SARS-CoV-2 and T-cell escape

	HLA	Epitope Mutant ²
		SHVYTMSL
		SHAYTMLL
		PHAYTMSL
		SHAYTMAL
	1107.01	SFLAYTMSL
	A*02:01	SIVAYTMSL.
		SILAYAMSL
		SHAYTMSF
		THAYTMSL
		SIIPYTMSL
		QYIKWPWYT
16	A*24:02	QHIKWPWYI
		QYIKWPWYS
		REQUIQUY
108	A*02:01	RLQSLQTYA
,	A*02:01	CLOPRTFLL

Infection and immunisation against SARS-CoV-2 is capable of generating specific neutralising antibodies and T-cells. However, this immunity may begin to fade due to evolutionary mutations of the virus (Read more here).

Recent studies have reported that the specific T-cell response to SARS-CoV-2 is robust and are relatively unaffected by the mutations seen in the variants of concern (VOCs). It must be said that a loss of CD8+ T-cell responses has been observed in a small group of individuals who have either recovered from infection or who are vaccinated against the SARS-CoV-2 Omicron variant.

The evolution of CD8+ T-cell epitopes has left a weaker T-cell response in some individuals, therefore compromising the protection established through vaccinations and/or infection.

A recent paper by Ahmed, et al., aimed to identify and screen the mutations of SARS-CoV-2 involved in CD8+ T-cell escape. The researchers looked at 753 distinct HLA-specific CD8+ T-cell epitopes and SARS-CoV-2 genetic sequence data.

In this present study they found 83 SARS-CoV-2 mutations of CD8+ T-cell epitopes which may result in an escape of the T-cell response (Table 1). In future, these mutations may become of concern as they may affect the ability of SARS-CoV-2 to evade the immune response in previously-infected and vaccinated individuals.

Table 1: List of SARS-CoV-2 immunoprevalent HLA-specific CD8+ T cell epitope mutants recommended for experimental investigation (Ahmed, et al., 2022).

Epitope ¹	HLA	Epitope Mutant ²	Count
	A*02:01	SIIVYTMSL	720
		SHAYTMLL	655
		PHAYTMSL	205
		SHAYTMAL	181
CHANCINACI		SFIAYTMSL	38
691SHAYTMSL699		SIVAYTMSL	24
		SHAYAMSL	7
		SHAYTMSF	7
		SILAYTMLL PILAYTMSL SILAYTMAL SELAYTMSL SIVAYTMSL SILAYAMSL	5
		SHFYTMSL	5
		QYIKWPWYT	314
208QYIKWPWYI ₁₂₁₆	A*24:02	QHIKWPWYI	15
		QYIKWPWYS	13
BLOGLOTPAL.		RFQSLQTYV	20
000RLQSLQTYV ₁₀₀₈	A*02:01	RLQSLQTYA	10
200 YLQPRTFLL277	A*02:01	CLOPRTFLL	6

Epitope ¹	HLA	Epitope Mutant ²	Count
	A*02:01	FLELTWICE	1478
		FLFLIWICI.	1394
		FLFLTCICL.	154
		LLFLTWICL	85
»FLFLTWICL»		FIFLTWICL	33
Streemers		CLELTWICL	17
		PVFLTWICL.	14
		FLFLTWICE	13
		VLFLTWICI.	
		FLLLBVICL.	7
	A*02501	KLDDKDSNF	633
		KLDNKDPNF	226
		KLNDKDPNF	177
		KEDDKDPNE	118
		KLDYKDPNE	79
		KLDHKDPNF	62
		KLGDKDPNF	38
SLDDKDPNF ₃₀		KLDDKDQNF	35
		KLDVKDPNF	13
		KLEDKDPNF	11
		KLDGKDPNF	10
		KLADKDPNF	6
		KLVDKDPNF	6
		KLDAKDPNF	6
		KLDDKDPNS	5
		KKFPPTEPK	300
	A*03:01	KTHPPTEPN	307
		KIRPPIEPE	57
KTEPPTEPK ₃₆₈		KREPPTEPK	30
		KIPPTEPE	16-
		KIEPPIEPT	14
		KTFPPTEPL	6
	A*11.01	KKFPPTEPK	300
		KTEPPTEPN	107
		KTHPPTEPI	57
KTFPPTEPK ₃₆₉		KREPPTEPK	30
		KTEPPTEPE	16
		KIEPPTEPT	14
		KTHPPTEPL	6

		VTBGALNTPK AABGALNTPK	1162 196
DeATEGALNTPK _{HO}	A*11:01	ANEGALNIPK	101
		APEGALNTPK	36
		TTEGALNTPK	27
		KKEPPTEPKK	300
361KTFPPTEPKK ₂₇₀	A*03:01	KREPPTEPKK	30
		KTEPSTEPKN	28
305 SPRWYFYYL 122	B*07:02	SSRWYFYYL	23
			54
			54
bble 1. Cont.			5.0
ible 1, Cont. Epilope ¹	HLA	Epitope Mutant ²	5 c
	HEA	Epitope Mutant ²	
Epitope ¹ ORF3a	HLA	Epitope Mutant ² VHFLOSINF	
Epitope ¹	HLA A*2402		Count
Epitope ¹ ORF3a		VHFLQSINF	Count 339
Epitope ¹ DRP3a 112VYFLQSINF ₁₂₀	A*24:02	VHFLQSINF VYFLQSINC	Count 339 112
Epitope ¹ ORF3a		VHFLQSINF VYFLQSINC VYFLQSINS	Count 339 112 50
Epitope ¹ DRP3a 112VYFLQSINF ₁₂₀	A*24:02	VHFLOSINF VYFLOSINC VYFLOSINS LIPYDANYFL	339 112 50 2582

NOTTEDPSFLGRYM _{INC}	ANIO	TINESPLCEOM	2336
		TIDESPLCINM	
		INDESEGRIM	902
			55
		TIDESPLGISM	5
		PERMITTY	141
HISPATTON STITTY CORE		PROPRITTY	13
		PEDNOTETH	33
$_{\rm 202} \rm LFTRFFYV_{\rm 200}$	A9201	DESTRUCTOR	675
		ILCTREPYV	33
	A501-01	HTTDPSEGRE	- 5

Journal article: Ahmed, S. F., et al., 2022. <u>Identification of Potential SARS-CoV-2 CD8+ T Cell Escape Mutants</u>. *Vaccines*.

Summary by Stefan Botha