February 2011
Vol. 39, No. 2

CONTENTS

SHORT COMMUNICATIONS

Delayed Elimination of SN-38 in Cancer Patients with Severe Renal Failure. Ken-ichi Fujita, Yu Su-nakawa, Keisuke Miwa, Yoko Akiyama, Minako Sugiyama, Kaori Kawara, Hiroo Ishida, Keishi Yama-shita, Keiko Mizuno, Shigehira Saji, Wataru Ichikawa, Wataru Yamamoto, Fumio Nagashima, Toshimichi Miya, Masaru Narabayashi, Yuichi Ando, Takashi Hirose, and Yasutsuna Sasaki

Altered Cytochrome P450 Expression in Mice during Pregnancy. Kwi Hye Koh, Hui Xie, Ai-Ming Yu, and Hyunyoung Jeong

Critique of the Two-Fold Measure of Prediction Success for Ratios: Application for the Assessment of Drug-Drug Interactions. Eleanor J. Guest, Leon Aarons, J. Brian Houston, Amin Rostami-Hodjegan, and Aleksandra Galetin

ARTICLES

Morphine Brain Pharmacokinetics at Very Low Concentrations Studied with Accelerator Mass Spectrometry and Liquid Chromatography-Tandem Mass Spectrometry. Muhammad Waqas Sadiq, Mehran Salehpour, Niklas Forsgard, Göran Possnert, and Margareta Hammarlund-Udenaes

Human CYP2S1 Metabolizes Cyclooxygenase- and Lipooxygenase-Derived Eicosanoids. Peter Bui, Satoshi Imazumi, Sudheer Reddy Beedanagari, Srinivasa T. Reddy, and Oliver Hankinson

CYP4F Enzymes Are Responsible for the Elimination of Fingolimod (FTY720), a Novel Treatment of Relapsing Multiple Sclerosis. Yi Jin, Markus Zollinger, Hubert Borell, Alfred Zimmerlin, and Christopher J. Patten

Absorption and Disposition of the Sphingosine 1-Phosphate Receptor Modulator Fingolimod (FTY720) in Healthy Volunteers: A Case of Xenobiotic Biotransformation Following Endogenous Metabolic Pathways. Markus Zollinger, Hans-Peter Gschwind, Yi Jin, Claudia Sayer, Frédéric Zécri, and Stefan Hartmann

Glutaredoxin and Thioredoxin Can Be Involved in Producing the Pharmacologically Active Metabolite of a Thienopyridine Antiplatelet Agent, Prasugrel. Katsunobu Haigihara, Miho Kazui, Atsushi Kurihara, Kazushi Kubota, and Toshihiko Ikeda

Disposition and Metabolism of [14C]SB-649868, an Orexin 1 and 2 Receptor Antagonist, in Humans. Cecilia Rczulli, Mike Nash, Mark Wright, Steven Thomas, Stefano Zamuner, Mario Pellegratto, Paolo Bettica, and Gary Boyle

Metabolism and Disposition of 14C-Labeled Peliglitzazor in Humans. Lefei Wang, Carey Munisick, Sean Chen, Samuel Bonacorsi, Peter T. Cheng, W. Griffith Humphreys, and Donglu Zhang

Effects of Ketoconazole on the In Vivo Biotransformation and Hepatobiliary Transport of the Thrombin Inhibitor AZD0837 in Pigs. Elin M. Mattson, Johan E. Palm, Ulf G. Eriksson, Per-nilla Bottner, Anna Lundahl, Lars Knutson, and Hans Lennernäs

Influence of Cremophor EL and Genetic Polymorphisms on the Pharmacokinetics of Paclitaxel and Its Metabolites Using a Mechanism-Based Model. Martin N. Fransson, Henrik Greén, Jan-Eric Lit-ton, and Lena E. Friberg

Tissue Distribution and Elimination of [14C]Apixaban in Rats. Lefei Wang, Kan He, Brad Maxwell, Scott

Continued on next page
Attenuation of Intestinal Absorption by Major Efflux Transporters: Quantitative Tools and Strategies Using a Caco-2 Model. Xuena Lin, Suzanne Skolnik, Xiaohui Chen, and Jianling Wang ........................................ 256

The Utility of In Vitro Data in Making Accurate Predictions of Human P-Glycoprotein-Mediated Drug-Drug Interactions: A Case Study for AZD5672. Robert Elsby, Michael Gillen, Caroline Butters, Gemma Imisson, Pradeep Sharma, Veronica Smith, and Dominic D. Surry ......................................................... 275

Tissue Distribution and Characterization of Drug-Related Material in Rats and Dogs after Repeated Oral Administration of Casopitant. Sabrina Pagliaruso, Silvia Martinucci, Ldicia Bordini, Lidia Miraglia, Domenico Cufari, Luca Ferrari, and Mario Pellegatti .............................................................. 283

A Pharmacokinetic Model for Evaluating the Impact of Hepatic and Intestinal First-Pass Loss of Saquinavir in the Rat. R. Lledó-García, A. Nácher, V. G. Casabó, and M. Merino-Sanjuań .............................................................. 294


Brain Tissue Binding of Drugs: Evaluation and Validation of Solid Supported Porcine Brain Membrane Vesicles (TRANSIL) as a Novel High-Throughput Method. Raffaele Longhi, Silvia Corbioli, Stefano Fontana, Federica Vinco, Simona Braggio, Lydia Helmdach, Jürgen Schiller, and Hinnerk Boriss .................................................. 312

Correlation between Bilirubin Glucuronidation and Estradiol-3-Gluronidation in the Presence of Model UDP-Glucuronosyltransferase 1A1 Substrates/Inhibitors. Jin Zhou, Timothy S. Tracy, and Rory P. Remmel ......................................................... 322


Identification and Validation of Novel Human Pregnan X Receptor Activators among Prescribed Drugs via Ligand-Based Virtual Screening. Yongmei Pan, Linhao Li, Gregory Kim, Sean Ekins, Hongbing Wang, and Peter W. Swaan ................................................ 337

Inactivation of Cytochrome P450 (P450) 3A4 but not P450 3A5 by OSI-930, a Thiophene-Containing Anticancer Drug. Hsia-lien Lin, Haoming Zhang, Christine Medower, Paul F. Hollenberg, and William W. Johnson ...................................................... 345

ERRATUM

Correction to “Identification of Clinically Utilized Drugs that Activate Pregnan X Receptors” . . . 351