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Related Download hscan smtp scanner » Smash Mouth · free download hscan smtp scanner. Related How to make a big pillow (2), free download hscan smtp scanner · Antibody-Drug Conjugates (ADCs) are chimeric molecules that combine the specificity of monoclonal antibodies with the therapeutic potential of cytotoxic agents. The most successful ADCs have structural similarity to the natural products maytansinoids. The maytansinoids are potent cytotoxins that are thought to enter cells via caveolae, clathrin-independent, lipid raft-containing vesicles. Once internalized, they appear to reach the lysosomal compartment after undergoing proteolytic cleavage within the acidic environment of the late endocytic pathway. The cytotoxicity of the maytansinoids is thought to result from the activation of a protease within the lysosome called lysosomal thiol-dependent cysteine

proteases or cathepsins. We hypothesize that the membrane transfer of a maytansinoid from early to late endosomes will provide a mechanistic basis for designing more effective ADCs that are targeting tumors that overexpress a particular antigen. Thus, Aim 1 of this proposal is to map the early events in the internalization and delivery of the maytansinoids. The goal of this aim is to determine what molecules or pathways are involved in the initial internalization of the maytansinoids, and we will use in vitro assays as well as in vivo imaging techniques to accomplish this goal. Aim 2