

**Additional File 2.** Comparison of plasma immune checkpoint proteins between subjects who spontaneously cleared HCV (SC group) versus controls (C group).

Marker	Un-adjusted			Adjusted		
	AMR (95%CI)	p-value	q-value	aAMR (95%CI)	p-value	q-value
BTLA	1.37 (1.16–1.63)	<b>&lt;0.001</b>	<b>0.004</b>	1.35 (1.14–1.60)	<b>&lt;0.001</b>	<b>0.004</b>
CD137(4-1BB)	1.46 (1.21–1.77)	<b>&lt;0.001</b>	<b>0.003</b>	1.46 (1.22–1.75)	<b>&lt;0.001</b>	<b>0.002</b>
CD152(CTLA4)	1.39 (1.02–1.88)	<b>0.041</b>	<b>0.083</b>	1.37 (1.00–1.88)	0.054	0.108
CD27	1.38 (1.05–1.81)	<b>0.024</b>	<b>0.057</b>	1.44 (1.11–1.87)	<b>0.008</b>	<b>0.022</b>
CD28	1.38 (1.04–1.82)	<b>0.027</b>	<b>0.059</b>	1.39 (1.06–1.84)	<b>0.023</b>	<b>0.048</b>
CD80	1.46 (1.15–1.85)	<b>0.003</b>	<b>0.011</b>	1.47 (1.17–1.86)	<b>0.002</b>	<b>0.007</b>
GITR	1.54 (1.22–1.94)	<b>&lt;0.001</b>	<b>0.004</b>	1.55 (1.24–1.96)	<b>&lt;0.001</b>	<b>0.003</b>
HVEM	1.33 (1.11–1.59)	<b>0.003</b>	<b>0.011</b>	1.38 (1.16–1.64)	<b>&lt;0.001</b>	<b>0.003</b>
IDO	1.46 (1.17–1.83)	<b>0.002</b>	<b>0.009</b>	1.47 (1.17–1.84)	<b>0.002</b>	<b>0.006</b>
LAG-3	1.46 (1.14–1.87)	<b>0.004</b>	<b>0.011</b>	1.44 (1.12–1.86)	<b>0.007</b>	<b>0.019</b>
PD-1	1.40 (1.13–1.75)	<b>0.004</b>	<b>0.011</b>	1.46 (1.19–1.81)	<b>&lt;0.001</b>	<b>0.004</b>
PD-L1	1.33 (1.06–1.65)	<b>0.015</b>	<b>0.038</b>	1.36 (1.08–1.70)	<b>0.010</b>	<b>0.024</b>
PD-L2	1.84 (1.46–1.33)	<b>&lt;0.001</b>	<b>&lt;0.001</b>	1.82 (1.49–1.23)	<b>&lt;0.001</b>	<b>&lt;0.001</b>
TIM-3	1.42 (1.13–1.78)	<b>0.004</b>	<b>0.011</b>	1.49 (1.20–1.84)	<b>&lt;0.001</b>	<b>0.003</b>
Arginase	0.97 (0.82–1.15)	0.724	0.751	0.96 (0.81–1.15)	0.690	0.743
E-Cadherin	1.31 (0.91–1.87)	0.147	0.229	1.23 (0.87–1.73)	0.251	0.351
MICA	1.11 (0.89–1.39)	0.355	0.452	1.03 (0.82–1.29)	0.773	0.801
MICB	1.10 (0.89–1.34)	0.384	0.461	1.11 (0.91–1.35)	0.324	0.413
NT5E(CD73)	1.15 (0.92–1.44)	0.220	0.307	1.13 (0.91–1.41)	0.282	0.376
Nectin-2(CD112)	1.28 (0.93–1.76)	0.131	0.215	1.25 (0.90–1.73)	0.183	0.285
PVR(CD155)	1.13 (0.89–1.44)	0.304	0.405	1.18 (0.94–1.48)	0.159	0.262
Perforin	1.33 (0.96–1.83)	0.088	0.164	1.31 (0.95–1.81)	0.111	0.195
Siglec-7	1.22 (0.95–1.55)	0.121	0.212	1.15 (1.92–1.44)	0.217	0.320
Siglec-9	1.22 (0.92–1.62)	0.174	0.256	1.30 (1.00–1.71)	0.060	0.111
Tactile(CD96)	0.88 (0.66–1.18)	0.395	0.461	0.91 (0.68–1.21)	0.509	0.594
ULBP-1	0.92 (0.67–1.26)	0.604	0.651	0.89 (0.64–1.24)	0.490	0.594
ULBP-3	0.98 (0.77–1.24)	0.860	0.860	0.97 (0.76–1.24)	0.817	0.817
ULBP-4	1.04 (0.91–1.18)	0.575	0.643	1.03 (0.90–1.19)	0.626	0.701

**Statistics:** Data were calculated by Generalized Linear Models (GLM) with a gamma distribution (log-link). Multivariable models were adjusted by age, sex, *IFN-λ<sub>3</sub>* genotype, and AST, previously selected by a stepwise method (forward) (see **Results Section**). The q-values represent p-values corrected for multiple testing using the False Discovery Rate (FDR). Significant differences are shown in bold.

**Abbreviations:** AMR, arithmetic mean ratio; aAMR, adjusted AMR; 95%CI, 95% of confidence interval; p, level of significance; q, corrected level of significance; BTLA, B and T lymphocyte attenuator; CD, cluster of differentiation; GITR, glucocorticoid-induced TNFR-related; HVEM, herpesvirus entry mediator; IDO, indoleamine 2,3-dioxygenase; LAG-3, lymphocyte activation gene-3; PD-1, programmed cell death protein 1; PD-L1, programmed death-ligand 1; PD-L2, programmed death-ligand 2; TIM-3, T-cell immunoglobulin and mucin-domain containing-3; MICA, MHC class I chain-related gene A; MICB, MHC class I chain-related gene B; NT5E, ecto-5'-nucleotidase; PVR, poliovirus receptor; Siglec, sialic acid-binding immunoglobulin-type lectin; ULBP, human ligand for binding protein.