

Supporting Material

Supplementary Methods

Immunosuppression and prophylaxis regimens

All kidney transplant recipients of organs from donors after circulatory death underwent induction therapy with intravenous (IV) rabbit antithymocyte globulin (ATG-Fresenius, 1.25 mg/Kg daily for 5-7 days), with delayed introduction of a calcineurin inhibitor (CNI) from post-transplant day 6. Recipients at high immunological risk also received ATG induction for 1-3 days with early CNI initiation from post-transplant day 0. Basiliximab (20 mg on days 0 and 4) with delayed CNI introduction from post-transplant day 5 was reserved to patients at high risk for CNI-related nephrotoxicity (i.e. older age or pre-transplant comorbidities). Maintenance immunosuppression consisted of tacrolimus (0.1 mg/Kg daily, adjusted to a target trough level of 10-15 ng/mL during the first month and 5-10 ng/mL thereafter); mycophenolic acid (360 mg twice daily); and prednisone (1 mg/Kg daily with progressive tapering). Conversion to mammalian target of rapamycin (mTOR) inhibitor-based regimens with reduced-dose tacrolimus (target trough level of 3-6 ng/mL) was performed on an individual basis for recipients experiencing severe CNI-related adverse effects, difficult-to-treat cytomegalovirus (CMV) infection or post-transplant cancer.

All patients received preoperatively a single dose of IV cefazolin (which was replaced with ciprofloxacin in those with β -lactam hypersensitivity). Prophylaxis for *Pneumocystis jirovecii* pneumonia with trimethoprim-sulfamethoxazole (160/800 mg three times weekly) or monthly pentamidine was given for 9 months. In patients at high-risk for CMV disease (serology mismatch [donor positive (D+) and recipient negative (R-)] or induction therapy with ATG), either IV ganciclovir (5 mg/Kg daily) or oral valganciclovir (900 mg daily) were administered as universal prophylaxis for 3 to 6 months. Intermediate-risk patients (R+ without T-cell-depleting therapy) were monitored every 2-4 weeks for CMV infection with a PCR-based assay (RealStar[®] CMV PCR kit, Altona Diagnostics, Hamburg, Germany), and preemptive therapy with IV ganciclovir (5

mg/Kg/12 hours) or oral valganciclovir (900 mg/12 hours) was administered for at least 2 weeks in presence of high-level (>1.000 IU/mL) or rapidly increasing viral load. (Val)ganciclovir dose was adjusted according to renal function when necessary [1,2].

References

1. Kotton CN, Kumar D, Caliendo AM, Asberg A, Chou S, Danziger-Isakov L, et al. Updated international consensus guidelines on the management of cytomegalovirus in solid-organ transplantation. *Transplantation* 2013; 96: 333-360.
2. Torre-Cisneros J, Aguado JM, Castón JJ, Almenar L, Alonso A, Cantisán S, et al. Management of cytomegalovirus infection in solid organ transplant recipients: SET/GESITRA-SEIMC/REIPI recommendations. *Transplant Rev (Orlando)* 2016; 30: 119-143.

Supplementary Results

Table S1. Genotypic frequencies of the candidate SNPs.

Gene (SNP database ID number)	Genotype	N	(%)
<i>CTL4</i> (rs5742909)	CC	180	81
	CT	36	16
	TT	5	2
<i>CTL4</i> (rs231775)	AA	112	51
	AG	88	40
	GG	21	10
<i>TLR3</i> (rs3775291)	CC	108	49
	CT	84	38
	TT	29	13
<i>TLR9</i> (rs5743836)	AA	160	72
	AG	51	23
	GG	10	5
<i>TLR9</i> (rs352139)	TT	64	29
	TC	96	43
	CC	61	28
<i>CD209</i> (rs735240)	GG	61	28
	GA	108	49
	AA	52	24
<i>CD209</i> (rs4804803)	AA	129	58
	AG	80	36
	GG	12	5
<i>IL28B</i> (rs12979860)	CC	106	48
	CT	86	39
	TT	32	14
<i>IL28B</i> (rs8099917)	TT	160	72
	TG	52	24
	GG	9	4
<i>TNF</i> (rs1800629)	GG	172	78
	GA	46	21
	AA	3	1
<i>IL10</i> (rs1800872)	TT	22	10
	TG	108	49
	GG	91	41
<i>IL10</i> (rs1878672)	GG	83	38
	GC	107	48
	CC	31	14

Redondo N, et al. Polymorphisms in genes orchestrating innate immune responses do not meaningfully impact replication kinetics of Torque teno virus in kidney transplant recipients

<i>IL12B</i> (rs3212227)	TT	110	50
	TG	93	42
	GG	18	8
<i>IL17A</i> (rs2275913)	GG	103	47
	GA	90	41
	AA	28	13

CTLA-4: cytotoxic T-lymphocyte antigen 4; IL: interleukin; SNP: single-nucleotide polymorphism; TLR: toll-like receptor; TNF: tumor necrosis factor.

Table S2. Comparison of demographics and clinical characteristics between evaluable recipients with or without undetectable TTV DNAemia at the baseline (pre-transplant) assessment (n = 187).

Variable	Undetectable TTV DNAemia at baseline		P-value
	No (n = 180)	Yes (n = 7)	
Age of recipient, years [mean ± SD]	54.4 ± 15.9	45.7 ± 6.6	0.012
Male gender of recipient [n (%)]	128 (71.1)	4 (57.1)	0.421
Current or prior smoking history [n (%)]	74 (41.1)	3 (42.9)	1.000
Pre-transplant chronic co-morbidities [n (%)]			
Hypertension	152 (84.9)	7 (100.0)	0.596
Diabetes mellitus	58 (32.2)	1 (14.3)	0.435
Chronic lung disease	26 (14.4)	0 (0.0)	0.596
Coronary heart disease	20 (11.1)	1 (14.3)	0.572
Other chronic heart disease	36 (20.0)	0 (0.0)	0.349
Peripheral arterial disease	15 (8.3)	1 (14.3)	0.471
Cerebrovascular disease	12 (6.7)	0 (0.0)	1.000
Previous solid organ transplantation [n (%)]	20 (11.2)	2 (28.6)	0.914
Underlying end-stage renal disease [n (%)]			
Glomerulonephritis	41 (22.8)	2 (28.6)	0.662
Diabetic nephropathy	36 (20.0)	1 (14.3)	1.000
Polycystic kidney disease	22 (12.2)	1 (14.3)	1.000
Chronic interstitial nephropathy	9 (5.0)	0 (0.0)	1.000
Unknown	22 (12.2)	1 (14.3)	1.000
Other	15 (8.3)	1 (14.3)	0.471
Positive CMV serostatus [n (%)]	152 (84.4)	5 (71.4)	0.312
Positive HCV serostatus [n (%)]	15 (8.3)	1 (14.3)	0.471
Positive HIV serostatus [n (%)]	2 (1.1)	0 (0.0)	1.000
Pre-transplant renal replacement therapy [n (%)]	158 (87.8)	4 (57.1)	0.052
Hemodialysis	132 (83.5)	4 (100.0)	1.000
Continuous ambulatory peritoneal dialysis	26 (16.5)	0 (0.0)	
Time on dialysis, months [median (IQR)]	17.6 (9.3 – 34.4)	43.7 (10.8 – 78.9)	0.356

CMV: cytomegalovirus; HCV: hepatitis C virus; HIV: human immunodeficiency virus; IQR: interquartile range; SD: standard deviation; TTV: torque teno virus.

Table S3. Areas under the curve for plasma TTV DNA through different post-transplant periods according to candidate SNPs.

SNP (ID number)	Model	Genotype	AUC for plasma TTV DNA, log ₁₀ copies/mL (mean ± SD)					
			First month	P-value	First 3 months	P-value	First 6 months	P-value
CTLA4 (rs5742909)	Dominant	CC	4.9 ± 1.7	0.303	6.7 ± 1.8	0.269	7.9 ± 1.6	0.448
		CT/TT	4.6 ± 1.6		6.2 ± 1.8		7.7 ± 1.7	
CTLA4 (rs5742909)	Recessive	CC/CT	4.8 ± 1.7	0.906	6.6 ± 1.8	0.685	7.8 ± 1.6	0.125
		TT	4.9 ± 1.7		6.2 ± 1.6		9.0 ± 1.2	
CTLA4 (rs231775)	Dominant	AA	4.8 ± 1.6	0.612	6.6 ± 1.8	0.628	7.8 ± 1.5	0.755
		AG/GG	4.9 ± 1.8		6.5 ± 1.8		7.9 ± 1.7	
CTLA4 (rs231775)	Recessive	AA/AG	4.8 ± 1.7	0.516	6.6 ± 1.7	0.168	7.9 ± 1.6	0.726
		GG	5.1 ± 1.6		5.9 ± 2.2		8.0 ± 2.0	
TLR3 (rs3775291)	Dominant	CC	4.8 ± 1.5	0.955	6.5 ± 2.0	0.784	8.0 ± 1.8	0.379
		CT/TT	4.8 ± 1.8		6.6 ± 1.6		7.8 ± 1.5	
TLR3 (rs3775291)	Recessive	CC/CT	4.8 ± 1.5	0.858	6.6 ± 1.8	0.448	7.9 ± 1.7	0.733
		TT	4.8 ± 2.5		6.3 ± 1.9		7.8 ± 1.5	
TLR9 (rs5743836)	Dominant	AA	4.8 ± 1.7	0.517	6.5 ± 1.8	0.164	7.8 ± 1.7	0.521
		AG/GG	4.7 ± 1.6		7.0 ± 1.7		8.0 ± 1.4	
TLR9 (rs5743836)	Recessive	AA/AG	4.8 ± 1.7	0.545	6.6 ± 1.8	0.896	7.9 ± 1.6	0.554
		GG	4.5 ± 1.4		6.5 ± 1.2		7.6 ± 1.8	
TLR9 (rs352139)	Dominant	TT	5.1 ± 1.9	0.164	7.1 ± 2.3	0.030	7.9 ± 1.8	0.944
		TC/CC	4.7 ± 1.6		6.4 ± 1.5		7.9 ± 1.6	
TLR9 (rs352139)	Recessive	TT/TC	4.9 ± 1.7	0.468	6.6 ± 1.9	0.835	7.9 ± 1.7	0.850
		CC	4.7 ± 1.6		6.5 ± 1.5		7.8 ± 1.4	
CD209 (rs735240)	Dominant	GG	4.7 ± 2.0	0.472	6.6 ± 1.9	0.935	7.7 ± 1.7	0.498

		GA/AA	4.9 ± 1.6		6.6 ± 1.7		7.9 ± 1.6	
	Recessive	GG/GA AA	4.7 ± 1.7 5.3 ± 1.4	0.034	6.5 ± 1.7 7.0 ± 2.0	0.149	7.8 ± 1.7 7.9 ± 1.4	0.748
CD209 (rs4804803)	Dominant	AA AG/GG	4.9 ± 1.6 4.8 ± 1.8	0.825	6.6 ± 1.8 6.5 ± 1.8	0.657	7.9 ± 1.5 7.8 ± 1.7	0.825
	Recessive	AA/AG GG	4.9 ± 1.6 4.1 ± 2.8	0.331	6.6 ± 1.8 7.0 ± 2.3	0.503	7.9 ± 1.6 8.0 ± 2.0	0.758
	Dominant	CC CT/TT	5.0 ± 1.6 4.6 ± 1.7	0.093	6.8 ± 1.7 6.4 ± 1.9	0.197	8.0 ± 1.6 7.7 ± 1.6	0.145
IFNL3 (rs12979860)	Recessive	CC/CT TT	4.8 ± 1.7 4.8 ± 1.6	0.939	6.7 ± 1.8 6.1 ± 1.7	0.249	7.9 ± 1.6 7.9 ± 1.6	0.981
	Dominant	TT TG/GG	4.9 ± 1.6 4.6 ± 1.9	0.181	6.7 ± 1.8 6.3 ± 1.8	0.205	7.9 ± 1.7 7.7 ± 1.5	0.331
IFNL3 (rs8099917)	Recessive	TT/TG GG	4.8 ± 1.7 5.1 ± 1.3	0.616	6.6 ± 1.8 7.2 ± 2.1	0.449	7.9 ± 1.6 8.2 ± 1.1	0.639
	Dominant	GG GA/AA	5.0 ± 1.6 4.4 ± 1.8	0.061	6.8 ± 1.7 5.8 ± 1.7	0.007	7.9 ± 1.6 7.8 ± 1.7	0.711
TNF (rs1800629)	Recessive	GG/GA AA	4.8 ± 1.7 5.4 ± 1.8	0.574	6.6 ± 1.8 5.7 ± 1.3	0.617	7.9 ± 1.6 8.3 ± 1.6	0.666
	Dominant	TT TG/GG	4.7 ± 1.4 4.8 ± 1.7	0.766	7.0 ± 1.4 6.5 ± 1.8	0.417	7.9 ± 1.4 7.9 ± 1.7	0.813
IL10 (rs1800872)	Recessive	TT/TG GG	4.8 ± 1.6 4.9 ± 1.8	0.808	6.6 ± 1.8 6.6 ± 1.7	0.937	7.8 ± 1.6 8.0 ± 1.7	0.291
	Dominant	GG GC/CC	4.8 ± 1.7 4.9 ± 1.7	0.822	6.6 ± 2.0 6.6 ± 1.6	0.884	7.8 ± 1.8 7.9 ± 1.5	0.907
IL10 (rs1878672)	Recessive	GG/GC	4.8 ± 1.6	0.705	6.6 ± 1.8	0.564	7.9 ± 1.6	0.900

		CC	4.7 ± 2.0		6.3 ± 1.7		7.8 ± 1.8	
IL12B (rs3212227)	Dominant	TT	4.9 ± 1.8	0.403	6.6 ± 2.0	0.959	7.7 ± 1.7	0.098
		TG/GG	4.7 ± 1.6		6.6 ± 1.6		8.0 ± 1.6	
	Recessive	TT/TG	4.8 ± 1.7	0.665	6.5 ± 1.8	0.322	7.9 ± 1.6	0.671
		GG	4.7 ± 1.3		7.1 ± 1.4		8.0 ± 1.9	
IL17A (rs2275913)	Dominant	GG	4.9 ± 1.6	0.586	6.8 ± 1.6	0.163	7.9 ± 1.5	0.946
		GA/AA	4.8 ± 1.8		6.4 ± 1.9		7.9 ± 1.7	
	Recessive	GG/GA	4.9 ± 1.6	0.404	6.6 ± 1.7	0.271	7.9 ± 1.7	0.831
		AA	4.6 ± 1.9		6.1 ± 2.0		7.8 ± 1.4	

AUC: area under the curve; CTLA-4: cytotoxic T-lymphocyte antigen 4; IL: interleukin; SNP: single-nucleotide polymorphism; TLR: toll-like receptor; TNF: tumor necrosis factor; TTV: torque teno virus.

Table S4. Increments (Δ) in TTV DNA levels from baseline to different points after transplantation according to candidate SNPs.

SNP (ID number)	Model	Genotype	Increment (Δ) in TTV DNA levels, log ₁₀ copies/mL (mean \pm SD)									
			Baseline \rightarrow day 15	P- value	Baseline \rightarrow month 1	P- value	Baseline \rightarrow moth 3	P- value	Baseline \rightarrow moth 6	P- value	Baseline \rightarrow moth 12	P- value
CTLA4 (rs5742909)	Dominant	CC CT/TT	2.4 \pm 0.9 2.7 \pm 1.5	0.344	3.5 \pm 1.2 3.2 \pm 0.7	0.487	4.5 \pm 1.8 4.5 \pm 2.1	0.894	4.9 \pm 2.1 5.0 \pm 1.4	0.921	4.3 \pm 1.9 4.6 \pm 1.8	0.537
	Recessive	CC/CT TT	2.5 \pm 1.1 2.0	0.626	3.5 \pm 1.1 2.9 \pm 0.8	0.516	4.5 \pm 1.8 4.2 \pm 2.6	0.850	4.9 \pm 2.0 5.1 \pm 1.7	0.873	4.4 \pm 1.9 4.3 \pm 2.0	0.971
CTLA4 (rs231775)	Dominant	AA AG/GG	2.5 \pm 1.1 2.5 \pm 1.0	0.863	3.5 \pm 1.1 3.4 \pm 1.2	0.819	4.4 \pm 1.8 4.5 \pm 1.8	0.700	4.9 \pm 1.8 5.0 \pm 2.1	0.922	4.4 \pm 1.7 4.4 \pm 2.0	0.936
	Recessive	AA/AG GG	2.5 \pm 1.1 2.4 \pm 0.5	0.802	3.5 \pm 1.2 3.3 \pm 0.8	0.708	4.5 \pm 1.8 3.8 \pm 1.3	0.270	4.9 \pm 1.9 5.4 \pm 2.2	0.428	4.4 \pm 1.9 4.4 \pm 1.8	0.975
TLR3 (rs3775291)	Dominant	CC CT/TT	2.4 \pm 1.1 2.6 \pm 1.0	0.333	3.3 \pm 1.1 3.5 \pm 1.2	0.473	4.6 \pm 1.7 4.3 \pm 1.9	0.442	5.2 \pm 2.1 4.7 \pm 1.8	0.094	4.5 \pm 2.0 4.2 \pm 1.8	0.302
	Recessive	CC/CT TT	2.5 \pm 1.1 2.4 \pm 1.0	0.781	3.5 \pm 1.1 3.3 \pm 1.4	0.666	4.6 \pm 1.9 3.8 \pm 1.3	0.118	5.0 \pm 1.9 4.4 \pm 2.2	0.200	4.4 \pm 1.8 4.1 \pm 2.0	0.543
TLR9 (rs5743836)	Dominant	AA AG/GG	2.4 \pm 1.0 2.7 \pm 1.2	0.257	3.6 \pm 1.1 3.1 \pm 1.1	0.084	4.2 \pm 1.6 5.2 \pm 2.3	0.058	4.8 \pm 1.9 5.2 \pm 2.1	0.257	4.3 \pm 1.9 4.6 \pm 1.9	0.317
	Recessive	AA/AG GG	2.5 \pm 1.0 2.8 \pm 1.5	0.465	3.5 \pm 1.1 2.6 \pm 1.1	0.131	4.5 \pm 1.8 3.8 \pm 1.3	0.390	5.0 \pm 2.0 4.8 \pm 1.8	0.853	4.4 \pm 1.9 3.8 \pm 1.9	0.398
TLR9 (rs352139)	Dominant	TT TC/CC	2.5 \pm 0.8 2.5 \pm 1.2	0.849	3.7 \pm 0.9 3.3 \pm 1.2	0.151	4.7 \pm 1.9 4.4 \pm 1.8	0.366	4.7 \pm 1.8 5.0 \pm 2.0	0.326	4.3 \pm 2.1 4.4 \pm 1.8	0.639
	Recessive	TT/TC CC	2.4 \pm 0.8 2.8 \pm 1.6	0.256	3.5 \pm 1.1 3.3 \pm 1.4	0.513	4.5 \pm 1.7 4.4 \pm 2.0	0.832	5.0 \pm 1.9 4.7 \pm 2.1	0.435	4.3 \pm 1.8 4.7 \pm 1.9	0.241

<i>CD209</i> (rs735240)	Dominant	GG GA/AA	2.4 ± 1.0 2.5 ± 1.1	0.605	3.6 ± 1.4 3.4 ± 1.1	0.475	4.5 ± 2.0 4.5 ± 1.7	0.996	5.0 ± 2.1 4.9 ± 1.9	0.849	4.5 ± 2.1 4.4 ± 1.8	0.759
	Recessive	GG/GA AA	2.4 ± 1.0 2.7 ± 1.4	0.406	3.4 ± 1.1 3.5 ± 1.1	0.731	4.4 ± 1.8 4.5 ± 1.9	0.840	5.0 ± 2.1 4.9 ± 1.6	0.794	4.4 ± 1.9 4.3 ± 1.9	0.696
<i>CD209</i> (rs4804803)	Dominant	AA AG/GG	2.6 ± 1.1 2.4 ± 1.0	0.347	3.4 ± 1.1 3.4 ± 1.2	0.976	4.5 ± 1.9 4.4 ± 1.7	0.628	4.8 ± 1.6 5.2 ± 2.4	0.264	4.1 ± 1.6 4.8 ± 2.1	0.030
	Recessive	AA/AG GG	2.5 ± 1.0 4.7 ± 1.0	0.003	3.4 ± 1.1 -	NA	4.4 ± 1.8 5.0 ± 2.5	0.386	4.9 ± 1.9 5.4 ± 3.1	0.533	4.3 ± 1.8 5.7 ± 4.0	0.535
<i>IFNL3</i> (rs12979860)	Dominant	CC CT/TT	2.5 ± 0.9 2.5 ± 1.2	0.709	3.5 ± 1.1 3.4 ± 1.2	0.851	4.6 ± 1.8 4.3 ± 1.8	0.456	4.9 ± 1.9 4.9 ± 2.0	0.979	4.3 ± 1.7 4.5 ± 2.0	0.510
	Recessive	CC/CT TT	2.6 ± 1.1 2.2 ± 0.7	0.207	3.5 ± 1.2 3.3 ± 0.9	0.602	4.6 ± 1.9 3.6 ± 1.0	0.004	5.0 ± 2.0 4.8 ± 1.9	0.610	4.4 ± 1.8 4.4 ± 2.2	0.923
<i>IFNL3</i> (rs8099917)	Dominant	TT TG/GG	2.4 ± 1.0 2.6 ± 1.2	0.393	3.5 ± 1.1 3.2 ± 1.3	0.237	4.6 ± 1.8 4.2 ± 1.8	0.294	5.0 ± 2.0 4.8 ± 1.9	0.463	4.4 ± 1.9 4.2 ± 1.7	0.560
	Recessive	TT/TG GG	2.5 ± 1.1 3.0 ± 0.3	0.364	3.4 ± 1.1 3.4 ± 1.3	0.881	4.4 ± 1.8 5.7 ± 2.8	0.230	5.0 ± 2.0 4.6 ± 1.3	0.625	4.4 ± 1.9 3.8 ± 1.1	0.515
<i>TNF</i> (rs1800629)	Dominant	GG GA/AA	2.5 ± 1.1 2.3 ± 0.9	0.427	3.5 ± 1.1 3.4 ± 1.1	0.782	4.6 ± 1.7 4.2 ± 2.1	0.379	4.8 ± 1.9 5.3 ± 2.1	0.259	4.3 ± 1.7 4.6 ± 2.3	0.363
	Recessive	GG/GA AA	2.5 ± 1.1 0	NA	3.5 ± 1.1 2.3	0.329	4.5 ± 1.8 4.2	0.862	5.0 ± 2.0 4.3 ± 0.8	0.666	4.4 ± 1.9 2.4 ± 1.2	0.132
<i>IL10</i> (rs1800872)	Dominant	TT TG/GG	2.9 ± 1.3 2.5 ± 1.0	0.168	3.2 ± 1.0 3.5 ± 1.1	0.456	4.9 ± 3.6 4.4 ± 1.6	0.705	4.6 ± 1.6 5.0 ± 2.0	0.433	4.8 ± 1.8 4.3 ± 1.9	0.384
	Recessive	TT/TG GG	2.6 ± 1.1 2.3 ± 1.1	0.272	3.5 ± 1.2 3.4 ± 1.1	0.706	4.6 ± 2.0 4.3 ± 1.5	0.368	4.7 ± 1.7 5.3 ± 2.2	0.076	4.1 ± 1.5 4.7 ± 2.2	0.056
<i>IL10</i> (rs1878672)	Dominant	GG GC/CC	2.6 ± 1.0 2.4 ± 1.1	0.372	3.2 ± 1.0 3.5 ± 1.2	0.290	5.0 ± 2.2 4.2 ± 1.5	0.034	5.1 ± 2.3 4.9 ± 1.8	0.523	4.4 ± 2.1 4.3 ± 1.7	0.799

	Recessive	GG/GC CC	2.5 ± 0.9 2.5 ± 1.6	0.992	3.3 ± 1.0 4.1 ± 1.4	0.018	4.4 ± 1.8 4.9 ± 2.0	0.481	4.8 ± 2.0 5.5 ± 1.9	0.164	4.3 ± 1.8 5.0 ± 2.1	0.082
	Dominant	TT TG/GG	2.6 ± 1.0 2.4 ± 1.1	0.459	3.7 ± 1.1 3.2 ± 1.1	0.105	4.6 ± 1.9 4.4 ± 1.7	0.549	4.6 ± 1.9 5.3 ± 1.9	0.030	4.5 ± 1.9 4.3 ± 1.9	0.548
IL12B (rs3212227)	Recessive	TT/TG GG	2.5 ± 1.1 2.4 ± 0.8	0.830	3.3 ± 1.1 4.4 ± 0.9	0.036	4.4 ± 1.8 5.2 ± 1.7	0.188	4.9 ± 1.9 6.0 ± 2.0	0.096	4.3 ± 1.8 5.0 ± 2.3	0.161
IL17A (rs2275913)	Dominant	GG GA/AA	2.5 ± 0.9 2.5 ± 1.2	0.781	3.6 ± 1.3 3.3 ± 1.0	0.200	4.5 ± 1.8 4.4 ± 1.8	0.708	4.8 ± 2.0 5.0 ± 1.9	0.514	4.4 ± 1.8 4.3 ± 1.9	0.842
	Recessive	GG/GA AA	2.4 ± 1.0 3.0 ± 1.1	0.078	3.4 ± 1.1 3.6 ± 1.0	0.704	4.4 ± 1.7 4.8 ± 2.4	0.460	4.9 ± 1.8 5.5 ± 2.8	0.383	4.4 ± 1.8 4.4 ± 2.0	0.876

CTLA-4: cytotoxic T-lymphocyte antigen 4; IL: interleukin; NA: not applicable; SNP: single-nucleotide polymorphism; TLR: toll-like receptor; TNF: tumor necrosis factor; TTV: torque teno virus.

Table S5. Peak plasma TTV DNA levels at different intervals after transplantation according to candidate SNPs.

SNP (ID number)	Model	Genotype	Peak plasma TTV DNA level, log ₁₀ copies/mL (mean ± SD)					
			First month	P-value	First 3 months	P-value	First 6 months	P-value
CTLA4 (rs5742909)	Dominant	CC	3.8 ± 1.7	0.155	4.9 ± 1.8	0.270	6.0 ± 1.7	0.143
		CT/TT	3.4 ± 1.9		4.6 ± 1.7		5.6 ± 1.7	
CTLA4 (rs5742909)	Recessive	CC/CT	3.7 ± 1.7	0.686	4.8 ± 1.8	0.843	5.9 ± 1.7	0.077
		TT	4.0 ± 1.4		5.0 ± 1.2		7.3 ± 1.3	
CTLA4 (rs231775)	Dominant	AA	3.8 ± 1.6	0.682	4.9 ± 1.7	0.667	5.9 ± 1.7	0.879
		AG/GG	3.7 ± 1.9		4.8 ± 1.8		5.9 ± 1.8	
CTLA4 (rs231775)	Recessive	AA/AG	3.7 ± 1.8	0.483	4.9 ± 1.7	0.335	5.9 ± 1.7	0.691
		GG	4.0 ± 1.4		4.5 ± 2.2		6.1 ± 2.1	
TLR3 (rs3775291)	Dominant	CC	3.7 ± 1.7	0.576	4.8 ± 1.8	0.877	6.1 ± 1.9	0.093
		CT/TT	3.8 ± 1.7		4.9 ± 1.7		5.7 ± 1.5	
TLR3 (rs3775291)	Recessive	CC/CT	3.7 ± 1.6	0.863	4.9 ± 1.7	0.429	6.0 ± 1.8	0.413
		TT	3.7 ± 2.4		4.6 ± 2.2		5.7 ± 1.5	
TLR9 (rs5743836)	Dominant	AA	3.8 ± 1.7	0.308	4.9 ± 1.8	0.640	5.8 ± 1.8	0.264
		AG/GG	3.5 ± 1.8		4.7 ± 1.7		6.1 ± 1.6	
TLR9 (rs5743836)	Recessive	AA/AG	3.7 ± 1.7	0.675	4.9 ± 1.8	0.200	6.0 ± 1.7	0.215
		GG	3.5 ± 1.4		4.1 ± 1.3		5.3 ± 1.5	
TLR9 (rs352139)	Dominant	TT	3.9 ± 2.1	0.547	5.2 ± 2.3	0.101	6.1 ± 1.9	0.516
		TC/CC	3.7 ± 1.5		4.7 ± 1.5		5.9 ± 1.7	
TLR9 (rs352139)	Recessive	TT/TC	3.7 ± 1.8	0.774	4.8 ± 1.9	0.943	5.6 ± 1.9	0.650
		CC	3.7 ± 1.5		4.8 ± 1.5		5.9 ± 1.4	

CD209 (rs735240)	Dominant	GG GA/AA	3.7 ± 2.0 3.7 ± 1.6	0.833	5.0 ± 1.7 4.8 ± 1.8	0.299	5.8 ± 1.7 6.0 ± 1.8	0.578
	Recessive	GG/GA AA	3.6 ± 1.8 4.0 ± 1.5	0.206	4.8 ± 1.8 5.1 ± 1.8	0.271	5.9 ± 1.7 5.9 ± 1.8	0.935
CD209 (rs4804803)	Dominant	AA AG/GG	3.8 ± 1.6 6.7 ± 1.9	0.599	4.9 ± 1.8 4.8 ± 1.7	0.886	5.9 ± 1.7 6.0 ± 1.8	0.673
	Recessive	AA/AG GG	3.8 ± 1.6 3.0 ± 2.8	0.366	4.8 ± 1.8 5.5 ± 1.9	0.211	5.9 ± 1.7 5.9 ± 2.1	0.993
IFNL3 (rs12979860)	Dominant	CC CT/TT	3.9 ± 1.6 3.6 ± 1.8	0.128	5.1 ± 1.6 4.6 ± 1.9	0.044	6.1 ± 1.7 5.8 ± 1.8	0.186
	Recessive	CC/CT TT	3.8 ± 1.7 3.6 ± 1.8	0.633	4.9 ± 1.8 4.4 ± 1.7	0.134	5.9 ± 1.7 6.0 ± 1.8	0.773
IFNL3 (rs8099917)	Dominant	TT TG/GG	3.8 ± 1.7 3.6 ± 1.8	0.379	4.9 ± 1.8 4.6 ± 1.7	0.183	6.0 ± 1.8 5.7 ± 1.7	0.237
	Recessive	TT/TG GG	3.7 ± 1.7 4.0 ± 1.3	0.582	4.8 ± 1.8 5.3 ± 1.7	0.380	5.9 ± 1.8 6.6 ± 1.2	0.235
TNF (rs1800629)	Dominant	GG GA/AA	3.8 ± 1.7 3.4 ± 1.6	0.111	5.0 ± 1.8 4.2 ± 1.5	0.008	6.0 ± 1.7 5.8 ± 1.8	0.590
	Recessive	GG/GA AA	3.7 ± 1.7 4.2 ± 1.9	0.611	4.8 ± 1.8 5.1 ± 0.4	0.799	5.9 ± 1.7 6.5 ± 1.6	0.574
IL10 (rs1800872)	Dominant	TT TG/GG	3.5 ± 1.4 3.8 ± 1.8	0.529	4.7 ± 1.5 4.9 ± 1.8	0.637	5.8 ± 1.3 5.9 ± 1.8	0.702
	Recessive	TT/TG GG	3.7 ± 1.7 3.8 ± 1.7	0.656	4.9 ± 1.7 4.7 ± 1.8	0.442	5.8 ± 1.8 6.2 ± 1.7	0.112
IL10 (rs1878672)	Dominant	GG GC/CC	3.6 ± 1.7 3.8 ± 1.7	0.506	4.9 ± 1.8 4.8 ± 1.8	0.814	5.9 ± 1.9 5.9 ± 1.6	0.926

IL12B (rs3212227)	Recessive	GG/GC CC	3.7 ± 1.7 3.7 ± 2.0	0.992	4.9 ± 1.8 4.5 ± 1.7	0.325	5.9 ± 1.7 5.9 ± 1.9	0.790
	Dominant	TT TG/GG	3.8 ± 1.8 3.7 ± 1.6	0.724	4.9 ± 1.9 4.8 ± 1.6	0.554	5.8 ± 1.8 6.1 ± 1.7	0.192
	Recessive	TT/TG GG	3.7 ± 1.8 3.7 ± 1.4	0.926	4.8 ± 1.8 4.8 ± 1.9	0.838	5.9 ± 1.7 6.0 ± 1.9	0.841
IL17A (rs2275913)	Dominant	GG GA/AA	3.8 ± 1.6 3.7 ± 1.8	0.489	5.0 ± 1.7 4.7 ± 1.8	0.367	5.9 ± 1.5 5.9 ± 1.9	0.951
	Recessive	GG/GA AA	3.8 ± 1.7 3.3 ± 2.1	0.156	4.9 ± 1.8 4.6 ± 1.7	0.481	5.9 ± 1.7 6.0 ± 2.2	0.914

CTLA-4: cytotoxic T-lymphocyte antigen 4; IL: interleukin; SNP: single-nucleotide polymorphism; TLR: toll-like receptor; TNF: tumor necrosis factor; TTV: torque teno virus.