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Correspondence

Sequence similarity suggests molecular mimicry-induced cardiovascular symptoms in multisystem inflammatory syndrome in children (MIS-C)



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To the Editor,

Cardiovascular symptoms are the hallmark of multisystem inflammatory syndrome in children (MIS-C) [1]. The pathophysiology includes cardiomyocyte invasion, endothelium injury and microvascular injury [2]. Molecular mimicry is a suspected mechanism of MIS-C pathogenesis after severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) infection. Several 6-mer or 7-mer viral peptides were revealed to mimic human proteins, which may result in autoimmune reaction in various tissues [3,4]. However, potential cardiovascular molecular mimicking targets of SARS-CoV-2 are rarely reported.

Group A streptococcus (GAS) is the pathogen with bacterial components mimicking epitopes of human cardiovascular system [5]. We compared target protein sequences of GAS with SARS-CoV-2 via PSI-BLAST (<https://www.ebi.ac.uk/Tools/sss/psiblast/>) UniProtKB COVID19 database with default parameters. Human myosin heavy chain 6 (MYH6) and proteins in human heart valve proteomes, which were studied in GAS autoimmune reaction [6,7], were included. The FASTA formatted protein sequences were obtained by UniProt (<https://www.uniprot.org/>). Based on this strategy, we identified eight 6-mer or 7-mer consecutively identical peptides (Table 1).

The comparison with Basic Local Alignment Search Tool (BLAST) in

SARS-CoV-2 studies needs more evidence to illustrate its autoimmune property [8]. We applied NetMHCcons-1.1 to confirm the class I human leukocyte antigen (HLA) affinity of the above-mentioned peptides [9]. The sequences with their three up- and three down- stream amino acids were input, and the “Peptide length” was set to “8–11mer peptides”. All alleles in “HLA supertype representative” were selected. As the results in Table 1, some peptides have a certain affinity to different class I HLAs.

The A02, B35, C04 allele group combination has been considered to increase MIS-C susceptibility [10,11]. In our study, the peptide EKMVSL mimicking MYH6 possesses strong binding affinity to the vast majority of the alleles in A02 and weak binding affinity to nearly 1/3 alleles in B35 and 3/4 alleles in C04 (Table 2), while other peptides do not show this property. Additionally, we found that the previously reported peptide IVDTVSA mimicking mitochondrial alanine-tRNA ligase (AARS2) also possesses a broad affinity to A02, B35, C04 allele group combination [4].

In conclusion, molecular mimicry may contribute to MIS-C cardiovascular symptoms similar to GAS targets.

Abbreviations: MIS-C, multisystem inflammatory syndrome in children; SARS-CoV-2, severe acute respiratory syndrome coronavirus 2; GAS, group A streptococcus; MYH6, human myosin heavy chain 6; BLAST, basic local alignment search tool; HLA, human leukocyte antigen; AARS2, mitochondrial alanine-tRNA ligase.

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Table 1

Affinity of potential molecular mimicking peptides of SARS-CoV-2 to class I HLA supertype representatives.

| Protein name human vs. SARS-CoV-2 | Peptide mimicking* | | | | | | | | | HLA binding [1-log50k; peptide]** | |
|--|--------------------|----------|----------|----------|----------|----------|----------|----------|----------|--|--|
| MYH6 [UniProt: P13533] vs. R1AB [UniProt: PODTD1] | 877 | E | K | M | V | S | L | L | 883 | HLA-A02:01 [0.671; KMVSLLSV] | |
| | 3909 | E | K | M | V | S | L | L | 3915 | HLA-A02:01 [0.672; KMVSLLSV] | |
| | | | | | | | | | | HLA-A26:01 [0.439; EAFEKMOVSL] [#] | |
| | | | | | | | | | | HLA-A26:01 [0.398; EAFEKMOVSL] [#] | |
| | | | | | | | | | | HLA-B08:01 [0.411; EAFEKMOVSL] [#] | |
| | | | | | | | | | | HLA-B39:01 [0.617; EKMVSLLSV] | |
| | | | | | | | | | | HLA-B39:01 [0.405; MVSLLSV] [#] | |
| | | | | | | | | | | HLA-B40:01 [0.656; FEKMOVSLLSV] | |
| | | | | | | | | | | HLA-B40:01 [0.605; FEKMOVSL] [#] | |
| | | | | | | | | | | HLA-B40:01 [0.390; FEKMOVSLLSV] [#] | |
| | | | | | | | | | | HLA-B15:01 [0.629; KMOVSLLSV] [#] | |
| | | | | | | | | | | HLA-B15:01 [0.436; KMOVSLLSV] [#] | |
| MYH6 [UniProt: P13533] vs. R1AB [UniProt: PODTD1] | 1804 | Q | I | A | L | K | G | G | K | 1811 | HLA-A03:01 [0.593; KIALKGGK] |
| | 2757 | K | I | A | L | K | G | G | K | 2764 | HLA-A03:01 [0.529; VTTKIALK] [#] |
| CAL1A1 [UniProt: P02452] vs. R1AB [UniProt: PODTD1] | 1116 | L | Q | G | P | P | G | | | 1121 | HLA-A03:01 [0.529; TLQPPGTGK] [#] |
| | 5604 | L | Q | G | P | P | G | | | 5609 | HLA-A03:01 [0.426; STLQPPGTGK] [#] |
| COL14A1 [UniProt: Q05707] vs. R1AB [UniProt: PODTD1] | 666 | T | E | E | V | V | L | K | | 672 | HLA-A01:01 [0.209; LTEEVVLK] [#] |
| | 753 | T | E | E | V | V | L | K | | 759 | HLA-A03:01 [0.488; VLTEEVVLK] [#] |
| | | | | | | | | | | | HLA-A26:01 [0.350; EVVLKTGDL] [#] |
| | | | | | | | | | | | HLA-B40:01 [0.377; EEVVLKTGDL] [#] |
| | | | | | | | | | | | HLA-B40:01 [0.509; TEEVVLKTGDL] [#] |
| COL14A1 [UniProt: Q05707] vs. R1AB [UniProt: PODTD1] | 1671 | Q | G | P | P | G | T | | | 1676 | HLA-A03:01 [0.529; TLQPPGTGK] [#] |
| | 5605 | Q | G | P | P | G | T | | | 5610 | HLA-A03:01 [0.426; STLQPPGTGK] [#] |
| COL6A2 [UniProt: P12110] vs. R1AB [UniProt: PODTD1] | 352 | G | P | D | G | Y | P | | | 357 | N/A |
| | 192 | G | P | D | G | Y | P | | | 197 | |
| CAL6A3 [UniProt: D9ZGF2] vs. R1AB [UniProt: PODTD1] | 912 | K | A | L | N | L | G | | | 917 | HLA-B58:01 [0.732; KALNLGETF] |
| | 701 | K | A | L | N | L | G | | | 706 | HLA-B15:01 [0.673; KLKALNLGETF] |
| | | | | | | | | | | | HLA-B15:01 [0.562; ALNLGETF] [#] |
| | | | | | | | | | | | HLA-B15:01 [0.462; KALNLGETF] [#] |
| CAL6A3 [UniProt: D9ZGF2] vs. NCAP [UniProt: PODTC9] | 2200 | F | G | R | R | G | P | | | 2213 | N/A |
| | 274 | F | G | R | R | G | P | | | 287 | |

* Sequence of Human and SARS-CoV-2 are placed on the upper and lower sides respectively.

** HLA binding affinity is scored by 1-log50k; 6-mer or 7-mer consecutively identical peptides (bold) with three upstream and three downstream amino acids were included in scoring.

Weak binding.

Table 2
Affinity of peptide EKMVSLL to class I HLAs in A02, B35, C04 combination.

| Allele groups | Alleles [1-log50k; peptide] * Strong binding | | Weak binding |
|---------------|---|--|--|
| A02 | HLA-A02:01 [0.672; KMVSLLSVL] [0.671; KMVSLLSV] | HLA-A02:139 [0.805; KMVSLLSV] [0.670; KMVSLLSVL] | HLA-A02:02 [0.618; KMVSLLSV] |
| | HLA-A02:02 [0.746; KMVSLLSVL] | HLA-A02:140 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:04 [0.474; KMVSLLSVL] |
| | HLA-A02:03 [0.789; KMVSLLSVL] [0.718; KMVSLLSV] | HLA-A02:141 [0.848; KMVSLLSV] [0.719; KMVSLLSVL] | HLA-A02:05 [0.564; KMVSLLSVL] |
| | HLA-A02:04 [0.667; KMVSLLSV] | HLA-A02:142 [0.822; KMVSLLSV] [0.662; KMVSLLSVL] | HLA-A02:06 [0.633; KMVSLLSVL] |
| | HLA-A02:05 [0.702; KMVSLLSV] | HLA-A02:144 [0.806; KMVSLLSV] | HLA-A02:07 [0.253; KMVSLLSVL] |
| | HLA-A02:06 [0.830; KMVSLLSV] | HLA-A02:145 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:08 [0.567; KMVSLLSV] [0.437; KMVSLLSVL] |
| | HLA-A02:07 [0.410; KMVSLLSV] | HLA-A02:146 [0.656; KMVSLLSV] | HLA-A02:10 [0.451; KMVSLLSVL] |
| | HLA-A02:09 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:147 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:14 [0.596; KMVSLLSVL] |
| | HLA-A02:10 [0.660; KMVSLLSV] | HLA-A02:148 [0.785; KMVSLLSV] [0.674; KMVSLLSVL] | HLA-A02:17 [0.366; KMVSLLSVL] |
| | HLA-A02:11 [0.940; KMVSLLSV] [0.863; KMVSLLSVL] | HLA-A02:149 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:18 [0.253; KMVSLLSVL] |
| | HLA-A02:12 [0.857; KMVSLLSV] [0.750; KMVSLLSVL] | HLA-A02:150 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:19 [0.544; KMVSLLSVL] |
| | HLA-A02:13 [0.851; KMVSLLSV] [0.746; KMVSLLSVL] | HLA-A02:151 [0.809; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:20 [0.568; KMVSLLSVL] |
| | HLA-A02:14 [0.739; KMVSLLSV] | HLA-A02:153 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:21 [0.609; KMVSLLSVL] |
| | HLA-A02:16 [0.869; KMVSLLSV] [0.738; KMVSLLSVL] | HLA-A02:154 [0.642; KMVSLLSV] | HLA-A02:28 [0.609; KMVSLLSVL] |
| | HLA-A02:17 [0.542; KMVSLLSV] | HLA-A02:155 [0.808; KMVSLLSV] [0.742; KMVSLLSVL] | HLA-A02:33 [0.486; KMVSLLSVL] |
| | HLA-A02:18 [0.410; KMVSLLSV] | HLA-A02:156 [0.665; KMVSLLSV] | HLA-A02:34 [0.567; KMVSLLSVL] |
| | HLA-A02:19 [0.777; KMVSLLSV] | HLA-A02:157 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:35 [0.574; KMVSLLSVL] |
| | HLA-A02:20 [0.753; KMVSLLSV] | HLA-A02:158 [0.876; KMVSLLSV] [0.770; KMVSLLSVL] | HLA-A02:36 [0.518; KMVSLLSVL] |
| | HLA-A02:21 [0.806; KMVSLLSV] | HLA-A02:159 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:37 [0.571; KMVSLLSVL] |
| | HLA-A02:22 [0.849; KMVSLLSV] [0.735; KMVSLLSVL] | HLA-A02:160 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:39 [0.547; KMVSLLSVL] |
| | HLA-A02:24 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:161 [0.842; KMVSLLSV] [0.720; KMVSLLSVL] | HLA-A02:45 [0.573; KMVSLLSVL] |
| | HLA-A02:25 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:162 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:46 [0.606; KMVSLLSVL] |
| | HLA-A02:26 [0.834; KMVSLLSV] [0.709; KMVSLLSVL] | HLA-A02:163 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:48 [0.615; KMVSLLSVL] |
| | HLA-A02:27 [0.810; KMVSLLSV] [0.677; KMVSLLSVL] | HLA-A02:164 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:51 [0.609; KMVSLLSVL] |
| | HLA-A02:28 [0.806; KMVSLLSV] | HLA-A02:165 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:54 [0.467; KMVSLLSV] |
| | HLA-A02:29 [0.804; KMVSLLSV] [0.650; KMVSLLSVL] | HLA-A02:166 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:55 [0.329; EAFERKMVSLL] [0.315; EAFERKMVSLL] |
| | HLA-A02:30 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:167 [0.829; KMVSLLSV] [0.701; KMVSLLSVL] | HLA-A02:56 [0.395; KMVSLLSVL] |
| | HLA-A02:31 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:168 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:57 [0.393; KMVSLLSVL] |
| | HLA-A02:33 [0.671; KMVSLLSV] | HLA-A02:170 [0.806; KMVSLLSV] | HLA-A02:60 [0.590; KMVSLLSVL] |
| | HLA-A02:34 [0.774; KMVSLLSV] | HLA-A02:171 [0.834; KMVSLLSV] [0.709; KMVSLLSVL] | HLA-A02:61 [0.609; KMVSLLSVL] |
| | HLA-A02:35 [0.790; KMVSLLSV] | HLA-A02:172 [0.702; KMVSLLSV] | HLA-A02:62 [0.465; KMVSLLSVL] |
| | HLA-A02:36 [0.741; KMVSLLSV] | HLA-A02:173 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:64 [0.578; KMVSLLSVL] |
| | HLA-A02:37 [0.770; KMVSLLSV] | HLA-A02:174 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:65 [0.367; KMVSLLSV] [0.342; KMVSLLSVL] |
| | HLA-A02:38 [0.825; KMVSLLSV] [0.724; KMVSLLSVL] | HLA-A02:175 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:72 [0.609; KMVSLLSVL] |
| | HLA-A02:39 [0.707; KMVSLLSV] | HLA-A02:176 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:78 [0.430; KMVSLLSVL] |
| | HLA-A02:40 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:177 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:79 [0.609; KMVSLLSVL] |
| | HLA-A02:41 [0.800; KMVSLLSV] [0.645; KMVSLLSVL] | HLA-A02:178 [0.693; KMVSLLSV] | HLA-A02:80 [0.513; KMVSLLSVL] |
| | HLA-A02:42 [0.814; KMVSLLSV] [0.687; KMVSLLSVL] | HLA-A02:179 [0.702; KMVSLLSV] | HLA-A02:84 [0.569; KMVSLLSVL] |
| | HLA-A02:44 [0.822; KMVSLLSV] [0.662; KMVSLLSVL] | HLA-A02:180 [0.806; KMVSLLSV] | HLA-A02:87 [0.593; KMVSLLSVL] [0.579; KMVSLLSV] |
| | HLA-A02:45 [0.697; KMVSLLSV] | HLA-A02:181 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:91 [0.609; KMVSLLSVL] |
| | HLA-A02:46 [0.718; KMVSLLSV] | HLA-A02:182 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:92 [0.606; KMVSLLSVL] |
| | HLA-A02:47 [0.819; KMVSLLSV] [0.758; KMVSLLSVL] | | HLA-A02:99 [0.635; KMVSLLSVL] |
| | HLA-A02:48 [0.745; KMVSLLSV] | | HLA-A02:101 [0.614; KMVSLLSVL] |
| | HLA-A02:49 [0.802; KMVSLLSV] [0.697; KMVSLLSVL] | | HLA-A02:103 [0.333; KMVSLLSVL] |
| | HLA-A02:50 [0.875; KMVSLLSV] [0.780; KMVSLLSVL] | | HLA-A02:106 [0.609; KMVSLLSVL] |
| | HLA-A02:51 [0.806; KMVSLLSV] | | HLA-A02:108 [0.338; KMVSLLSVL] |
| | HLA-A02:52 [0.806; KMVSLLSV] [0.684; KMVSLLSVL] | | HLA-A02:110 [0.462; KMVSLLSVL] |
| | HLA-A02:54 [0.719; KMVSLLSV] | | HLA-A02:112 [0.424; KMVSLLSVL] [0.381; KMVSLLSV] |
| | HLA-A02:56 [0.567; KMVSLLSV] | | HLA-A02:114 [0.461; KMVSLLSVL] |
| | | | HLA-A02:117 [0.558; KMVSLLSVL] |
| | | | HLA-A02:122 [0.615; KMVSLLSVL] |
| | | | HLA-A02:126 [0.609; KMVSLLSVL] |
| | | | HLA-A02:127 [0.601; KMVSLLSVL] |
| | | | HLA-A02:129 [0.531; KMVSLLSVL] [0.500; KMVSLLSV] |
| | | | HLA-A02:130 [0.253; KMVSLLSVL] |
| | | | HLA-A02:136 [0.582; KMVSLLSV] |
| | | HLA-A02:137 [0.609; KMVSLLSVL] | |
| | | HLA-A02:143 [0.571; KMVSLLSV] | |
| | | HLA-A02:144 [0.609; KMVSLLSVL] | |
| | | HLA-A02:146 [0.504; KMVSLLSVL] | |
| | | HLA-A02:152 [0.367; KMVSLLSV] [0.342; KMVSLLSVL] | |
| | | HLA-A02:154 [0.494; KMVSLLSVL] | |
| | | HLA-A02:156 [0.499; KMVSLLSVL] | |
| | | HLA-A02:169 [0.620; KMVSLLSV] [0.434; KMVSLLSVL] | |
| | | HLA-A02:170 [0.609; KMVSLLSVL] | |

(continued on next page)

Table 2 (continued)

| Allele groups | Alleles [1-log50k; peptide] * | Weak binding |
|---------------|--|--|
| | HLA-A02:57 [0.532; KMVSLLSV] | HLA-A02:172 [0.564; KMVSLLSVL] |
| | HLA-A02:58 [0.798; KMVSLLSV] [0.703; KMVSLLSVL] | HLA-A02:178 [0.488; KMVSLLSVL] |
| | HLA-A02:59 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:179 [0.564; KMVSLLSVL] |
| | HLA-A02:60 [0.764; KMVSLLSV] | HLA-A02:180 [0.609; KMVSLLSVL] |
| | HLA-A02:61 [0.806; KMVSLLSV] | HLA-A02:184 [0.626; KMVSLLSVL] |
| | HLA-A02:62 [0.702; KMVSLLSV] | HLA-A02:188 [0.548; KMVSLLSVL] |
| | HLA-A02:63 [0.808; KMVSLLSV] [0.742; KMVSLLSVL] | HLA-A02:191 [0.344; KMVSLLSVL] |
| | HLA-A02:64 [0.747; KMVSLLSV] | HLA-A02:195 [0.465; KMVSLLSVL] |
| | HLA-A02:66 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:217 [0.398; KMVSLLSVL] |
| | HLA-A02:67 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:219 [0.253; KMVSLLSVL] |
| | HLA-A02:68 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:224 [0.488; KMVSLLSVL] |
| | HLA-A02:69 [0.935; KMVSLLSV] [0.860; KMVSLLSVL] | HLA-A02:229 [0.581; KMVSLLSVL] |
| | HLA-A02:70 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:232 [0.564; KMVSLLSVL] |
| | HLA-A02:71 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:233 [0.635; KMVSLLSVL] |
| | HLA-A02:72 [0.806; KMVSLLSV] | HLA-A02:242 [0.547; KMVSLLSVL] |
| | HLA-A02:73 [0.837; KMVSLLSV] [0.725; KMVSLLSVL] | HLA-A02:244 [0.451; KMVSLLSVL] |
| | HLA-A02:74 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:246 [0.461; KMVSLLSVL] |
| | HLA-A02:75 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:247 [0.512; KMVSLLSVL] |
| | HLA-A02:76 [0.810; KMVSLLSV] [0.666; KMVSLLSVL] | HLA-A02:248 [0.609; KMVSLLSVL] |
| | HLA-A02:77 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | HLA-A02:249 [0.576; KMVSLLSVL] |
| | HLA-A02:78 [0.654; KMVSLLSV] | HLA-A02:254 [0.605; KMVSLLSVL] |
| | HLA-A02:79 [0.806; KMVSLLSV] | HLA-A02:255 [0.160; KMVSLLSVL] |
| | HLA-A02:80 [0.677; KMVSLLSV] | HLA-A02:259 [0.609; KMVSLLSVL] |
| | HLA-A02:81 [0.688; KMVSLLSVL] [0.653; KMVSLLSV] | HLA-A02:261 [0.235; KMVSLLSVL] |
| | HLA-A02:84 [0.754; KMVSLLSV] | HLA-A02:264 [0.253; KMVSLLSVL] |
| | HLA-A02:85 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:86 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:89 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:90 [0.844; KMVSLLSV] [0.695; KMVSLLSVL] | |
| | HLA-A02:91 [0.806; KMVSLLSV] | |
| | HLA-A02:92 [0.718; KMVSLLSV] | |
| | HLA-A02:93 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:95 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:96 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:97 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:99 [0.791; KMVSLLSV] | |
| | HLA-A02:101 [0.779; KMVSLLSV] | |
| | HLA-A02:102 [0.808; KMVSLLSV] [0.742; KMVSLLSVL] | |
| | HLA-A02:103 [0.402; KMVSLLSV] | |
| | HLA-A02:104 [0.849; KMVSLLSV] [0.735; KMVSLLSVL] | |
| | HLA-A02:105 [0.835; KMVSLLSV] [0.721; KMVSLLSVL] | |
| | HLA-A02:105 [0.806; KMVSLLSV] | |
| | HLA-A02:107 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:108 [0.442; KMVSLLSV] | |
| | HLA-A02:109 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:110 [0.548; KMVSLLSV] | |
| | HLA-A02:111 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:183 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:184 [0.757; KMVSLLSV] | |
| | HLA-A02:185 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:186 [0.808; KMVSLLSV] [0.742; KMVSLLSVL] | |
| | HLA-A02:187 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:188 [0.704; KMVSLLSV] | |
| | HLA-A02:189 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:190 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:191 [0.475; KMVSLLSV] | |
| | HLA-A02:192 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:193 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:194 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:195 [0.608; KMVSLLSV] | |
| | HLA-A02:196 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:197 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:198 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:199 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:200 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:201 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:202 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:203 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:204 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:205 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:206 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:207 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:208 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:209 [0.808; KMVSLLSV] [0.742; KMVSLLSVL] | |
| | HLA-A02:210 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:211 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:212 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:213 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:214 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:215 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:216 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:217 [0.553; KMVSLLSV] | |
| | HLA-A02:218 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:219 [0.410; KMVSLLSV] | |
| | HLA-A02:220 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:221 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |
| | HLA-A02:224 [0.719; KMVSLLSV] | |
| | HLA-A02:228 [0.828; KMVSLLSV] [0.691; KMVSLLSVL] | |

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Table 2 (continued)

| Allele groups | Alleles [1-log50k; peptide] * | Weak binding |
|---------------|---|--|
| | KMVSLLSVL] | HLA-A02:229 [0.701; KMVSLLSV] |
| | HLA-A02:114 [0.640; KMVSLLSV] | HLA-A02:230 [0.868; KMVSLLSV] [0.774; KMVSLLSV] |
| | HLA-A02:115 [0.808; KMVSLLSV] [0.742; KMVSLLSV] | HLA-A02:231 [0.690; KMVSLLSV] [0.499; KMVSLLSV] |
| | HLA-A02:116 [0.817; KMVSLLSV] [0.680; KMVSLLSV] | HLA-A02:232 [0.702; KMVSLLSV] |
| | HLA-A02:117 [0.692; KMVSLLSV] | HLA-A02:233 [0.812; KMVSLLSV] |
| | HLA-A02:118 [0.828; KMVSLLSV] [0.691; KMVSLLSV] | HLA-A02:234 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | HLA-A02:119 [0.828; KMVSLLSV] [0.691; KMVSLLSV] | HLA-A02:235 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | HLA-A02:120 [0.828; KMVSLLSV] [0.691; KMVSLLSV] | HLA-A02:236 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | HLA-A02:121 [0.828; KMVSLLSV] [0.691; KMVSLLSV] | HLA-A02:237 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | HLA-A02:122 [0.781; KMVSLLSV] | HLA-A02:238 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | HLA-A02:123 [0.828; KMVSLLSV] [0.691; KMVSLLSV] | HLA-A02:239 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | HLA-A02:124 [0.688; KMVSLLSV] [0.653; KMVSLLSV] | HLA-A02:240 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | HLA-A02:126 [0.806; KMVSLLSV] | HLA-A02:241 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | HLA-A02:127 [0.793; KMVSLLSV] | HLA-A02:242 [0.707; KMVSLLSV] |
| | HLA-A02:128 [0.890; KMVSLLSV] [0.758; KMVSLLSV] | HLA-A02:243 [0.814; KMVSLLSV] [0.668; KMVSLLSV] |
| | HLA-A02:130 [0.410; KMVSLLSV] | HLA-A02:244 [0.660; KMVSLLSV] |
| | HLA-A02:131 [0.870; KMVSLLSV] [0.772; KMVSLLSV] | HLA-A02:245 [0.837; KMVSLLSV] [0.725; KMVSLLSV] |
| | HLA-A02:132 [0.828; KMVSLLSV] [0.691; KMVSLLSV] | HLA-A02:246 [0.640; KMVSLLSV] |
| | HLA-A02:133 [0.828; KMVSLLSV] [0.691; KMVSLLSV] | HLA-A02:247 [0.646; KMVSLLSV] |
| | HLA-A02:134 [0.828; KMVSLLSV] [0.691; KMVSLLSV] | HLA-A02:248 [0.806; KMVSLLSV] |
| | HLA-A02:135 [0.594; KMVSLLSV] [0.594; KMVSLLSV] | HLA-A02:249 [0.708; KMVSLLSV] |
| | HLA-A02:136 [0.621; KMVSLLSV] | HLA-A02:251 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | HLA-A02:137 [0.806; KMVSLLSV] | HLA-A02:252 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | HLA-A02:138 [0.828; KMVSLLSV] [0.691; KMVSLLSV] | HLA-A02:253 [0.868; KMVSLLSV] [0.774; KMVSLLSV] |
| | | HLA-A02:254 [0.759; KMVSLLSV] |
| | | HLA-A02:255 [0.319; KMVSLLSV] |
| | | HLA-A02:256 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | | HLA-A02:257 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | | HLA-A02:258 [0.868; KMVSLLSV] [0.774; KMVSLLSV] |
| | | HLA-A02:259 [0.806; KMVSLLSV] |
| | | HLA-A02:260 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| | | HLA-A02:261 [0.369; KMVSLLSV] |
| | | HLA-A02:262 [0.830; KMVSLLSV] [0.702; KMVSLLSV] |
| | | HLA-A02:263 [0.839; KMVSLLSV] [0.705; KMVSLLSV] |
| | | HLA-A02:264 [0.868; KMVSLLSV] [0.774; KMVSLLSV] |
| | | HLA-A02:265 [0.410; KMVSLLSV] |
| | | HLA-A02:266 [0.828; KMVSLLSV] [0.691; KMVSLLSV] |
| B35 | N/A | HLA-B35:02 [0.341; EAFKMSVSL] [0.293; EAFKMSVSL] |
| | | HLA-B35:03 [0.171; EAFKMSVSL] |
| | | HLA-B35:04 [0.341; EAFKMSVSL] [0.293; EAFKMSVSL] |
| | | HLA-B35:06 [0.265; EAFKMSVSL] [0.220; EAFKMSVSL] |
| | | HLA-B35:09 [0.341; EAFKMSVSL] [0.293; EAFKMSVSL] |
| | | HLA-B35:11 [0.448; EAFKMSVSL] |
| | | HLA-B35:12 [0.341; EAFKMSVSL] [0.293; EAFKMSVSL] |
| | | HLA-B35:18 [0.225; EAFKMSVSL] |
| | | HLA-B35:09 [0.465; EAFKMSVSL] [0.429; EAFKMSVSL] |

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Table 2 (continued)

| Allele groups | Alleles [1-log50k; peptide] * Strong binding | Weak binding |
|---------------|---|---|
| | | HLA-B35:22 [0.376; EAFEKMVSL] [0.296; EAFEKMVSL] |
| | | HLA-B35:25 [0.430; EAFEKMVSL] |
| | | HLA-B35:26 [0.344; EKMVSLLSVL] |
| | | HLA-B35:28 [0.499; KMVSLLSVL] |
| | | HLA-B35:29 [0.508; EAFEKMVSL] [0.435; EAFEKMVSL] |
| | | HLA-B35:31 [0.395; EAFEKMVSL] |
| | | HLA-B35:33 [0.258; EAFEKMVSL] [0.197; EAFEKMVSL] |
| | | HLA-B35:34 [0.332; EAFEKMVSL] [0.271; EAFEKMVSL] |
| | | HLA-B35:35 [0.471; EAFEKMVSL] [0.430; EAFEKMVSL] |
| | | HLA-B35:36 [0.256; EAFEKMVSL] [0.196; EAFEKMVSL] |
| | | HLA-B35:38 [0.205; EAFEKMVSL] |
| | | HLA-B35:39 [0.332; EAFEKMVSL] [0.271; EAFEKMVSL] |
| | | HLA-B35:44 [0.385; EAFEKMVSL] [0.323; EAFEKMVSL] |
| | | HLA-B35:47 [0.465; EAFEKMVSL] [0.429; EAFEKMVSL] |
| | | HLA-B35:49 [0.492; EAFEKMVSL] [0.442; EAFEKMVSL] |
| | | HLA-B35:55 [0.256; EAFEKMVSL] [0.196; EAFEKMVSL] |
| | | HLA-B35:56 [0.215; EAFEKMVSL] |
| | | HLA-B35:59 [0.265; EAFEKMVSL] [0.220; EAFEKMVSL] |
| | | HLA-B35:63 [0.531; FEKMVSLLSVL] [0.479; FEKMVSL] |
| | | [0.421; FEKMVSLLSV] |
| | | HLA-B35:69 [0.519; EAFEKMVSL] [0.459; EAFEKMVSL] |
| | | HLA-B35:70 [0.256; EAFEKMVSL] [0.196; EAFEKMVSL] |
| | | HLA-B35:74 [0.247; EAFEKMVSL] [0.196; EAFEKMVSL] |
| | | HLA-B35:75 [0.282; EAFEKMVSL] [0.220; EAFEKMVSL] |
| | | HLA-B35:81 [0.355; EAFEKMVSL] [0.309; EAFEKMVSL] |
| | | HLA-B35:83 [0.341; EAFEKMVSL] [0.293; EAFEKMVSL] |
| | | HLA-B35:84 [0.248; EAFEKMVSL] [0.200; EKMVSLLSVL] |
| | | [0.196; FEKMVSLLSVL] [0.192; EAFEKMVSL] |
| | | HLA-B35:85 [0.135; EAFEKMVSL] [0.125; EAFEKMVSL] |
| | | HLA-B35:87 [0.373; EAFEKMVSL] |
| | | HLA-B35:88 [0.336; EAFEKMVSL] [0.284; EAFEKMVSL] |
| | | HLA-B35:92 [0.456; EAFEKMVSL] [0.391; EAFEKMVSL] |
| | | HLA-B35:93 [0.447; EAFEKMVSL] |
| | | HLA-B35:95 [0.208; EKMVSLLSVL] [0.190; EAFEKMVSL] |
| | | [0.177; EAFEKMVSL] |
| | | HLA-B35:96 [0.321; EAFEKMVSL] [0.238; EAFEKMVSL] |
| | | HLA-B35:98 [0.256; EAFEKMVSL] [0.196; EAFEKMVSL] |
| | | HLA-B35:106 [0.256; EAFEKMVSL] [0.196; EAFEKMVSL] |
| | | HLA-B35:109 [0.287; EAFEKMVSL] [0.226; EAFEKMVSL] |
| | | HLA-B35:117 [0.381; EAFEKMVSL] |
| | | HLA-B35:127 [0.256; EAFEKMVSL] [0.196; EAFEKMVSL] |
| | | HLA-B35:128 [0.256; EAFEKMVSL] [0.196; EAFEKMVSL] |
| | | HLA-B35:136 [0.412; EAFEKMVSL] [0.310; EAFEKMVSL] |

C04

N/A

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Table 2 (continued)

| Allele groups | Alleles [1-log50k; peptide] * Strong binding | Weak binding |
|---------------|---|---|
| | | HLA-C04:01 [0.197; AFEKMSVLL] [0.172; FEKMSVLL] |
| | | HLA-C04:05 [0.163; AFEKMSVLL] |
| | | HLA-C04:07 [0.163; AFEKMSVLL] |
| | | HLA-C04:08 [0.168; AFEKMSVLL] |
| | | HLA-C04:10 [0.177; AFEKMSVLL] |
| | | HLA-C04:12 [0.163; AFEKMSVLL] |
| | | HLA-C04:14 [0.125; AFEKMSVLL] |
| | | HLA-C04:18 [0.163; AFEKMSVLL] |
| | | HLA-C04:19 [0.163; AFEKMSVLL] |
| | | HLA-C04:20 [0.163; AFEKMSVLL] |
| | | HLA-C04:23 [0.141; AFEKMSVLL] |
| | | HLA-C04:24 [0.163; AFEKMSVLL] |
| | | HLA-C04:25 [0.163; AFEKMSVLL] |
| | | HLA-C04:26 [0.163; AFEKMSVLL] |
| | | HLA-C04:27 [0.163; AFEKMSVLL] |
| | | HLA-C04:28 [0.163; AFEKMSVLL] |
| | | HLA-C04:30 [0.163; AFEKMSVLL] |
| | | HLA-C04:32 [0.180; AFEKMSVLL] |
| | | HLA-C04:33 [0.163; AFEKMSVLL] |
| | | HLA-C04:35 [0.163; AFEKMSVLL] |
| | | HLA-C04:37 [0.221; AFEKMSVLL] |
| | | HLA-C04:38 [0.163; AFEKMSVLL] |
| | | HLA-C04:39 [0.163; AFEKMSVLL] |
| | | HLA-C04:40 [0.141; AFEKMSVLL] |
| | | HLA-C04:41 [0.163; AFEKMSVLL] |
| | | HLA-C04:42 [0.151; AFEKMSVLL] |
| | | HLA-C04:43 [0.163; AFEKMSVLL] |
| | | HLA-C04:44 [0.163; AFEKMSVLL] |
| | | HLA-C04:45 [0.163; AFEKMSVLL] |
| | | HLA-C04:46 [0.163; AFEKMSVLL] |
| | | HLA-C04:47 [0.163; AFEKMSVLL] |
| | | HLA-C04:48 [0.178; AFEKMSVLL] |
| | | HLA-C04:49 [0.198; AFEKMSVLL] |
| | | HLA-C04:50 [0.163; AFEKMSVLL] |
| | | HLA-C04:51 [0.163; AFEKMSVLL] |
| | | HLA-C04:52 [0.163; AFEKMSVLL] |
| | | HLA-C04:53 [0.163; AFEKMSVLL] |
| | | HLA-C04:54 [0.277; AFEKMSVLL] |
| | | HLA-C04:56 [0.163; AFEKMSVLL] |
| | | HLA-C04:57 [0.163; AFEKMSVLL] |
| | | HLA-C04:60 [0.163; AFEKMSVLL] |
| | | HLA-C04:61 [0.154; AFEKMSVLL] |
| | | HLA-C04:62 [0.163; AFEKMSVLL] |
| | | HLA-C04:63 [0.163; AFEKMSVLL] |
| | | HLA-C04:64 [0.163; AFEKMSVLL] |
| | | HLA-C04:65 [0.163; AFEKMSVLL] |
| | | HLA-C04:66 [0.163; AFEKMSVLL] |
| | | HLA-C04:67 [0.163; AFEKMSVLL] |
| | | HLA-C04:69 [0.203; KMSVLLSV] |
| | | HLA-C04:70 [0.163; AFEKMSVLL] |

* HLA binding affinity is scored by 1-log50k; EKMVSLL (bold) with three upstream and three downstream amino acids were included in scoring. Records were excluded when amino acid length in EKMVSLL is shorter than 6.

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Declaration of Competing Interest

None.

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- Heng Wang^a, Gangning Wu^b, Yan Yang^c, Feng Lian^a, Song Xue^{a,*}
^a *Shanghai Jiao Tong University School of Medicine Affiliated Renji Hospital, China*
^b *Shanghai Jiao Tong University College of Basic Medicine, China*
^c *Shanghai Jiao Tong University School of Medicine Affiliated Ninth People's Hospital, China*
- * Corresponding author.
E-mail address: xuesong64@163.com (S. Xue).