# Clinical Chemistry IMPC\_CBC\_002

#### **Purpose**

Clinical chemistry determines biochemical parameters in plasma including enzymatic activity, specific substrates and electrolytes.

Ontological description: MP:0001545 – blood physiology abnormalities.

#### **Experimental Design**

• Minimum number of animals: 7M + 7F

• Age at test: Week 16

• Sex: We would expect the results of this test to show sexual dimorphism

#### **Equipment**

- 1. Clinical chemistry analyser
- 2. Vortex
- 3. Refrigerated centrifuge
- 4. Eppendorf tubes
- 5. Pipettes (200-1000 ul)

#### **Procedure**

Set up the clinical chemistry analyser and perform QC analyses of the control reagents in accordance with the equipment guidelines.

#### Sample collection and preparation:

- a. Collect the appropriate volume of blood required (160-200l of plasma), for the clinical chemistry analyser being used for assessment, in gel tube containing lithium Heparin with the relevant blood collection procedure (see IMPC protocol Blood collection by retro-orbital puncture). Time of day for collection is in the morning, starting no earlier than 07:30.
- b. Keep whole blood samples in a bag on wet ice until centrifugation. Centrifuge for 10 minutes at 5000 x g in a refrigerated centrifuge set at 8°C. If plasma samples cannot be analysed immediately, keep them in the fridge until analysis.
- c. Analysis of samples is optimally done on the day of collection. When not possible the plasma samples can be stored at 2-8°C. If samples require storage for > 48 hours,

- freeze plasma at -20 °C in single aliquots. All samples are allowed to come to room temperature prior to analysis.
- d. Use plasma samples undiluted or diluted to a ratio of 1:2 with deionised water if the volume is insufficient.
- e. Plasma samples that were frozen or stored in the fridge should be vortexed briefly and centrifuged again at ~5000 x g for 2-3 minutes immediately prior to analysis. If necessary, remove fibrin clots using a wooden applicator.

#### Analysis:

Samples that produce results that lie outside the linear range for a specific assay have to be re-tested. In some cases it may be necessary to dilute samples with water to bring test results into range.

#### **Notes**

Blood collection for Clinical Chemistry and Hematology is usually performed as a non-fasting, terminal procedure but can be performed as a non-terminal procedure under certain circumstances. Mice from the terminal procedure may be used for subsequent gross pathology and other procedures included in terminal assessments. Whole blood (for Hematology) and plasma (for Clinical Chemistry) require different collection tubes so two independent samples are required from each mouse.

The information about the date of the experiment, that is the date when the measurement is performed, is an important parameter which is to be submitted in the Experiment xml file (dateOfExperiment="2013-02-28").

**Dilution**. Dilution of blood is highly discouraged, but is allowed when the total necessary amount is not obtained. If dilution is necessary then the assays should be done in one run.

Hemolysis. Two fields currently exist to capture metadata information about the hemolysis status in the clinical chemistry plasma samples. The first is the LIH Hemolysis severity score which can only be performed by clinics who run one of the Beckman Coulter AU-series of analysers. Such clinics are encouraged to capture and submit the hemolysis score of the LIH test in this field. Clinics who do not have an AU analyser are encouraged to use the second /alternative field which is simply titled Hemolysis. Simply enter "slight", "moderate", or "marked" based on whether the sample is visibly haemolysed or not. Provision of this information is not compulsory and it is suggested that any clinic completes at least one field or the other (not both).

#### Data QC

- 1. Plasma samples must be free of Fibrin clots in order to be analysed.
- 2. Badly haemolysed samples should be discarded.
- 3. Each morning, all parameters are tested with control sera (see ESLIM\_015\_001\_Annex\_3: Controls for biochemistry on AU400). Some parameters are tested with control serum level 1 (Beckman Coulter System Reagent, ODC0003)

- and control serum level 2 (Beckman Coulter System Reagent, ODC0004), which consists of lyophilised human plasma with a normal and a pathological concentration. Other parameters are tested with specific controls from other suppliers.
- 4. Controls are thawed and vortexed before utilisation and loaded according to the analyser's display. Control values must lie within the acceptable range indicated by the manufacturer, otherwise the specific tests must be recalibrated and specific measurements repeated. Controls can be stored in 200l aliquots at -20°C for up to 1 week.

#### **Metadata and examples**

Metadata	Example
Equipment ID	ID of the machine used when more than 1 is used having same model and manufacturer. E. g. machine 1, machine 2, machine Minnie, machine Mickey Mouse, etc.
Equipment manufacturer	Manufacturer of the equipment. E.g. Olympus Diagnostics.
Equipment model	Model of the equipment. E.g. AU400
Blood collection tubes	The tubes used for blood collection. E.g. Sarstedt Li-Heparin gel tubes or Kabe Labortechnik Lithium heparin coated tubes.
Anaesthesia used for blood collection	The drug used for anaesthesia during blood collection. E. g. Isofluorane.
Method of blood collection	Concise description of the method used for blood collection. E.g. retro-orbital puncture.
Anticoagulant	Anticoagulant drug used for blood collection. E. g. Li-Heparin.
Samples kept on ice between collection and analysis	Yes/No.

Storage temperature from blood collection till measurement	E.g. 2°C
Sample status	Indicate if the sample were frozen (analysis on the same day of collection not possible) or fresh (analysis on the same day of collection). E.g Fresh/Frozen.
Plasma dilution	Dilution is highly discouraged but if necessary indicate here. E.g. "No dilution" or 1:2. Note that results submitted to DCC are assumed to be already corrected for any dilutions made.
ID of blood collection SOP	ID of the protocol followed for blood collection. Can be a center specific protocol. E.g. ESLIM_024_001.
Date and time of blood collection	Time of day for collection is in the morning, starting no earlier than 07:30. E.g. Year, month, day, time.
Date of measurement	The day of blood analysis. Year, month, day.
Hemolysis status	If no AU analyser score is provided, indicate here the gauged degree of hemolysis. E.g. slight/moderate/marked.
Blood collection experimenter ID	An ID of any format to be used coherently both inside the same procedure and for all procedures indicating the experimenter who collected the blood. E.g. Harw_001, or 1/2/3.
Blood analysis experimenter ID	An ID of any format to be used coherently both inside the same procedure and for all procedures indicating the experimenter who analyzed the blood. E.g. Harw_001, or 1/2/3.
Date equipment last calibrated	Most recent date in which the equipment (or any part of) used in the procedure was subject to a calibration event.
Date and time of sacrifice	The date and time when the mouse is sacrified.

#### **Parameters and Metadata**

#### **Sodium** IMPC\_CBC\_001\_001 | v1.3

simpleParameter

Req. Analysis: false Req. Upload: false Is Annotated: true

Unit Measured: mmol/l

**Description:** sodium

## Potassium IMPC\_CBC\_002\_001 | v1.3

simpleParameter

Req. Analysis: false Req. Upload: false Is Annotated: true

Unit Measured: mmol/l

**Description:** potassium

#### Chloride IMPC\_CBC\_003\_001 | v1.4

simpleParameter

Req. Analysis: false Req. Upload: false Is Annotated: true

Unit Measured: mmol/l		
Description: chloride		
Urea (Blood Urea simpleParameter	a Nitrogen - BUN)	MPC_CBC_004_001   v1.5
Req. Analysis: false	Req. Upload: true	Is Annotated: true
Unit Measured: mg/dl		
Description: urea		
Creatinine IMPC_CI simpleParameter	BC_005_001   v1.5	
Req. Analysis: false	Req. Upload: true	Is Annotated: true
Unit Measured: mg/dl		
<b>Description:</b> creatinine_er	nzymatic_method_preferred_	

# Total protein IMPC\_CBC\_006\_001 | v1.2

simpleParameter

Req. Analysis: false Req. Upload: true Is Annotated: true

Unit Measured: g/l				
Description: total_protein				
Albumin IMPC_CBC_007_001   v1.2 simpleParameter				
Req. Analysis: false	Req. Upload: true	Is Annotated: true		
Unit Measured: g/l				
Description: albumin				
Total bilirubin IMPC_CBC_008_001   v1.4 simpleParameter				
Req. Analysis: false	Req. Upload: true	Is Annotated: true		
Unit Measured: mg/dl				
Description: total_bilirubin				

# Calcium IMPC\_CBC\_009\_001 | v1.5

simpleParameter

Req. Analysis: false Req. Upload: true Is Annotated: true

Unit Measured: mg/dl		
Description: calcium		
Phosphorus IMPC_CR	BC_010_001   v1.6	
simpleParameter	'	
Req. Analysis: false	Req. Upload: true	Is Annotated: true
Unit Measured: mg/dl		
Description: phosphate		
Iron IMPC_CBC_011_001 simpleParameter	v1.5	
Req. Analysis: false	Req. Upload: false	Is Annotated: true
Unit Measured: mg/dl		
Description: iron		

# Aspartate aminotransferase IMPC\_CBC\_012\_001 | v1.2

simpleParameter

Req. Analysis: false Req. Upload: true Is Annotated: true Unit Measured: U/I **Description:** aspartate\_aminotransferase Alanine aminotransferase IMPC\_CBC\_013\_001 | v1.2 simpleParameter Req. Analysis: false Req. Upload: true Is Annotated: true Unit Measured: U/L **Description:** alanine\_aminotransferase Alkaline phosphatase IMPC\_CBC\_014\_001 | v1.2 simpleParameter Req. Analysis: false Req. Upload: true Is Annotated: true Unit Measured: U/I Description: alkaline\_phosphatase

Req. Analysis: false Req. Upload: true Is Annotated: true

Unit Measured: mg/dl

**Description:** total\_cholesterol

#### HDL-cholesterol IMPC\_CBC\_016\_001 | v1.4

simpleParameter

Req. Analysis: false Req. Upload: true Is Annotated: true

Unit Measured: mg/dl

**Description:** hdl\_cholesterol

## Triglycerides IMPC\_CBC\_017\_001 | v1.4

simpleParameter

Req. Analysis: false Req. Upload: true Is Annotated: true

Unit Measured: mg/dl

**Description:** triglycerides

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Glucose IMPC\_CBC\_018\_001 | v1.5

**Reg. Analysis:** false **Reg. Upload:** true Is Annotated: true Unit Measured: mg/dl **Description:** glucose LIH (Hemolysis Severity - available on AU analysers) IMPC\_ CBC\_019\_001 | v1.3 simpleParameter Reg. Analysis: false Reg. Upload: false Is Annotated: false **Description:** lih\_hemolysis\_severity\_available\_on\_au\_analysers\_ Fructosamine IMPC\_CBC\_020\_001 | v1.2 simpleParameter Req. Analysis: false Req. Upload: false Is Annotated: true Unit Measured: umol/l **Description:** fructosamine

Req. Analysis: false Req. Upload: false Is Annotated: true Unit Measured: U/I **Description:** lipase Lactate dehydrogenase IMPC\_CBC\_022\_001 | v1.2 simpleParameter Req. Analysis: false Req. Upload: false Is Annotated: true Unit Measured: U/I **Description:** lactate\_dehydrogenase Alpha-amylase IMPC\_CBC\_023\_001 | v1.2 simpleParameter Req. Analysis: false Req. Upload: false Is Annotated: true Unit Measured: U/I **Description:** alpha\_amylase

# **UIBC (unsaturated iron binding capacity)** IMPC\_CBC\_024\_001 | v1

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simpleParameter

Req. Analysis: false Req. Upload: false Is Annotated: true

Unit Measured: umol/l

**Description:** uibc\_unsaturated\_iron\_binding\_capacity\_

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#### LDL-cholesterol IMPC\_CBC\_025\_001 | v1.3

simpleParameter

Req. Analysis: false Req. Upload: false Is Annotated: true

Unit Measured: mg/dl

**Description:** Idl\_cholesterol

## Free fatty acids IMPC\_CBC\_026\_001 | v1.4

simpleParameter

Req. Analysis: false Req. Upload: false Is Annotated: true

Unit Measured: mmol/l

Description: free fatty acids

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#### Glycerol IMPC\_CBC\_027\_001 | v1.4

simpleParameter

Unit Measured: umol/l

Description: uric\_acid

Req. Analysis: false Req. Upload: false Is Annotated: true Unit Measured: mmol/l **Description:** glycerol Creatine kinase IMPC\_CBC\_028\_001 | v1.2 simpleParameter Req. Analysis: false Req. Upload: false Is Annotated: true Unit Measured: U/I **Description:** creatine\_kinase Uric acid IMPC\_CBC\_029\_001 | v1.2 simpleParameter Req. Analysis: false Req. Upload: false Is Annotated: true

#### Ferritin IMPC\_CBC\_030\_001 | v1.3

simpleParameter

Req. Analysis: false Req. Upload: false Is Annotated: true

Unit Measured: ng/ml

**Description:** ferritin

## Transferrin IMPC\_CBC\_031\_001 | v1.2

simpleParameter

Req. Analysis: false Req. Upload: false Is Annotated: true

Unit Measured: mg/dl

**Description:** transferrin

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## C-reactive protein IMPC\_CBC\_032\_001 | v1.0

simpleParameter

Reg. Analysis: false Reg. Upload: false Is Annotated: true

Unit Measured: mg/l

**Description:** c reactive protein

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## Equipment ID IMPC\_CBC\_033\_001 | v1.0

procedureMetadata

Req. Analysis: false Req. Upload: true Is Annotated: false

**Description:** equipment\_name

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## Equipment manufacturer IMPC\_CBC\_034\_001 | v1.0

procedureMetadata

Req. Analysis: true Req. Upload: true Is Annotated: false

**Description:** equipment\_manufacturer

Options: Cobas, Olympus Diagnostics, Beckman Coulter, Hitachi, JEOL (Siemens), Roche,

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## Equipment model IMPC\_CBC\_035\_001 | v1.0

procedureMetadata

Req. Analysis: true Req. Upload: true Is Annotated: false

**Description:** equipment\_model

Options: Integra 400 Plus, AU 400, AU 480, 7020, JCA-BM2250 (Advia 2400), Hitachi 917,

AU 680, UniCel 600 Pro, JCA-BM6070,

## Anesthesia used for blood collection IMPC\_CBC\_036\_001 | v1.0

procedureMetadata

Req. Analysis: true Req. Upload: true Is Annotated: false

**Description:** anesthesia used for blood collection

Options: Gas anaesthesia with Isofluorane,

Injection narcosis with Ketamine (100mg/kg)/Xylazine (10mg/kg),

Injection narcosis with Ketamine (100mg/kg)/ Xylazine (10mg/kg)/Antipamezole (Antisedan, 1mg/kg),

Injection narcosis with Ketamine (110mg/kg)/Xylazine (11mg/kg),

Injection narcosis with Ketamine (110mg/kg)/Xylazine (11mg/kg)/ Antipamezole (Antisedan, 1mg/kg),

Injection narcosis with Tribromoethanol (Avertin),

Injection narcosis with Sodium Pentobarbital (Pentobarb, 0.1ml),

Injection narcosis with Sodium Pentobarbital (Euthatal), No,

Injection narcosis with Ketamine (137mg/kg)/Xylazine (6.6mg/kg),

Injection narcosis with Sodium Pentobarbital (Somnopentyl),

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#### Method of blood collection IMPC\_CBC\_037\_001 | v1.0

procedureMetadata

Req. Analysis: true Req. Upload: true Is Annotated: false

**Description:** method\_of\_blood\_collection

Options: Cardiac puncture, Retro-orbital puncture, Heart puncture, Jugular vein, Tail vein,

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#### Anticoagulant IMPC\_CBC\_038\_001 | v1.1

procedureMetadata

Req. Analysis: false Req. Upload: true Is Annotated: false

**Description:** anticoagulant

Options: No, Lithium Heparin, Sodium Heparin, Heparine,

# Samples kept on ice between collection and analysis IMPC\_

CBC\_042\_001 | v1.1

procedureMetadata

Req. Analysis: true Req. Upload: true Is Annotated: false

**Description:** samples\_kept\_on\_ice\_between\_collection\_and\_analysis

Options: Yes, No,

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#### Sample status IMPC\_CBC\_043\_001 | v1.1

procedureMetadata

Req. Analysis: false Req. Upload: true Is Annotated: false

**Description:** sample\_status

Options: Fresh, Frozen, Fresh and frozen,

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## Sample dilution IMPC\_CBC\_044\_001 | v1.2

procedureMetadata

Req. Analysis: false Req. Upload: true Is Annotated: false

**Description:** plasma dilution

Options: Neat plasma, 1:2, Yes (by Equipment, automatically), Varies, Neat serum, 1:3, 1:4,

1:5,

#### ID of blood collection SOP IMPC CBC 045 001 | v1.1

procedureMetadata

Req. Analysis: false Req. Upload: true Is Annotated: false

**Description:** id\_of\_blood\_collection\_sop

Options: ESLIM\_024\_001, sop.inv.019, RIKENMPP\_004a\_003, PHENO\_CBC, sop.inv.063,

#### Date and time of blood collection IMPC CBC 046 001 | v1.2

procedureMetadata

Req. Analysis: false Req. Upload: true Is Annotated: false

**Description:** date\_and\_time\_of\_blood\_collection

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#### Hemolysis status IMPC\_CBC\_048\_001 | v1.1

procedureMetadata

Reg. Analysis: false Reg. Upload: false Is Annotated: false **Description:** hemolysis\_status **Options:** Slight, Moderate, Marked, None, Blood collection experimenter ID IMPC\_CBC\_049\_001 | v1.1 procedureMetadata Req. Analysis: false Req. Upload: true Is Annotated: false Date equipment last calibrated IMPC\_CBC\_050\_001 | v1.2 procedureMetadata Req. Analysis: false Req. Upload: false Is Annotated: false

#### Blood collection tubes IMPC\_CBC\_039\_001 | v1.1

procedureMetadata

Req. Analysis: false Req. Upload: false Is Annotated: false

Options: Sarstedt Li-Heparin gel tubes, Kabe Labortechnik Lithium heparin coated tubes,

Kabe Labortechnik 1000ul Lithium Heparin,

BD Microtainer Lithium Heparin/PST Gel Blood Tube,

TERUMO CAPIJECT Lithium heparin coated tubes, Eppendorf 1.7ml,

BD Microtainer Lithium Heparin Tube,

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#### Date and time of sacrifice IMPC\_CBC\_040\_001 | v1.1

procedureMetadata

Req. Analysis: false Req. Upload: true Is Annotated: false

# Storage temperature from blood collection till measurement IMPC\_CBC\_041\_001 | v1.3

procedureMetadata

Req. Analysis: true Req. Upload: true Is Annotated: false

Unit Measured: C

Options: 2, 18-22, 4, -80,

#### Blood analysis experimenter ID IMPC\_CBC\_051\_001 | v1.0

procedureMetadata

Req. Analysis: false Req. Upload: true Is Annotated: false

Glycosilated hemoglobin A1c (HbA1c) IMPC\_CBC\_052\_001 | v1.3 simpleParameter Req. Analysis: false Req. Upload: false Is Annotated: true Unit Measured: % Thyroxine IMPC\_CBC\_053\_001 | v1.2 simpleParameter Req. Analysis: false Req. Upload: false Is Annotated: true Unit Measured: ug/dl Magnesium IMPC\_CBC\_054\_001 | v1.5 simpleParameter Req. Analysis: false Req. Upload: false Is Annotated: true Unit Measured: mg/dl

Difficult bleed IMPC\_CBC\_055\_001 | v1.0

#### procedureMetadata