SMART frameworks for bioreactor assessment

PhD student:	Héctor Aroldo Maldonado de León
Email:	h.a.maldonadodeleon@tudelft.nl
Supervisor:	Dr. Cees Haringa
Promotor:	Dr. Adrie Straathof
Project term:	September 2022 – August 2026



Description

Computational Fluid Dynamics (CFD) has proved to be an effective tool for engineers to design and assess the performance of different pieces of equipment. In biotechnology, this tool coupled with a kinetic model allows us to estimate not only the flow patterns but the gradients of chemical species within bioreactors (i.e., Computational Reaction Dynamics) [1-3]. These gradients may play an important role in determining the efficiency of the process since the microorganisms will adapt themselves upon the fluctuations within their environment [4, 5]. Mitigating those fluctuations is the basis of the down-top design approach. However, its extensive implementation is hindered by the inherent demands of time and computational resources of the traditional CFD-CRD software packages. Various methods have been proposed as CFD substitutes, from compartment models to neural networks [6 - 10].



Figure 1. Iterative design process aided by a fast reconstruction of fields within a bioreactor.

In my research project, I explore how computationally less demanding models can be employed towards studying bioprocesses. We will develop a framework in which hydrodynamic- and kinetic models are combined for rapid assessment of different bioreactor configurations. Once established, the developed framework will be applied towards development and downscaling of novel bioprocesses in the framework of <u>the zero emission</u> <u>biotechnology program</u>.

References

- [1] C. Haringa, A. T. Deshmukh, R. F. Mudde, and H. J. Noorman, "Euler-Lagrange analysis towards representative down-scaling of a 22m3 aerobic S. cerevisiae fermentation," Chemical Engineering Science, vol. 170, pp. 653–669, 2017, doi: 10.1016/j.ces.2017.01.014.
- [2] C. Haringa et al., "Computational fluid dynamics simulation of an industrial P. chrysogenum fermentation with a coupled 9-pool metabolic model: Towards rational scale-down and design optimization," Chemical Engineering Science, vol. 175, pp. 12–24, Jan. 2018, Accessed: Apr. 03, 2018. [Online]. Available: https://www.eciapsedirect.com/science/article/pii/S00002E001720E742

https://www.sciencedirect.com/science/article/pii/S0009250917305742

[3] Puiman, L., Almeida Benalcázar, E., Picioreanu, C., Noorman, H. J., & Haringa, C. (2023). Downscaling Industrial-Scale Syngas Fermentation to Simulate Frequent and Irregular Dissolved Gas Concentration Shocks. Bioengineering, 10(5), 518. MDPI AG. Retrieved from http://dx.doi.org/10.3390/bioengineering10050518

- [4] Lara, A.R.; Galindo, E.; Ramírez, O.T.; Palomares, L.A. Living with heterogeneities in bioreactors. *Mol. Biotechnol.* 2006, *34*, 355–381
- [5] Haringa, C., Mudde, R.F., Noorman, H.J., "From industrial fermentor to CFD-guided downscaling: what have we learned?", Biochemical Engineering Journal, vol. 140, pp. 57-71, Dec. 2018, https://doi.org/10.1016/j.bej.2018.09.001
- [6] M. Raissi, P. Perdikaris, and G. E. Karniadakis, "Physics-informed neural networks: A deep learning framework for solving forward and inverse problems involving nonlinear partial differential equations," Journal of Computational Physics, vol. 378, pp. 686–707, Feb. 2019, doi: 10.1016/J.JCP.2018.10.045.
- [7] Hennigh, O. et al. (2021). NVIDIA SimNet[™]: An AI-Accelerated Multi-Physics Simulation Framework. In: Paszynski, M., Kranzlmüller, D., Krzhizhanovskaya, V.V., Dongarra, J.J., Sloot, P.M. (eds) Computational Science – ICCS 2021. ICCS 2021. Lecture Notes in Computer Science(), vol 12746. Springer, Cham. https://doi.org/10.1007/978-3-030-77977-1_36
- [8] A. Delafosse et al., "CFD-based compartment model for description of mixing in bioreactors," Chemical Engineering Science, vol. 106, pp. 76–85, 2014, Accessed: Jan. 26, 2014. [Online]. Available: <u>http://www.sciencedirect.com/science/article/pii/S0009250913007690</u>
- [9] T. Tajsoleiman, R. Spann, C. Bach, K. v. Gernaey, J. K. Huusom, and U. Krühne, "A CFD based automatic method for compartment model development," Computers and Chemical Engineering, vol. 123, pp. 236–245, Apr. 2019, doi: 10.1016/j.compchemeng.2018.12.015.
- [10] G. Nadal-Rey et al., "Development of dynamic compartment models for industrial aerobic fedbatch fermentation processes," Chemical Engineering Journal, p. 130402, May 2021, doi: 10.1016/j.cej.2021.130402.