

Silico analysis of Impact of SARS-CoV-2 Variants

to Flowflex SARS-CoV-2 Antigen Rapid Test

The product *Flowflex* SARS-CoV-2 Antigen Rapid Test was designed to detect the nucleocapsid protein antigen from SARS-CoV-2 in human nasal and nasopharyngeal swab specimens. The current variants of concern and corresponding changes in known antigen targets have several mutations in the spike protein and a few mutations in the nucleocapsid protein. The mutation on the nucleocapsid protein for each variant is listed below:

| No. | Variant designation | Mutation on the nucleocapsid protein |
|-----|-------------------------------|--------------------------------------|
| 1 | Alpha (B.1.1.7) | D3L, S235F |
| 2 | Beta (B.1.351) / Mu (B.1.621) | T205I |
| 3 | Gamma (P.1) | P80R |
| 4 | B.1.617 | R203K+D377Y |
| 5 | Kappa (B.1.617.1) | R203M+D377Y |
| 6 | Delta (B.1.617.2) | D63G+D377Y+R203M |
| 7 | B.1.617.3 | P67S+D377Y+R203M |
| 8 | B.1.618 | A119S, A217S, E367Q, G18S, M234I |
| 9 | Epsilon (B.1.427/B.1.429) | T205I |
| 10 | Zeta (P.2) | A119S+M234I+RG203KR |
| 11 | Eta (B.1.525) | A12G+T205I |
| 12 | Theta (P.3) | RG203KR |

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| 13 | Iota (B.1.526) | M234I+P199L |
| 14 | B.1.616 | T325I |
| 15 | A.23.1 | S202N |
| 16 | Lambda (C.37) | P13L+R203K+G204R+G214C |
| 17 | Delta Plus | D63G, R203M, G215C, D377Y |
| 18 | Omicron (B1.1.529) | P13L, E31-, R32-,S33-, R203K, G204R |

The antibodies used in Flowflex SARS-CoV-2 Antigen Rapid Test is target to the region of 209 – 232aa of the nucleocapsid protein. There is no any mutation in this range for different variants, except B. 1.618, Lambda (C.37) and Delta Plus. And these three variants have only one mutation site in this range. Base on the in silico analysis, we do not anticipate that nucleocapsid protein mutations out of the region of 209 – 232aa will have any effect on test performance.

And the wet analysis study were conducted with recombination nucleocapsid proteins for all the variants, except the latest variant of Omicron (B1.1.529). No obvious difference observed when testing with different recombination nucleocapsid protein of different variants of SARS-CoV-2.

And we will conduct a wet analysis study as soon as we get the recombination nucleocapsid protein for Omicron (B1.1.529).