## Unique Enzymatic Reaction Depending on Iron-Sulfur Cluster and Its Maturation

<u>Nobuyoshi Esaki</u>, Hisaaki Mihara, Tatsuo Kurihara <sup>1</sup>Institute for Chemical Research, Kyoto University, Uji, Kyoto 611-0011, Japan E-mail; esakin@scl.kyoto-u.ac.jp

Iron-sulfur (Fe-S) clusters are versatile cofactors of proteins that are involved in electron transport, enzyme catalysis and regulation of gene expression. 2-Hydroxyacyl-CoA dehydratases are two-component systems composed of an activator, Archerase, and the actual dehydratase, component D. Archerase, a homodimer with one [4Fe–4S] cluster, transfers an electron to component D, a heterodimer with 1-2 [4Fe–4S] clusters and FMN, concomitant with hydrolysis of two ATP. From component D the electron is further transferred to the substrate, where it facilitates elimination of the hydroxyl group. After elimination the electron is handed-over to the next incoming substrate without further hydrolysis of ATP (Figure 1).

The synthesis of Fe-S clusters and their insertion into apoproteins involves the function of complex cellular machineries. Cysteine desulfurase is a pyridoxal-5'-phosphate-dependent enzymes that cleave sulfur from cysteine via an enzyme cysteinyl persulfide intermediate and provides sulfur for the biosynthesis of Fe-S proteins. A similar but different mechanism is employed in the degradation of selenocysteine by selenocysteine lyase, a selenium-specific enzyme participating in selenium metabolism.

Properties and function of unique enzymes relating to Fe-S cluster will be presented.



Figure 1: Crystal structure of Archerase (left) and the reaction catalyzed by 2-hydroxyacyl-CoA dehydratase.

- J. Kim, M. Hetzel, C. D. Boiangiu, and W. Bückel, *FEMS Microbiol. Rev.* 28 (2004) 455.
- [2] K. P. Locher, M. Hans, A. P. Yeh, B. Schmid, W. Bückel, and D. C. Rees, J. Mol. Biol. 307 (2001) 297.
- [3] H. Mihara, and N. Esaki, *Appl. Microbiol. Biotechnol.*, **60** (2002) 12.
- [4] N. Esaki, T. Nakamura, H. Tanaka, and K. Soda, J.Biol. Chem. 257 (1982) 4386.