynck, M. *et al. RNA* **21**, (2015).