## **European Respiratory Society Annual Congress 2012**

**Abstract Number: 2018** 

**Publication Number: P2648** 

**Abstract Group:** 10.2. Tuberculosis

Keyword 1: Tuberculosis - mechanism Keyword 2: Immunology Keyword 3: Monocyte / Macrophage

**Title:** Role of the macrophage-inducible C-type lectin Mincle in the lung host defense against mycobacterial infections in mice

Ms. Friederike 12110 Behler Behler.Friederike@mh-hannover.de <sup>1</sup>, Dr. Kathrin 12111 Steinwede Steinwede.Kathrin@mh-hannover.de <sup>1</sup>, Mrs. Regina 12112 Maus Maus.Regina@mh-hannover.de <sup>1</sup>, Ms. Jennifer 12113 Bohling Bohling.Jennifer@mh-hannover.de <sup>1</sup>, Prof. Dr Tobias 12114 Welte Welte.Tobias@mh-hannover.de MD <sup>2</sup> and Prof. Dr Ulrich 12116 Maus Maus.Ulrich@mh-hannover.de <sup>1</sup>. <sup>1</sup> Department of Experimental Pneumology, Hannover Medical School, Hannover, Germany, 30625 and <sup>2</sup> Clinic for Pneumology, Hannover Medical School, Hannover, Germany, 30625 .

**Body:** The macrophage-inducible C-type lectin Mincle has been identified as receptor for the mycobacterial cell wall component trehalose dimycolate (TDM) of M. tuberculosis. We here examined the role of Mincle in lung protective immunity against mycobacterial pathogens in mice. We found that mice infected with M. bovis BCG responded with a delayed expression of Mincle on alveolar macrophages by days 14-21 post-challenge. In line with this finding, we observed that Mincle KO mice showed significantly reduced proinflammatory cytokine release and alveolar leukocyte recruitment as well as increased mycobacterial loads particularly in lung draining lymph nodes and spleens relative to wild-type mice infected with M. bovis BCG. Importantly, flow-sorted alveolar macrophages of wild-type mice responded with substantially greater proinflammatory TNF-a, KC, CCL2 and CCL5 mRNA levels to infection with M. bovis BCG relative to alveolar macrophages of BCG-infected Mincle KO mice. Together, the current study shows that Mincle exhibits delayed cell surface expression kinetics on alveolar macrophages upon M. bovis BCG challenge, thus acting as a 'delayed-type' regulator of proinflammatory macrophage activation during mycobacterial infections.