



TOX

Toxicology Proficiency Testing Scheme

Scheme Description

LGC Standards Proficiency Testing

1 Chamberhall Business Park
Chamberhall Green
Bury
Lancashire
BL9 0AP
United Kingdom

Telephone: +44 (0) 161 762 2500
Fax: +44 (0) 161 762 2501
Email: tox@lgcgroup.com
Website: www.lgcstandards.com



0001

Toxicology Scheme Description

Record of issue status and modifications

ISSUE	ISSUE DATE	DETAILS	AUTHORISED BY
1	Feb 2012	Version 1 issued.	K. Morgan
2	Aug 2012	UKAS logo added to first page.	N. Stephenson
3	Jan 2013	2013 version of Scheme Description. Contains 2013 Quant Tox information	K Morgan
4	Feb 2014	2014-15 Quant Tox information included. New Toxic alcohol (including ethylene glycol) sample added. Addition of a lyophilised GHB in human urine sample. Expansion to Assigned Value Information in Appendix A. Addition of 'Methods' section.	K Morgan
5	August 2014	New sample information for Benzodiazepine Mix, Barbiturate Mix, Z-Drug Mix and Fentanyl	K Morgan
6	December 2014	2015-2016 Quant Tox information included and volume 10ml. Acetaminophen added to paracetamol listing. Spiking range of Carboxyhaemoglobin updated. Inclusion of subcontracting information in 'Test Materials' section.	K Morgan
7	June 2015	Removal of Barbiturate Mix and Fentanyl from sample types.	K Morgan
8	Jan 2016	2016-2017 Quant Tox information included. Removed Hard copy report information	K Morgan A McCarthy
9	Jan 2017	2017-2018 Quant Tox information included and updates made to concentration ranges	K Morgan
10	Jan 2018	2018-2019 Quant Tox information included. Addition of new Benzodiazepine sample (BNZB)	K Morgan
11	Sep 2018	Addition of a new sample for toxic alcohols in serum (TAS)	K Morgan

Notes:

Where this document has been translated, the English version shall remain the definitive version

Scheme Aims and Organisation

The primary aim of the Toxicology proficiency testing scheme (TOX) is to enable laboratories, performing the analysis of biological specimens for the range of compounds as stated under “test materials”, to monitor their performance and compare it with that of their peers. The TOX scheme also aims to provide information to participants on technical issues and methodologies relating to these analyses.

The TOX scheme year operates from April to March. Further information about TOX, including test material availability, round despatch dates and reporting deadlines, are available on the current TOX application form.

The operation of all schemes is supported by an Advisory Group consisting of members of the professional bodies, scheme participants, and others experienced in the field. The scheme reports on the performance of U.K. participants to the National Quality Assurance Advisory Panels for Chemical Pathology.

Test Materials

Details of test materials available in TOX are given in Appendix A. The test parameters are continually reviewed to ensure they meet the needs of current laboratory testing and regulatory requirements.

Samples are prepared using pre-screened human serum, pre-screened human blood and urine from donors.

The human serum is pooled and is obtained from donors who have verbally declared themselves drug free. The serum has been sterile-filtered, tested and found negative for:

- HEP B antigen
- HEP C antigen
- Combo HIV 1 and 2
- Syphilis
- Alanine transferase

Certificates of Analysis of the serum are retained at LGC.

The blood is obtained from the Welsh Blood Transfusion Service and is pre-screened.

The urine is supplied from donors.

Note: All test materials provided are intended for use as proficiency testing materials only and are not to be used for any other purposes.

Some aspects of the scheme, such as test material production, homogeneity testing and stability assessment, can from time to time be subcontracted. When subcontracting occurs, it is placed with a competent subcontractor and LGC is responsible for this work. The planning of the scheme, the evaluation of performance and the authorisation of the final report will never be subcontracted.

Statistical Analysis

Information on the statistics used in TOX can be found in the General Protocol and in the Scheme Report. Methods for determining assigned values and the values for SDPA used for individual samples are given in Appendix A.

Methods

Methods are listed in PORTAL. Please select the most appropriate method from the list. If none of the methods are appropriate, then please report your method as 'Other' and record a brief description in the Comments Section in PORTAL.

Results and Reports

TOX results are returned through our electronic reporting software, PORTAL, full instructions for which are provided by email. However, participants may request result submission forms on which to report and return results if they are unable to report through electronic means. This will incur an additional charge.

TOX reports will be available on the website within 10 working days of round closure. Participants will be emailed a link to the report when it is available.

Reports for case studies will be issued within 6 weeks of round closure.

APPENDIX A - Description of abbreviations used

Assigned Value (AV)

The assigned value may be derived in the following ways:

- From the robust mean (RMean). This is the median of participant results after the removal of test results that are inappropriate for statistical evaluation, e.g. miscalculations, transpositions and other gross errors. Generally, the assigned value will be set using results from all methods, unless the measurement is considered method-dependant, in which case the assigned value will be set by method as illustrated in the report tables.
For some analytes, where there is a recognised reference method for that type of measurement, this may be used as the assigned value for a particular analyte i.e. it would be applied to results obtained by any method.

Traceability: Assigned values which are derived from the participant results, or a subset of the results are not traceable to an international measurement standard. The uncertainty of assigned values derived in this way is estimated from the participant results, according to ISO 13528.

- From a formulation value (Formulation). This denotes the use of an assigned value derived from sample preparation details, where known and exact quantities of analyte have been used to prepare the sample.

Traceability: Assigned values calculated from the formulation of the test sample are traceable, via an unbroken metrological traceability chain, to an international measurement standard. The measurement uncertainty of the assigned value is calculated using the contributions from each calibration in the traceability chain.

- From a qualitative formulation (Qual Form). This applies to qualitative tests where the assigned value is simply based on the presence/absence of the analyte in the test material.

Traceability: Assigned values calculated from the qualitative formulation of the test sample are traceable to a certified reference standard or a microbiological reference strain.

- From expert labs (Expert). The assigned value for the analyte is provided by an 'expert' laboratory.

Traceability: Assigned values provided by an 'expert' laboratory may be traceable to an international measurement standard, according to the laboratory and the method used. The uncertainty of measurement for an assigned value produced in this way will be provided by the laboratory undertaking the analysis. Details of traceability and the associated uncertainty will be provided in the report for the scheme/round.

Range

This indicates the concentration range at which the analyte may be present in the test material.

SDPA

SDPA represents the 'standard deviation for proficiency assessment' which is used to assess participant performance for the measurement of each analyte. For all samples with the exception of the quantitative toxicology and case study the SDPA is based on a concentration dependent model derived from historic data. For the quantitative toxicology and case study the SDPA is based upon the RobustSD.

Units

This indicates the units used for the assessment of data. These are the units in which participants should report their results. For some analytes in some schemes participants may have a choice of which units to report their results, however, the units stipulated in this scheme description are the default units to which any results reported using allowable alternative results will be converted to.

DP

This indicates the number of decimal places to which participants should report their measurement results.

CDM

Concentration Dependent Model.

Toxicology**Sample: SM Quantitative - serum**

Participants will receive: 3 x 1.7 ml of human serum. Issued monthly. The serum contains 0.01% Bronidox and is filtered at 0.2µm.

Analyte	Method	Range	AV	SDPA	Units	DP
Ethanol	All	0 to 500	RMean	Fixed from CDM	mg per 100ml (mg%)	1
Paracetamol (Acetaminophen)	All	0 to 500	RMean	Fixed from CDM	mg/L	1
Salicylic Acid	All	0 to 1000	RMean	Fixed from CDM	mg/L	1

Sample: BLD Quantitative - blood

Participants will receive: 1 x 1.7ml of human blood. Issued monthly. The blood contains 0.1% gentamicin, 0.1% penicillin and 0.01% Bronidox.

Analyte	Method	Range	AV	SDPA	Units	DP
Ethanol	All	0 to 500	RMean	Fixed from CDM	mg%	1
Paracetamol (Acetaminophen)	All	0 to 500	RMean	Fixed from CDM	mg/L	1
Salicylic Acid	All	0 to 1000	RMean	Fixed from CDM	mg/L	1
Carboxyhaemoglobin	All	0 to 75	RMean	Fixed from CDM	%	1

Sample: URN Quantitative - urine

Participants will receive: 1 x 1.7ml of human urine. Issued monthly. The urine is heat treated at 60 degrees Centigrade for 1.5 hours prior to use. The urine contains 0.01% Bronidox.

Analyte	Method	Range	AV	SDPA	Units	DP
Ethanol	All	0 to 500	RMean	Fixed from CDM	mg%	1

Sample: CAS Toxicology case studies

Participants will receive: 1 x 7ml sample of serum and 1 x 20ml sample of urine accompanied by a short clinical or forensic scenario. The samples contain 1% sodium fluoride. Issued quarterly.

Laboratories are requested to analyse the specimens, providing information regarding qualitative and quantitative data. Participants are also required to make suitable comments regarding the interpretation and advice offered. The data reported and interpretation is assessed by an independent panel of experts and scores awarded. Comments are then included in the final reports. The aim of this sample is educational.

The Assigned Value will be based upon the Qual Form or RMean. For quantitative data the SPDA is based upon the Robust SD.

The default reporting units in the report will be those most suitable for the analyte. Participants will, however, have a choice of units in which to submit their results. The number of decimal places will be dependent upon the analyte and concentration. The Assigned Value is based upon the RMean and the SPDA is based upon the Robust SD.

Sample: TAK Quantitative Sample: Toxic alcohols

Participants will receive: 1 x 1.7 ml sample of whole blood containing 1% sodium fluoride. Two rounds per year.

Analyte	Method	Range	AV	SDPA	Units	DP
Ethanol	All	0 to 5000	RMean	RobustSD	mg/L	0
Methanol	All	0 to 3000	RMean	RobustSD	mg/L	0
Isopropylalcohol (IPA)	All	0 to 1500	RMean	RobustSD	mg/L	0
Acetone	All	0 to 5500	RMean	RobustSD	mg/L	0
Ethylene Glycol	All	0 to 8000	RMean	RobustSD	mg/L	0

Toxicology Scheme Description

Sample: TAS * **Quantitative Sample: Toxic alcohols in serum**

Participants will receive: 1 x 1.7 ml sample of serum containing 1% sodium fluoride. Two samples, two rounds per year so that a range can be covered each round.

Analyte	Method	Range	AV	SDPA	Units	DP
Ethanol	All	0 to 5000	RMean	RobustSD	mg/L	0
Methanol	All	0 to 3000	RMean	RobustSD	mg/L	0
Isopropylalcohol (IPA)	All	0 to 1500	RMean	RobustSD	mg/L	0
Acetone	All	0 to 5500	RMean	RobustSD	mg/L	0
Ethylene Glycol	All	0 to 8000	RMean	RobustSD	mg/L	0

*not currently included in LGC's UKAS Scope of Accreditation

Sample: GHB **Quantitative Sample: Gammahydroxybutyrate**

Participants will receive: 3 x 2ml lyophilised human urine. Two rounds per year.

Analyte	Method	Range	AV	SDPA	Units	DP
Gammahydroxybutyrate	All	0 to 500	RMean	RobustSD	mg/L	1

Toxicology Scheme Description

Sample: QT

Whole blood quantitative toxicology

Participants will receive: 2 x 10ml blood samples. Issued quarterly. Samples contain analytes which are pre-defined utilising our Advisory Group. The blood contains 1% sodium fluoride. The samples will be comprised as follows:

APRIL 2018	
QT1	QT2
Benzoyllecgonine	Diazepam
Cocaine	Midazolam
Free morphine	Bromazepam
Free codeine	Methadone
Pregabalin	EDDP
JULY 2018	
QT1	QT2
Sertraline	Hydroxychloroquine
Citalopram	Promethazine
Desmethylcitalopram	Tramadol
Fluoxetine	O-Desmethyltramadol
Quetapine	Sildenafil
OCTOBER 2018	
QT1	QT2
THC	Nordiazepam
THC-COOH	Oxazepam
Amfetamine	Lorazepam
Methamphetamine	Chlordiazepoxide
MDMA	Mephedrone
JANUARY 2019	
QT1	QT2
Amfetamine	Trazodone
Diphenhydramine	Amitriptyline
MDMA	Nortriptyline
MDA	Clomipramine
Fentanyl	Olanzapine

The default reporting units in the report are **µg/L**. Participants will, however, have a choice of units in which to submit their results. The number of decimal places will be dependent upon the analyte and concentration. The Assigned Value is based upon the RMean and the SPDA is based upon the Robust SD.

Many of the analytes included in the scheme are there at the request of participants.

Toxicology Scheme Description

Sample: BNZ*

Quantitative Sample: Benzodiazepine Mix

Participants will receive: 2 x 4 ml samples of lyophilised human serum. Four rounds per year.

Analyte	Method	Range	AV	SDPA	Units	DP
Diazepam	All	0 to 5000	RMean	RobustSD	µg/L	0
Nordazepam	All	0 to 3000	RMean	RobustSD	µg/L	0
Temazepam	All	0 to 2000	RMean	RobustSD	µg/L	0
Oxazepam	All	0 to 2000	RMean	RobustSD	µg/L	0
Nitrazepam	All	0 to 3000	RMean	RobustSD	µg/L	0

*not currently included in LGC's UKAS Scope of Accreditation

Sample: BNZB*

Quantitative Sample: Benzodiazepine Mix B

Participants will receive: 2 x 4 ml samples of lyophilised human serum. Four rounds per year.

Analyte	Method	Range	AV	SDPA	Units	DP
Alprazolam	All	0 to 500	RMean	RobustSD	µg/L	0
Bromazepam	All	0 to 500	RMean	RobustSD	µg/L	0
Clonazepam	All	0 to 200	RMean	RobustSD	µg/L	0
Lorazepam	All	0 to 800	RMean	RobustSD	µg/L	0
Midazolam	All	0 to 500	RMean	RobustSD	µg/L	0

*not currently included in LGC's UKAS Scope of Accreditation

Sample: ZMIX*

Quantitative Sample: Z-Drug Mix

Participants will receive: 2 x 4 ml samples of lyophilised human serum. Four rounds per year.

Analyte	Method	Range	AV	SDPA	Units	DP
Zolpidem	All	0 to 1000	RMean	RobustSD	µg/L	0
Zaleplon	All	0 to 250	RMean	RobustSD	µg/L	0
Zopiclone	All	0 to 1000	RMean	RobustSD	µg/L	0

*not currently included in LGC's UKAS Scope of Accreditation