Howlett, M. G.; Engwerda, A. H. J.; Scanes, R. J. H.; Fletcher, S. P. An Autonomously Oscillating Supramolecular Self-Replicator. Nat. Chem. 2022, 1–6. https://doi.org/10.1038/s41557-022-00949-6.



**Figure 1** | Reaction cycle describing the chemical steps and feedback processes involved in the oscillations: formation, self-assembly, thiol uptake, thiol release, destruction and oxidation.

## Who are the corresponding authors and what are their research areas?

Stephen P. Fletcher - Professor of Chemistry at the University of Oxford, Oxford, UK. His research interests include asymmetric catalysis, origins of life, and synthesis.

## What is the main claim of the article?

This study presents a purely synthetic biomimetic system of supramolecular self-replicators that achieves autonomous, controllable, and sustained oscillations of supramolecular compartments (micellar species). This dynamic behavior (oscillation) of a system across different length scales (coupling molecular and supramolecular feedback processes) is considered to be an important piece in the chemical evolutionary path to cellular biochemical machinery.

### How is it demonstrated?

The existence of oscillations at the molecular level was demonstrated by the concentration profile of the species **1-4** in the aqueous phase (the system was followed by aliquots collected regularly and analyzed by UPLC). In addition, it has been demonstrated that there are oscillations in the quantity of self-assembled micelles in time (determined via interferometric scattering spectroscopy - iSCAT) and that both oscillations (of molecular and supramolecular entities) occur in a concerted manner (**Figure 2**).



Figure 2 | (Top) Molecular-level kinetics, showing oscillations in the concentration of species 3, and a representation of the stages of an oscillation. (Bottom) Supramolecular kinetics using iSCAT images of the oscillating system reveal that the number of micelles oscillate in phase to the concentration of 3.

#### What are the typical experimental conditions?

The biphasic system consists of a stirred aqueous solution of compound 1 (0.058 mmol) and DMAP in the TRIS buffer (0.5 M) and the organic phase (2, 2.87 mmol). Addition of hydrogen peroxide was done by syringe pump.

# Which are the key related papers?

(1) Howlett, M. G.; Scanes, R. J. H.; Fletcher, S. P. Selection between Competing Self-Reproducing

Lipids: Succession and Dynamic Activation. *JACS Au* **2021**, *1* (9), 1355–1361. <u>https://doi.org/10.1021/jacsau.1c00138</u>.

(2) Morrow, S. M.; Colomer, I.; Fletcher, S. P. A Chemically Fuelled Self-Replicator. *Nat. Commun.* 2019, *10* (1), 1011. <u>https://doi.org/10.1038/s41467-019-08885-9</u>.

(3) Leira-Iglesias, J.; Tassoni, A.; Adachi, T.; Stich, M.; Hermans, T. M. Oscillations, Travelling Fronts and Patterns in a Supramolecular System. *Nat. Nanotechnol.* **2018**, *13* (11), 1021–1027. https://doi.org/10.1038/s41565-018-0270-4.