Table St. Generation of WO-1 leaders from plasma and peoplesed bland economiclar cells of 18 prospectively sample
Subpress St. Section of St.

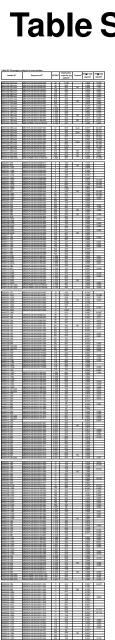


Table S3. Number of viral isolates from ART suppressed individuals with and without treatment interruption

Virus	Subjects ¹	Gender ²	Years on ART before ATI ³	Sample time point	Viral load (copies/μl)	CD4 count (cells/µl)	Sample ⁴	Number of Limiting dilution	n Study Number	References	
						<u> </u>		virus Isolates ⁵			
VOA	B199	M	4	second leukapheresis	<20	200	QVOA Supernatant	1	MNU-0628	Lorenzi et el 2016	
VOA	B106	M	7	first and second leukapheresis	<20	390	QVOA Supernatant	2	MINU-0020	Lorenzi et al., 2016	
VOA	9242	M	2	during ATI (12 weeks) before rebound	<20	967	QVOA Supernatant	2			
VOA	9243	M	5	during ATI (12 weeks) before rebound	<20	545	QVOA Supernatant	1	1		
VOA	9244	M	5	2 weeks pre ATI	<20	796	QVOA Supernatant	4			
VOA	9244	M	5	during ATI (12 weeks) before rebound	<20	796	QVOA Supernatant	2			
VOA	9241	M	5	during ATI (12 weeks) before rebound	<20	709	QVOA Supernatant	1	NCT02825797	Mandaza et al. 2019	
VOA	9241	M	5	2 weeks pre ATI	<20		QVOA Supernatant	4	NC102023131	Mendoza et al., 2018	
Rebound	9242	M	2	16 weeks post ATI	65,520	450	Plasma	5			
Rebound	9243	M	5	21 weeks post ATI	10,110	350	Plasma	1	1		
Rebound	9244	M	5	22 weeks post ATI	120,330	730	Plasma	5			
Rebound	9241	M	5	22 weeks post ATI	13,250	500	Plasma	1			
i	-			·							
VOA	A06	M	>3.6	2.5 weeks pre ATI	<50	>470	QVOA Supernatant and cryopreserved PBMC	6			
VOA	A06	M	>3.6	10.5 months post ATI	<50		QVOA Supernatant	3			
VOA	A08	M	>3.6	4 weeks pre ATI	<50	>470	QVOA Supernatant	4	1		
VOA	A08	M	>3.6	11 months post ATI	<50		QVOA Supernatant and cryopreserved PBMC	16	NCT02463227	Bar et al., 2016;	
VOA	A09	M	>3.6	1 week pre ATI	<50	>470	QVOA Supernatant	4	NC102403221	Salantes et al., 2018	
VOA	A09	M	>3.6	12 months post ATI	<50		QVOA Supernatant	2			
Rebound	A09	M	>3.6	3 weeks post ATI	2,330	>350	Plasma	8			
Rebound	A08	M	>3.6	5 weeks post ATI	20,000	>350	Plasma	17	1		
Rebound	S22	M	8	during 2nd ATI	750,000	449	Plasma	5	1		
Rebound	S23	М	5	during 2nd ATI	556,372	351	Plasma	2	NCT00051818	Papasavvas et al., 2004	
Rebound	S30	М	5	during 3rd ATI	517,473	467	Plasma	4	1		
i	-			-						•	
Rebound	004	M	>1	4 weeks post ATI (week 12 of study)	33,812	794	Plasma	4			
Rebound	030	M	>1	4 weeks post ATI (week 12 of study)	18,056	377	Plasma	4	NCT02227277	Azzoni et al, 2020	
Rebound	044	F	>1	4 weeks post ATI (week 12 of study)	20,171	496	Plasma	4	1		
Rebound	601	M	20	5 weeks post ATI (week 29 of study)	112,160	707	Plasma	6	NCT02588586	Cohen et al., 2018	

All subjects were recruited in the US and had subtype B infections, except for subject A09 who also had a subtype A strain (only subtype B viruses were isolated)

²M, male; F, female.

³ATI, analytical treatment interruption

⁴QVOA, quantitative viral outgrowth assay

⁵Viral isolates were confirmed by sequence analysis to be single virion-derived (Genbank accession number are shown in Table S X)

Table S4. Biological characterization of viral outgrowth and rebound isolates										
Type ¹	Isolate ID ²	Study number	Replicative capacity (ng p24/ml) ³	Isolate tropism ⁴	IFN ₀ 2 IC ₅₀ (pg/ml) ⁵	IFNβ IC ₅₀ (pg/ml) ⁵	sCD4 IC50 (ug/ml) ⁶	CD4-lg IC50 (ug/ml) ⁷	Infectivity (IU/pg RT) ⁸	p24 Particle release (%) ⁹
VOA VOA	B199.2.C.BF.1 B106.1.C.3.8 ¹¹	MNU-0628	80 100		0.04	0.12 3.12				
VOA	B106.2.C.D6.S8 ¹¹	11110 0020	195		0.16	1.96				
VOA - week 12 ¹⁰	9242.VOA.12F23		386	R5	0.02	0.04	>20	>20	39.33	
VOA - week 12 ¹⁰ VOA - week 12 ¹⁰	9242.VOA.12M6 9243.VOA.12Y10		641 679	R5 R5	0.01	0.01	>20 >20	17.43 17.03	79.48 76.88	
VOA - week 12 ¹⁰	9244.VOA.12I10 9244.VOA.12J17		492 483	R5 Dual	0.02	0.04	>20 >20	1.73 >20	14.14 30.82	
VOA - week 12 ¹⁰ VOA - week -2 ¹⁰	9244.VOA.G12		694	R5	0.01	0.10	>20	>20	17.29	
VOA - week -2 ¹⁰ VOA - week -2 ¹⁰	9244.VOA.H1 9244.VOA.K2	-	496 404	R5 Dual	0.02	0.06 0.15	>20	>20	45.21 46.79	48%
VOA - week -210	9244.VOA.P11		423	R5	0.02	0.11	>20	10.97	139.08	
VOA - week 12 ¹⁰ VOA - week -2 ¹⁰	9241.VOA.12AU7 9241.VOA.D2		360 445	Dual R5	0.03	0.25			174.31 2.54	56% 42%
VOA - week -2 ¹⁰ VOA - week -2 ¹⁰	9241.VOA.A10 9241.VOA.C19		315 282		0.04 0.04	0.13				
VOA - week -2 ¹⁰	9241.VOA.C8	NCT02825797	278		0.04	0.10				
Rebound Rebound	9242.REB.6B8 9242.REB.6A7 12		299 309	R5 R5	1.13	45.04 21.99			4.59 5.63	53%
Rebound Rebound	9242.REB.6B2 ¹² 9242.REB.13B1 ¹²		286 547	R5 R5	1.93 1.26	30.49 61.88			15.69 9.93	
Rebound	9242.REB.13B8 12		418	R5	1.70	58.24			9.68	
IMC Rebound	9242.REB.r1 ¹² 9243.REB.13F7		108 554	R5 R5	2.08 1.56	46.67 49.26			3.54 10.58	
Rebound Rebound	9244.REB.7A8 9244.REB.7A1		499 459	R5 R5	1.53 1.59	54.25 31.67			6.41 6.05	81%
Rebound Rebound	9244.REB.7C1 9244.REB.9D3		567 578	R5 R5	1.72 1.35	34.43 48.06	>20	10.21	4.14 27.08	61%
Rebound	9244.REB.9E6		535	R5	1.70	77.03	>20	>20	136.72	81%
Rebound	9241.REB.7D4	1	626	R5	1.62	82.64	>20	>20	4.82	91%
VOA - Pre ATI VOA - Pre ATI	A06.VOA.M5 A06.VOA.M10	1	139 144	R5 R5	0.15 0.11	1.70 1.57	>20 >20	>20 >20	71.20 42.78	52% 22%
VOA - Pre ATI VOA - Pre ATI	A06.VOA.M30 A06.VOA.M28	-	158 97	R5 R5	0.09	1.81	>20	>20	62.60 83.36	28%
VOA - Pre ATI VOA - Pre ATI	A06.VOA.M19 A06.VOA.M29	1	150 124	R5 R5	0.08	1.40			54.54 128.33	38%
VOA - Post ATI	A06.VOA.M29 A06.VOA.2B A06.VOA.M20		143 126	R5	0.07	0.85 1.67			128.52	51% 48%
VOA - Post ATI VOA - Post ATI	A06.VOA.M3		117	R5	0.09	1.10			121.20	48%
VOA - Pre ATI VOA - Pre ATI	A08.VOA.1E2 A08.VOA.M1		99 1548	X4 Dual	0.04	0.32	>20	>20	75.20 14.96	
VOA - Pre ATI VOA - Pre ATI	A08.VOA.M4 A08.VOA.M5		965 665	X4 X4	0.01 0.01	0.04			43.40 16.78	49% 24%
VOA - Post ATI VOA - Post ATI	A08.VOA.M17 A08.VOA.M20		977 620	Dual X4	0.01	0.60			20.23 8.64	31%
VOA - Post ATI	A08.VOA.1B5		122	X4	0.03	0.90	>20	>20	8.29	3170
VOA - Post ATI VOA - Post ATI	A08.VOA.1C8 A08.VOA.1F8		152 163	X4 Dual	0.03	0.08			4.35 4.44	
VOA - Post ATI VOA - Post ATI	A08.VOA.3D8 A08.VOA.4A1	-	65 103	R5 X4	1.47 0.02	5.17 0.15	>20	>20	6.04 13.91	
VOA - Post ATI VOA - Post ATI	A08.VOA.4E4 A08.VOA.5C3		120 83	X4 X4	0.02 0.03	0.09	>20	>20	15.37 2.93	
VOA - Post ATI VOA - Post ATI	A08.VOA.5E2 A08.VOA.5E4		142 119	X4 Dual	0.02 0.04	0.03 0.49	>20	>20	13.11 1.83	
VOA - Post ATI VOA - Post ATI	A08.VOA.6D3 A08.VOA.6E2		120 86	X4 X4	0.03	0.04			11.00	
VOA - Post ATI	A08.VOA.6F6		66	R5	1.23	8.24			8.84	
VOA - Post ATI VOA - Post ATI	A08.VOA.7F8 A08.VOA.8E8		150 50	Dual Dual	0.04 0.03	0.14 0.03			3.36 2.65	
Rebound Rebound	A08.REB.1A3 A08.REB.1D1		451 496	Dual Dual	1.06 1.45	83.13 51.37			2.50 5.98	94% 87%
Rebound Rebound	A08.REB.5C2 A08.REB.5D2	NCT02463227	125 250	R5 X4	1.22	80.62 45.44			47.47 29.40	
Rebound Rebound	A08.REB.6B3 A08.REB.6D6		125 728	R5 X4	0.85 1.22	55.84 46.83	>20	>20	59.56 40.83	
Rebound Rebound	A08.REB.7A3 A08.REB.7C1		504 707	X4 X4	1.51	75.85 68.06			24.00 24.61	
Rebound	A08.REB.7C2		176	R5	2.07	44.85	- 00	- 00	24.26	
Rebound Rebound	A08.REB.7D3 A08.REB.8A5		309 768	X4 X4	1.65 1.59	38.78 41.98	>20 >20	>20 >20	25.16 39.00	
Rebound Rebound	A08.REB.8B3 A08.REB.8D1		763 425	X4 X4	1.61	41.17 31.24	>20	>20	46.30 29.05	
Rebound Rebound	A08.REB.6B5 A08.REB.8B5	-	510 510	X4 X4	2.28 1.94	64.49 69.78		_	142.66 224.92	
Rebound IMC	A08.REB.2F4 A08.REB.2F4	1	420 66	R5	3.01	92.25 117.61			3.82	77%
Rebound IMC	A08.REB.1A5		506	X4	1.69	42.34			1.31	84%
VOA - Pre ATI	A08.REB.1A5 A09.VOA.M25	1	106 194	X4 R5	2.70 0.26	49.23 1.29			82.33	
VOA - Pre ATI VOA - Pre ATI	A09.VOA.M22 A09.VOA.M2	1	206 254	R5 R5	0.05 0.05	0.19 0.46			115.31 82.42	23%
VOA - Pre ATI VOA - Post ATI	A09.VOA.M6 A09.VOA.M9	1	171 171	R5 R5	0.06 0.21	0.70			110.34 45.72	49%
VOA - Post ATI Rebound	A09.VOA.M16 A09.REB.1B5		285 275	R5	0.05 3.16	0.69 58.67			74.11	
Rebound Rebound	A09.REB.1D5 A09.REB.2B2	1	101 171	R5	2.19	51.20 26.36			1.05	
Rebound	A09.REB.1A1		376	R5	2.40	46.03			7.73	73%
Rebound Rebound	A09.REB.1A3 A09.REB.1A4		227 239	R5 R5	1.78 2.48	58.57 70.36			3.04 3.01	79%
Rebound Rebound	A09.REB.1A5 A09.REB.1A2	1	249 221	R5 R5	2.55 2.54	61.27 83.45	>20	>20	11.68 5.30	74%
IMC	A09.REB.1A2		104	R5	1.63	142.15				
Rebound Rebound	S22.REB.24C4 S22.REB.1E2		399 389	R5 R5	1.64	90.59 61.14			5.50 8.59	74%
Rebound Rebound	S22.REB.1F5 S22.REB.3A6	1	374 487	R5 R5	1.62	93.32	>20	>20 9.977	12.95 21.40	74%
Rebound	S22.REB.3B6		378	R5	1.00	75.65	>20	>20	13.32	74% 65%
Rebound Rebound	S23.REB.3D1 S23.REB.24A5	NCT00051818	358 475	R5 R5	1.70 1.25	66.56 76.30	>20	11.63	21.68 29.05	
Rebound Rebound	S30.REB.8D1 S30.REB.2D1	-	407 303	-	1.21 2.14	84.20 87.20		_	-	
Rebound Rebound	S30.REB.2D2 S30.REB.7D1	1	350 336	R5 R5	1.54 1.72	70.82 72.69			18.95 3.42	71% 85%
		1								
Rebound Rebound	004.REB.4D4 004.REB.4E6		413 263	R5 R5	2.58 2.54	93.58 63.21	>20 >20	>20 >20	18.74 15.19	85% 88%
Rebound Rebound	004.REB.4D1 004.REB.4E3	1	259 252	R5 R5	2.65 5.55	97.87 111.80	>20 >20	>20 >20	32.26 8.66	80%
Rebound Rebound	030.REB.3E1 030.REB.2E1	NOTO	371 396	R5 R5	2.99 2.84	56.49 47.24			24.35 30.22	68% 93%
Rebound Rebound	030.REB.4E1 030.REB.8E1	NCT02227277	427 308	R5 R5	3.48	70.43 69.21			30.93 32.36	74% 87%
Rebound	044.REB.6F1		283	R5	3.42	79.99			39.42	74%
Rebound Rebound	044.REB.5F1 044.REB.7F1		348 329	R5 R5	2.75 2.80	69.41 72.71			30.89 16.85	86% 81%
Rebound	044.REB.8F1		355	R5	2.48	91.45		<u> </u>	37.08	92%

¹²Infectious molecular clones (IMCs) 9242.REB.r1 and 601.REB.r1 were synthesized based on the consensus sequences of four closely related rebound isolates, including 9242.REB.6A7, 9242.REB.6B2, 9242.REB.13B1 and 9242.REB.13B8 as well as 601.REB.4B4, 601.REB.4A8, 601.REB.4A7 and 601.REB.4C1, respectively.

Table S5. Virus replication in macrophages from different donors.

					•			Donors							
ID	Type	Virus	Source	Virus Tropism	sCD4 IC50 (ug/ml)	CD4-lg IC50 (ug/ml)	ZB620	ZB624	ZB710	ZB28	ZB31	ZB722	ZB725	ZB668	
	- 75-	•			, ,										
MCST	Chronic	IMC	Parrish et al., PNAS 2013	R5			no	yes	yes	yes	no				
STCOr1	Chronic	IMC	Parrish et al., PNAS 2013	Dual			no	yes	no	no	yes				
THRO	Transmitted founder	IMC	Salazar-Gonzalez et al., J Virol 2009	R5			no	yes	yes	yes	yes		ļ		
TYBE*** UG021***	Chronic***	IMC IMC	Yi et al., J Virol 1999	X4						yes	yes		<u> </u>		
WITO	Chronic*** Transmitted founder	IMC	Gao et al., J Virol 1996 Salazar-Gonzalez et al., J Virol 2009	X4 R5			no	yes	yes	yes yes	yes yes	-			
YU2***	Chronic***	IMC	Li et al., J Virol 1991	R5	0.9224	0.08674	no	yes	yes	ves	ves				
REJO	Transmitted founder	IMC	Salazar-Gonzalez et al., J Virol 2009	R5	0.0224	0.00014	no	yes	yes	yes	ves		-		
RHGA	Chronic	IMC	Parrish et al., PNAS 2013	R5			no	yes	yes	no	no				
RHPA	Transmitted founder	IMC	Salazar-Gonzalez et al., J Virol 2009	R5			no	yes	yes	yes	yes				
CH058	Transmitted founder	IMC	Salazar-Gonzalez et al., J Virol 2009	R5			no	yes	yes	yes	yes	yes	no	yes	
CH077	Transmitted founder	IMC	Salazar-Gonzalez et al., J Virol 2009	Dual			no	yes	yes	yes	yes				
CH141	Chronic	IMC	Parrish et al., PNAS 2013	R5			no	yes	yes	yes	yes				
CH167	Chronic	IMC	Parrish et al., PNAS 2013	R5			no	yes	yes	yes	yes		ļ		
CH042	Transmitted founder	IMC	Parrish et al., PNAS 2013	R5			no	yes	yes	yes	yes		<u> </u>		
CH470 CH492	Transmitted founder Chronic	IMC IMC	Parrish et al., PNAS 2013 lyer et al., PNAS 2017	R5 R5						yes no	yes no	yes	yes no	yes no	
MM33.TF	Transmitted founder	IMC	This study	R5			-			no	no	no	110	no	
MM33.13	Chronic	IMC	This study	R5						yes	yes				
MM33.17	Chronic	IMC	This study	R5						no	no				
								ı	ı			ı			
9244.VOA.G12	VOA (week -2)	Isolate	Mendoza et al., 2018	R5	>20	>20	no	yes		yes	yes				
9244.VOA.K2	VOA (week -2)	Isolate	Mendoza et al., 2018	Dual	>20	>20	no	yes		yes	yes				
9244.VOA.P11	VOA (week -2)	Isolate	Mendoza et al., 2018	R5	>20	10.97	no	yes		yes	yes		<u> </u>		
9244.VOA.12J17 9244.REB.9D3	VOA (week 12) Rebound	Isolate	Mendoza et al., 2018 This study	Dual R5	>20	>20	no	yes		yes	yes				
9244.REB.9E6	Rebound	Isolate Isolate	This study This study	R5	>20	>20	yes no	yes yes		yes yes	yes yes				
9241.REB.7D4	Rebound	Isolate	This study	R5	>20	>20	no	yes		yes	yes				
9242.REB.r1	Rebound	IMC	This study	R5	- 20	- 20	no	yes	yes	yes	yes	ves	yes	yes	
								,,,,,		, ,	. ,	, ,		,	
A08.VOA.1B5	VOA (Post ATI)	Isolate	This study	X4	>20	>20				no	no	no	no	no	
A08.VOA.1C8	VOA (Post ATI)	Isolate	This study	X4								no	no		
A08.VOA.1F8	VOA (Post ATI)	Isolate	This study	Dual								no	no		
A08.VOA.3D8	VOA (Post ATI)	Isolate	This study	R5								no	no	no	
A08.VOA.4E4	VOA (Post ATI)	Isolate	This study	X4	>20	>20				no	no	no	no	no	
A08.VOA.5E4	VOA (Post ATI)	Isolate	This study	Dual								no	no	no	
A08.VOA.6D3 A08.VOA.6F6	VOA (Post ATI)	Isolate	This study This study	X4 R5								no	no	no	
A08.VOA.5E2	VOA (Post ATI) VOA (Post ATI)	Isolate Isolate	This study This study	X4	>20	>20	-					no no	no no	no no	
A08.VOA.7F8	VOA (Post ATI)	Isolate	This study This study	Dual	-20	720						no	no	110	
A08.VOA.7C1	VOA (Post ATI)	Isolate	This study	X4								no	no		
A08.VOA.8E8	VOA (Post ATI)	Isolate	This study	Dual								no	no		
A08.REB.5C2	Rebound	Isolate	This study	R5						yes	yes	yes	no	yes	
A08.REB.5D2	Rebound	Isolate	This study	X4						no	no	no	no	no	
A08.REB.6B3	Rebound	Isolate	This study	R5						yes	yes	no	no	no	
A08.REB.6D6	Rebound	Isolate	This study	X4	>20	>20				no	no	no	no	no	
A08.REB.7C1	Rebound	Isolate	This study	X4	-				1	no	no	no	no	no	
A08.REB.7C2 A08.REB.7D3	Rebound	Isolate	This study This study	R5 X4	>20	>20	-			no	yes	yes	no	no	
A08.REB.7D3 A08.REB.8D1	Rebound Rebound	Isolate Isolate	This study This study	X4 X4	>20	>20	-	-		no	no	no no	no	no	
A08.REB.8A5	Rebound	Isolate	This study This study	X4	>20	>20			 			no	no no	no no	
A08.REB.1A5	Rebound	IMC	This study This study	X4			no	no	no	no	no	no	no	no	
A08.REB.2F4	Rebound	IMC	This study	R5			no	yes	yes	no	no	yes	yes	yes	
A09.REB.1A5	Rebound	Isolate	This study	R5	>20	>20	no	yes	1	yes	yes	1 ,		1	
A09.REB.1A2	Rebound	IMC	This study	R5			no	yes	yes	yes	yes	yes	yes	yes	
			•												
S22.REB.1F5	Rebound	Isolate	This study	R5	>20	>20	no	yes		yes	yes				
S22.REB.3B6	Rebound	Isolate	This study	R5	>20	>20	no	yes		yes	yes				
S23.REB.3D1	Rebound	Isolate	This study	R5	>20	11.63	no	yes		yes	yes		Ь		
004 DED 4D4	Debeund	I to all to	This start.	Dr.	- 00	- 00		1	T	T		T			
004.REB.4D1	Rebound	Isolate	This study	R5	>20	>20 >20	no	yes	1	yes	yes	1	↓		
004.REB.4E3 004.REB.4E6	Rebound Rebound	Isolate Isolate	This study This study	R5 R5	>20 >20	>20	no no	yes	 	yes	yes	 	 	<u> </u>	
004.NEB.4E0	Venonin	ISUIDIE	This study	сл	~20	~20	110	yes	I	yes	yes	l			
601.REB.r1	Rebound	IMC	This study	R5		ĺ	no	yes	yes	yes	yes	yes	yes	yes	
	obouilu		study	110	i .	i .		, , , , ,	, , , , ,	, , , , ,	, ,	,,,,,	, ,,,,,	,	

^{*}Using CD14 selection, moncytes were isolates from peripheral blood mononuclear cells of 8 healthy donors, and cultured in M-CSF containing media to allow differentiation into macrophages. Virus replication was monitored for 20 days by measuring p24 antigen in culture supernatants. "yes" denotes p24 production above 0.5ng/ml.