

Responsible Research and Innovation Exploration of the transcriptional landscapes of culturable and nonculturable bacteria using in vitro transcription emulators

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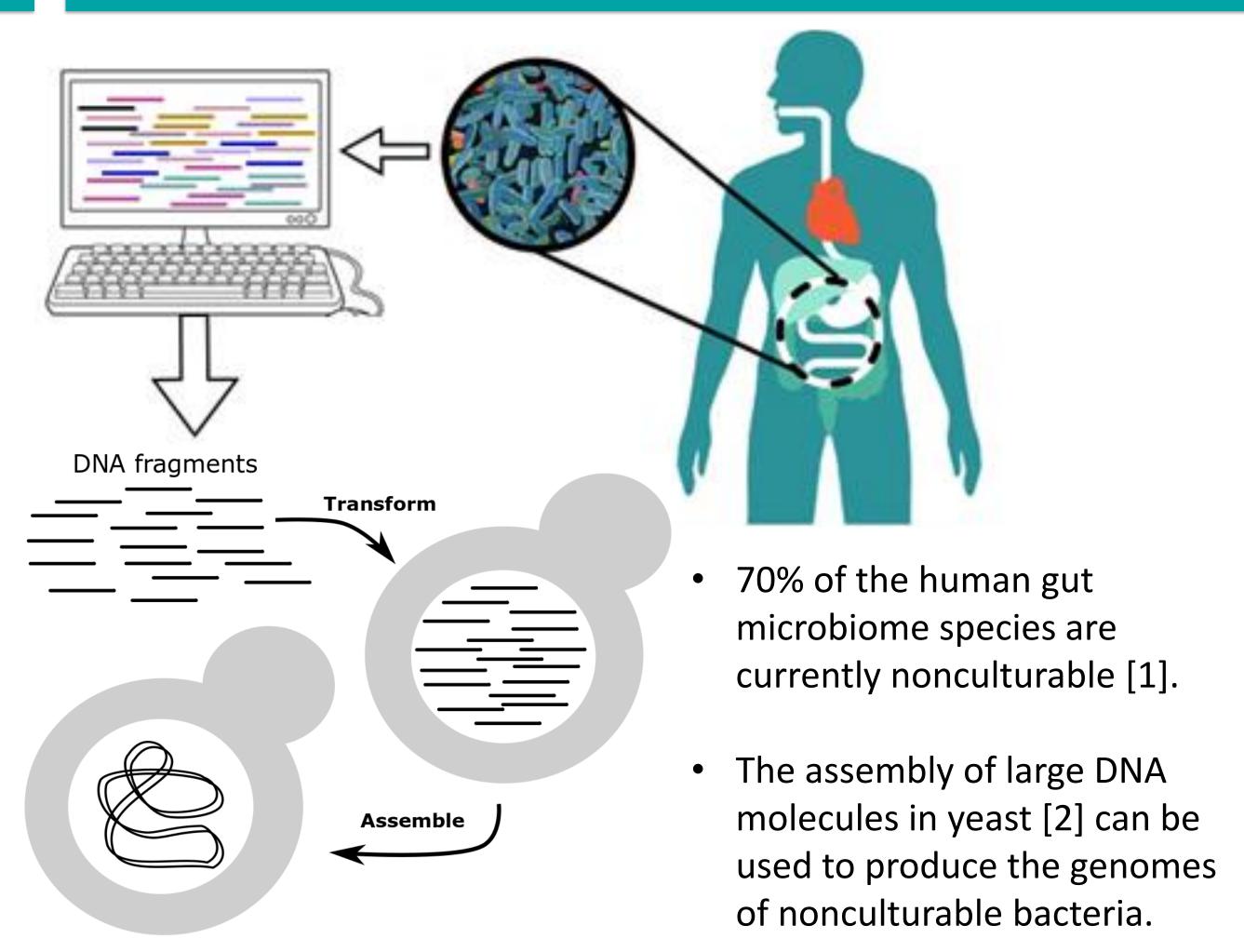


Exploration of the transcriptional landscapes of culturable and nonculturable bacteria using in vitro transcription emulators

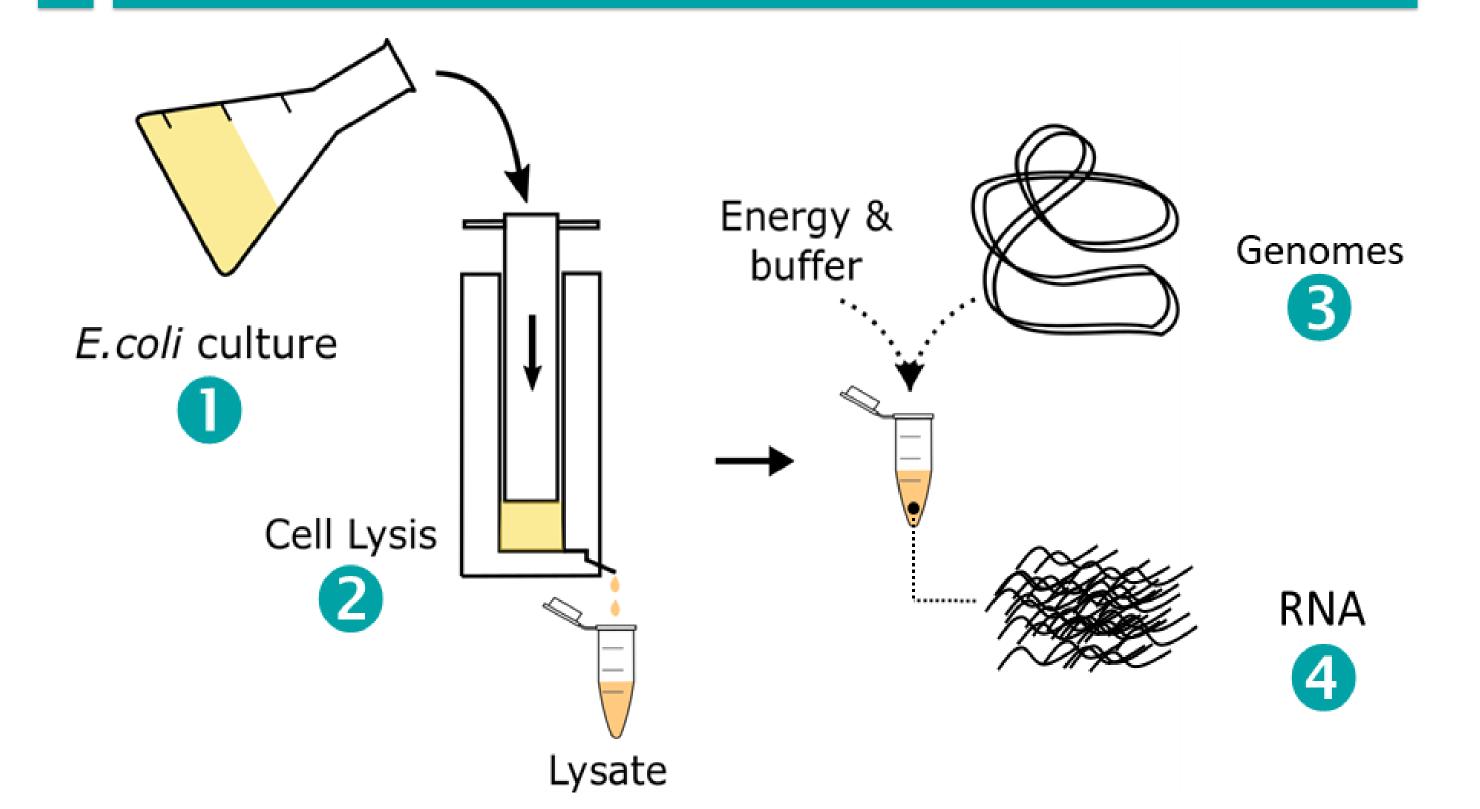
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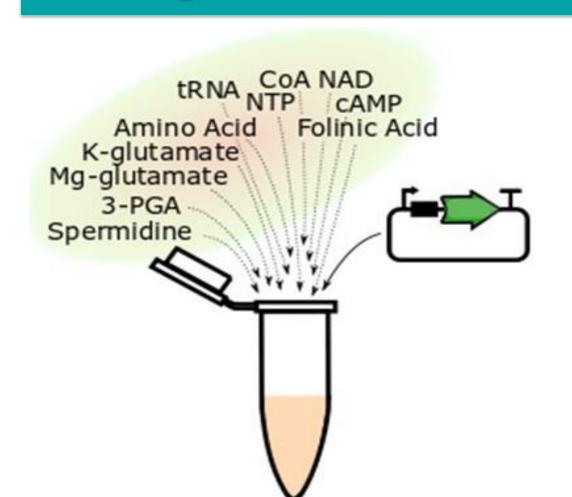
Background: Human gut microbiome



Genome expression in cell-free

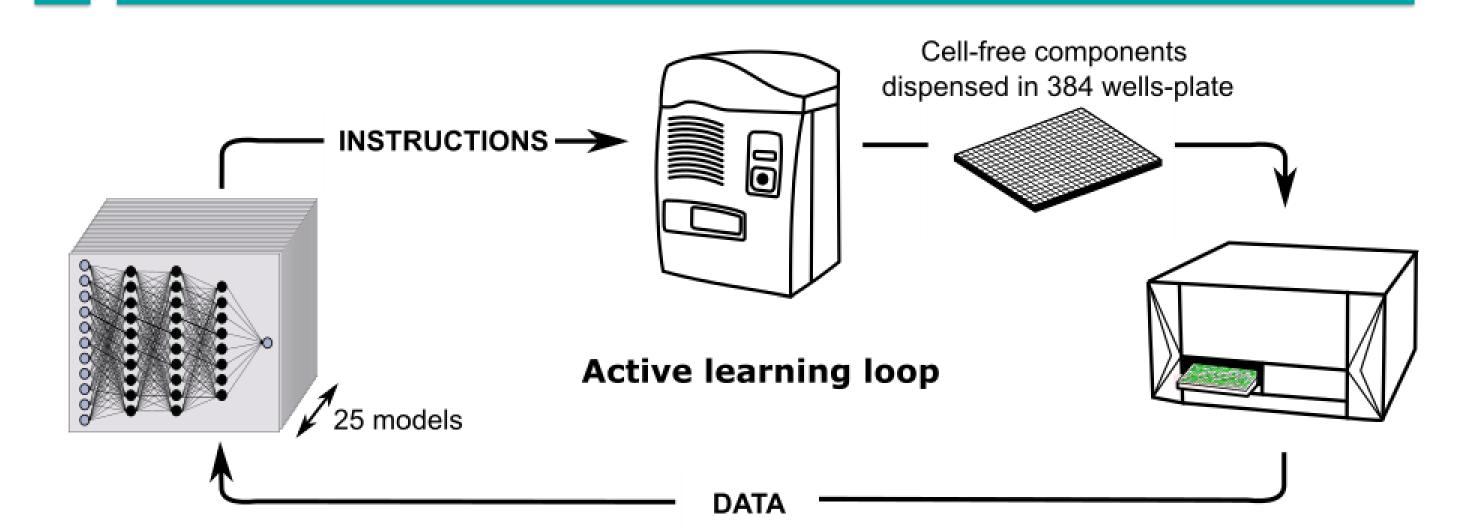


Background: cell-free systems



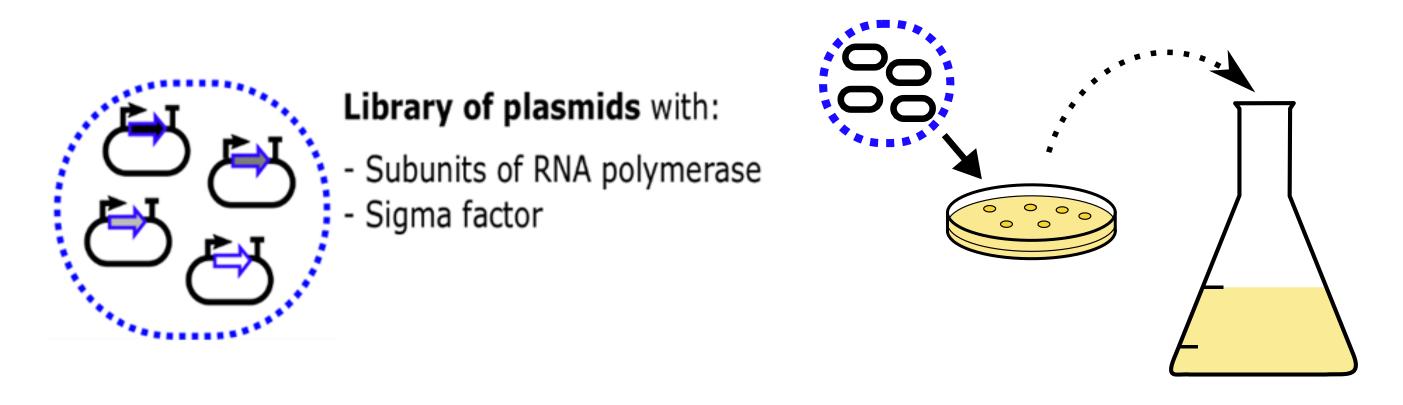
- Cell-free systems are based on cell lysates to express genes of interest by adding DNA.
- The lysate is supplemented with multiple chemical compounds to maximize protein production [3].

Active learning for cell-free optimization



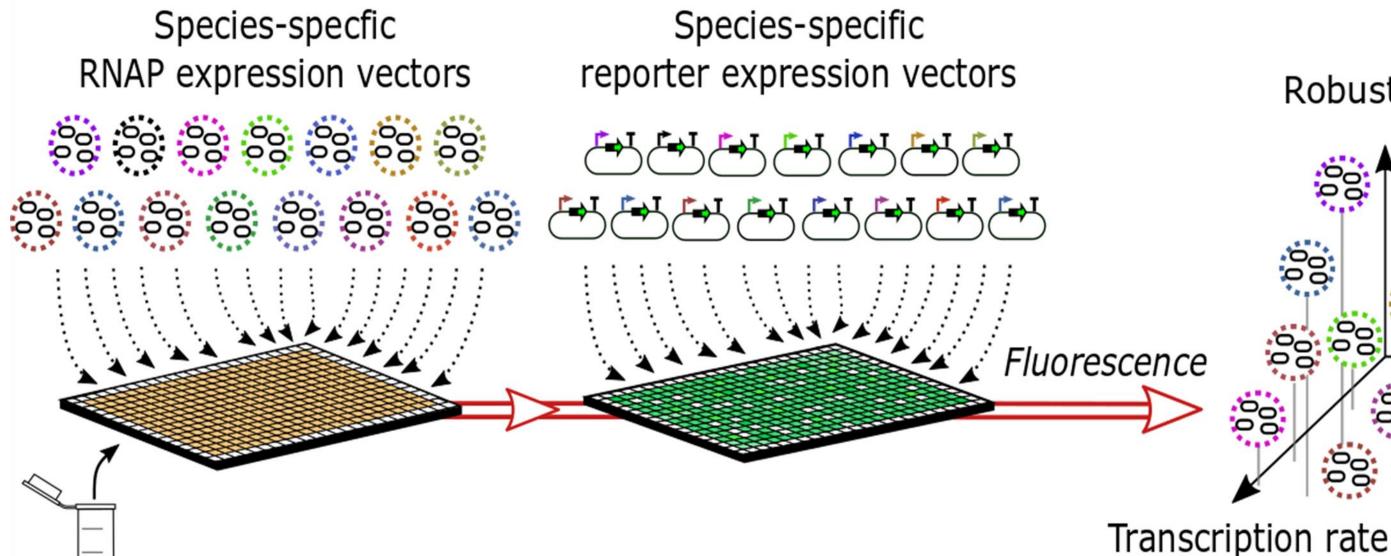
- A machine learning approach coupled with a liquid-handling robot has been used to maximized protein production in cell-free [4].
- RNA production can also be maximized by exploring a combinatorial space of ~1,000,000 cell-free compositions.

Production of RNAP from nonculturable bacteria

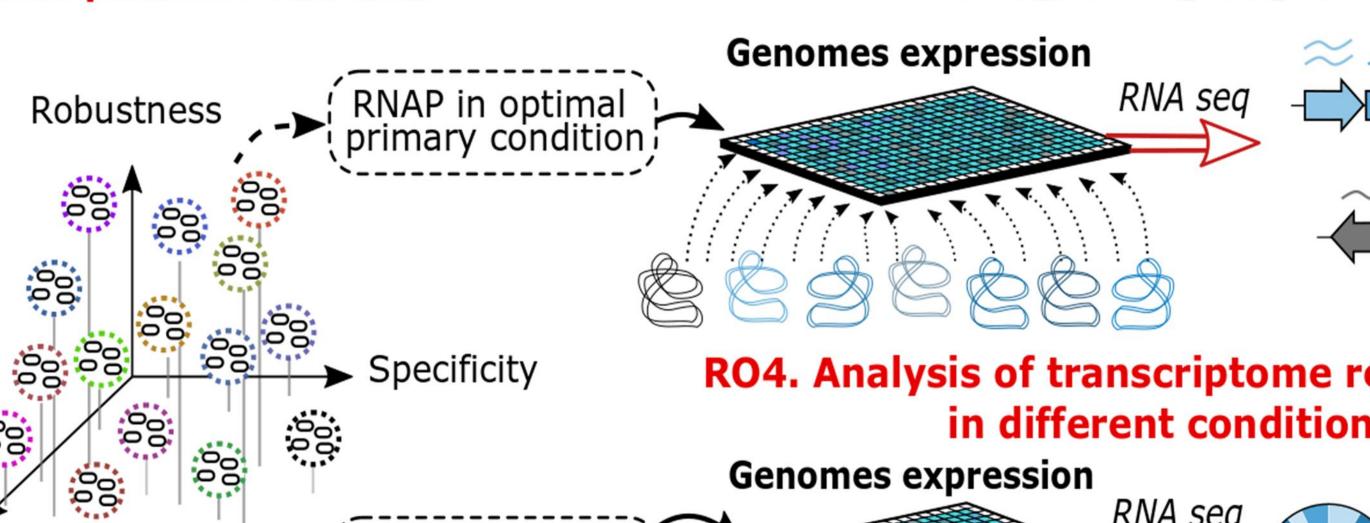


Research objectives of the project

RO1. Development of an in vitro platform for species-specific gene expression



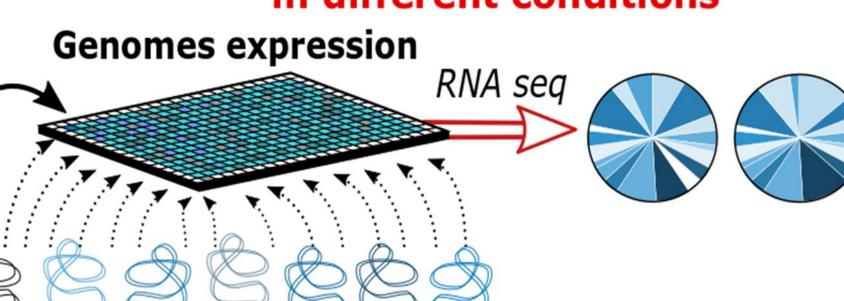
RO2. Characterization of transcription machineries



RO4. Analysis of transcriptome reorganisation in different conditions

RO3. Identification of new transcription units

and regulatory sequences



- ☐ What are the full **transcriptional landscapes of nonculturable** bacteria of the human gut?
- ☐ To what extent nonculturable bacteria in the human gut can reorganize their genomes activity?
- ☐ Which **regulations and molecular mechanisms** are involved in such reorganization?

References

- Almeida, A. et al., 2021. A unified catalog of 204,938 reference genomes from the human gut microbiome. Nature biotechnology, 39(1), pp.105-114.
- 2. Gibson, D.G., et al., 2008. One-step assembly in yeast of 25 overlapping DNA fragments to form a complete synthetic Mycoplasma genitalium genome. Proceedings of the National Academy of Sciences, 105(51), pp.20404-20409
- 3. Sun, Z.Z. et al., 2013. Protocols for implementing an Escherichia coli based TX-TL cell-free expression system for synthetic biology. JoVE (Journal of Visualized Experiments), (79), p.e50762.
- Borkowski, O. et al., 2020. Large scale active-learning-guided exploration for in vitro protein production optimization. *Nature communications*, 11(1), p.1872.