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## Systems biology of the host response to severe infection

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### PhD PORTFOLIO

### **Hina Naz Khan**

**PhD period:** August 2015 – October 2019

PhD supervisors: Prof. Dr. T. van der Poll, Prof. Dr. A.H.Zwindermaan

and Dr. Brendon P. Scicluna

Courses	Year
- Genetic epidemiology	2016
- Scientific writing	2016
- Unix	2016
- System medicine	2016
- Practical biostatistics	2017
- E-Science	2017
- MIAGE course (Multi-omics integrative analysis of gene	2018
expression), CIPF, Valencia, Spain	
Seminars, workshops, retreats and master classes	
- Annual Amsterdam Infection and immunity institute (AI & II)	2016
PhD retreat	
- ARTeC seminar AMC/VUMC	2017
- BBMRI-omics workshop, Utrecht	2017
- Masterclass Prof. Douglas Golenbock	2017
- CEMM Journal Club	2015-2019
- CEMMinars	2015-2019
- Infectious diseases retreat (congress center Mennorode in	2017
Elspeet)	
- Kickoff symposium (Microbiota center Amsterdam)	2018
(Inter)national conferences	
- Summers Frontiers 2016: System Biology of innate immunity.	2016
Nijmegen, The Netherlands	
- 17 <sup>th</sup> International Congress of European Shock society. Paris,	2017
France	2017
- Sepsis 2017: New Successes, New challenges. Paris, France	2017
- New Frontiers in Innate immunity and Inflammation. Cluj-	2018
Napoca, Romania	

#### **Oral Presentations**

- "The circulatory small non-coding RNA landscape in health 2017 and community-acquired pneumonia."

17<sup>th</sup> International Congress of European Shock society. Paris, France

- "Identification of blood transcriptional networks dependent 2017 on lipopolysaccharide dose in human endotoxemia."
   AII&I PhD Retreat 2017.
- "Canonical and non-canonical RNA splicing in specific peripheral blood mononuclear cells of critically ill patients 2018 with sepsis".

New Frontiers in Innate immunity and Inflammation. Cluj-Napoca, Romania

#### Poster Presentations

- "The circulatory small non-coding RNA landscape in health 2016 and community-acquired pneumonia."
   Summers Frontiers 2016: System Biology of innate immunity.
   Poster presentation
- "Identification of blood transcriptional networks dependent 2017 on lipopolysaccharide dose in human endotoxemia."
   AII&I PhD Retreat 2017.
- "The circulatory small non-coding RNA landscape in health 2017 and community-acquired pneumonia."
  - Sepsis 2017: New Successes, New challenges. Paris, France
- "Canonical and non-canonical RNA splicing in specific 2018 peripheral blood mononuclear cells of critically ill patients with sepsis".

New Frontiers in Innate immunity and Inflammation. Cluj-Napoca, Romania

### **Tutoring, Mentoring**

- Student monitoring in "Identification of blood transcriptional networks dependent on lipopolysaccharide dose in human endotoxemia" project

#### List of publications

- Khan HN, Perlee D, Schoenmaker L, van der Meer AJ, Franitza M, Toliat MR, Nürnberg P, Zwinderman AH, van der Poll T, Scicluna BP. Leukocyte transcriptional signatures dependent on LPS dosage in human endotoxemia. J Leukoc Biol. 2019 Nov;106(5):1153-1160. doi: 10.1002/JLB.4A0219-050R. Epub 2019 Jul 7. PMID: 31280495; PMCID: PMC6852106.
- Khan HN, Brands X, Aufiero S, Hoogendijk AJ, Klarenbeek AM, van Engelen TSR, Haak BW, van Vught LA, Horn J, Schultz MJ, Zwinderman AH, van der Poll T, Scicluna BP. The circular RNA landscape in specific peripheral blood mononuclear cells of critically ill patients with sepsis. Crit Care. 2020 Jul 13;24(1):423. doi: 10.1186/s13054-020-03146-4. PMID: 32660590; PMCID: PMC7359566.
- Khan HN, Jongejan A, van Vught LA, Horn J, Schultz MJ, Zwinderman AH, Cremer OL, Bonten MJ, van der Poll T, Scicluna BP. The circulatory small non-coding RNA landscape in community-acquired pneumonia on intensive care unit admission. J Cell Mol Med. 2021 Aug;25(16):7621-7630. doi: 10.1111/jcmm.16406. Epub 2021 Jul 17. PMID: 34272809; PMCID: PMC8358855.
- 4. Matsumoto H, Scicluna BP, Jim KK, Falahi F, Qin W, Gürkan B, Malmström E, Meijer MT, Butler JM, Khan HN, Takagi T, Ishii S, Schultz MJ, van de Beek D, de Vos AF, van 't Veer C, van der Poll T. HIVEP1 Is a Negative Regulator of NF-κB That Inhibits Systemic Inflammation in Sepsis. Front Immunol. 2021 Nov 5;12:744358. doi: 10.3389/fimmu.2021.744358. PMID: 34804025; PMCID: PMC8602905.
- Ykema BLM, Hoefnagel SJM, Rigter LS, Kodach LL, Meijer GA, van Leeuwen FE, Khan HN, Snaebjornsson P, Aleman BMP, Broeks A, Meijer SL, Wang KK, Carvalho B, Krishnadath KK, van Leerdam ME; GIOCA-OES. Gene expression profiles of esophageal squamous cell cancers in Hodgkin lymphoma survivors versus sporadic cases. PLoS One. 2020 Dec 21;15(12):e0243178. doi: 10.1371/journal.pone.0243178. PMID: 33347497; PMCID: PMC7751872.

Kloek AT, Khan HN, Valls Seron M, Jongejan A, Zwinderman AH, Baas F, van der Ende A, van de Beek D, Ferwerda B, Brouwer MC. Variation in coagulation and fibrinolysis genes evaluated for their contribution to cerebrovascular complications in adults with bacterial meningitis in the Netherlands. J Infect. 2018 Jul;77(1):54-59. doi: 10.1016/j.jinf.2018.03.007. Epub 2018 May 7. PMID: 29746949.

#### **Submitted and in preparation Manuscripts:**

- 1. **Khan HN**, Brouwer MC, Geldhoff M, Scicluna BP, Zwinderman AH, van de Beek D., Bart Ferwerda. Protein quantitative trait locus of cerebrospinal fluid inflammatory mediators in patients with pneumococcal meningitis.
- 2. Malmström E, **Khan HN**, van't Veer C, Stunnenberg M, Meijer MT, Matsumoto H, Otto NA, Geijtenbeek TBH, Alex F. de Vos, van der Poll T and Scicluna BP. The long non-coding antisense RNA JHDM1D-AS1 regulates inflammatory responses in human monocytes
- 3. Hoefnagel SJM, Koemans WJ, **Khan HN**, Koster J, Meijer SL, van Dieren JM, Kodach LL, van Sandick JW, Calpe S, del Sancho-Serra CM, Correia AC, Van Laarhoven HWM, Van Berge Henegouwen MI, Gisbertz SS, Hulshof MCCM, Krishnadath KK. Subgroup analyses and predictive RNA signatures for response to chemoradiotherapy in esophageal adenocarcinoma: an aid towards personalized therapy.
- 4. Uhel F, Scicluna BP, Peters-Sengers H, Butler J, **Khan HN**, van Vught LA, Cremer OL, Bonten MJ, Schultz MJ, and van der Poll T. Comparative analysis of the host response to septic and non-septic shock.

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Hina Naz Khan March 2022, Barendrecht

#### About the author



Hina Naz Khan was born on 12<sup>th</sup> may 1984 in Rawalpindi, Pakistan. She received her primary and secondary education at different schools in Pakistan. In 2007 she obtained her Bachelor's degree in bioinformatics. Then in 2010 she attained her Master's degree in Bioinformatics and composed her master thesis entitled "Identification of an antiepileptic

compound with a more selective activity using *In-silico* approach". She has worked as a lecturer for O levels biology in Beaconhouse educational complex and also worked as lab instructor and a lecturer in Mohammad Ali Jinnah University, Pakistan.

In 2012 she moved to Netherlands. She pursed her education and received another master's bioinformatics degree in 2015 from Vrij Universiteit Amsterdam. She spent 8 months in Erasmus MC as a graduate trainee under the supervison of Dr. Andrew Stubbs and Dr. Pim French. She studied the application and validation of RNA-Seq for Glioblastoma patients' stratification in a clinical trial setting.

In August 2015, she started her PhD at Amsterdam UMC, location AMC, University of Amsterdam, under the supervision of Prof. dr. Tom van der Poll, Prof. dr. A.H. Zwinderman and Dr. Brendon P. Scicluna. During her PhD, she presented a number of transcriptomic signatures and cellular signaling pathways that may play a role in the pathophysiology of severe infections. The results of these studies are described and discussed in this thesis.

After her PhD, Hina continued her research and working as postdoctoral researcher (Bioinformatics) in the department of molecular genetics, Erasmus MC. 198