



WADA External Quality Assessment Scheme (EQAS) and its new statistical framework

Eurachem's 10th Workshop on Proficiency Testing in Analytical Chemistry, Microbiology and Laboratory Medicine

Dr Juris MEIJA
Research Officer, Metrology
National Research Council Canada

Dr Osquel BARROSO
WADA Senior Associate Director Science
& Medicine, Laboratories.

26-Sep-2023
Windsor, UK

WADA External Quality Assessment Scheme



WADA regularly distributes urine and blood test samples to Laboratories to continuously monitor their proficiency.

The **International Standard for Laboratories (ISL)** requires satisfactory EQAS performance in order to obtain and maintain WADA accreditation.

WADA EQAS: Main Objectives



To evaluate laboratory proficiency



To improve test result uniformity between laboratories



To provide educational opportunities

Regular WADA External Quality Assessment Schemes

Blind EQAS



- Labs are aware that the samples are EQAS samples
- Samples delivered by WADA EQAS sample provider
- 3 rounds of 5 samples per year

Double-Blind EQAS



- Samples are indistinguishable from routine athlete samples
- Samples delivered by Anti-Doping Organizations
- total of 5 samples per year

Educational EQAS



- Samples are provided in a variety of forms
- For educational purposes or data gathering
- 2-3 samples per year

Samples involving threshold substances are typically obtained from excretion studies. These are value-assigned, tested for batch homogeneity and storage/transportation stability.

Other WADA External Quality Assessment Schemes

EQAS for Athlete Biological Passport (ABP) blood samples

Conducted monthly in collaboration with the Quality Control Center Switzerland

EQAS for Laboratory accreditation decisions

Pre-probationary Test • Final Accreditation Test

Lifting of suspensions or analytical testing restrictions

EQAS for Major Events

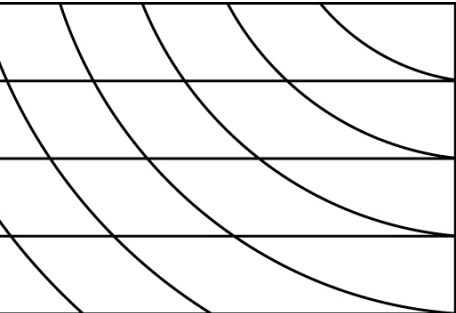
Olympic Games • Paralympic Games

EQAS for Laboratory on-site assessments

EQAS for Laboratory Investigations

Statistical evaluation of the quantitative results from **WADA EQAS** is going to change

Why revise the current statistical evaluation approach?

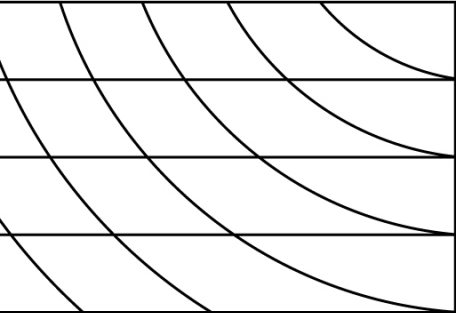
- | | |
|---|---|
|  | <p>❶ Currently, the calculations rely on empirical ‘robust’ statistical methods (ISO 13528:2015) instead of explicit statistical measurement models</p> |
|---|---|

$$s^* = 1,134 \sqrt{\sum_{i=1}^p (x_i^* - x^*)^2 / (p - 1)}$$

$$s^* = \frac{1}{0,798 \times p} \sum_{i=1}^p |x_i - med(x)|$$

$$r_p = \begin{cases} \frac{1}{p} \left[1,6019 + \frac{1}{p} \left(-2,128 - \frac{5,172}{p} \right) \right] & p \text{ odd} \\ \frac{1}{p} \left[3,6756 + \frac{1}{p} \left(1,965 + \frac{1}{p} \left(6,987 - \frac{77}{p} \right) \right) \right] & p \text{ even} \end{cases}$$

Why revise the current statistical evaluation approach?



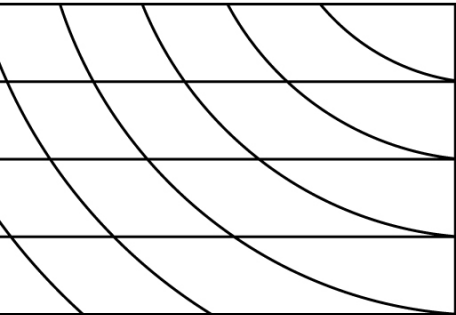
② Current evaluations do not always recognize the finer structure of the reported results

Carbon isotope ratio measurements (GC/C/IRMS) are reported with the associated measurement uncertainties, which are ignored

Steroid Profile and Threshold Substance measurements have proportional errors (most standard models assume constant errors)

The rounding of the specific gravity results that is imposed on the laboratories is not taken into account

Why revise the current statistical evaluation approach?



- ③ The decision making relies on the use of maximum allowed measurement uncertainty, not the reproducibility uncertainty

The EQAS 2023-1 Round provided a total of 1350 quantitative results with **zero cases of $|z| \geq 3.0$** , indicating the conservative nature of current data evaluation methods.

Mixed Effects Measurement Model

INSTITUTE OF PHYSICS PUBLISHING

Metrologia 41 (2004) 132–136

METROLOGIA

PII: S0026-1394(04)73994-8

Combining information from interlaboratory evaluations using a random effects model

Raghu N Kacker

National Institute of Standards and Technology, Gaithersburg, MD 20899-8910, USA

E-mail: raghu.kacker@nist.gov

Computational Statistics and Data Analysis 55 (2011) 1815–1827



ELSEVIER

Contents lists available at ScienceDirect

Computational Statistics and Data Analysis

journal homepage: www.elsevier.com/locate/csda

Laplace random effects models for interlaboratory studies

Andrew L. Rukhin*, Antonio Possolo

Statistical Engineering Division, National Institute of Standards and Technology, Gaithersburg, MD 20899, USA

We employ linear mixed effects measurement model which provides a coherent statistical framework for all results of a given analyte

– across all Laboratories and all EQAS Samples

Mixed Effects Measurement Model Example

Specific Gravity of Urine

Reported result by
each laboratory (*lab*)
for each sample (*s*)

Laboratory
bias

Measurement
error *

Rounding
error

$$X_{lab,s} = \mu_s + L_{lab} + E_{lab,s} + R_{lab,s}$$

Sample-specific
consensus value

* Estimated by combining the
results from multiple samples

Laplace
distribution

Gaussian
distribution

Uniform
distribution

Bayesian Framework for Interlaboratory Comparisons

IOP Publishing | Bureau International des Poids et Mesures
Metrologia 54 (2017) S34–S62

Metrologia

<https://doi.org/10.1088/1681-7575/aa6c0e>

Consensus building for interlaboratory studies, key comparisons, and meta-analysis

Amanda Koepke¹, Thomas Lafarge², Antonio Possolo² and Blaza Toman²

¹ National Institute of Standards and Technology, US Department of Commerce, Boulder, CO, United States of America

² National Institute of Standards and Technology, US Department of Commerce, Gaithersburg, MD, United States of America

Accred Qual Assur (2017) 22:1–19
DOI 10.1007/s00769-017-1247-y

GENERAL PAPER

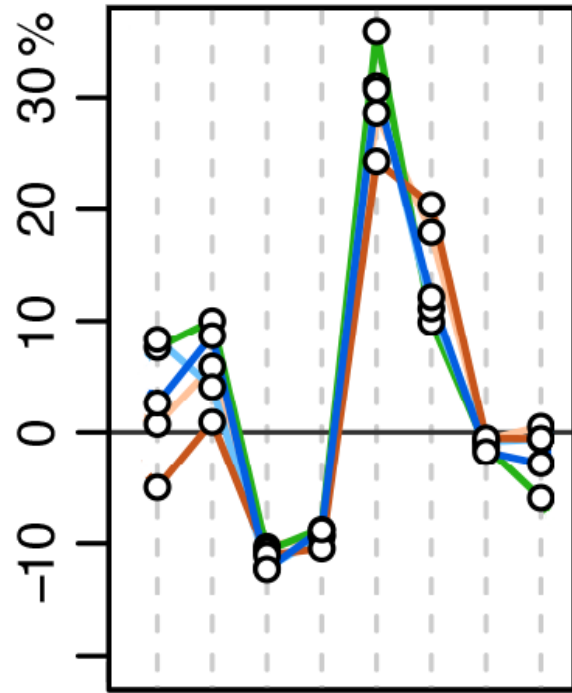
Bayesian framework for proficiency tests using auxiliary information on laboratories

Séverine Demeyer¹  · Nicolas Fischer¹

- ① provides flexible, model-based approach to data reduction
- ② allows to incorporate the expected measurement uncertainties ($u_c < u_{c_Max}$)
- ③ provides uncertainty distributions for all model parameters
- ④ naturally recognizes correlations between model parameters, for example, when evaluating the Laboratory bias

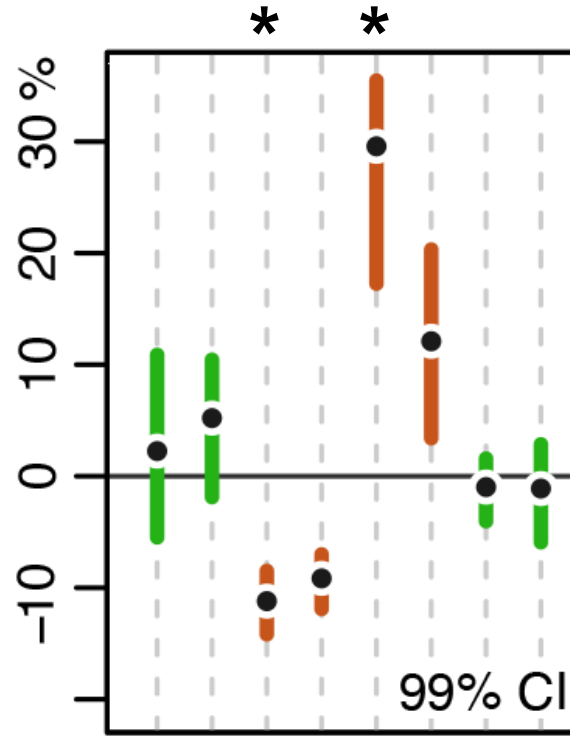
Example Analysis: Steroids in Urine

EQAS 2023-1
5 β -androstanediol



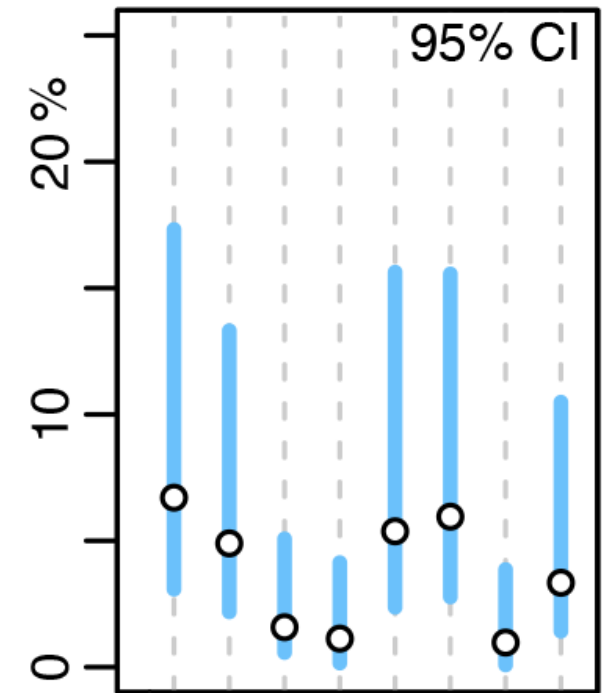
RESULTS FROM FIVE SAMPLES
(median centered)

=



LABORATORY BIASES

(4 Labs show significant bias)



MEASUREMENT ERRORS

Results of 8 out of 30 Labs are shown here

* Only 2 of these Labs are flagged by the conventional Rank Sum analysis

Conventional analysis does not provide MU evaluation

Custom Software for EQAS Data Reduction



EQAS Calculator (v.1)

Upload EQAS XLSX datafile

Browse... No file selected

Type of calculation

preliminary final

Fit model & make plots from datafile

Make plots from a complete datafile

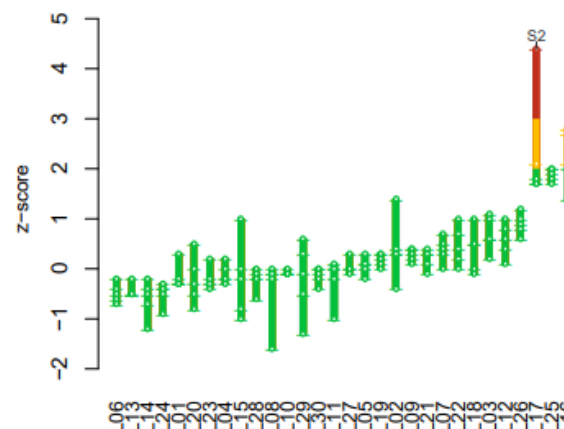
WADA EQAS Calculator (2023), v.1

wadaama

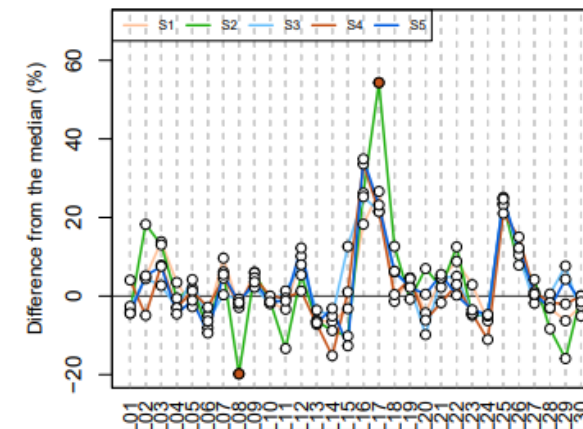


- 1 z-scores are calculated using model-based predictive standard deviations of the results
- 2 PDF and XLSX reports are generated programatically

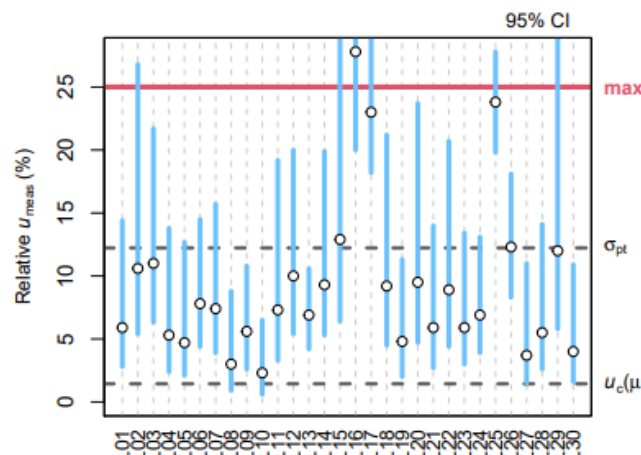
z-scores for all samples



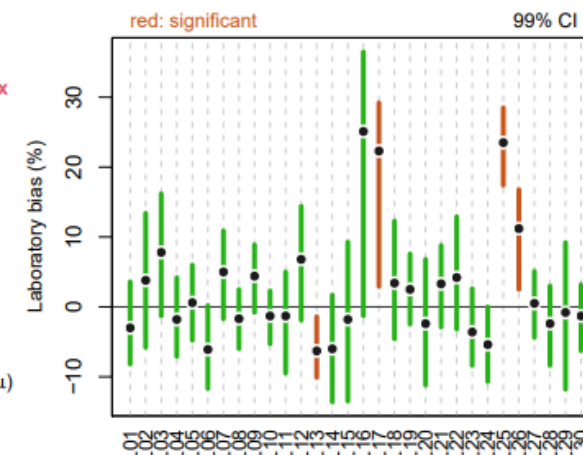
Normalized reported results



Evaluation of measurement uncertainty



Evaluation of laboratory bias



**The use of modern
statistical methods
and tools provides**

**more information from the same data
more advanced statistical modelling
more realistic uncertainty evaluations**

play true