

SHORT COMMUNICATION

Transcriptomic signature of CD4-expressing T-cell abundance developed in healthy peripheral blood predicts strong anti-retroviral therapeutic response in HIV-1: A retrospective and proof-of-concept study

Supplementary File
Table S1. Subject demographics by gene-expression cohort

| | Discovery population | Application population |
|---------------|----------------------|------------------------|
| n | 340 | 24 |
| Age (mean±SD) | 41.79±12.96 | 36.79±10.22 |
| Sex (%) | | |
| Female | 251 (73.8) | 0 (0.0) |
| Male | 89 (26.2) | 24 (100.0) |
| Race (%) | | |
| Caucasian | 318 (93.5) | 7 (29.2) |
| Other | 22 (6.5) | 17 (70.8) |

Abbreviation: SD: Standard deviation.

Table S2. Performance of final LASSO model in a 10-fold cross-validation experiment

| | RMSE | RMSE (original unit) | Pearson's correlation | R ² |
|---------|-------|-------------------------|--------------------------|----------------|
| Fold_1 | 0.070 | 0.163 | 0.936 | 0.876 |
| Fold_2 | 0.072 | 0.167 | 0.878 | 0.771 |
| Fold_3 | 0.077 | 0.179 | 0.833 | 0.693 |
| Fold_4 | 0.083 | 0.193 | 0.890 | 0.792 |
| Fold_5 | 0.071 | 0.165 | 0.920 | 0.846 |
| Fold_6 | 0.076 | 0.176 | 0.870 | 0.757 |
| Fold_7 | 0.069 | 0.159 | 0.889 | 0.791 |
| Fold_8 | 0.069 | 0.161 | 0.915 | 0.837 |
| Fold_9 | 0.072 | 0.168 | 0.867 | 0.751 |
| Fold_10 | 0.070 | 0.162 | 0.903 | 0.815 |
| Avg | 0.073 | 0.169 | 0.890 | 0.793 |

Abbreviations: RMSE: Root mean-squared error; LASSO: Least Absolute Shrinkage and Selection Operator.

Table S3. Gene list with non-zero LASSO coefficients comprising the CD4T gene signature

| Gene | Coefficient | Association with CD4T abundance |
|-----------------|-------------|---------------------------------|
| <i>ABCA1</i> | -0.00264 | Negative |
| <i>ACLY</i> | -0.00133 | Negative |
| <i>AHDC1</i> | -0.00202 | Negative |
| <i>ARMETL1</i> | -0.00136 | Negative |
| <i>BRD8</i> | -0.00603 | Negative |
| <i>BRPF1</i> | -0.00283 | Negative |
| <i>C21orf63</i> | -0.00311 | Negative |
| <i>C21orf70</i> | -0.00671 | Negative |
| <i>CAT</i> | -0.00197 | Negative |
| <i>CBWD5</i> | -0.00142 | Negative |
| <i>CCDC28A</i> | -0.00202 | Negative |
| <i>CCDC55</i> | -0.00535 | Negative |
| <i>CD19</i> | -0.00137 | Negative |
| <i>CD276</i> | -0.00418 | Negative |
| <i>CD8A</i> | -0.01616 | Negative |
| <i>CD8B</i> | -0.00552 | Negative |
| <i>CDC2L6</i> | -0.00597 | Negative |
| <i>CDCA7L</i> | -0.00574 | Negative |
| <i>CMTM2</i> | -0.00254 | Negative |
| <i>CPNE2</i> | -0.00758 | Negative |
| <i>CR2</i> | -0.00743 | Negative |
| <i>CSF3R</i> | -0.00167 | Negative |
| <i>DENND3</i> | -0.00380 | Negative |
| <i>DGKQ</i> | -0.00594 | Negative |
| <i>DKK3</i> | -0.00614 | Negative |
| <i>DNAJC1</i> | -0.00330 | Negative |
| <i>DNHD2</i> | -0.00413 | Negative |
| <i>EIF2C4</i> | -0.01041 | Negative |
| <i>EIF4G3</i> | -0.00319 | Negative |
| <i>FAM33A</i> | -0.00189 | Negative |
| <i>FLJ20712</i> | -0.00393 | Negative |
| <i>FLJ45445</i> | -0.00211 | Negative |
| <i>GAB3</i> | -0.00823 | Negative |
| <i>GBE1</i> | -0.00643 | Negative |
| <i>GCM1</i> | -0.00870 | Negative |
| <i>HCG27</i> | -0.00204 | Negative |
| <i>HCLSI</i> | -0.00807 | Negative |
| <i>HOXC4</i> | -0.00532 | Negative |
| <i>HTR3A</i> | -0.00348 | Negative |
| <i>INPP4A</i> | -0.00649 | Negative |
| <i>KIAA1271</i> | -0.00650 | Negative |

Table S3. (Continued)

| Gene | Coefficient | Association with CD4T abundance |
|------------------|-------------|---------------------------------|
| <i>KIF15</i> | -0.00894 | Negative |
| <i>KLRC3</i> | -0.00361 | Negative |
| <i>LAG3</i> | -0.00207 | Negative |
| <i>LIPC</i> | -0.00239 | Negative |
| <i>LOC221136</i> | -0.01179 | Negative |
| <i>LOC401089</i> | -0.00565 | Negative |
| <i>LOC642183</i> | -0.00477 | Negative |
| <i>LOC642756</i> | -0.00450 | Negative |
| <i>LOC643492</i> | -0.00264 | Negative |
| <i>LOC644033</i> | -0.00228 | Negative |
| <i>LOC644644</i> | -0.00357 | Negative |
| <i>LOC645636</i> | -0.00173 | Negative |
| <i>LOC646494</i> | -0.00572 | Negative |
| <i>LOC652264</i> | -0.00242 | Negative |
| <i>LOC653113</i> | -0.00171 | Negative |
| <i>LOC653303</i> | -0.00146 | Negative |
| <i>MYO9B</i> | -0.00127 | Negative |
| <i>NDN</i> | -0.00417 | Negative |
| <i>NFKBIL2</i> | -0.00643 | Negative |
| <i>NSUN7</i> | -0.00157 | Negative |
| <i>NUDT2</i> | -0.00177 | Negative |
| <i>OPRK1</i> | -0.00800 | Negative |
| <i>OTOF</i> | -0.00608 | Negative |
| <i>PDE6G</i> | -0.00581 | Negative |
| <i>PDGFC</i> | -0.00206 | Negative |
| <i>PIGL</i> | -0.00136 | Negative |
| <i>PIGT</i> | -0.00238 | Negative |
| <i>PLCG2</i> | -0.00682 | Negative |
| <i>PMCHL2</i> | -0.00905 | Negative |
| <i>POLA1</i> | -0.00385 | Negative |
| <i>POLR3D</i> | -0.00719 | Negative |
| <i>PPP1R15B</i> | -0.00230 | Negative |
| <i>PPP2R5E</i> | -0.00466 | Negative |
| <i>PRR13</i> | -0.00661 | Negative |
| <i>RAPGEFL1</i> | -0.00153 | Negative |
| <i>RUNDCL</i> | -0.00605 | Negative |
| <i>SCARB1</i> | -0.00318 | Negative |
| <i>SLC25A4</i> | -0.00139 | Negative |
| <i>SLC29A3</i> | -0.00242 | Negative |
| <i>SLC45A4</i> | -0.00384 | Negative |
| <i>SLC1</i> | -0.00272 | Negative |
| <i>SUCNR1</i> | -0.00184 | Negative |

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Table S3. (Continued)

| Gene | Coefficient | Association with CD4T abundance |
|---------------------|-------------|---------------------------------|
| <i>TBC1D5</i> | -0.00385 | Negative |
| <i>TCEA2</i> | -0.00388 | Negative |
| <i>TLR10</i> | -0.00511 | Negative |
| <i>TMEM16H</i> | -0.00332 | Negative |
| <i>TMEM35</i> | -0.00776 | Negative |
| <i>TNFRSF6B</i> | -0.00607 | Negative |
| <i>TRAF4</i> | -0.00328 | Negative |
| <i>TTC16</i> | -0.00792 | Negative |
| <i>UBE2J1</i> | -0.00174 | Negative |
| <i>ZNF285A</i> | -0.00345 | Negative |
| <i>ACOT1</i> | 0.00376 | Positive |
| <i>ACPL2</i> | 0.00266 | Positive |
| <i>AK2</i> | 0.00827 | Positive |
| <i>AP3M2</i> | 0.00472 | Positive |
| <i>ARHGAP11A</i> | 0.00228 | Positive |
| <i>ARMCX4</i> | 0.00464 | Positive |
| <i>ASAHL</i> | 0.00354 | Positive |
| <i>BAG3</i> | 0.00960 | Positive |
| <i>BGLAP</i> | 0.00150 | Positive |
| <i>BIRC3</i> | 0.00352 | Positive |
| <i>BRD1</i> | 0.00622 | Positive |
| <i>C16orf30</i> | 0.00262 | Positive |
| <i>C19orf21</i> | 0.00536 | Positive |
| <i>C6orf105</i> | 0.00178 | Positive |
| <i>CACNA1I</i> | 0.01344 | Positive |
| <i>CAPN2</i> | 0.00293 | Positive |
| <i>CD28</i> | 0.01101 | Positive |
| <i>CD36</i> | 0.00252 | Positive |
| <i>CD40LG</i> | 0.01145 | Positive |
| <i>CDC14A</i> | 0.00663 | Positive |
| <i>CEP135</i> | 0.00244 | Positive |
| <i>CETN2</i> | 0.00175 | Positive |
| <i>CHCHD6</i> | 0.00347 | Positive |
| <i>CHCHD7</i> | 0.00279 | Positive |
| <i>COMM6</i> | 0.00704 | Positive |
| <i>CTLA4</i> | 0.01113 | Positive |
| <i>DKFZp434N035</i> | 0.00423 | Positive |
| <i>DPP7</i> | 0.00262 | Positive |
| <i>EDA</i> | 0.01913 | Positive |
| <i>EPB41L2</i> | 0.00334 | Positive |
| <i>EPHA4</i> | 0.00814 | Positive |
| <i>FBXO22</i> | 0.00427 | Positive |

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Table S3. (Continued)

| Gene | Coefficient | Association with CD4T abundance |
|------------------|-------------|---------------------------------|
| <i>FHIT</i> | 0.00202 | Positive |
| <i>FLJ22374</i> | 0.00464 | Positive |
| <i>GATA3</i> | 0.00543 | Positive |
| <i>GOLGA8A</i> | 0.00156 | Positive |
| <i>GPR153</i> | 0.00745 | Positive |
| <i>HMG A2</i> | 0.00254 | Positive |
| <i>ICOS</i> | 0.00636 | Positive |
| <i>IL11RA</i> | 0.00734 | Positive |
| <i>IL7R</i> | 0.00358 | Positive |
| <i>IMPAD1</i> | 0.00670 | Positive |
| <i>INPP4B</i> | 0.00238 | Positive |
| <i>ISG20L1</i> | 0.00169 | Positive |
| <i>KLF11</i> | 0.00453 | Positive |
| <i>KRTAP10-2</i> | 0.00157 | Positive |
| <i>LEFTY1</i> | 0.00422 | Positive |
| <i>LOC158830</i> | 0.00588 | Positive |
| <i>LOC283683</i> | 0.00260 | Positive |
| <i>LOC441151</i> | 0.00571 | Positive |
| <i>LOC642210</i> | 0.00320 | Positive |
| <i>LOC642576</i> | 0.00509 | Positive |
| <i>LOC644297</i> | 0.00308 | Positive |
| <i>LOC644634</i> | 0.00541 | Positive |
| <i>LOC644919</i> | 0.00498 | Positive |
| <i>LOC645052</i> | 0.00274 | Positive |
| <i>LOC645464</i> | 0.00147 | Positive |
| <i>LOC645968</i> | 0.00424 | Positive |
| <i>LOC649270</i> | 0.00556 | Positive |
| <i>LOC652589</i> | 0.00474 | Positive |
| <i>LOC652659</i> | 0.00943 | Positive |
| <i>LOC653316</i> | 0.00532 | Positive |
| <i>LOC653663</i> | 0.00653 | Positive |
| <i>LRIG1</i> | 0.00217 | Positive |
| <i>MEIS3</i> | 0.00132 | Positive |
| <i>MFHAS1</i> | 0.00254 | Positive |
| <i>MGC35361</i> | 0.00401 | Positive |
| <i>MMP14</i> | 0.00416 | Positive |
| <i>MORC2</i> | 0.00814 | Positive |
| <i>MYH4</i> | 0.00258 | Positive |
| <i>NOX1</i> | 0.00152 | Positive |
| <i>NOXA1</i> | 0.00499 | Positive |
| <i>NPCDR1</i> | 0.00381 | Positive |
| <i>NRBP2</i> | 0.00160 | Positive |

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Table S3. (Continued)

| Gene | Coefficient | Association with CD4T abundance |
|------------------|-------------|---------------------------------|
| <i>NT5DC3</i> | 0.00739 | Positive |
| <i>NTSR1</i> | 0.00180 | Positive |
| <i>OR1L8</i> | 0.00763 | Positive |
| <i>OR2H2</i> | 0.00814 | Positive |
| <i>P2RX1</i> | 0.00456 | Positive |
| <i>P2RY6</i> | 0.01526 | Positive |
| <i>PANK1</i> | 0.01224 | Positive |
| <i>PBX4</i> | 0.00320 | Positive |
| <i>PLAG1</i> | 0.00198 | Positive |
| <i>POLE2</i> | 0.00179 | Positive |
| <i>PRIM2A</i> | 0.00135 | Positive |
| <i>PRKCA</i> | 0.00332 | Positive |
| <i>PRKY</i> | 0.00401 | Positive |
| <i>PURA</i> | 0.00602 | Positive |
| <i>RPL23AP13</i> | 0.00126 | Positive |
| <i>RPL35</i> | 0.00284 | Positive |
| <i>SCRN2</i> | 0.00433 | Positive |
| <i>SLC22A17</i> | 0.00153 | Positive |
| <i>SLC35E2</i> | 0.00751 | Positive |
| <i>SNAPC3</i> | 0.00485 | Positive |

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Table S3. (Continued)

| Gene | Coefficient | Association with CD4T abundance |
|-----------------|-------------|---------------------------------|
| <i>SPOCK2</i> | 0.00747 | Positive |
| <i>SUSD3</i> | 0.00523 | Positive |
| <i>TACSTD2</i> | 0.00224 | Positive |
| <i>TBC1D4</i> | 0.01202 | Positive |
| <i>TCEA3</i> | 0.00365 | Positive |
| <i>TIAM1</i> | 0.00537 | Positive |
| <i>TMED4</i> | 0.00463 | Positive |
| <i>TNFRSF25</i> | 0.00510 | Positive |
| <i>TOE1</i> | 0.00514 | Positive |
| <i>TRAF6</i> | 0.00149 | Positive |
| <i>TRAT1</i> | 0.00170 | Positive |
| <i>UBASH3A</i> | 0.00401 | Positive |
| <i>USP37</i> | 0.00471 | Positive |
| <i>VPS72</i> | 0.00355 | Positive |
| <i>ZBTB7B</i> | 0.00160 | Positive |
| <i>ZNF179</i> | 0.00616 | Positive |
| <i>ZNF37A</i> | 0.00624 | Positive |
| <i>ZNF432</i> | 0.00411 | Positive |
| <i>ZNF623</i> | 0.00820 | Positive |
| <i>ZNF71</i> | 0.00212 | Positive |

Abbreviation: LASSO: Least Absolute Shrinkage and Selection Operator.