Program: List of Speakers with titles

<u>June 21 (Sunday)</u>

14:00~ Registration

15:30~15:50 <Ocean Hall>

Opening

Opening Remarks: **F. Peter Guengerich** (Vanderbilt University School of Medicine) 15:50-16:45 Plenary Lecture <Ocean Hall>

Shigeaki Kato (University of Tokyo): Nuclear vitamin D receptor-regulated expression of the human CYP27B1 gene mediates the DNA methylation/demetylation

17:00~19:00 <Ocean Hall>

Session 1: Bioinformatics

Chairs: Osamu Gotoh, Daniele Werck-Reichhart

S1-L1:

17:00-17:30 **Osamu Gotoh** (Kyoto University): Birth and death of P450 genes in metazoan linage

S1-L2:

- 17:30-18:00 **Daniele Werck-Reichhart** (CNRS-University, de Strasbourg): P450 evolution and neofunctionalization in plants: emergence of a novel phenolic pathway in Brassicaceae
- S1-L3:
- 18:00-18:30 **Jongsun Park** (Seoul National University): Integrated bioinformatics platforms for identifying and analyzing cytochrome P450s from diverse genomic resources

S1-L4:

18:30-19:00 **Masanori Arita** (University of Tokyo): Data management of cross-disciplinary information using wiki

19:00~ < Summit Hall >

Welcome Reception

June 22 (Monday)

9:00~13:00 < Summit Hall >

Session 2 : P450s in metabolism and drug discovery

Chairs: Hiroshi Yamazaki, Steven J. Leeder

S2-L1:

9:00-9:30 **Steven J. Leeder** (Children's Mercy Hospital and Clinics): Ontogeny and pediatric pharmacogenetics in drug development

S2-L2:

9:30-10:00 Caroline Lee (Pfizer Global Research & Development): Novel application of cytochrome P450 tools to potentially impact drug design and evaluate drug interactions
S2-L3:

10:00-10:30 Yasuhiro Yamada (Mitsubishi Tanabe Pharma Corp.): A strategic approach to positioning of cytochrome P450 studies and risk assessment for drug-drug interaction in drug discovery

10:30-11:00 (Coffee Break)

S2-L4:

11:00-11:30 Hiroshi Yamazaki (Showa Pharmaceutical University): Remarked properties of human cytochrome P450 1A2 and 3A5 in function and cooperativity for drug development

P-2 (S2-S1):

11:30-11:45 Mariko Nakano (University of Washington): *Tissue distribution and metabolism of arachidonic acid by the orphan P450, CYP4V2*

P-3 (S2-S2):

11:45-12:00 Shinichi Ikushiro (Toyama Prefectural University): Functional co-expression of xenobiotic metabolizing enzymes, human cytochromes P450 and UDP-glucuronosyltransferases, in yeast microsomes

P-4 (S2-S3):

12:00-12:15 Takahiro Murai (University of Utah): CYP3A5-specific mechanism-based inactivation by fluticasone propionate, an inhaled glucocorticoid

P-5 (S2-S4):

- 12:15-12:30 Sasitorn Aueviriyavit (Chiba University): Humanized P450 mouse: (2) Functional expression of human CYP3a isoforms in Cyp3a-Hac mice and their application in drug-drug interaction study via mechanism-based inactivation P-6 (S2-S5):
- 12:30-12:45 Xinxin Ding (Wadsworth Center, New York State Department of Health): Role of mouse CYP2A5 in nicotine clearance and testosterone homeostasis

P-7 (S2-S6):

12:45-13:00 Danielle Sevior (RMIT University): Using the rapid N-in-one cocktail to screen commercially available herbal products for CYP inhibition

9:00~13:00 <Ocean Hall>

Session 3 : Structure and function of P450s

Chairs: Andrew Munro, Eric F. Johnson

S3-L1:

9:00-9:30 Eric F. Johnson (The Scripps Research Institute): Structural flexibility underlying the broad substrate specificity of human P450 2C9: Conformational changes upon substrate binding as determined by X-ray crystallography

S3-L2:

9:30-10:00 **David Leys** (The University of Manchester): Crystal structures of Mycobacterium tuberculosis P450s

S3-L3:

10:00-10:30 Emily E. Scott (The University of Kansas): Structures of human cytochrome P450 2E1

10:30-11:00 (Coffee Break)

S3-L4:

11:00-11:30 **Shingo Nagano** (RIKEN SPring-8 Center): Cytochrome P450 in biosynthesis of indolocarbazoles

P-32 (S3-S1):

- 11:30-11:45 Stephen G. Bell (University of Oxford): The CYP199A2 Class I P450 system from the metabolically diverse bacterium Rhodopseudomonas palustris
 P-33 (S3-S2):
- 11:45-12:00 Natallia Strushkevich (SGC, University of Toronto): Crystal structure of human lanosterol 14alpha-demethylase (CYP51) in complex with ketoconazole
 P-34 (S3-S3):
- 12:00-12:15 Michael R. Wester (Pfizer Global Research & Development): Crystal structures of CYP3A4 in complex with bromocriptine and clotrimazole: Evidence of structural plasticity in the active site

P-35 (S3-S4):

12:15-12:30 Dmitri R. Davydov (University of California, San Diego): Allosteric transitions in cytochrome P450 3A4: A multi-step substrate binding mechanism revealed with novel fluorescent ligands

P-36 (S3-S5):

12:30-12:45 Lianhua Xu (University of Tokyo): Crystal structures of filipin hydroxylases (CYP105P1 and CYP105D6) from Streptomyces avermitilis

13:00~14:30 ~Lunch~

- 14:30~18:30 < Summit Hall >
 - Session 4 : Post-transcriptional, post-translational, and epigenetic events
 - Chairs: Tsuyoshi Yokoi, Daniel W. Nebert

S4-L1:

- 14:30-15:00 Maria Almira Correia (University of California, San Francisco): Post-translational CYP3A phosphorylation/ubiquitination: Means to a degradative end S4-L2:
- 15:00-15:30 **Daniel W. Nebert** (University Cincinnati Medical Center): *Knock-in mouse lines expressing either mitochondrial or microsomal CYP1A1*

S4-L3:

15:30-16:00 Alvaro Puga (University of Cincinnati College of Medicine): *Epigenetic* regulation of drug metabolism Genes

S4-L4:

16:00-16:30 **Miki Nakajima** (Kanazawa University): *MicroRNA as a regulator of human cytochrome P450s and nuclear receptors*

16:30-17:00 (Coffee Break)

P-50 (S4-S1):

17:00-17:15 Takuya Mohri (Kanazawa University): MicroRNA regulates the expression of human CYP2E1

P-51 (S4-S2):

17:15-17:30 Tsuyoshi Yokoi (Kanazawa University): miR-24 represses human HNF4 α expression

P-52 (S4-S3):

17:30-17:45 Johan Lundqvist (Uppsala University): Regulation of key steroidogenic CYP enzymes – a potential novel role for 1α , 25-dihydroxyvitamin D_3

P-53 (S4-S4):

17:45-18:00 Markus Storvik (University of Kuopio): The effects of tobacco-smoke and dietary compounds on Phase I and Phase II xenobiotic metabolism enzyme genes through AH-receptor and tissue-specific transcription factors

S4-L5:

18:00-18:30 Negishi Masahiko (NIEHS, NIH): Signal-mediated mechanism of CAR activation

14:30~18:00 <Ocean Hall>

Session 5 : New mechanisms of P450 reactions

Chair: F. Peter Guengerich

S5-L1:

14:30-15:00 F. Peter Guengerich (Vanderbilt University School of Medicine): *Kinetics of sequential reactions catalyzed by P450 enzymes*

S5-L2:

15:00-15:30 Fumihiko Sato (Kyoto University): Novel P450s in isoquinoline alkaloid biosynthesis

S5-L3:

15:30-16:00 **Andrew Munro** (University of Manchester): Characterization of P450 redox systems in the human pathogen Mycobacterium tuberculosis

S5-L4:

16:00-16:30 **Paul Ortiz de Montellano** (University of California, San Francisco): Cytochrome P450 conformation and catalysis probed by unnatural amino acid substitutions

16:30-17:00 (Coffee Break)

P-54 (S5-S1):

17:00-17:15 Arti Singh (The University of Queensland): Cytochrome P450-mediated fatty acid oxidation: Mechanistic investigations

P-55 (S5-S2):

17:15-17:30 Tatyana Spolitak (University of Michigan): Evidence for catalytic intermediates involved in generation of the chromopyrrolic acid scaffold of rebeccamycin by joint RebO and RebD action. The views on the role of P450s in oxidative coupling reactions

18:30~21:00 Poster Session I (Posters of Sessions $1 \cdot 5$)

<u>June 23 (Tuesday)</u>

9:00~13:00 < Summit Hall >

Session 6 : Transcriptional regulation

Chairs: Masahiko Negishi, Frank J. Gonzalez

S6-L1:

9:00-9:30 Frank J. Gonzalez (National Cancer Institute): Regulation of human P450s: Studies with humanized mice

P-60 (S6-S1):

9:30-9:45 Nico Scheer (TaconicArtemis): Use of humanized mouse models to study the transcriptional regulation of Cytochrome P450 genes
P-61 (S6-S2):

- 9:45-10:00 Yoshihiro Konno (NIEHS, NIH): Nuclear xenobiotic receptor PXR locks co-repressor SMRT onto the CYP24A1 promoter to attenuate vitamin D₃ activation P-62 (S6-S3):
- 10:00-10:15 Jin-ding Huang (National Cheng Kung University): Transcriptional inhibition of CYP3A4 as novel mechanism of drug-drug interaction
- P-63 (S6-S4):
- 10:15-10:30 Kaoru Kobayashi (Chiba University): A compound in fetal bovine serum modulates the activation of pregnane X receptor
- 10:30-11:00 (Coffee Break)

S6-L2:

11:00-11:30 **Oliver Hankinson** (University of California, Los Angeles): *Transcriptional* regulation of CYP1A1, CYP1B1, and CYP2S1

S6-L3:

11:30-12:00 Akinori Ohta (University of Tokyo): Control of cytochromes P450 production by an Opi1-family transcription factor in yeast Yarrowia lipolytica

S6-L4:

12:00-12:30 **Graham Robertson** (The University of Sydney): *Transcriptional repression of hepatic drug clearance pathways by tumour-derived cytokines*

P-64 (S6-S5):

- 12:30-12:45 Jae-Gook Shin (Inje University College of Medicine): Pharmacogenomics of transcription regulation in drug metabolism and pharmacokinetics
 P-65 (S6-S6):
- 12:45-13:00 Uli Zanger (Dr. Margarete Fischer-Bosch Institute of Clinical Pharmacology): Pathway-oriented pharmacogenomics approach to probe transcriptional regulation of CYP3A4

9:00~13:00 <Ocean Hall>

Session 7: Heme-thiolate proteins

Chair: Toru Shimizu, Huiying Li

S7-L1:

9:00-9:30 Hofrichter Martin (International Graduate School of Zittau): Aromatic peroxygenases from mushrooms: extracellular heme-thiolate proteins of a new enzyme sub-subclass?

S7-L2:

9:30-10:00 **Toru Shimizu** (Tohoku University): The critical role of the thiolate-heme complex in the functioning of heme-sensor proteins

S7-L3:

- 10:00-10:30 **Huiying Li** (University of California Irvine): *Engineering a P450BM3-like red FMN semiquinone into neuronal nitric oxide synthase*
- 10:30-11:00 (Coffee Break)

S7-L4:

11:00-11:30 **Shigetoshi Aono** (National Institutes of Natural Sciences): *Physiological role* of thiolate coordination to the heme in CooA from R. rubrum

P-72 (S7-S1):

11:30-11:45 Hirofumi Shoun (University of Tokyo): NADH-peroxidase activity of

multi-functional detoxifying enzyme, P450nor

P-73 (S7-S2):

11:45-12:00 Yasuhiro Mie (AIST): Electrochemically-driven drug metabolism by human cytochrome P450 immobilized on hydrophobic electrode surface

P-74 (S7-S3):

- 12:00-12:15 Kelath M. Manoj (Vellore Institute of Technology): Kinetics of one-electron oxidations by chloroperoxidase: What is the analogous take-home lesson for CYPs?
 P-75 (S7-S4):
- 12:15-12:30 Jotaro Igarashi (Tohoku University): The heme-binding site of the Heme-regulated Inhibitor (HRI), and the role of the heme regulatory motif in heme sensing

 $13:00 \sim Excursion$

19:00~ Banquet (at KARIYUSHI BEACH HOTEL)

June 24 (Wednesday)

9:00~13:00 < Summit Hall >

Session 8 : Biotechnology I

Chairs: Rita Bernhardt, Toshiyuki Sakaki

S8-L1:

9:00-9:30 **Rita Bernhardt** (Saarland University): Engineering CYP106A2 for changed substrate specificity and characterization of new substrates

S8-L2:

9:30-10:00 James R. Halpert (University of California, San Diego): Engineering of mammalian cytochromes P450 2B and 3A by site-directed mutagenesis and directed evolution

S8-L3:

10:00-10:30 **Toshiyuki Sakaki** (Toyama Prefectural University): Construction of a highly active vitamin D hydroxylase based on crystal structure of CYP105A1 and its application to production of 1α,25-dihydroxyvitamin D₃

10:30-11:00 (Coffee Break)

S8-L4:

11:00-11:30 Akira Arisawa (Mercian Corporation): Actinomycete cytochrome P450 from Pseudonocardia autotrophica that catalyzes vitamin D₃ hydroxylation
P-80 (S8-S1):

11:30-11:45 Hirofumi Ichinose (Kyushu University): Functional characterization of cytochrome P450 from the white-rot fungus Phanerochaete chrysosporium
P-81 (S8-S2):

11:45-12:00 Sun-Ha Park (Chonnam National University): Engineering of bacterial P450 BM3 for human P450 2A6 activity with indigo formation

P-82 (S8-S3):

12:00-12:15 Shinya Fushinobu (University of Tokyo): Altering the substrate specificity of P450foxy

P-83 (S8-S4):

12:15-12:30 Hidehiko Hirakawa (University of Tokyo): Chimeric self-sufficient P450 system with a branched structure

P-84 (S8-S5):

12:30-12:45 Sjef Cornelissen (TU Dortmund): A cytochrome P450 based bioprocess for limonene hydroxylation to perillyl alcohol

9:00~13:00 <Ocean Hall>

<u>Session 9 : New aspects of P450 functions</u> *Chairs: Hirofumi Shoun, John H. Dawson* S9-L1:

9:00-9:30 John H. Dawson (University of South Carolina): Generation and reactivity of transient cytochrome P450 oxygen-containing intermediates

S9-L2:

9:30-10:00 Andreas Daiber (Johannes Gutenberg University Mainz, School of Medicine): Reaction mechanisms of P450nor

S9-L3:

10:00-10:30 **Neil C. Bruce** (University of York): Insight into the structure and function of XplA: a unique explosives degrading cytochrome P450

10:30-11:00 (Coffee Break)

S9-L4:

11:00-11:30 **Debashis Ghosh** (Hauptman-Woodward Medical Research Institute): Structure and function of aromatase, the estrogen factory

P-94 (S9-S1):

- 11:30-11:45 Kenneth Jensen (University of Copenhagen): *Utilizing the power of light* P-95 (S9-S2):
- 11:45-12:00 Hisakazu Yamane (University of Tokyo): The presence of diterpenoid phytoalexin biosynthetic gene clusters containing cytochrome P450 monooxygenase genes in rice

P-96 (S9-S3):

12:00-12:15 Naoki Takaya (University of Tsukuba): Production of hydroxy-fatty acids and alkanediols by Escherichia coli cells producing fungal cytochrome P450foxy

13:00~14:30 ~Lunch~

14:30~18:30 < Summit Hall >

Session 10: Biotechnology II

Chairs: Hideo Ohkawa, Frances H. Arnold

S10-L1:

14:30-15:15 Frances H. Arnold (California Institute of Technology): In Darwin's honor: artificial selection of cytochrome P450s

S10-L2:

15:15-15:45 **Birger Lindberg Meller** (University of Copenhagen): *The cytochrome P450s in cyanogenic glucoside synthesis*

S10-L3:

15:45-16:15 Hideo Ohkawa (Fukuyama University): How are recombinant P450s and

AhRs useful for phytomonitoring and phytoremediation of persistent organic pollutants of environmental chemicals?

P-100 (S10-S1):

- 16:15-16:30 Dong-Hyun Kim (Chonnam National University): Generation of the human metabolite piceatannol from the anti-cancer preventive agent resveratrol by bacterial cytochrome P450 BM3
- 16:30-17:00 (Coffee Break)

S10-L4:

- 17:00-17:30 **Yoshikazu Tanaka** (Suntory Holdings Ltd): *Flower colour modification by modulating expression of cytochrome P450 genes*
- P-101 (S10-S2):
- 17:30-17:45 Kwon-Young Choi (Seoul National University): Functional expression of Nocardia farcinica IFM10152 P450s in E. coli for A-ring specific monohydroxylation of daidzein

P-102 (S10-S3):

17:45-18:00 Hyun-Jun Kim (Suntory Institute For Bioorganic Research): Metabolic engineering of lignan biosynthesis in Forsythia cell suspension cultures by expression of a Sesamum CYP81Q1

P-103 (S10-S4):

18:00-18:15 Satoshi Iwakami (Kyoto University): Isolation of cytochrome P450 genes and their expression in the multiple-herbicide resistant biotype of late watergrass (Echinochloa phyllopogon)

14:30~18:30 <Ocean Hall>

<u>Session 11 : Biophysical methods & nanotechnology</u>

Chairs: Alexander Archakov, William M. Atkins

14:30-15:00 Ilia G Denisov (University of Illinois at Urbana-Champaign): Common properties and specific features of cytochromes P450

S11-L2:

15:00-15:30 **Paul M. Champion** (Northeastern University): Vibrational coherence spectroscopy investigations of heme proteins with sulphur ligation

S11-L3:

15:30-16:00 Alexander Archakov (Russian Academy of Medical Sciences): Nanotechnology for visualization, counting and kinetic studies of cytochrome P450 complexes

S11-L4:

- 16:00-16:30 William M. Atkins (University of Washington): Ligand-based NMR and single molecule spectroscopy as probes of heterotropic effects in cytochrome P450s
- 16:30-17:00 (Coffee Break)
- P-105 (S11-S1):
- 17:00-17:15 Wang Yuqin (Swansea University): Quantitative screening of cytochrome P450s by mass spectrometry: Specificity and sensitivity

S11-L1:

18:30~21:00 Poster Session II (Posters of Sessions 6 - 13)

June 25 (Thursday)

9:00~13:00 < Summit Hall >

Session 12 : Functional genomics

Chairs: Damjana Rozman, Shigehiro Ohdo

NOTE: The order of lectures changed!

S12-L4:

9:00-9:30 Katrin Marcus (Ruhr-University Bochum): Mass spectrometry- based assay for absolute quantification of cytochrome P450s in human liver

S12-L2:

9:30-10:00 **Shigehiro Ohdo** (Kyushu University): *Molecular clock mechanisms of drug metabolism*

S12-L3:

- 10:00-10:30 William Griffiths (Swansea University): Discovering new products of CYP catalysed reactions in brain, CSF and plasma
- 10:30-11:00 (Coffee Break)

S12-L1:

- 11:00-11:30 **Damjana Rozman** (University of Ljubljana, Faculty of Medicine): *The* cross-talk of CYPs in cholesterol homeostasis and drug metabolism
- P-107 (S12-S1):
- 11:30-11:45 Simon Horvat (University of Ljubljana): Generation of Cyp51 conditional knockout mice to study its function during embryo and male germ cell development
 P-108 (S12-S2):
- 11:45-12:00 Hajime Takeuchi (Tokushima Bunri University): Cyp26b1 regulates the expression of the gut-homing receptor CCR9 in T cells

P-109 (S12-S3):

12:00-12:15 Bhagavatula Moorthy (Baylor College of Medicine): Mice deficient in the gene for cytochrome P450 (CYP)1A2 display increased levels of F2-isoprostanes, oxidative DNA adducts, and augmented susceptibility to oxygen-mediated lung injury in vivo

9:00~13:00 <Ocean Hall>

Session 13 : Steroidogenesis

Chairs: Michael R. Waterman, Nobuhiro Harada

S13-L1:

9:00-9:30 **Kjell Wikvall** (Uppsala University): Regulation of CYP enzymes in steroidogenesis with particular focus on cholesterol metabolism and vitamin D-related processes

S13-L2:

- 9:30-10:00 Marion Sewer (Georgia Institute of Technology): Protein kinase C regulates CYP17 transcription by mediating the phosphorylation of steroidogenic factor-1
 S13-L3:
- 10:00-10:30 **Michael R. Waterman** (Vanderbilt University School of Medicine): Development of CYP51 inhibitors in protozoa

10:30-11:00 (Coffee Break)

S13-L4:

11:00-10:30 Nobuhiro Harada (School of Medicine, Fujita Health University): Multiple transcription factors regulate a brain-specific promoter activity of mouse aromatase (CYP19A1) gene

S13-L5:

- 10:30-12:00 **Evan Simpson** (Prince Henry's Institute of Medical Research): *Obesity, aromatase and breast cancer*
- P-112 (S13-S1):
- 12:00-12:15 Hiroshi Kataoka (University of Tokyo): Cytochrome P450s involved in biosynthesis of ecdysteroids in insects; regulation of gene expression by neuropeptides
 P-113 (S13-S2):
- 12:15-12:30 Robert Tuckey (University of Western Australia): Metabolism of substrates incorporated into phospholipid vesicles by 25-hydroxyvitamin D3 1α-hydroxylase (CYP27B1)

P-114 (S13-S3):

12:30-12:45 Tadashi Ogishima (Kyushu University): Local steroidogenesis in pancreatic beta-cells

P-115 (S13-S4):

12:45-13:00 Maamar Souidi (Institute for Radiological Protection and Nuclear Safety): *Testicular steroidogenesis in adult rats following chronic exposure to 137 Cesium since uterine life or birth* Oral presentation canceled

13:00~14:30 ~Lunch~

14:30~15:30 ~Poster award and Closing~