

Innovative R&D Solutions for Respiratory Diseases and Chemicals Testing

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Problems



- ☐ There is no *in vitro* model to test the long term effets of particles/chemicals on airways (today, only animal models are avalaible)
- ☐ The chemical industry is progressively forced to test the toxicity of chemicals without using animals (enforced by New European REACH LEGISLATION)
- ☐ The bio-pharmaceutical industry needs for faster and more efficient new drug entities discovery, screening and testing techniques

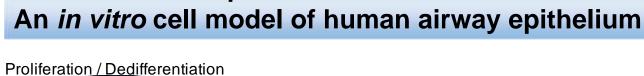


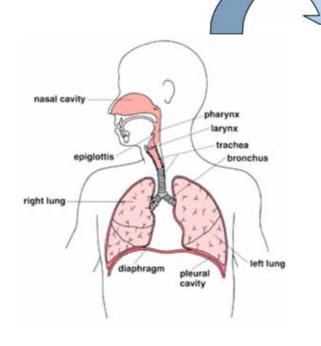


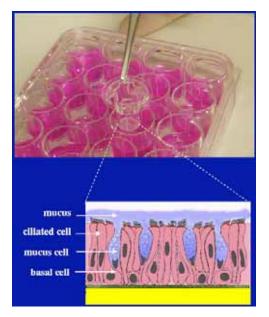
Our solution : MucilAir

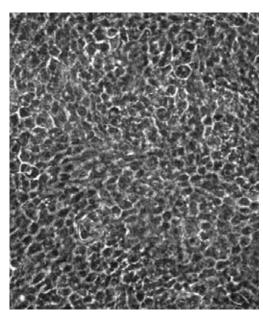


We produces and sells





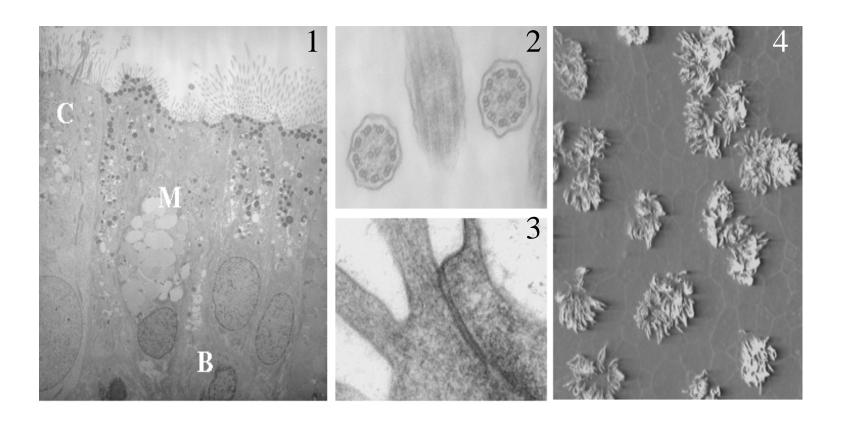








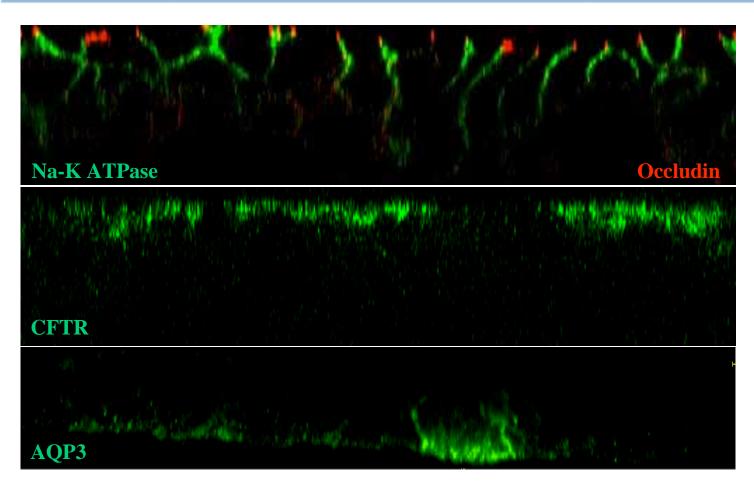
Similar, if not identical, ultra-structures as in the native tissues



Immunohistochemistry

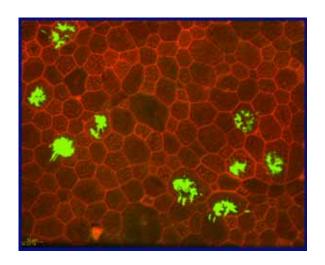


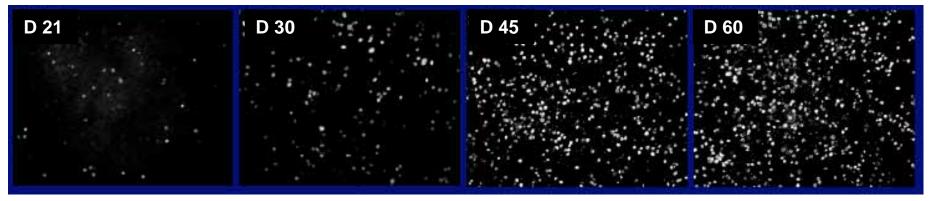
The distribution of markers and ions channels is identical to that observed in vivo





A time course of cilia formation monitored by anti-β-tubilin antibody

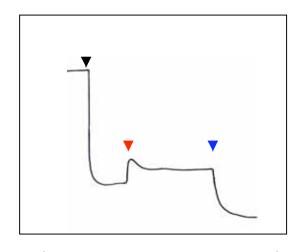






MUCILAIR show typical electrophysiological responses to ion channel inhibitors and activators

Current measurement in Ussing chamber



- **▼** Amiloride
- **▼** Isoproterenol
- **▼** Bumetanide

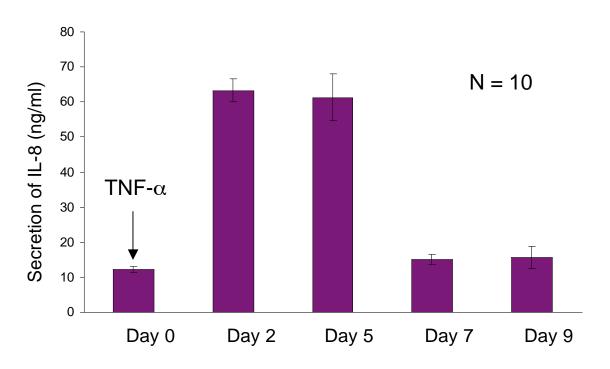
$\mathbf{R}(\Omega.\mathbf{cm}^2)$
(resistance)

 438 ± 18 (N=203)

- Amiloride (sodium channel inhibitor) reduces the current.
- Isoproterenol, a CFTR activator (chloride channel) increases current.
- Bumetanide, a general channel blocker, abrogates the total channel current activities.



MUCILAIR responds to pro-inflammatory mediators (TNF- α) in a physiological manner



A time course of IL-8 secretion (an inflammatory marker) after 30 minutes of TNF- α stimulation on a 3 months old epithelium. Note that this same epithelium can be reused several days later for other experiments

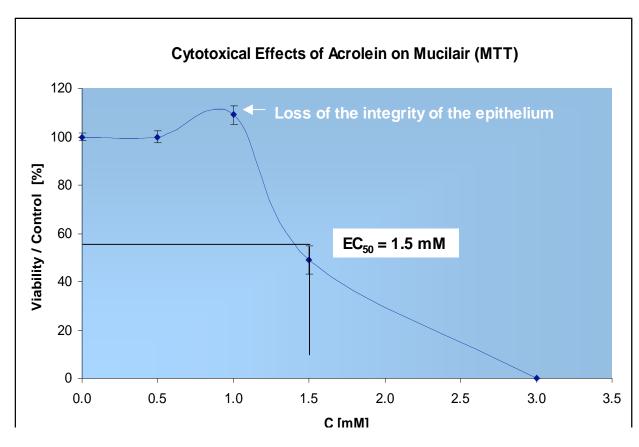
The main characteristics of MucilAir



- It closely mimics the morphology and functions of the normal human airway epithelium.
- It has a unique shelf-life of one year.
- It is easy to handle and maintain.
- It can be adapted to high-throughput screening and testing.

Cytotoxicity effect of Acrolein on Mucilair





N = 24

Incubation time: 24h

■ «dose-effect» response: 0.5-30 mM

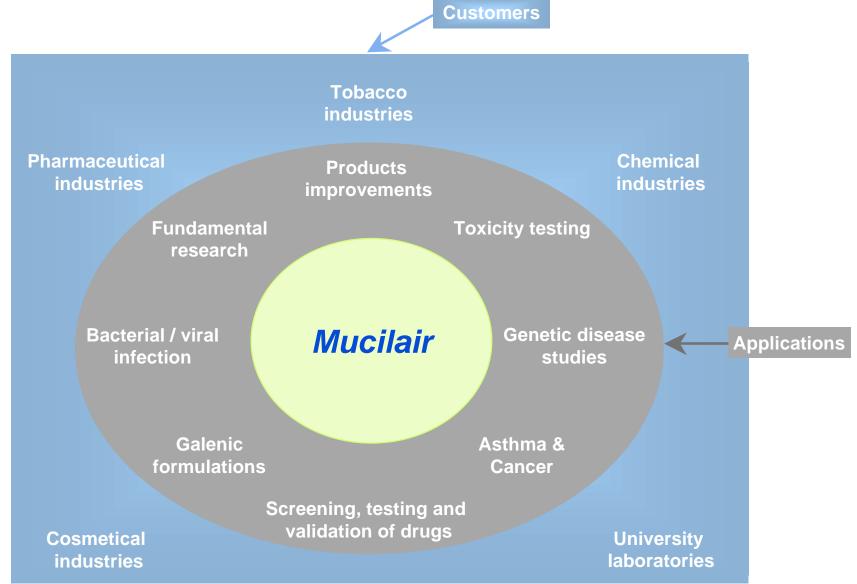
Indicators of Toxicity



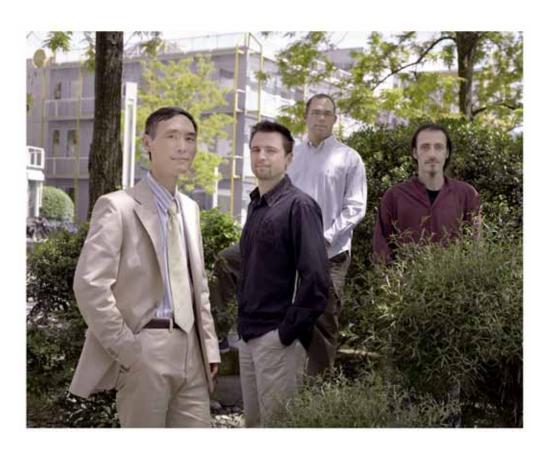
- Barrier function: TER
- LDH release
- MTT test
- Inflammatory mediator release
- Cilia beating
- Mucin secretion
- Intracellular ATP/ADP content
- etc.

Many valuable applications















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