

1 **Supplemental Methods:**

2 T cell counts were performed using Tritest Reagent with Trucount Tubes and analyzed on a
3 FACSCalibur platform (BD Biosciences).

4 Viral Loads: Plasma RNA was extracted using the Abbott Sample Preparation System, amplified
5 using Real Time HIV-1 Amplification Reagent Kits (Abbott Molecular, Des Plaines, IL), and run
6 on the M2000 Realtime System (Abbott) with a lower limit of detection of 40 copies/ml.

7 QVOA: rCD4 (CD4+/CD69-/CD25-/HLA-DR-) were isolated from PBMCs using two steps of
8 negative-selection, plated at limiting dilution, and activated with PHA and γ -irradiated allogenic
9 PBMCs. If the number of CD4+ cells were low after the first isolation, total CD4 counts were
10 plated. Released virus was expanded using MOLT-4 CCR5+ (NIH AIDS reagent program).
11 Culture supernatants were examined for the presence of actively growing virus using a p24
12 ELISA (PerkinElmer) after 14-21 days (1). Infectious units per million rCD4 cells (IUPM) were
13 calculated using IUPMStats (2). If all culture wells were negative for viral outgrowth, the median
14 posterior estimate of infected cell frequency was used.

15 Total HIV DNA was quantified by quantitative PCR (qPCR) for a segment of the gag gene, using
16 a previously published protocol adapted for qPCR (3, 4). DNA was extracted from rCD4 cells
17 using the AllPrep DNA/RNA minikit (Qiagen, Valencia, CA) and amplified by qPCR using the
18 TaqMan Fast Advanced Master Mix (Thermo Fisher Scientific, Waltham, MA) and gag targeted
19 primers with the following cycling conditions: 2 minute UNG incubation at 50°C, 20 second
20 enzyme activation at 95°C, 50 cycles of a 1 second denaturation at 95°C followed by 20
21 seconds of annealing/extension at 60°C. The host cell RNase P/MRP 30-kDa-subunit gene
22 (RPP30) was amplified using the same conditions with respective primers. *gag* copy numbers

23 were calculated as the mean of triplicate qPCR measurements and were normalized to 1x10⁶
24 rCD4, as determined by RPP30 (which appears in 2 copies/cell) (4).

25 HIV-1 subtyping was determined from historical data or performed by next-generation amplicon
26 sequencing of viral RNA from wells positive for p24 in the QVOA, using reverse transcriptase
27 (RT) and gp41-targeted primers, as described previously (5). Subtypes for reverse transcriptase
28 (RT) and gp41 were determined phylogenetically, or, if phylogenetic analyses were unclear,
29 using the NCBI HIV subtyping tool
30 (<https://www.ncbi.nlm.nih.gov/projects/genotyping/formpage.cgi>). Individuals with both RT and
31 gp41 regions identified as the same subtype were classified with a pure subtype, and those with
32 divergent calls were classified as recombinants.

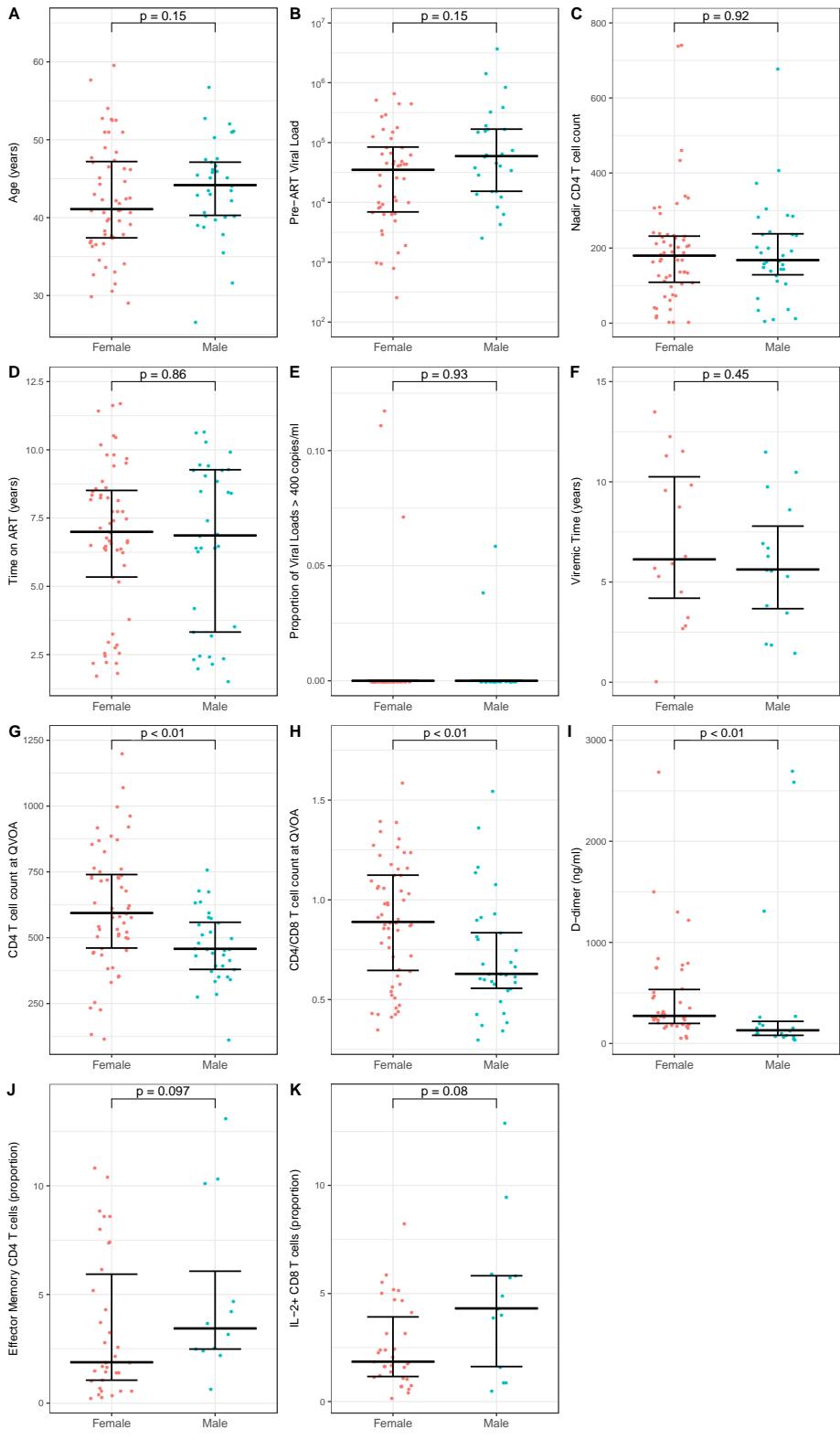
33 Soluble Immune Analytes: An 8-plex custom Human Ultra-Sensitive kit was used to quantify:
34 IFN-2α (lower limit of quantification (LLOQ) 4.88pg/ml), IL10 (LLOQ 0.62pg/ml), IL15 (LLOQ
35 1.5pg/ml), IL6 (LLOQ 0.28pg/ml), IL7 (LLOQ 0.42pg/ml), MIG/CXCL9 (78.12pg/ml LLOQ),
36 MIP3 α /CCL19 (19.54pg/ml LLOQ), and MIP3 β /CCL20 (LLOQ 4.88pg/ml). Additional analytes
37 were measured individually: D-dimer (LLOQ 47.5ng/ml), sCD14 (LLOQ 1.25ng/ml), IP10 (LLOQ
38 7.8pg/ml), CRP (LLOQ 78.0ng/ml), and CCL21 (LLOQ 32.76pg/ml). Plates were imaged using
39 the Sector Imager 2400A platform.

40 Flow Cytometry: Antibodies and fluorophores (clone, manufacturer) used were as follows: CD3
41 APC-H7 (Sk7, BD Biosciences), CD4 APC-R700 (RPA-T4, BD Biosciences), CD45RO PerCP-
42 Cy5.5 (UCHL1, BD Biosciences), CD28 PE-CF594 (CD28-2, BD Biosciences), CCR7 BV711
43 (G043H7, Biolegend), CD57 BB515 (NK-1, BD Biosciences), HLA-DR BV510 (G46-6, BD
44 Biosciences), CD38 BV605 (HB7, BD Biosciences), CD69 BV421 (FN50 BD Biosciences), Ki-67
45 eFluor660 (20Raj1, eBioscienes), FoxP3 PE (PCH101, eBiosciences, CD25 BV786 (M-A251,
46 BD Biosciences); and one to assess T cell effector function, including markers of T cell

47 exhaustion (CD3, CD4, CD8 BB515 (RPA-T8, BD Biosciences), CD107a AF647 (H4A3, BD
48 Biosciences), Perforin BV421 (B-D48, Biolegend), IFN γ PerCP-Cy5.5 (B27, BD Biosciences),
49 TNF BV785 (Mab11, Biolegend), IL2 BV510 (5344-111, BD Biosciences), Tim3 PE (344823,
50 R&D Systems), CTLA4 PE-CF594 (BNI3, BD Biosciences), PD1 BV605 (EH12-1, BD
51 Biosciences). Both panels included a viability dye (LIVE/DEAD Fixable Blue Dead Cell Stain Kit,
52 Invitrogen).

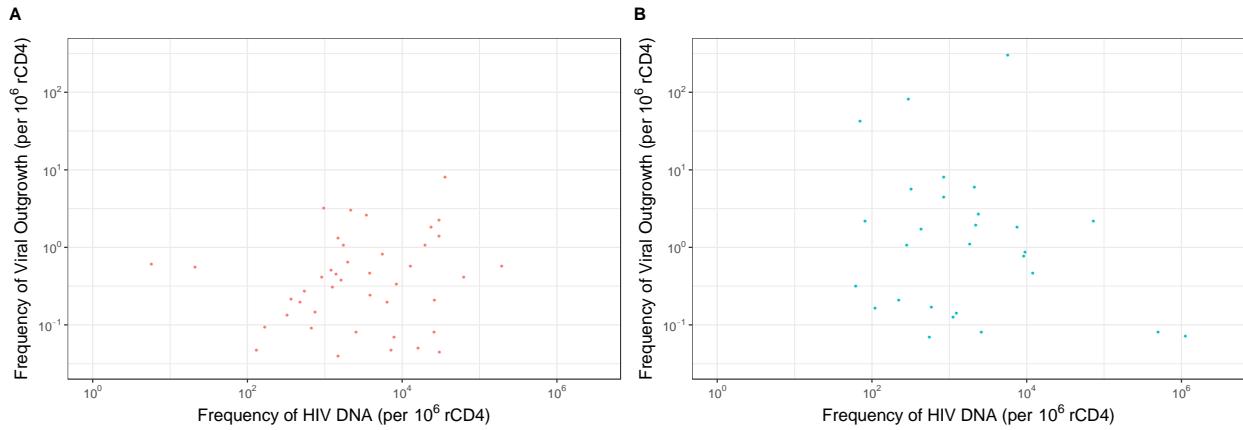
53 PBMCs were thawed, resuspended at 1x10⁶ viable cells/ml (trypan blue) in R10 (RPMI 1640
54 media supplemented with: 10% heat-inactivated FBS, 10U/ml penicillin, 10 μ g/ml streptomycin,
55 250ng/ml amphotericin B, and 2mM L-Glutamine) and rested overnight under normal growth
56 conditions (37°C, 5% CO₂, humidified atmosphere). The following morning 100 μ l cell
57 suspension was plated in a round bottom 96-well plate for stimulation and/or staining. To assess
58 cytokine production, one well per participant was stimulated with 1ng/ml phorbol-12-myristate-
59 13-acetate (PMA) and 1 μ g/ml ionomycin and one well with vehicle (0.1% DMSO), both for 3
60 hours under normal growth conditions and in the presence of 10 μ g/ml Brefeldin A, 0.7 μ g/ml
61 monensin, and anti-CD107a antibody. Cells for CD4 T cell phenotyping were left unstimulated.
62 All samples were then washed with PBS, stained with viability dye for 30 minutes at room
63 temperature, and washed twice more. Cells were then stained with fluorochrome-labeled
64 antibodies against surface antigens in the presence of 2% FBS in PBS for 30 minutes at room
65 temperature. Excess surface antibody was removed by washing with 2% FBS in PBS. Samples
66 for the assessment of T cell phenotype were permeabilized using the eBioscience
67 fixation/permeabilization solution for Treg identification, while samples for the assessment of
68 effector functions were permeabilized with the BD Cytofix/Cytoperm solution (BD Biosciences).
69 Cells were washed in the appropriate permeabilization wash buffer and stained with antibodies
70 against intracellular antigens.

71 Samples were acquired using an LSRII flow cytometer (BD Systems) and data analysis
72 performed using FlowJo (version 10·6·1, Becton Dickinson). Representative gating images are
73 shown in Supplementary Figure 3.

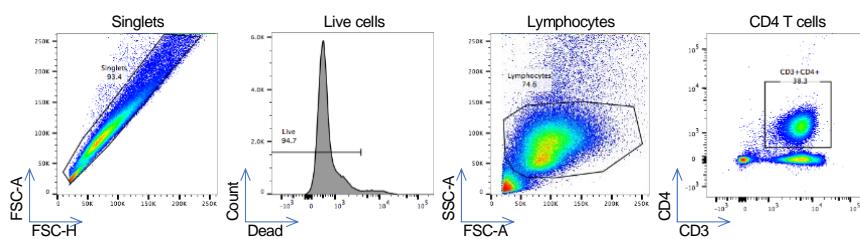
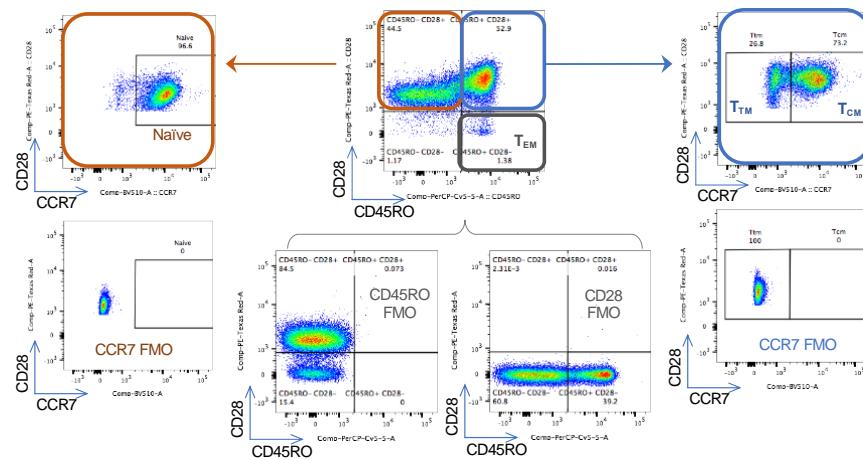
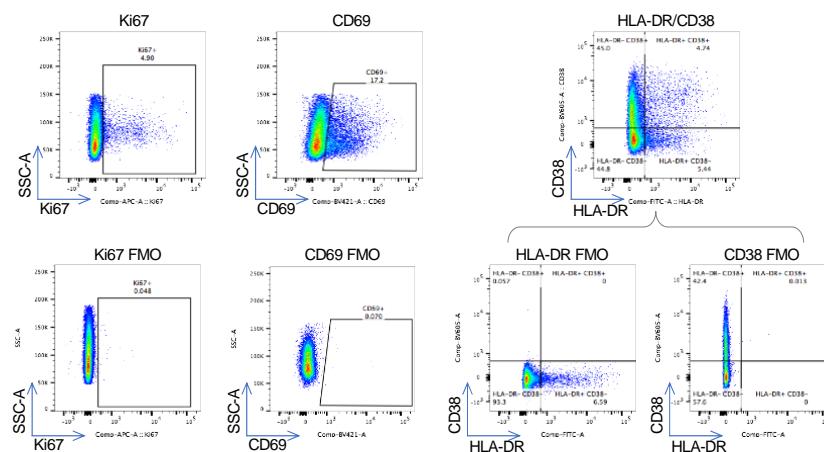
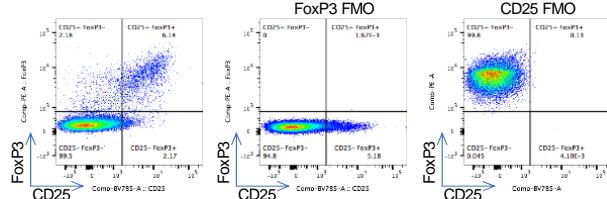


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75 **Supplemental Figure 1:** Participant clinical and immunological characteristics, by sex.

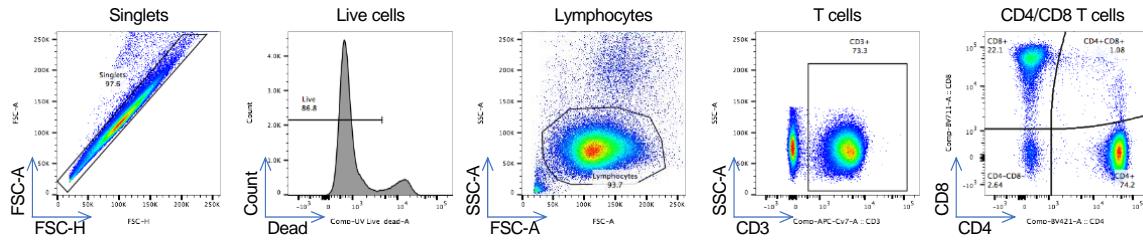


76
 77 **Supplemental Figure 2:** No correlation between QVOA and qPCR estimates of the frequency
 78 of persistently infected resting CD4 T cells (rCD4), in either females (**A**) or males (**B**). QVOA:
 79 Quantitative Viral Outgrowth Assay (QVOA); qPCR: quantitative PCR for total HIV DNA.

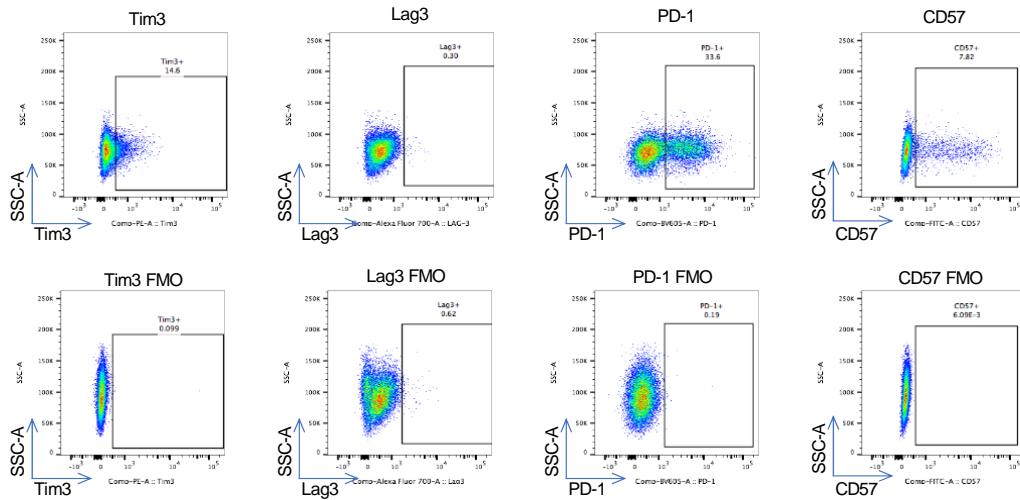
Supplemental Figure 3: Flow cytometry gating strategy and representative plots**A. Flow cytometry panel 1: CD4 T cell subsets and activation status****CD4 T Cell Identification****CD4 T Cell Memory Subsets****CD4 T Cell Activation****Regulatory T cells (Treg)**

B. Flow cytometry panel 2: CD4 and CD8 T cell immune exhaustion (using unstimulated cells)

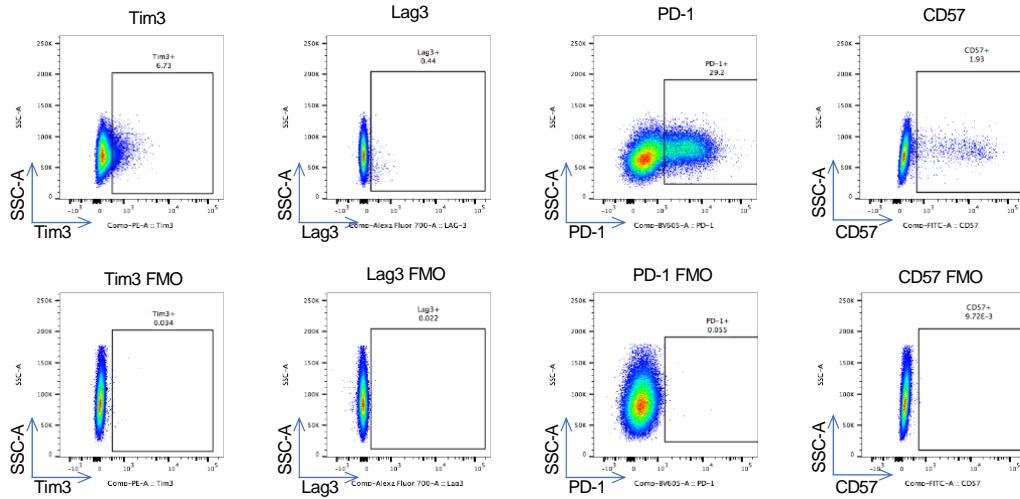
CD4 and CD8 T Cell Identification



CD8 T Cell Exhaustion

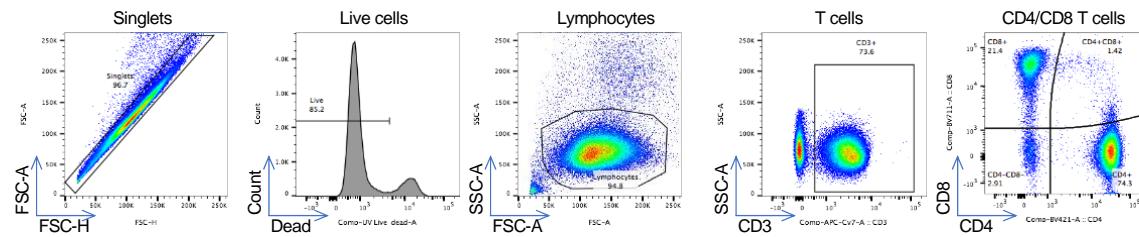


CD4 T Cell Exhaustion

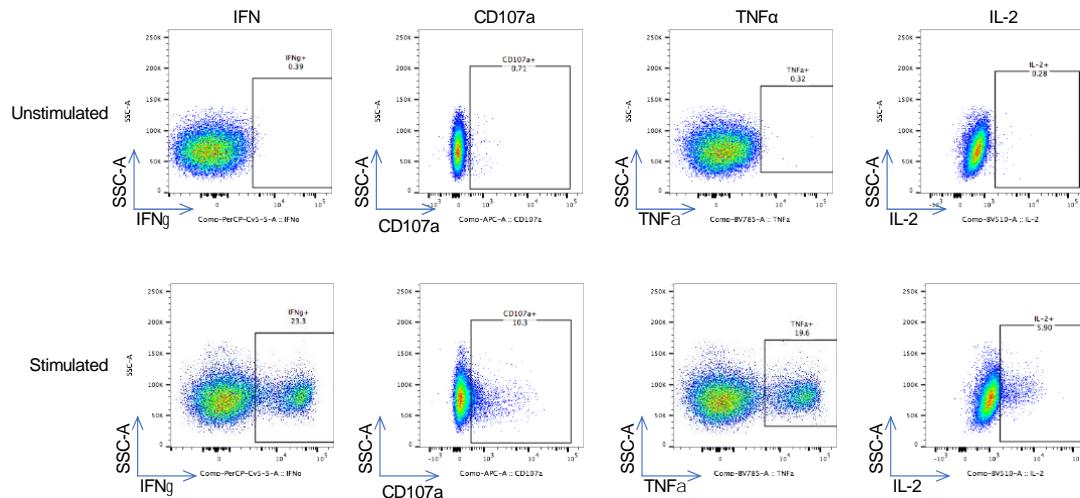


C. Flow cytometry panel 2: CD4 and CD8 T cell immune function

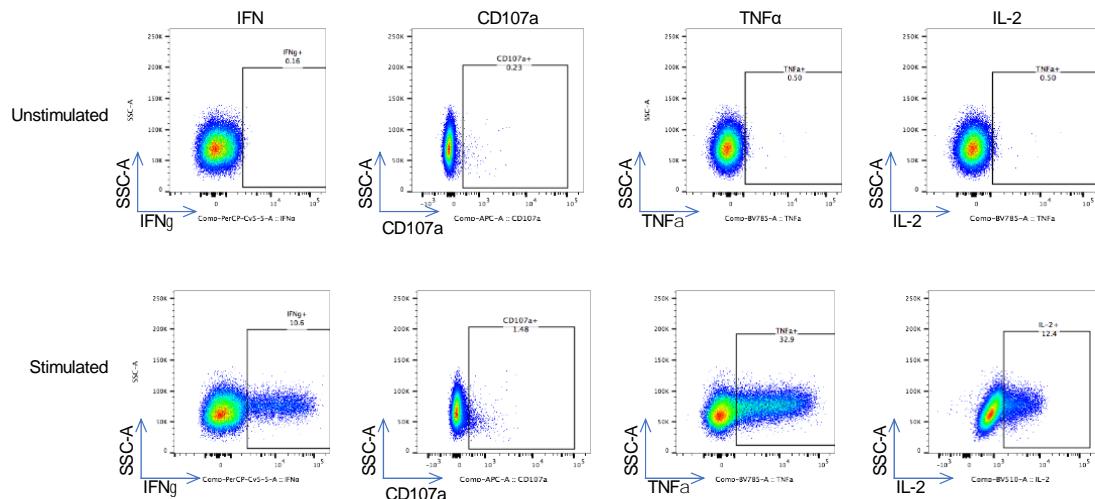
CD4 and CD8 T Cell Identification (3hrs PMA/iono stimulation)



CD8 T Cell Effector Function



CD4 T Cell Effector Function



87 **Supplementary Table 1.** Univariate regression analysis of immunological variables on reservoir
 88 outgrowth (IUPM), including both males and females.

	Immune Characteristic	Median (IQR)	Regression Coefficient	P-value
Soluble Immune Analytes (\log_{10} pg/ml, unless stated otherwise)	IL-7	0.6 (0.4, 0.8)	0.20	0.44
	IL-15	0.5 (0.4, 0.5)	-0.31	0.61
	CRP (\log_{10} ng/ml)	3.2 (2.7, 3.6)	-0.05	0.70
	D-dimer (\log_{10} ng/ml)	2.4 (2.2, 2.7)	-0.07	0.72
	IL-10 (binary)	0.3 (0.2, 0.6)	0.00	1.00
	IL-6	-0.1 (-0.3, 0.0)	-0.15	0.63
	IP10	2.0 (1.9, 2.2)	-0.00	0.99
	CCL21	2.3 (1.5, 2.7)	0.07	0.63
	MIG	2.6 (2.4, 2.8)	-0.11	0.70
	MIP-3 α (binary)	0.8 (0.7, 1.0)	0.07	0.67
	MIP-3 β	1.9 (1.8, 2.0)	-0.17	0.74
T cell subsets (proportion of CD3+ cells)	CD4+	51.7 (42.5, 60.4)	-0.00	0.72
	CD8+	41.1 (33.3, 51.9)	0.00	0.57
CD4 T cell memory subsets (proportion of CD4 T cells)	Naïve (CD28+/CD45RO-/CCR7+)	45.3 (32.3, 54.9)	-0.00	0.69
	Central (CD28+/CD45RO+/CCR7+)	34.0 (24.7, 42.6)	0.00	0.80
	Effector (CD28-/CD45RO+/CCR7-)	2.5 (1.4, 5.9)	0.02	0.49
	Transitional (CD28+/CD45RO+/CCR7-)	7.7 (5.5, 13.0)	0.02	0.22
CD4 T cell activation (proportion of CD4 T cells)	CD69	15.8 (11.6, 19.6)	0.01	0.39
	HLA-DR+/CD38+	2.3 (1.5, 3.4)	0.00	0.97
	CD69+/Ki-67+	0.3 (0.2, 0.4)	0.41	0.26
	CD69- /Ki-67+	1.7 (1.1, 2.2)	0.03	0.62

	Ki-67+	1.9 (1.5, 2.6)	0.03	0.55
	T regulatory cells	1.9 (1.4, 2.7)	0.06	0.46
T cell exhaustion: summary subsets (proportion of parent)	CD57+ CD4 T cells	2.8 (1.6, 5.6)	0.04	0.18
	Lag-3+ CD4 T cells	1.6 (1.0, 2.8)	-0.04	0.54
	PD-1+ CD4 T cells	27.8 (19.5, 33.4)	0.01	0.12
	Tim-3+ CD4 T cells	2.7 (1.8, 3.9)	-0.03	0.47
	CD57+ CD8 T cells	27.6 (15.7, 35.4)	0.01	0.13
	Lag-3+ CD8 T cells	0.5 (0.3, 0.7)	0.30	0.25
	PD-1+ CD8 T cells	21.1 (14.9, 25.8)	-0.01	0.63
	Tim-3+ CD8 T cells	10.4 (6.7, 15.3)	0.01	0.46
T cell exhaustion: specific subsets (proportion of parent)	CD57-/Lag3-/PD1-/Tim3- CD4 T cells	68.3 (61.9, 75.5)	-0.01	0.13
	CD57-/Lag3-/PD1-/Tim3+ CD4 T cells	1.4 (0.7, 2.0)	-0.07	0.21
	CD57-/Lag3-/PD1+/Tim3- CD4 T cells	22.2 (16.8, 27.8)	0.01	0.14
	CD57-/Lag3-/PD1+/Tim3+ CD4 T cells	0.7 (0.5, 1.2)	0.05	0.75
	CD57-/Lag3+/PD1-/Tim3- CD4 T cells	1.4 (0.7, 2.2)	-0.03	0.58
	CD57-/Lag3+/PD1-/Tim3+ CD4 T cells	0.01 (0.00, 0.01)	-9.13	0.41
	CD57-/Lag3+/PD1+/Tim3- CD4 T cells	0.2 (0.1, 0.3)	-0.12	0.86
	CD57-/Lag3+/PD1+/Tim3+ CD4 T cells	0.02 (0.01, 0.03)	-0.47	0.62
	CD57+/Lag3-/PD1-/Tim3- CD4 T cells	0.5 (0.3, 1.4)	0.02	0.78
	CD57+/Lag3-/PD1-/Tim3+ CD4 T cells	0.04 (0.02, 0.08)	0.05	0.93
	CD57+/Lag3-/PD1+/Tim3- CD4 T cells	1.6 (1.1, 4.0)	0.06	0.10
	CD57+/Lag3-/PD1+/Tim3+ CD4 T cells	0.08 (0.03, 0.21)	0.12	0.50
	CD57+/Lag3+/PD1-/Tim3- CD4 T cells	0.01 (0.00, 0.02)	-4.50	0.18
	CD57+/Lag3+/PD1-/Tim3+ CD4 T cells	0.00 (0.00, 0.00)	-6.93	0.79
	CD57+/Lag3+/PD1+/Tim3- CD4 T cells	0.01 (0.00, 0.02)	-0.11	0.97

	CD57+/Lag3+/PD1+/Tim3+ CD4 T cells	0.00 (0.00, 0.01)	-4.06	0.81
	CD57-/Lag3-/PD1-/Tim3- CD8 T cells	51.2 (41.9, 61.0)	-0.01	0.27
	CD57-/Lag3-/PD1-/Tim3+ CD8 T cells	5.3 (2.8, 9.8)	0.00	0.91
	CD57-/Lag3-/PD1+/Tim3- CD8 T cells	12.2 (9.1, 16.3)	-0.03	0.12
	CD57-/Lag3-/PD1+/Tim3+ CD8 T cells	0.8 (0.5, 1.5)	0.11	0.32
	CD57-/Lag3+/PD1-/Tim3- CD8 T cells	0.2 (0.1, 0.3)	0.21	0.62
	CD57-/Lag3+/PD1-/Tim3+ CD8 T cells	0.03 (0.01, 0.04)	3.05	0.40
	CD57-/Lag3+/PD1+/Tim3- CD8 T cells	0.09 (0.05, 0.17)	0.85	0.33
	CD57-/Lag3+/PD1+/Tim3+ CD8 T cells	0.01 (0.00, 0.02)	12.78	0.07
	CD57+/Lag3-/PD1-/Tim3- CD8 T cells	17.1 (9.6, 22.9)	0.01	0.13
	CD57+/Lag3-/PD1-/Tim3+ CD8 T cells	2.0 (0.8, 3.3)	0.02	0.48
	CD57+/Lag3-/PD1+/Tim3- CD8 T cells	5.2 (3.0, 7.5)	0.02	0.42
	CD57+/Lag3-/PD1+/Tim3+ CD8 T cells	0.5 (0.3, 1.3)	0.05	0.57
	CD57+/Lag3+/PD1-/Tim3- CD8 T cells	0.05 (0.02, 0.08)	2.79	0.09
	CD57+/Lag3+/PD1-/Tim3+ CD8 T cells	0.00 (0.00, 0.02)	11.23	0.17
	CD57+/Lag3+/PD1+/Tim3- CD8 T cells	0.04 (0.02, 0.06)	2.19	0.31
	CD57+/Lag3+/PD1+/Tim3+ CD8 T cells	0.00 (0.00, 0.01)	8.11	0.36
T cell effector functions: summary subsets (proportion of parent)	CD107+ CD4 T cells	4.4 (2.4, 9.3)	0.02	0.22
	IFN γ + CD4 T cells	6.7 (2.9, 10.6)	0.02	0.13
	IL-2+ CD4 T cells	8.4 (4.7, 15.7)	0.02	0.23
	TNF+ CD4 T cells	20.1 (9.5, 33.5)	0.01	0.22
	CD107+ CD8 T cells	6.9 (3.7, 11.5)	0.00	0.86
	IFN γ + CD8 T cells	15.4 (8.1, 22.8)	0.00	0.55
	IL-2+ CD8 T cells	2.3 (1.2, 4.8)	0.00	0.89
	TNF+ CD8 T cells	9.0 (3.3, 19.5)	0.01	0.45

T cell effector functions: specific subsets (proportion of parent)	CD107-/IFN γ -/IL2-/TNF- CD4 T cells	71.7 (59.9, 77.7)	-0.01	0.11
	CD107-/IFN γ -/IL2-/TNF+ CD4 T cells	8.9 (4.6, 13.7)	0.01	0.45
	CD107-/IFN γ -/IL2+/TNF- CD4 T cells	1.4 (0.8, 1.9)	0.16	0.15
	CD107-/IFN γ -/IL2+/TNF+ CD4 T cells	5.2 (2.3, 10.0)	0.02	0.43
	CD107-/IFN γ +/IL2-/TNF- CD4 T cells	1.0 (0.4, 1.9)	0.04	0.46
	CD107-/IFN γ +/IL2-/TNF+ CD4 T cells	2.7 (1.2, 4.9)	0.03	0.20
	CD107-/IFN γ +/IL2+/TNF- CD4 T cells	0.08 (0.03, 0.15)	0.30	0.56
	CD107-/IFN γ +/IL2+/TNF+ CD4 T cells	1.3 (0.5, 2.6)	0.06	0.20
	CD107+/IFN γ -/IL2-/TNF- CD4 T cells	3.2 (1.9, 59)	0.02	0.47
	CD107+/IFN γ -/IL2-/TNF+ CD4 T cells	0.3 (0.1, 0.9)	0.29	0.04
	CD107+/IFN γ -/IL2+/TNF- CD4 T cells	0.1 (0.07, 0.3)	-0.00	1.00
	CD107+/IFN γ -/IL2+/TNF+ CD4 T cells	0.1 (0.06, 0.4)	0.27	0.21
	CD107+/IFN α +/IL2-/TNF- CD4 T cells	0.04 (0.02, 0.09)	1.30	0.33
	CD107+/IFN γ +/IL2-/TNF+ CD4 T cells	0.1 (0.04, 0.4)	0.29	0.09
	CD107+/IFN γ +/IL2+/TNF- CD4 T cells	0.01 (0.00, 0.01)	-1.54	0.74
	CD107+/IFN γ +/IL2+/TNF+ CD4 T cells	0.05 (0.02, 0.15)	0.55	0.12
	CD107-/IFN γ -/IL2-/TNF- CD8 T cells	72.8 (66.5, 83.8)	-0.00	0.53
	CD107-/IFN γ -/IL2-/TNF+ CD8 T cells	1.2 (0.8, 2.8)	0.04	0.43
	CD107-/IFN γ -/IL2+/TNF- CD8 T cells	0.8 (0.5, 1.4)	0.09	0.50
	CD107-/IFN γ -/IL2+/TNF+ CD8 T cells	0.4 (0.1, 0.9)	0.04	0.72
	CD107-/IFN γ +/IL2-/TNF- CD8 T cells	5.8 (2.7, 8.6)	0.00	0.95
	CD107-/IFN γ +/IL2-/TNF+ CD8 T cells	4.8 (1.6, 10.1)	0.01	0.54
	CD107-/IFN γ +/IL2+/TNF- CD8 T cells	0.3 (0.1, 0.4)	0.20	0.53
	CD107-/IFN γ +/IL2+/TNF+ CD8 T cells	0.5 (0.2, 1.1)	-0.01	0.91
	CD107+/IFN γ -/IL2-/TNF- CD8 T cells	4.6 (2.5, 7.8)	-0.00	0.86

	CD107+/IFN γ -/IL2-/TNF+ CD8 T cells	0.06 (0.02, 0.22)	0.25	0.62
	CD107+/IFN γ -/IL2+/TNF- CD8 T cells	0.08 (0.05, 0.12)	-0.17	0.43
	CD107+/IFN γ -/IL2+/TNF+ CD8 T cells	0.01 (0.00, 0.03)	-0.29	0.76
	CD107+/IFN γ +/IL2-/TNF- CD8 T cells	0.4 (0.2, 0.8)	0.15	0.42
	CD107+/IFN γ +/IL2-/TNF+ CD8 T cells	0.4 (0.2, 1.9)	0.07	0.25
	CD107+/IFN γ +/IL2+/TNF- CD8 T cells	0.03 (0.01, 0.04)	-0.73	0.66
	CD107+/IFN γ +/IL2+/TNF+ CD8 T cells	0.01 (0.00, 0.16)	-0.03	0.85

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90 **Supplementary Table 2.** Univariate regression analysis of immunological variables on reservoir

91 outgrowth (IUPM), among females only.

	Immune Characteristic	Median (IQR)	Regression Coefficient	P-value
Soluble Immune Analytes (\log_{10} pg/ml, unless otherwise stated)	IL-7	0.6 (0.4, 0.8)	0.47	0.09
	IL-15	0.5 (0.4, 0.5)	-0.29	0.66
	CRP (\log_{10} ng/ml)	3.0 (2.6, 3.6)	-0.09	0.48
	D-dimer (\log_{10} ng/ml)	2.4 (2.3, 2.8)	0.05	0.85
	IL-10 (binary)	0.3 (0.2, 0.6)	-0.16	0.43
	IL-6	-0.1 (-0.3, 0.1)	-0.14	0.67
	IP10	2.0 (1.9, 2.1)	0.50	0.25
	CCL21	2.3 (1.5, 2.6)	-0.07	0.65
	MIG	2.6 (2.4, 2.7)	0.01	0.98
	MIP-3 α (binary)	5.8 (3.7, 10.0)	0.03	0.85
T cell subsets (proportion CD3+ cells)	CD4+	51.7 (42.7, 59.8)	0.01	0.48
	CD8	42.2 (32.8, 51.9)	0.00	0.53
CD4 T cell memory	Naïve (CD28+/CD45RO-/CCR7+)	47.1 (32.6, 58.2)	0.00	0.97

subsets (proportion CD4 T cells)	Central (CD28+/CD45RO+/CCR7+)	33.3 (22.0, 42.6)	0.00	0.88
	Effector (CD28-/CD45RO+/CCR7-)	1.88 (1.1, 5.9)	0.03	0.34
	Transitional (CD28+/CD45RO+/CCR7-)	6.97 (5.6, 11.5)	0.01	0.62
CD4 T cell activation (proportion CD4 T cells)	CD69	15.8 (12.3, 19.3)	-0.01	0.66
	HLA-DR+/CD38+	2.3 (1.6, 3.1)	0.13	0.11
	CD69+/Ki-67+	0.3 (0.2, 0.4)	-0.02	0.94
	CD69- /Ki-67+	1.7 (1.1, 2.2)	0.03	0.57
	Ki-67+	1.9 (1.5, 2.5)	0.03	0.63
	T regulatory cells	1.9 (1.5, 2.7)	0.06	0.51
T cell exhaustion: summary subsets (proportion of parent)	CD57+ CD4 T cells	2.3 (1.3, 5.6)	0.03	0.25
	Lag-3+ CD4 T cells	1.7 (1.0, 2.6)	-0.03	0.57
	PD-1+ CD4 T cells	27.8 (19.5, 33.5)	0.00	0.55
	Tim-3+ CD4 T cells	2.9 (1.7, 4.5)	-0.04	0.36
	CD57+ CD8 T cells	26.1 (16.4, 36.8)	0.00	0.67
	Lag-3+ CD8 T cells	0.4 (0.3, 0.6)	0.36	0.15
	PD-1+ CD8 T cells	22.0 (14.9, 26.9)	-0.01	0.17
	Tim-3+ CD8 T cells	12.4 (6.7, 16.6)	0.01	0.38
T cell exhaustion: specific subsets (proportion of parent)	CD57-/Lag3-/PD1-/Tim3- CD4 T cells	68.4 (62.1, 76.9)	0.00	0.63
	CD57-/Lag3-/PD1-/Tim3+ CD4 T cells	1.5 (0.7, 2.6)	-0.06	0.23
	CD57-/Lag3-/PD1+/Tim3- CD4 T cells	21.6 (16.8, 27.8)	0.00	0.67
	CD57-/Lag3-/PD1+/Tim3+ CD4 T cells	0.8 (0.6, 1.3)	-0.10	0.55
	CD57-/Lag3+/PD1-/Tim3- CD4 T cells	1.5 (0.6, 2.0)	-0.03	0.64
	CD57-/Lag3+/PD1-/Tim3+ CD4 T cells	0.01 (0.00, 0.01)	-10.79	0.30
	CD57-/Lag3+/PD1+/Tim3- CD4 T cells	0.2 (0.1, 0.3)	-0.53	0.44
	CD57-/Lag3+/PD1+/Tim3+ CD4 T cells	0.03 (0.01, 0.04)	-0.44	0.60

	CD57+/Lag3-/PD1-/Tim3- CD4 T cells	0.5 (0.3, 1.4)	0.04	0.73
	CD57+/Lag3-/PD1-/Tim3+ CD4 T cells	0.04 (0.02, 0.07)	-0.88	0.43
	CD57+/Lag3-/PD1+/Tim3- CD4 T cells	1.5 (0.7, 4.0)	0.04	0.20
	CD57+/Lag3-/PD1+/Tim3+ CD4 T cells	0.08 (0.03, 0.23)	0.13	0.41
	CD57+/Lag3+/PD1-/Tim3- CD4 T cells	0.01 (0.00, 0.02)	-3.70	0.21
	CD57+/Lag3+/PD1-/Tim3+ CD4 T cells	0.000 (0.000, 0.002)	51.47	0.29
	CD57+/Lag3+/PD1+/Tim3- CD4 T cells	0.01 (0.00, 0.02)	-0.08	0.98
	CD57+/Lag3+/PD1+/Tim3+ CD4 T cells	0.00 (0.00, 0.01)	-14.03	0.36
	CD57-/Lag3-/PD1-/Tim3- CD8 T cells	50.0 (41.4, 61.8)	0.00	0.89
	CD57-/Lag3-/PD1-/Tim3+ CD8 T cells	6.6 (2.6, 9.6)	0.01	0.59
	CD57-/Lag3-/PD1+/Tim3- CD8 T cells	13.1 (8.5, 18.4)	-0.03	0.04
	CD57-/Lag3-/PD1+/Tim3+ CD8 T cells	0.8 (0.4, 1.5)	0.02	0.90
	CD57-/Lag3+/PD1-/Tim3- CD8 T cells	0.2 (0.1, 0.3)	0.41	0.28
	CD57-/Lag3+/PD1-/Tim3+ CD8 T cells	0.03 (0.01, 0.04)	5.04	0.16
	CD57-/Lag3+/PD1+/Tim3- CD8 T cells	0.09 (0.05, 0.14)	0.79	0.43
	CD57-/Lag3+/PD1+/Tim3+ CD8 T cells	0.007 (0.003, 0.011)	19.45	0.04
	CD57+/Lag3-/PD1-/Tim3- CD8 T cells	16.9 (9.6, 22.5)	0.00	0.68
	CD57+/Lag3-/PD1-/Tim3+ CD8 T cells	2.1 (0.8, 3.9)	0.02	0.46
	CD57+/Lag3-/PD1+/Tim3- CD8 T cells	5.2 (3.0, 7.6)	-0.01	0.75
	CD57+/Lag3-/PD1+/Tim3+ CD8 T cells	0.4 (0.3, 1.2)	0.05	0.55
	CD57+/Lag3+/PD1-/Tim3- CD8 T cells	0.04 (0.02, 0.07)	3.12	0.09
	CD57+/Lag3+/PD1-/Tim3+ CD8 T cells	0.005 (0.000, 0.014)	13.70	0.17
	CD57+/Lag3+/PD1+/Tim3- CD8 T cells	0.04 (0.02, 0.05)	1.97	0.38
	CD57+/Lag3+/PD1+/Tim3+ CD8 T cells	0.002 (0.000, 0.009)	10.61	0.32
T cell effector function:	CD107+ CD4 T cells	4.2 (2.7, 9.0)	0.03	0.17

summary subsets (proportion of parent)	IFN γ + CD4 T cells	6.6 (2.7, 10.4)	0.02	0.08
	IL-2+ CD4 T cells	8.4 (2.9, 13.1)	0.03	0.05
	TNF+ CD4 T cells	20.0 (7.6, 31.9)	0.12	0.01
	CD107+ CD8 T cells	7.7 (3.8, 12.1)	0.02	0.20
	IFN γ + CD8 T cells	15.0 (7.8, 23.1)	0.01	0.31
	IL-2+ CD8 T cells	1.9 (1.2, 3.9)	0.12	0.01
	TNF+ CD8 T cells	8.5 (3.2, 19.2)	0.02	0.05
	CD107-/IFN γ -/IL2-/TNF- CD4 T cells	72.6 (61.6, 79.9)	-0.01	0.02
T cell effector functions: specific subsets (proportion of parent)	CD107-/IFN γ -/IL2-/TNF+ CD4 T cells	8.9 (4.2, 13.7)	0.03	0.07
	CD107-/IFN γ -/IL2+/TNF- CD4 T cells	1.1 (0.7, 1.9)	0.17	0.13
	CD107-/IFN γ -/IL2+/TNF+ CD4 T cells	5.2 (1.3, 8.1)	0.03	0.11
	CD107-/IFN γ +/IL2-/TNF- CD4 T cells	1.0 (0.5, 1.9)	-0.01	0.85
	CD107-/IFN γ +/IL2-/TNF+ CD4 T cells	2.8 (0.1, 4.6)	0.04	0.08
	CD107-/IFN γ +/IL2+/TNF- CD4 T cells	0.08 (0.03, 0.14)	-0.16	0.77
	CD107-/IFN γ +/IL2+/TNF+ CD4 T cells	1.3 (0.4, 2.6)	0.10	0.06
	CD107+/IFN γ -/IL2-/TNF- CD4 T cells	3.2 (2.1, 5.5)	0.01	0.65
	CD107+/IFN γ -/IL2-/TNF+ CD4 T cells	0.3 (0.1, 0.9)	0.35	0.01
	CD107+/IFN γ -/IL2+/TNF- CD4 T cells	0.1 (0.1, 0.2)	1.69	0.05
	CD107+/IFN γ -/IL2+/TNF+ CD4 T cells	0.1 (0.0, 0.4)	0.43	0.04
	CD107+/IFN γ +/IL2-/TNF- CD4 T cells	0.04 (0.01, 0.10)	-0.16	0.90
	CD107+/IFN γ +/IL2-/TNF+ CD4 T cells	0.15 (0.04, 0.42)	0.35	0.02
	CD107+/IFN γ +/IL2+/TNF- CD4 T cells	0.007 (0.002, 0.013)	-3.84	0.73
	CD107+/IFN γ +/IL2+/TNF+ CD4 T cells	0.04 (0.02, 0.21)	0.67	0.03
	CD107-/IFN γ -/IL2-/TNF- CD8 T cells	74.9 (67.3, 85.1)	-0.01	0.13
	CD107-/IFN γ -/IL2-/TNF+ CD8 T cells	1.2 (0.8, 2.5)	0.15	0.02

	CD107-/IFN γ -/IL2+/TNF- CD8 T cells	0.7 (0.5, 1.1)	0.29	0.04
	CD107-/IFN γ -/IL2+/TNF+ CD8 T cells	0.3 (0.1, 0.7)	0.26	0.04
	CD107-/IFN γ +/IL2-/TNF- CD8 T cells	6.0 (2.5, 8.8)	-0.01	0.52
	CD107-/IFN γ +/IL2-/TNF+ CD8 T cells	4.6 (1.7, 9.7)	0.02	0.17
	CD107-/IFN γ +/IL2+/TNF- CD8 T cells	0.3 (0.1, 0.4)	0.42	0.30
	CD107-/IFN γ +/IL2+/TNF+ CD8 T cells	0.5 (0.1, 1.0)	0.33	0.04
	CD107+/IFN γ -/IL2-/TNF- CD8 T cells	4.5 (2.6, 7.9)	0.01	0.67
	CD107+/IFN γ -/IL2-/TNF+ CD8 T cells	0.06 (0.02, 0.19)	0.78	0.13
	CD107+/IFN γ -/IL2+/TNF- CD8 T cells	0.08 (0.05, 0.11)	0.77	0.62
	CD107+/IFN γ -/IL2+/TNF+ CD8 T cells	0.01 (0.00, 0.02)	4.42	0.09
	CD107+/IFN γ +/IL2-/TNF- CD8 T cells	0.4 (0.2, 1.0)	0.17	0.33
	CD107+/IFN γ +/IL2-/TNF+ CD8 T cells	0.3 (0.1, 1.7)	0.10	0.04
	CD107+/IFN γ +/IL2+/TNF- CD8 T cells	0.02 (0.01, 0.04)	1.94	0.46
	CD107+/IFN γ +/IL2+/TNF+ CD8 T cells	0.04 (0.01, 0.16)	0.63	0.08

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93 **Supplementary Table 3.** Univariate regression analysis of immunological variables on reservoir
 94 outgrowth (IUPM), among males only.

	Immune Characteristic	Median	Regression Coefficient	P-value
Soluble Immune Analytes (\log_{10} pg/ml, unless stated otherwise)	IL-7	0.6 (0.3, 0.8)	-0.28	0.60
	IL-15	0.5 (0.4, 0.5)	-0.55	0.66
	CRP (\log_{10} ng/ml)	3.3 (2.8, 3.5)	-0.02	0.96
	D-dimer (\log_{10} ng/ml)	2.1 (1.9, 2.3)	0.04	0.92
	IL-10 (binary)	0.5 (0.2, 0.7)	0.17	0.64
	IL-6	-0.1 (-0.3, 0.0)	-0.18	0.82
	IP10	2.0 (1.9, 2.2)	-1.03	0.23

	CCL21	2.3 (1.5, 2.8)	0.34	0.24
	MIG	2.6 (2.4, 2.8)	-0.30	0.57
	MIP-3 α (binary)	6.4 (4.8, 12.8)	0.08	0.82
	MIP-3 β	1.9 (1.8, 2.0)	-0.50	0.64
T cell subsets (proportion CD3+ cells)	CD4+	52.2 (42.6, 63.4)	-0.02	0.20
	CD8	39.2 (33.4, 51.3)	0.03	0.11
CD4 T cell memory subsets (proportion CD4 T cells)	Naïve (CD28+/CD45RO-/CCR7+)	40.3 (23.2, 47.3)	0.00	0.94
	Central (CD28+/CD45RO+/CCR7+)	34.6 (30.4, 39.1)	0.01	0.73
	Effector (CD28-/CD45RO+/CCR7-)	3.4 (2.5, 6.1)	-0.03	0.64
	Transitional (CD28+/CD45RO+/CCR7-)	9.8 (7.3, 15.5)	0.03	0.44
CD4 T cell activation (proportion CD4 T cells)	CD69	14.9 (9.9, 19.5)	0.03	0.20
	HLA-DR+/CD38+	2.2 (1.5, 3.7)	-0.28	0.09
	CD69+/Ki-67+	0.3 (0.2, 0.6)	1.90	0.04
	CD69- /Ki-67+	1.7 (1.2, 2.4)	0.02	0.93
	Ki-67+	2.0 (1.4, 3.0)	0.07	0.69
	T regulatory cells	1.8 (1.4, 2.7)	0.01	0.96
T cell exhaustion: summary subsets (proportion of parent)	CD57+ CD4 T cells	4.1 (2.4, 5.5)	0.04	0.64
	Lag-3+ CD4 T cells	1.5 (1.0, 3.2)	-0.03	0.87
	PD-1+ CD4 T cells	28.2 (21.0, 32.7)	0.04	0.03
	Tim-3+ CD4 T cells	2.6 (1.9, 3.2)	0.24	0.31
	CD57+ CD8 T cells	29.7 (15.2, 33.6)	0.02	0.10
	Lag-3+ CD8 T cells	0.6 (0.4, 0.9)	-0.15	0.84
	PD-1+ CD8 T cells	20.0 (16.1, 22.0)	0.05	0.13
	Tim-3+ CD8 T cells	8.6 (7.4, 10.2)	0.02	0.58
T cell exhaustion:	CD57-/Lag3-/PD1-/Tim3- CD4 T cells	66.4 (61.4, 74.4)	-0.04	0.03

specific subsets (proportion of parent)	CD57-/Lag3-/PD1-/Tim3+ CD4 T cells	1.3 (1.0, 1.7)	0.09	0.84
	CD57-/Lag3-/PD1+/Tim3- CD4 T cells	24.3 (18.9, 27.7)	0.04	0.04
	CD57-/Lag3-/PD1+/Tim3+ CD4 T cells	0.9 (0.4, 1.0)	0.63	0.13
	CD57-/Lag3+/PD1-/Tim3- CD4 T cells	1.1 (0.8, 2.7)	-0.05	0.80
	CD57-/Lag3+/PD1-/Tim3+ CD4 T cells	0.003 (0.000, 0.011)	38.85	0.41
	CD57-/Lag3+/PD1+/Tim3- CD4 T cells	0.2 (0.1, 0.3)	1.79	0.42
	CD57-/Lag3+/PD1+/Tim3+ CD4 T cells	0.01 (0.00, 0.02)	5.90	0.51
	CD57+/Lag3-/PD1-/Tim3- CD4 T cells	1.0 (0.5, 1.4)	-0.06	0.69
	CD57+/Lag3-/PD1-/Tim3+ CD4 T cells	0.06 (0.02, 0.12)	-0.05	0.95
	CD57+/Lag3-/PD1+/Tim3- CD4 T cells	3.0 (1.6, 3.9)	0.29	0.10
	CD57+/Lag3-/PD1+/Tim3+ CD4 T cells	0.08 (0.04, 0.15)	1.08	0.41
	CD57+/Lag3+/PD1-/Tim3- CD4 T cells	0.01 (0.007, 0.02)	-15.44	0.46
	CD57+/Lag3+/PD1-/Tim3+ CD4 T cells	0 (0, 0.001)	-32.64	0.40
	CD57+/Lag3+/PD1+/Tim3- CD4 T cells	0.01 (0.01, 0.02)	11.76	0.61
	CD57+/Lag3+/PD1+/Tim3+ CD4 T cells	0.002 (0.000, 0.005)	101.14	0.10
	CD57-/Lag3-/PD1-/Tim3- CD8 T cells	53.0 (49.7, 57.1)	-0.04	0.02
	CD57-/Lag3-/PD1-/Tim3+ CD8 T cells	4.42 (3.4, 5.0)	0.04	0.75
	CD57-/Lag3-/PD1+/Tim3- CD8 T cells	10.9 (9.8, 13.9)	0.01	0.76
	CD57-/Lag3-/PD1+/Tim3+ CD8 T cells	0.9 (0.6, 1.4)	0.27	0.29
	CD57-/Lag3+/PD1-/Tim3- CD8 T cells	0.2 (0.1, 0.3)	-1.47	0.37
	CD57-/Lag3+/PD1-/Tim3+ CD8 T cells	0.03 (0.01, 0.05)	-5.52	0.59
	CD57-/Lag3+/PD1+/Tim3- CD8 T cells	0.09 (0.05, 0.24)	0.56	0.75
	CD57-/Lag3+/PD1+/Tim3+ CD8 T cells	0.01 (0.00, 0.03)	4.53	0.72
	CD57+/Lag3-/PD1-/Tim3- CD8 T cells	20.1 (9.9, 23.9)	0.03	0.14
	CD57+/Lag3-/PD1-/Tim3+ CD8 T cells	1.9 (1.3, 2.9)	0.01	0.80

	CD57+/Lag3-/PD1+/Tim3- CD8 T cells	5.1 (3.8, 7.0)	0.14	0.04
	CD57+/Lag3-/PD1+/Tim3+ CD8 T cells	0.7 (0.4, 1.4)	0.16	0.72
	CD57+/Lag3+/PD1-/Tim3- CD8 T cells	0.07 (0.05, 0.14)	0.84	0.83
	CD57+/Lag3+/PD1-/Tim3+ CD8 T cells	0.002 (0.000, 0.017)	6.58	0.66
	CD57+/Lag3+/PD1+/Tim3- CD8 T cells	0.05 (0.03, 0.09)	0.71	0.90
	CD57+/Lag3+/PD1+/Tim3+ CD8 T cells	0.004 (0.000, 0.007)	3.19	0.85
T cell effector functions: summary subsets (proportion of parent)	CD107+ CD4 T cells	4.4 (2.2, 9.4)	0.01	0.88
	IFN γ + CD4 T cells	6.8 (4.1, 11.0)	0.01	0.75
	IL-2+ CD4 T cells	11.1 (6.6, 17.7)	-0.03	0.35
	TNF+ CD4 T cells	24.6 (10.9, 36.7)	-0.01	0.44
	CD107+ CD8 T cells	6.6 (3.8, 8.8)	-0.03	0.35
	IFN γ + CD8 T cells	15.5 (8.2, 21.7)	-0.01	0.72
	IL-2+ CD8 T cells	4.3 (1.6, 5.8)	-0.13	0.02
	TNF+ CD8 T cells	14.3 (3.7, 24.3)	-0.02	0.28
	CD107-/IFN γ -/IL2-/TNF- CD4 T cells	69.0 (54.8, 74.3)	0.01	0.52
T cell effector functions: specific subsets (proportion of parent)	CD107-/IFN γ -/IL2-/TNF+ CD4 T cells	9.5 (5.8, 13.0)	-0.03	0.39
	CD107-/IFN γ -/IL2+/TNF- CD4 T cells	1.6 (1.0, 1.8)	-0.03	0.94
	CD107-/IFN γ -/IL2+/TNF+ CD4 T cells	7.1 (2.9, 11.8)	-0.05	0.27
	CD107-/IFN γ +/IL2-/TNF- CD4 T cells	0.9 (0.3, 1.9)	0.30	0.05
	CD107-/IFN γ +/IL2-/TNF+ CD4 T cells	2.5 (1.3, 5.9)	-0.01	0.87
	CD107-/IFN γ +/IL2+/TNF- CD4 T cells	0.09 (0.02, 0.14)	1.07	0.35
	CD107-/IFN γ +/IL2+/TNF+ CD4 T cells	1.0 (0.5, 2.6)	-0.01	0.90
	CD107+/IFN γ -/IL2-/TNF- CD4 T cells	3.0 (1.5, 7.5)	0.01	0.78
	CD107+/IFN γ -/IL2-/TNF+ CD4 T cells	0.4 (0.2, 0.6)	0.11	0.77
	CD107+/IFN γ -/IL2+/TNF- CD4 T cells	0.3 (0.1, 0.3)	-0.62	0.28

	CD107+/IFN γ -/IL2+/TNF+ CD4 T cells	0.2 (0.1, 0.4)	-0.48	0.43
	CD107+/IFN γ +/IL2-/TNF- CD4 T cells	0.05 (0.03, 0.06)	6.07	0.07
	CD107+/IFN γ +/IL2-/TNF+ CD4 T cells	0.12 (0.04, 0.35)	0.12	0.92
	CD107+/IFN γ +/IL2+/TNF- CD4 T cells	0.01 (0.01, 0.016)	-3.89	0.56
	CD107+/IFN γ +/IL2+/TNF+ CD4 T cells	0.07 (0.04, 0.11)	-0.47	0.89
	CD107-/IFN γ -/IL2-/TNF- CD8 T cells	67.6 (66.0, 82.6)	0.02	0.27
	CD107-/IFN γ -/IL2-/TNF+ CD8 T cells	2.3 (0.8, 3.2)	-0.11	0.25
	CD107-/IFN γ -/IL2+/TNF- CD8 T cells	1.0 (0.5, 2.0)	-0.42	0.11
	CD107-/IFN γ -/IL2+/TNF+ CD8 T cells	0.6 (0.2, 0.9)	-0.31	0.11
	CD107-/IFN γ +/IL2-/TNF- CD8 T cells	5.2 (3.9, 6.9)	0.04	0.37
	CD107-/IFN γ +/IL2-/TNF+ CD8 T cells	6.4 (1.4, 12.1)	-0.01	0.61
	CD107-/IFN γ +/IL2+/TNF- CD8 T cells	0.4 (0.1, 0.6)	-0.22	0.69
	CD107-/IFN γ +/IL2+/TNF+ CD8 T cells	0.8 (0.4, 1.7)	-0.40	0.04
	CD107+/IFN γ -/IL2-/TNF- CD8 T cells	4.7 (1.7, 7.2)	-0.03	0.47
	CD107+/IFN γ -/IL2-/TNF+ CD8 T cells	0.09 (0.04, 0.23)	-1.19	0.29
	CD107+/IFN γ -/IL2+/TNF- CD8 T cells	0.10 (0.05, 0.31)	-0.32	0.25
	CD107+/IFN γ -/IL2+/TNF+ CD8 T cells	0.03 (0.01, 0.05)	-1.36	0.29
	CD107+/IFN γ +/IL2-/TNF- CD8 T cells	0.3 (0.2, 0.7)	0.40	0.59
	CD107+/IFN γ +/IL2-/TNF+ CD8 T cells	0.6 (0.2, 2.1)	-0.20	0.35
	CD107+/IFN γ +/IL2+/TNF- CD8 T cells	0.02 (0.02, 0.06)	-3.20	0.20
	CD107+/IFN γ +/IL2+/TNF+ CD8 T cells	0.08 (0.04, 0.27)	-0.25	0.30

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