DILIsym Services, a Simulations Plus company, presents:
The DILIsym® 3-Day Training Workshop

March 5 - 7, 2018 in San Diego, California

DILIsym Services, a Simulations Plus company, will be hosting its "DILIsym® 3-Day Training Workshop" in San Diego, California on March 5 - 7, 2018.

This three-day, hands-on course provides a working knowledge of the basic theories and application of state-of-the-art quantitative system toxicology (QST) modeling and simulation software (DILIsym®) for the analysis of the potential for drugs and drug candidates to cause drug-induced liver injury (DILI), or for the investigation of mechanisms involved in a documented DILI event. A combination of presentations and interactive examples using DILIsym will illustrate how to gather the appropriate *in vitro* toxicity data, derive DILIsym related toxicity parameters, conduct simulations of expected outcomes and analyze results for impacting decisions.

Attendance for this event is limited, so register today!

Who should attend?

This workshop is appropriate for pharmaceutical/biotechnology scientists and engineers in the areas of toxicology, clinical pharmacology, pharmacovigilance, DMPK and ADME. This is a beginner's course - prior experience with DILIsym is not required. The course will use DILIsym for all case studies, but the guiding principles will be taught in a software-independent manner. Class size is limited to encourage interaction with the course instructors and among attendees. Interaction and networking among industry, government, and academic scientists is an important and valuable part of the experience!

What will you learn?

Upon completion of this course, you should have a solid understanding of the interactions that exist among the various mechanistic phenomena affecting drug actions on the liver that could result in a safety risk to patients.

You will understand the following important aspects of liver safety investigation:
- primary mechanisms often involved in DILI events, including mitochondrial dysfunction, oxidative stress production, lipotoxicity and bile acid transporter inhibition
- *in vitro* assay design, execution and stage-wise approach considerations for:
    o high content imaging techniques to assess oxidative stress, GSH content, ATP, etc.
    o the Seahorse XF analyzer assay for a detailed analysis of mitochondrial effects
    o bile acid transporter inhibition IC50 and Ki studies
- pharmacokinetics including prediction of liver concentrations and how this impacts DILI predictions
- differences between early (discovery) stage approaches and late stage (clinical) approaches
- accepted and novel liver safety biomarkers
- general concepts accepted by regulators related to DILI monitoring and detection

You will gain basic experience with:
- translating *in vitro* data into DILIsym parameter values
- simulating expected DILI outcomes for humans, rats, mice and dogs
- utilizing simulated populations (SimPops™) to:
    o assess expected outcomes on a population level, including rare responses expected as the treated population grows
    o understand how disease populations may respond differently from healthy volunteers
- understanding and using the:
    o DILIsym optimization feature
    o DILIsym clinical monitoring feature capable of allowing dynamic dose titration or cessation depending on liver safety outcomes in real time
    o DILIsym Data Comparisons feature for validating key concepts represented in DILIsym
    o DILIsym Specified Data feature that allows for direct input of PK information to drive DILI predictions directly from GastroPlus™ and other simulators
    o DILIsym sensitivity analysis capabilities for determining uncertainty around predictions prior to making decisions
- examples of DILIsym use within the regulatory arena where development programs were impacted through QST
- using simulated outcomes to help better design mechanistic *in vitro* toxicology studies

How will the workshop operate?

All presentation files and data sets needed to run case studies will be available in electronic format. All attendees will be responsible for bringing their own laptop computers.

Continental breakfast, refreshment breaks, and lunch will be provided each day.

Where will this be held?
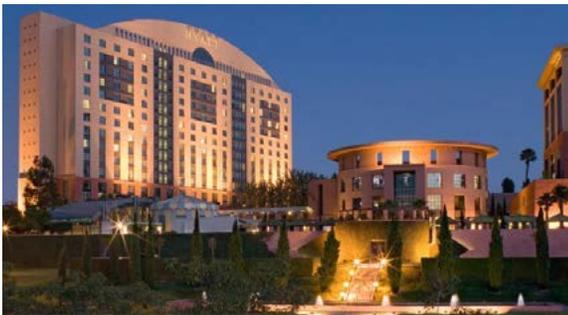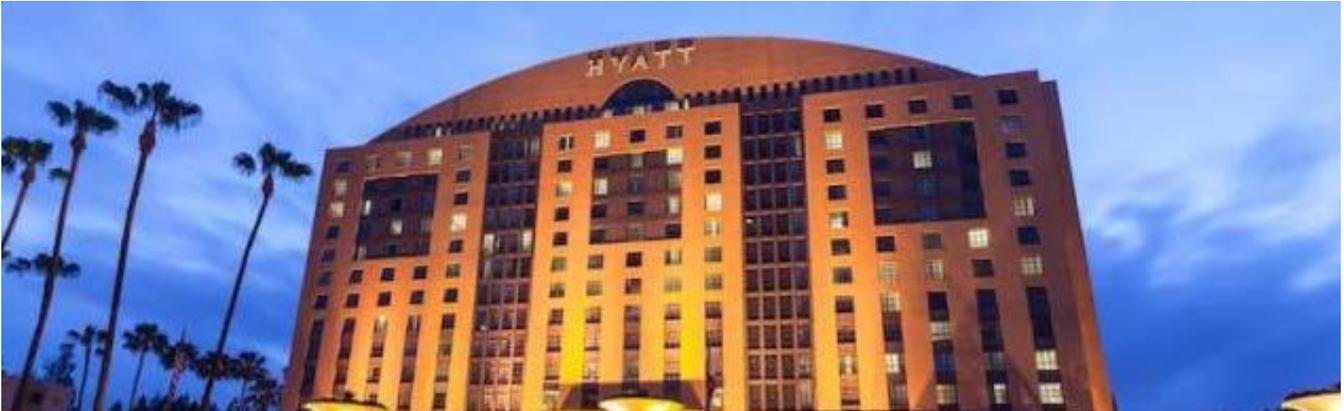
Hyatt Regency La Jolla at Aventine
3777 La Jolla Village Drive
San Diego, California, USA, 92122
Tel: +1 858 552 1234
https://lajolla.regency.hyatt.com/en/hotel/home.html

Please fill in this form and return to Renee Bouche (renee@simulations-plus.com); Fax: +1-661-723-5524
To register by phone, please call Renee at +1-661-723-7723 ext. 227

The DILIsym® 3-Day Training Workshop
March 5 - 7, 2018 in San Diego, California

Title: Professor / Dr. / Mr. / Mrs. / Miss / Ms.

FIRST NAME:

LAST NAME: COMPANY:

POSITION: DEPARTMENT:

ADDRESS:

TELEPHONE: EMAIL:

PURCHASE ORDER NO. (if applicable):

Cost for the workshop is USD $1200 per person (industry non DILI-sim members), $996 (DILI-sim Tier 2 members), $828 DILI-sim Tier 1 members), and $600 (government/academic) and includes all workshop materials, continental breakfast, refreshment breaks, and lunch each day.

Hotel accommodations are not included with registration.

Method of payment (Please check one)

☐ Credit card (a confirmation message will be sent to the email address provided)

Name on card: _____ Email: _____ Tel: _____

Card billing address: _____

_____ Zip/Post Code: _____

_____

Type of card: ☐ Visa ☐ MasterCard ☐ AMEX Card Number: _____

Expires: _____ Security code: _____

☐ Payment by check (you will be invoiced upon receipt of your completed registration form)

☐ Payment by wire transfer (you will receive wire transfer information upon receipt of your completed registration form)

☐ Payment online (you will be redirected to the payment portal when registering online at simulations-plus.com/workshops)

Terms and Conditions
Cancellation Policy: Cancellations made prior to February 28, 2018 will be eligible for an 80% refund. Refunds for cancellations will be honored up to 45 days after the date of payment for credit card transactions. Substitutions are allowed up to 10 days before the event.

Payment Terms: Following completion and return of the registration form, the total fee must be paid within 30 days of receipt of invoice. All fees must be paid in full prior to the start of the workshop.