

Program

Natural Products Discovery and Production: New Challenges; New Opportunities

June 4-8, 2006

Hotel Santa Fe
1501 Paseo de Peralta, Santa Fe, NM 87501

Co-Chairs

Beth Junker

Senior Director, Merck & Co., Inc. USA

Arnold Demain

Fellow, Charles A. Dana Research Institute,
Drew University, Madison, NJ

Organizing Committee

David Hopwood (John Innes Centre)

Jens Nielsen (Technical University of Denmark)

Satoshi Omura (Kitasato University)

Daniel Wang (Massachusetts Institute of Technology)

Stefano Donadio (KtedoGen)

ECI

Engineering Conferences International

6 MetroTech Center

Brooklyn, NY 11201, USA

Phone: 1-718-260-3743, Fax: 1-718-260-3754

www.engconfintl.org - info@eci.poly.edu

Engineering Conferences International (ECI) is a global engineering conferences program, originally established in 1962, that provides opportunities for the exploration of problems and issues of concern to engineers and scientists from many disciplines. ECI is a not-for-profit partnership between the Engineering Conferences Foundation (ECF) and Polytechnic University.

ECF BOARD MEMBERS

Barry C. Buckland
Mark Green
Allen I. Laskin
Raymond McCabe
Eli Pearce
David K. Robinson
P. Somasundaran

Chair of ECI Conferences Committee: Jules Routbort

ECI Technical Liaison for this conference: Allen Laskin

Polytechnic University President: Jerry Hultin

Polytechnic Liaison to ECF: T.C. Westcott

ECI Director: Barbara K. Hickernell

ECI Assistant Director: Kevin Korpics

The chairs and organizers would like to thank the following for their financial support of this conference:

Merck & Co., Inc.

Eli Lilly and Co.

Astellas

DSM

The Kitasato Institute

Sandoz

Wyeth

Merck KGaA

Martek BioSciences Corporation

Sunday, June 4, 2006

04:00 p.m. – 06:00 p.m.	Registration
06:00 p.m. – 07:20 p.m.	Dinner
07:20 p.m. – 07:30 p.m.	Speaker Introduction: Stephen Donadio, KtedoGen
07:30 p.m. – 09:00 p.m.	Keynote Speaker: Natural products status: present and future paradigms and projections <i>Steve Projan, Wyeth</i>
09:00 p.m. – 10:00 p.m.	Evening Social

NOTES

- Please observe “No Smoking” at ECI technical sessions, meals and social hours.
- Speakers should allow time at the end of their presentation for questions and discussion.
- Poster presenters should stay by their poster for questions and discussion during the poster session in which their poster is scheduled.
- Please silence your cell phone during technical sessions.

- All technical sessions are in Kiva ABC.

- All poster sessions are on the patio in the tent.

- All meals are on the patio with the exception of Sunday dinner and Wednesday banquet. These will be in Kiva ABC.

Monday, June 5, 2006

- 07:30 a.m. – 08:15 a.m. Breakfast
- 08:15 a.m. – 08:30 a.m. **Opening Remarks:** Beth Junker, Merck Research Laboratories;
Allen Laskin, ECI
- Technical Session 1 – Product discovery and modification**
- 08:30 a.m. – 08:35 a.m. **Introduction:** Hans-Peter Fiedler, Universitat Tubingen
- 08:35 a.m. – 09:20 a.m. **Pharmaceutical profiling in drug discovery**
Guy Carter, Wyeth
- 09:20 a.m. – 10:05 a.m. **Discovery of novel inhibitors of bacterial fatty acid synthesis**
Olga Genilloud, CIBE-Merck
- 10:05 a.m. – 10:25 a.m. Coffee Break
- 10:25 a.m. – 11:10 a.m. **Combinatorial biosynthesis by de novo design of modular polyketide synthesis genes**
Hugo Menzella, Kosan
- 11:10 a.m. – 11:55 a.m. **New natural products and biosynthetic pathways from *Streptomyces* genome mining**
Greg Challis, University of Warwick
- 11:55 a.m. – 12:00 p.m. **Summary:** Leonard Katz
- 12:00 p.m. – 04:30 p.m. Lunch on own in town and afternoon free-time
- 04:30 p.m. – 06:00 p.m. **Poster Session A** and refreshments
Ray Lam, Nereus; Allen Laskin, Drew University
- Special Session I – Reduction to practice**
- 06:00 p.m. – 06:05 p.m. **Introduction:** Max Kennedy, IRL Biopharm
- 06:05 p.m. – 06:20 p.m. **Difficult industrial fermentations: Special requirements for producing cytotoxins and growing marine microbes**
Max Kennedy, IRL Biopharm
- 06:20 p.m. – 06:35 p.m. **Applying metabolic flux analysis in strain improvement**
Anna Eliasson Lantz, Biocentrum-DTU
- 06:35 p.m. – 06:50 p.m. **Productivity improvement in industrial fermentations with mycelial morphology.**
Sanjay Tiwari, Biocon
- 06:50 p.m. – 07:05 p.m. **Production of exopolysaccharides by *Claviseps* spp. and possible scale up of this process**
Ales Prell, Institute of Microbiology Czech Academy of Sciences
- 07:05 p.m. – 07:20 p.m. **Control of mycelial morphology: Practical approach**
Sergei Braun, Hebrew University

Monday, June 5, 2006 (continued)

07:20 p.m. – 07:25 p.m. **Summary:** Janet Westpheling, University of Georgia

07:30 p.m. – 09:00 p.m. Dinner

09:00 p.m. – 10:00 p.m. Evening Social

Tuesday, June 6, 2006

07:30 a.m. – 08:30 a.m.

Breakfast

Technical Session 2 – Biodiversity and strain improvement

08:30 a.m. – 08:35 a.m.

Introduction: Margo Haygood, Oregon Health and Science University

08:35 a.m. – 09:20 a.m.

Deep sea actinomycetes: New opportunities for natural product search and discovery
Alan Bull, University of Kent

09:20 a.m. – 10:05 a.m.

Uncultivated symbionts
Eric Schmidt, University of Utah

10:05 a.m. – 10:25 a.m.

Coffee Break

10:25 a.m. – 11:10 a.m.

Secondary metabolite potential of a biological control agent, *Bacillus cereus*, UW85
Michael Thomas, University of Wisconsin

11:10 a.m. – 11:55 a.m.

Combinatorial biosynthesis of lipopeptide antibiotics
Richard Baltz, Cubist

11:55 a.m. – 12:00 p.m.

Summary: Richard Baltz, Cubist

12:00 p.m. – 01:00 p.m.

Lunch

Special Session II – “Off the beaten path” approaches

01:00 p.m. – 01:05 p.m.

Introduction: Kim Lewis, Novobiotic/Northeastern University

01:05 p.m. – 01:20 p.m.

***In silico* biosynthesis: A potent new approach to natural product discovery**
James McAlpine, Ecopia

01:20 p.m. – 01:35 p.m.

From pathway engineering to production: The role of process development
Jorge Galazzo, Kosan

01:35 p.m. – 01:50 p.m.

Semi-synthetic production of the antimalarial natural product Artemisinin
Kinkead Reiling, Arymis

01:50 p.m. – 02:05 p.m.

Growing “unculturable” bacteria for natural product drug discovery
Kim Lewis, Novobiotic/Northeastern University

02:05 p.m. – 02:10 p.m.

Summary: Kim Lewis, Novobiotic/Northeastern University

Tuesday, June 6, 2006 (continued)

Special Session III– “Visions for the future” - panel discussion

02:15 p.m. – 02:20 p.m.

Introduction: Steven Giovannoni, Oregon State University

02:20 p.m. – 02:25 p.m.

Bradley Moore, Scripps Institution of Oceanography

02:25 p.m. – 02:30 p.m.

Juan Asenjo, University of Chile

02:30 p.m. – 02:35 p.m.

Rowan Morris, Novartis

02:35 p.m. – 02:40 p.m.

William Fenical, Scripps Institution of Oceanography

02:40 p.m. – 03:10 p.m.

Debate on the future of natural products

03:10 p.m. – 03:15 p.m.

Summary: William Fenical, Scripps Institution of Oceanography

03:15 p.m.

Mid afternoon and evening free-time – dinner on own in town

Wednesday, June 7, 2006

- 07:30 a.m. - 08:30 a.m. Breakfast
- Technical Session 3 – Scale up/scale down (well plates, shake flasks, bioreactors)**
- 08:30 a.m. – 08:35 a.m. **Introduction:** Wouter Duetz, Enzyscreen
- 08:35 a.m. – 09:20 a.m. **Microtiter plate wells as mini-bioreactors**
Wouter Duetz, Enzyscreen
- 09:20 a.m. – 10:05 a.m. **Microplates and shake-flasks with dissolved oxygen sensors for microbial screening**
Elmar Heinzle, Saarland University
- 10:05 a.m. – 10:25 a.m. Coffee Break
- 10:25 a.m. – 11:10 a.m. **Industrialized strain and fermentation development at Sandoz**
Helmut Haecker, Sandoz
- 11:10 a.m. – 11:55 a.m. **Scale-up of a viscous fungal fermentation for pneumocandins from pilot scale to manufacturing**
Robert Stieber, Merck
- 11:55 a.m. – 12:00 p.m. **Summary:** Heinrich Scherfler, Sandoz
- 12:00 p.m. – 04:30 p.m. Lunch on own in town and afternoon free time
- 04:30 p.m. – 06:00 p.m. **Poster Session B** and refreshments
Ray Lam, Nereus; Allen Laskin, Drew University
- 06:00 p.m. – 07:30 p.m. **Banquet**
- 07:30 pm. – 07:35 p.m. **Remarks:** Allen Laskin, ECI
- 07:35 p.m. – 07:40 p.m. **Poster Awards**
- 07:40 p.m. – 07:45 p.m. **Speaker Introduction:** Leonard Katz
- 07:45 p.m. – 09:15 p.m. **Banquet Speaker**
Misconception, Mistreatment, Misunderstanding and Microbes
Julian Davies, UBC
- 09:15 p.m. – 10:15 p.m. Evening Social

Thursday, June 8, 2006

- 07:30 a.m. - 08:30 a.m. Breakfast
- Technical Session 4 – Cell environment optimization**
- 08:30 a.m. – 08:35 a.m. **Introduction:** Andrew Russell, Eli Lilly
- 08:35 a.m. – 09:20 a.m. **Construction of a novel secretion pathway for intracellular proteins in *Aspergillus niger***
Cees M. J. Sagt, DSM Life Science Products, Food Specialties RD&T
- 09:20 a.m. – 10:05 a.m. **Understanding global regulation of antibiotic biosynthesis in *Streptomyces***
David Sherman, University of Michigan
- 10:05 a.m. – 10:25 a.m. Coffee Break
- 10:25 a.m. – 11:10 a.m. **Cell environment optimization with stochastic search strategies**
Dirk Weuster-Bolz, Technical University of Munich
- 11:10 a.m. – 11:55 a.m. **Comprehensive engineering of metabolic networks to enhance microbial production systems**
Kevin Madden, Microbia
- 11:55 a.m. – 12:00 p.m. **Summary:** Jackie Shanks, Iowa State University
- 12:00 p.m. – 12:15 p.m. **Closing Remarks** – Arnold Demain, Drew University
- 12:15 p.m. Box Lunch distribution and departure

POSTERS – Session A

Natural product production, diversity, regulation and scale-up

1. **Moriniafungin, a novel sordarin derivative produced by the endophytic fungus *Morinia pestalozzioides***
Javier Collado, Centro de Investigación Básica, Merck Research Laboratories, Merck, Sharpe and Dohme de España
2. **YM-254890, a novel cyclic depsipeptide with heterotrimeric G protein Gq/11 inhibitory activity**
Masatoshi Taniguchi, Astellas Pharma Inc.
3. **New anti-infective compounds of microbial origin**
Hiroshi Tomoda, Kitasato University
4. **Abyssomicins, novel polycyclic polyketide antibiotics active against multiresistant Gram-positive pathogens**
H. -P. Fiedler, Universität Tübingen
5. **NADH-fumarate reductase inhibitors produced by fungi**
Kazuro Shiomi, Kitasato Institute for Life Sciences, Kitasato University
6. **Actinohivin, a promising anti-HIV protein of microbial origin**
Haruo Tanaka, The Kitasato Institute
7. **Lycopene and beta-carotene accumulation by a soil bacterium**
Frederik van Keulen, Centro de Engenharia Biológica e Química, Instituto Superior Técnico
8. **Psychrophilic lipases of Antarctic origin**
Barbara Andrews, University of Chile
9. **Isolation of novel cryophilic proteases from Antarctic microorganisms**
Juan A. Asenjo, University of Chile
10. **Searching for novel antibiotic-producing bacteria**
Stefano Donadio, KtedoGen
11. **Biological and chemical diversity: success factors for industrial natural product research**
Rowan Morris, Novartis Institutes
12. **Scale-up of bioreactors: physiological approach contra geometrical similarity**
Ales Prell, Institute of Microbiology Czech Academy of Sciences
13. **Scale-down and parallel operation of a riboflavin production process with *Bacillus subtilis***
Dirk Weuster-Botz, Institute of Biochemical Engineering, Technical University of Munich, Germany
14. **Microalgal fermentation scale-up**
Ray Gladue, Martek Biosciences Corporation

15. **Large scale saline fermentation of an obligate marine actinomycete to produce the proteasome inhibitor NPI-0052**
Ray Lam, Nereus Pharmaceuticals
16. **Dbv4 is a DNA-binding protein regulating expression of OxyA during A40926 biosynthesis**
Rosa Alduina, Dipartimento di Biologia Cellulare e dello Sviluppo, Università degli Studi di Palermo
17. **The BCDH gene clusters of *Streptomyces* differentially regulate antibiotic production and morphogenesis and are the exclusive source of some of the precursors for both type I and type II polyketide biosynthesis in this genus**
Janet Westpheling, University of Georgia
18. **NADPH manipulation in *E. coli* for alteration of pattern of metabolic products**
G. Bennett, Rice University
19. **Pathways for central carbon metabolism in *Actinobacter***
Anna Eliasson Lantz, Center for Microbial Biotechnology, BioCentrum-DTU
20. **Connected antioxidant enzymes in controlled protection of the cell environment**
A.V. Maksimenko, E.G. Tischenko, A.V. Vavaev, Russian Cardiology Research and Production Complex
21. **Metabolic effect of a natural-based mixture in patients with Type 2 diabetes**
M. Kidron, I. Raz, E. Khayat, M. Sinai, H. Bar-on, Hadassah Hospital

POSTERS – Session B

Natural product screening, modeling, pathway engineering and strain improvement

22. **Genomic and phenotypic diversity of *Lactococcus lactis* from dairy and non-dairy origin**
Johan E.T. van Hylckama Vlieg, NIZO Food Research
23. **Functional metagenomics to survey β -lactam antibiotic resistance in a remote Alaskan soil**
Heather K. Allen, University of Wisconsin-Madison
24. ***Saccharomyces cerevisiae* as heterologous host for polyketide production**
Songsak Wattanachaisaereekul, Center for Microbial Biotechnology, BioCentrum, Technical University of Denmark
25. ***Aspergillus* as an efficient cell factory: Metabolic engineering strategies for efficient production of the acetyl-CoA precursor molecule**
Gianni Panagiotou, Center of Microbial Biotechnology-Technical University of Denmark
26. ***Amycolatopsis balhimycina* as a model organism for improved glycopeptides production**
Anna Eliasson Lantz, Center for Microbial Biotechnology-Technical University of Denmark
27. **Re-engineering of our natural products library**
Maya Singh, Wyeth Research
28. **Microtiter plate-based high-throughput fermentation and chemical processing for natural product discovery**
Dwight Baker, Cubist Pharmaceuticals
29. **Microtiter based screening - finding big things in little places**
William Vassiliou, Eli Lilly and Company
30. **Classical strain improvement or finding a needle in a haystack**
Richard DeMaio, Eli Lilly and Company
31. **A 24 deep-well plate cultivation format for classical strain improvement of natural products**
Diane Vesey, Kathleen McLaughlin, Andre Walker, Beth Junker and Neal Connors, Merck & Co.
32. **Breeding of FK506 industrial strain**
Shiho Shimizu, Astellas Pharma Inc.
33. **Identification of novel analogues targeting the nocathiacin I binding site from natural product extracts utilizing a GFP-tagged sensitive/resistant pair**
Samantha Samaras, Merck and Co.
34. **Heterologous expression of compactin gene cluster in *Aspergillus nidulans***
Kanchana Rueksomtawin, Technical University of Denmark
35. **Metabolic engineering of flavonoid biosynthesis in *Saccharomyces cerevisiae* and *Escherichia coli***
Mattheos Koffas, University at Buffalo/The State University of New York

36. **Metabolic engineering bacteria to produce human-like drug metabolites of sirolimus**
Jamie E. Prior, University of Colorado
37. **Metabolic engineering of *Catharanthus roseus* hairy roots for production of indole alkaloids**
Jacqueline V. Shanks, Iowa State University
38. **Engineering of flavonoid biosynthesis in yeast**
John A. Morgan, Purdue University
39. **Laboratory scale modeling of large-scale sterilization cycles prior to technical transfer**
Jean Crinean, Wyeth
40. **A putative membrane protein and an ABC-transporter of *Planobispora rosea* stimulate antibacterial activity in *Streptomyces lividans***
Anna Giardina, Università degli Studi di Palermo
41. **Optimize and improve bacterial polysaccharide production using design of experiment (DOE) and traditional research processes**
Brian Bahler, Wyeth
42. **Improvement of the reproducibility of fermentation processes by robust process operational design and control: an answer on FDA's Pat -Initiative**
Andreas Lubbert, Martin-Luther-University