THE OBESITY-METABOLISM LINK

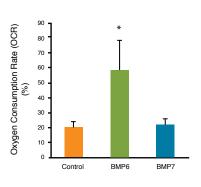
Cell Metabolism Assays for Obesity & Diabetes Research



GOLD STANDARD ASSAYS FOR MEASURING METABOLIC REPROGRAMMING

METABOLISM AND BROWNING

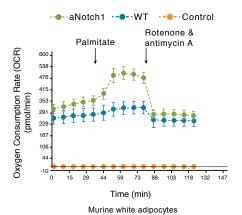
Excess white adipose tissue causes deleterious health effects, while brown adipose tissue is beneficial to overall health. Inducible brown adipocytes, also known as beige, brown-in-white, or brite adipocytes, present a promising treatment for obesity, diabetes, and metabolic disorders. Researchers are using XF Technology for relevant assay testing conditions and parameters. The XF Cell Mito Stress Test measures the key parameters of mitochondrial function: basal respiration, ATP-linked respiration, proton leak, maximal respiration, and spare respiratory capacity. The XF Glycolysis Stress Test measures the key parameters of glycolytic function: glycolysis, glycolytic capacity, and glycolytic reserve.



Adipocytes derived from C2C12 cells

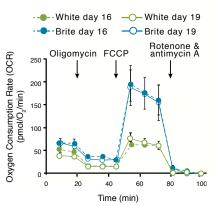
Sharma A et al., (2014) PLoS One.

XF Technology reveals a significant increase in exogenous palmitate oxidation in bone morphogenetic protein 6 (BMP)-stimulated cells.



Bi P et al., (2014) Nat Med.

XF Technology reveals Notch signaling inhibition increases fatty acid oxidation.



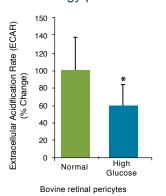
Rosiglitazone-deprived white and brite hMADs adipocytes

Loft A et al., (2015) Genes. Dev.

XF Cell Mito Stress Test reveals a link between rosiglitazone stimulation and metabolic profile in hMADs-derived brite adipocytes.

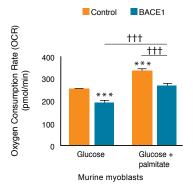
METABOLISM AND SUBSTRATE UTILIZATION

Nutrients are critical to maintaining cellular homeostasis and have a significant effect on metabolism. Metabolic disorders can cause abnormal processing of various nutrients leading to metabolic stress. XF Technology provides the capability to test, analyze, and understand the collected data.



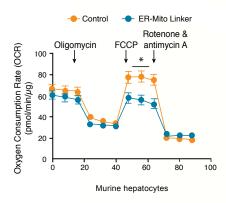
Trudeau K et al., (2011) Invest Ophthalmol Vis Sci.

XF Technology reveals decreased glycolytic activity in response to high glucose.



Hamilton DL, et al., (2014) Diabetologia.

XF Technology reveals Beta-site APP-clearing enzyme 1 (BACE1)-induced reduction in glucose oxidation.



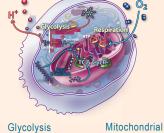
Arruda AP et al., (2014) Nat Med.

XF Cell Mito Stress Test reveals an inhibition of spare respiratory capacity due to increased ER-mitochondrial interaction.

THE WORLD'S MOST ADVANCED METABOLIC ANALYZERS

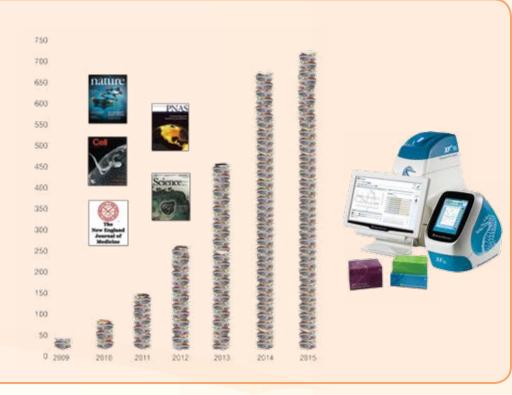
PROVEN TECHNOLOGY FOR CUTTING EDGE RESEARCH

There are over 2,000 references utilizing XF Technology published in leading journals such as Nature and Cell. Scientists are embracing XF Technology to identify metabolic phenotypes and reprogramming to target metabolic pathways for therapeutic purposes.



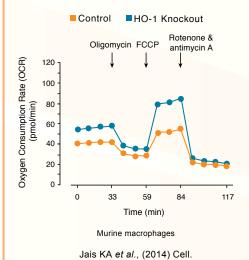
ECAR (Extracellular Acidification Rate)

Respiration
OCR (Oxygen
Consumption Rate)

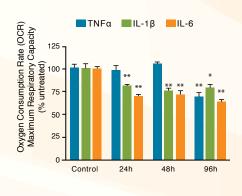


METABOLISM AND INFLAMMATION

Chronic inflammation and other immunological effects are linked to metabolic disorders. Researchers are using XF Technology to further their research into the connection between metabolic disorders and chronic inflammation.



XF Cell Mito Stress Test identifies heme oxygenase-1 (HO-1) gene requirement for metabolic programming of naïve macrophages.

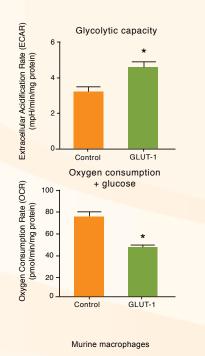


3T3-L1 adipocytes

Hahn WS et al., (2014)

Am J Physiol Endrocrinol Metab.

XF Technology demonstrates the reduction of respiratory capacity in the presence of proinflammatory cytokines.



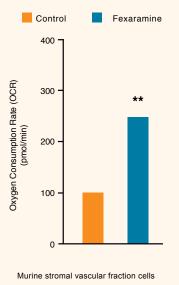
Freemerman AJ et al., (2014) J Biol Chem.

XF Glycolysis Stress Test reveals that glucose transporter 1 (GLUT1) overexpression increases macrophage glycolytic capacity while reducing mitochondrial respiration.

MEASURING THE KEY PARAMETERS OF CELL METABOLISM

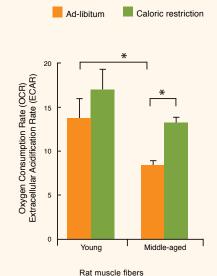
TRANSLATIONAL DIABETES AND THERAPEUTICS

Research into promising therapeutics may provide much needed cures for obesity, diabetes, and other metabolic disorders. XF Technology provides researchers the tools necessary for comprehensive investigations into disease progression and therapeutic candidates.



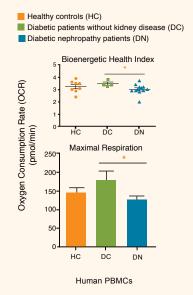
Fang S et al., (2015) Nat Med.

XF Technology reveals treatment with Fexaramine, a farnesoid X receptor agonist, increased mitochondrial respiration.



Chen CN et al., (2015) Am J Physiol Endocrinol Metab.

XF Technology reveals that caloric restriction improves metabolism in muscle from middle-aged rats.

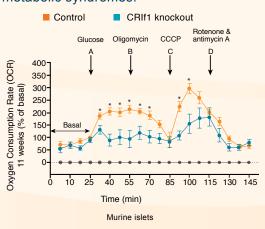


Czajka A et al., (2015) EbioMedicine.

XF Cell Mito Stress Test reveals a decreased maximal respiration in patients with diabetic nephropathy and correlates with patient Bioenergetic Health Index (BHI).

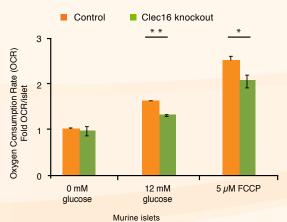
METABOLISM AND ISLETS

Using a cell line model to mimic the *in vivo* environment is crucial to any experiment. Researchers are using XF Technology with various cell lines and types to best represent the *in vivo* conditions associated with metabolic syndromes.



Kim YK et al., (2015) Diabetologia

XF Technology reveals a mitochondrial inner membrane protein (CRIf1) requirement for OXPHOS function in isolated islets.



Soleimanpour SA, et al., (2014) Cell.

XF Technology identifies a diabetes susceptibility gene requirement, Clec16a, for normal glucose-utilization in islets.

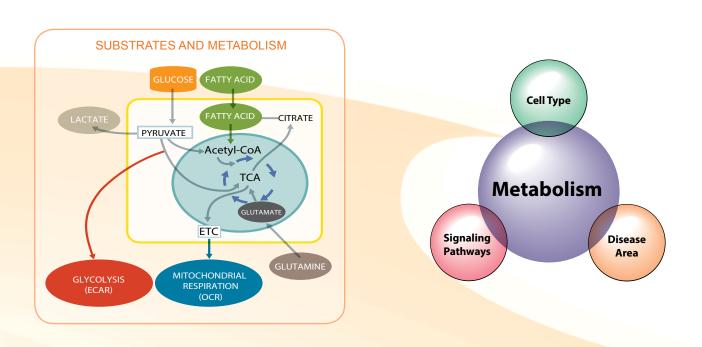
FUNCTIONAL XF METABOLIC ASSAYS

THE METABOLIC SYNDROME AND METABOLISM LINK

Metabolic syndromes are currently at pandemic levels and are a significant health risk to the general population. As one of the world's leading causes of death and disability worldwide, these syndromes include insulin resistance, obesity, non-alcoholic fatty liver disease, cardiovascular disease, and inflammation. The connection between metabolism and metabolic syndromes is evident, regardless of whether the research focuses on a specific cell type or tissue origin, disease area, or signaling pathway.

Mitochondrial dysfunction has emerged as a common thread amongst metabolic syndromes. Research into metabolic profiles and changes provides insight into browning, substrate and nutrient utilization, and inflammation. These metabolic paradigms are leading to opportunities for translational and therapeutic candidates.

XF Technology provides the capability to examine the mechanisms that link functional metabolism with the abnormalities that result in clinical disease.

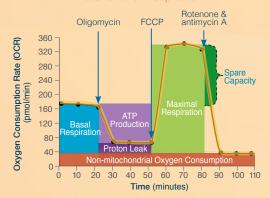


GOLD STANDARD METABOLIC ASSAYS

MEASURING THE KEY PARAMETERS OF CELL METABOLISM

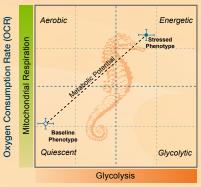
XF Cell Mito Stress Test Profile

Mitochondrial Respiration



XF Cell Energy Phenotype Test

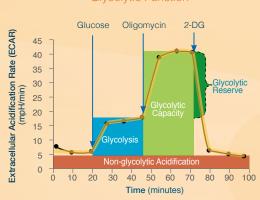
Metabolic Phenotype & Potential



Extracellular Acidification Rate (ECAR)

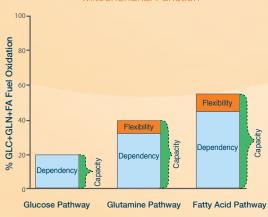
XF Glycolysis Stress Test Profile

Glycolytic Function



XF Mito Fuel Flex Test Profile

Mitochondrial Function





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