

1 **Supplementary table 1.** TREC results for the neonatal SCID screening. Triplicate dried blood spots were  
2 used for TREC and  $\beta$ -actin detection.

	TREC (copies $\mu\text{L}^{-1}$ )	$\beta$ -actin (copies $\mu\text{L}^{-1}$ )	Result
Controls*			
No TREC control	0	600	NA
Low control	120	20	NA
High control	670	600	NA
SCID patient			
Test 1	0	123	TREC negative
Test 2	0	184	TREC negative
Test 4	12	127	TREC negative

3 TREC  $<40$  copies  $\mu\text{L}^{-1}$  was defined as TREC negative<sup>23</sup>.

4 NA indicates not available.

5 \*The values for controls were provided from the manufacturer (Perkin Elmer).

**Supplementary table 2.** Laboratory findings after hematopoietic cell transplantation

Age		238D	269D	308D	338D	353D	16M
Lab	Day after HCT	Day 40	Day 71	Day 110	Day 140	D153	D364
	Normal						
WBC	6000-14,000, $\mu\text{L}^{-1}$	5310	6400	4950	2490	4940	5580
ALC	1500-3000, $\mu\text{L}^{-1}$	3650	4520	4010	1230	2260	3260
Hb	10.5-14.0, g $\text{dL}^{-1}$	9.6	11.2	13.7	7.0	9.6	12.6
PLT	150-400, $\times 10^3 \mu\text{L}^{-1}$	288	396	405	61	364	274
CRP	0-8, mg $\text{dL}^{-1}$	<0.3	<0.3	<0.3	<0.3	0.5	-
TB	0.2-0.8, mg $\text{dL}^{-1}$	0.1	0.1	0.2	1.5	0.4	0.3
AST	13-34, IU $\text{L}^{-1}$	118	65	114	501	57	35
ALT	5-46, IU $\text{L}^{-1}$	114	56	103	475	37	35
CD3 <sup>+</sup>	2284-4776, $\mu\text{L}^{-1}$	3500	4571	4014	1784	-	1417
CD4 <sup>+</sup>	1523-3472, $\mu\text{L}^{-1}$	7	32	269	585	-	585
CD8 <sup>+</sup>	524-1583, $\mu\text{L}^{-1}$	1536	2115	2414	832	-	832
CD19 <sup>+</sup>	776-2238, $\mu\text{L}^{-1}$	1886	2267	-	242	-	228
CD56 <sup>+</sup>	230-801, $\mu\text{L}^{-1}$	28	64	76	21	-	22
IgG	176-601, mg $\text{dL}^{-1}$	843 (on IGRT)		1137 (on IGRT)			254 (without IGRT)
IgA	4.4-84, mg $\text{dL}^{-1}$	Not detected		Not detected			Not detected
IgM	17-105, mg $\text{dL}^{-1}$	8.5		17.5			42.7
CMV	<500, copies $\text{mL}^{-1}$	3055	1375	Not detected	Not detected	Not detected	Not detected

WBC indicates white blood cell; ALC, absolute lymphocyte count; Hb, hemoglobin; PLT, platelet count; CRP, C-reactive protein; TB, total bilirubin; AST, aspartate aminotransferase; ALT, alanine aminotransferase; LDH, lactate dehydrogenase; Ig, immunoglobulin; D, day; M, month; IGRT; immunoglobulin G replacement therapy..

**Supplementary table 3.** Main clinical features associated with the patient

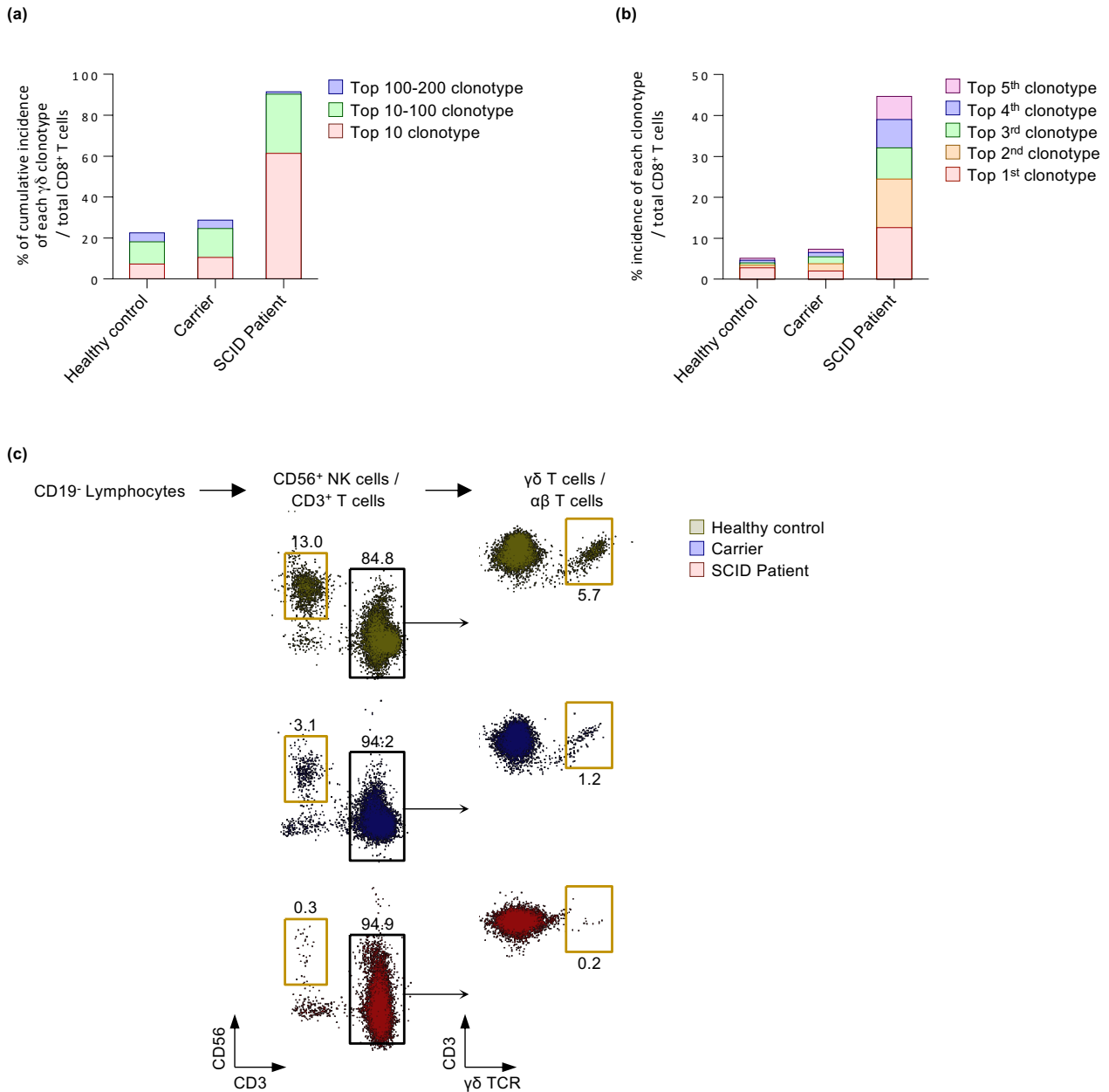
Clinical features		Age at onset
Fever of unknown origin	No	-
Skin rash	No	-
Lymphadenopathy (mesenteric)	<b>Yes</b>	<b>4.5 months</b>
Hepatomegaly	No	-
Splenomegaly	<b>Yes</b>	<b>4.5 months</b>
Neuropathy	No	-
Bronchiectasis	No	-
Enteropathy	No	-
Autoimmune cytopenia	No	-
Vasculitis	No	-
Malignancy	No	-
Developmental delay	No	-
Failure to thrive	No	-
Diarrhea	<b>Yes</b>	<b>6 months</b>
Sentinel infections		
CMV	<b>CMV hepatitis</b>	<b>3 months (1<sup>st</sup>), 5.5 months (2<sup>nd</sup>), 7 months (3<sup>rd</sup>), 8 months (4<sup>th</sup>)</b>
VZV	No	-
HSV	No	-
PJP	No	-
Rotavirus	No	-
BCG infection	<b>Probable</b>	<b>4.5 months</b>
Fungus	No	-
Other infections 1	<b><i>E. coli</i> urinary tract infection</b>	<b>1.5 months</b>
Other infections 2	<b>Noroviral enteritis</b>	<b>6 months</b>
Other infections 3	<b>Lt, 1<sup>st</sup> toe cellulitis</b>	<b>5 months</b>

CMV indicates cytomegalovirus; VZV, varicella zoster virus; HSV, herpes simplex virus; PJP, *Pneumocystis jirovecii*; BCG, Bacillus Calmette–Guérin.

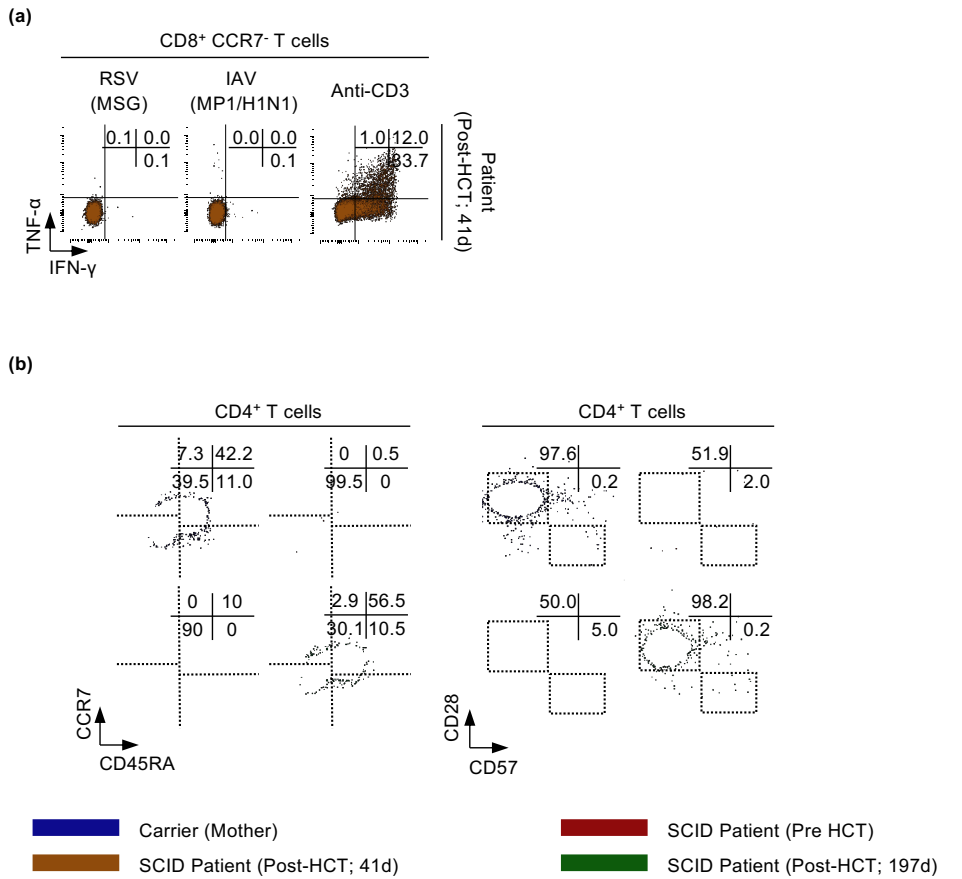
**Supplementary table 4. Key resources table**

REAGENT or RESOURCE	SOURCE	IDENTIFIER
<b>Antibodies</b>		
BV421 anti-human CD132 (clone AG184)	BD Biosciences	Cat# 566222
PE anti-human CD132 (clone AG184)	BD Biosciences	Cat# 555900
BV711 anti-human CD8 (clone RPA-T8)	BD Biosciences	Cat# 563677
BV786 anti-human CD56 (clone NCAM16.2)	BD Biosciences	Cat# 564058
AlexaFluor® 647 anti-human CD16 (clone 3G8)	BD Biosciences	Cat# 557710
PerCP-Cy™5.5 anti-human CD4 (clone RPA-T4)	BD Biosciences	Cat# 560650
APC anti-human CD3 (clone UCHT1)	Biolegend	Cat# 300412
Alexa Fluor® 700 anti-human CD19 (clone HIB19)	BD Biosciences	Cat# 557921
APC/Cyanine7 anti-human HLA-DR (clone L243)	Biolegend	Cat# 307618
BV421 anti-human CD4 (clone RPA-T4)	BD Biosciences	Cat# 562424
BV510 anti-human CD3 (clone UCHT1)	BD Biosciences	Cat# 563109
VioBright515 anti-human CD56 (clone REA196)	Miltenyi Biotec	Cat# 130-108-990
AlexaFluor® 647 anti-human Stat5 (clone pY694)	BD Biosciences	Cat# 612599
PE-eFluor610 anti-human CD14 (clone 61D3)	eBioscience	Cat# 61-0149-42
PE-eFluor610 anti-human CD19 (clone HIB19)	eBioscience	Cat# 61-0199-42
PE-Cy™7 anti-human CD8 (clone RPA-T8)	BD Biosciences	Cat# 557746
PerCP-Cy™5.5 anti-human CD197 (clone 150503)	BD Biosciences	Cat# 561144
APC-H7 anti-human CD3 (clone SK7)	BD Biosciences	Cat# 560176
BV421 anti-human CD28 (clone CD28.2)	BD Biosciences	Cat# 562613
BV510 anti-human CD3 (clone HIT3 $\alpha$ )	BD Biosciences	Cat# 564713
FITC anti-human CD4 (clone RPA-T4)	BD Biosciences	Cat# 561842
PE anti-human CD57 (clone NK-1)	BD Biosciences	Cat# 560844
APC/Cyanine7 anti-human CD45RA (clone HI100)	Biolegend	Cat# 304128
PE anti-human IFN- $\gamma$ (clone B27)	BD Biosciences	Cat# 554701
APC anti-human TNF- $\alpha$ (clone Mab11)	Invitrogen	Cat# 17-7349-82
CD3 pure-functional grade, human (clone OKT3)	Miltenyi Biotec	Cat# 130-093-387
<b>Chemicals, peptides, and recombinant proteins</b>		
PepMix™ EBV (BARF1)	JPT	Cat# PM-EBV-BARF1
PepMix™ Influenza A (MP1 /California (H1N1))	JPT	Cat# PM-INFA-MP1-

		H1N1
PepMix™ HRSVA (major surface glycoprotein G)	JPT	Cat# PM-HRSVA-MSG
PepMix™ HCMVA (pp65) (>90%)	JPT	Cat# PM-PP65-2
PepMix™ HCMVA (IE-1)	JPT	Cat# PM-IE1
Purified protein derivative (PPD)	Dr. Brennan at Aeras (Rockville, MD).	
Recombinant human IL-2	Peptotech	200-2
Brilliant stain buffer	BD Biosciences	Cat# 563794
Dimethyl sulfoxide	Sigma-Aldrich	Car# 276855
Ficoll-paque PREMIUM	GE Healthcare Life Science	Cat# 17-5442-02
<b>Critical commercial assays</b>		
Brefeldin A protein transport inhibitor	BD Biosciences	Cat# 555029
Monensin protein transport inhibitor	BD Biosciences	Cat# 554724
eBioscience™ IC fixation buffer	Invitrogen	Cat# 00-8222-49
eBioscience™ Foxp3/transcription factor staining buffer set	Invitrogen	Cat# 00-5523-00
LIVE/DEAD fixable red dead cell stain kit	Invitrogen	Cat# L23102
LIVE/DEAD fixable aqua dead cell stain kit	Invitrogen	Cat# L34957
Methanol	Biosesang	Cat# M1066



**Supplementary figure 1. (a, b)**  $\gamma\delta$ -TCR repertoire analysis of CD8<sup>+</sup> T cells from a healthy donor, carrier, and the SCID patient showed vigorous expansion of the oligoclonotypes of engrafted CD8<sup>+</sup> T cells from the patient. **(c)** Proportion of  $\gamma\delta$ -T cells among CD3<sup>+</sup> T cells from a healthy donor, carrier, and the patient.



**Supplementary figure 2.** (a) Engrafted CD8<sup>+</sup> T cells did not respond to viral antigen after HCT. (b) Immunophenotyping analysis revealed an increased frequency of effector/effector memory (CD45RA<sup>-</sup>CCR7<sup>-</sup>) and replicative senescent (CD28<sup>-</sup>CD57<sup>+</sup>) CD4<sup>+</sup> T cells from the patient pre-HCT. The percentage of naïve CD4<sup>+</sup> T cells was increased after HCT.