# **Respiratory Toxicology**

Using Ex Vivo Precision Cut Long Slices and Reconstructed Human Airway Models

- Established Acute Toxicology Assays in 2D and 3D models
- Customized programs to address your specific respiratory questions
- Evaluate hazards of simple to complex aerosols
- Acute and chronic airway and oral mucosal tissue toxicology
- Dedicated laboratory with specialized dosing and exposure systems



Culturing Human Precision Cut Lung Slices into Roller Chambers

### **Human Precision Cut Lung Slices (hPCLS)**

- Human lung tissues cultured long-term at Air-Liquid Interface (ALI), submerged or in roller culture
- Cytotoxicity and Viability Endpoints -Lactate Dehydrogenase, Adenylate Kinase and WST-8/MTT conversion assays
- Airway Contractility Responsiveness

- Histological Evaluations
  - Alveolar vs. small airway changes
  - H&E architectural and cellular changes
  - Masson's Trichrome (collagen deposition)
  - Custom IHC staining (e.g., macrophages)
- Fibrosis Evaluations
  - Inflammatory Cytokine Analyses
  - Tissue collagen content by ELISA
- Activated macrophage quantitation

hPCLS are available for purchase. Contact <u>clientservices@iivs.org</u> for more information.

## **Reconstructed Human Airway Models**

- Human tissue models from oral cavity, nasopharynx, trachea, bronchi, small airway and alveolar spaces cultured longterm at ALI
- Cytotoxicity Endpoints Lactate Dehydrogenase (LDH) or Adenylate Kinase (AK) release assays
- Viability Endpoint WST-8 or MTT conversion assays (for GHS acute inhalation classification)
- Mucociliary clearance by ciliary beat frequency (analyses with SAVA)
- Inflammatory Cytokine Analyses by ELISA or Multi-plex
- Mucin production
- Goblet Cell hyperplasia evaluation



### **Exposure Systems and Other Endpoints**

#### **Exposure Systems:**

- Vitrocell® VC-1
  - Climate Chamber with 12/12 or 12/6 and
  - Exposure Modules
- VitroCell® CLOUD
- Tecan D300 Digital Dispenser

#### **Other Endpoints:**

- Oxidative Stress
- TEER
- Protein Content
- RNA Analysis
- Proteomics



VitroCell® Cloud system for airway exposures generously donated by PETA Science Consortium International (PSCI)

Molecular
<b>Initiating</b>
Events

Cellular **Key Events** 

Tissue/ Organ Key **Events** 

Aqueous - Solvent- Aerosol		· (100, 10)	
Test System	2D Colle	2D DIII.	an knoue
Biomarkers/Events	2D Cells	3D RHuA	3D hPCLS
Oxidative Stress	<b>√</b>	<b>√</b>	
DNA Binding/Strand Breaks	✓	<b>✓</b>	_
Mitotoxicity	✓	✓ /	
Cytotoxicity	<b>✓</b>	✓	<b>✓</b>
Vitability	✓	✓	✓/
Macrophage Activation			Ø
Cytokine/Chemokine Response	<b>√</b>	✓	<b>√</b>
Tight Junction Integrity		₫	
ECM Deposition			$\Box$
Mucociliary Clearance		Ø	
Airway Contractility			Ø
Goblet Cell Increase		Ø	
Mucin Expression		✓ ☑ >—	
Chronic Inflamation		<b>/</b>	<b>✓</b>
Chronic Tissue Health		<b>✓</b>	$\square$

☑ = unique to Model

For more information, contact Client Services at clientservices@iivs.org.

