ADVANCED COURSE Microbial Physiology and Fermentation Technology

> Course board: Pascale Daran-Lapujade Henk Noorman

AIM OF THE COURSE

13 - 24 January

The aim of the course is to familiarize participants with the integrated, interdisciplinary approach required in modern biotechnology. The course will cover intensive and in-depth presentations of the state of the art. At the same time, the course provides the necessary link between, on one hand, fundamental subjects (thermodynamics, stoichiometry, kinetics, genetics, system biology, fermentor transport and modeling) and, on the other hand, practical aspects (cultivation, (¹³C-)metabolomics, measurements) and applications in large-scale biotechnological processes.

The course provides lectures in fundamental aspects, followed by extensive exercises, made in groups, to stimulate interdisciplinary teamwork.

Subsequently the integration of biological and engineering concepts will be experienced in a design task.

It will be possible to visit research projects and discuss topics of interest (technical aspects of fermentation, sampling, analytics) with scientific/technical staff.

At the end of the course you have learned to integrate life science and technology to achieve effective development/optimisation of new and existing fermentation processes.





COURSE DESCRIPTION

This 10 day course is taught in English and has intensive days. To ensure active participation by those attending, a combination of theoretical (lectures) and practical (exercises, computer simulations, design case study) work is offered. Some online preparatory materials will be given to facilitate all participants to have the same basic knowledge.

LECTURES

The lectures are mainly scheduled in the mornings and sometimes the early evenings. In the lectures, attention will be paid to the following themes:

- Thermodynamics and balances
- Kinetics and stoichiometry (process reaction), membrane transport
- Regulation of metabolism by environmental parameters
- High-cell-density fed batch fermentation
- · Metabolic networks: modelling, regulation and stoichiometry
- Case study, Rate based design
- Metabolic engineering, strategy and applications

EXERCISES, DESIGN CASE STUDY, COMPUTER SIMULATIONS

- The exercises by hand cover thermodynamics, balances, kinetics, stochiometry analysis of fermentation data, metabolomics pathway stoichiometry.
- Tutorials in setting-up and simulating computer models of metabolic networks.
- The case study is on design of a syngas fermentation

CONTACT WITH FACULTY STAFF

- There will be possibilities to visit the research projects of the Department of Biotechnology of Delft University of Technology.
- There will be possibilities to make appointments with faculty staff. Please contact us before hand.

WHO SHOULD ATTEND?

This Advanced Course is aimed at professionals (MSc, PhD or equivalent experience) in microbiology, biochemistry or biochemical engineering with a basic working knowledge of the two other disciplines. Also, molecular biologists with a microbial background may apply. The course is primarily aimed at those already employed in industry and academia who wish to update their theoretical knowledge and practical insight in this field. In addition, this Advanced Course is an option in the two-year postgraduate programs of Delft University of Technology.



COURSE BOARD Pascale Daran-Lapujade

Industrial Biotechnology Experimental Systems Biology/Synthetic Biology Henk Noormán **Bioprocess Engineering** dsm-firmenich

FACULTY STAFF Industrial Biotechnology

Jack Pronk Djordje Bajić Bioprocess Engineering Marieke Klijn Cees Haringa Ludovic Jourdin Environmental Biotechnology Robbert Kleerebezem Martin Pabst Rebeca Gonzalez Cabaleiro

Curator of the Beijerinck Museum Lesley Robertson Cell Systems Engineering Sef Heijnen

COURSE COORDINATION

Yvonne van Gameren Jenifer Baptiste BioTech Delft, Delft University of Technology Department of Biotechnology, the Netherlands

ECTURERS Wouter van Winden dsm-firmenich Delft, the Netherlands

Bjorn Heijstra LanzaTech B.V. Amsterdam, the Netherlands

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David Humbird DWH Process Consulting Denver, USA

Nico Callewaert VIB-Ugent center for medical Biotechnology Zwijnaarde, Belgium

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Frank Bruggeman Vrije Universiteit Amsterdam Amserdam, the Netherlands

Katelijne Bekers/Luisa Cruz MicroHarvest Hamburg, Germany

Chris Mehrer San Diego, USA

Ismahane Remmonay Strassbourg, France

University of California / DTU, Denmark San Diego, USA Adam Feist

Sarah Lieder dsm-firmenich Delft, the Netherlands

PROGRAM

MONDAY 13 JANUARY 2025

Theme: Beginning with the end in mind, balances, conversion rates modes of fermenter operation Coordinator Pascale Daran-Lapujade

- **08.45** Registration **09.00** Outline of the course
- Pascale Daran-Lapujade Microbial Physiology vs. Fermentation Technology Jack Pronk / Henk Noorman 09.15
- 10.30 Developing scalable bioprocesses: Lessons and examples from industry Chris Mehrer
- 11.30 Balances and rates
- Sef Heijnen Lunch & Picture
- 12.30
- q: Biomass specific conversion rates 13.30 Sef Heiinen
- 14.15 Excercises on balances and biomass specific conversion rates Sef Heijnen
- 16.45 Balances and modes of fermentation operation
- Marieke Klijn Social drink and buffet 17.40

TUESDAY 14 JANUARY 2025

Theme: Thermodynamics, stoichiometry and kinetics pf growth, product formation and process design Coordinator Sef Heijnen

09:00 Basic energetics of microbial metabolism Robbert Kleerebezem

- 10:00 Exercises on energetics of microbial metabolism
- Robbert Kleerebezem
- Lunch 12.30 Implications and applications of the thermodynamics of microbial metabolism *Ton van Maris* Black Box model: kinetics/parametrization 13.15
- 14:15 Sef Heijnen
- 15.30 Black Box model: stoichiometry/process reaction for design
- Sef Heijner 17:00 Cellulosic Ethanol a Key Intermediate for Sustainable Mobility and Chemical Production Henning Marckmann
- 18.00
- End of the day

WEDNESDAY 15 JANUARY 2025

Theme: Fermentation: design, transport and data analysis Coordinator Sef Heijnen

- Introduction to bioprocess design using the process reaction 09.00
- Sef Heiinen 10:00 Excercises on bioprocess reactions in design
- Sef Heijnen
- 12.30 Lunch
- 13.30 Fermentor O2/CO2 transport and fed-batch design Sef Heijnen Exercises on data analysis of fermentation processes 15.15
- Sef Heiinen
- 17.40 End of the day

THURSDAY 16 JANUARY 2025

Theme: Pathway stoichiometry, pathway-based models and glutamate fed-batch demo

- Coordinator Wouter van Winden 09.00
- Metabolic pathway stoichiometry: for energy, product, biomass Djordje Bajić Exercise: Stoichiometry pathway model for glutamate fed-batch demo Djordje Bajić /Wouter van Winden / Sef Heijnen 10.45
- 12.30 Lunch
- Fed Batch: Glutamate Fed-batch in silico demo Wouter van Winden / Sef Heijnen 13.15
- 17.00 Fermentation feedstocks: beyond (pure) carbohydrates
- Wouter van Winden 18.00 End of the day

FRIDAY 17 JANUARY 2025

Theme: Metabolic networks: model-based fed-batch, modelling of large-scale dynamics

oordinator Wouter van Winden

- Metabolic network models, ME-models, resource allocation Jens Nielsen 09.00
- 10.00 Metabolic studies in the industrial contexts Jens Nielsen
- Fed Batch: Feed phase: Interactive quantification of knowledge for microorganisms Wouter van Winden / Sef Heijnen 11.15 12.45
- Fed Batch: Model-based optimization of the glutumate fed-batch Wouter van Winden / Sef Heijnen Challenges of large scale fermentation 13.30
- 16.15
- Cees Haringa
- Multi-scale modelling of process dynamics in large-scale bioreactors 17.00 es Haringa 18.15 Social drink

MONDAY 20 JANUARY 2025

- Theme: Case study: Ethanol from syngas Coordinator Henk Noorman 09.00
- Rate-based design of biosystems Sef Heijnen
- Case study: Ethanol from syngas Henk Noorman / Wouter van Winden / Sef Heijnen 09.30
- Lunch 12.45
- (optional) Visit fermentation lab 14.00
- Continuation Case Study Henk Noorman / Wouter van Winden / Sef Heijnen
- Gas Fermentation: a path to low carbon fuel and chemical production with impact 17.15 Biorn Heiistra
- 18.15 End of the day

TUESDAY 21 JANUARY 2025

Theme: Metabolic engineering: Strategies and applications

- Coordinator Henk Noorman 09.00 Metabolic engineering strategies for reducing costs
- Sef Heijnen 10.30 Exercises on metabolic engineering strategies for reducing costs
- Sef Heijnen 12.30 Lunch
- 12 55
- (optional) Visit fermentation lab Metabolic Engineering: from bench to business 13.30
- Irina Borodina Evolutionary Engineering highlighting Adaptive Laboratory Evolution 14.45 Adam Feist
- Engineering membrane transport Jack Pronk 15.30
- 16.30
- Fermentation Intensification Henk Noorman
- 17.30 End of the day

WEDNESDAY 22 JANUARY 2025

Theme: Scale down: Single cell studies and heterogeneity, physiology in the screening stage Coordinator Pascale Daran-Lapujade 09.00 Single cell studies of micro-organisms / Microfluidics

- Frank Bruggeman Population heterogeneity in large-scale bioprocesses
- 10.00 Anna-Lena Heins
- 11.15 Scale-down for understanding large scale performance and robust bioprocessing Cees Haringa / Anna-Lena Heins
- 12.15 12.40 Lunch
- (optional) Visit fermentation lab Transcription factor-based biosensors for strain development 13.15
- Jan Marienhagen Spatio-temporal single-cell analysis in picoliter reactors 14.15 Dietrich Kohlheye
- 15.30 High-throughput strain construction and phenotype testing Sarah Lieder Advances and developments in microbial proteomics
- 16.30 Martin Pabsi
- End of the day 17.30

THURSDAY 23 JANUARY 2025

Theme: Microbial protein production in an industrial context

and bioenergy Robbert Kleerebezem / Rebeca Gonzalez Cabaleiro

Synthetic consortia: new options for industrial biotechnology Djordje Bajić

The Vanishing Link Between Van Leeuwenhoek's Animalcules and Disease before the 19th Century

11111

Lesley Robertson The Megatrends that could re"shape"" the Biotechnology Ismahane Remonnay

Electrobiotechnology: production of chemicals using renewable

- oordinator Henk Noorman Microbial protein production, a complex system in an industrial context 09.00
- Cees Sagt Microbial production of therapeutic proteins Nico Callewaert Scale-up considerations for cultivated meat 10.15
- 11.30
- David Humbird
- 12.30 Lunch
- Microbial proteins for food applications Katelijne Bekers/Luisa Cruz 13.30

electricity as energy source Ludovic Jourdin Lunch

What have we learned? Jack Pronk / Henk Noorman / Pascale Daran-Lapujade

LOCATION

Van der Maasweg 9

http://bt.tudelft.nl

The course will be held at the

Delft University of Technology Department of Biotechnology

2629 HZ Delft, The Netherlands

- 14.30 19.00 Field trip Course dinner

10.15

11.15

12.15

13.15

14.15

15.00

FRIDAY 24 JANUARY 2025 Theme: Looking ahead: Microbial communities, Electrobiotechnology, Megatrends. Looking back: Antonie van Leeuwenhoek, MPFT 2024 Coordinator Pacale Daran-Lapujade 09.00 Microsbial community engineering for production of chemicals

COURSE REGISTRATION

Please register via the website to attend the course. Deadline for application is 23 December 2024. Applications will be handled in order of the date of receipt.

COURSE FEE

€ 4.250 in case of registration before 4 November 2024 or
€ 4.500 in case of registration after this date. In the event of cancellation before 18 November 2024, a full refund will be granted, after this date, a 25% fee charge can be made.

To facilitate enrolment of young PhD-students from universities, a limited number of fellowships is available. The course fee with fellowship is \in 2.100. To apply, please include a copy of your registration as a PhD-student from your university.

The fee includes course materials, lunches, the buffet on Monday and the course dinner on Thursday. The fee does not cover other meals and lodging.

When the number of participants is too low to have a fruitful course, Biotech Delft will cancel the event no later than six weeks before the start of the course. The course fee will be reimbursed within three weeks after cancellation.

In case a speaker will not be able to present his/her lecture due to unforeseen circumstances, Biotech Delft will arrange an equivalent replacement.

Preparatory texts will be sent after receipt of the course fee, a month before the start of the course. The complete digital course book will be supplied at the start of the course.

Biotech Delft organises biotechnology education at postgraduate level. Biotech Delft closely cooperates with the department of Biotechnology of Delft University of Technology. Since its foundation, in 1987, Biotech Delft has very successfully organised various types of postdoctoral education.

Currently Biotech Delft offers Advanced Courses given each year, covering the multidisciplinary spectrum of biotechnology. The courses have a long track-record dating back to 1988.

- Microbial Physiology and Fermentation Technology (1988)
- Downstream Processing (1989)
- Environmental Biotechnology (1993)
- Biocatalysis and Protein Engineering (1999)
- Bioprocess Design (2014)
- Modelling and Computation for Microorganisms in Bioprocesses (2018)
- Integrated Multi-Omics approaches for
 Improvement of Industrial Microbes (2020)

FURTHER INFORMATION

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