Tat System of *Escherichia coli*: Zn²⁺-bound Structures of tatD, ycfH and yjjV

<u>Vladimir N. Malashkevich</u>^a, Dao Feng^b, Frank M. Raushel^b, Steven C. Almo^a, ^aDepartment of Biochemistry, Albert Einstein College of Medicine, Bronx, USA. ^bDepartment of Chemistry, Texas A&M University, College Station, USA. E-mail: vladimir@medusa.aecom.yu.edu

The Escherichia coli Tat system mediates Sec-independent export of protein precursors bearing twin arginine signal peptides. Genes known to be involved in this process include *tatA*, *tatB*, and *tatC* that form an operon with a fourth gene, tatD. The tatD gene product has two homologues in *E. coli* coded by the unlinked *ycfH* and *yjjV* genes. The actual role of these enzymes and their substrates are not yet known, however, it was suggested that they might posses Zn²⁺dependent amidohydrolase activity. Significant number of amidohydrolases share TIM-barrel fold. The diversity of the catalytic mechanisms and substrate specificities is achieved through sequence and structural variation within the loop area. As a part of large-scale genomic effort to establish structure-function relationships within the amidohydrolase family, we determined high-resolution X-ray structures of tatD, ycfH and yjjV. Despite relatively low sequence identity of 24-29%, all three structures share similar overall fold, however, the number of Zn^{2+} ions in the active sites and their coordination differ significantly in three enzymes. Despite proposed deoxyribonuclease activity for tatD, none of the structures demonstrates nucleotide binding when co-crystallized with short DNA fragment. The potential functional roles of these enzymes will be discussed in the light of the structural and scarce biochemical data.

 Wexler M., Sargent F., Jack R.L., Stanley, R.L., Bogsch E.G., Robinson K., Berks B.C., Palmer T., *J. Biol. Chem.*, 2000, **275**, 16717.
Keywords: amidohydrolases, Zn²⁺-enzymes, tatD