Thermococcus strain KS-1 由来プレフォルディン β サブユニットの結晶構造 Crystal Structure of prefoldin beta subunit from *Thermococcus* strain KS-1

木田 宗志¹, 菅野 由利², 飯塚 怜^{2,3}, 藤橋 雅宏¹, 養王田 正文², 三木 邦夫¹ 京大・院理・化学, ²農工大・院工・生命工, ³東大・院薬

Prefoldin (PFD) is a heterohexameric molecular chaperone that is found in eukaryotic cytosol and archaea. PFD is composed of α and β subunits and forms a "jellyfish-like" structure. PFD binds and stabilizes nascent polypeptide chains and transfers them to group II chaperonins for completion of their folding. Recently, a whole genome from *Thermococcus* kodakaraensis KOD1 was reported and shown to contain the genes of two α and two β subunits of PFD. The genome of *Thermococcus* strain KS-1 also possesses the two sets of α $(\alpha 1 \text{ and } \alpha 2) \text{ and } \beta \text{ (}\beta 1 \text{ and } \beta 2) \text{ subunits of PFD (TsPFD)}. However, the functions and roles of$ each of these PFD subunit have not been well investigated. We crystallized the TsPFD α 2- β 1 complex. The obtained crystals belong to the space group I422 with unit cell dimensions of a=b=71Å, c=114Å and diffracted to 1.9Å resolution. The SIRAS phased electron density map showed clear peaks corresponding to the β 1 subunit, whereas the α 2 subunit was not observed. The refinement of the structure composed only of the $\beta 1$ subunit reduced the crystallographic R_{work} and R_{free} factors to 17.7% and 19.7%, respectively. TsPFD β 1 subunits form a tetramer with four coiled-coil tentacles resembling the "jellyfish-like" structure of heterohexameric PFD. β hairpin linkers of β 1 subunits assemble to a β barrel "body" around a central four-fold axis. Size exclusion chromatography and multi-angle light scattering analysis shows that the β1 subunits form a tetramer at pH 6.8. The tetrameric β1 subunits can protect against aggregation of a relatively small proteins, such as insulin and lysozyme. The structural and biochemical analyses imply that PFD \(\beta \)1 subunits act as a molecular chaperone in living cells of some archaea.

Figure The "jellyfish-like" structure of the TsPFD $\beta1$ subunits tetramer (a) The side view of the TsPFD $\beta1$ subunits tetramer generated by the crystallographic four-fold symmetry. The four subunits are colored blue, green, red and orange, respectively. (c) The monomer structure of the TsPFD $\beta1$ subunit.

