Origins and Phases

Yagmur Denizhan

Electrical and Electronics Engg. Dept., Bogazici University, Turkey

Vefa Karatay

Molecular biologist, independent researcher



Gilbert Simondon (1924-1989)

a Processual Theory of Ontogenesis

metastable preindividual being no phases

Process of Individuation

Phase Separation("se déphaser")

"In a theory of the phases of being, becoming is ... a perpetuated and renewed resolution, an amplifying, incorporating resolution that proceeds via crises, such that its sense is in each of its phases, not at its origin or its end alone."

Individuation in Light of Notions of Form and Information, Gilbert Simondon

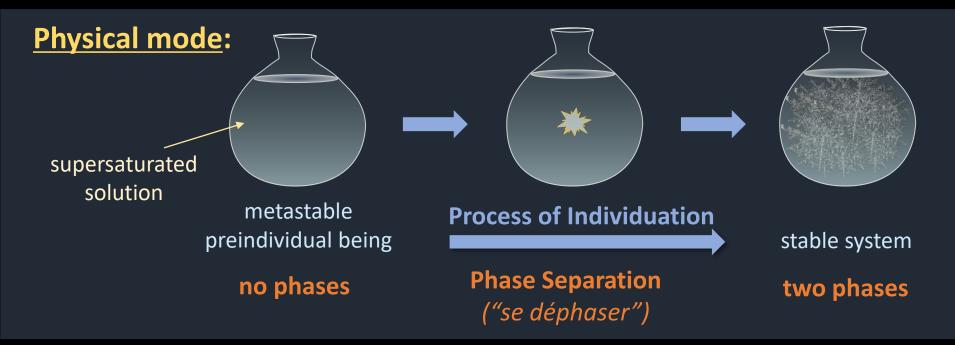


Gilbert Simondon (1924-1989)

a Processual Theory of Ontogenesis

three modes of the process of individuation:

- Physical
- Vital
- Psycho-collective





Gilbert Simondon (1924-1989)

A Processual Theory of Ontogenesis

three modes of the process of individuation:

- **Physical**
- Vital
- **Psycho-collective**

Physical mode: metastable preindividual being

no phases

Process of Individuation

Phase Separation

stable individuated being distinct phases

Vital mode:

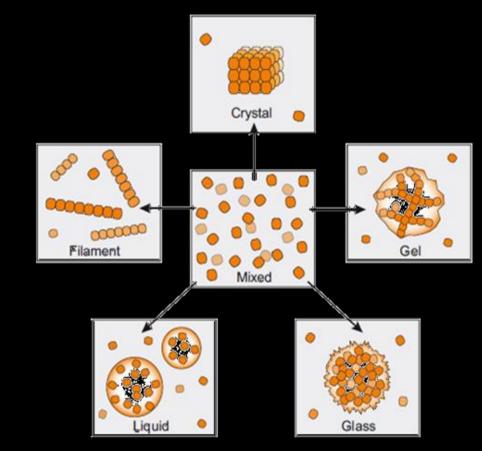
How to extend the notion of phase to living systems?

Psycho-collective mode:

- Dominantly fluidic environment
- More refined categories of phases

From Alberti:

The wisdom of crowds: regulating cell function through condensed states of living matter *Journal of Cell Science* (Review Article) **2017** Sep 1; 130(17), pp.2789-2796



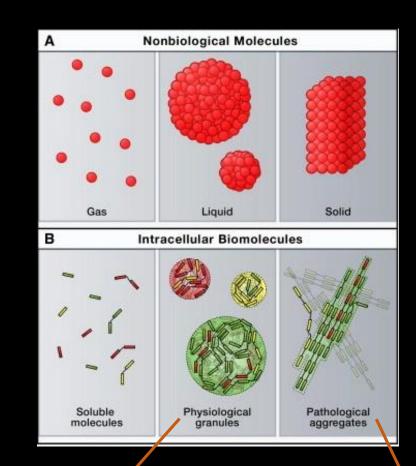
Different condensed states of proteins

- Dominantly fluidic environment
- More refined categories of phases
- Physiological and pathological structures

From Weber and Brangwynne:

Getting RNA and Protein in Phase

Cell, 2012 Jun 8; 149(6), pp. 1188-91



RNPs, membraneless organelles, biomolecular condensates...

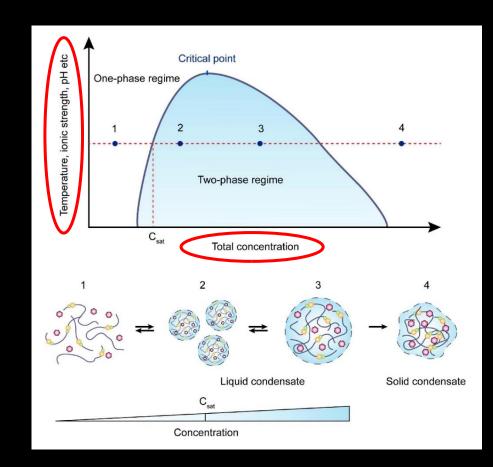
Prions, amyloids ...

- Dominantly fluidic environment
- More refined categories of phases
- Physiological and pathological structures

- Biomolecular condensates
- Membraneless organelles
- Prion-Like Mechanisms (PriLiM)
- Coacervates: dense liquid droplets of macromolecules
- Liquid-Liquid Phase Separation (LLPS)
- Phase Separation (PS)

- Dominantly fluidic environment
- More refined categories of phases
- Physiological and pathological structures
- Several other variables than only temperature and pressure

From Gao, Zhang, Chang, Zhang, Yang, Zhao:
Liquid-Liquid Phase Separation:
Unraveling the Enigma of Biomolecular
Condensates in Microbial Cells
Frontiers of Microbiology (Review Article)
2021 Oct 25; 12:751880



- Dominantly fluidic environment
- More refined categories of phases
- Physiological and pathological structures
- Several other variables than only temperature and pressure
- Feedback-based phase tuning and functionality

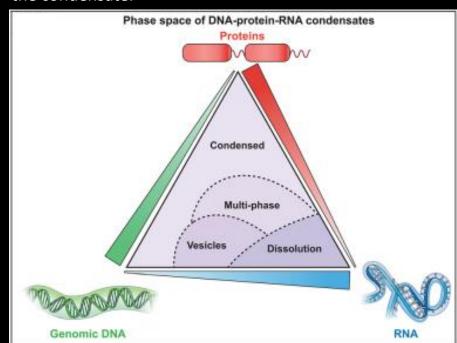
From Feric, Marina and Misteli:

Function Moves Biomolecular Condensates in Phase Space

BioEssays: news and reviews in molecular, cellular and developmental biology **2022**, Vol. 44; 5: e2200001

Structure of transcriptional condensates arising from (im)miscibility of their components.

Ternary DNA-protein-RNA phase diagram indicating various phases depending on the relative composition of the condensate.



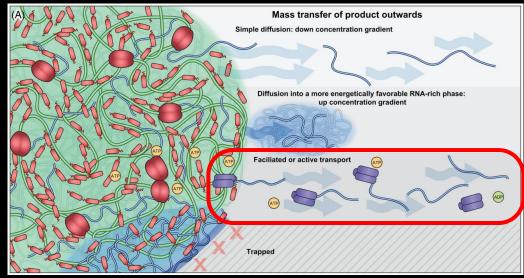
- Dominantly fluidic environment
- More refined categories of phases
- Physiological and pathological structures
- Several other variables than only temperature and pressure
- Feedback-based phase tuning and functionality
- Non-equilibrium activities and active mechanisms

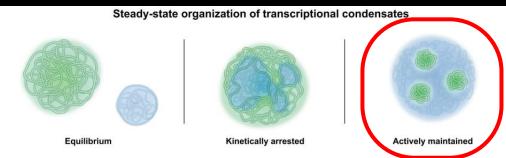
From Feric, Marina and Misteli:

Function Moves Biomolecular Condensates in

Phase Space

BioEssays: news and reviews in molecular, cellular and developmental biology **2022**, Vol. 44; 5: e2200001

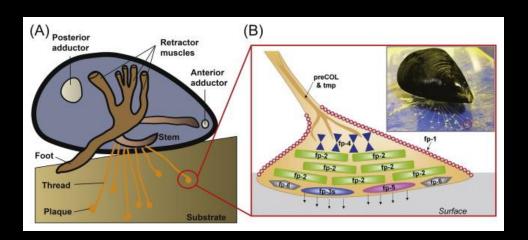




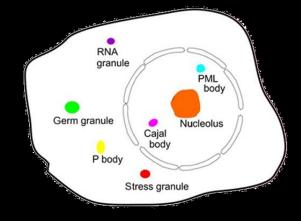
From Astoricchio, Alfano, Rajendran, Temussi, Pastore:

The Wide World of Coacervates: From the Sea to Neurodegeneration

Trends in Biochemical Sciences **2020**; Vol. 45, Issue 8, pp. 706-717



Marine organisms often produce extracellular coacervates to solve the problem of achieving and maintaining strong adhesion on polar surfaces underwater (surface adhesion).



Intracellular coacervates or membraneless organelles have also been observed in a wide spectrum of cell types.

From Wen, Ma:

Phase Separation Drives the Formation of Biomolecular Condensates in the Immune System

Frontiers in Immunology **2022** Nov. 10, 13 986589

cGAMP NF-kB activation Pro-inflammatory Type I interferon **DNA** virus N/P-virus DNA Free mRNA Stress granules STING concentrate ZMYND8/p65 concentrate Viral factories cGAS/DNA concentrate RNA virus N/P-virus RNA

Phase separation in innate immune and inflammatory responses.

Thriving Biophysics Research on Phase Separation

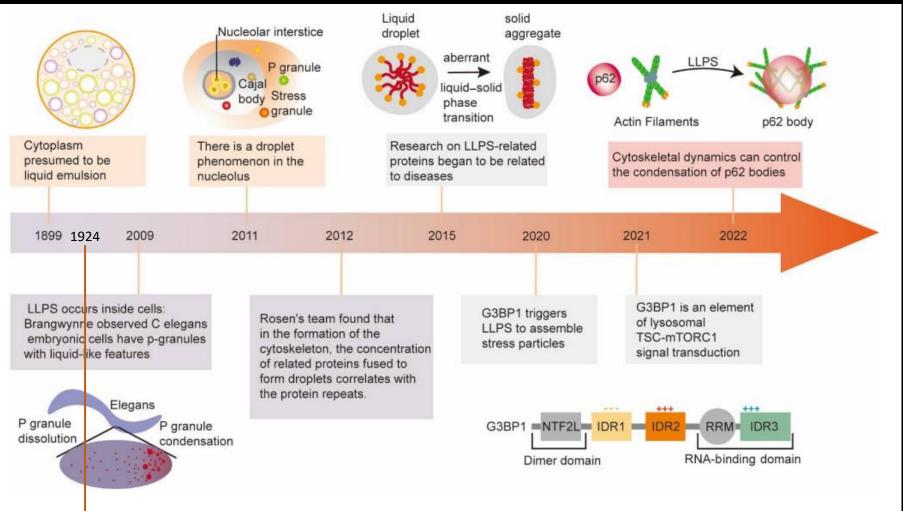


Figure 1. The development history and discovery of the amazing and vital roles of LLPS in biology. Representative milestones sparking tremendous development of LLPS are enumerated in the figure.

Oparin: abiogenesis, LLPS, coacervates

From: Chen, Huai, Mao, Wang, Ru, Qian, Yang: Liquid—Liquid Phase Separation of Biomacromolecules and Its Roles in Metabolic Diseases. *Cells*, 2022, 11, 3023.

From Ghosh, Bose, Tang:

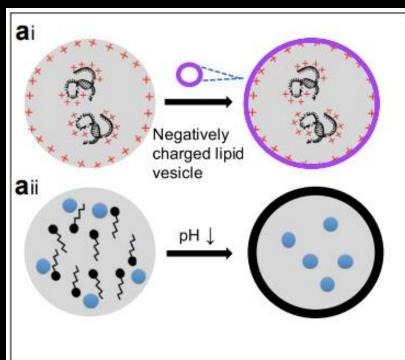
Can coacervation unify disparate hypotheses in the origin of cellular life?

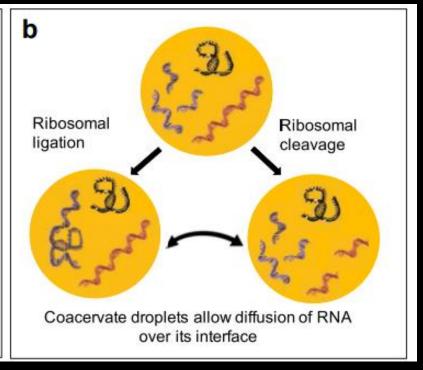
Colloid & Interface Science (Current Opinion) **2021**; Volume 52, 101415, ISSN 1359-0294

Figure 3. The role of coacervation in hypothetical origin of life scenarios:

(a) Coacervation and the lipid world.

(b) Coacervates and the RNA world.



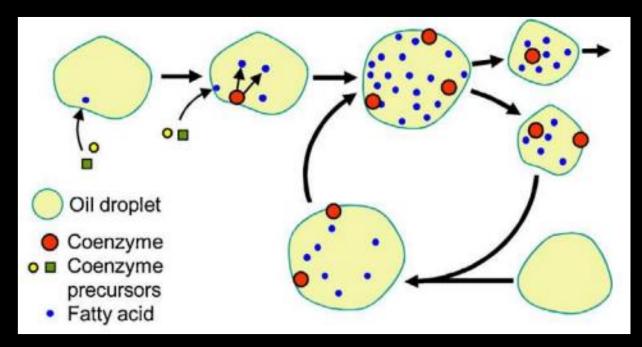


From Sharov:

Coenzyme world model of the origin of life

Biosystems **2016**; 144; pp. 8-17

Scenario of coenzyme self-reproduction on oil droplets: a coenzyme molecule makes the surface hydrophilic via oxidation of hydrocarbons; this change facilitates synthesis of coenzymes from precursors on the surface. Hydrophilic oil droplets easily divide and may coalesce with new oil droplets (i.e., capture new oil resource)



From Ghosh, Bose, Tang:

Can coacervation unify disparate hypotheses in the origin of cellular life?

Colloid & Interface Science (Current Opinion) **2021**; Volume 52, 101415, ISSN 1359-0294

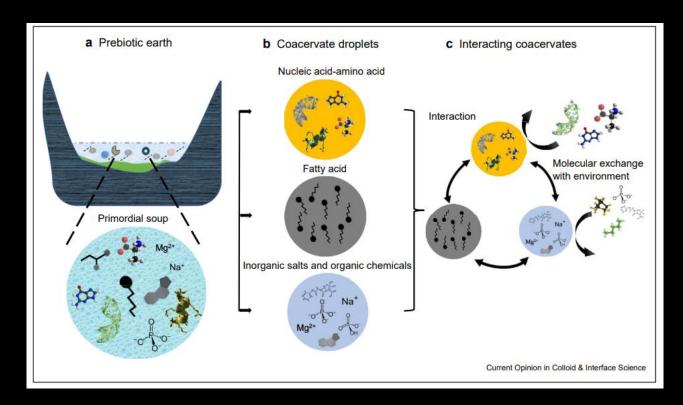


Figure 1. Coacervate formation on prebiotic Earth.

- (a) A primordial soup on prebiotic Earth may have contained biomolecular precursors.
- (b) Coacervation, a phase separation process, could have enriched the biomolecular precursors in the membrane-free coacervate droplets to facilitate reactions.
- (c) Coacervates can concentrate reactants, support enzyme reactions and allow the exchange of ions and small molecules with surrounding media and other compartments.

Matsuo and Kurihara:

Proliferating coacervate droplets as the missing link between chemistry and biology in the origins of life

Nature communications **2021**, **12**, 5487



Gilbert Simondon (1924-1989)

"... the vital mode intervenes by decelerating the physical individuation..."

a prolonged transient state capacity for ongoing origination

Physical mode:

metastable preindividual being

no phases

Process of Individuation

Phase Separation

stable individuated being

distinct phases

Vital mode:

tensions only partially resolved, "preindividual charge" not exhausted

How to extend the notion of phase to living systems?

Psycho-collective mode:



"... becoming is a perpetuated and renewed resolution, an amplifying, incorporating resolution that proceeds via crises..."

Gilbert Simondon (1924-1989)

Physical mode: metastable preindividual being

no phases

Process of Individuation

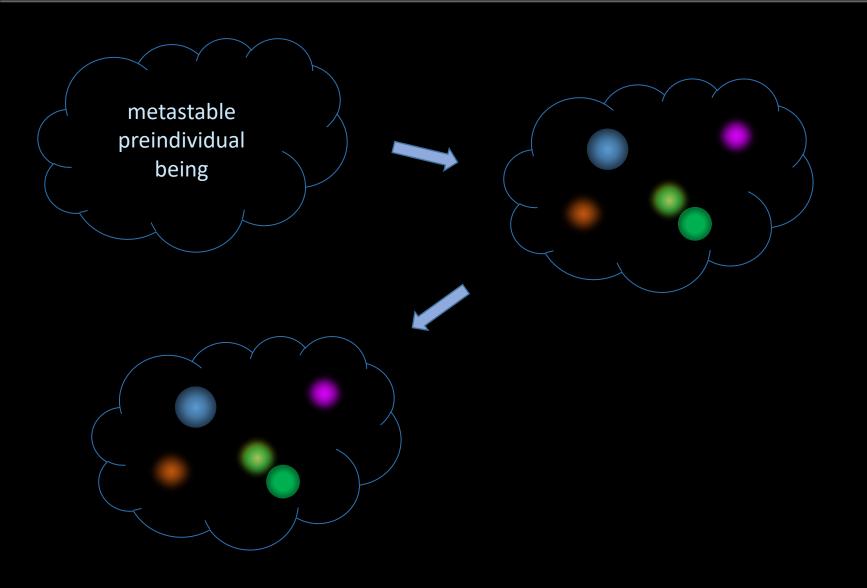
Phase Separation

stable individuated being distinct phases

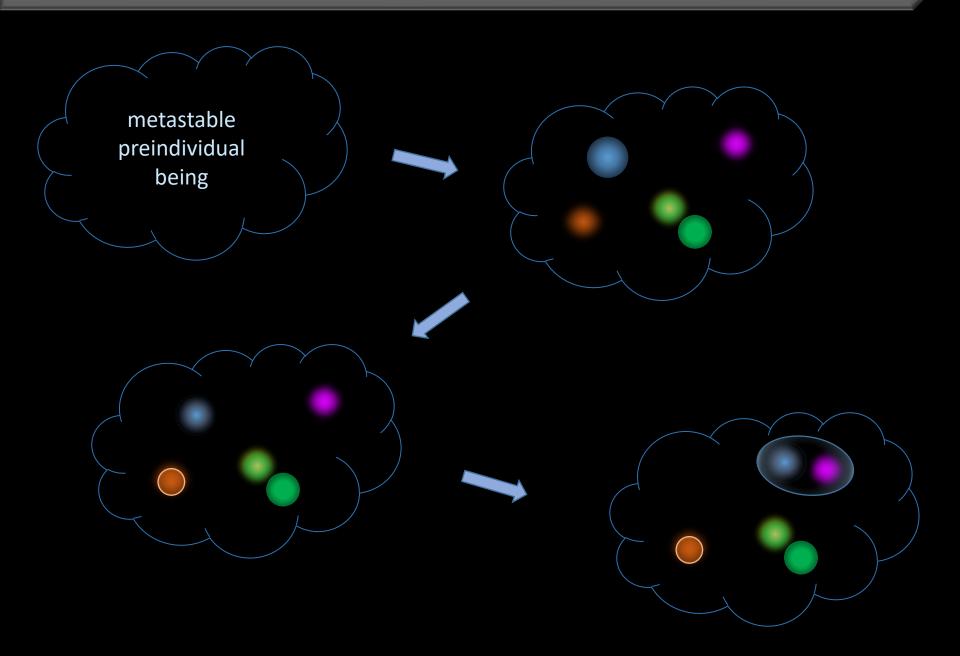
Vital mode: tensions only partially resolved, "preindividual charge" not exhausted How to extend the notion of phase to living systems?

Psycho-collective mode:

Partial Resolutions ...



Partial Resolutions ...



... Code Plurality



Jesper Hoffmeyer & Claus Emmeche

"... the chain of events which sets life apart from non-life.... needs at least two codes: one code for action (behaviour) and one code for memory."

Code-duality and the semiotics of nature, (1991)

Jesper Hoffmeyer & Claus Emmeche

Thank you!

... how different types of codes may have emerged via Phase Separation and Phase-Tuning at successive leaps of vital individuation...

