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Biphasic activation of complement and fibrinolysis during the human nasal allergic response

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## ACCEPTED MANUSCRIPT

1 Biphasic activation of complement and fibrinolysis during the human nasal allergic 2 response Ryan S Thwaites, PhD,<sup>1</sup>, Natasha C Gunawardana, MBBS MA MRCP,<sup>1</sup>, Verena Broich, MSc,<sup>1</sup>, Elizabeth 3 H Mann, PhD,<sup>2</sup>, Josefin Ahnström, PhD,<sup>3</sup>, Gaynor A Campbell, PhD,<sup>1</sup>, Sarah Lindsley<sup>1</sup>, Nehmat Singh, 4 MBBS, <sup>1</sup>, Tanushree Tunstall, MSc, <sup>1</sup>, David A Lane, PhD, <sup>3</sup>, Peter J Openshaw, MBBS FRCP PhD FRSB, <sup>1</sup>, 5 Catherine M Hawrylowicz, PhD, 2, Trevor T Hansel, MBBCh FRCPath PhD, 1,4 6 7 Affiliations: 8 <sup>1</sup> National Heart and Lung Institute, Faculty of Medicine, Imperial College London, W2 1PG, United 9 <sup>2</sup> MRC and Asthma UK Centre for Allergic Mechanisms of Asthma, King's College London, Guy's 10 Hospital, London, SE1 9RT, United Kingdom 11 <sup>3</sup> Centre for Haematology, Faculty of Medicine, Imperial College London, W12 0NN, United Kingdom 12 <sup>4</sup> MRC and Asthma UK Centre, Imperial College London, W2 1NY, United Kingdom 13 14 Funding: This work was supported by funding from the Imperial National Institute for Health 15 16 Research (NIHR) biomedical research centre, the NIHR Health Protection Research unit in 17 Respiratory Infections at Imperial College London in partnership with Public Health England, the 18 NIHR Imperial Patient Safety Translational Research centre and the British Heart Foundation (grant 19 FS/12/60/28974). The views expressed in this publication are those of the author(s) and not 20 necessarily those of the NHS, the National Institute for Health Research, or the Department of 21 Health. E.H.M was supported by a Medical Research Council (MRC) funded PhD studentship within 22 the MRC and Asthma UK Centre for Allergic Mechanisms of Asthma. 23 24 Acknowledgements: T.T.H, C.M.H and P.J.O conceived and designed the study. R.S.T., N.C.G., V.B., 25 E.H.M., J.A. and G.A.C. acquired the data. N.C.G., S.L. and N.S. collected study samples. R.S.T., N.C.G., 26 V.B., E.H.M., J.A., G.A.C., T.T., D.A.L., P.J.O., C.M.H. and T.T.H analysed and interpreted the data. 27 R.S.T. and T.T.H. drafted the manuscript which was critically revised for important intellectual 28 content and approved by all authors. R.S.T and T.T.H are the guarantors. 29 30 Abstract: Complement, coagulation and fibrinolysis contribute to the pathology of many respiratory 31 diseases. Here we detail the biphasic activation of these pathways following nasal allergen challenge. 32 Understanding these mechanisms may lead to therapeutic insight in common respiratory diseases. 33 34 Email Address for Correspondence: Trevor Hansel: t.hansel@imperial.ac.uk

35

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