# Characterising Innate Immune Responses and the Role of PD-1 in Patients with COPD

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B.BiomedSci (Honours)

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#### **Statement of Originality**

The thesis contains no material which has been accepted for the award of any other degree or diploma in any university or other tertiary institution and, to the best of my knowledge and belief, contains no material previously published or written by another person, except where due reference has been made in the text. I give consent to the final version of my thesis being made available worldwide when deposited in the University's Digital Repository\*\*, subject to the provisions of the Copyright Act 1968.

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## Conference Presentations and Publications Related to this Research

#### **Conference Presentations**

- 1. See H, Oldham R, Timmins N, Hansbro P, Wark P, Gibson P. Rhinovirus (RV) induced inflammatory response in mononuclear cells is enhanced by coculture with airway epithelium. Respirology 2007;[TSANZ Abstracts, Auckland] (E3).
- 2. See H, Oldham R, Timmins N, Hansbro P, Wark P, Gibson P. Response by peripheral blood monocytes (PBMCs) to rhinovirus (RV) is greater in cells exposed to infected epithelium in coculture than exposed to RV alone. Am J Respir Crit Care Med. 2007;[ATS 2007; San Francisco] (E3).
- Wark PAB, See HV, Simpson JL, Forbes RL, Hansbro PM. Peripheral blood monocytes (PBMCs) display innate antiviral response to rhinovirus (RV) infected bronchial epithelial cells. J Immunol. 2009;[Seattle, WASH] (E3).
- See HV, Simpson JL, Hansbro PM, Wark PAB. COPD patients PBMCs have an impaired immune response to rhinovirus-infected bronchila epithelium. Respirology 2011;[TSANZ Abstracts, Perth] (E3).
- See HV, Simpson JL, Hansbro PM, Wark PAB. Stable COPD patients have less rhinovirusinduced intracellular innate cytokines detected in PBMCs compared to healthy adults.
  Respirology 2012;[TSANZ Abstracts, Canberra] (E3).

#### **Journal Articles**

 See H, Wark P. Innate immune response to viral infection of the lungs. Paediatr Respir Rev. 2008 Dec;9(4):243-50. PubMed PMID: 19026365. eng.

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#### **List of Abbreviations**

7-AAD 7-Aminoactinomycin D

APC antigen presenting cell

ATCC American Type Culture Collection

BAL bronchial alveolar lavage

BEC bronchial epithelial cell

CAT COPD Assessment Test

CBA Cytometric Bead Array

CCQ Common Cold Questionnaire

CD cluster of differentiation

CFSE carboxyfluorescein succinimidyl ester

cfu colony forming units

COPD Chronic Obstructive Pulmonary Disease

CSE cigarette smoke extract

CXCL chemokine (C-X-C motif) ligand

CXCR chemokine (C-X-C motif) receptor

dsRNA double-stranded ribonucleic acid

ELISA enzyme linked immunosorbent assay

ELIspot Enzyme-Linked ImmunoSpot

FACs Fluorescence-activated cell sorting

FCS fetal calf serum

FEV<sub>1</sub> forced expiratory volume in one second

FSC forward scatter

FVC forced vital capacity

GOLD Global Initiative of Chronic Obstructive Lung Disease

H1N1 haemagglutinin 1 neuraminidase 1

HLA human leukocyte antigen

HMRI Hunter Medical Research Institute

hr hour

ICS inhaled corticosteroids

IFN interferon

IL interleukin

IQR interquartile range

JHH John Hunter Hospital

LPS lipopolysaccharide

mDC myeloid dendritic cells

MHC major histocompatibility complex

MFI median intensity fluorescence

MOI multiplicity of infection

NK natural killer

NTHi non-typeable haemophilus influenzae

OCS oral corticosteroids

pam3cys4K palmitoyl-3-cystein

pBEC primary bronchial epithelial cell

PBMC peripheral blood mononucleocytes

PBS phosphate-buffered saline

PC physical containment

PD-1 programmed death-1

pDC plasmacytoid dendritic cells

PD-L1 programmed death-ligand 1

pfu plaque-forming units

PHA phytohaemagglutinin

PolyI:C polyinosinic-polycytidylic acid

PRR pattern recognition receptor

RPMI Roswell Park Memorial Institute

RNA ribonucleic acid

RSV respiratory syncytial virus

RV rhinovirus

Spn streptococcus pneumoniae

SSC side scatter

ssRNA single-stranded ribonucleic acid

TCID<sub>50</sub> tissue culture infectious dose 50

TCM T lymphocyte central memory cell

TCR T-cell receptor

TEM T lymphocyte effector memory cell

TLR Toll-like receptor

TNF tumour necrosis factor

UV ultra-violet

WBC white blood cell