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Development of Polymeric Particles for Cancer Nanomedicine

Abstract: Controlled release technology is expected to have a profound impact in many medical fields including oncology. The incorporation of chemotherapeutic agents in nanoparticle delivery vehicles has improved drug solubility, reduced clearance, and enhanced therapeutic effectiveness. With controlled release nanoparticle systems, a single dose can sustain drug levels within the desired therapeutic range for long periods in various diseases. However, currently approved nanomedicines provide modest survival benefits for patients, in part because of poor tumor penetration. Physiological barriers, such as the dense interstitial collagen matrix, hinder the delivery of drugs throughout the entire tumor. In addition, many nanoparticulate systems often suffer from translational problem through bench to clinical trials, due in part to uncontrolled formulation (e.g. low drug loading) that leads to modest preclinical efficacy.

In this talk, I will present efforts that address some of those critical problems in nanomedicine. First, a controlled nanoparticulate formulation using drug-polymer conjugates will be introduced: multifunctional drugs are conjugated onto the polymer through one specific functional group with high drug loading up to 30 wt% for various drugs with controlled particle sizes. Secondly, I will present a photo-switching nanoparticulate system that uses light as the remote means of triggering both on-demand drug release and reversible changes in particle volume (from 130 to 40 nm) to enhance tissue penetration. These photo-switchable nanoparticles provide spatio-temporal control of drug release and enhanced tissue penetration, useful properties in many disease states including cancers. For the first time, the remotely light-triggered nanoparticulate drug delivery system shows preclinical benefits that enable repetitive dosing from a single administration with improved delivery efficacy throughout tumors.