Oral Contributions

[MS9-02] Crystal structures of fungal velvet regulators reveal an unexpected similarity to NF-κB <u>Ralf Ficner</u>

Department of Molecular Structural Biology, Georg-August-University Göttingen, 37077 Göttingen, Germany Email: rficner@uni-goettingen.de

The morphological development of fungi and their production of secondary metabolites are mainly coordinated by velvet proteins containing a yet functionally and structurally uncharacterized velvet domain. We determined the crystal structure of the velvet domain of VosA from Aspergillus nidulans by means of SAD using a crystalsoaked with potassium iodine. The structure was refined at a resolution of 1.8 Å and revealed an unforeseen structural similarity with the Rel homology domain (RHD) of the mammalian transcription factor NF-kB. Based on this structural similarity several conserved amino acid residues present in all velvet domains were identified, and we demonstrate that these residues are essential for the DNA binding activity of VosA. The velvet domain also contains the dimerization interface of the VosA homo-dimer. Velvet proteins have been known to form also hetero-dimers. Therefore we crystallized the VosA-VelB hetero-dimer, and its crystal structure was refined at 2.2 Å resolution. The fold of the VelB velvet domain closely resembles that of VosA. However, the subunit arrangement is different with respect to the VosA homo-dimer, which has a significant impact on the DNAbinding surface. The structural similarity of the velvet domains and NF-kB suggests that defense mechanisms of both fungi and animals might be controlled by structurally related DNA-binding transcription factors.

Keywords: Transcription factor; DNA-binding domain