

METABOLOMIX⁺ : A NON-INVASIVE PERSONALIZED ASSESSMENT

Metabolomix⁺ is a urinary nutritional evaluation that includes key organic acids and amino acids to evaluate the functional need for antioxidants, B-vitamins, minerals, digestive support, and amino acids.

Metabolomix⁺ also includes the option of additional Essential & Metabolic Fatty Acids, Toxic and Nutrient Elements, plus Genomics.

The Metabolomix⁺ provides targeted nutrient therapeutics designed to give insight into:

- Mood disorders^{1,2}
- Cardiovascular disease^{3,4}
- Metabolic syndrome^{5,6}
- Fatigue^{7,8}
- Obesity and weight issues^{9,10}
- Cognitive decline^{11,12}
- Athletic optimization^{13,14}
- Malnutrition¹⁵

Metabolomix⁺ consists of:

- **Organic Acids** – providing insight into nutritional cofactor needs, digestive issues, cellular energy production, neurotransmitter metabolism, detoxification, **and now oxalates.**
- **Amino Acid Analysis** – features 37 urine amino acids. This assesses nutritionally essential and non-essential amino acids, as well as intermediary metabolites that augment an understanding of B vitamin need, and need for support of protein digestion and absorption.
- **Oxidative Stress Analysis** – highlights the body's current state of oxidative stress and reserve capacity.

Why Use Metabolomix⁺?

Studies suggest that even a balanced diet may not provide all essential nutrients. Metabolomix + can help to support you and your patients by:

- Identifying nutritional insufficiencies that may be at the root cause of complex chronic conditions
- Providing at-home specimen collection
- Offering an easy-to-use "Interpretation At-A-Glance" that provides patients with valuable information about the function of nutrients, their dietary sources, and the causes and complications of their deficiencies
- Giving insight into potential digestion and absorption abnormalities

The Metabolomix⁺ report offers functional pillars with a built-in scoring system to guide therapy in a systems-based fashion. The report also contains dynamic biochemical pathway charts for clearer understanding

• Add-on Components

- Bloodspot Essential Metabolic Fatty Acids
- Urine Nutrient & Toxic Elements
- Genomic SNPs



FATIGUE



MOOD DISORDERS



WEIGHT ISSUES

● Results Overview



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© Genova Diagnostics

Metabolomix+

3200 Metabolomix+ - FMV Urine

Results Overview









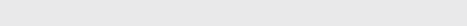






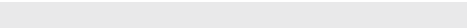







Functional Imbalance Scores

Key **0-4** : Minimal Need for Support **5-7** : Moderate Need for Support **8-10** : High Need for Support

| Need for Antioxidant Support | Need for Mitochondrial Support | Need for Inflammation Support | Need for Reduced Exposure | Need for Methylation Support |
|---|---|---|---|--|
| Oxidative Stress <div>5</div> <ul style="list-style-type: none"> Cystine ▼ Cysteine ● Lipid Peroxides ● 8-OHdG ● Taurine ▼ Citric Acid ▲ cis-Aconitic Acid ● | Mitochondrial Dysfunction <div>0</div> <ul style="list-style-type: none"> Magnesium ▼ FIGLU ● Methylmalonic Acid ● Glutaric Acid ▲ Lactic Acid ● Pyruvic Acid ▲ Citric Acid ▲ cis-Aconitic Acid ● Isocitric Acid ● α-Ketoglutaric Acid ● Succinic Acid ▼ Malic Acid ▲ Adipic Acid ● Suberic Acid ● Manganese ▲ | Omega Imbalance <div>8</div> <ul style="list-style-type: none"> Omega-3 Index ▼ Omega 6/3 Ratio ● α-Linolenic Acid ▼ Arachidonic Acid ▲ Linoleic Acid ▼ γ-Linolenic Acid ▼ Dihomo-γ-linolenic Acid ▼ <p>OMEGA IMBALANCE AVAILABLE WITH OPTIONAL ADD-ONS</p> | Toxic Exposure <div>7</div> <ul style="list-style-type: none"> Lead ▲ Mercury ● α-Hydroxyisobutyric Acid ● α-Ketophenylacetic Acid ● Arsenic ● Cadmium ▲ Pyroglutamic Acid ● Orotic Acid ● Citric Acid ▲ cis-Aconitic Acid ● Isocitric Acid ● Glutaric Acid ▲ | Methylation Imbalance <div>7</div> <ul style="list-style-type: none"> Methylmalonic Acid ● Methionine ▲ FIGLU ● Sarcosine ▲ Vanilmandelic Acid ● Arginine ▼ Glycine ● Serine ▲ Creatinine ▼ |

● Suggested Supplement Schedule

Nutrient Need Overview

| | Nutrient Need | | | | | | | | | | | DRI | Suggested Recommendations | Provider Recommendations |
|-------------------------------|---|---|---|---|---|---|---|---|---|---|----|----------|---------------------------|--------------------------|
| | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 10 | | | |
| Antioxidants | | | | | | | | | | | | | | |
| Vitamin A |  | | | | | | | | | | | 3,000 IU | <div>3,000 IU</div> | |
| Vitamin C |  | | | | | | | | | | | 90 mg | <div>250 mg</div> | |
| Vitamin E / Tocopherols |  | | | | | | | | | | | 22 IU | <div>100 IU</div> | |
| α-Lipoic Acid |  | | | | | | | | | | | | <div>100 mg</div> | |
| CoQ10 |  | | | | | | | | | | | | <div>30 mg</div> | |
| Glutathione |  | | | | | | | | | | | | | |
| Plant-based Antioxidants |  | | | | | | | | | | | | | |
| B-Vitamins | | | | | | | | | | | | | | |
| Thiamin - B1 |  | | | | | | | | | | | 1.2 mg | <div>50 mg</div> | |
| Riboflavin - B2 |  | | | | | | | | | | | 1.3 mg | <div>25 mg</div> | |
| Niacin - B3 |  | | | | | | | | | | | 16 mg | <div>30 mg</div> | |
| Pyridoxine - B6 |  | | | | | | | | | | | 1.7 mg | <div>25 mg</div> | |
| Biotin - B7 |  | | | | | | | | | | | 30 mcg | <div>100 mcg</div> | |
| Folic Acid - B9 |  | | | | | | | | | | | 400 mcg | <div>800 mcg</div> | |
| Cobalamin - B12 |  | | | | | | | | | | | 2.4 mcg | <div>100 mcg</div> | |
| Minerals | | | | | | | | | | | | | | |
| Magnesium |  | | | | | | | | | | | 420 mg | <div>800 mg</div> | |
| Manganese |  | | | | | | | | | | | 2.3 mg | <div>3.0 mg</div> | |
| Molybdenum |  | | | | | | | | | | | 45 mcg | <div>75 mcg</div> | |
| Zinc |  | | | | | | | | | | | 11 mg | <div>20 mg</div> | |
| Essential Fatty Acids | | | | | | | | | | | | | | |
| Omega-3 Fatty Acids |  | | | | | | | | | | | 500 mg | <div>2,000 mg</div> | |
| GI Support | | | | | | | | | | | | | | |
| Digestive Support/Enzymes |  | | | | | | | | | | | | <div>10,000 IU</div> | |
| Microbiome Support/Probiotics |  | | | | | | | | | | | | <div>50 billion CFU</div> | |

Amino Acids (mg/day)

| | | | |
|------------|-------|---------------|-------|
| Arginine | 1,460 | Methionine | 0 |
| Asparagine | 0 | Phenylalanine | 0 |
| Cysteine | 0 | Serine | 0 |
| Glutamine | 0 | Taurine | 1,212 |
| Glycine | 0 | Threonine | 0 |
| Histidine | 2,190 | Tryptophan | 0 |
| Isoleucine | 1,168 | Tyrosine | 0 |
| Leucine | 1,308 | Valine | 10 |
| Lysine | 1,730 | | |

Recommendations for age and gender-specific supplementation are set by comparing levels of nutrient functional need to optimal levels as described in the peer-reviewed literature. They are provided as guidance for short-term support of nutritional deficiencies only.

The Nutrient Need Overview is provided at the request of the ordering practitioner. Any application of it as a therapeutic intervention is to be determined by the ordering practitioner.

Interpretation At-A-Glance

Antioxidant Needs

Vitamin A



4

- Beta-carotene & other carotenoids are converted to vitamin A (retinol), involved in vision, antioxidant & immune function, gene expression & cell growth.
- Vitamin A deficiency may occur with chronic alcoholism, zinc deficiency, hypothyroidism, or oral contraceptives containing estrogen & progestin.
- Deficiency may result in night blindness, impaired immunity, healing & tissue regeneration, increased risk of infection, leukoplakia or keratosis.
- Food sources include cod liver oil, fortified cereals & milk, eggs, sweet potato, pumpkin, carrot, cantaloupe, mango, spinach, broccoli, kale & butternut squash.

Vitamin C



0

- Vitamin C is an antioxidant (also used in the regeneration of other antioxidants). It is involved in cholesterol metabolism, the production & function of WBCs and antibodies, and the synthesis of collagen, norepinephrine and carnitine.
- Deficiency may occur with oral contraceptives, aspirin, diuretics or NSAIDs.
- Deficiency can result in scurvy, swollen gingiva, periodontal destruction, loose teeth, sore mouth, soft tissue ulcerations, or increased risk of infection.
- Food sources include oranges, grapefruit, strawberries, tomato, sweet red pepper, broccoli and potato.

Vitamin E / Tocopherols



4

- Alpha-tocopherol (body's main form of vitamin E) functions as an antioxidant, regulates cell signaling, influences immune function and inhibits coagulation.
- Deficiency may occur with malabsorption, cholestyramine, colestipol, isoniazid, orlistat, olestra and certain anti-convulsants (e.g., phenobarbital, phenytoin).
- Deficiency may result in peripheral neuropathy, ataxia, muscle weakness, retinopathy, and increased risk of CVD, prostate cancer and cataracts.
- Food sources include oils (olive, soy, corn, canola, safflower, sunflower), eggs, nuts, seeds, spinach, carrots, avocado, dark leafy greens and wheat germ.

α-Lipoic Acid



6

- α-Lipoic acid plays an important role in energy production, antioxidant activity (including the regeneration of vitamin C and glutathione), insulin signaling, cell signaling and the catabolism of α-keto acids and amino acids.
- High biotin intake can compete with lipoic acid for cell membrane entry.
- Optimal levels of α-lipoic acid may improve glucose utilization and protect against diabetic neuropathy, vascular disease and age-related cognitive decline.
- Main food sources include organ meats, spinach and broccoli. Lesser sources include tomato, peas, Brussels sprouts and brewer's yeast.

CoQ10



0

- CoQ10 is a powerful antioxidant that is synthesized in the body and contained in cell membranes. CoQ10 is also essential for energy production & pH regulation.
- CoQ10 deficiency may occur with HMG-CoA reductase inhibitors (statins), several anti-diabetic medication classes (biguanides, sulfonylureas) or beta-blockers.
- Low levels may aggravate oxidative stress, diabetes, cancer, congestive heart failure, cardiac arrhythmias, gingivitis and neurologic diseases.
- Main food sources include meat, poultry, fish, soybean, canola oil, nuts and whole grains. Moderate sources include fruits, vegetables, eggs and dairy.

Glutathione



2

- Glutathione (GSH) is composed of cysteine, glutamine & glycine. GSH is a source of sulfate and plays a key role in antioxidant activity and detoxification of toxins.
- GSH requirement is increased with high-fat diets, cigarette smoke, cystinuria, chronic alcoholism, chronic acetaminophen use, infection, inflammation and toxic exposure.
- Deficiency may result in oxidative stress & damage, impaired detoxification, altered immunity, macular degeneration and increased risk of chronic illness.
- Food sources of GSH precursors include meats, poultry, fish, soy, corn, nuts, seeds, wheat germ, milk and cheese.

Plant-based Antioxidants



5

- Oxidative stress is the imbalance between the production of free radicals and the body's ability to readily detoxify these reactive species and/or repair the resulting damage with anti-oxidants.
- Oxidative stress can be endogenous (energy production and inflammation) or exogenous (exercise, exposure to environmental toxins).
- Oxidative stress has been implicated clinically in the development of neurodegenerative diseases, cardiovascular diseases and chronic fatigue syndrome.
- Antioxidants may be found in whole food sources (e.g., brightly colored fruits & vegetables, green tea, turmeric) as well as nutraceuticals (e.g., resveratrol, EGCG, lutein, lycopene, ginkgo, milk thistle, etc.).

KEY

- Function of Nutrient
- Cause of Deficiency
- Complications of Deficiency
- Food Sources of Nutrient

Interpretation At-A-Glance

B-Vitamin Needs

Thiamin - B1



- B1 is a required cofactor for enzymes involved in energy production from food, and for the synthesis of ATP, GTP, DNA, RNA and NADPH.
- Low B1 can result from chronic alcoholism, diuretics, digoxin, oral contraceptives and HRT, or large amounts of tea & coffee (contain anti-B1 factors).
- B1 deficiency may lead to dry beriberi (e.g., neuropathy, muscle weakness), wet beriberi (e.g., cardiac problems, edema), encephalopathy or dementia.
- Food sources include lentils, whole grains, wheat germ, Brazil nuts, peas, organ meats, brewer's yeast, blackstrap molasses, spinach, milk & eggs.

Riboflavin - B2



- B2 is a key component of enzymes involved in antioxidant function, energy production, detoxification, methionine metabolism and vitamin activation.
- Low B2 may result from chronic alcoholism, some anti-psychotic medications, oral contraceptives, tricyclic antidepressants, quinacrine or adriamycin.
- B2 deficiency may result in oxidative stress, mitochondrial dysfunction, low uric acid, low B3 or B6, high homocysteine, anemia or oral & throat inflammation.
- Food sources include milk, cheese, eggs, whole grains, beef, chicken, wheat germ, fish, broccoli, asparagus, spinach, mushrooms and almonds.

Niacin - B3



- B3 is used to form NAD and NADP, involved in energy production from food, fatty acid & cholesterol synthesis, cell signaling, DNA repair & cell differentiation.
- Low B3 may result from deficiencies of tryptophan (B3 precursor), B6, B2 or Fe (cofactors in B3 production), or from long-term isoniazid or oral contraceptive use.
- B3 deficiency may result in pellagra (dermatitis, diarrhea, dementia), neurologic symptoms (e.g., depression, memory loss), bright red tongue or fatigue.
- Food sources include poultry, beef, organ meats, fish, whole grains, peanuts, seeds, lentils, brewer's yeast and lima beans.

Pyridoxine - B6



- B6 (as P5P) is a cofactor for enzymes involved in glycogenolysis & gluconeogenesis, and synthesis of neurotransmitters, heme, B3, RBCs and nucleic acids.
- Low B6 may result from chronic alcoholism, long-term diuretics, estrogens (oral contraceptives and HRT), anti-TB meds, penicillamine, L-DOPA or digoxin.
- B6 deficiency may result in neurologic symptoms (e.g., irritability, depression, seizures), oral inflammation, impaired immunity or increased homocysteine.
- Food sources include poultry, beef, beef liver, fish, whole grains, wheat germ, soybean, lentils, nuts & seeds, potato, spinach and carrots.

Biotin - B7



- Biotin is a cofactor for enzymes involved in functions such as fatty acid synthesis, mitochondrial FA oxidation, gluconeogenesis and DNA replication & transcription.
- Deficiency may result from certain inborn errors, chronic intake of raw egg whites, long-term TPN, anticonvulsants, high-dose B5, sulfa drugs & other antibiotics.
- Low levels may result in neurologic symptoms (e.g., paresthesias, depression), hair loss, scaly rash on face or genitals or impaired immunity.
- Food sources include yeast, whole grains, wheat germ, eggs, cheese, liver, meats, fish, wheat, nuts & seeds, avocado, raspberries, sweet potato and cauliflower.

Folic Acid - B9



- Folic acid plays a key role in coenzymes involved in DNA and SAME synthesis, methylation, nucleic acids & amino acid metabolism and RBC production.
- Low folate may result from alcoholism, high-dose NSAIDs, diabetic meds, H2 blockers, some diuretics and anti-convulsants, SSRIs, methotrexate, trimethoprim, pyrimethamine, triamterene, sulfasalazine or cholestyramine.
- Folate deficiency can result in anemia, fatigue, low methionine, increased homocysteine, impaired immunity, heart disease, birth defects and CA risk.
- Food sources include fortified grains, green vegetables, beans & legumes.

Cobalamin - B12



- B12 plays important roles in energy production from fats & proteins, methylation, synthesis of hemoglobin & RBCs, and maintenance of nerve cells, DNA & RNA.
- Low B12 may result from alcoholism, malabsorption, hypochlorhydria (e.g., from atrophic gastritis, H. pylori infection, pernicious anemia, H2 blockers, PPIs), vegan diets, diabetic meds, cholestyramine, chloramphenicol, neomycin or colchicine.
- B12 deficiency can lead to anemia, fatigue, neurologic symptoms (e.g., paresthesias, memory loss, depression, dementia), methylation defects or chromosome breaks.
- Food sources include shellfish, red meat, poultry, fish, eggs, milk and cheese.

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Interpretation At-A-Glance

Mineral Needs

Magnesium



- Magnesium is involved in >300 metabolic reactions. Key areas include energy production, bone & ATP formation, muscle & nerve conduction and cell signaling.
- Deficiency may occur with malabsorption, alcoholism, hyperparathyroidism, renal disorders (wasting), diabetes, diuretics, digoxin or high doses of zinc.
- Low Mg may result in muscle weakness/spasm, constipation, depression, hypertension, arrhythmias, hypocalcemia, hypokalemia or personality changes.
- Food sources include dark leafy greens, oatmeal, buckwheat, unpolished grains, chocolate, milk, nuts & seeds, lima beans and molasses.

Manganese



- Manganese plays an important role in antioxidant function, gluconeogenesis, the urea cycle, cartilage & bone formation, energy production and digestion.
- Impaired absorption of Mn may occur with excess intake of Fe, Ca, Cu, folic acid, or phosphorous compounds, or use of long-term TPN, Mg-containing antacids or laxatives.
- Deficiency may result in impaired bone/connective tissue growth, glucose & lipid dysregulation, infertility, oxidative stress, inflammation or hyperammonemia.
- Food sources include whole grains, legumes, dried fruits, nuts, dark green leafy vegetables, liver, kidney and tea.

Molybdenum



- Molybdenum is a cofactor for enzymes that convert sulfites to sulfate, and nucleotides to uric acid, and that help metabolize aldehydes & other toxins.
- Low Mo levels may result from long-term TPN that does not include Mo.
- Mo deficiency may result in increased sulfite, decreased plasma uric acid (and antioxidant function), deficient sulfate, impaired sulfation (detoxification), neurologic disorders or brain damage (if severe deficiency).
- Food sources include buckwheat, beans, grains, nuts, beans, lentils, meats and vegetables (although Mo content of plants depends on soil content).

Zinc



- Zinc plays a vital role in immunity, protein metabolism, heme synthesis, growth & development, reproduction, digestion and antioxidant function.
- Low levels may occur with malabsorption, alcoholism, chronic diarrhea, diabetes, excess Cu or Fe, diuretics, ACE inhibitors, H2 blockers or digoxin.
- Deficiency can result in hair loss and skin rashes, also impairments in growth & healing, immunity, sexual function, taste & smell and digestion.
- Food sources include oysters, organ meats, soybean, wheat germ, seeds, nuts, red meat, chicken, herring, milk, yeast, leafy and root vegetables.

Essential Fatty Acid Needs

Need for Essential Fatty Acids



- Omega-3 (O3) and Omega-6 (O6) fatty acids are polyunsaturated fatty acids that cannot be synthesized by the human body. They are classified as essential nutrients and must be obtained from dietary sources.
- The standard American diet is much higher in O6 than O3 fatty acids. Deficiency of EFAs may result from poor dietary intake and/or poor conversion from food sources.
- EFA deficiency is associated with decreased growth & development of infants and children, dry skin/rash, poor wound healing, and increased risk of infection, cardiovascular and inflammatory diseases.
- Dietary sources of the O6 Linoleic Acid (LA) include vegetable oils, nuts, seeds and some vegetables. Dietary sources of the O3 α-Linolenic Acid (ALA) include flaxseeds, walnuts, and their oils. Fish (mackerel, salmon, sardines) are the major dietary sources of the O3 fatty acids EPA and DHA.

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Interpretation At-A-Glance

Microbiome & Digestive Support

Need for Probiotics



- Probiotics have many functions. These include: production of some B vitamins and vitamin K; enhance digestion & absorption; decrease severity of diarrheal illness; modulate of immune function & intestinal permeability.
- Alterations of gastrointestinal microflora may result from C-section delivery, antibiotic use, improved sanitation, decreased consumption of fermented foods and use of certain drugs.
- Some of the diseases associated with microflora imbalances include: IBS, IBD, fibromyalgia, chronic fatigue syndrome, obesity, atopic illness, colic and cancer.
- Food sources rich in probiotics are yogurt, kefir and fermented foods.

Need for Pancreatic Enzymes



- Pancreatic enzymes are secreted by the exocrine glands of the pancreas and include protease/peptidase, lipase and amylase.
- Pancreatic exocrine insufficiency may be primary or secondary in nature. Any indication of insufficiency warrants further evaluation for underlying cause (i.e., celiac disease, small intestine villous atrophy, small bowel bacterial overgrowth).
- A high functional need for digestive enzymes suggests that there is an impairment related to digestive capacity.
- Determining the strength of the pancreatic enzyme support depends on the degree of functional impairment. Supplement potency is based on the lipase units present in both prescriptive and non-prescriptive agents.

Functional Imbalances

Mitochondrial Dysfunction



- Mitochondria are a primary site of generation of reactive oxygen species. Oxidative damage is considered an important factor in decline of physiologic function that occurs with aging and stress.
- Mitochondrial defects have been identified in cardiovascular disease, fatigue syndromes, neurologic disorders such as Parkinson's and Alzheimer's disease, as well as a variety of genetic conditions. Common nutritional deficiencies can impair mitochondrial efficiency.

Need for Methylation



- Methylation is an enzymatic process that is critical for both synthesis and inactivation. DNA, estrogen and neurotransmitter metabolism are all dependent on appropriate methylation activity.
- B vitamins and other nutrients (methionine, magnesium, selenium) functionally support catechol-O-methyltransferase (COMT), the enzyme responsible for methylation.

Toxic Exposure



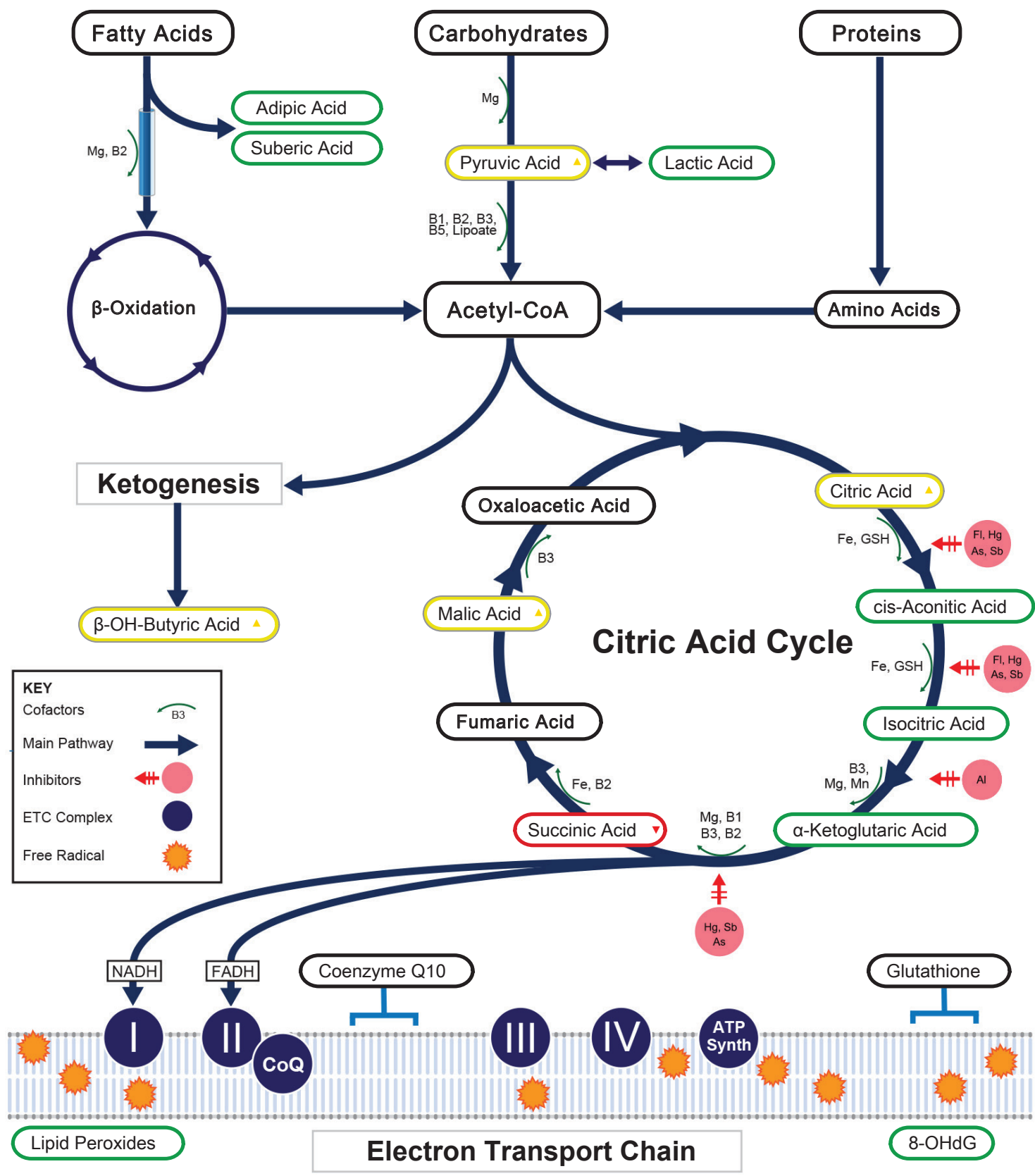
- Methyl tert-Butyl Ether (MTBE) is a common gasoline additive used to increase octane ratings, and has been found to contaminate ground water supplies where gasoline is stored. Inhalation of MTBE may cause nose and throat irritation, as well as headaches, nausea, dizziness and mental confusion. Animal studies suggest that drinking MTBE may cause gastrointestinal irritation, liver and kidney damage and nervous system effects.
- Styrene is classified by the US EPA as a "potential human carcinogen," and is found widely distributed in commercial products such as rubber, plastic, insulation, fiberglass, pipes, food containers and carpet backing.
- Levels of these toxic substances should be examined within the context of the body's functional capacity for methylation and need for glutathione.

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Oxidative Stress & Mitochondrial Dysfunction



Organic Acids

| Organic Acids | | | |
|--|-----------------|--|-----------------|
| Malabsorption & Dysbiosis Markers | | Vitamin Markers | |
| Malabsorption Markers | | Branched-Chain Catabolites (B1, B2, B3, ALA) | |
| | Reference Range | | Reference Range |
| Indoleacetic Acid | 3.1 | α-Ketoadipic Acid | 0.7 |
| Phenylacetic Acid | 0.16 | α-Ketoisovaleric Acid | 0.95 |
| Dysbiosis Markers | | α-Ketoisocaproic Acid | 0.64 |
| Dihydroxyphenylpropionic Acid (DHPPA) | 5.8 | α-Keto-β-Methylvaleric Acid | 1.6 |
| 3-Hydroxyphenylacetic Acid | <dl | Glutaric Acid | 0.47 |
| 4-Hydroxyphenylacetic Acid | <dl | Isovalerylglycine | <dl |
| Benzoic Acid | 0.07 | Methylation Markers (Folate, B12) | |
| Hippuric Acid | 361 | Formiminoglutamic Acid (FIGlu) | <dl |
| Yeast / Fungal Dysbiosis Markers | | Methylmalonic Acid | 1.3 |
| D-Arabinitol | 12 | Biotin Markers | |
| Citramalic Acid | 5.1 | 3-Hydroxypropionic Acid | 7 |
| Tartaric Acid | <dl | 3-Hydroxyisovaleric Acid | <dl |
| Cellular Energy & Mitochondrial Markers | | Neurotransmitter Metabolites | |
| Fatty Acid Metabolism | | Kynurenine Markers (Vitamin B6) | |
| | Reference Range | | Reference Range |
| Adipic Acid | <dl | Kynurenic Acid | <dl |
| Suberic Acid | <dl | Quinolinic Acid | 3.3 |
| Carbohydrate Metabolism | | Kynurenic / Quinolinic Ratio | NR |
| Pyruvic Acid | 27 | Xanthurenic Acid | <dl |
| Lactic Acid | 6.8 | Catecholamine Markers | |
| α-Hydroxybutyric Acid | 44.65 | Homovanillic Acid | 2.0 |
| β-OH-Butyric Acid | 2.1 | Vanilmandelic Acid | 1.6 |
| β-OH-β-Methylglutaric Acid | <dl | 3-Methyl-4-OH-phenylglycol | 0.10 |
| Energy Metabolism | | Serotonin Markers | |
| Citric Acid | 404 | 5-OH-indoleacetic Acid | 11.2 |
| cis-Aconitic Acid | 19 | Toxin & Detoxification Markers | |
| Isocitric Acid | 50 | | Reference Range |
| α-Ketoglutaric Acid | 15 | Pyroglutamic Acid | 21 |
| Succinic Acid | <dl | α-Ketophenylacetic Acid (from Styrene) | 0.17 |
| Malic Acid | 2.9 | α-Hydroxyisobutyric Acid (from MTBE) | 3.9 |
| | | Orotic Acid | 0.54 |

Methodology: GC/MS. LC/MS/MS. Alkaline Picrate. Colorimetric

Organic Acid Reference Ranges are Age Specific



Organic Acids

Oxalate Markers

Reference
Range



Creatinine Concentration

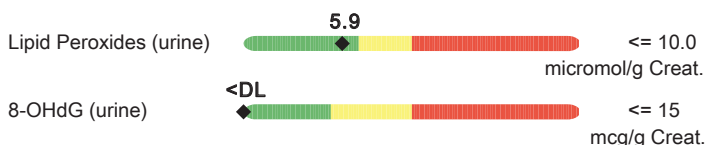
Reference
Range



Oxidative Stress Markers

Oxidative Damage

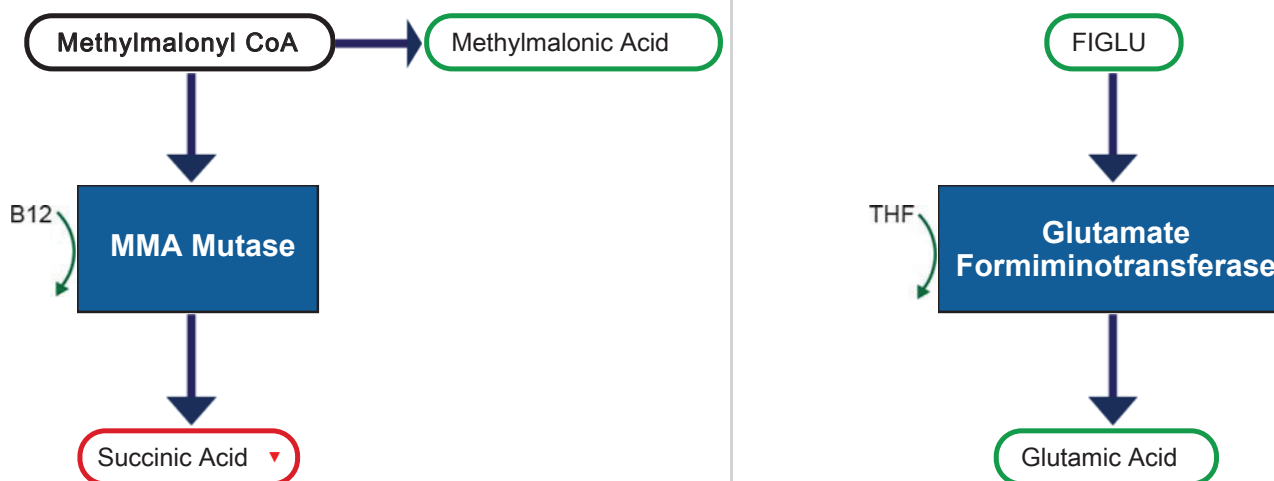
Reference
Range



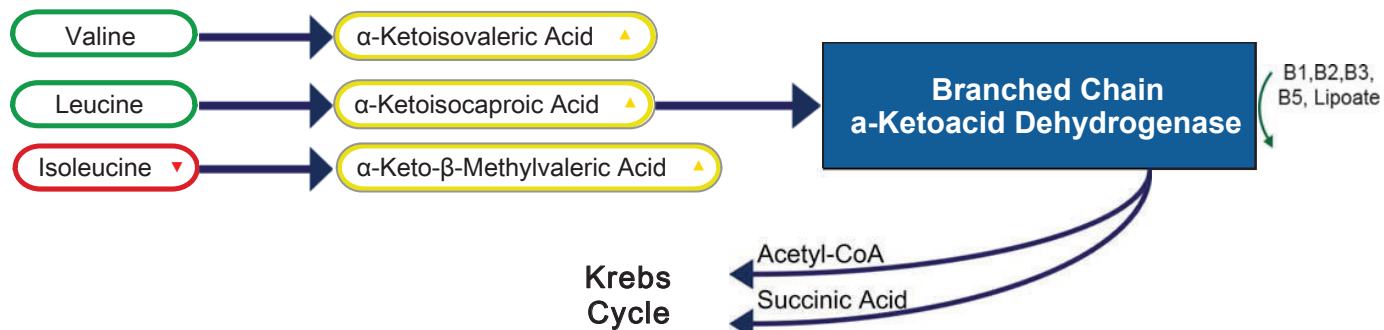
The Oxidative Stress reference ranges are based on an adult population.

Pathways

Methylation Markers



Branch-Chain Amino Acid Metabolism



Amino Acids (FMV)

| Nutritionally Essential Amino Acids | | | Intermediary Metabolites | | |
|-------------------------------------|-----|-----------------|----------------------------|-----|-------------------|
| Amino Acid | | Reference Range | B-Vitamin Markers | | Reference Range |
| Arginine | <dl | 3-43 | α-Aminoadipic Acid | 19 | 6-56 |
| Histidine | <dl | 102-763 | α-Amino-N-butyric Acid | 12 | 2-21 |
| Isoleucine | <dl | 3-25 | β-Aminoisobutyric Acid | 25 | 4-194 |
| Leucine | 7 | 6-61 | Cystathionine | <dl | 4-48 |
| Lysine | 15 | 15-231 | Urea Cycle Markers | | |
| Methionine | 11 | 2-16 | Citrulline | 6.0 | 0.7-3.4 |
| Phenylalanine | 25 | 7-92 | Ornithine | 10 | 3-17 |
| Taurine | 21 | 39-568 | Urea ♦ | 149 | 150-380 |
| Threonine | 51 | 9-97 | | | mmol/g creatinine |
| Tryptophan | 24 | 8-58 | Glycine/Serine Metabolites | | |
| Valine | 11 | 5-43 | Glycine | 161 | 47-435 |

Nonessential Protein Amino Acids

| Amino Acid | | Reference Range | | | Reference Range |
|---------------------|-----|-----------------|---------------------------------|-----|-----------------|
| Alanine | 87 | 26-275 | Serine | 149 | 24-140 |
| Asparagine | 75 | 12-115 | Ethanolamine | 349 | 40-226 |
| Aspartic Acid | <dl | <= 9 | Phosphoethanolamine | 5 | 1-9 |
| Cysteine | 27 | 9-60 | Phosphoserine | <dl | 2-13 |
| Cystine | <dl | 10-116 | Sarcosine | 1.9 | <= 1.0 |
| γ-Aminobutyric Acid | 2 | <= 3 | Dietary Peptide Related Markers | | |
| Glutamic Acid | 12 | 2-16 | Anserine (dipeptide) | 0.6 | 0.7-76.1 |
| Glutamine | 200 | 85-518 | Carnosine (dipeptide) | 5 | 1-32 |
| Proline | 4 | 1-9 | 1-Methylhistidine | <dl | 18-887 |
| Tyrosine | 40 | 19-135 | 3-Methylhistidine | <dl | 47-232 |
| | | | β-Alanine | <dl | <= 18 |

Creatinine Concentration

| | | Reference Range |
|--------------|-----|-----------------|
| Creatinine ♦ | 1.9 | 3.1-19.5 mmol/L |

Amino Acid reference ranges are age specific.

Methodology: LC/MS/MS, Alkaline Picrate

OPTIONAL ADD-ON

3202 Add-on Bloodspot Essential & Metabolic Fatty Acids – Bloodspot

Methodology: GCMS

Essential & Metabolic Fatty Acids Markers (RBCs)

| Omega-3 Fatty Acids | | | | Omega-6 Fatty Acids | | | |
|------------------------------------|------|-----------------|----------------|--|------|-----------------|----------------|
| Analyte | | Reference Range | | Analyte | | Reference Range | |
| (cold water fish, flax, walnut) | | | | (vegetable oil, grains, most meats, dairy) | | | |
| α-Linolenic (ALA) 18:3 n3 | 0.11 | | >= 0.28 wt % | Linoleic (LA) 18:2 n6 | 15.0 | | 18.8-28.3 wt % |
| Eicosapentaenoic (EPA) 20:5 n3 | 0.14 | | >= 0.12 wt % | γ-Linolenic (GLA) 18:3 n6 | 0.14 | | 0.15-0.54 wt % |
| Docosapentaenoic (DPA) 22:5 n3 | 1.09 | | >= 0.34 wt % | Dihomo-γ-linolenic (DGLA) 20:3 n6 | 1.17 | | >= 1.02 wt % |
| Docosahexaenoic (DHA) 22:6 n3 | 1.7 | | >= 0.8 wt % | Arachidonic (AA) 20:4 n6 | 17 | | 7-12 wt % |
| % Omega-3s | 3.0 | | >= 1.6 | Docosatetraenoic (DTA) 22:4 n6 | 2.74 | | 0.45-1.25 wt % |
| Omega-9 Fatty Acids | | | | Eicosadienoic 20:2 n6 | 0.39 | | <= 0.26 wt % |
| Analyte | | Reference Range | | % Omega-6s | 36.4 | | 30.5-39.7 |
| (olive oil) | | | | Monounsaturated Fatty Acids | | | |
| Oleic 18:1 n9 | 13 | | 14-21 wt % | Omega-7 Fatty Acids | | | |
| Nervonic 24:1 n9 | 3.0 | | 1.1-1.8 wt % | Palmitoleic 16:1 n7 | 0.29 | | <= 2.58 wt % |
| % Omega-9s | 16.6 | | 17.3-22.5 | Vaccenic 18:1 n7 | 1.23 | | <= 1.65 wt % |
| Saturated Fatty Acids | | | | Trans Fats | | | |
| Analyte | | Reference Range | | | | | |
| (meat, dairy, coconuts, palm oils) | | | | Elaidic 18:1 n9t | 0.15 | | <= 0.59 wt % |
| Palmitic C16:0 | 21 | | 19-27 wt % | Delta-6-Desaturase Activity | | | |
| Stearic C18:0 | 17 | | 9-12 wt % | Upregulated Functional Impaired | | | |
| Arachidic C20:0 | 0.23 | | 0.24-0.40 wt % | Linoleic / DGLA 18:2 n6 / 20:3 n6 | 12.8 | | 12.6-31.5 |
| Behenic C22:0 | 0.95 | | 0.88-1.61 wt % | Cardiovascular Risk | | | |
| Tricosanoic C23:0 | 0.14 | | 0.19-0.26 wt % | | | | |
| Lignoceric C24:0 | 2.8 | | 1.1-1.9 wt % | Analyte | | Reference Range | |
| Pentadecanoic C15:0 | 0.04 | | 0.14-0.30 wt % | Omega-6s / Omega-3s | 11.9 | | 8.5-27.4 |
| Margaric C17:0 | 0.23 | | 0.24-0.45 wt % | AA / EPA 20:4 n6 / 20:5 n3 | 118 | | 10-86 |
| % Saturated Fats | 42.3 | | 39.8-43.6 | Omega-3 Index | 4.6 | | >= 4.0 |

The Essential Fatty Acid reference ranges are based on an adult population.

OPTIONAL ADD-ON

Page 12

Fatty Acid Metabolism

Omega-3 Metabolism

Enzyme

 α -Linolenic Acid ▼

Stearidonic Acid

Eicosatetraenoic Acid

Eicosapentaenoic Acid

Anti-Inflammatory
Eicosanoids

Docosapentaenoic Acid

Docosahexaenoic Acid

Delta-6-Desaturase

Important Regulators:
B2, B3, B6, Vitamin C,
Insulin, Zn, Mg

Elongase

Important Regulators:
B3, B5, B6, Biotin,
Vitamin C

Delta-5-Desaturase

Important Regulators:
B2, B3, B6, Vitamin C,
Insulin, Zn, Mg

Elongase

Important Regulators:
B3, B5, B6, Biotin,
Vitamin C

Elongase

Delta-6-Desaturase

Omega-6 Metabolism

Linoleic Acid ▼

 γ -Linolenic Acid ▼Dihomo- γ -Linolenic Acid ▼Anti-Inflammatory
Series 1 Prostaglandins

Arachidonic Acid ▲

Pro-Inflammatory
Eicosanoids

Docosatetraenoic Acid ▲

OPTIONAL ADD-ON

3204 Add - on Comprehensive Urine Elements - FMV Urine

Methodology: ICP-MS and Alkaline Picrate

| Elemental Markers | | | | | |
|----------------------------|-------|-----------------|----------------------------|-----------------|-----------|
| Toxic Elements | | | Nutrient Elements | | |
| Element | | Reference Range | Element | Reference Range | |
| Results in ug/g creatinine | | | Results in ug/g creatinine | | |
| Lead | 5.6 | <= 1.4 | Chromium | 0.6 | 0.6-9.4 |
| Mercury | 0.28 | <= 2.19 | Cobalt | 1.50 | 0.01-2.60 |
| Aluminum | 5.0 | <= 22.3 | Copper | 121.0 | 4.0-11.4 |
| Antimony | 0.130 | <= 0.149 | Iron | 5 | 5-64 |
| Arsenic | 1 | <= 50 | Lithium | 14 | 9-129 |
| Barium | 3.4 | <= 6.7 | Manganese | 11.20 | 0.03-1.16 |
| Bismuth | 2.00 | <= 2.28 | Molybdenum | 15 | 15-175 |
| Cadmium | 0.71 | <= 0.64 | Selenium | 274 | 32-333 |
| Cesium | 5.0 | <= 10.5 | Strontium | 275 | 47-346 |
| Gadolinium | 0.015 | <= 0.019 | Vanadium | 2.0 | 0.1-3.2 |
| Gallium | 0.020 | <= 0.028 | Zinc | 84 | 63-688 |
| Nickel | 1.20 | <= 3.88 | Results in mg/g creatinine | | |
| Niobium | 0.050 | <= 0.084 | Calcium | 250 | 37-313 |
| Platinum | 0.025 | <= 0.033 | Magnesium | 37 | 41-267 |
| Rubidium | 14 | <= 2,263 | Potassium | 2,656 | 759-4,653 |
| Thallium | 0.220 | <= 0.298 | Sulfur | 1,000 | 367-1,328 |
| Thorium | 3.500 | <= 4.189 | Creatinine Concentration | | |
| Tin | 5.22 | <= 2.04 | Reference Range | | |
| Tungsten | 0.150 | <= 0.211 | Urine Creatinine ♦ | | |
| Uranium | 0.010 | <= 0.026 | mmol/L | | |

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| Metabolomix+ PROFILE | | | | | | | | | |
|---|---|-------------------------------------|---|----------------------------|---|-------------------------------|--|--------------------|---|
| Analytes reported | | Analytes reported | | Analytes reported | | Analytes reported | | Analytes reported | |
| Organic Acids | | Organic Acids | | Amino Acids | | Add-on Fatty Acids | | Add-on Elements | |
| Malabsorption and Dysbiosis | | Vitamin Markers | | Creatinine Concentration | | Saturated Fatty Acids | | Nutrient Elements* | |
| Malabsorption Markers | | a-Ketoadipic Acid | • | Creatinine | • | Behenic Acid | • | Chromium* | • |
| Indoleacetic Acid (AA) | • | a-Ketoisovaleric Acid | • | Intermediary Metabolites | | Tricosanoic Acid | • | Cobalt* | • |
| Phenylacetic Acid (PAA) | • | a-Ketoisocaproic Acid | • | B Vitamin Markers | | Lignoceric Acid | • | Copper* | • |
| Bacterial Dysbiosis Markers | | a-Keto-β-Methylvaleric Acid | • | α-Aminoadipic Acid | • | Pentadecanoic Acid | • | Iron* | • |
| Dihydroxyphenylpropionic Acid | • | Formiminoglutamic Acid | • | α-Amino-N-Butyric Acid | • | Margaric Acid | • | Lithium* | • |
| 3-Hydroxypropionic Acid | • | Glutaric Acid | • | Urea Cycle Markers | | % Saturated Fats | • | Manganese* | • |
| 4-Hydroxyphenylpyruvic Acid | • | Isovalerylglycine | • | Citrulline | • | Omega 6 Fatty Acids | | Molybdenum* | • |
| Benzoic Acid | • | Methylmalonic Acid | • | Ornithine | • | Linoleic Acid | • | Selenium* | • |
| Hippuric Acid | • | Xanthurenic Acid | • | Urea | • | γ-Linolenic Acid | • | Strontium* | • |
| Yeast/Fungal Dysbiosis Markers | | 3-Hydroxypropionic Acid | • | Glycine/Serine Metabolites | | Dihomo-γ-linolenic Acid | • | Vanadium* | • |
| D-Arabinitol | • | 3-Hydroxyisovaleric Acid | • | Glycine | • | Arachidonic | • | Zinc* | • |
| Citramalic Acid | • | Toxin and Detoxification Markers | | Serine | • | Docosatetraenoic Acid | • | Calcium* | • |
| Tartaric Acid | • | a-Ketophenylacetic Acid | • | Ethanolamine | • | Eicosadienoic Acid | • | Magnesium* | • |
| Cellular Energy and Mitochondrial Metabolites | | a-Hydroxyisobutyric Acid | • | Phenylalanine | • | % Omega 6s | • | Potassium* | • |
| Carbohydrate Metabolism | | Orotic Acid | • | Phosphoethanolamine | • | Monounsaturated Fats | | Sulfur | • |
| Lactic Acid | • | Pyroglutamic Acid | • | Phospherserine | • | Omega 7 Fats | | Toxic Elements | |
| Pyruvic Acid | • | Oxalates | | Sarcosine | • | Palmitoleic Acid | • | Lead | • |
| α-Hydroxybutyric Acid | • | Glyceric Acid | • | Dietary Peptide | | Vaccenic Acid | • | Mercury | • |
| β-Hydroxybutyric Acid | • | Glycolic Acid | • | Anserine | • | Trans Fat | | Aluminium | • |
| Energy Metabolism | | Oxalic Acid | • | Carnosine | • | Elaidic Acid | • | Antimony | • |
| Citric Acid | • | Amino Acids | | 1-Methylhistidine | • | Delta - 6 Desaturase Activity | | Arsenic | • |
| cis-Aconitic Acid | • | Nutritionally Essential Amino Acids | | 3-Methylhistidine | • | Linoleic/DGLA ratio | • | Barium | • |
| Isocitric Acid | • | Arginine | • | β-Alanine | • | Add-on Genomic Markers | | Bismuth | • |
| α-Ketoglutaric Acid (AKG) | • | Histidine | • | Oxidative Stress | | APO E (C112R + R158C) | • | Cadmium | • |
| Succinic Acid | • | Isoleucine | • | Lipid Peroxides (urine) | • | COMT (V158M) | • | Cesium | • |
| Malic Acid | • | Leucine | • | 8-OHDg (urine) | • | MTHFR Combined | • | Gadolinium | • |
| β-OH-β-Methylglutaric Acid (HMG) | • | Lysine | • | Add-on Fatty Acids | | TNFA | • | Nickel | • |
| Fatty Acid Metabolism | | Methionine | • | Omega 3 Fatty Acids | | | Niobium | • | |
| Adipic Acid | • | Phenylalanine | • | α-Linolenic Acid | • | | Platinum | • | |
| Suberic Acid | • | Taurine | • | Eicosapentaenoic Acid | • | | Rubidium | • | |
| Creatinine Concentration | | Threonine | • | Docosapentaenoic Acid | • | | Thallium | • | |
| Creatinine | • | Tryptophan | • | Docosahexaenoic Acid | • | | Thorium | • | |
| Neurotransmitter Metabolism Markers | | Valine | • | % Omega 3s | • | | Tin | • | |
| Vanilmandelic Acid | • | Nonessential Protein Amino Acids | | Omega 9 Fatty Acids | | | Tungsten | • | |
| Homovanillic Acid | • | Alanine | • | Oleic Acid | • | | Uranium | • | |
| 5-OH-indoleacetic Acid | • | Asparagine | • | Nervonic Acid | • | | *NUTRIENT ELEMENTS AVAILABLE ONLY IF ADD-ON COMPREHENSIVE URINE ELEMENT PROFILE #3203 IS ORDERED | | |
| 3-Methyl-4-OH-phenylglycol | • | Aspartic Acid | • | % Omega 9s | • | | | | |
| Kynurenic Acid | • | Cysteine | • | Saturated Fatty Acids | | | | | |
| Quinolinic Acid | • | Cystine | • | Palmitic Acid | • | | | | |
| Kynurenic / Quinolinic Ratio | • | Gamma-Aminobutyric Acid | • | Stearic Acid | • | | | | |
| | | Glutamic Acid | • | Arachidic Acid | • | | | | |
| | | Glutamine | • | | | | | | |
| | | Proline | • | | | | | | |
| | | Tyrosine | • | | | | | | |

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