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Single Cell Response to Time-dependent Stimuli using a Microfluidic Bioreactor¹ ERIC M. JOHNSON-CHAVARRIA, Center for Biophysics and Computational Biology, University of Illinois at Urbana-Champaign, UTSAV AGRAWAL, MELIKHAN TANYERI, Department of Chemical and Biomolecular Engineering, University of Illinois at Urbana-Champaign, THOMAS E. KUHLMAN, Department of Physics, University of Illinois at Urbana-Champaign, CHARLES M. SCHROEDER, Department of Chemical and Biomolecular Engineering, University of Illinois at Urbana-Champaign — Cellular adaptation is critical for survival under uncertain or dynamic environmental conditions. Recent studies have reported the ability of biological systems to implement low-pass filters to distinguish high frequency noise in environmental stimuli from lower frequency input signals, yet we still lack a complete understanding of this phenomenon. In this work, we report a microfluidic-based platform for single cell analysis that provides dynamic control over periodic, time-dependent culture media. Single cells are confined in free solution by the sole action of gentle fluid flow, thereby enabling non-perturbative trapping of cells for long time scales. In this way, our microfluidic-based technique provides the ability to control external stimuli with precise methods while observing nonadherent cells over long timescales. Using this approach, we observed intranucleoid diffusion of genetic repressor proteins released from a chromosomal binding array. Overall, this microfluidic approach provides a direct method for sustaining periodic environmental conditions, measuring growth rates, and detecting gene expression of single cells in free solution.

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