A HUMAN PROTEIN ATLAS FOR NORMAL AND CANCER TISSUES

Peter Nilsson, Sophia Hober, Erik Björling, Fredrik Pontén and Mathias Uhlén

Dept. of Proteomics, School of Biotechnology, Albanova University Center, KTH -Royal Institute of Technology, Stockholm, Sweden

Antibody-based proteomics provides a powerful approach for the functional study of the human proteome involving the systematic generation of protein-specific affinity reagents. We have used this strategy to construct a comprehensive, antibody-based protein atlas for expression and localization profiles in 48 normal human tissues and 20 different cancers (1). The Human Protein Atlas is publicly available (<u>www.proteinatlas.org</u>) and contains, at present, approximately 1,2 million high-resolution images corresponding to more than 1500 antibodies towards human proteins. Each image has been annotated by certified pathologists to provide a knowledge base for functional studies and to allow queries about protein profiles in normal and disease tissues (2, 3). We have recently (4) used the antibody reagents for validation of potential biomarkers to Mantle Cell Lymphoma (MCL) as well as high-throughput biomarker analysis of thousands of serum samples using a reverse array format (5) and multiplex assays (6). Our results suggest it should be possible to extend this analysis to the majority of all human proteins thus providing a valuable tool for medical and biological research. in particular for biomarker analysis in various patient cohorts.

References:

(1) Uhlen and Ponten (2005). Antibody-based Proteomics for Human Tissue Profiling. *Mol Cell Proteomics* 4(4):384-393.

(2) Nilsson, Paavilainen, Larsson, Ödling, Sundberg, Andersson, Kampf, Persson, Szigyarto, Ottosson, Björling, Hober, Wernerus, Wester, Ponten and Uhlen (2005). Towards a Human Protein Atlas: High-throughput generation of mono-specific antibodies for tissue profiling. *Proteomics* 5(17):4327-4337.

(3) Uhlen *et al* (2005) A human protein atlas for normal and cancer tissues, *Mol Cell Proteomics* 4(12):1920-1932.

(4) Ek, Andreasson, Hober, Kampf, Ponten, Uhlen, Merz and Borrebaeck (2006). From gene expression analysis to tissue microarrays - a rational approach to identify therapeutic and diagnostic targets in lymphoid malignancies. *Mol Cell Proteomics* 5(6):1072-1081.

(5) Janzi, Ödling, Pan-Hammarström, Sundberg, Lundeberg, Uhlen, Hammarström and Nilsson (2005). Serum microarrays for large scale screening of protein levels. *Mol Cell Proteomics* 4(12):1942-1947.

(6) Schwenk, Lindberg, Sundberg, Uhlén and Nilsson (2007). Determination of binding specificities in highly multiplex bead based assays for antibody proteomics. *Mol Cell Proteomics* 6(1):125-132.