

# Pulmonary and Systemic Immune Response to Inhaled Oil Condensates

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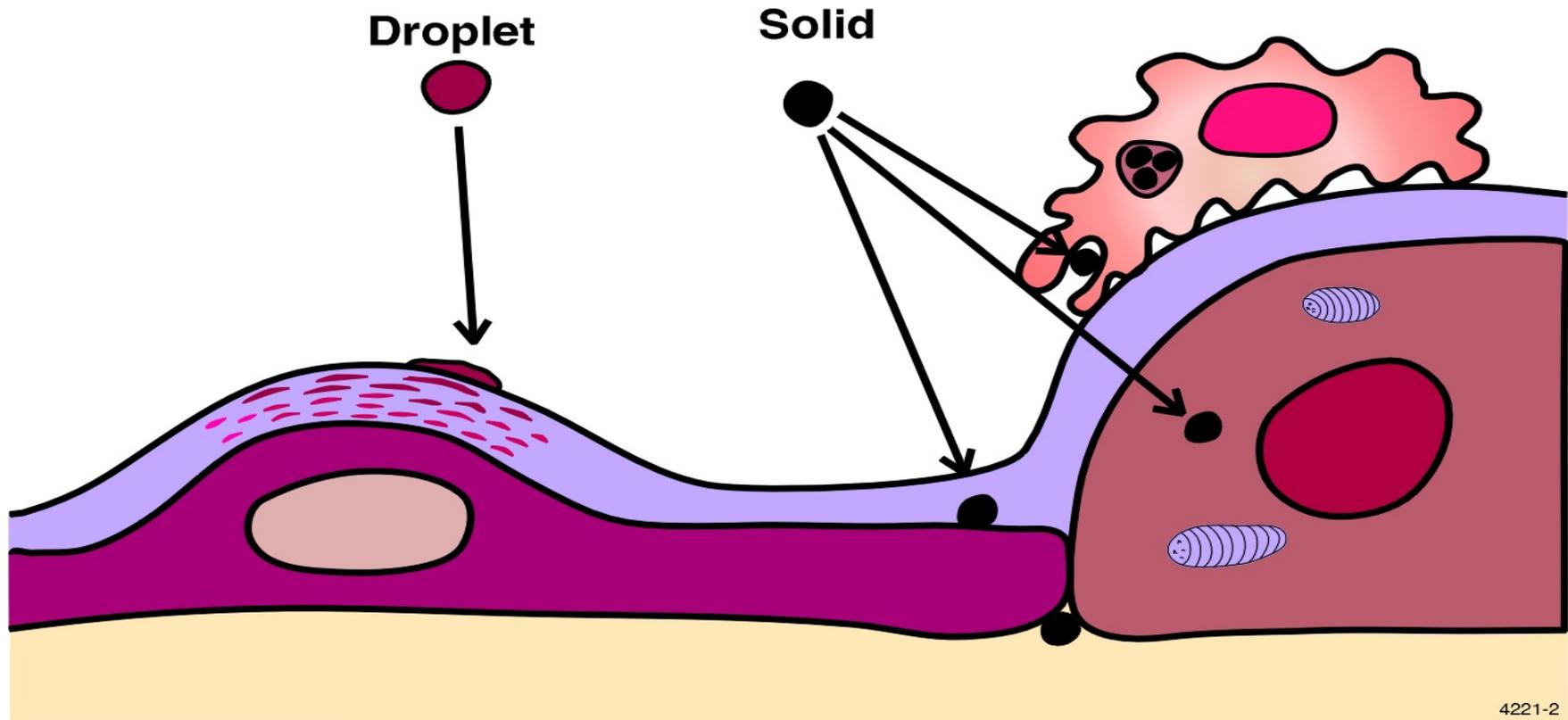


# Nanoparticles are a Topic of Interest

- Interest driven by a few studies, mostly with non-environmental particles
- Consideration of particle number standards
- Particle mass going down (thanks to you)
- Not certain if nanocondensates will be removed as quickly as solid “soot”
  - Limited information on hazard of this material

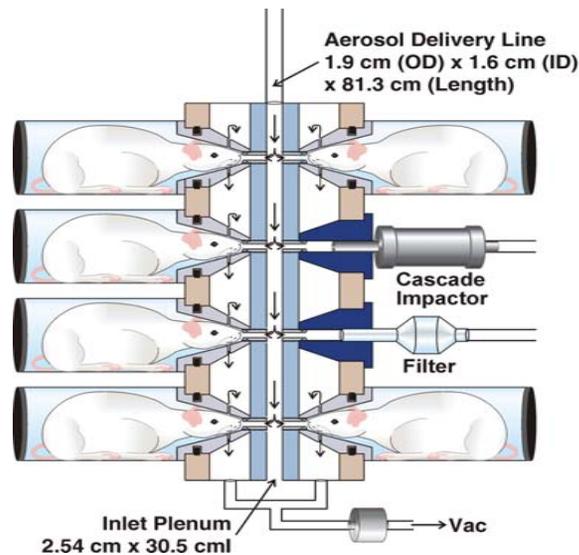
# Solid and liquid particles may “behave” differently at lung surface

## ULTRAFINES/NANOPARTICLES

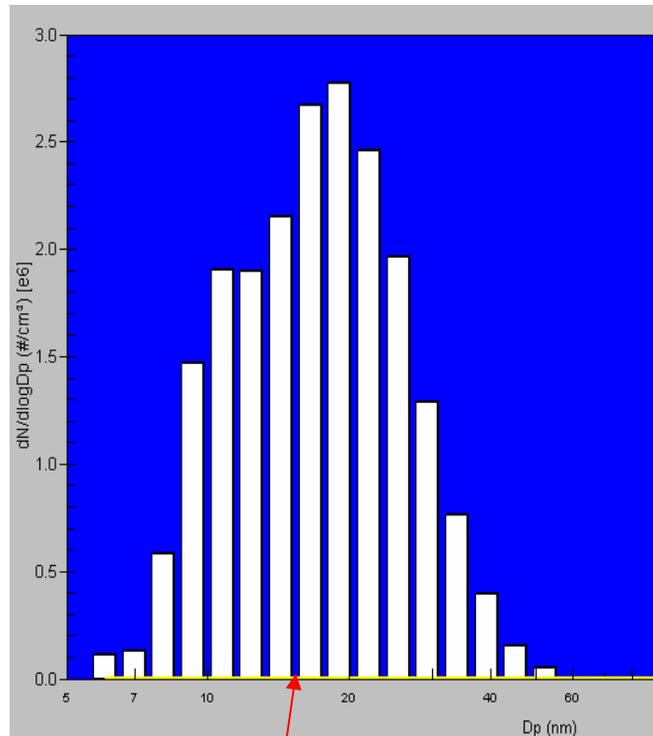


# Aerosol Exposures

- Mice exposed by nose-only inhalation 6 hr/day for 7 days (*match diesel protocols*)
- Shell Rotella-T15w-40 oil from 200 hr change in Cummins 5.9L engine on FTP cycle
- Oil nanocondensates generated by evaporation/condensation
- Diluted to target of  $10^6$  particles/cm<sup>3</sup> at a <20 nm particle size



# Exposure Atmosphere



17 nm

- Median diameter: 17 nm
- Particle Mass:  $300 \mu\text{g/m}^3$
- Particle Count:  $10^6 \text{ p/cc}$
- Surface Area:  $25 \text{ nm}^2/\text{cc}$
- Vapor HC: 0.6 ppm
- CO: 0.3 ppm
- NO<sub>x</sub>: < 40 ppb
- Additional analyses underway

Calculated dose, assuming 50 % deposition, =  $2 \mu\text{g/day}$

# Measurements of Response

Mice exposed to clean air or oil

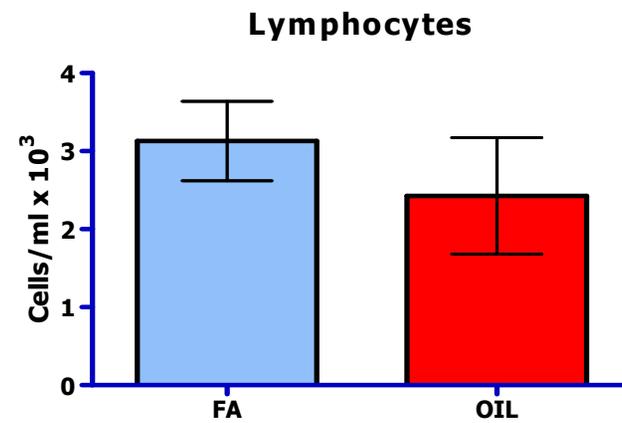
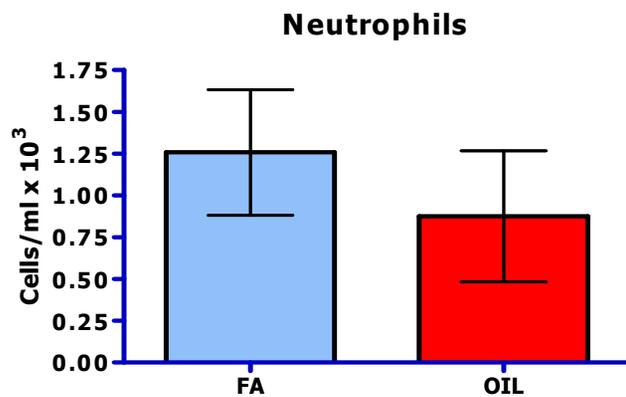
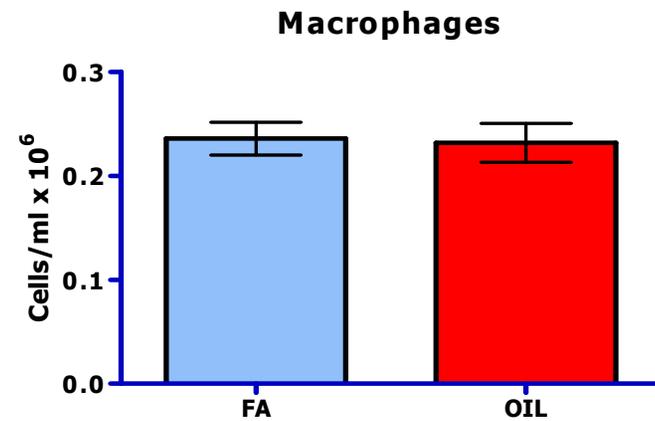
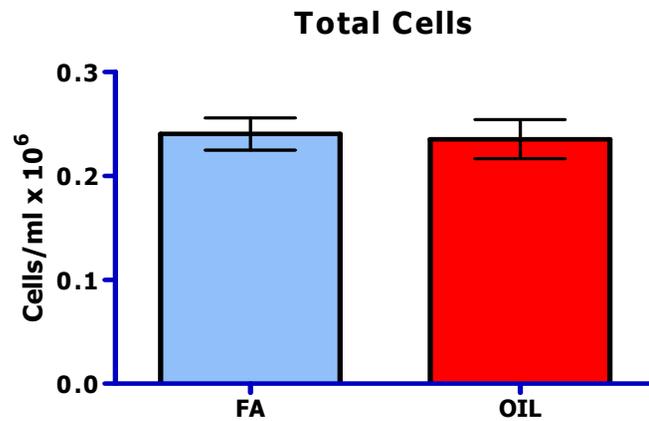
- Change in inflammatory cells in lung
- Change in biochemical indicators of injury, inflammation or chemical stress in lung or serum
- Presence of particles or lung tissue damage assessed by light microscopy
- Change in systemic immune response

# Does exposure increase response to a known irritant?

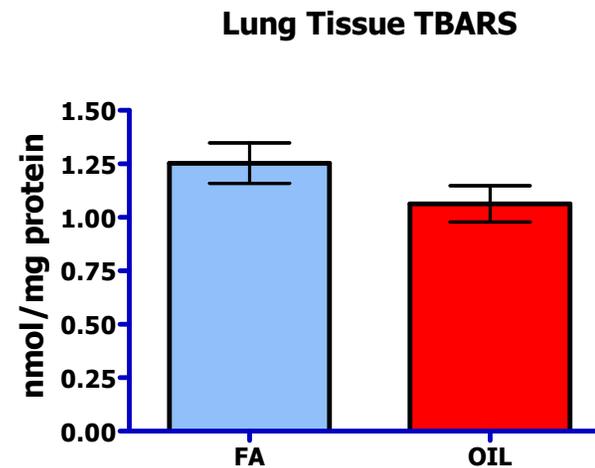
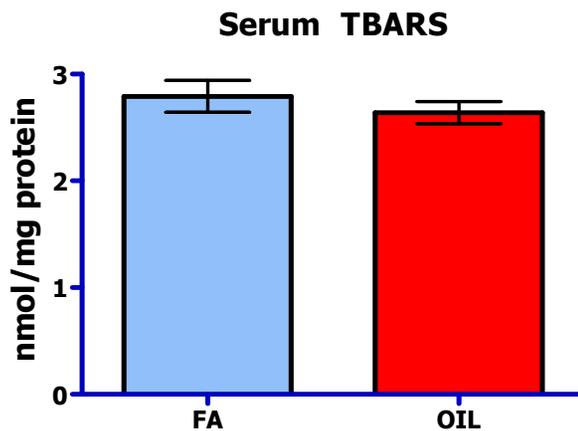
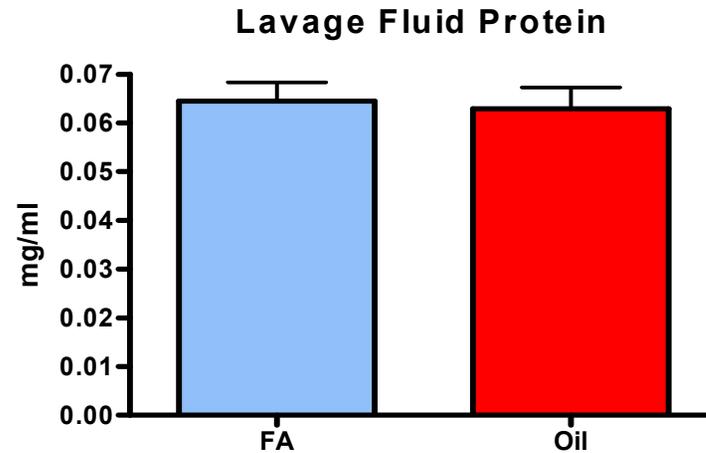
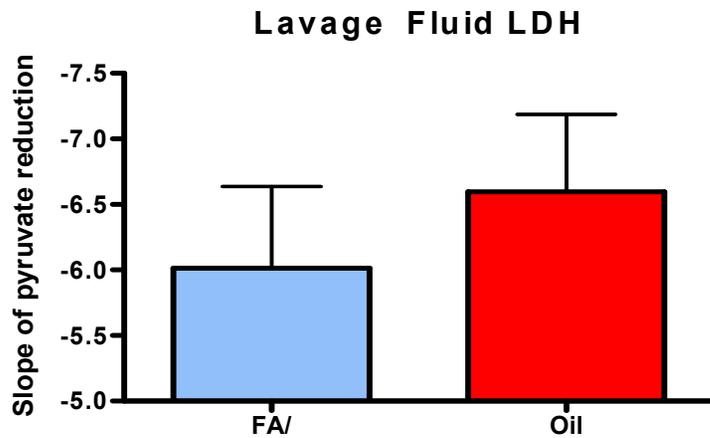
- Separate group of mice exposed (on day 4) by inhalation to ~5  $\mu\text{g}$  inhaled dose of bacterial cell walls to induce inflammation
  - Similar to “bronchitis”
- Measured same endpoints as non-LPS

# No increase in inflammatory cells

## Lavage Cells



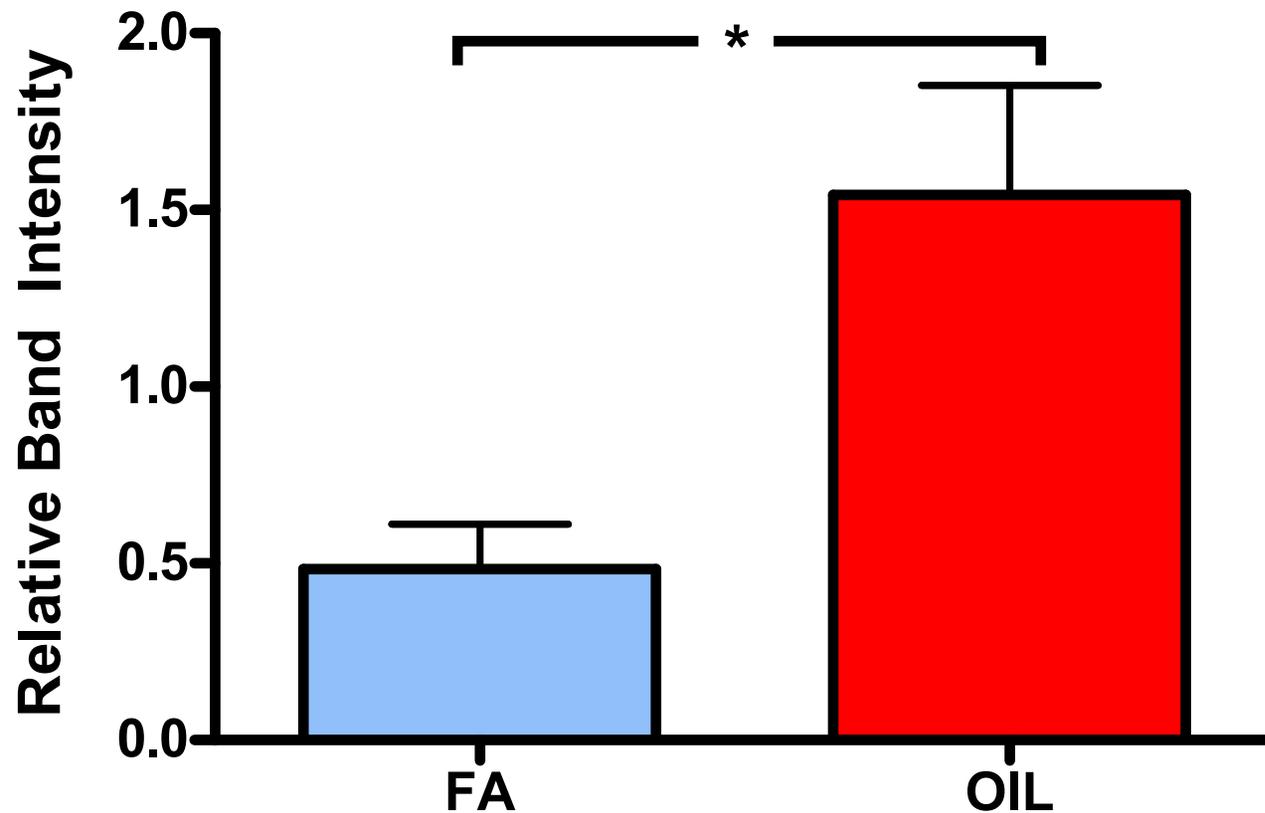
# Most Biochemical Indicators Did Not Increase with Exposure



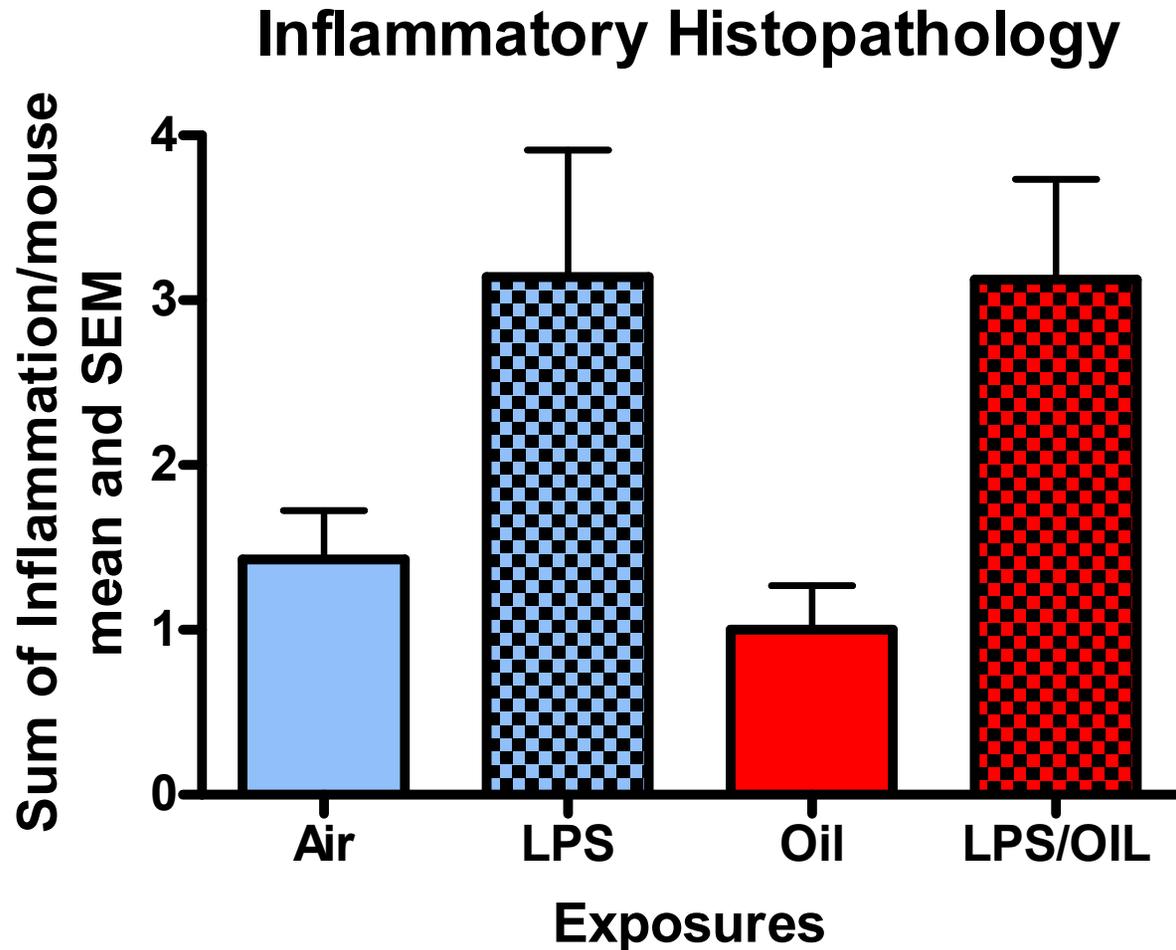
Also did not see increase in inflammatory cytokines

# A Sensitive Indicator of Oxidant Stress *D/D* Increase

## Heme Oxygenase-1 Protein

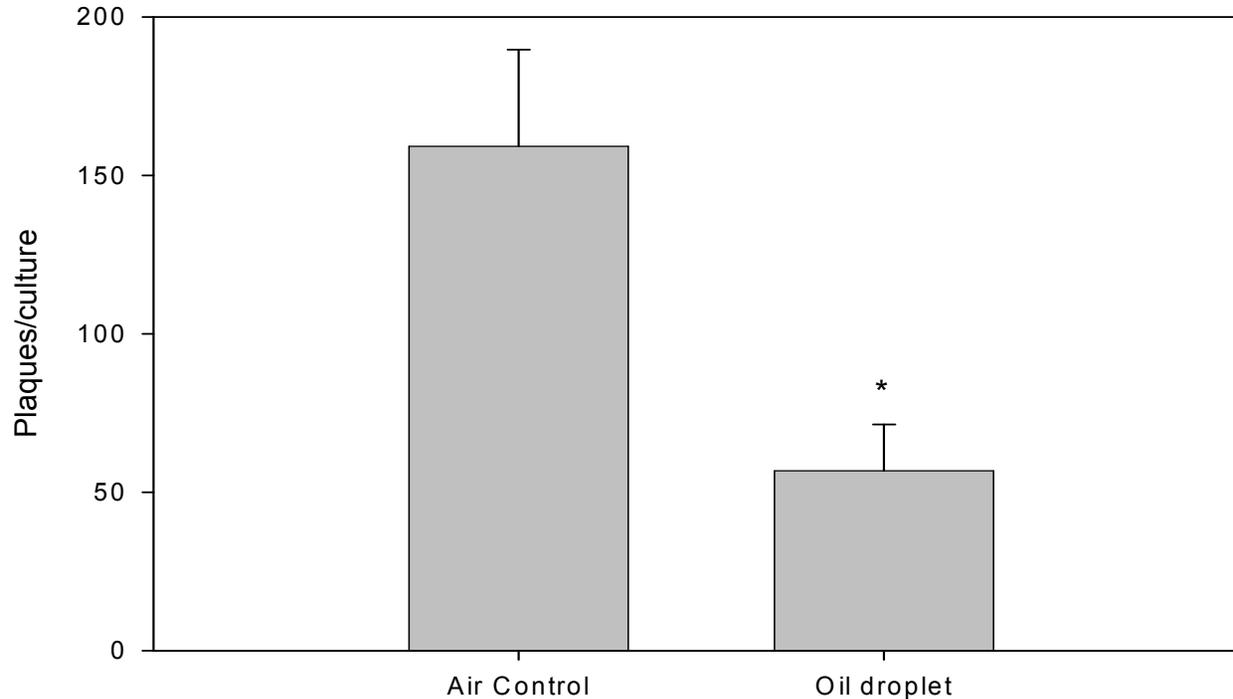


# LPS induced injury/inflammation not exacerbated by nanocondensates



LPS/OIL significantly greater than from FA/OIL

# Immune Response Decreased After Nanocondensate Exposure



\* statistically significant difference with air control ( $P < 0.05$ )  
n=7 mice/group

# Summary and Future Directions

- Inhaled nanocondensates showed very limited response for pathology, inflammatory cells, and biochemical indicators.
- While oil may have driven response to instilled material in previous studies with rats, it appears to be only mildly toxic at lower and inhaled doses in mice
- Only most sensitive indicators (oxidant stress/systemic immunity) showed physiological response to inhaled nanocondensates
  - These results to be confirmed and extended to no-effects levels
  - Results to also be compared with new oil, oil obtained from gasoline engines and nanosulfate for comparison.

IS NANOTOX = NANOHYPER??

# Acknowledgements

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