

DEEPWEP

FEASIBILITY ANALYSIS OF A NON STIRRED MINIATURE BIOREACTOR

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Motivation / Introduction

The scale-up of bioprocesses is a **time-intensive** task and constitutes a major part in bioprocess development. To successfully transfer knowledge from the lab-scale to the pilot- and industrial scale, **reliable scale-up parameters** are required. The **volumetric oxygen- and energy inputs** represent such critical scale-up parameters [1-3]. In the lab-scale, high-throughput minibioreactor systems are used for bioprocess development. A high number of parallel reactors coupled with automated liquid handling and small volumes enable a **cost-efficient** generation of relevant process data.

Modern mini-bioreactor systems rely on impellers, magnetic stirrers or agitation to supply oxygen to the culture medium. Disadvantages of these systems include **temperature shifts** caused by changes of stirrer speed, **shear force gradients** and **high energy inputs** [4]. To overcome these drawbacks, the aim of this work was to assess the miniaturization capabilities of a **non-stirred 3D-printed bioreactor system** and its use in 12 mL scale.

Concept and Results

3D-printed non-stirred



conventional stirred

COMPARABLE $k_L a$ -VALUES

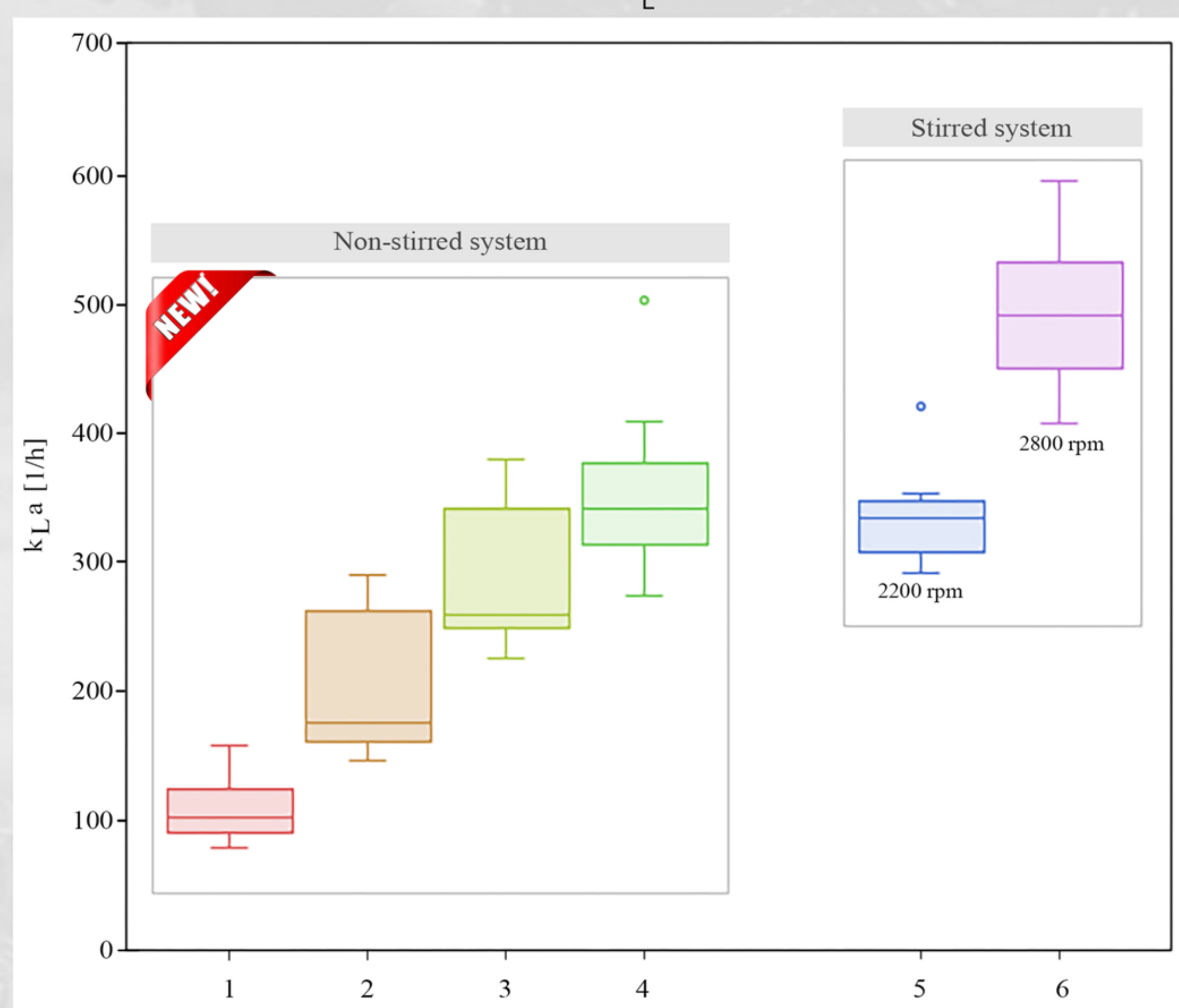


Figure 1. Comparative presentation of $k_L a$ -values for non-stirred and stirred systems with aeration of ambient air in a 12 mL minibioreactor. Each boxplot represents a total of nine measurements. The results are grouped in experiments from 1 - 6.

Non-stirred System: (1) Mean $k_L a$ of $109 \pm 22 \text{ h}^{-1}$; (2) Mean $k_L a$ of $203 \pm 53 \text{ h}^{-1}$; (3) Mean $k_L a$ of $287 \pm 53 \text{ h}^{-1}$; (4) Mean $k_L a$ of $352 \pm 56 \text{ h}^{-1}$; Stirred system: (5) 2200 rpm with a mean $k_L a$ of $333 \pm 29 \text{ h}^{-1}$; (6) 2800 rpm with a mean $k_L a$ of $443 \pm 38 \text{ h}^{-1}$.

LOWER ENERGY INPUT

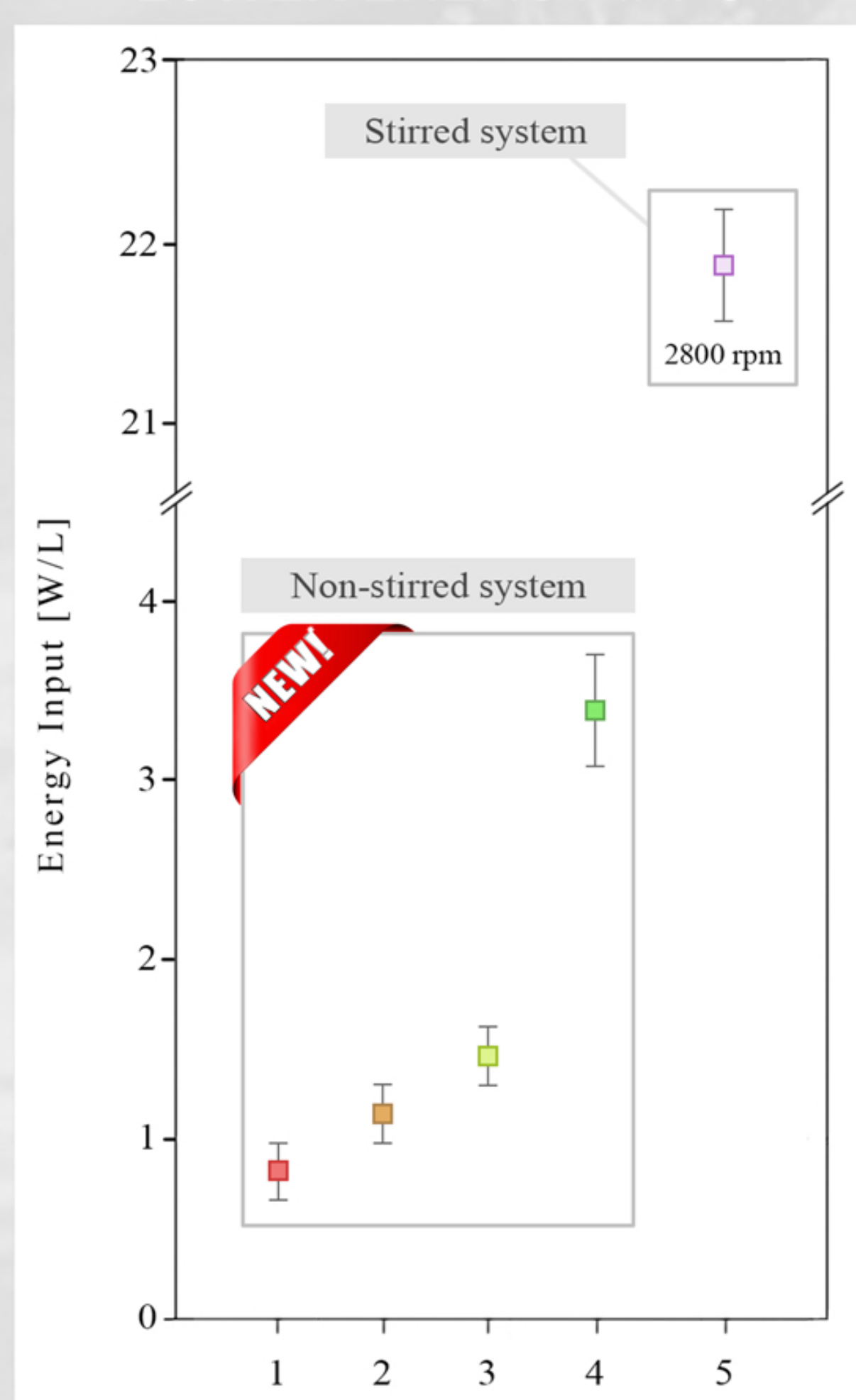


Figure 2. Comparative presentation of volumetric energy input for non-stirred and stirred systems in a 12 mL minibioreactor. Each data point represents a total of three measurements. The results are grouped in experiments from 1 - 5.

Non-stirred System: (1) Mean energy input of $0.81 \pm 0.15 \text{ W L}^{-1}$; (2) Mean energy input of $1.13 \pm 0.16 \text{ W L}^{-1}$; (3) Mean energy input of $1.45 \pm 0.17 \text{ W L}^{-1}$; (4) Mean energy input of $3.37 \pm 0.31 \text{ W L}^{-1}$; Stirred system: (5) 2800 rpm with a mean energy input of $22.00 \pm 1.68 \text{ W L}^{-1}$.

LOWER TEMPERATURE FLUCTUATIONS

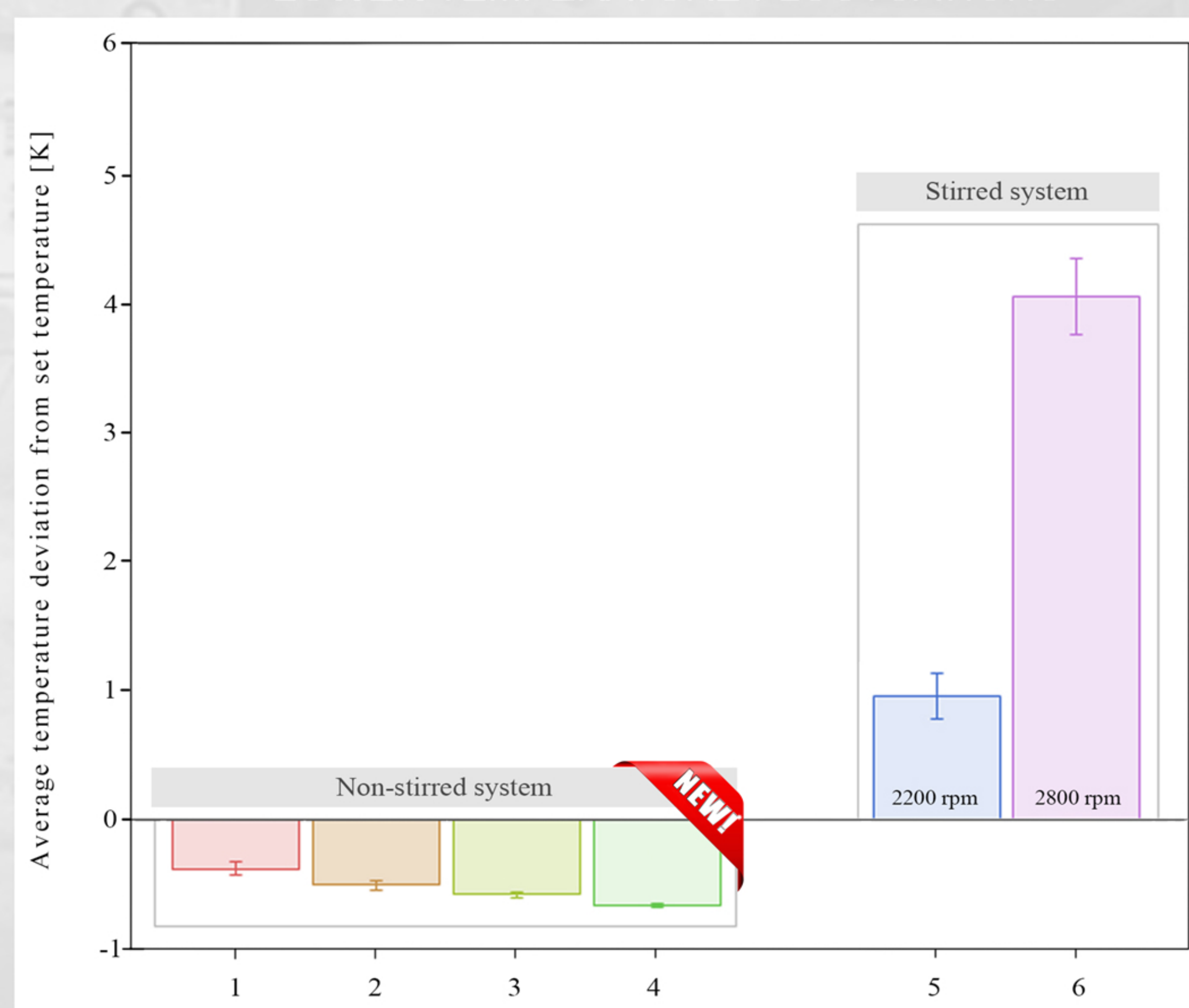


Figure 3. Average temperature deviation from set-temperature of a non-stirred system and a stirred system during $k_L a$ -experiments in a 12 mL minibioreactor. The results are grouped in experiments from 1 - 6.

Each group represents a total of nine measurements. Non-stirred System: (1) Temperature deviation of $-0.38 \pm 0.05 \text{ K}$; (2) Temperature deviation of $-0.51 \pm 0.03 \text{ K}$; (3) Temperature deviation of $-0.58 \pm 0.02 \text{ K}$; (4) Temperature deviation of $-0.66 \pm 0.01 \text{ K}$; Stirred system: (5) Temperature deviation of $0.96 \pm 0.17 \text{ K}$; (6) Temperature deviation of $4.04 \pm 0.29 \text{ K}$.

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Let's bring this system to market.

Conclusion and Outlook

In this study, the feasibility of a non-stirred minibioreactor could be demonstrated. Temperature profiles and volumetric energy inputs of the non-stirred system indicate its beneficial use in high-throughput systems compared to stirred systems. To validate these results, a cultivation in the developed system is proposed for future research. This work contributes to the development of **novel minibioreactor systems** and increases the variety of reactor types which can be used in high-throughput systems.

Quickie? Advance minibioreactors in 8 min!



<https://sprw.io/stt-a2ef30>

Acknowledgments / References

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- [3] Lübbert, A.; Jørgensen, S.B. Bioreactor performance: a more scientific approach for practice. *Journal of Biotechnology* 2001, 85, 187 – 212.
- [4] Bareither, R.; Pollard, D. A review of advanced small-scale parallel bioreactor technology for accelerated process development: Current state and future need. *Biotechnology Progress* 2011, 27, 2–14.