RecJ エキソヌクレアーゼの一本鎖 DNA 特異性の構造的基盤の解明

Structure of RecJ exonuclease definces its specificity for single-stranded DNA

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RecJ single-stranded DNA (ssDNA) specific 5'-3' is exonucelease and deoxyribophosphodiesterase (dRPase), functions in DNA repair and recombination systems in almost bacteria and arhaea. As far, our group have reported the crystal structure of the catalytic core domain of RecJ from Thermus thermophilus HB8 (cd-ttRecJ) containing motif I-IV and DHHA1 motif bound to one Mn²⁺ and showed that cd-ttRecJ folded into two domains. However, none of the intact RecJ structure had been reported so far. To elucidate how RecJ achieves its high specificity for ssDNA, we determined the 2.15 to 2.50 Å resolution crystal structures of the intact ttRecJ alone, in complex with Mg^{2+} , and with Mn^{2+} . The entire RecJ consists of four domains which forms a molecule with an O-like structure (Fig. 1). One of two newly identified domains had structural similarities to an oligonucleotide/oligosaccharide-binding (OB)-fold (Fig. 2). The OB-fold domain alone could bind to DNA, indicating that this domain is a novel member of the OB-fold superfamily. The truncated RecJ containing only the core domain exhibited much lower affinity for the ssDNA substrate compared with intact RecJ. These results support the hypothesis that these structural features allow specific binding of RecJ to ssDNA. In addition, the structure of RecJ-Mn²⁺ complex suggests that the hydrolysis reaction catalyzed by RecJ proceeds through a two-metal ion mechanism.



Fig. 1 Overall structure of ttRecJ

Fig. 2 Structural comparision between ttRecJ-OB domain and the OB-fold domains of other proteins