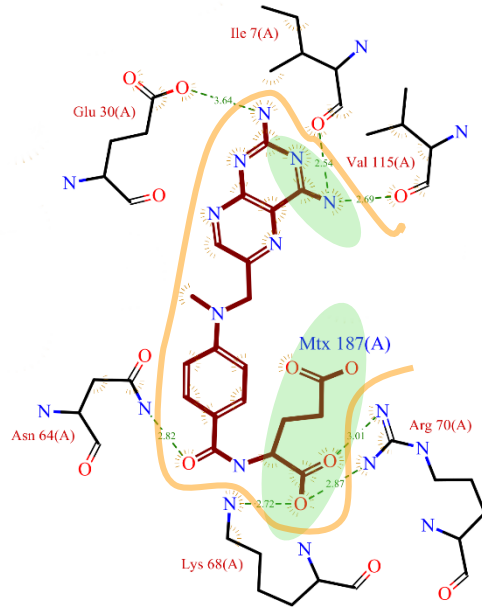
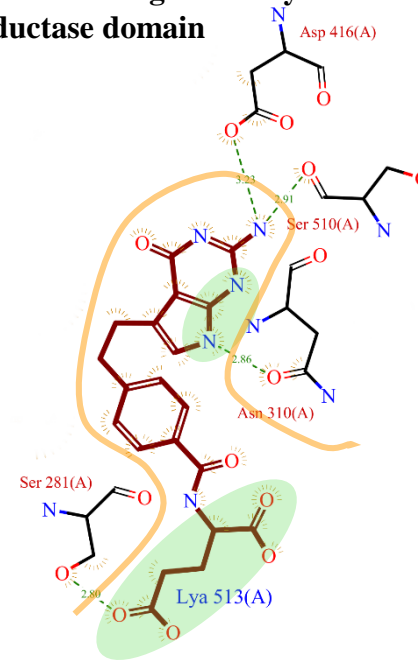


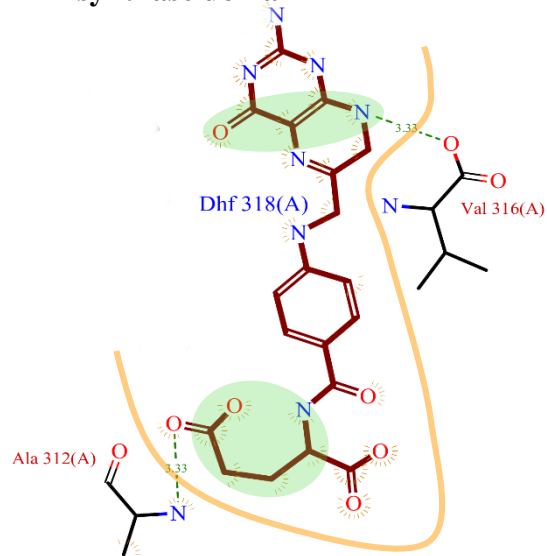
A. Methotrexate against dihydrofolate reductase domain



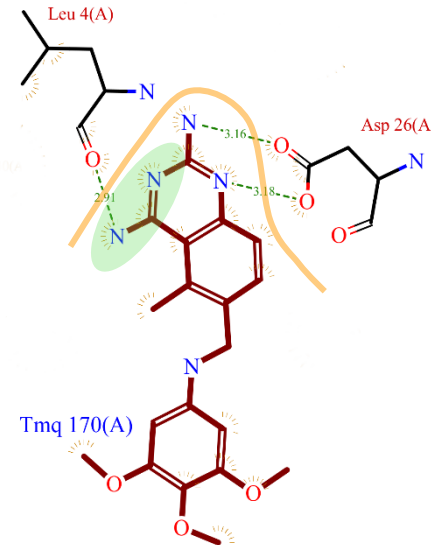
B. Pemetrexed against dihydrofolate reductase domain








C. DHF against thymidylate synthase domain



D. Trimetrexate against dihydrofolate reductase domain



-  Predicted substructures targeting the domains
-  Hydrogen bond and its length
-  Hydrophobic contact
-  Compound bond
-  Non-compound bond