
Poor Sakura Vol 6 2 48

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in the search for novel cancer treatment options, the drug delivery approach has gained increasing attention, and nano-based drug delivery strategies have become increasingly popular. thus, the aim of the present study was to evaluate the potential of the hap-based formulation for delivery of poorly water-soluble drugs. the aim of the present study was to evaluate the potential of the hap-based formulation for delivery of poorly water-soluble drugs. the nanoparticles were designed to enhance the solubility and bioavailability of poorly water-soluble drugs and to selectively target the tumorous sites. this strategy has been termed 'site-specific drug delivery (ssd)' [3, 4, 5, 6, 7, 8, 9]. in an earlier study, we had demonstrated that the hap-based formulation can be used to enhance the solubility and bioavailability of the poorly water-soluble drug bezafibrate [15]. in the present study, however, although the bcs class iv drug az was uniformly coated with nano-hap particles, its solubility in dts 2 was not significantly improved. az in solution is reported to be only weakly acidic, with a pka of 7.2 [20], and this weak acidity might conceivably have precluded any positive buffering effect of hap. we therefore next conducted an experiment using the ussing chamber as a test for intracellular transport. in contrast to the in vitro single cell-culture caco-2 model, the ex vivo ussing method uses fresh intestinal epithelium consisting of various functional cells, offering more complex physiological interaction [66]. another advantage of this method is that in addition to a sample of intestinal epithelium containing principally villi, a sample containing principally peyers patch tissue can also be used [33, 67]. in the case of poorly water-soluble drugs, there is some debate about the possibility that drug solids not completely dissolved in the intestinal tract after oral ingestion may be absorbed as they are [5, 68, 69], with transport by intracellular pathways or the lymphatic system considered to be the most probable routes.

Poor Sakura Vol 6 2 48

they can also provide a solution for the poor solubility of the expensive, labile, and unstable pvp va64 particles, the pvp of which produces toxic insoluble particles in solution. this problem could be mitigated by adjusting the ph of the solution to a value near neutral, which can be achieved by using a buffer, such as hap. for example, although the solubility of pvp va64 in dts 2 solution was only 1.6%, the solubility was significantly increased to 2.5% by using a hap formulation [15]. the target of this hap formulation is poorly water-soluble drugs, namely bcs class ii and class iv drugs. in the case of bcs class ii drugs, which often show acidity, adjusting the ph of the drug in solution to near neutral is known to increase the solubility of the drug and therefore its absorption from the intestinal tract [10,11]. hap is known to function as a buffer in an acidic ph range [44]. in an earlier study, we succeeded in increasing the auc of the bcs class ii drug bezafibrate by using a hap formulation [15], and we postulated that this buffer function of hap may play a part in improving the drugs solubility. in the present study, however, although the bcs class iv drug az was uniformly coated with nano-hap particles, its solubility in dts 2 was not significantly improved. az in solution is reported to be only weakly acidic, with a pka of 7.2 [20], and this weak acidity might conceivably have precluded any positive buffering effect of hap. for drawing and sketching, the sakura koi brush pens are just okay. its weaknesses arent uncommon for the category. the 6-piece grey set is \$14.94 msrp, which comes out to \$2.49/marker, which is what blick has for the price of open stock. few other retailers have open stock available, though the sets are often available for cheaper than msrp, including at blick. i got mine for \$11.50 a while back, so just under \$2/marker. 5ec8ef588b

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