

Internal Seminar

Structural characterization of a cataract-active mutant of human γ S Crystallin

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Cataract, or opacification of the eye lens, is the leading cause of blindness world over. While age-related cataract is the result of accumulation of environmental and metabolic effects, congenital cataract, seen in newborn children, is essentially genetic in origin. A mutant of human γ S-crystallin, G57W, has been recently reported in a Chinese family wherein a young boy and his mother were found to have cataract in the center of the lens. We attempted to structurally characterize the G57W mutant and study its dynamics by solution NMR. In this endeavour, a suite of heteronuclear 2D and 3D NMR experiments with uniformly $^{13}\text{C}/^{15}\text{N}$ -labelled γ S-G57W has enabled almost complete sequence-specific ^1H , ^{13}C and ^{15}N resonance assignments. Unfolding kinetics of γ S-G57W upon heating and addition of chemical denaturants revealed γ S-G57W to be less stable compared to its wild-type. Efforts are on to study the inter-domain interactions in γ S-G57W and to understand the mechanism of cataractogenesis, since it is not clear as to how a single point mutation in γ S-crystallin compromises eye lens transparency and packing, and finally causes cataracts.

Tuesday, Oct 3rd 2017

02:00 PM (Tea/Coffee at 01:45 PM)

Auditorium, TIFR-H (FReT-B)