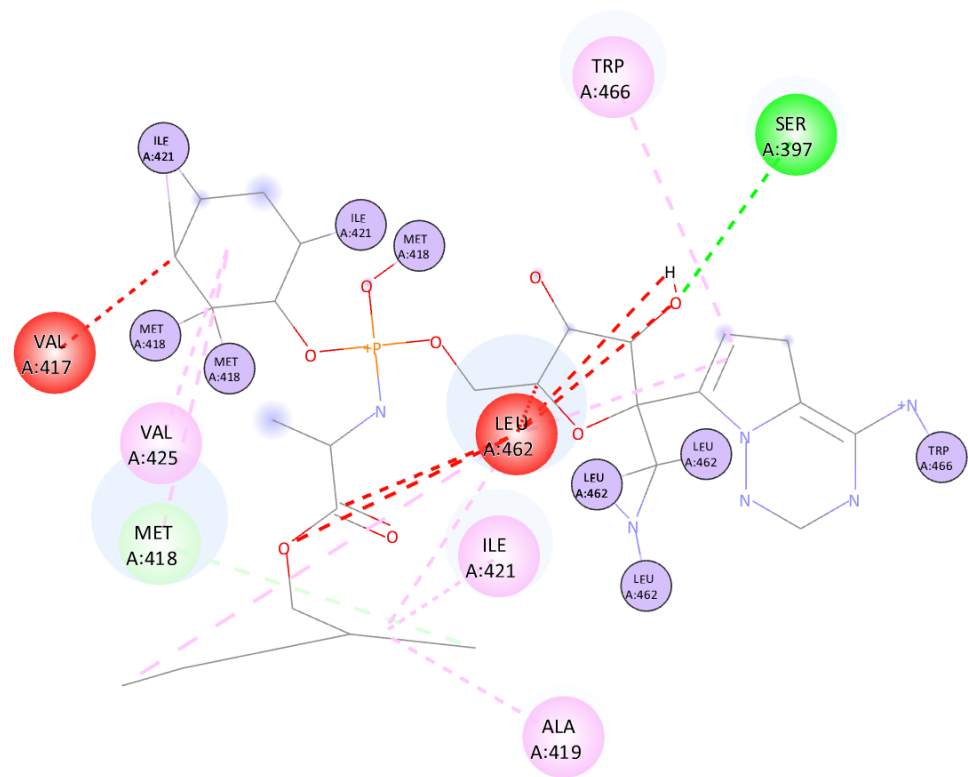
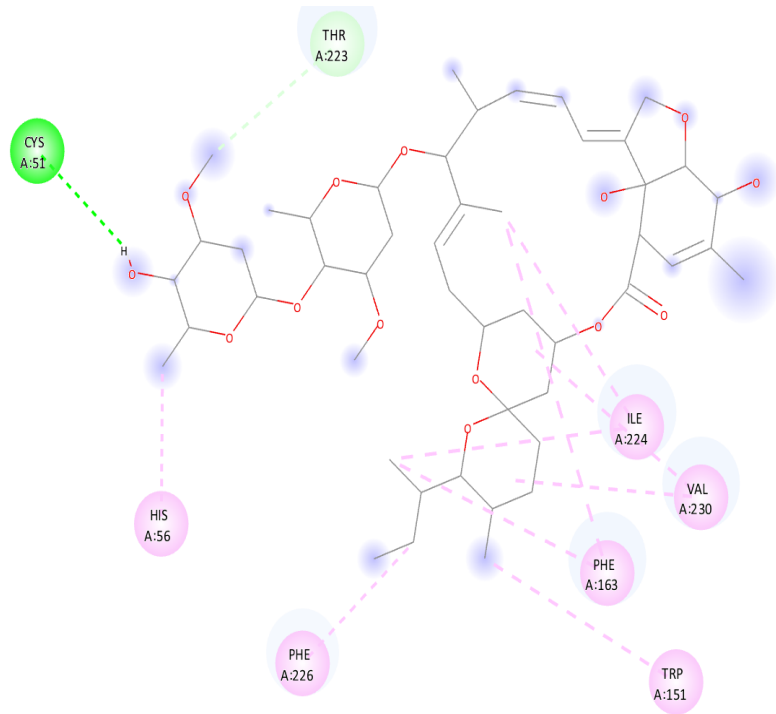


Wild type I120_Remdesivir (NSP1)

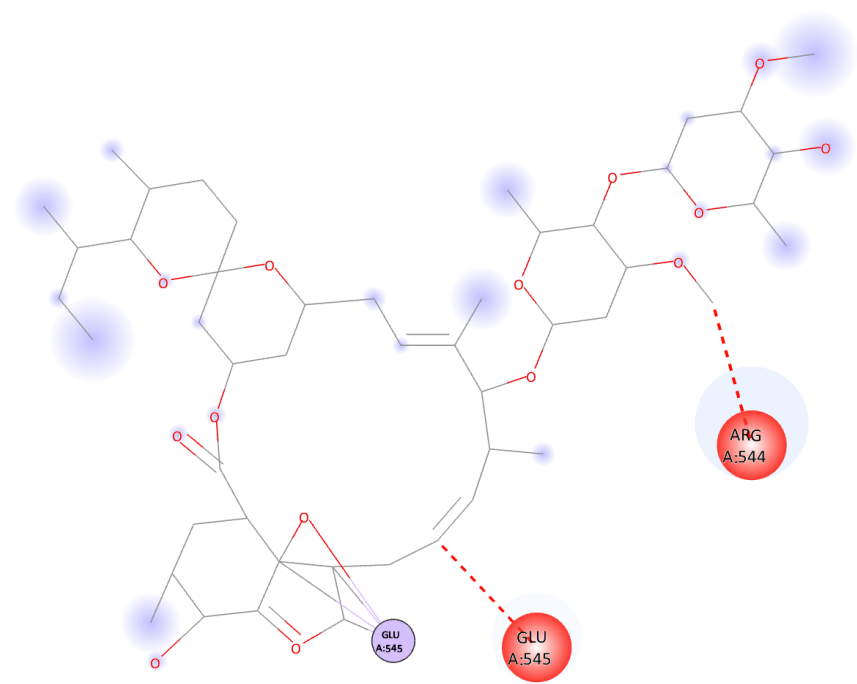


Mutant F120_Remdesivir (NSP1)

Supplementary Figure 1: Drug-binding interactions

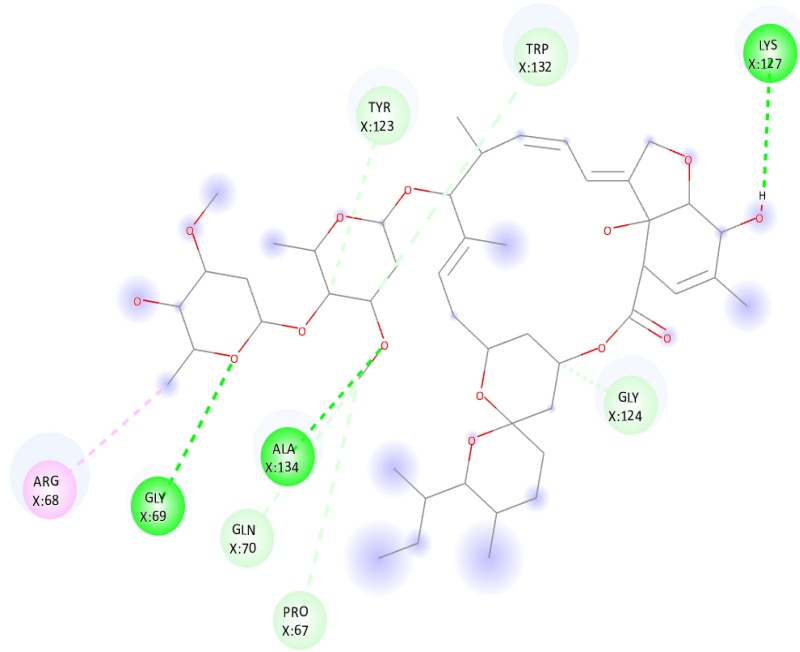


Wildtype I120_Ivermectin (NSP1)

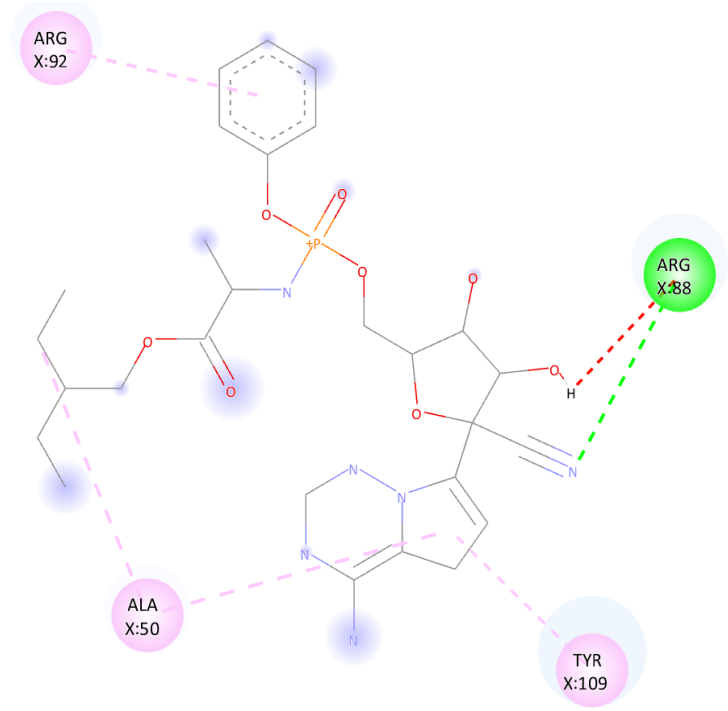


Mutant F120_Ivermectin(NSP1)

Supplementary Figure 1: Drug-binding interactions

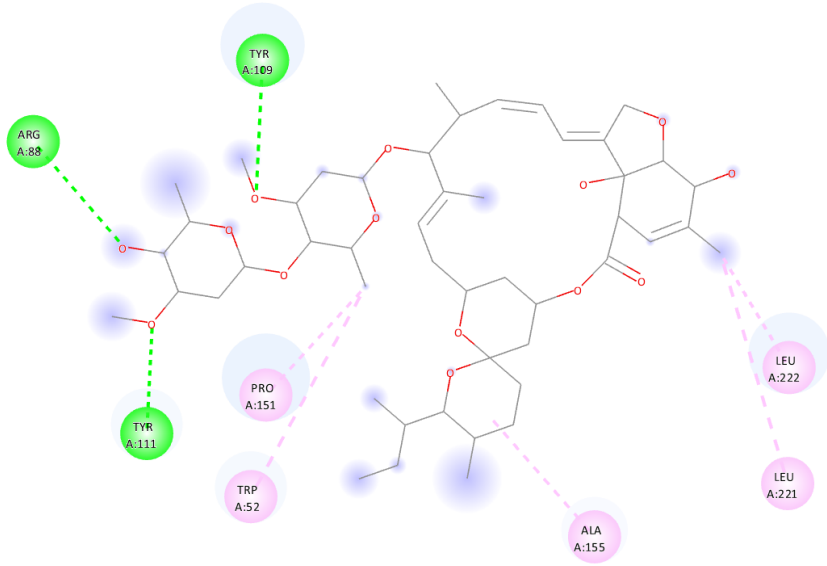


Wild type 202_203_204_Ivermectin (N)

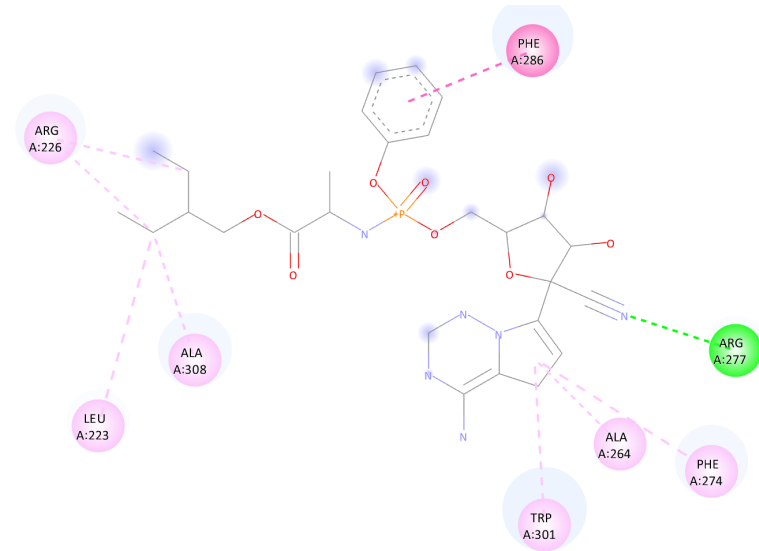


Wild type 202_203_204_Remdesivir (N)

Supplementary Figure 1: Drug-binding interactions

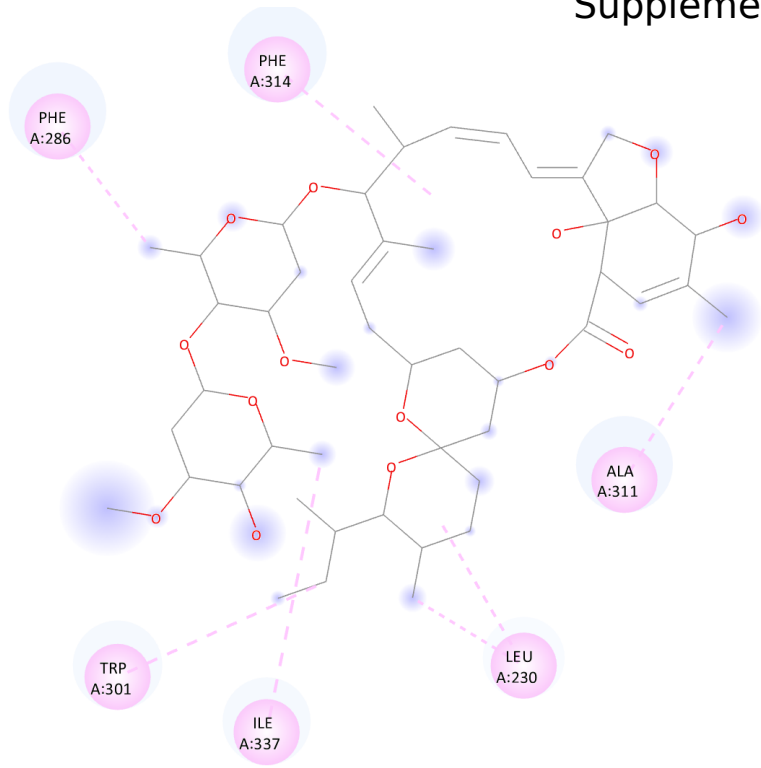


Mutant_203_Ivermectin (N)

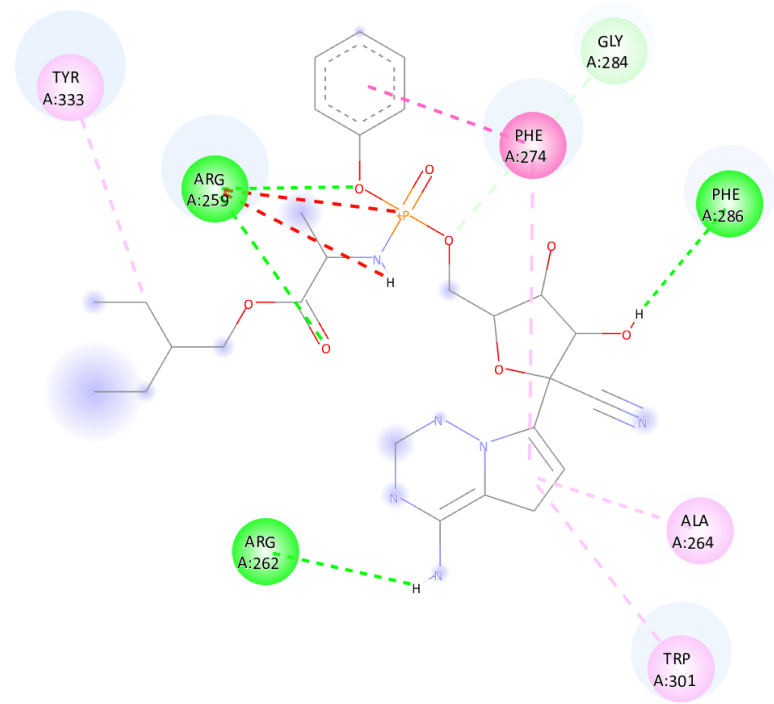


Mutant_203_Remdesivir (N)

Supplementary Figure 1: Drug-binding interactions

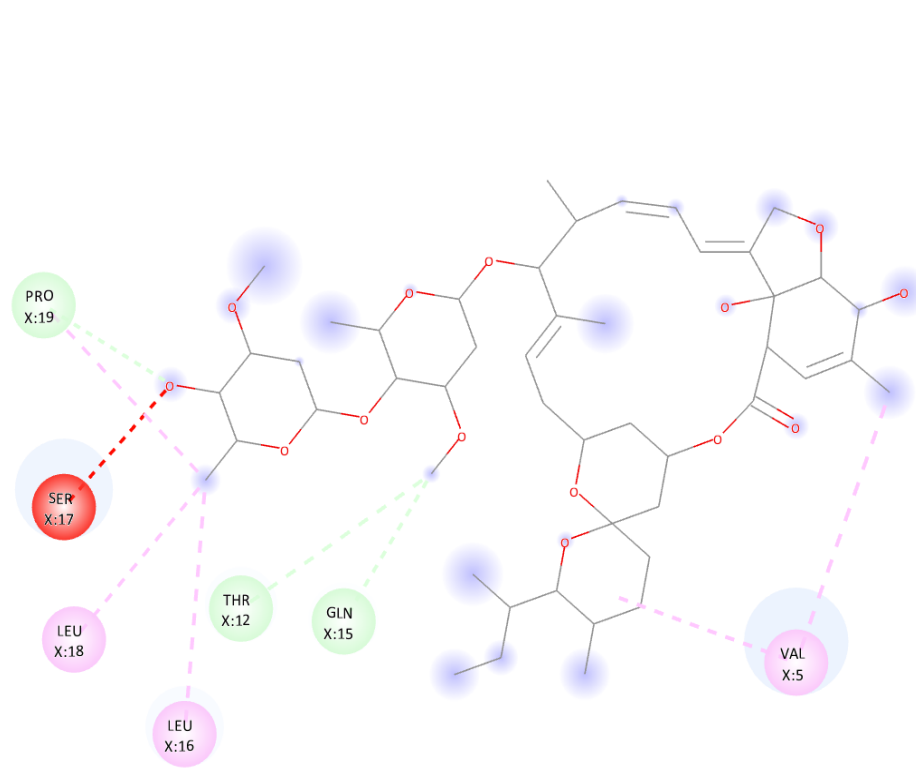


Mutant_204_Ivermectin (N)

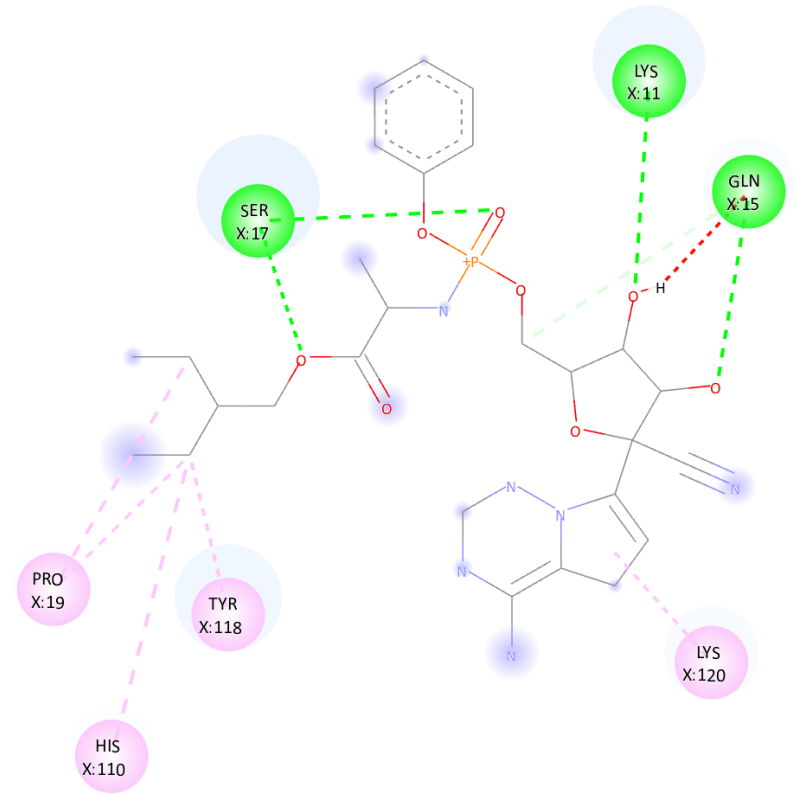


Mutant_204_Remdesivir (N)

Supplementary Figure 1: Drug-binding interactions

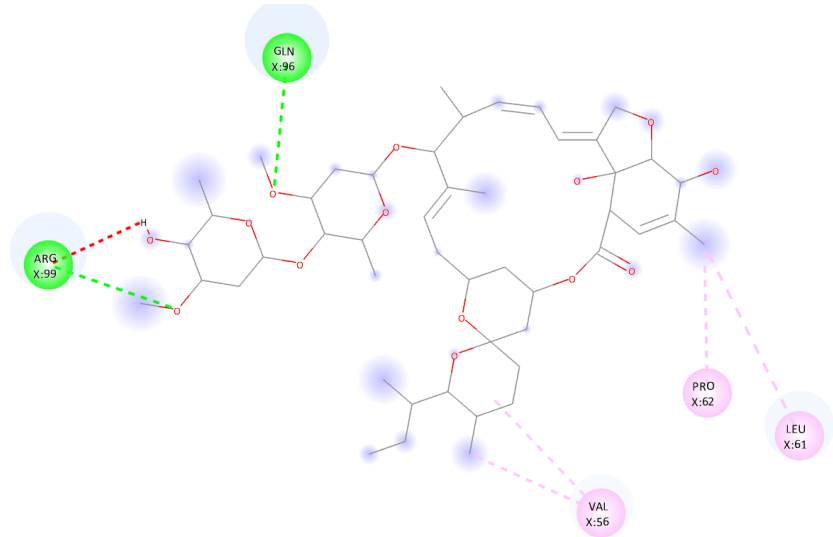


Wild type Valine121_Ivermectin (NSP1)

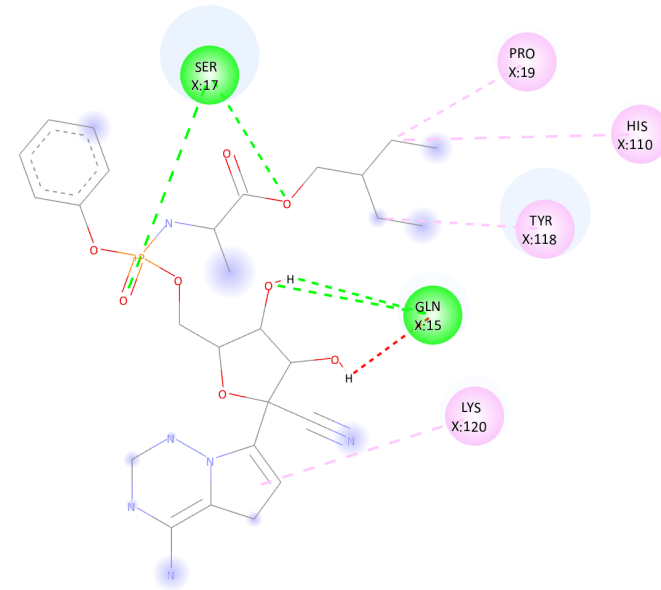


Wild type Valine_121_Remdesivir (NSP1)

Supplementary Figure 1: Drug-binding interactions

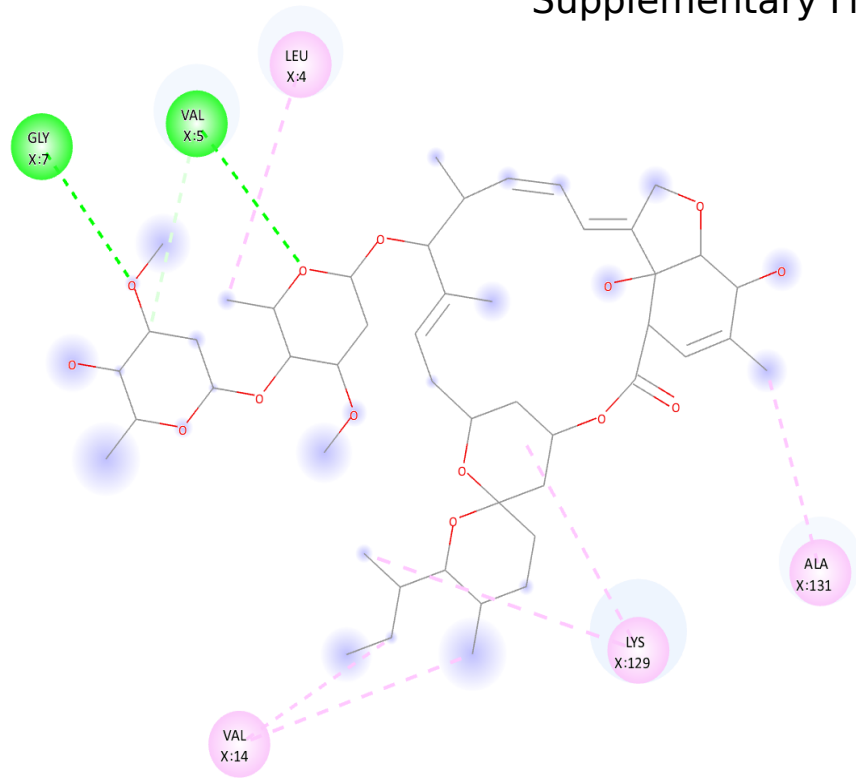


Wild type L122__Ivermectin (NSP1)

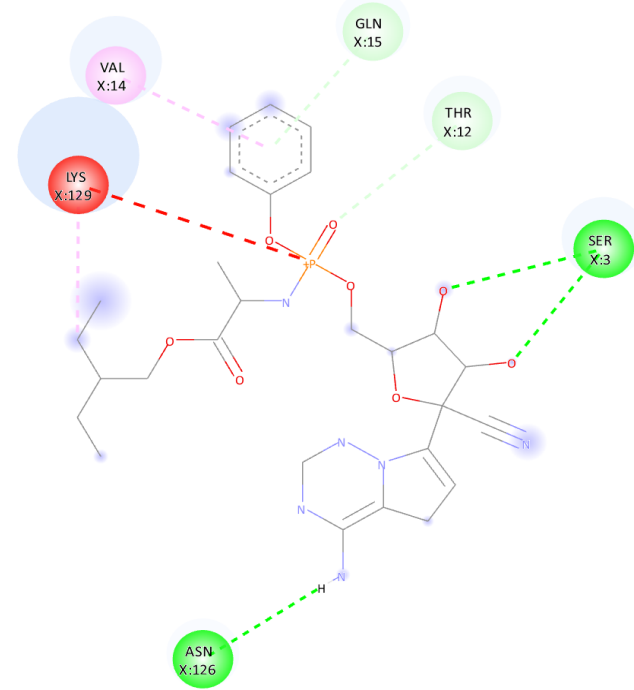


Wild type L122_Remdesivir (NSP1)

Supplementary Figure 1: Drug-binding interactions

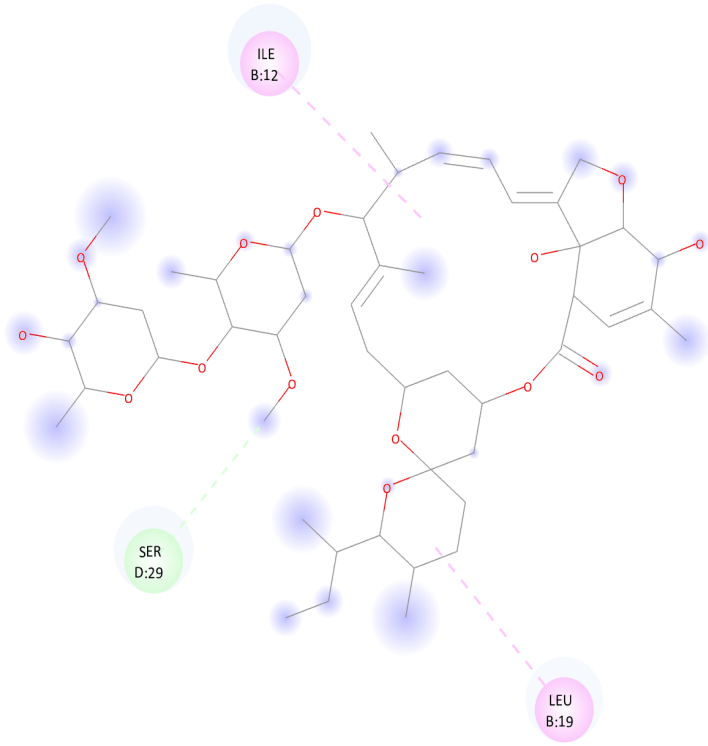


Mutant Ile122_Ivermectin (NSP1)

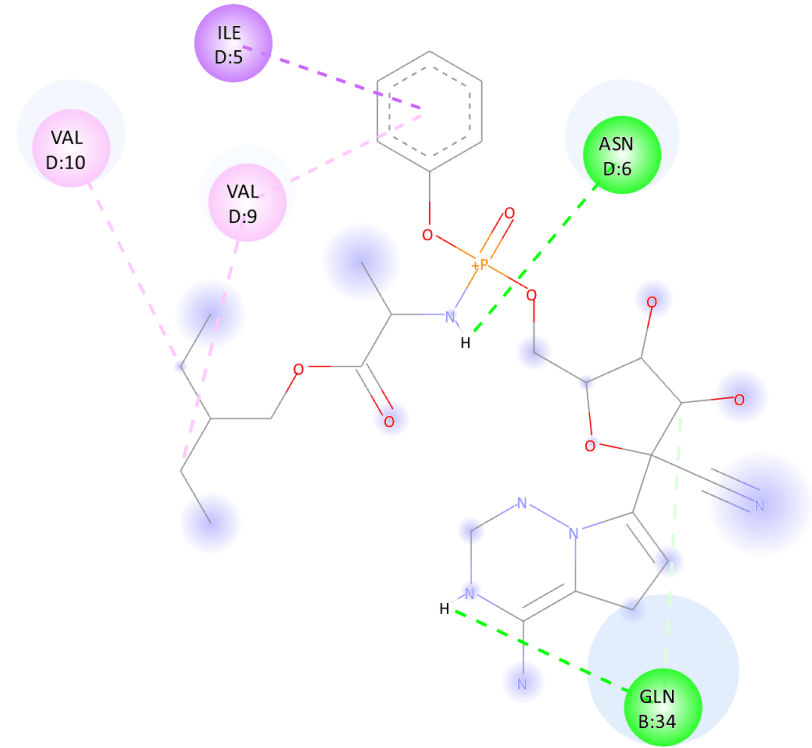


Mutant Ile122_Remdesivir (NSP1)

Supplementary Figure 1: Drug-binding interactions

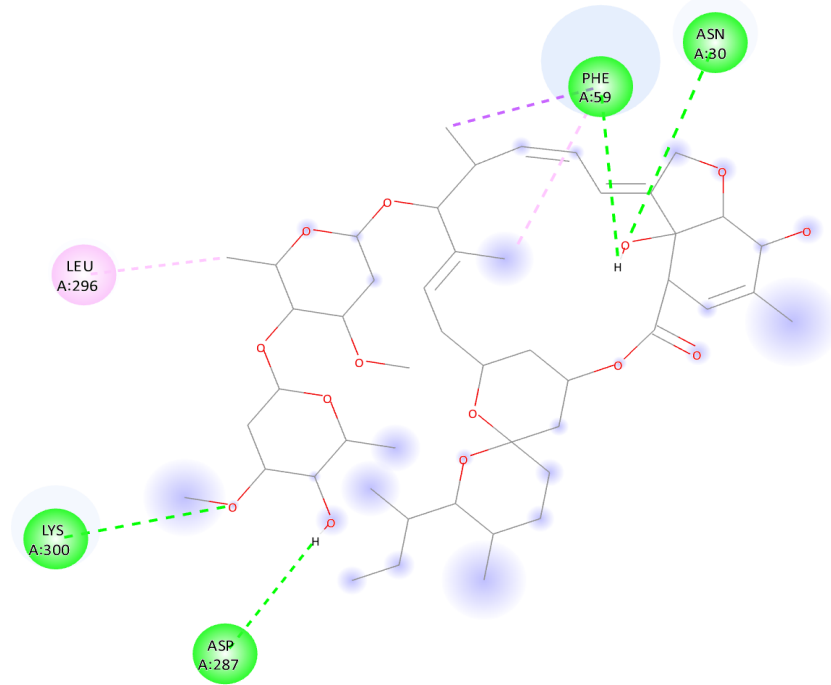


Wild ASP614_Ivermectin (S)

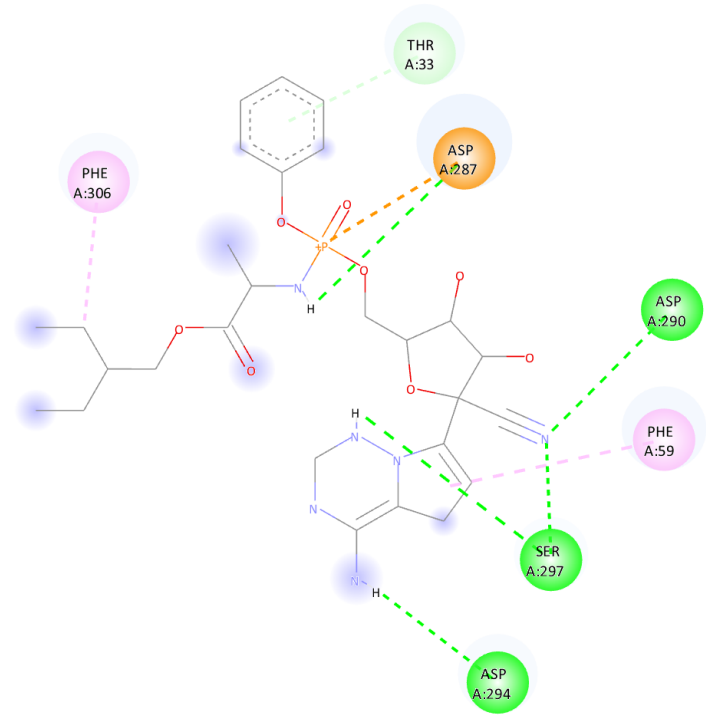


Wild ASP614_Remdesivir (S)

Supplementary Figure 1: Drug-binding interactions

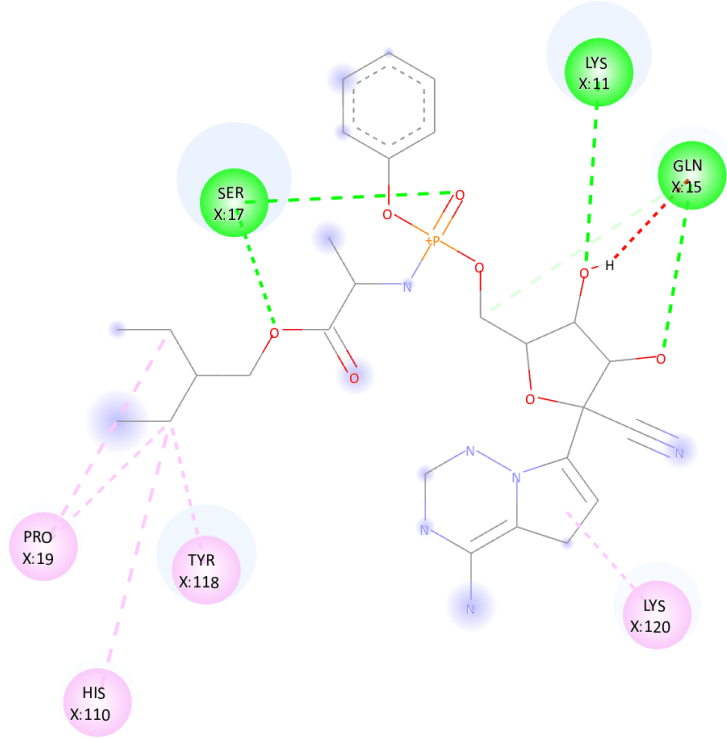


Mutant D614G_Ivermectin (S)

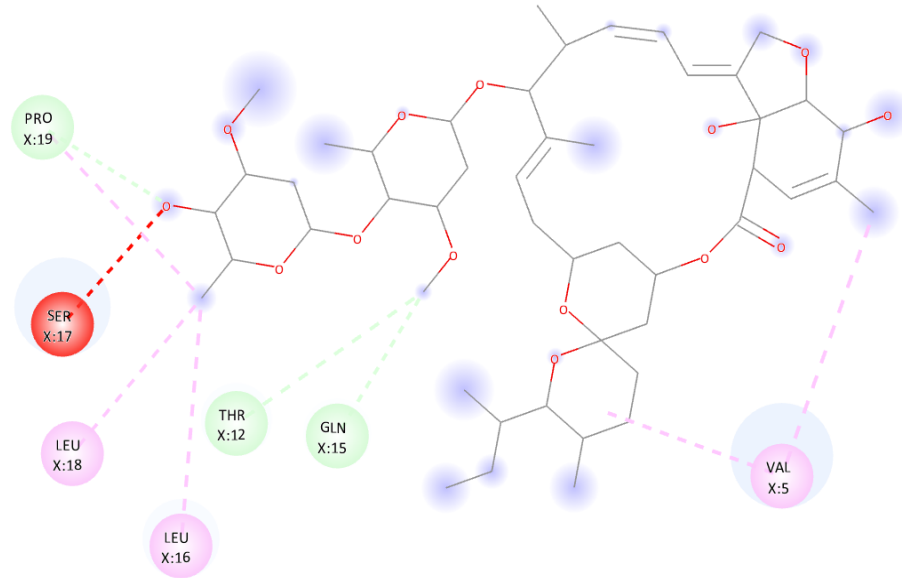


Mutant D614G_Remdesivir (S)

Supplementary Figure 1: Drug-binding interactions

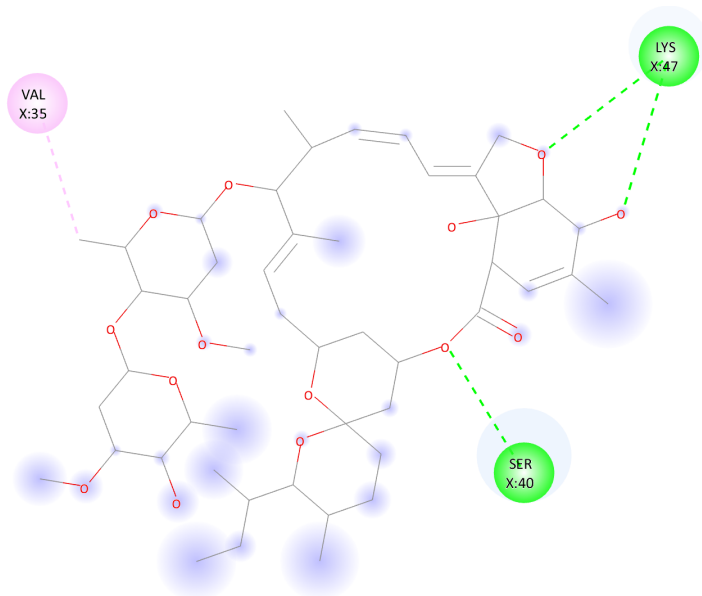


Wild type V_121_Remdesivir (NSP1)

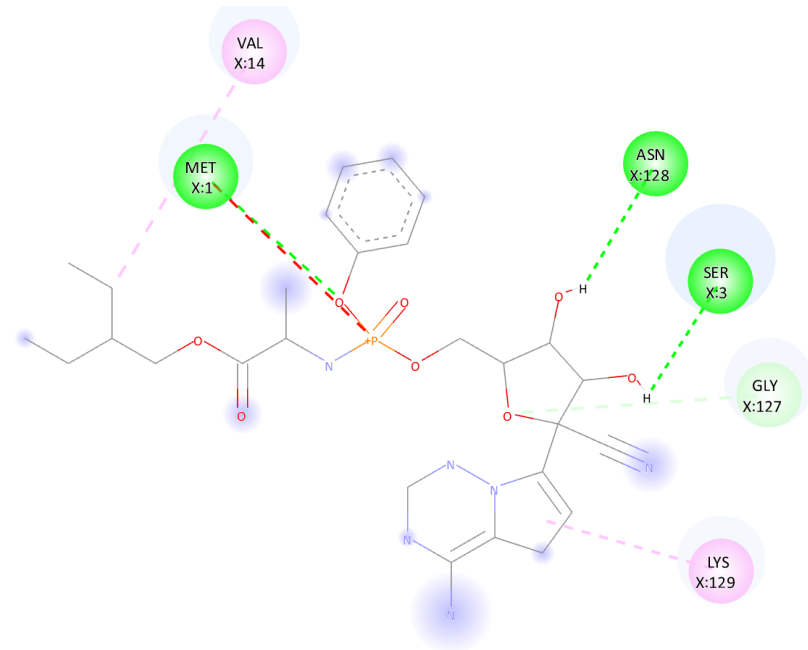


Wild type V_121_Ivermectin (NSP1)

Supplementary Figure 1: Drug-binding interactions

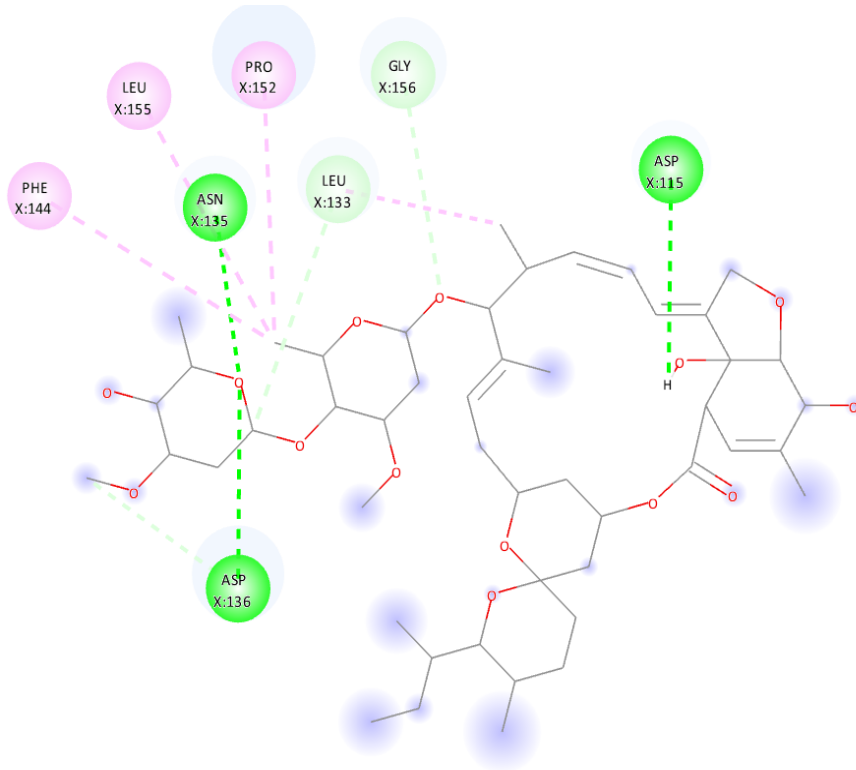


Mutant D_121_Ivermectin (NSP1)

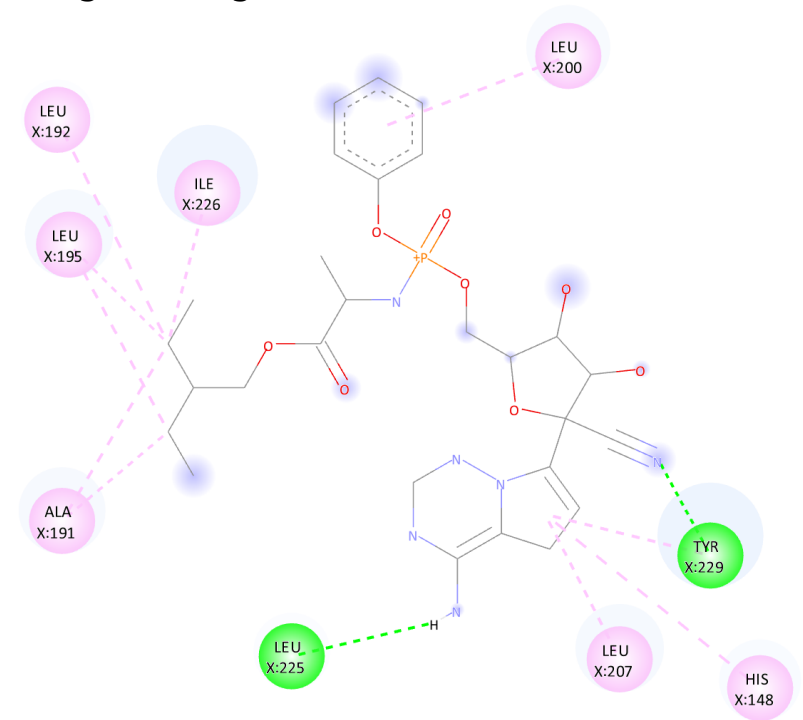


Mutant D_121_Remdesivir (NSP1)

Supplementary Figure 1: Drug-binding interactions

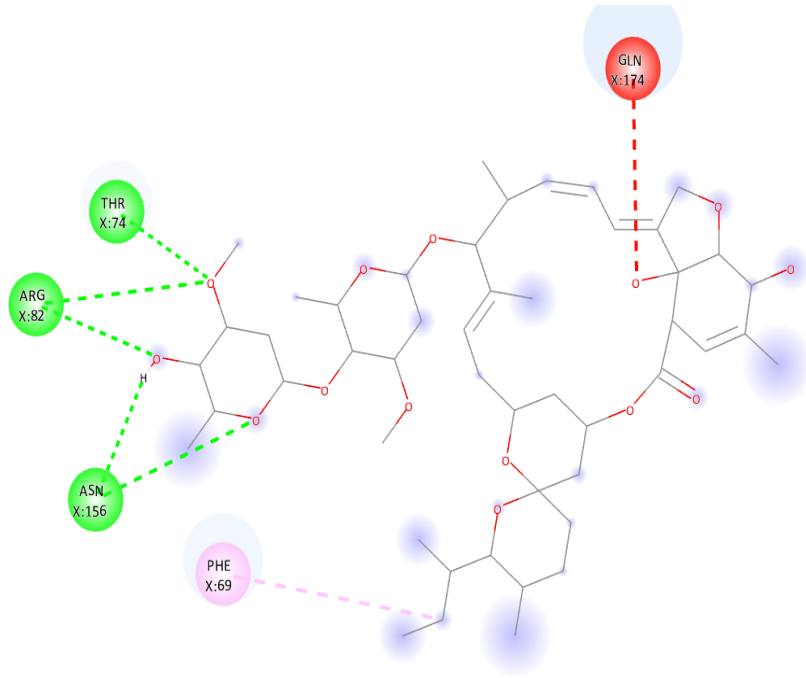


Wild type_843, 889_Ivermectin (NSP3)

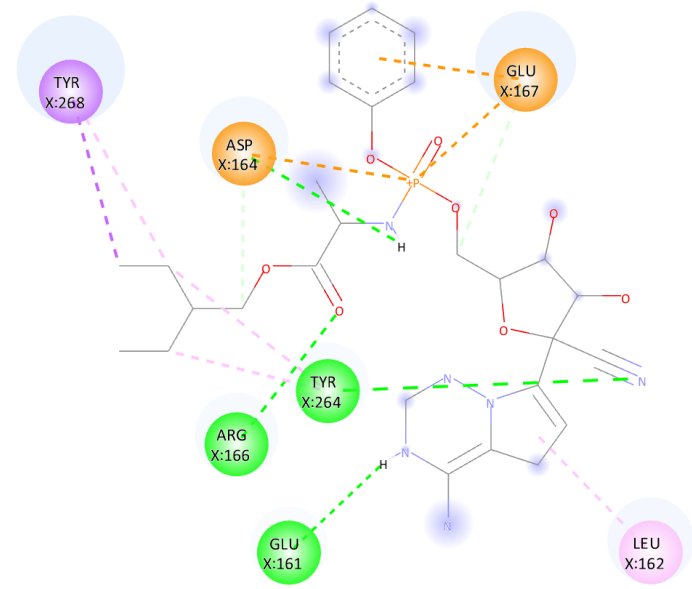


Wild type_843, 889_Remdesivir (NSP3)

Supplementary Figure 1: Drug-binding interactions

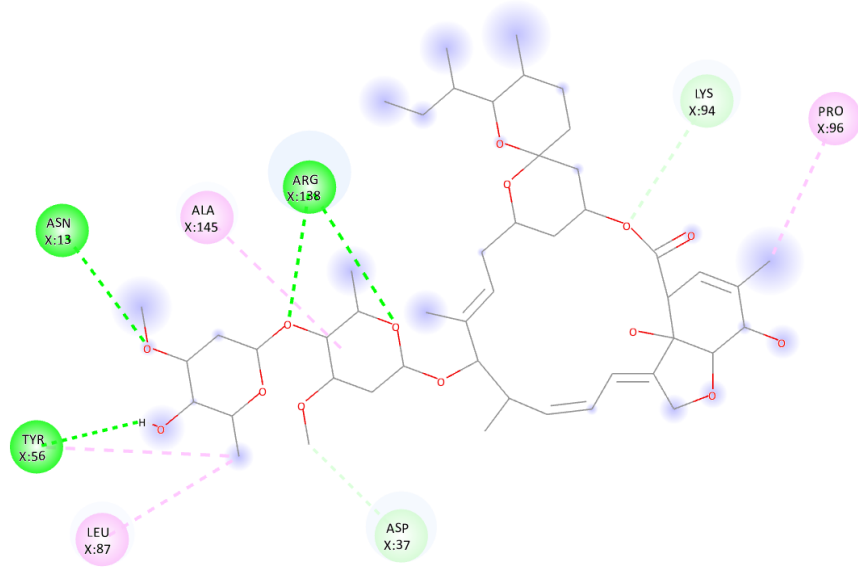


Mutant V843F_Ivermectin (NSP3)

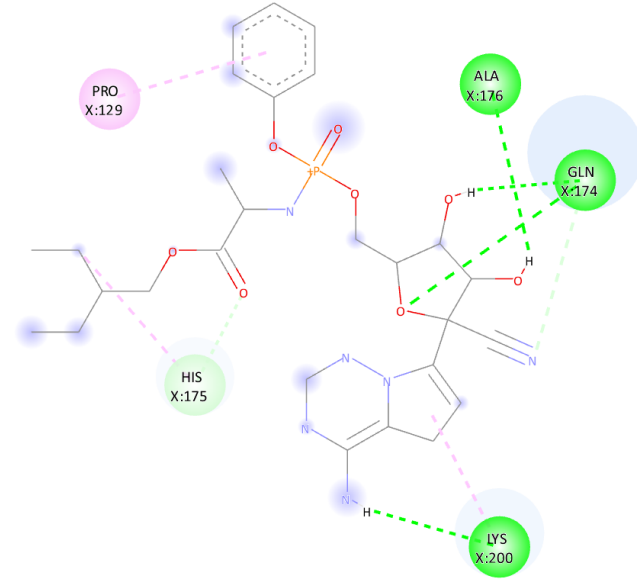


Mutant V843F_Remdesivir (NSP3)

Supplementary Figure 1: Drug-binding interactions



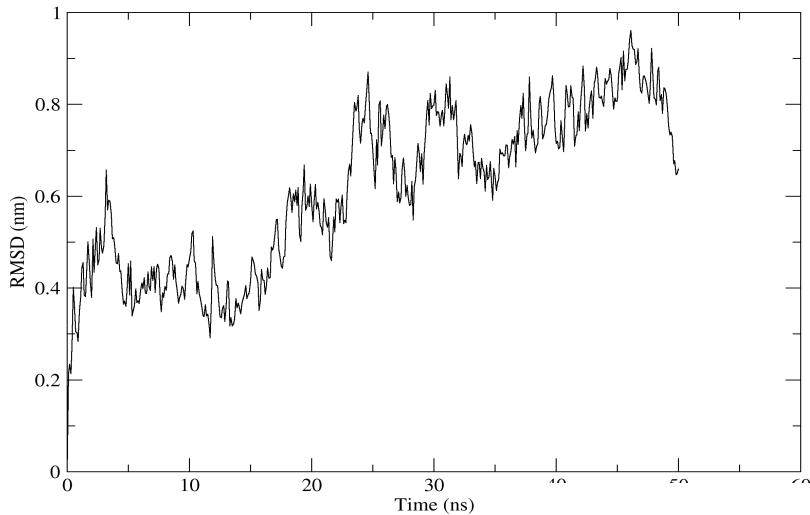
Mutant type A889V Ivermectin (NSP3)



Mutant type A889V Remdesivir (NSP3)

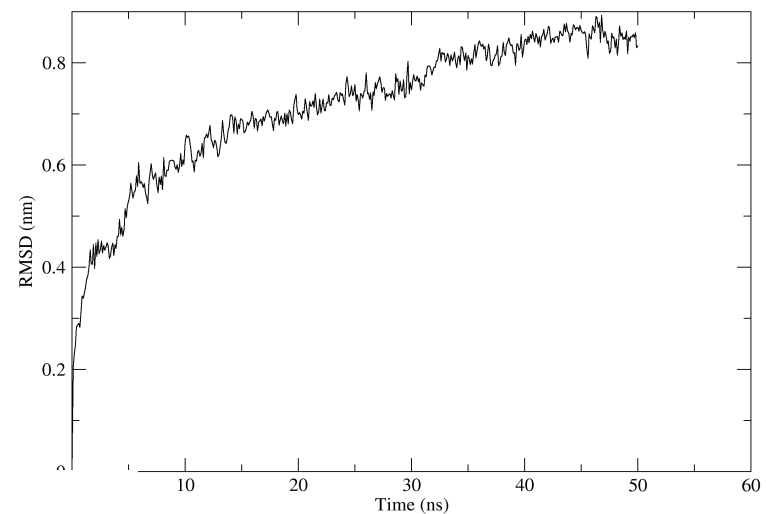
Supplementary Figure 2: Results of Molecular Dynamic

RMSD Simulation
Backbone after lsq fit to Backbone



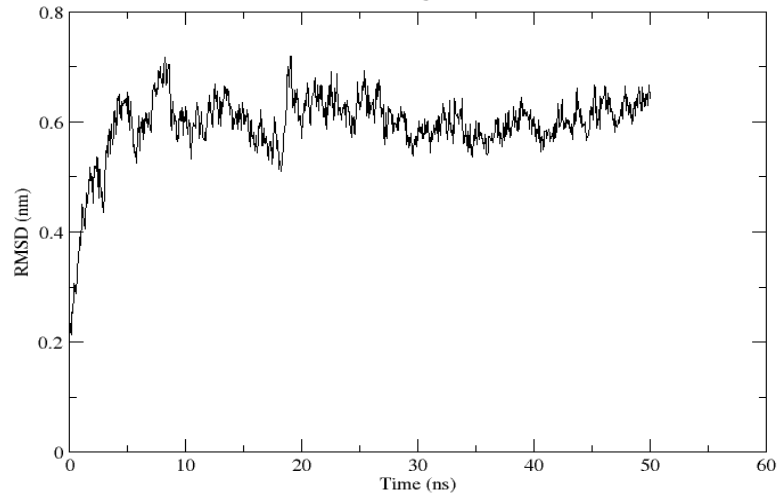
NSP3 wild

RMSD
Backbone after lsq fit to Backbone



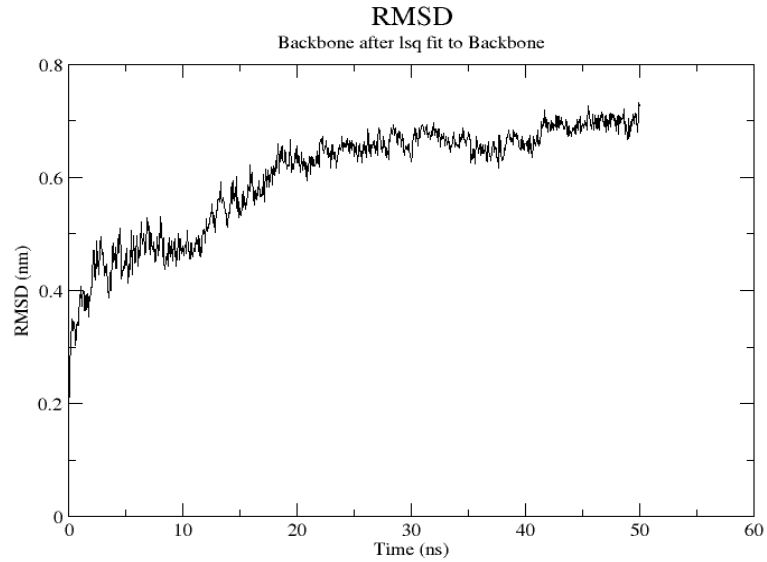
NSP3 V843F

RMSD
Backbone after lsq fit to Backbone

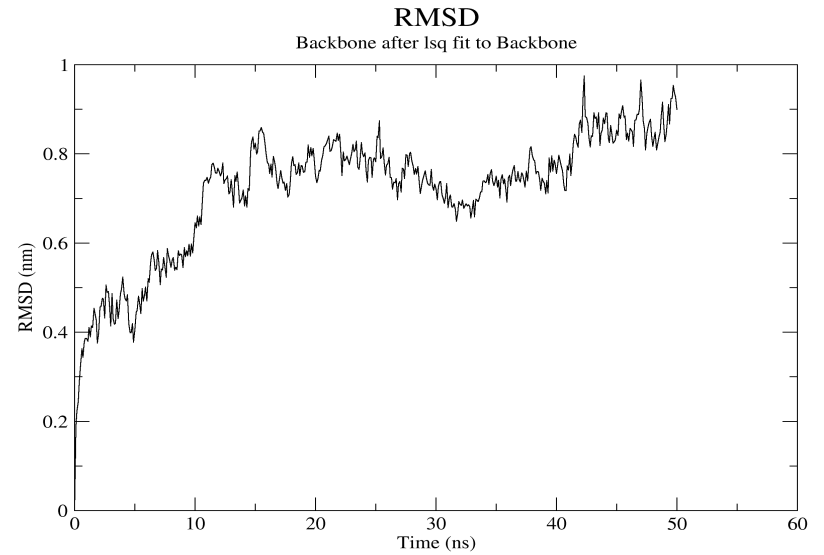


NSP3 A889V

Supplementary Figure 2: Results of Molecular Dynamic Simulation



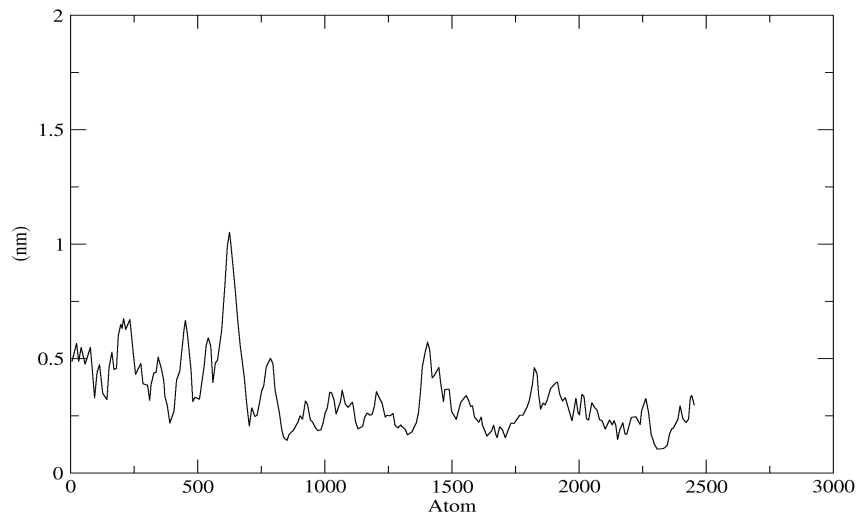
NSP1 wild



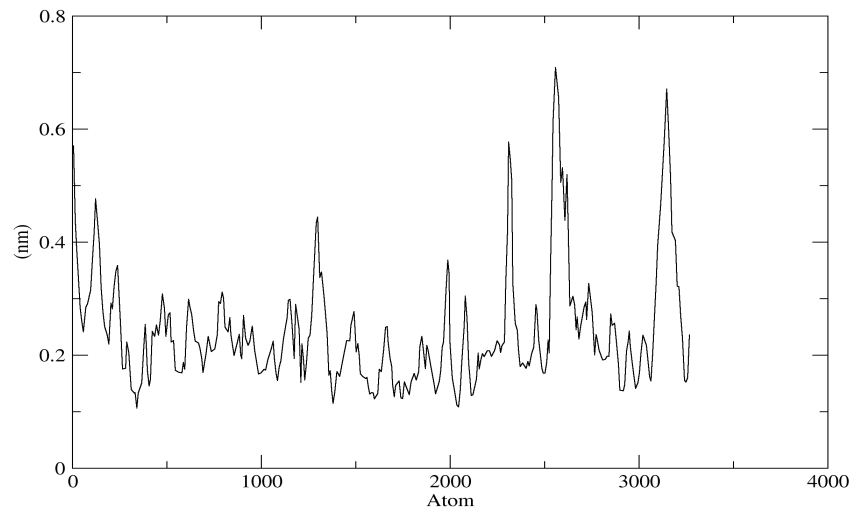
NSP1 V121D

Supplementary Figure 2: Results of Molecular Dynamic

RMS fluctuation

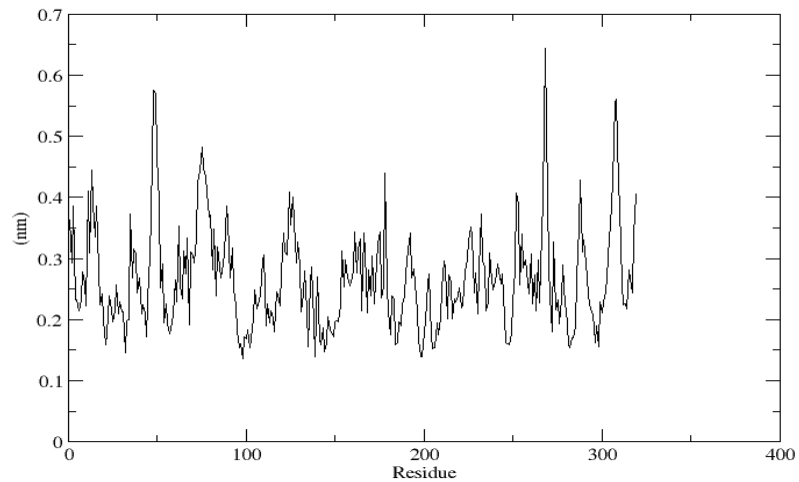


RMS fluctuation



NSP3 wild

RMS fluctuation

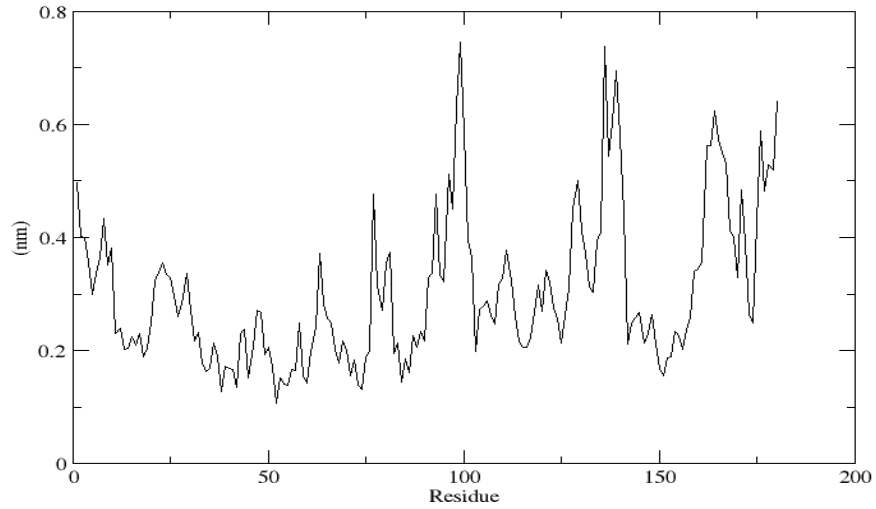


NSP3 V843F

NSP3 A889V

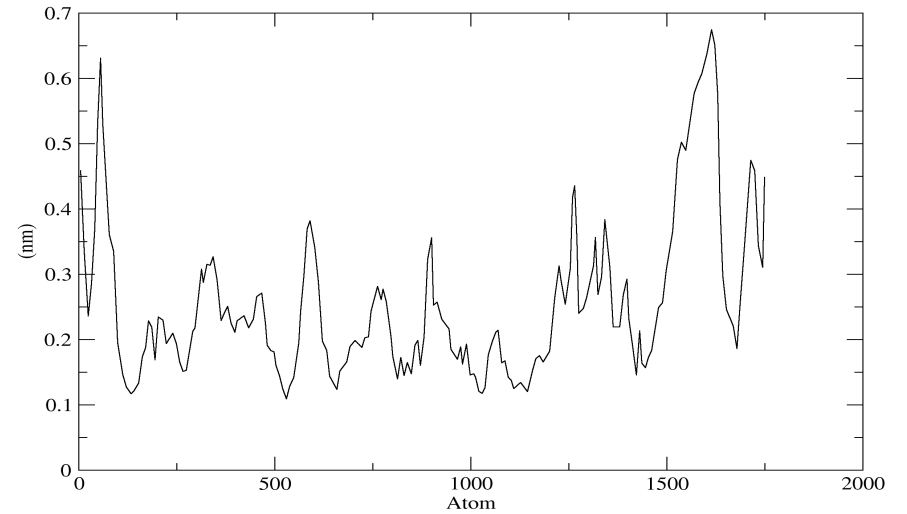
Supplementary Figure 2: Results of Molecular Dynamic Simulation

RMS fluctuation



NSP1 wild

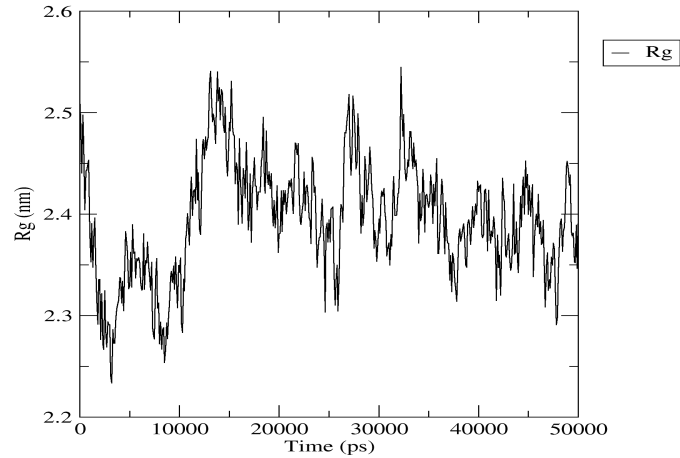
RMS fluctuation



NSP1 V121D

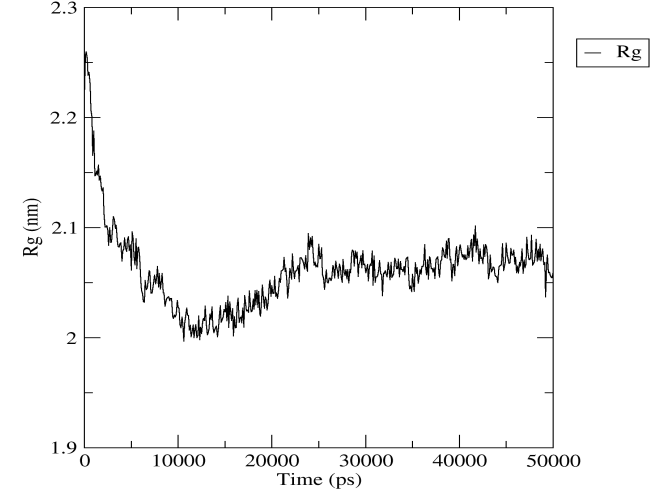
Supplementary Figure 2: Results of Molecular Dynamic Simulation

Radius of gyration (total and around axes)



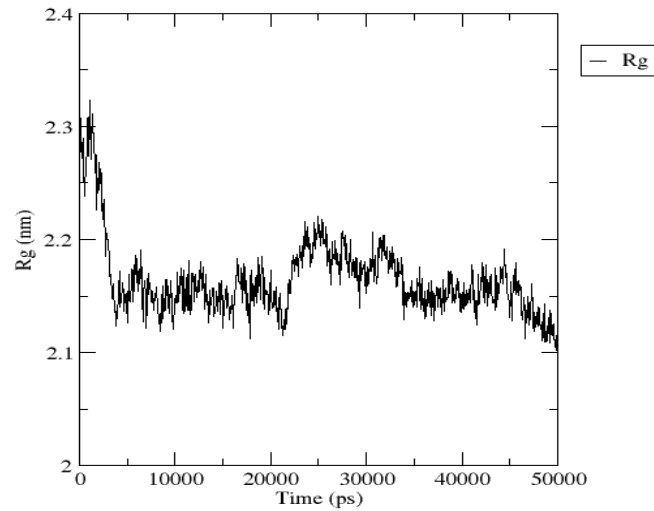
NSP3 wild

Radius of gyration (total and around axes)



NSP3 V843F

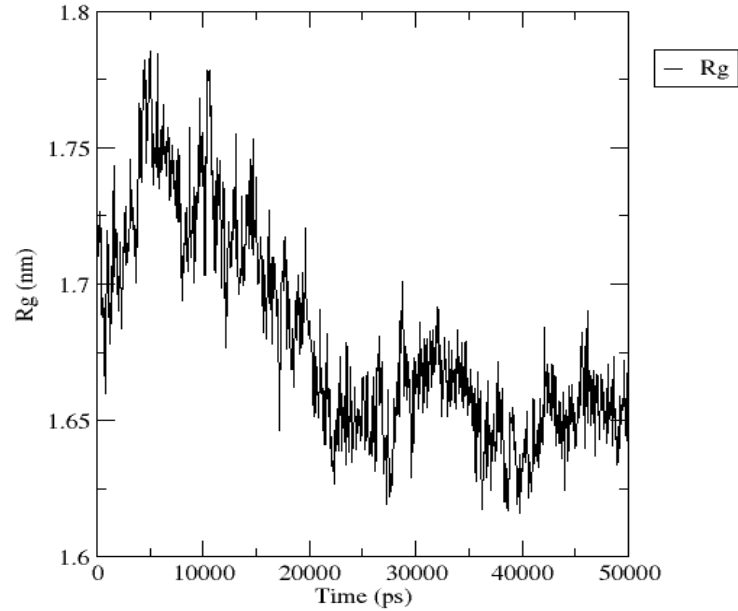
f gyration (total and around axes)



NSP3 A889V

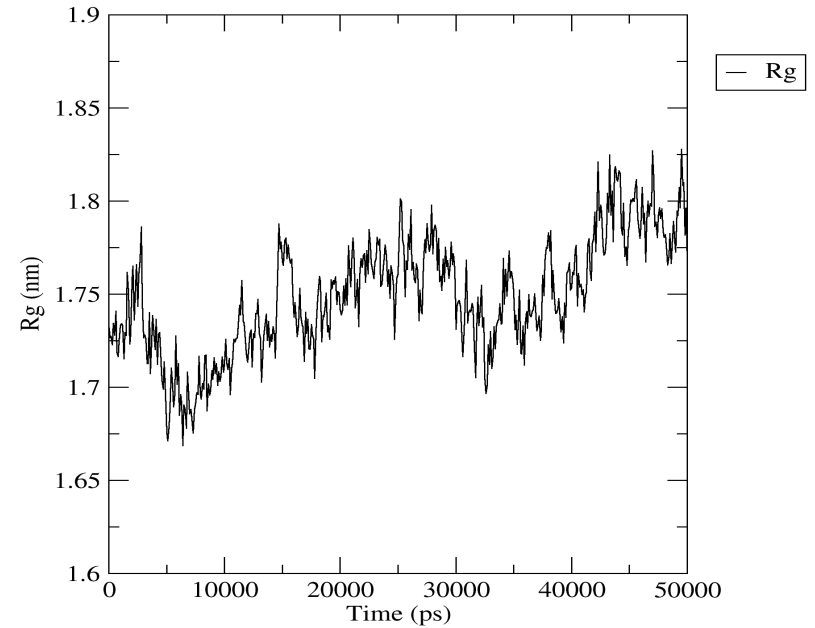
Supplementary Figure 2: Results of Molecular Dynamic Simulation

Radius of gyration (total and around axes)



NSP1 wild V121

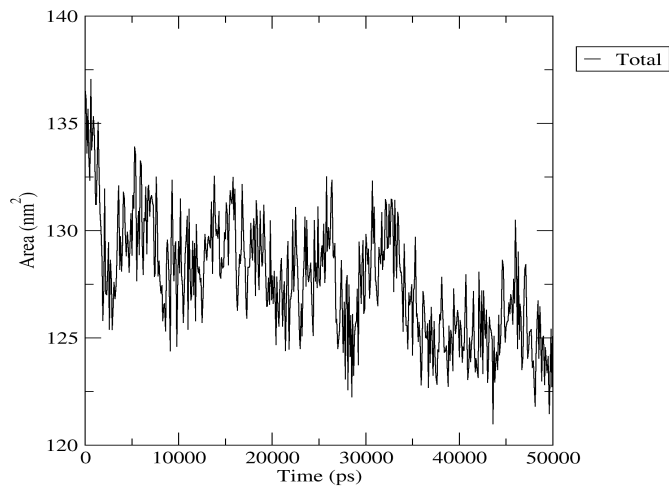
Radius of gyration (total and around axes)



NSP1 V121D

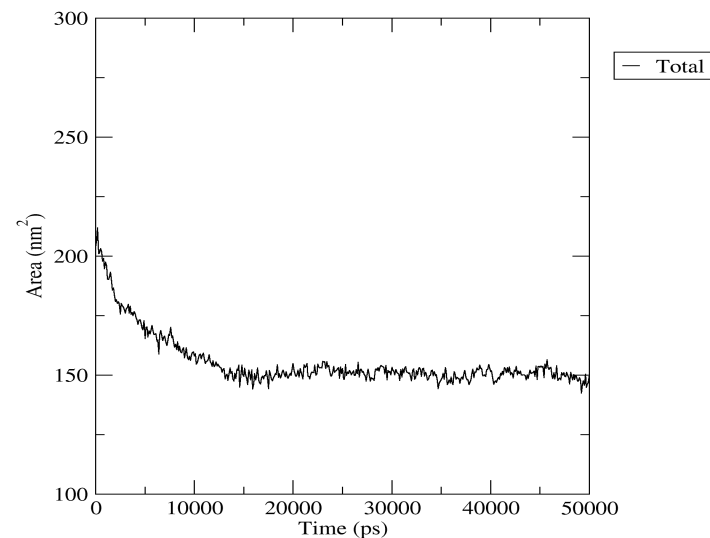
Supplementary Figure 2: Results of Molecular Dynamic Simulation

Solvent Accessible Surface



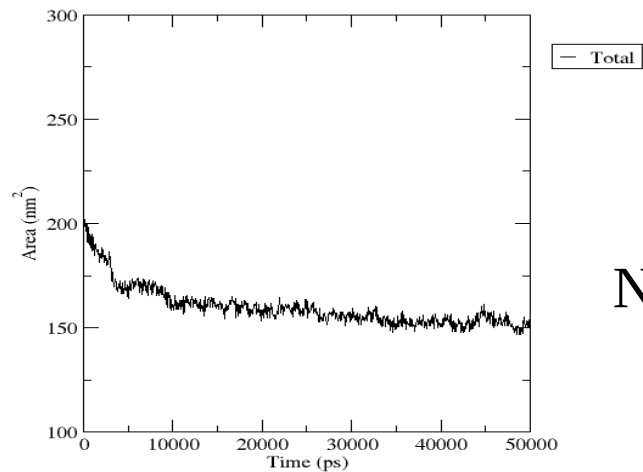
NSP3 wild

Solvent Accessible Surface



NSP3 V843F

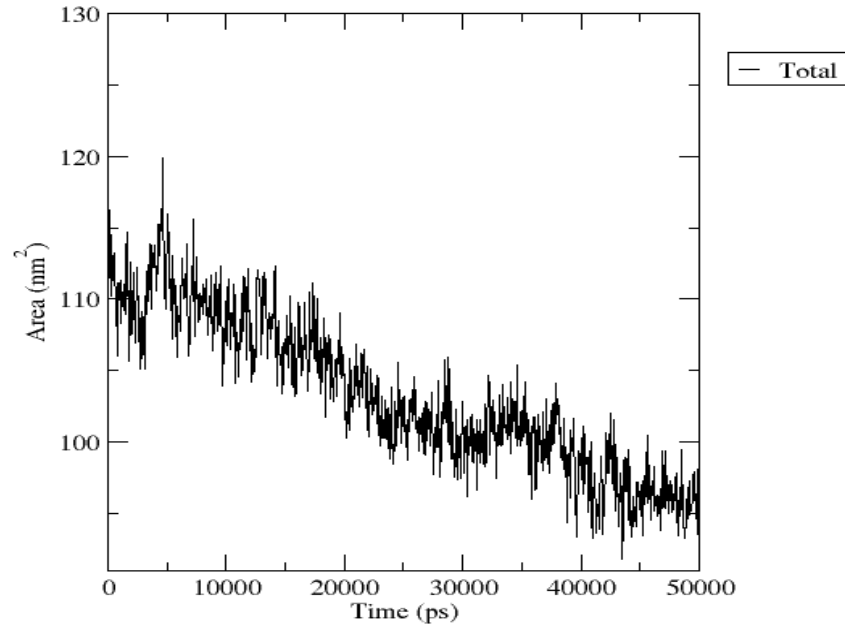
Solvent Accessible Surface



NSP3 A889V

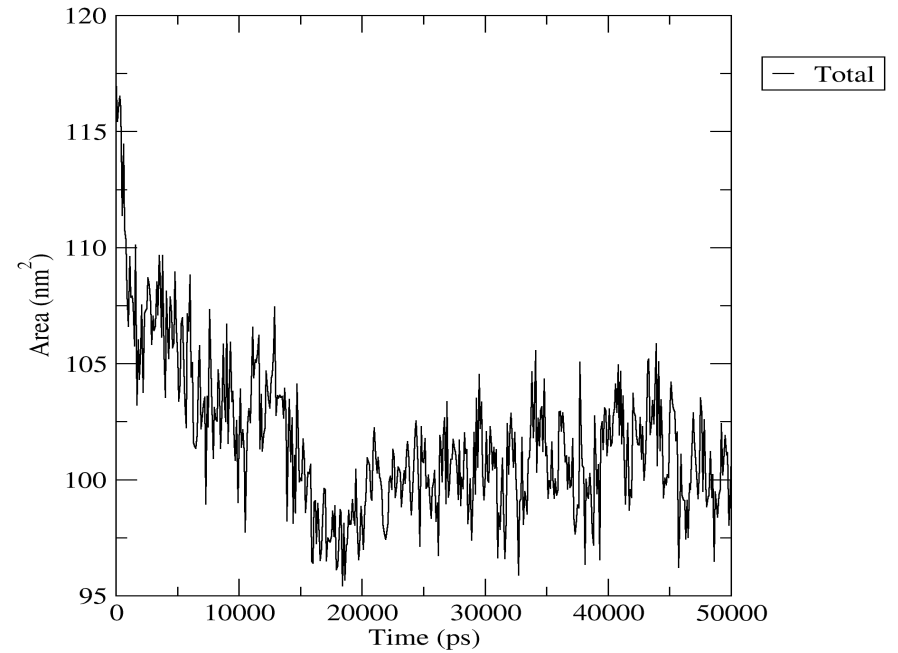
Supplementary Figure 2: Results of Molecular Dynamic Simulation

Solvent Accessible Surface



NSP1 Wild V121

Solvent Accessible Surface



NSP1 V121D