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*promising treatments of many brain diseases are often thwarted due to their inability to cross the blood brain barrier convection enhanced drug delivery ced uses direct infusion of drug containing solutions into tissue to circumvent the bloodbrain barrier the aim of this thesis was to combine models of transport in porous media with microfabricated devices to develop novel methods for controlling the distribution of infused drugs we used a poroelastic model of the brain to explore the effect of infusion induced dilation on transport we calculated that during infusions at flow rates greater than one microliter minute the effective pore size of the extracellular matrix was doubled by dilation from approximately fifty nanometers to one hundred nanometers a computational fluid dynamic model of the rat brain determined the perturbation of the flow field between white and gray matter we found no further perturbation of the flow field when the ratio of permeabilities between white and gray matter exceeded one hundred the results of our models suggest that the material properties of a targeted tissue region dictate the transport of infused solutions to better control and manipulate drug distribution we developed a novel microfluidic platform for delivering drug solutions at flow rates relevant for ced the microfluidic devices consisted of parylene channels with a cross section area of fifty microns by ten microns on a silicon structure with a cross sectional area of one hundred microns by one hundred microns these probes were tested in the normal rat brain and demonstrated performance advantages over standard needles including no channel occlusion and attenuation of backflow we expanded on the simple single channel device to implement more advanced strategies using multiple channels a two channel device was fabricated for infusing an enzyme or mannitol solution prior to infusing polystyrene nanoparticles by pre treating the targeted tissue region with enzymes or mannitol we increased the effective pore size of the extracellular matrix which resulted in a doubling of the distribution volume of nanoparticles abstract*

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