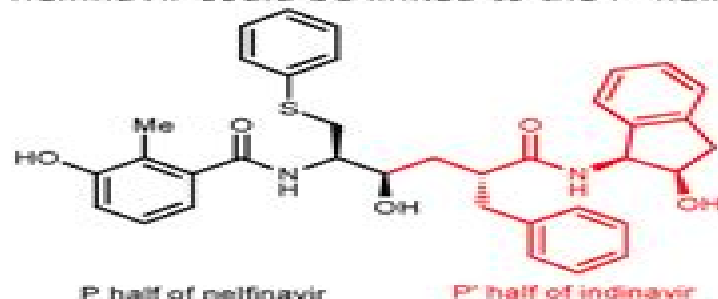
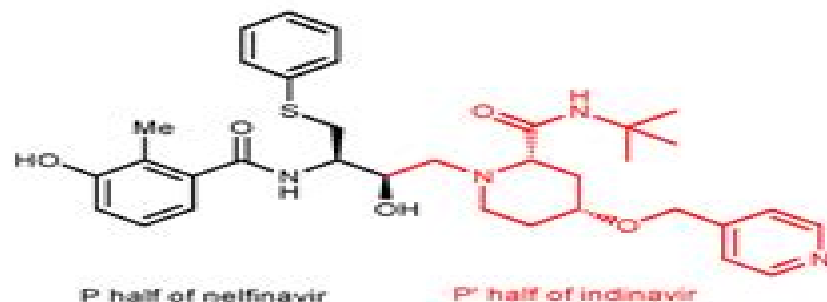


Answers to End of the Chapter Questions Chapter 20

1) There are various possibilities where one could link one half of a protease inhibitor to one half of a different inhibitor. There is no guarantee that they would be active, but they would be worth testing. For example, the P half of nelfinavir could be linked to the P' half of indinavir.



2) There are several possibilities where one could modify a known protease inhibitor with a suitably extended subsite ligand. Again, there is no guarantee that such an analogue would be effective. One possibility is to modify the P1' moiety of nelfinavir in a similar fashion to palinivir. This structure can also be seen as a hybrid structure consisting of the P half of nelfinavir and the P' half of palinivir.



3) Cidofovir is an analogue of deoxycytidine 5-monophosphate where the sugar and phosphate groups have been replaced by an acyclic group and a phosphonomethylene group respectively. The latter group acts as a bioisostere for the phosphate group and can be phosphorylated twice by cellular kinases to give the active compound. Since a phosphate bioisostere is present, the compound is active against viral strains lacking viral thymidine kinase. This enzyme normally catalyses the phosphorylation of antiviral

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