

Translational Metabolomics Shared Resource Redox and Bioenergetics Instrumentation and Scientific Examples

Instrumentation

The Translational Metabolomics Shared Resource (TraMSR) Redox and Bioenergetics unit provides state-of-the-art instrumentation, cutting-edge techniques, and sophisticated expertise dedicated to investigating cancer cell metabolism and redox signaling, and for designing custom probes for reactive oxygen species.



Agilent Seahorse XF96, Seahorse XFe96, Seahorse XFe24

Seahorse XF instruments enable researchers to interrogate cellular bioenergetics function in real time by simultaneously monitoring oxygen consumption and extracellular acidification rates in a 24- or 96-well plate format.



Shimadzu UHPLC LCMS8030 system

The Shimadzu LCMS8030 is a liquid chromatograph coupled with UV-Vis absorption and mass spectrometry detection. It allows selective detection and absolute quantification of small molecules (endogenous metabolites, drugs, chemical probes) in chemical mixtures and cell or tissue extracts.

Scientific Examples

Mitochondrial stress assay

Analysis of the protective effects of carvedilol against doxorubicininduced loss of mitochondrial function in iPSC-derived cardiomyocytes

In: Carvedilol Phenocopies PGC-1α Overexpression to Alleviate Oxidative Stress, Mitochondrial Dysfunction and Prevent Doxorubicin-Induced Toxicity in Human iPSC-Derived Cardiomyocytes. <u>https://doi.org/10.3390/antiox12081585</u>

Glycolysis stress assay

Role of p38y MAPK in aerobic glycolysis (Warburg effect) in KPC cells.

In: p38γ MAPK Is Essential for Aerobic Glycolysis and Pancreatic Tumorigenesis. <u>https://doi.org/10.1158/0008-5472.CAN-19-3281</u>





Ο.5 μΜ

Mito-MGN

HDH

Vehicle

10

20

30

ECAR (mpH/min)

50

OCR (pmol/min)

350

Δ2 μM

Mito-MGN

нĀн

40

Bioenergetic status

Mitochondria-targeted magnolol improved bioenergetic function in T-cells mostly via increased glycolytic (ECAR) function.

In: Synchronous effects of targeted mitochondrial complex I inhibitors on tumor and immune cells abrogate melanoma progression. https://doi.org/10.1016/j.isci.2021.102653

Superoxide (O₂^{•–}) measurements

Stimulation of mitochondrial superoxide production by mitochondria-targeted lonidamine.

In: Targeting lonidamine to mitochondria mitigates lung tumorigenesis and brain metastasis. <u>https://doi.org/10.1038/s41467-019-10042-1</u>



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Hydrogen peroxide (H₂O₂) measurements

Role of Notch4 in controlling H_2O_2 levels in blood vessels.

In: Nitric oxide synthase and reduced arterial tone contribute to arteriovenous malformation. https://doi.org/10.1126/sciadv.ade7280



в

100 M

NADPH oxidase 2-mediated oxygen consumption

Identification of NADPH oxidase-2 inhibitors by monitoring Nox2-dependent oxygen consumption.

In: Novel NADPH Oxidase-2 Inhibitors as Potential Anti-Inflammatory and Neuroprotective Agents. https://doi.org/10.3390/antiox12091660

Peroxynitrite (ONOO⁻) measurements

Development of two-photon fluorescent probe for peroxynitrite.

In: Two-photon fluorescent probe for cellular peroxynitrite: Fluorescence detection, imaging, and identification of peroxynitrite-specific products. https://doi.org/10.1016/j.freeradbiomed.2021.04.011



Control

Δ

OCR (pmol O₂/min) 0 05 00

- Η Ο μΜ - Φ 5 μΜ

100

Time (min)

32

200

IFN + LPS + PMA

0⁶0.1 1 10 10 TG15-132 (μM)

100