

Table S1. Infection details, clinical parameters and neutralizing antibody responses in rhesus macaques

Animal ID	Inoculum	Env Subtype	Route	Depletion of CD8 ⁺ cells	Weeks of Followup	VL Peak (copies/ml)	VL Setpoint (copies/ml)	Clinical AIDS	Peak autologous NAb Titer	bNAb Induction	bNAb Specificities
RM6069	SHIV CH505	C	IV	Yes*	40	19,400,000	988,110	Yes	1396	No	
RM6070	SHIV CH505	C	IV	Yes*	36	62,600,000	736,483	Yes	7837	Yes	V2 Apex
RM6072	SHIV CH505	C	IV	Yes*	184	15,700,000	20,405	No	3544	No	
RM6697	SHIV CH505	C	IV	No	112	812,436	1,638	No	918	No	
RM6698	SHIV CH505	C	IV	No	112	943,718	526	No	277	No	
RM6699	SHIV CH505	C	IV	No	112	1,155,342	506	No	422	No	
RM6701	SHIV CH505	C	IV	No	112	302,350	<250	No	<20	No	
RM6702	SHIV CH505	C	IV	No	96	1,007,432	<250	No	514	No	
RM6703	SHIV CH505	C	IV	No	112	5,257,396	2,546	No	787	No	
RM5695	SHIV CH505#	C	IV	No	65	4,791,344	12,110	Yes	573	Yes	V2 Apex
RM6163	SHIV CH848	C	IV	Yes*	176	24,870,060	38,409	No	544	Yes	V3 glycan
RM6167	SHIV CH848	C	IV	Yes*	136	116,025,900	117,295	Yes	558	Yes	V3 glycan
RM6700	SHIV CH848	C	IV	No	112	1,271,384	13,760	No	418	No	
RM6713	SHIV CH848	C	IV	No	89	17,682,010	45,008	Yes	732	No	
RM6714	SHIV CH848	C	IV	No	112	832,110	15,114	No	655	No	
RM6720	SHIV CH848	C	IV	No	112	1,323,796	<250	No	187	No	
RM40591	SHIV CAP256SU	C	IV	No	129	2,540,090	24,676	No	3067	Yes	V2 Apex
RM42056	SHIV CAP256SU	C	IV	Yes^	88	55,885,850	8,130	No	347	Yes	V2 Apex
RM6727	SHIV CAP256SU	C	IV	Yes^	60	5,282,264	67,392	Yes	665	Yes	Unknown
RM43037	SHIV CAP256SU	C	IV	Yes^	88	11,662,030	10,280	No	492	No	
RM40547	SHIV CAP256SU	C	IV	No	88	825,090	<250	No	<20	No	
RM40624	SHIV CAP256SU	C	IV	No	88	670,330	<250	No	<20	No	

Plasma from RMs 6069 (wk10 & wk20), 6070 (wk10 & wk20), 6072 (wk4, wk10 & wk20)

* Anti-CD8 α antibody (MT807R1, NHP Reagent Resource), 50 mg/kg by slow intravenous push administered on day 0 with respect to SHIV inoculation.

^ Anti-CD8 β antibody (CD8beta255R1, NHP Reagent Resource), 25mg/kg by slow intravenous push administered on day -7 with respect to SHIV inoculation.

Table S3. Neutralization breadth of DH650.8 against 118-strain panel

Virus ID	Clade*	Titer (µg/ml)
		DH650.8
		IC50
6535.3	B	>50
QH0692.42	B	>50
SC422661.8	B	>50
PVO.4	B	>50
TRO.11	B	>50
AC10.0.29	B	>50
RHPA4259.7	B	>50
THRO4156.18	B	>50
REJO4541.67	B	>50
TRJO4551.58	B	>50
WITO4160.33	B	>50
CAAN5342.A2	B	>50
WEAU_d15_410_787	B (T/F)	>50
1006_11_C3_1601	B (T/F)	>50
1054_07_TC4_1499	B (T/F)	>50
1056_10_TA11_1826	B (T/F)	>50
1012_11_TC21_3257	B (T/F)	>50
6240_08_TA5_4622	B (T/F)	>50
6244_13_B5_4576	B (T/F)	>50
62357_14_D3_4589	B (T/F)	>50
SC05_8C11_2344	B (T/F)	>50
Du156.12	C	>50
Du172.17	C	>50
Du422.1	C	>50
ZM197M.PB7	C	>50
ZM214M.PL15	C	>50
ZM233M.PB6	C	>50
ZM249M.PL1	C	>50
ZM53M.PB12	C	>50
ZM109F.PB4	C	>50
ZM135M.PL10a	C	>50
CAP45.2.00.G3	C	>50
CAP210.2.00.E8	C	>50
HIV-001428-2.42	C	>50
HIV-0013095-2.11	C	>50
HIV-16055-2.3	C	>50
HIV-16845-2.22	C	>50
Ce1086_B2	C (T/F)	>50
Ce0393_C3	C (T/F)	>50
Ce1176_A3	C (T/F)	>50
Ce2010_F5	C (T/F)	>50
Ce0682_E4	C (T/F)	>50
Ce1172_H1	C (T/F)	>50
Ce2060_G9	C (T/F)	>50
Ce703010054_2A2	C (T/F)	>50
BF1266.431a	C (T/F)	>50
246F C1G	C (T/F)	>50
249M B10	C (T/F)	>50
ZM247v1(Rev-)	C (T/F)	>50
7030102001E5(Rev-)	C (T/F)	>50
1394C9G1(Rev-)	C (T/F)	>50
Ce704809221_1B3	C (T/F)	>50
CNE19	BC	>50
CNE20	BC	>50
CNE21	BC	>50
CNE17	BC	>50
CNE30	BC	>50
CNE52	BC	>50
CNE53	BC	>50
CNE58	BC	>50
MS208.A1	A	>50
Q23.17	A	>50
Q461.e2	A	>50
Q769.d22	A	>50
Q259.d2.17	A	>50
Q842.d12	A	>50
0260.v5.c36	A	>50
3415.v1.c1	A	>50
0330.v4.c3	A	>50
191955_A11	A (T/F)	>50
191084_B7-19	A (T/F)	>50
9004SS_A3_4	A (T/F)	>50
T257-31	CRF02_AG	>50
928-28	CRF02_AG	>50
263-8	CRF02_AG	>50
T250-4	CRF02_AG	>50
T251-18	CRF02_AG	>50
T278-50	CRF02_AG	>50
T255-34	CRF02_AG	>50
211-9	CRF02_AG	>50
235-47	CRF02_AG	>50
620345.c01	CRF01_AE	>50
CNE8	CRF01_AE	>50
C1080.c03	CRF01_AE	>50
R2184.c04	CRF01_AE	>50
R1166.c01	CRF01_AE	>50
R3265.c06	CRF01_AE	Not tested
C2101.c01	CRF01_AE	>50
C3347.c11	CRF01_AE	>50
C4118.c09	CRF01_AE	>50
CNE5	CRF01_AE	>50
BJOX009000.02.4	CRF01_AE	>50
BJOX015000.11.5	CRF01_AE (T/F)	>50
BJOX010000.06.2	CRF01_AE (T/F)	>50
BJOX025000.01.1	CRF01_AE (T/F)	>50
BJOX028000.10.3	CRF01_AE (T/F)	>50
X1193_c1	G	>50
P0402_c2_11	G	>50
X1254_c3	G	>50
X2088_c9	G	>50
X2131_C1_B5	G	>50
P1981_C5_3	G	>50
X1632_S2_B10	G	>50
3016.v5.c45	D	>50
A07412M1.vrc12	D	>50
231965.c01	D	>50
231966.c02	D	>50
3817.v2.c59	CD	>50
6480.v4.c25	CD	>50
6952.v1.c20	CD	>50
6811.v7.c18	CD	>50
89-F1_2_25	CD	>50
3301.v1.c24	AC	>50
6041.v3.c23	AC	>50
6540.v4.c1	AC	>50
6545.v4.c1	AC	>50
0815.v3.c3	ACD	>50
3103.v3.c10	ACD	>50

* (T/F): Transmitted / Founder Virus

Table S4. Diffraction and Refinement Statistics

DH650 + GP120	
PDB ID	6XCJ
Diffraction data	
Space group	<i>P2₁2₁2</i>
Cell dimensions	
Lengths (Å) a, b, c	138.5 122.1 53.4
Angles (°) α, β, γ	90.0 90.0 90.0
d _{min} (Å)	2.80 (2.97-2.80) ^a
R _{merge} ^b	0.13 (1.52)
CC _{1/2}	99.7 (64.4)
Avg I/ I ₁	13.99 (1.28)
Completeness (%)	99.8 (99.9)
Average redundancy	5.5 (5.6)
Refinement	
Data range (Å)	49.9-2.80
Reflections	20,060
R/R _{free} ^c	0.23/0.28
RMS deviations	
Bond (Å)	0.008
Angles (°)	1.02
Avg B-factor (Å ²)	
Protein	54.2
Water	33.32
Number of Atoms	
Protein	5,565
Water	21

^a Values in parentheses are for the outermost shell of data.

^b $R_{\text{merge}} = \frac{\sum |I_i - \langle I \rangle|}{\sum I_i}$, where I_i is the intensity of the i^{th} observation and $\langle I \rangle$ is the mean intensity. Sums are taken over all reflections.

^c $R = \frac{\sum |F_o| - |F_c|}{\sum |F_o|}$. R_{free} is calculated for a 5% subset of the data

Table S5. RM5695 personalized Ig germline repertoire

Heavy			Lambda		
IGHV	GenBank Accession*	IGHJ	IGLV	GenBank Accession*	IGLJ
IGHV1-AAU-U*01		JH-1*01	IGLV1-ABB*01_S4519	MT656217	JL1*01
IGHV1-AAU-U*02		JH-2P*01	IGLV1-ABB*01_S6715	MT656218	JL2*01
IGHV1-ABW-S*01_S9171	MT656172	JH-3*01	IGLV1-ABB*01_S8237	MT656219	JL3*01
IGHV1-AEP*01		JH-4*01	IGLV1-ABN*02_S2133	MT656220	JL6*01_S7212
IGHV1-AFS*01		JH-5-1*01_S8786	IGLV1-ABN*02_S8168	MT656221	JL7*02
IGHV1-AGS*02_S4535	MT656173	JH-5-2*01	IGLV1-ABS*01_S5111	MT656222	
IGHV1-AGS*02_S6849	MT656174	JH-6*01	IGLV1-ACN*02		
IGHV2-ABG-S*01			IGLV1-ACR*01		
IGHV2-ABG-S*02_S2032	MT656175		IGLV1-ACV*01_S2830	MT656223	
IGHV2-ABU-S*01_S6524	MT656176		IGLV1-ACV*01_S4108	MT656224	
IGHV2-AEY-S*01			IGLV1-ACW*02		
IGHV2-AEY-S*01_S8940	MT656177		IGLV1-ADA*01		
IGHV3-AAB*01			IGLV1-ADL*01		
IGHV3-AAB*02			IGLV1-ADU*01_S4142	MT656225	
IGHV3-ABA*01			IGLV1-ADU*01_S8066	MT656226	
IGHV3-ABA*01_S4331	MT656178		IGLV2-ABE*01_S2946	MT656227	
IGHV3-ABJ*01_S1922	MT656179		IGLV2-ABE*01_S9768	MT656228	
IGHV3-ABK*01_S3779	MT656180		IGLV2-ABI*01_S7244	MT656229	
IGHV3-ABY-S*01			IGLV2-ABI*01_S8018	MT656230	
IGHV3-ABY-S*01_S8824	MT656181		IGLV2-ABJ*01_S9052	MT656231	
IGHV3-ABY-S*03_S0204	MT656182		IGLV2-ABU*01_S3471	MT656232	
IGHV3-ACA*01			IGLV2-ABX*01_S5497	MT656233	
IGHV3-ACN*01_S7808	MT656183		IGLV2-ACE*01_S4286	MT656234	
IGHV3-ACZ*03			IGLV2-ACE*01_S7743	MT656235	
IGHV3-ADF*01_S4250	MT656184		IGLV2-ADC*01		
IGHV3-ADL-S*01_S4190	MT656185		IGLV2-ADC*01_S5785	MT656236	
IGHV3-ADO*01			IGLV2-ADK*01		
IGHV3-ADR*01			IGLV3-AAB*01_S3980	MT656237	
IGHV3-ADR*03_S0556	MT656186		IGLV3-AAB*01_S5697	MT656238	
IGHV3-ADR*03_S5688	MT656187		IGLV3-AAI*01_S0428	MT656239	
IGHV3-ADX*01_S5510	MT656188		IGLV3-AAV*01		
IGHV3-AEH*01_S5714	MT656189		IGLV3-AAV*04		
IGHV3-AEH*01_S6567	MT656190		IGLV3-ACB*01_S7967	MT656240	
IGHV3-AET*01_S5460	MT656191		IGLV3-ADT*01_S0731	MT656241	
IGHV3-AEW-S*03			IGLV3-AEB-X*02		
IGHV3-AEW-S*03_S2686	MT656192		IGLV3-AEB-X*03_S6646	MT656242	
IGHV3-AFC*01_S2394	MT656193		IGLV3-AEC-X*01		
IGHV3-AFE*01_S4619	MT656194		IGLV3-AED-X*01_S4855	MT656243	
IGHV3-AFR*01_S0643	MT656195		IGLV3-AED-X*01_S6916	MT656244	
IGHV3-AFW*01_S5271	MT656196		IGLV3-AED-X*06_S3217	MT656245	
IGHV3-AFY-S*02			IGLV3-AED-X*06_S4240	MT656246	
IGHV3-AFY-S*06			IGLV4-ACF*01_S1875	MT656247	
IGHV3-AGQ-S*02_S3249	MT656197		IGLV4-ACF*02		
IGHV4-ABB-S*01_S0511	MT656198		IGLV5-AAM*01		
IGHV4-ABB-S*01_S8200	MT656199		IGLV5-AAP*01		
IGHV4-ADD*01_S6003	MT656200		IGLV5-AAP*01_S8201	MT656248	
IGHV4-ADD*01_S9501	MT656201		IGLV5-AAX*01		
IGHV4-ADG-U*01_S1296	MT656202		IGLV5-ABG*01_S1697	MT656249	
IGHV4-AEB*01_S7180	MT656203		IGLV5-ABL*01		
IGHV4-AEX-S*01			IGLV5-ABT*01_S4414	MT656250	
IGHV4-AEX-S*01_S0860	MT656204		IGLV5-ACY*01		
IGHV4-AEX-S*01_S4729	MT656205		IGLV5-ADQ*01_S3524	MT656251	
IGHV4-AFB-S*01			IGLV6-AAC*01_S4139	MT656252	
IGHV4-AFB-S*01_S8393	MT656206		IGLV6-ADD*01_S2475	MT656253	
IGHV4-AFI*01_S1365	MT656207		IGLV6-ADW*01		
IGHV4-AFL-U*01			IGLV7-AAK*01		
IGHV4-AFQ-U*01			IGLV7-ABO*01		
IGHV4-AFQ-U*01_S2532	MT656208		IGLV7-ADB*01		
IGHV4-AFU-U*01			IGLV8-ABK*01		
IGHV4-AFU-U*01_S9450	MT656209		IGLV10-AAO*01		
IGHV4-AGL*01_S3100	MT656210		IGLV11-AAY*01		
IGHV4-AGR*01					
IGHV4-AGU*01					
IGHV4-AGU*01_S1118	MT656211				
IGHV4-AGU*01_S2780	MT656212				
IGHV5-ABI*01_S2502	MT656213				
IGHV5-ABI*02_S3096	MT656214				
IGHV7-ABS*01_S4281	MT656215				
IGHV7-AGO-S*01					
IGHV7-AGO-S*03_S8272	MT656216				

*Sequences with GenBank accession numbers represent novel alleles identified by IgDiscover analysis of Ig gene transcripts from IgM+IgD+IgG- B cells from animal RM5695

Table S6. Cryo-EM Data Collection and Refinement Statistics

RHA1.V2.01 in complex with HIV-1 Env BG505 DS-SOSIP	
EMDB ID	22295
PDB ID	6XRT
<u>Data Collection</u>	
Microscope	FEI Titan Krios
Voltage (kV)	300
Electron dose (e ⁻ /Å ²)	71.06
Detector	Gatan K2 Summit
Pixel Size (Å)	1.073
Defocus Range (μm)	-0.1 to -4.1
Magnification	22500
<u>Reconstruction</u>	
Software	cryoSparcV2.12
Particles	21,480
Symmetry	C1
Box size (pix)	288
Resolution (Å) (FSC _{0.143})	3.9
<u>Refinement</u>	
Software	Phenix 1.18
Protein residues	1947
Chimera CC	0.87
EMRinger Score	1.49
R.m.s. deviations	
Bond lengths (Å)	0.005
Bond angles (°)	1.114
<u>Validation</u>	
Molprobit score	1.61
Clash score	4.97
Favored rotamers (%)	99.1
Ramachandran	
Favored regions (%)	95.06
Disallowed regions (%)	0.0