

Basic approach to design and optimize a PDO fermentation process

Technology for biobased products

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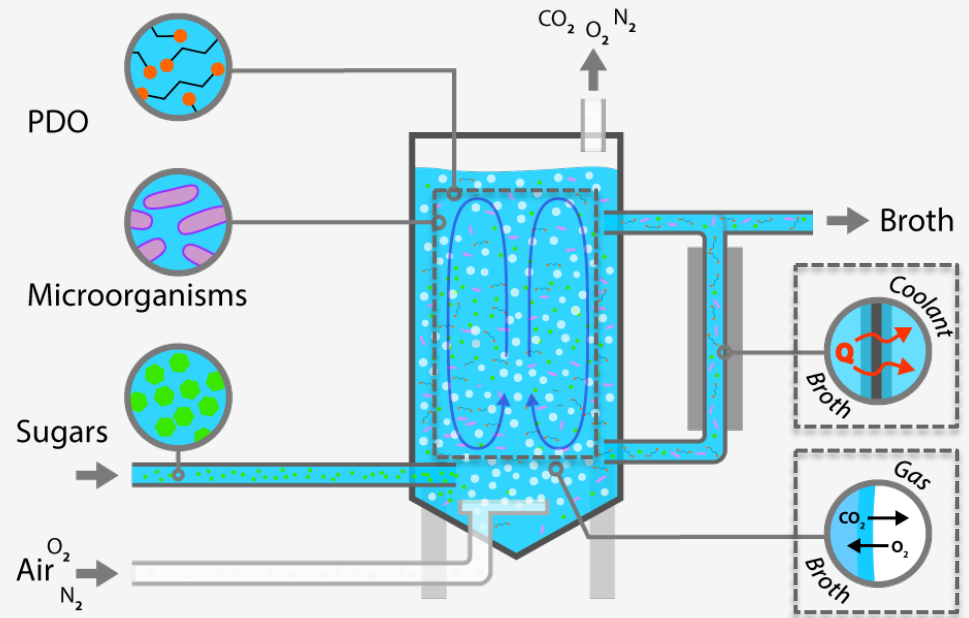
Design approach

- Process reaction based on black box model (week 3)
- Bioreactor type and mode of operation
 - STR, BC or ALR
 - Batch, continuous or fed-batch
- Reactor volume is determined by rate-limiting transport step

Large-scale design: Comparison of the transport phenomena

What is the rate-limiting transport step?

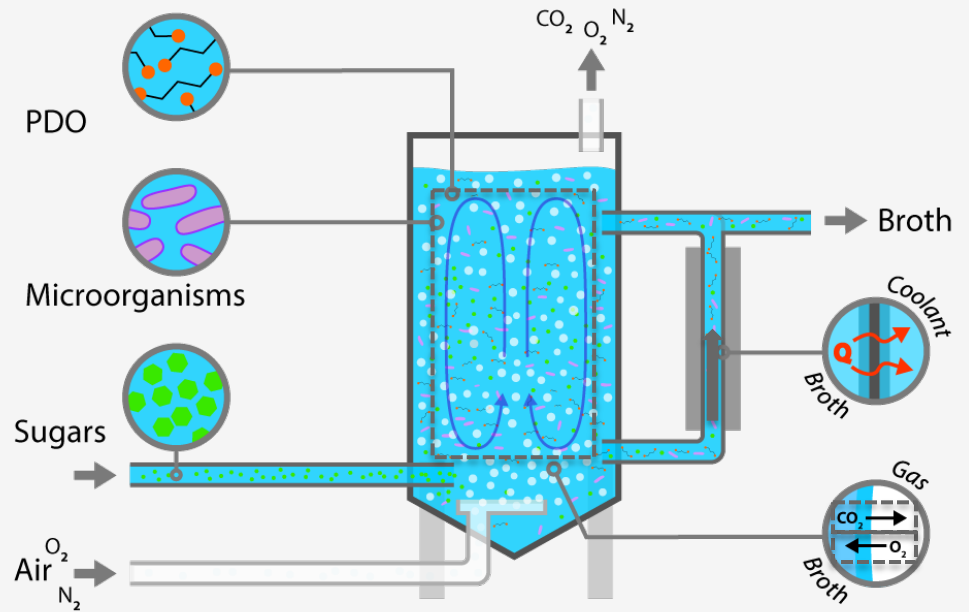
- Oxygen transfer
→ minimize c_o gradients
- CO_2 removal
→ minimize CO_2 inhibition
- Heat removal
→ minimize T shocks
- Mixing
→ minimize c_s gradients



Defining what is most likely limiting factor

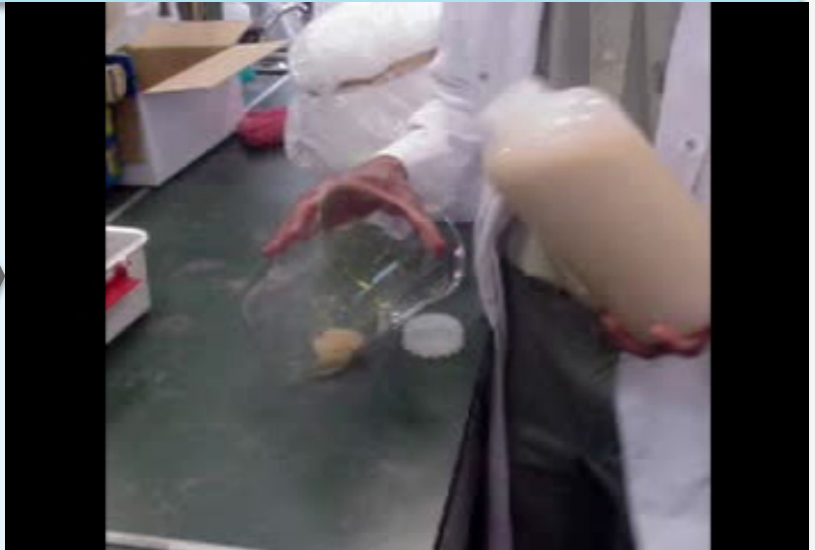
PDO Case, transport terms determined in previous units:

- Required $T_{N,0} = 193 \text{ mol}/(\text{m}^3 \cdot \text{h})$
Actual $T_{N,0} = 135 \text{ mol}/(\text{m}^3 \cdot \text{h})$
- $c_c = 7.0 \text{ mol}/\text{m}^3$
- $F_{\text{loop}} = 2.46 \text{ m}^3/\text{s}$
- $t_{\text{mix}} = 69 \text{ s}$



Note: this all assumes low viscosity

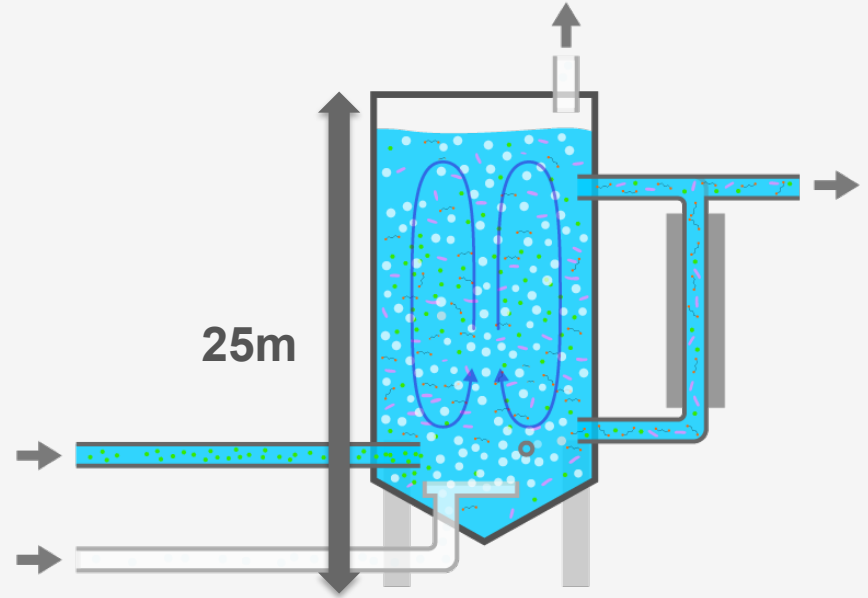
Always design/select non-filamentous microorganisms so that the broth viscosity stays low. If not, transport rates can be 10-100× lower



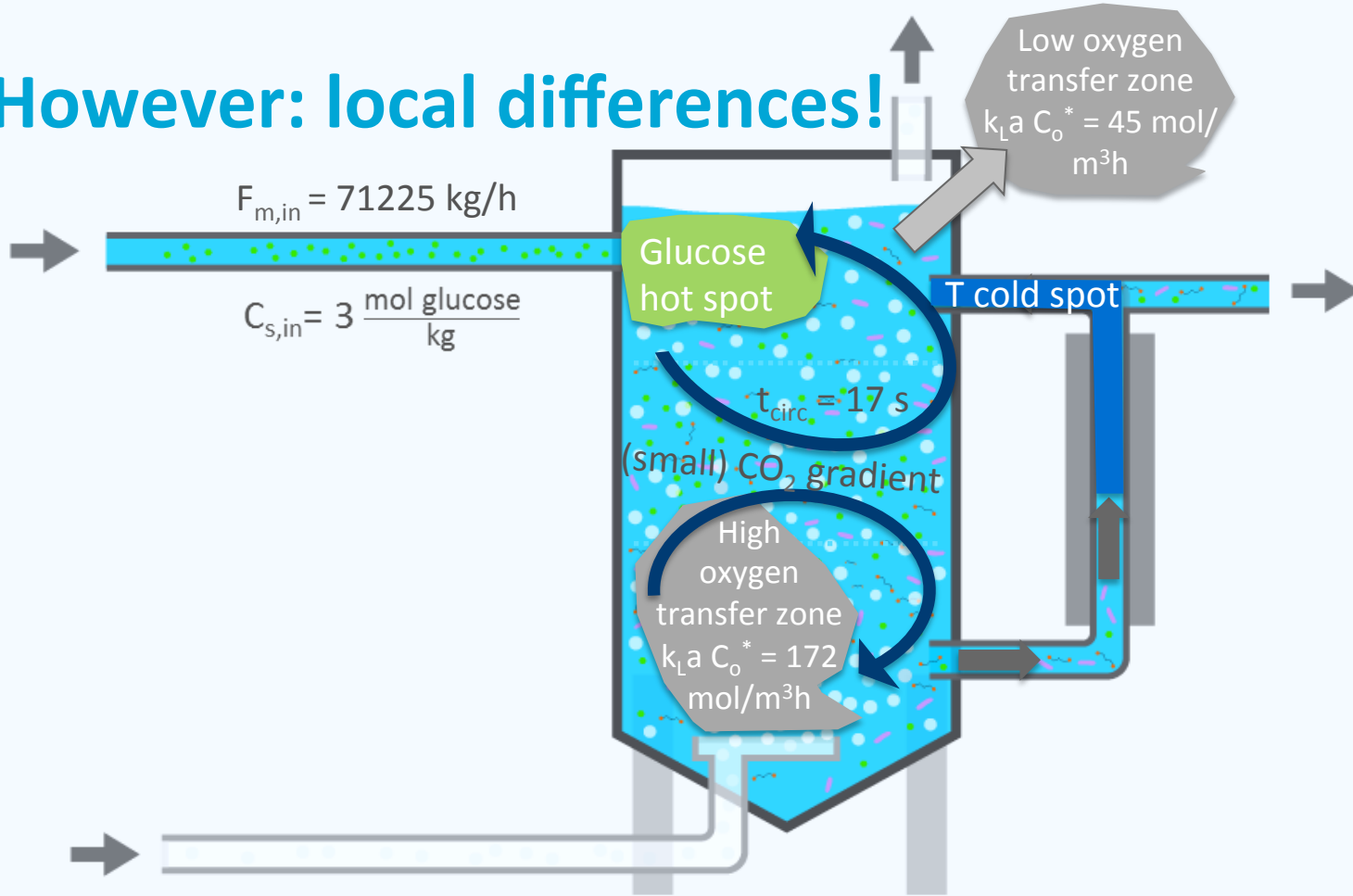
PDO performance

Economy performance indicators

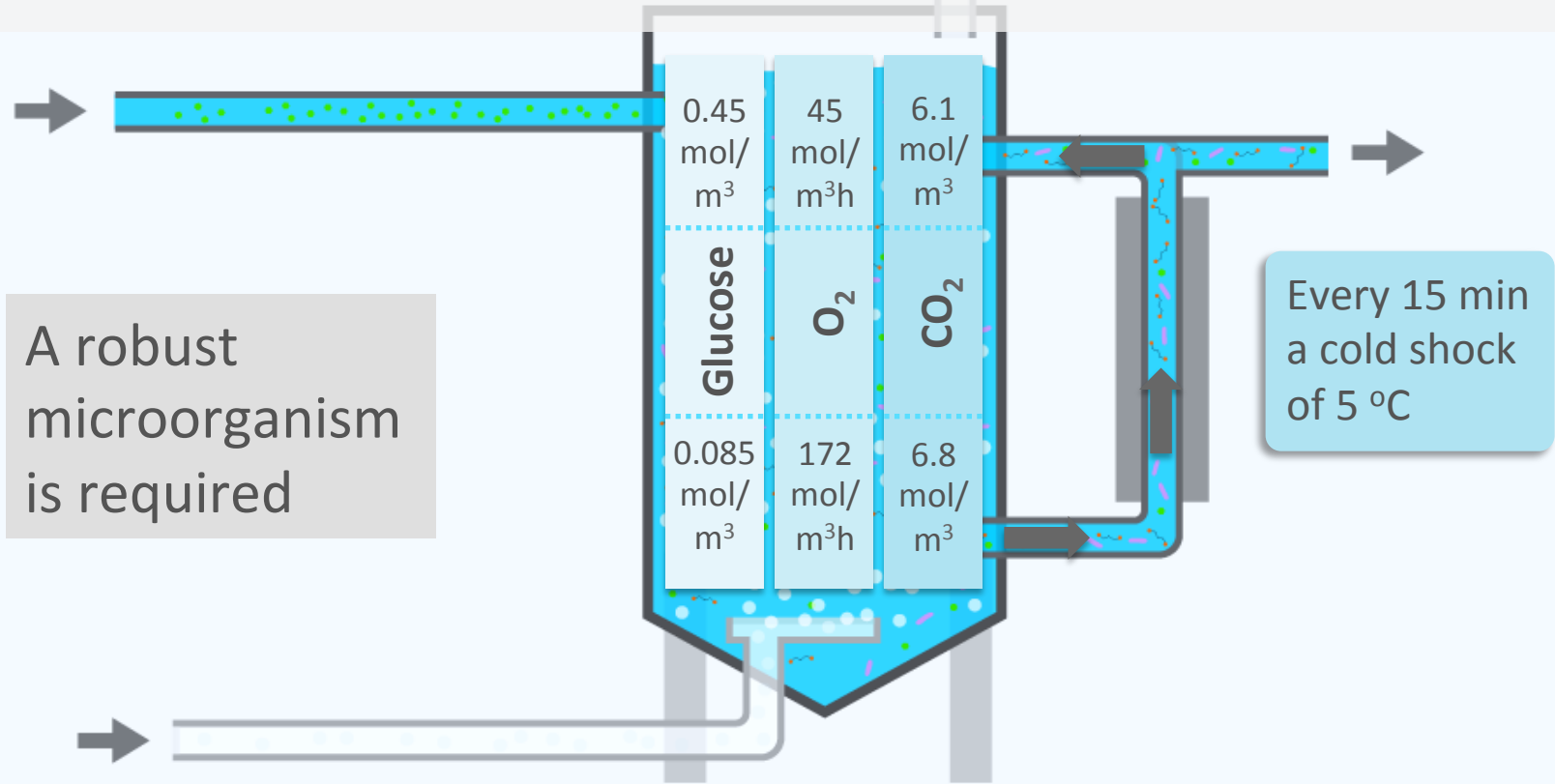
Titer kg PDO / tonne broth	228
Rate kg PDO/h / tonne broth	5.6
Yield kg PDO / kg glucose	0.33
(theoretical maximal)	(0.528)



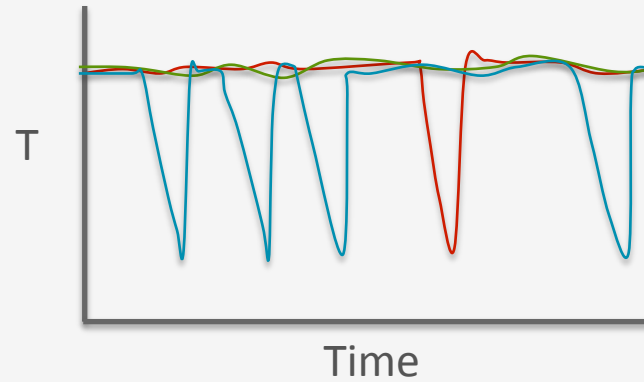
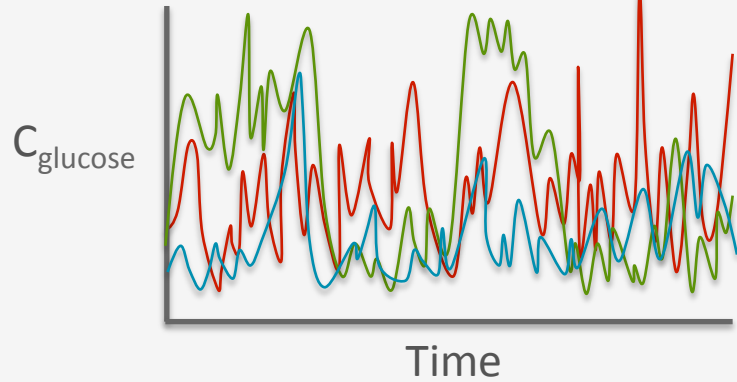
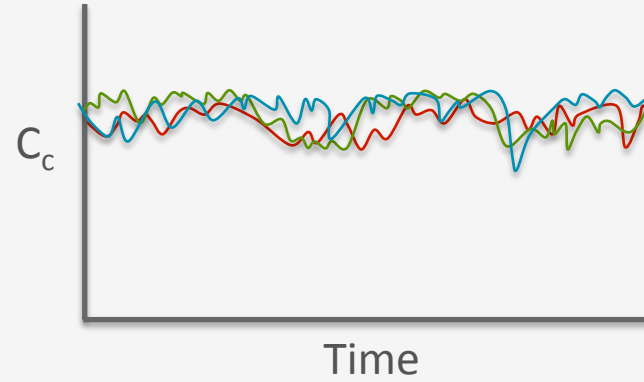
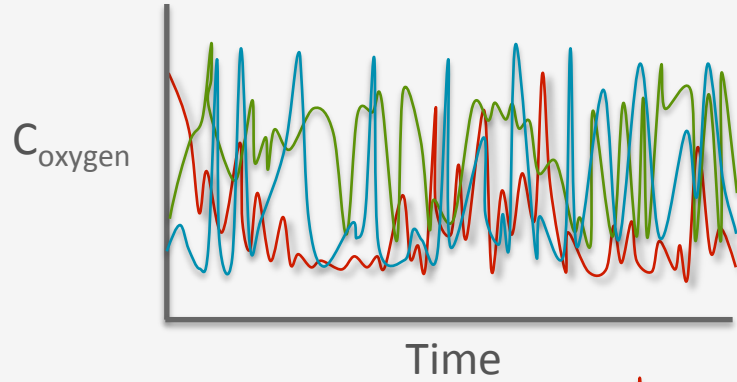
However: local differences!



Cells are facing non-ideal conditions

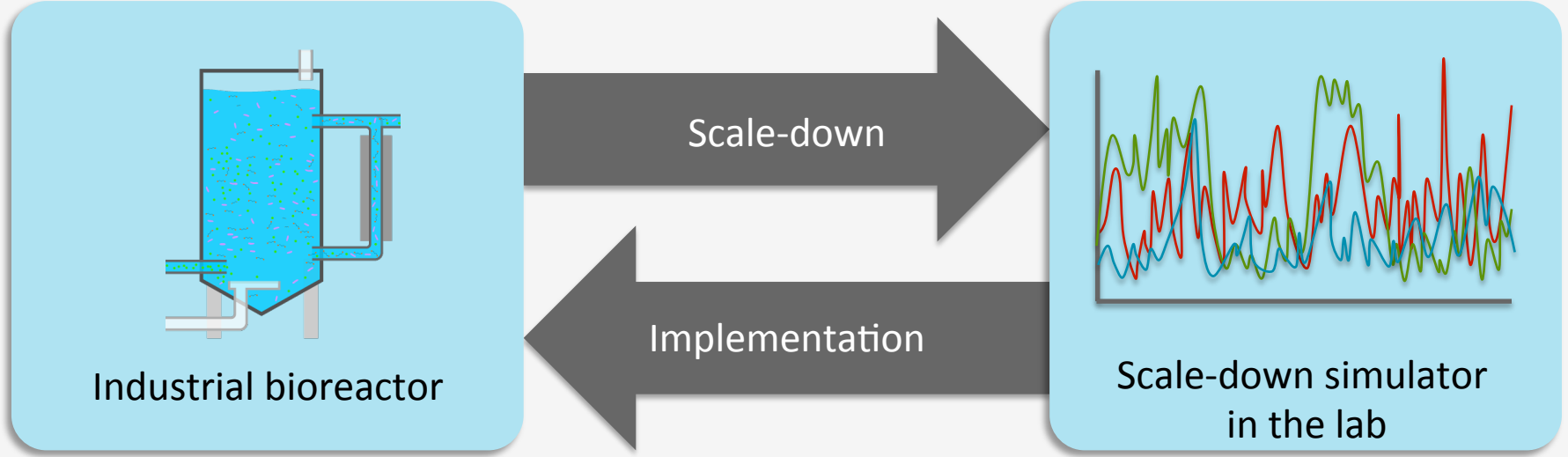


Simulation of cell life-lines of 3 individual cells



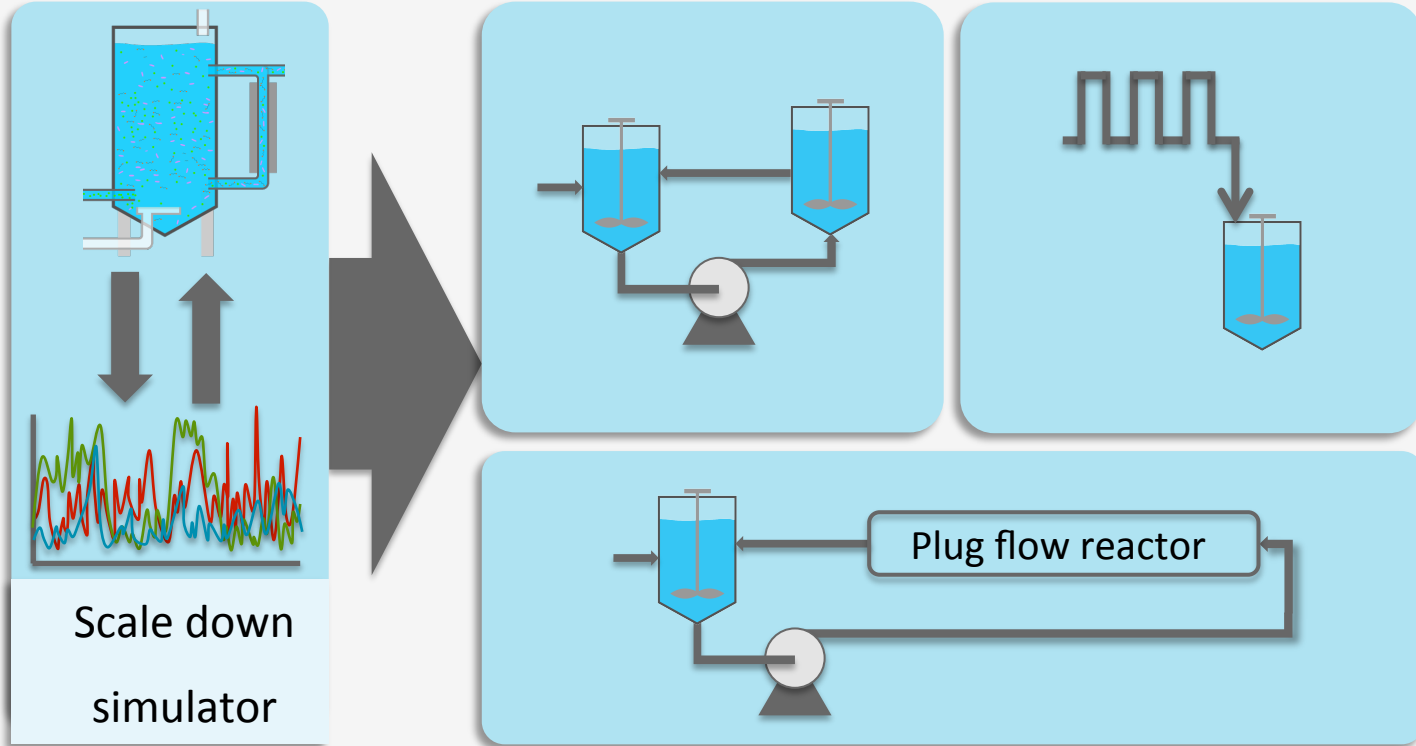
Is the microorganism robust enough?

→ Scale-down, not scale-up



Is the microorganism robust enough?

→ Scale-down, not scale-up



General ways to improve the process:

Micro-organism

- Selection of robust microorganisms
- Metabolic Engineering

Physiology

- Use black box model to better control critical process parameters

Reactor engineering

- Hardware adjustments

Conclusion

Integral design requires quantification and comparison of:

- O₂ transfer, CO₂ transfer, Heat transfer, Mixing to determine
1. the transport bottleneck
 2. the full scale environment of the microorganism

Best rational optimization method:

Scale-down rather than scale-up

See you in the next unit!