

Fast Track to Metabolic Biomarker Signatures:

Absolute/DQ® p180 Kit



Targeted Metabolomics Designed for Clinical Utility

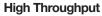


The Fast and Reproducible Way to Targeted Metabolomics Results

The Absolute IDQ® p180 Kit quantifies a broad metabolite spectrum, reflecting diverse pathophysiological processes such as inflammation, mitochondrial dysfunction, or cell cycle control. In hundreds of research projects, the kit has provided circulatory biomarker signatures for diagnosis, prognosis or prediction in multi-factorial diseases.

This **standardized** solution is designed to support targeted metabolomics studies in large-scale epidemiological and clinical cohorts. It is ideal for research projects where long-term reliability and throughput matter. The minimal sample volume needed enables longitudinal analyses in small animal models.

















Minimal Sample



Why Metabolomics? Why Choose a Targeted Approach?

The metabolome integrates an individual's genetic background, aging, lifestyle and environmental factors. As it closely reflects the phenotype, the metabolome can provide important information about the state of a cell, organ, or organism. Most metabolic pathways are identical in humans and animal models, also making metabolomics ideally suited for translational research approaches.

The biochemical diversity of metabolic substances makes it impossible to cover the full metabolome with a single analytical method targeted coverage of carefully selected pathways is the key to successful metabolomics projects. Focusing on pre-defined analytes allows analyzing analytically challenging metabolites confidently & quantitatively.

Covered Metabolite Classes and Relevance in Main Application Areas

The up to 188 analytes covered by the Absolute/DQ® p180 Kit span four main areas: amino acid (21 amino acids and 21 biogenic amines), glucose (sum of hexoses), fatty acid (40 acylcarnitines) and lipid metabolism (90 glycerophospholipids and 15 sphingolipids). Those

reflect a vast number of biological functions and rate limiting enzymes. The table below exemplarily shows the relevance of selected metabolites in main application areas.

Indication/ Metabolites	Amino Acids and Biogenic Amines	Acylcarnitines	Glycerophospho- and Sphingolipids	Hexoses
Oncology	Serine: link between glycolysis, lipid and nucleotide metabolism Leucine, arginine: mTOR activation	Tumor metabolism: increased usage of fatty acids as nutrients	Growth and proliferation: lipids required as building blocks Cancer risk: lipid metabolism modulates probability of future disease	Glycolysis: strongly upregulated in many cancers
Cardiology	Phenylalanine to tyrosine ratio: marker for endothelial function NO signaling: involved in angiogen- esis and cardiomyocite homeostasis	Cardiomyocytes: fatty acid oxidation is the main source of energy	Atherosclerotic plaque inflammation: lipid breakdown by PLA2 involved	Insulin resistance: major risk factor for cardiovascular disease
Diabetes	Branched chain amino acids: involved iin insulin resistance Tryptophan, kynurenine: role of immune pathways in diabetes	Mitochondrial function: affects inflammation and lipid accumulation in the liver	Lipoproteins: Metformin alters LDL cholesterol levels via lipid metabolism Low-grade inflammation: lipids are inflammatory mediators	Disturbed glucose homeostasis: major hallmark
+ + (Neurology	Neurotransmitter function: e.g. glutamate, dopamine, serotonin Urea cycle deregulation: probable involvement in neurodegeneration	Neurodegeneration: evidence for involvement of mitochondrial metabolism	Myelin sheaths: sphingo- lipids are a major compo- nent	Insulin resistance: contributes to devel- opment of Alzheimer's disease

Table: Indications and relevant Metabolites (relevant literature available upon request)



Key Benefits

- Comprehensive information about the physiological state of an organism
 - Due to broad analyte coverage (up to 188 metabolites)
- **Highly reproducible** due to standardization and quality control
 - Proven in a multinational ring trial
- Quantitative easy to handle data format
- **High throughput** (up to 500 samples per week and instrument)
- Saving precious sample material
 - Only 10 μL needed for most biofluids, fully validated for human plasma
 - Strong experience with many biological matrices
- Ready to use no method development required



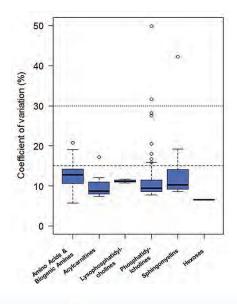
Standardization, Throughput and Proven Reproducibility

Building upon previous studies is drastically simplified by reproducibility of analytical procedures, and the possibility to acquire data quickly.

The Absolute/DQ® p180 Kit is a widely used targeted metabolomics solution. Having proven inter-platform, inter-laboratory reproducibility and providing high throughput, it can significantly increase research output and confidence in the findings.

The Absolute IDQ® p180 Kit has been used in hundreds of metabolomics projects, spanning all major fields of healthcare spending. We will be happy to inform you about available results from research questions related to yours.

Figure: Inter-Platform Reproducibility of the Absolute/DQ® p180 Kit (internal reproducibility data; unpublished)



How to Generate Data

The Kit relies on triple quadrupole mass spectrometry – a technology that is in routine clinical use, for example in newborn screening, pharmacology and toxicology. The table below shows differ-

ent options how data can be generated. Contact us to discuss how to move forward.

Use of Kits in Your Laboratory Collaboration **Metabolic Phenotyping Services** • You have a suitable mass spectrometer. The full analytical workflow is performed at You intend to do measurements through a Biocrates' laboratories. partner laboratory. • Biocrates provides the Kit with validated · Biocrates can provide kits and technical sup-· Biocrates ensures short turnaround times and method, reagents, and required software. port directly to your collaborator. can provide biostatistical support. Contact us for a detailed list of requirements. We will be pleased to connect you with potential Contact us for further information.

Workflow (for measurement in your own laboratory)

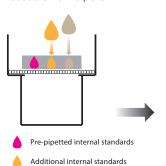
Sample Preparation (3-5 hours)

First Steps

Register samples in Met/DQ® software (provided with the Kit) and use Testmix to ensure suitable system conditions.

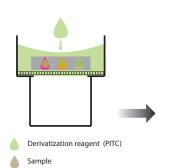
Loading

In addition to pre-pipetted internal standards, further internal standards and the sample are loaded onto the plate.



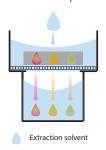
Derivatization

Analyzing biogenic amines and amino reliably requires derivatization.



Extraction

Metabolites are extracted with organic solvent, for subsequent mass spectrometric analysis in two steps.



Sample Processing (24-34 hours)*

Run Samples

First, the liquid chromatography (LC-MS) part for amino acids and biogenic amines is run, followed by the flow injection (FIA-MS) part for all other metabolites.

* Depending on analytical platform

Quantify

Quantitative LC-MS data is importet into Met**IDQ**® software, FIA-MS data is quantified by Biocrates' software.

Data Analysis (1 hour)**

Quality Control DQ® performs an

Met*IDQ*® performs an automated quality assessment. Additional manual checks can be made.

Statistics and Export

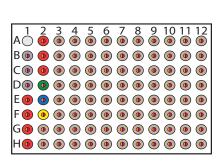
Met/DQ® supports basic statistics, as well as export of data. Upgrades for Met/DQ® are available, enabling extended data analysis within the software package.

Plate Layout

The Absolute $IDQ^{@}$ p180 Kit comes in a convenient 96 well format, for the measurement of up to 82 samples.

A minimum of 16 wells per plate is reserved for zero sample, 3 blank samples, 7 calibration standard levels (for amino acids and biogenic amines), and a minimum of 3 quality control samples. In addition, a series of internal standards are added to every well (except zero sample). These quality assurance provisions, together with a patented plate design, are major contributors to analytical precision and reproducibility.





The Absolute IDQ® p180 Kit: Fast Track to Metabolic Biomarker Signatures!

Products

Product Description	Product Number
Absolute /DQ ® p180 Kit - (96)	Contact us
Met <i>IDQ</i> ™ StatPack Module	9120052120189
Met /DQ ™ Ratio Explorer Module	9120052120899

The Absolute \mbox{IDQ}^{\otimes} p180 Kit is designed for research use only. Not for in-vitro diagnostic use

If you want to learn more, contact us!

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^{**} minimum time for quality control and export of data