Potential role of pharmacological STAT3 inhibitors for cancer therapy

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Conventional anti-cancer therapeutic strategies involve the use of chemically synthesized drugs and/or administration of high-energy radiation to circumvent tumor growth. However, these strategies are generally poorly tolerated, often resulting in adverse side effect. As such, there is an urgent need to develop novel anti-cancer therapeutic agents that not only overcome the chemoresistance barricade, but also elicit minimal side effects and is well tolerated amongst patients of diverse demographics. Plant-based natural drugs contributes to primary health care in approximately 80% of the world’s population, with uses dating back to the ancient times as traditional herbal medicine by physicians such as Hippocrates. Our group is currently exploring the role of STATs family of cytoplasmic transcription factors that transmit signals, mediate intracellular signaling usually generated at cell surface receptors and transmitted to the nucleus. There is a strong evidence to suggest that aberrant STAT3 signaling promotes development and progression of human cancers by either inhibiting apoptosis or inducing inflammation, cell proliferation, angiogenesis, invasion, and metastasis. Suppression of activation of STAT3 results in the induction of apoptosis in tumor cells, and accordingly its inhibition by approaches such as tyrosine kinase inhibitors, antisense oligonucleotides, decoy nucleotides, dominant negative proteins, RNA interference and chemopreventive agents have been employed to suppress the tumorigenicity. However, the development of novel drugs for the targeting STAT3 that is both safe and efficacious remains an important scientific and clinical challenge. My talk will provide the evidence for critical roles of STAT3 in oncogenesis and discusses the potential for development of novel cancer therapies based on mechanistic understanding of STAT3 signaling.

Biography
Gautam Sethi has completed his Postdoctoral training at University of Texas MD Anderson Cancer Center and joined the Department of Pharmacology, Yong Loo Lin School of Medicine, National University of Singapore in 2008 as an Assistant Professor and was promoted to Associate Professor in 2015. The focus of his research over the past few years has been to elucidate the mechanisms of activation of oncogenic transcription factors such as NF KB/STAT3 by carcinogens and inflammatory agents and the identification of novel inhibitors of these proteins for prevention of and therapy for cancer. From traditional Chinese and Indian medicinal plants, his group has identified numerous small molecules that can suppress various pro-tumorigenic signaling cascades involved in cancer initiation and promotion. The novel findings of his research work have so far resulted in more than one 50 scientific publications in high impact factor peer reviewed journals and several international awards. He currently serves as an Academic Editor for prestigious PLOS ONE Journal and ad-hoc Reviewer for several other international journals.

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