Speakers:

Dr Christopher Burgess
Chairman of the Analytical Quality Control Working Group

Stephen Young
Head of Analytical Science Inspection, Enforcement and Standards Division, MHRA, UK

Dr Peter Rauenbuehler
Roche, USA

Dr Bernd Renger
Member of the Analytical Quality Control Working Group

Dr Lance Smallshaw
UCB Biopharma sprl, Belgium

Dr Bianca Teodorescu
UCB Biopharma sprl, Belgium

Every participant will receive the Draft Version of ECA's SOP on OOE and OOT Results!

ECA ACADEMY

International Conference

OOT Forum 2014

SOP Out of Expectation (OOE) and Out of Trend (OOT) Results

Compiled by ECA's Analytical Quality Control Working Group

Prague, Czech Republic, 22 - 23 October 2014

Highlights

- Methods and Approaches for Detecting
  - Out of Expectation (OOE) Data
  - Out of Trend (OOT) Data, where no Trend is Expected
  - Out of Trend (OOT) Data, where a Trend is Expected, e.g. for Stability Testing
- Become Part of the Peer Review Group to have a direct impact on the contents of this new document:
  All participants will have the opportunity to provide input to the contents of this guidance document during the interactive 'critique of the proposals' sessions for each of these topics.

Objective

The **ECA Working Group on Analytical Quality Control** was set up in 2010 in order to generate a harmonised SOP on managing analytical deviations within the laboratory including OOS, OOE and OOT results.

The result was a framework **SOP Version 1 which was launched at the OOS Forum in 2012** as a basis for broader discussion. In the meantime Version 2 of this ECA OOS SOP is available for all ECA members. At the same time it was decided that the handling of OOT results should be addressed in a separate **OOT SOP**, since there is a lack of knowledge in the industry and a lack of guidance for trend analysis from the regulators. In the EU there have been recent updates to Chapter 6 and Chapter 1 of the EU GMP for trend requirements.

The aim of ECA's QC Working Group is to address these issues by developing a new guideline - OOT SOP - aimed at QC and other quality groups to encourage the application of a consistent and scientifically sound approaches to trend analysis as part of a QMS for assuring data integrity. There are three planned components:

1. methods and approaches for detecting out of expectation (OOE) data within an analytical sequence which could be based on process capability of analytical procedures
2. methods and approaches to detecting out of trend (OOT) data between analytical sequences where no trend is expected. This could be a variety Statistical Process Control approaches and
3. methods and approaches to detecting out of trend (OOT) data between analytical sequences where a trend is expected as is the case for Stability Testing.

At this ECA OOT Forum in Prague we want to discuss and explore these methods and approaches some of which will be simple and others more complex. Furthermore it is our desire to position the regulatory needs and inspection expectations to put all the technical aspects in context.

Participants will have the opportunity to provide input to the contents of the guidance document during the interactive 'critique of the proposals' sessions for each of the topics.

The ECA QC Working Group's goal is to have a basic global framework for OOT within R&D, production and QC laboratories acceptable to the authorities which individual companies may adapt to fit their particular Quality Management System.

Target Group

This conference is intended for all levels of technical and managerial personnel dealing with out-of-trend results, including R&D, production, analytical laboratories, contract laboratories, microbiological laboratories and employees in the Quality Assurance/Quality Control.

Forum Moderator

Dr Christopher Burgess,
Burgess Analytical Consultancy, UK,
Chairman of the Analytical QC Working Group

Social Event

In the evening of the first course day, all participants are invited to a guided sight-seeing tour of Prague and a dinner in the city of Prague afterwards. This is an excellent opportunity to share your own views and experiences with colleagues from other companies in a relaxed and casual atmosphere

INTRODUCTION / REGULATORY SESSION

Introduction to ECA's Analytical QC Working Group and the OOT Process
- Aims & Objectives
- Lessons learned from the OOS SOP process
- Structure of the OOT process
- Technical aspects of OOT
- Outline of OOT Forum; aims & objectives

Dr Christopher Burgess, UK, Chairman of the Analytical QC Working Group

Regulatory Importance of Trend Analysis under the EU GMPs
- Inspectors expectations
- Trending within/across techniques
- Challenges for trending non-routine tests

Stephen Young, MHRA, UK

SESSION 2 — OUT OF EXPECTATION RESULTS (OOE)

Position Talk: Out of Expectation Results (OOE) and Recommended Methods of Detection with Examples
- Single Out of Expectation Results – how to detect?
- Stable processes – unstable processes - does SPC apply?
- Nested Out of Expectation Results
- What level of investigation is necessary and appropriate?

Dr Bernd Renger, Bernd Renger Consulting, Germany
Member of the Analytical QC Working Group

SESSION 3 — TRENDING FOR PROCESS CONTROL OF VARIABLES (OOT 1)

Position Talk: Trending for Process Control of Variables (OOT 1)
Types of Control Charts that can be applied for Continuous Data Monitoring for manufactured batches and for analytical test control samples
- Turning data into information
- Types of run rules for Control Charts
- Periodic Review of Control Charts
- Out of Control, when should a CAPA be raised

Dr Lance Smallshaw, UCB Biopharma sprl, Belgium

Basics of SPC for Variables
- Statistical Basis of the Charts
- Setting the control limits
- Process Capability
- What if data are not normally distributed?

Dr Bianca Teodorescu, UCB Biopharma sprl, Belgium

SPC for Variables – Recommended Methods with Examples
- Types of charts
- Which chart to use and when?

Dr Bianca Teodorescu, UCB Biopharma sprl, Belgium

SESSION 4 — TRENDING FOR PROCESS CONTROL OF ATTRIBUTES (OOT 2)

Position Talk: Trending for Process Control of Attributes (OOT 2)
- Basic differences between attributes and variables
- Distributional requirements
- Control charts for attributes
- Applications for attribute data

Dr Christopher Burgess, UK, Chairman of the Analytical QC Working Group

Trending for Process Control of Attributes (OOT 2) – Recommended Methods with Examples
Speaker to be named

SESSION 5 — TRENDING FOR STABILITY DATA (OOT 3)

Position Talk: Trending for Stability Data (and Differences from SPC)

- Generation and maintenance of trend limits for stability test results
  - Calculated from existing historical stability data
- Simplified linear regression approach
  - Focus on stability indicating assays
  - Minimum data requirements
- More sophisticated random coefficients statistical model for analyzing more complex stability data sets
  - Calculating the 99% prediction interval
- Periodic review of trend limits
  - Reassessment of trend limits

Dr Peter Rauenbuehler, Roche, USA

Trending for Stability Data - Recommended Methods with Examples
N.N. OR
Dr Peter Rauenbuehler, Roche, USA

SESSION 6 — PULLING IT ALL TOGETHER / NEXT STEPS

Summary of the OOT Forum / Next Steps

- Review of Forum activities
- Challenges remaining
- Capturing the outputs
- Process of developing the OOT SOP

Dr Christopher Burgess, Chairman of the Analytical QC Working Group, UK

Dr Christopher Burgess, Burgess Analytical Consultancy Limited, UK - Chairman of the Analytical QC Working Group
Dr Burgess is a "Qualified Person" and a member of the European QP Association advisory board. He has been appointed to the United States Pharmacopoeia's Council of Experts 2010 to 2015 In addition, he is the chairman of the ECA Analytical Quality Control Group and a member of the Executive committee of European Compliance Academy.

Stephen Young, Head of Analytical Science Inspection, Enforcement and Standards Division, MHRA, UK
Stephen Young worked for ten years in the UK Pharmaceutical Industry, in various roles including analytical development, product stability and manufacturing technical support. He joined the Inspection, Enforcement and Standards division of MHRA in 2003 and currently provides leadership to the Agencies physico-chemical laboratories, which include the Pharmacopoeia and Regulatory Laboratory functions.

Dr Peter Rauenbuehler, Roche, USA
Peter Rauenbuehler, Ph.D., is a Senior Principal Technical Advisor, within Global Quality Systems & Processes at Genentech focused on laboratory policies.

Dr Bernd Renger, Bernd Renger Consulting, Germany Member of the Analytical QC Working Group
Dr Bernd Renger started at Hoechst AG. Since then, he has held several quality management positions at Mundipharma, Byk Gulden (now Takeda) and Baxter BioScience in Vienna and Vetter Pharma-Fertigung. He is a member of the European Compliance Academy (ECA) Advisory Board and Immediate Past Chair of the European QP Association.

Dr Lance Smallshaw, UCB Biopharma sprl, Belgium
Lance Smallshaw is Global Director of Analytical Strategy for NBEs at UCB Biopharma sprl in Belgium. Before that he was Senior Scientist at Eli Lilly and Company, having nearly 30 years experience in Analytical Development and QC Laboratories. He is one of the original conception members of the UK Pharmaceutical Analytical Science Group (Pasg) Biopharm. Working Group and currently is their honorary secretary..

Dr Bianca Teodorescu, UCB Biopharma sprl, Belgium
Bianca Teodorescu is Principal Statistician CMC Development in the Technical Operations department at UCB Biopharma sprl in charge of the non-clinical statistical team supporting the development department for biological and chemical entities (analytical, process, pharma), as well as the QC and manufacturing department.

Organisation / Contact

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Special Offer with Lufthansa – Discounted Travel for OOT Forum 2014 Attendees

As an ECA course or conference attendee, you will receive **up to 20% discounted travel fares** (according to availability). And as Lufthansa German Airlines offers a comprehensive global route network linking major cities around the world you will most likely be able to benefit from these special prices and conditions. And this is how it works: Once you registered for a course or conference you will receive a link together with your registration confirmation. Opening that link will take you to the Mobility Partner Program website where you can enter a code in the "Access to Event Booking" area you will also receive. This will take you into an online booking platform* that will automatically calculate the discount offered or provide you with an even better offer if another promotional fare is available. We look forward to welcoming at one of our next events – and we already wish you a pleasant flight!

*Please note that you may have to enable pop-ups on this site – otherwise the booking platform window will not open.

About CONCEPT HEIDELBERG

Founded in 1978, CONCEPT HEIDELBERG is the leading organiser of seminars on pharmaceutical production, quality control, quality assurance and GMP in Europe. This year more than 240 events will be organised by CONCEPT HEIDELBERG. The ECA Academy has entrusted CONCEPT HEIDELBERG with the organisation of its events.

ECA Education Course
Reduced Sampling/Reduced Testing
20-21 October 2014, Prague, Czech Republic

Directly before the OOT Forum on 20-21 October 2014, the ECA Education Course Reduced Sampling/Reduced Testing will take place with these topics:

- Regulatory Requirements for Sampling Procedures
- Design and Qualification of Sampling Areas for Incoming Goods Products
- Supplier Qualification: an important Prerequisite for Reduced Sampling and Reduced Testing
- How to Deal with Divergent Compendial Method Requirements (EP, USP, JP)
- Case Study I: How to Define Inspection Procedures for Packaging Materials (Primary and Secondary) in the Incoming Goods Control
- Case Study II: How to Define and Optimise Sampling and Testing Procedures for APIs and Excipients in the Incoming Goods Control
- Sampling and Documentation to make the Supplier liable for Defect Products

Further details will be discussed in a parallel session with 3 workshops.

Further information about this course can be received at www.gmp-compliance.org.

Participants who register simultaneously for both courses will receive a 350 € discount (not valid for EU GMP Inspectorates).

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Date
Wednesday, 22 October 2014, 09.00 – 18.30 h
(Registration and coffee 08.30 – 09.00 h)
Thursday, 23 October 2014, 08.30-16.00 h

Venue
Corinthia Hotel Prague
Kongresova 1
14069 Prague 4, Czech Republic
Phone + 420 261 191 111
Fax + 420 261 225 011

Conference Fees (per delegate plus VAT)
ECA Members EUR 1,590
APIC Members EUR 1,690
Non-ECA Members EUR 1,790
EU GMP Inspectorates EUR 895

The conference fee is payable in advance after receipt of invoice and includes conference documentation, dinner on the first day, lunch on both days and all refreshments. VAT is reclaimable.

Would you like to safe money?
If you register for the ECA Education Course "Reduced Sampling/Reduced Testing" from 20-21 October 2014 at the same time, you will receive a € 350 discount. This is not valid for EU GMP Inspectorates.

Conference Language
The official conference language will be English.

Accommodation
CONCEPT HEIDELBERG has reserved a limited number of rooms in the conference hotel. You will receive a room reservation form when you have registered for the event. Please use this form for your room reservation to receive the specially negotiated rate for the duration of your stay. Reservation should be made directly with the hotel. Early reservation is recommended.

Registration
Via the attached reservation form, by e-mail to info@concept-heidelberg.de by fax to +49 6221 / 84 44 34 .
Or you register online at www.gmp-compliance.org

If the bill-to-address deviates from the specification to the right, please fill out here:

CONCEPT HEIDELBERG
Postfach 10 17 64
Fax 06221/84 44 34

D-69007 Heidelberg

Reservation Form (Please complete in full)

+ 49 6221 84 44 34

☐ **OOT Forum 2014**
22 - 23 October 2014, Prague, Czech Republic

☐ Yes, I also want to participate in the ECA Education Course **Reduced Sampling/ Reduced Testing on 20-21 October 2014**, Prague, Czech Republic

☐ Mr ☐ Ms

Title, first name, surname

Company Department

Important: Please indicate your company's VAT ID Number **P.O. Number (if applicable)**

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General Terms of Business
If you cannot attend the conference you have two options:
1. We are happy to welcome a substitute colleague at any time.
2. If you have to cancel entirely, we must charge the following processing fees: Cancellation
- until 2 weeks prior to the conference 10 % of the registration fee.
- until 1 week prior to the conference 50 % of the registration fee.
- within 1 week prior to the conference 100 % of the registration fee.

CONCEPT HEIDELBERG reserves the right to change the materials, instructors, or speakers without notice or to cancel an event. If the event must be cancelled, registrants will be notified as soon as possible and will receive a full refund of fees paid.
CONCEPT HEIDELBERG will not be responsible for discount airfare penalties or other costs incurred due to a cancellation.
Terms of payment: Payable without deductions within 10 days after receipt of invoice.

Important: This is a binding registration and above fees are due in case of cancellation or non-appearance. If you cannot take part, you have to inform us in writing. The cancellation fee will then be calculated according to the point of time at which we receive your message. In case you do not appear at the event without having informed us, you will have to pay the full registration fee even if you have not made the payment yet. You are not entitled to participate in the conference until we have received your payment (receipt of payment will not be confirmed) (As of January 2012)