MIMETAS

3D Blood-Brain Barrier Toxicity Assay

Human brain microvascular endothelial cells (HBMEC)

OrganoService

- Kickstart drug toxicity studies in a physiologically relevant 3D human BBB model
 - Assess potential barrier-disrupting effects of your compounds in a perfusable 3D human brain microvascular endothelial model
- Multiple time point measurements
 - High sensitivity of compoundinduced barrier disruption with optimized time point measurements
- Data reporting in one go
 Receive a clear data report and raw data sheets

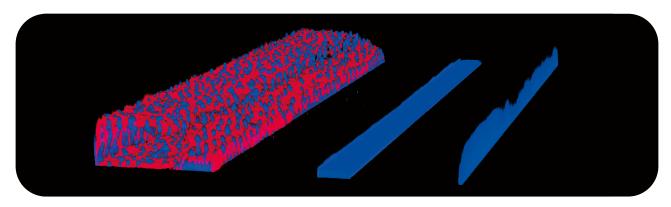
- Robust and reproducible assay
 Automated workflow ensures
 consistent performance and data
 quality
- TEER evaluation under physiological conditions
 Barrier integrity TEER data are obtained with the OrganoTEER®
- Complement your BBB toxicity studies with additional drug permeability studies

Study transport kinetics of your compounds across the BBB with our OrganoService BBB Small Molecule Transport Assay

About OrganoService

Blood-Brain Barrier Toxicity Assay

Profiling and screening of your compounds with a high-throughput barrier integrity assay on our established HBMEC BBB model.



Robust HBMEC BBB model in the OrganoPlate®

The role of the blood- brain barrier in drug development

The blood-brain barrier (BBB) ensures a homeostatic environment for the central nervous system (CNS) and is essential for brain function. When developing new therapies and assessing drug toxicity, on- and off-target BBB drug toxicity is a rate-limiting step, whereby assessing toxicity forms a critical part of the drug development quality control process. A common way of determining toxicity in the context of the BBB is via barrier integrity assessment, which up until now has been performed with 2D in vitro models that do not fully recapitulate key aspects of the BBB.

To assess the barrier in a sensitive andreliable manner, Trans Endothelial Electrical Re-

sistance (TEER) measurements are considered the gold standard.

OrganoService Blood-Brain Barrier Toxicity Assay

To study the BBB in vitro and assess barrier integrity in a high-throughput manner, we developed a robust primary HBMEC BBB model (OrganoReady® BBB) combined with a validated fluorescent barrier integrity assay and usage of the OrganoTEER®, which rapidly measures the transepithelial electrical resistance (TEER) of 64 endothelial cultures at once. By offering this as an off-the-shelf service, we allow you to focus on your compounds directly, while our scientist focuses on the biology and experiments.

What will you get:

- Screening of your compounds in a validated, physiologically-relevant BBB model
- Screening of up to 13 compounds in one concentration per OrganoPlate
- 4 technical replicates and 3 control compound
- Raw and normalized TEER values in a clear data report
- Extend the compound screen with more plates, in case a larger screen is desired

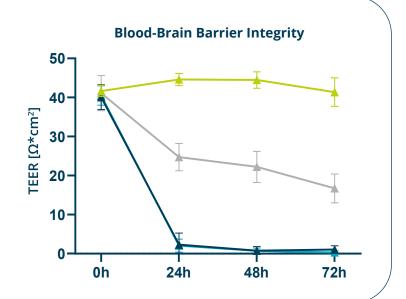
Results

Blood-Brain Barrier Toxicity Assay

Raw TEER values

Concentration-dependent disruption of blood-brain barrier integrity

- ★ Vehicle (DMSO)
- ★ Staurosporine 33 nM
- → TNFα & IL1β at 0.33 ng/mL
- \rightarrow TNF α & IL1 β at 10 ng/mL



Service details

Total compounds per plate

Number of concentrations/compound

Exposure

Number of technical replicates

Number of controls

Analysis method

Assay time

Time points

Data delivery

Data points/compound

Test article volume requirement

Test article solvent

Turn-around time

Compound testing

13

1

Apical

4

3 (staurosporine, TNFα and vehicle/medium)

TEER

2 days

0h, 24h, 48h

Raw & normalized TEER values

12

 $50 \mu L$ of 1,000X stock solution

DMSO, PBS, Water

4-6 weeks