

# In vitro air-liquid interface (ALI) exposure method to simulate in vivo inhalation exposure

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## Contact person

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## Organisation

**Name of the organisation** Vlaamse Instelling voor Technologisch Onderzoek (VITO)

**Department** Health

**Country** Belgium

**Geographical Area** Flemish Region

## Partners and collaborations

Vlaamse Instelling voor Technologisch Onderzoek (VITO)

## SCOPE OF THE METHOD

<b>The Method relates to</b>	Environment, Human health, Other: Alternative method for in vivo/human inhalation exposure. ALI exposure systems can be used to screen for human health impact of the following cases: Nano/ultrafine particles, Environmental/occupational compounds, Petroleum-derived substances,
<b>The Method is situated in</b>	Basic Research, Translational - Applied Research
<b>Type of method</b>	In vitro - Ex vivo
<b>Specify the type of cells/tissues/organs</b>	Lung cells

## DESCRIPTION

## **Method keywords**

in vitro and in vivo tool

in vitro

cell culture

lung

simulation

aerosol

(nano)particle

vapour

gas

## **Scientific area keywords**

in vitro

lung disease

respiratory toxicology

toxicity

Biomarkers

inflammation

aerosol

vapour

gas

(nano)particle

## **Method description**

VITO can offer expertise in animal-free methods for inhalation testing. VITO has an Air-Liquid Interface (ALI) platform with three ALI exposure modules: two commercial systems for bronchial studies and one in-house developed system for lower airway studies. At VITO ALI exposure modules, aerosol generation and online characterization instruments, and a battery of biological assays (e.g. TransEpithelial Electrical Resistance, cell viability/cytotoxicity, oxidative stress, and inflammatory response) can be used for screening of human health impact of e.g. Nano/ultrafine particles, Environmental/occupational compounds, Petroleum-derived substances, consumer products, pharmaceuticals (toxicity, efficacy, pharmacokinetics).

## Lab equipment

Air-liquid interface exposure system(s) ;

(Nano)aerosol generation facilities :

- Condensation Monodisperse Aerosol generator,
- Electrospray Aerosol generator,
- Single and Six Jet atomizers,
- Solid Aerosol generator,
- and a Soot generator;

Dedicated cell culture laboratories and assay facilities for biological endpoint measurements.

## Method status

History of use

Internally validated

Published in peer reviewed journal

## PROS, CONS & FUTURE POTENTIAL

### Advantages

Inhalation toxicity testing has traditionally been conducted using animals. Concerns related to the weak predictive ability as well as the use of animals, cost, time, and technical difficulty of *in vivo* inhalation resulted in the development of *in vitro* efficient and accurate, human-relevant lung cell-based methods to assess the potential hazards associated with xenobiotic exposure. The first *in vitro* inhalation studies were performed with submerged lung cell cultures. While these cultures are still widely used in *in vitro* pulmonary toxicity studies due to relative ease of handling, dispersion exposures are poorly representative of aerosol inhalation in humans. To overcome this, *in vitro* systems are developed for airborne exposure of lung cells at the air-liquid interface.

### Challenges

There are different ALI exposure systems on the market. No harmonized protocols are available.

## Modifications

Yes.

## Future & Other applications

Yes, ALI technology might be of relevant for biotech/pharma sector.

## REFERENCES, ASSOCIATED DOCUMENTS AND OTHER INFORMATION

### References

Frijns E et al. 2017. A Novel Exposure System Termed NAVETTA for In Vitro Laminar Flow Electrodeposition of Nanoaerosol and Evaluation of Immune Effects in Human Lung Reporter Cells. *Environmental Science & Technology*, 51 (9),

DOI:10.1021/acs.est.7b00493

Patent application for NAVETTA product; Flatbed air-liquid interface exposure module and methods (EP16200571.4; 2016, CN201780072888.1; 2019)

### Links

<http://www.piscltd.org.uk/vitrocell-prize>

<https://vito.be/nl/nieuws/vito-sterk-ontwikkeling-van-dierproefvervangende-test...>

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