

शुभिनी अ सराफ एम. फार्म, पीएचडी निदेशक Shubhini A Saraf M. Pharm, PhD Director



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Subject: Citation (summary) of outstanding research work of **Dr Prabhat Ranjan Mishra** for SUN Pharma Research Award-2024 nomination in Pharmaceutical Sciences.

Dr Mishra has significantly contributed and established a niche area of translational research and actively involved in developing bio-functionalized nano-biomaterials for therapeutics & diagnostic applications and elucidating the mechanistic aspect of its cellular translocation to achieve a higher therapeutic index with the low toxicity of drugs. Major contributions are summarized below:

Knowledge generation and development: In brief, he has discovered that breast cancer cell derived exosomes facilitates receptor mediated fusion & intracellular trafficking of quantum dots resulting spatial targeting to tumors at significantly lower dose (J. Controlled Rel 2024; I.F. 11.43) while acylated pyridoxine exerts proton sponge effect when tethered on nanoparticles that helps in escaping lysosomal degradation to facilitate intracellular drug release resulting fifteen-fold reduction in the therapeutic dose of doxorubicin with enhanced antitumor efficacy (ACS Appl. Mater. Interfaces 2016; I.F. 10.38). He elucidated key mechanism of Aggregation Induced Emission demonstrating image-guided chemotherapy showing enhanced antitumor efficacy using anisamide anchored lyotropic nano-liquid crystals (ACS Appl. Mater. Interfaces 2018; I.F. 10.38) while the tumor microenvironment was modulated using endogenous stimuli sensitive nano-systems through synergistic amplification of Mn⁺² with betulinic acid to achieve comprehensive cancer nano-theranostics without external stimulus (Acta Biomaterialia 2020; I.F. 10.63). He also demonstrated that lipid bilayercamouflaged mesoporous silica nanoparticles deliver metformin and topotecan in a synchronized manner that promotes apoptosis via mitochondrial membrane depolarization to achieve 100-fold higher drug concentration in tumors (Adv. Healthcare Mater 2018 I.F. 11.12) while redox-sensitive micellar system bearing aldose reductase inhibitor epalrestat and doxorubicin gets disrupted in tumor redox environment to promote synergistic tumor suppression (Biomater.Sci. 2019; IF 7.59). He elucidated that pegylated chondroitin and hyaluronate anchored taxanes expedite CD44 receptor-mediated endocytosis to exhibit higher tumor regression through the mitochondrial-lysosomotropic pathway (ACS Appl. Mater. Interfaces 2018; I.F. 10.38) while enhancing oral bioavailability through reversible P-gp modulation a with a significant reduction in tumor growth ensuing patient-friendly "chemotherapy at home" (Acta Biomaterialia 2015; I.F. 10.63). Over the years he has published more than 145 research papers (with h-index of 43 and citation of more than 6500) in peer reviewed international journals.

Translational contribution and commercialization: He successfully developed layer-by-layer (LBL) and SMEDDS technology that impacted product development in the area of bone-related disorders. He has been instrumental in patenting 27 technologies, out of which FIVE have been licensed to Industries while TWO are commercialized as Joint Fresh™ and Reunion™ [(Granted US patent 8,496,964; AU Patent; 2010217238A; EP patent 2400957 B1) Other products for the treatment of osteoarthritis and bone loss are also developed (Granted US Patent 10,596,115; AU Patent 2014291615; US patent 10265297)]. In addition, he has been actively involved in developing Umifenovir and its formulation under repurposing for COVID patients whose Phase III clinical trial has been completed.

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(Prof. Shubhini A. Saraf)