## Reparatory Toxicity of Cigarette Smoke and ENDS Aerosols in MucilAir™ Airway Model

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### **OBJECTIVE**

To assess the reduced harm potential of aerosols from a commercially available ENDS product in comparison to 1R6F reference cigarette smoke by evaluating acute effects and the reversibility of damage after a 7day recovery period using MucilAir<sup>™</sup> Bronchial tissue model paired with Vitrocell® exposure system

### **APPROACH**

**AEROSOL CHARACTERIZATION EXPOSURE SYSTEM TESTED PRODUCTS** & DOSIMETRY ASSESSMENT & EXPOSURE REGIMEN 1R6F: nicotine, carbonyls **ENDS**: nicotine, carbonyls 1R6F smoke (ISO 20778) Culture **BIOLOGICAL ASSESSMENT** insert 64 min (N = 3)MucilAir™ • 10 → 1 L/min Tissue: Tissue • Cytotoxicity: WST-8 Microporous • Tissue integrity: TEER Membrane **Exposure** 24 h & • Ciliary Function: AA & CBF 7 days Mucociliary clearance\* Culture Medium recovery Histology: H&E\* Immunohistochemistry\* **ENDS** aerosols Supernatant: (ISO 20768) • Cytotoxicity: LDH MucilAir<sup>™</sup> Bronchial tissue 240 min, 6  $\rightarrow$  0.25 L/min • Oxidative Stress: 8-Isoprostane Vitrocell® HTP2.0+ 24-well module 180 min, undiluted • Inflammation: Cytokines Vitrocell® VC10® smoking robot

AA = cilia active area; CBF = cilia beat frequency; H&E = Hematoxylin and Eosin staining; LDH = Lactate Dehydrogenase Assay; TEER = transepithelial electrical resistance

\*Not presented here. Pending analysis.

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### MAIN RESULTS AND CONCLUSION

- 1R6F exhibited greater toxicity than ENDS with more severe and persistent effects across all endpoints evaluated, while ENDS demonstrated overall better recovery over time
- The reduction in the toxicity profile of ENDS aligns with their substantially lower levels of HPHCs compared to 1R6F

#### **IMPACT**

- This study provides strong in vitro evidence supporting ENDS as a less harmful alternative to combustible cigarettes
- Highlights the potential for ENDS to offer a reduced-harm alternative for adult smokers
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