

Dist. Professor Magdalena Plebanski,

**Director:** Biomedical & Health Innovation Enabling Impact Platform (BHI-EIP)

**Head:** Translational Immunology & Nanotechnology Theme & Cancer Ageing and Vaccines (CAVA) Research Group School of Health & Biomedical Sciences, College of STEM



Growing the circular economy for plastics

Solving Plastic Waste

Cooperative Research Centre



RMIT University acknowledges the people of the Woi wurrung and Boon wurrung language groups of the eastern Kulin Nations on whose unceded lands we conduct the business of the University. RMIT University respectfully acknowledges their Ancestors and Elders, past and present. RMIT also acknowledges the Traditional Custodians and their Ancestors of the lands and waters across Australia where we conduct our business.



## Nano/microplastics and the lung

### Distinguished Professor Magdalena Plebanski

BScHon (Virology), Dip. St. (Psychology), MBA (Business), PhD (Immunology)

**Director:** Biomedical & Health Innovation Enabling Impact Platform (BHI-EIP)

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School of Health & Biomedical Sciences, College of STEM Human trials, Biomarkers, LoC Vaccines, Nanoparticles

What's next...

## Cancer Ageing and Vaccines Research Group

## Accelerator of Translational Research in Clinical Trials (ATRACT) Centre



School of Health and Biomedical Sciences, College of STEM

# CAVA Lab (key directly involved in research shown)

Dr. Kirsty Wilson Dr. Jennifer Boer

#### **Previous PhD Students**

Dr. Amlan Chakraborty Dr. Li Tan Prof. Rohimah Mohamud

## Prof. Magdalena Plebanski

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**FINANCIAL SUPPORT** NHMRC, ARC, CRC for Asthma KEY COLLABORATORS Only on Research Mentioned Robyn O'Hehir Jenny Rolland (Emeritus) Charles Hardy Cordelia Selomulya Simon Royce Jean-Pierre Scheerlinck Deanne Greenwood Rob Bischoff Tim Moss Deborah Glass Martina Deanekamp Vasso Apostolopoulos

- 1 in 7 Australians suffer from lung inflammation
- Chronic obstructive pulmonary disease (COPD) & asthma
- Third leading causes of death, >4 million deaths.
- Tobacco smoking and household air pollution major risk factors
  Ultrafine nanoparticles <100 nm major contributing factor</li>





Influenced by size, shape, charge, chemistry, adsorbed contaminants...

Are nano-plastics a risk factor?

Current Drug Delivery 46:176-90, 2014; Occup Environ Med. 74:868 , 2017; Expert Rev Respir Med. 12:941, 2018; Hum Immunol. 81:634, 2020 ; https://www.who.int/news-room/fact-sheets/detail/chronic-obstructive-pulmonary-disease-(copd)





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#### Polystyrene (PS) is a common environmental plastic

- PS microparticles (PSMP) 3 particles/m<sup>3</sup>
- PS-nanoparticles (PSNP) 13 particles/m<sup>3</sup>
- In vitro positively charged PSNPs in human alveolar macrophages and show toxicity
- Intranasal administration positively charged PSNPs facilitates brain entry Gaston et al., 2020, Roshanzadeh et al., 2020; Liu et al., 2022

#### Negatively charged PSNPs

- Inert and non-toxic in mice when injected, in sheep (in-house) and rats (ICP Firefly)
- No inflammatory ERK or TNF pathway induction on murine DC Karlson et al. (2013)



NO. OF CONSUMER PRODUCTS

# The global **nanomaterials market** valued at USD **11.99 billion in 2022** and USD **61.96 billion by 2032**

# Already in human medical products (PSMP&PSNP in human bone cement)

## Non-toxic Non-inflammatory



## Effect of negatively charged 50 nm PSNPs or 500 nm PSMPs on the lung



Sci Rep 7:14704, 2017 (sheep) Occup Environ Med. 74:868 , 2017 (humans)

## No lung inflammation or ROS production even at high doses PSNP or PSMP



# Nanoparticles are primarily taken up by dendritic cells in the lung and microparticles by macrophages



Hardy et al., Journal of Immunology, 2013





50 nm but not 500 nm **down-regulated** MAPK, Jak-STAT and PI3K **pro-inflammatory** signalling pathways

Dr. Ros Chapman (MIMR) (Tan et al., unpublished)



## But what about downstream consequences of exposure?





## **Decreased lung inflammation at time of allergen challenge**



### **Decrease lung inflammation ....and enhance clearance of influenza virus**

.



**PSNP** pre-treatment increases TNFR2+ Treg/T effector ratio upon allergen challenge

**PSNP CD103 DC activation** correlates with TNFR2+Treg expansion



Lung Ki67<sup>+</sup> of TNFR2<sup>+</sup>Foxp3<sup>+</sup> Treg (%)

Mohamud et al., Front. Immunol., 2017

# Novel homeostatic mechanism

Negatively charged PSNPs in the lung:

- imprint in the lung for resilience to allergic airways inflammation
- potentially non-specifically enhance immunity against viruses



## Debate

# Speakers



Dist. Professor Magdalena Plebanski, RMIT University



Professor Deborah Glass, Monash University A/Professor Mayur Garg, University of Melbourne



Dr Kirsty Wilson, RMIT University Health implications of plastics in our ecosystem

Ending Plastic Waste Symposium - Tuesday 6 August 2024

Facilitated by Professor Magdalena Plebanski (RMIT University)



Growing the circular economy for plastics

