

Structural Basis for Regulation of RpoS Turnover by Phosphorylation and ClpXP Anti-Adaptors

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In enterobacteria such as *Escherichia coli*, the general stress response is mediated by RpoS, the stationary phase dissociable promoter specificity subunit of RNA polymerase. RpoS is degraded by ClpXP during active growth in a process dependent on the RssB adaptor, which is thought to be stimulated by phosphorylation of a conserved aspartate in its N-terminal receiver domain and inhibited by stress-inducible and structurally distinct anti-adaptor proteins, IraD, IraM and IraP. Here we will present the crystal structure of full-length RssB bound to a beryllofluoride phosphomimic as well as structures of IraD:RssB and IraM:RssB complexes together with hydrogen-deuterium exchange and functional data in vitro and in vivo. Strikingly, in all of these three structures, RssB assumes a different conformation, heavily dependent on the plasticity of its inter-domain segmented helical linker. Our study emphasizes the importance of the 4-5-5 face of the RssB receiver domain in binding to RpoS and in serving as a landing platform for the IraD and IraM anti-adaptors, thereby excluding RpoS and inhibiting its delivery to ClpXP.