# Prof. Dr. Ulrich Johannes Steinhoff

Philipps-Universität Marburg Institut für med. Mikrobiologie Hans-Meerwein-Str.2 35043 Marburg Germany

Email: <u>ulrich.steinhoff@staff.uni-marburg.de</u> Phone: ++49-(0)6421-2866134 (Foto: https://www.uni-marburg.de/fb20/medmikrobio/personal/leitung.html)

### **Personal Statement**



Forschungs-

My scientific training in the Max-Planck-Institutes and in the lab of the Nobel-laureate Zinkernagel was in infection and inflammation. In 1999, I dropped into mucosal immunology by showing that Mycobacteria-crossreactive T cells induce tissue-specific pathology in the small intestine (Immunity, 1999). This finding sparked my interest to study mechanisms of inflammatory responses within the small and large bowel. Our discovery that intestinal tissues differ in their proteasome subunit composition was a breakthrough for the understanding of immunological and inflammatory reactions within the small intestine and the colon (J Exp. Med. 2002). This finding was confirmed for the human gut (Inflamm. Bowel Dis., 2009) and studies with IBD patients further revealed that expression of immunproteasomes is massively enhanced in inflamed tissues sites (JCI 2006). Molecular studies identified the immunoproteasome ß5i subunit as crucial catalytic center that controls immunoproteasome assembly as well as inflammatory signaling in IBD patients. Further, modulation of  $\beta$ 5i leads to massively reduced inflammatory reaction in experimental IBD (Gut 2010; Mucosal Immunology 2012). Consequently, we will now test  $\beta$ 5i-specific "new generation" proteasome inhibitors for their therapeutic potential and safety in chronic colitis and inflammation induced colon cancer. In 2010 I got a tenure track professorship at the university of Marburg for installing a clinical/basic science research group for mucosal immunology.

## **Positions and Employment**

| 1995-1999 | Scientist at the Max-Planck-Institute for Infection Biology, Berlin, FRG |
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| 1999-2002 | Group leader for mucosal immunology, Max-Planck-Institute for Infection  |
|           | Biology, Berlin, FRG   |
| 2003-2010 | Associate director, Max-Plank-Institute for Infection Biology, Dept. of  |

- Immunology, Berlin
- 2010-present Full professor, University of Marburg, Dept. of Medical Microbiology and Hygiene, Marburg

## **Other Experience and Professional Memberships**

1992 Member of the German society of Immunology (DGFI),



| 1995    | Member of the German society of Hygiene and Microbiology (DGHM)  |
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| 1996    | Member of the WHO advisory board (TDR)   |
| 2001    | Member of the Society for Mucosal Immunology (SMI)   |
| 2001    | Ad hoc grant reviewer for the Boehringer Ingelheim Fond (Germany), grant agency of the academy of sciences of the Czech Republic and Deutsche Forschungsgemeinschaft (DFG) |
| 2002    | Venia Legendi in Immunology (Habilitation) at the Humboldt University, Berlin  |
| 2004    | Consultant Immunologist DGFI (German Society of Immunology)  |
| 2004-08 | Coordinator of EU 6th Frame Work "Muvapred" (Mucosal vaccines against poverty related diseases)  |
| 2005    | Opponent and lecturer at the University of Gothenburg  |
| 2006    | Peer Review for the following journals: JEM, EJI, I&I, Int. Immunol, Gut, Mucosal Immunology   |
| 2010    | Associate Editor for European Journal of Immunology and Microbiology   |
| Honors  |  |
| 1984-85 | Fulbright fellowship for graduate studies at the University of<br>Massachusetts at Amherst, MA   |
| 1988    | Boehringer Ingelheim stipend   |
| 1991    | Research stipend for infectious disease, German Ministry for Education and Science   |
| 1992-94 | EMBO scholarship   |
| 2002    | Scientific paper of the year 2002 for contribution to the understanding of organ-specific autoimmunity, Paul Ehrlich Club of the University of Ireland                     |
| 2006-09 | Three manuscript were evaluated as scientifically outstanding by the faculty 1000  |
| 2010    | Excellence in Teaching, medical faculty of Marburg Publications  |

## Selected Publications

Müller U, **Steinhoff U**, Reis L, Hemmi S L, Pavlowitch J, Zinkernagel R (1993). Fuctional inactivation of the murine type I IFN-receptor abrogates antiviral defense. *Science* 264: 1918-1921

**Steinhoff U**, Brinkmann V, Klemm U, Aichele P, Seiler P, Prinz I, Zügel U and Kaufmann SHE (1999). Autoimmune intestinal pathology induced by hsp60-specific CD8 T cells. *Immunity* 11: 349-358



Kuckelkorn U, Ruppert T, Jungblut P, Zimny-Arndt U, Lamers S, Prinz I, Drung I, Kloetzel PM, Kaufmann SHE and **Steinhoff U** (2002). A link between autoimmunity and organ-specific antigen processing by 20S proteasomes. *J. Exp. Med*.195:983-990

Visekruna A, Joeris T, Zeitz M, Kaufmann SHE, Loddenkemper C, Kroesen A, Slavova N and **Steinhoff U** (2006). Proteasome-mediated degradation of  $I\kappa B\alpha$  and processing of p105 in Crohn disease and ulcerative colitis. *J. Clin. Invest.*, 116: 3195-3203 (faculty 1000 paper) Di Marco Barros R, Roberts NA, Dart RJ, Vantourout P, Jandke A, Nussbaumer O, Deban L, Cipolat S, Hart R, Iannitto ML, Laing A, Spencer-Dene B, East P, Gibbons D, Irving PM, Pereira P, **Steinhoff U**, Hayday A. (2016). Epithelia Use Butyrophilin-like Molecules to Shape Organ-Specific  $\gamma\delta$  T Cell Compartments. *Cell* 22:203-218.