

SARS-CoV-2 and T-cell escape

HLA	Epitope Mutant ²		
61	A*0201	SIIVYTMRL SIAYTMRL FIAYTMRL SFIAYTMAL SFIAYTMSL SIVAYTMRL SIIAYAMSL SIIAYTMRF TIAYTMRL SIIPTMRL	
	1226	A*2402	QVIKWPWYT QIEKWPWYT QVIKWPWYS
		488	A*0201
	27		A*0201

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
S			
691SIAYTMSL ₆₉₉	A*02:01	SIIVYTMSL	720
		SIAYTMLL	655
		PIIAYTMSL	205
		SIAYTMAL	181
		SFIAYTMSL	38
		SIVAYTMSL	24
		SIAYAMSL	7
		SIAYTMSF	7
		THIAYTMSL	5
SIIFYTMSL	5		
1288QYIKWPWYI ₁₂₁₆	A*24:02	QYIKWPWYT QHKKWPWYI QYIKWPWYS	314 15 13
1000RLQSLQTYV ₁₀₀₈	A*02:01	RFQSLQTYV RLQSLQTYA	20 10
269YLQPRIFLL ₂₇₇	A*02:01	CLQPRIFLL	6

Table 1. Cont.

Epitope ¹	HLA	Epitope Mutant ²	Count
M			
25FLLTWKCL ₃₄	A*02:01	FLLTWKCF	1479
		FLLTWKCL	1384
		FLLTWKLL	154
		LLFLLTWKCL	85
		FFLLTWKCL	33
		CLFLLTWKCL	17
		IFVLLTWKCL	14
		FLLTWKCI	13
VLLTWKCL	8		
FLLLWKCL	7		
N			
128KLDKDPNF ₁₃₆	A*02:01	KLDKDPNF	633
		KLDKDPNF	226
		KLDKDPNF	177
		KFDKDPNF	118
		KLDKDPNF	78
		KLDKDPNF	62
		KLDKDPNF	58
		KLDKDPNF	35
		KLDKDPNF	13
		KLDKDPNF	11
		KLDKDPNF	10
		KLDKDPNF	6
		KLDKDPNF	6
KLDKDPNF	5		
341KTFPTTEPK ₃₅₀	A*03:01	KKTFPTTEPK	300
		KTFPTTEPK	307
		KTFPTTEPK	57
		KKTFPTTEPK	30
		KTFPTTEPK	16
		KTFPTTEPK	14
341KTFPTTEPK ₃₅₀	A*11:01	KKTFPTTEPK	300
		KTFPTTEPK	307
		KTFPTTEPK	57
		KKTFPTTEPK	30
		KTFPTTEPK	16
		KTFPTTEPK	14

156ATEGALNTPK ₁₆₃	A*11:01	AIEGALNTPK	9685
		VIEGALNTPK	1162
		AAEGALNTPK	196
		ANEALNTPK	101
		APEGALNTPK	36
TIEGALNTPK	27		
361KTFPTTEPK ₃₇₀	A*03:01	KKTFPTTEPK	300
		KKTFPTTEPK	30
		KTFPTTEPKN	28
305SPRWYFYLL ₃₁₁	F*07:02	SRWYFYLL	23

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Table 1. Cont.

Epitope ¹	HLA	Epitope Mutant ²	Count
O0F3a			
102VYFLQSINF ₁₂₀	A*24:02	VHFLQSINF	339
		VYFLQSNIC	112
		VYFLQSNIS	50
138LLYDANYFL ₁₄₇	A*02:01	LIFDANYFL	2582
		LLYDANYFF	1276
207FTSDVYQLY ₂₁₅	A*01:01	FTSDVYQLC FTSDVYQLH	121 64

HLA	Epitope	Count
A*02:01	SIIVYTMSL	720
	SIAYTMLL	655
	PIIAYTMSL	205
A*03:01	KKTFPTTEPK	300
	KKTFPTTEPK	30
	KTFPTTEPKN	28
A*11:01	KKTFPTTEPK	300
	KTFPTTEPK	307
	KTFPTTEPK	57
A*24:02	QYIKWPWYT	314
	QHKKWPWYI	15
	QYIKWPWYS	13
A*02:01	RFQSLQTYV	20
	RLQSLQTYA	10
	CLQPRIFLL	6
A*02:01	FLLTWKCF	1479
	FLLTWKCL	1384
	FLLTWKLL	154
	LLFLLTWKCL	85
	FFLLTWKCL	33
	CLFLLTWKCL	17
	IFVLLTWKCL	14
	FLLTWKCI	13
VLLTWKCL	8	
FLLLWKCL	7	
A*02:01	KLDKDPNF	633
	KLDKDPNF	226
	KLDKDPNF	177
	KFDKDPNF	118
	KLDKDPNF	78
	KLDKDPNF	62
	KLDKDPNF	58
	KLDKDPNF	35
	KLDKDPNF	13
	KLDKDPNF	11
	KLDKDPNF	10
	KLDKDPNF	6
	KLDKDPNF	6
KLDKDPNF	5	
A*03:01	KKTFPTTEPK	300
	KTFPTTEPK	307
	KTFPTTEPK	57
	KKTFPTTEPK	30
	KTFPTTEPK	16
	KTFPTTEPK	14
A*11:01	KKTFPTTEPK	300
	KTFPTTEPK	307
	KTFPTTEPK	57
	KKTFPTTEPK	30
	KTFPTTEPK	16
	KTFPTTEPK	14
A*24:02	VHFLQSINF	339
	VYFLQSNIC	112
	VYFLQSNIS	50
A*02:01	LIFDANYFL	2582
	LLYDANYFF	1276
A*01:01	FTSDVYQLC	121
	FTSDVYQLH	64

Journal article: Ahmed, S. F., et al., 2022. [Identification of Potential SARS-CoV-2 CD8+ T Cell Escape Mutants](#). *Vaccines*.

Summary by Stefan Botha