Mitochondrial, Immunological, & Surfactant-Based Biosignatures to Assess Stress in Airmen Training Activities and Combat Environments

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1. Occupational Exposure

Airmen are exposed to a aerosols which can lead to inflamed airways and potentially longterm adverse outcomes. These experiments simulate aerosol exposures and identify stress markers that lead to impaired health or inhibit active-duty functions.



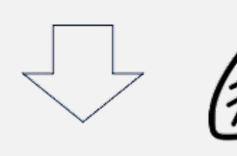


Concentration

Molecular Initiating Event

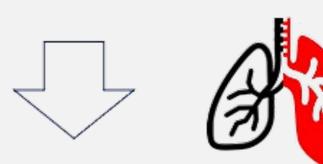


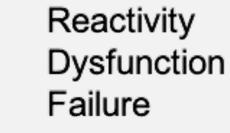
Lung Response



Structure

Diminished Lung Function





Decreased Airmen Performance

2. System Controls

Alveolar Immunological Model Controls			
Chemical Structure	Chemical	Туре	
PMA Ionomycin CH ₃ (CH ₂) _{1,1} CH ₂ CH ₃ CH ₃ CH ₃ CH ₃ H ₃ C H ₃ C H ₄ CH ₃ CH ₃ H ₄ CH ₃	Phorbol 12- myristate 13- acetate (PMA), lonomycin	Positive Control for Cell Activation	
H ₃ C NCO	Isophorone Diisocyanate (IPDI)	Positive Control for Sensitization	

Surfactant Drop	let Model Co	et Model Controls	
Chamical Structure	Chamical	Tree	

Chemical Structure	Chemical	Туре
CH ₂ OH OH OH OH	Lactose	Negative Control for Surfactant Inhibition
	Albumin	Positive Control for Surfactant

Inhibition

Work Flow

Mitochondrial

flux indicates

metabolic

Health &

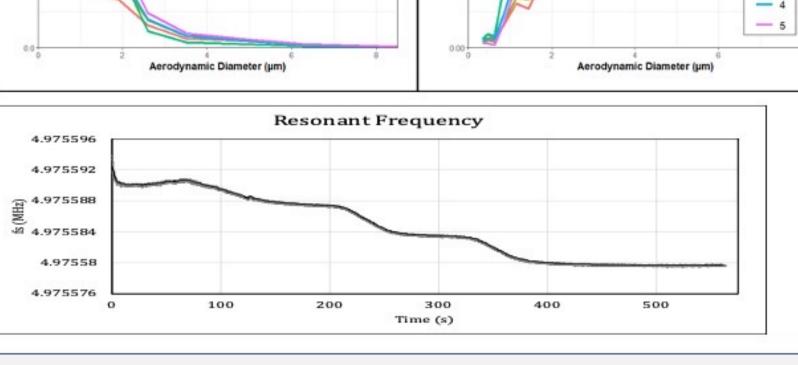
dependent on

mitochondria

Our Approach

3. Functional *In Vitro* Lung Models

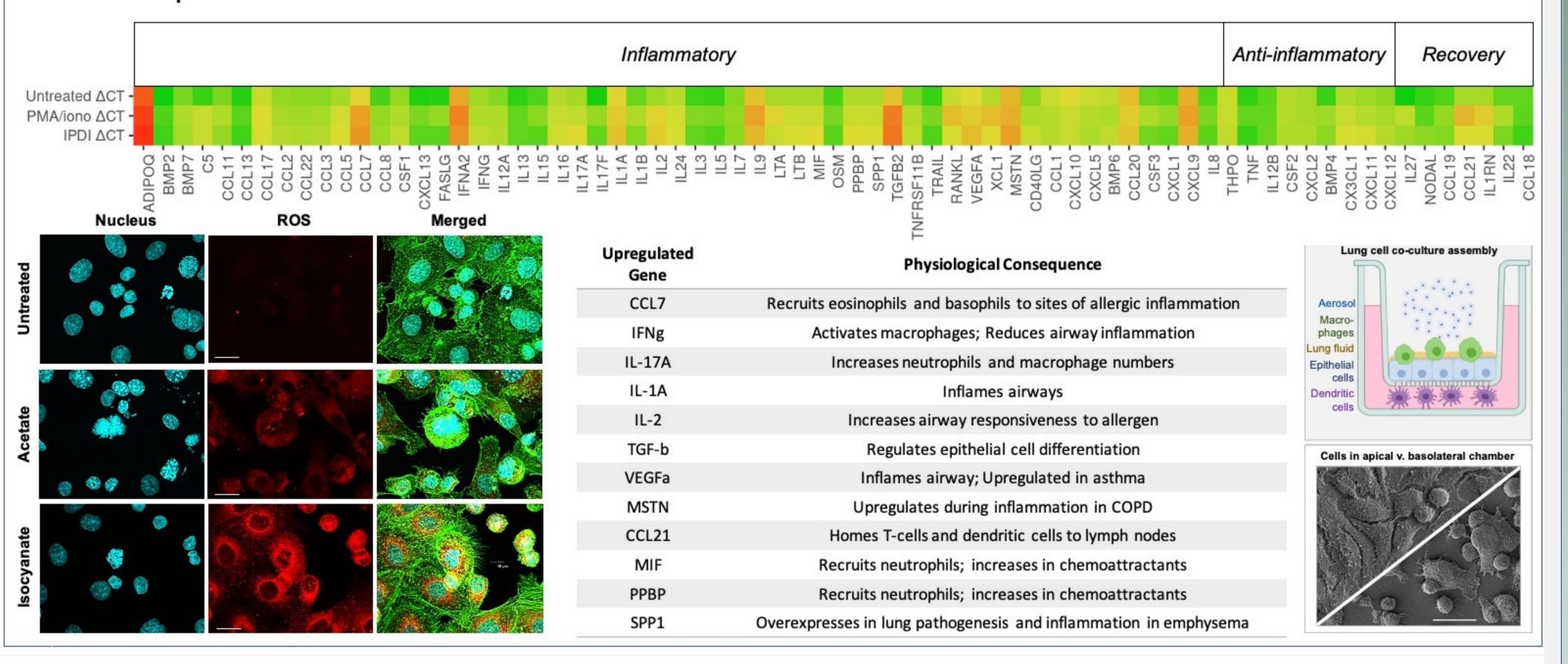
Four distinct cell-based models are used to assess immunological effects after exposure to Air Force are designed to produce data as input to predictive adverse health & performance effects.



Impact to Military Health

6. Improving READINESS through Performance

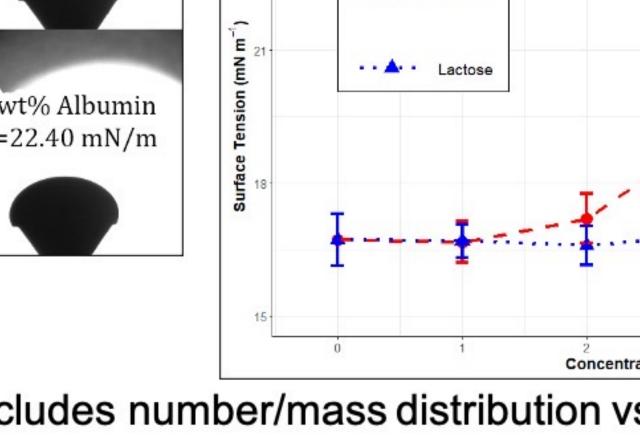
Compared to untreated cells, sensitizer treated cells have significant upregulation related to oxidative stress and show significant upregulation of F-actin. Known sensitizers lead to increases in mitochondrial activity as measured by oxygen species and increases in general cell activity as measured by F-actin. The physiological consequence may be an Increased risk for obstructive airway diseases and conditions which can inhibit performance.



7. Addressing RESILIENCE after Exposure

Aerosols are characterized by particle size distribution. Dose is measured by quartz crystal microbalance (QCM). Surface tension is calculated based on droplet shape analysis.

5wt% Lactose $\gamma = 16.95 \, \text{mN/m}$ - - Albumin 5wt% Albumin $\gamma = 22.40 \text{ mN/m}$

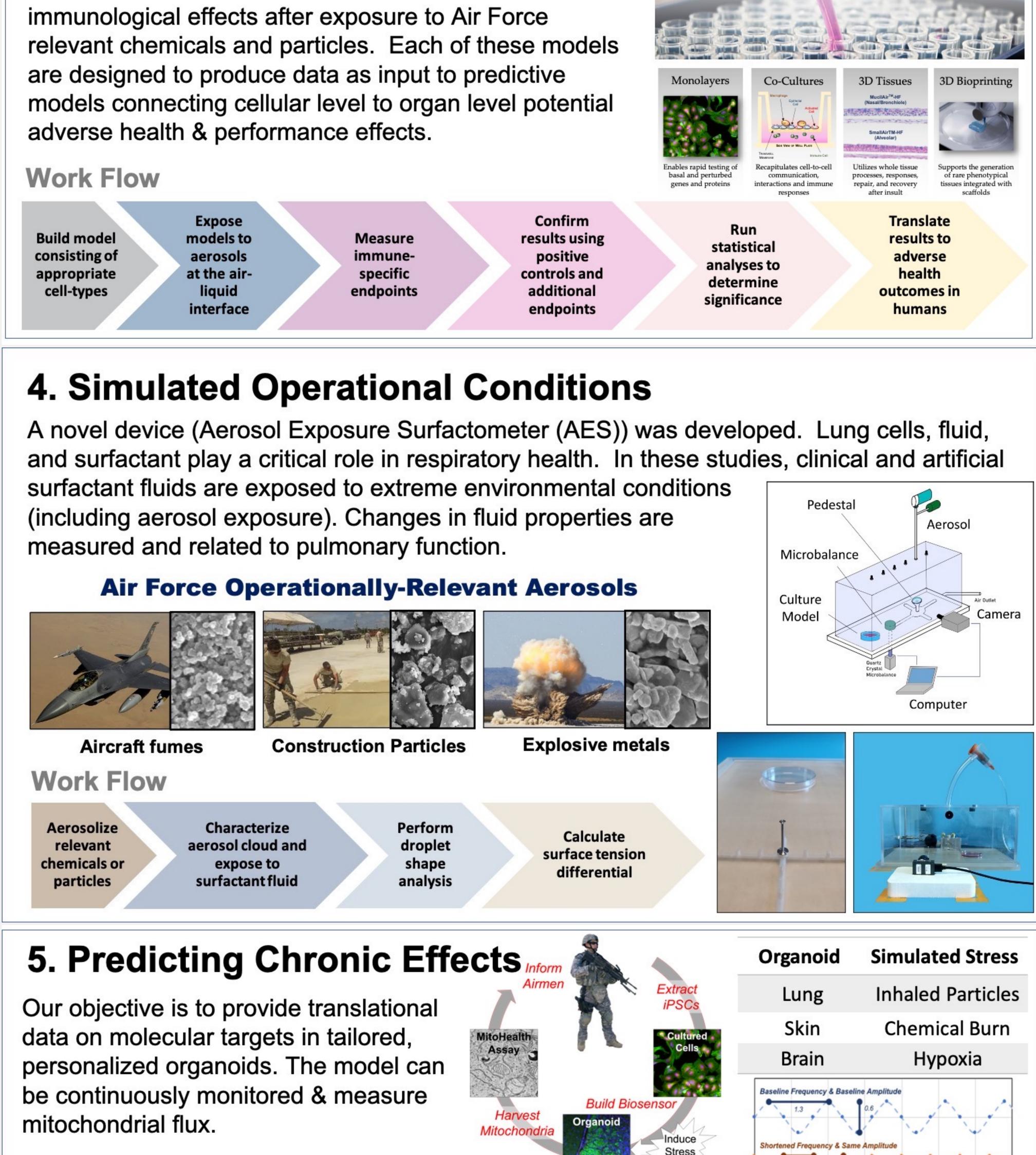


Data includes number/mass distribution vs. particle size, dose via QCM, droplet images, shape analysis, and surface tension



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Mitochondrial

flux is

measured

through

measurable

fingerprints

Data enables insight

into cell structure and

function and can

inform Airman stasis

before & after

induced stress