

Associations between socioeconomic status and bispecific LV20.19 CAR T-cell therapy outcomes

Jennifer M. Knight,^{1,2*} Edward Hackett,^{3*} Aniko Szabo,⁴ Ruizhe Wu,⁴ Garrett Sauber,⁵ Bryon Johnson,⁶ Rachel N. Cusatis,⁷ Elizabeth Aughey,³ Steve W. Cole,⁸ Cecilia J. Hillard^{5#} and Nirav N. Shah^{6#}

¹Department of Psychiatry, Medical College of Wisconsin, Milwaukee, WI; ²Departments of Medicine and Microbiology & Immunology, Medical College of Wisconsin, Milwaukee, WI; ³Medical College of Wisconsin; Milwaukee, WI; ⁴Division of Biostatistics; Medical College of Wisconsin, Milwaukee, WI; ⁵Department of Pharmacology and Toxicology and Neuroscience Research Center, Medical College of Wisconsin, Milwaukee, WI; ⁶BMT and Cellular Therapy Program, Division of Hematology and Oncology, Medical College of Wisconsin, Milwaukee, WI; ⁷Department of Medicine, Division of Hematology and Oncology, Medical College of Wisconsin, Milwaukee, WI and ⁸Departments of Psychiatry and Biobehavioral Sciences and Medicine, Division of Hematology-Oncology, David Geffen School of Medicine at UCLA, Los Angeles, CA, USA

*JMK and EH contributed equally as co-first authors.

#CJH and NNS contributed equally as co-senior authors.

Correspondence: J. M. KNIGHT - jmknight@mcw.edu

<https://doi.org/10.3324/haematol.2022.281957>

Supplementary Tables and Figures

Supplementary Table 1. Peak cytokines

	Low SES Median	High SES Median	Fold Change Ratio	CI	p-value	SES p-value adjusted for LDH
BCA-1	16.78	9.87	6.9	0.9-3.23	0.0941	0.382
Fractalkine	13.33	10.59	2.7	0.57-2.78	0.5476	0.937
G-CSF	14.15	6.36	7.8	1.05-4.73	0.0389	0.245
I-309	8.00	4.62	3.4	1.06-2.82	0.0312	0.208
IFNg	12.18	6.62	5.6	0.69-4.95	0.2046	0.533
IL-6	10.59	5.26	5.3	0.88-4.61	0.0883	0.283
IL-8	9.39	5.26	4.1	1.09-2.94	0.0254	0.152
IP-10	38.09	22.65	15.4	1.12-2.57	0.0198	0.0247
MCP-2	9.97	6.55	3.4	1.03-2.27	0.0391	0.119
TNFa	14.30	7.54	6.8	1.18-3.06	0.0118	0.1218

SE, standard error; SES, socioeconomic status; LDH, baseline lactate dehydrogenase; BCA-1, B-lymphocyte chemoattractant 1; G-CSF, granulocyte colony stimulating factor; IFNg, interferon gamma; IL-6, interleukin 6; IL-8, interleukin 8; IP-10, interferon gamma-induced protein 10; MCP-2, monocyte chemoattractant protein 2; TNFa, tumor necrosis factor alpha. Cytokine mean differences and standard error reported as raw values and t-test was derived using log-transformed values.

Supplementary Table 2. Kynurenine metabolites

	Baseline			D14			D28			D90		
	Ratio* (CI)	p.value	p.value w/LDH**									
TRP	0.869 (0.9-1.47)	0.25	0.261	0.882 (0.87-1.47)	0.332	0.3411	1 (0.78-1.29)	0.999	0.937	0.920 (0.84-1.41)	0.523	0.505
KYN	1.208 (0.64-1.07)	0.145	0.229	1.119 (0.68-1.18)	0.414	0.554	1.117 (0.69-1.17)	0.408	0.523	1.093 (0.69-1.20)	0.514	0.612
KA	0.902 (0.75-1.65)	0.602	0.398	1.027 (0.64-1.48)	0.898	0.776	1.079 (0.62-1.40)	0.711	0.974	0.867 (0.76-1.75)	0.496	0.337
3-HK	1.36 (0.43-1.25)	0.24	0.643	1.324 (0.44-1.30)	0.296	0.745	1.418 (0.41-1.21)	0.193	0.56	1.029 (0.56-1.68)	0.913	0.622
3-HAA	1.766 (0.25-1.28)	0.164	0.438	2.002 (0.21-1.16)	0.103	0.332	2.442 (0.18-0.94)	0.037	0.152	1.486 (0.29-1.57)	0.345	0.688
QA	2.081 (0.30-0.76)	0.004	0.012	2.509 (0.23-0.70)	0.004	0.021	1.939 (0.30-0.88)	0.018	0.052	1.597 (0.33-1.18)	0.135	0.148

TRP, Tryptophan; KYN, Kynurenine; KA, Kynurenic acid; 3-HK, 3-Hydroxylkynurenine; 3-HAA, 3-Hydroxyanthranilate; QA, Quinolinic acid; SES, Socioeconomic Status; LDH, Lactate Dehydrogenase.

Kynurenine metabolites were analyzed using linear mixed effects model and estimated marginal means (least-square means) was done to compare kynurenine metabolites between income groups at each timepoint.

*Ratios are estimated as low SES/high SES

**Linear mixed effect models with estimated means were used to compare groups and a separate analysis was done using LDH as an adjustment to examine potential influence.

Supplementary Table 3. Patient reported outcomes

	Baseline				D14				D28				D90			
	Low SES (SE)	High SES (SE)	p.value	p.value w/LDH	Low SES (SE)	High SES (SE)	p.value	p.value w/LDH	Low SES (SE)	High SES (SE)	p.value	p.value w/LDH	Low SES (SE)	High SES (SE)	p.value	p.value w/LDH
Depression	37.4 (3.73)	36.6 (3.99)	0.884	0.784	32.0 (5.7)	41.1 (3.99)	0.199	0.327	40.0 (3.95)	36.2 (4.25)	0.526	0.489	32.3 (4.54)	30.4 (4.25)	0.77	0.71
Anxiety	16.6 (2.08)	16.1 (2.23)	0.876	0.688	17.2 (2.86)	17.4 (2.23)	0.953	0.83	18.7 (2.17)	19.7 (2.32)	0.759	0.979	17.7 (2.39)	16 (2.32)	0.611	0.979
PSQI	6.25 (1.19)	3.71 (1.28)	0.156	0.121	5.60 (1.9)	3.90 (1.38)	0.472	0.346	6.11 (1.27)	4.43 (1.5)	0.398	0.308	7.11 (1.49)	3.03 (1.5)	0.061	0.05
FSII	3.91 (0.675)	2.68 (0.721)	0.224	0.177	2.72 (1.01)	4.07 (0.721)	0.285	0.502	3.44 (0.711)	3.08 (0.765)	0.73	0.571	3.45 (0.813)	2.66 (0.765)	0.487	0.387
FSIF	2.43 (0.705)	1.63 (0.754)	0.446	0.384	1.28 (1.126)	2.67 (0.754)	0.311	0.483	3.12 (0.751)	1.4 (0.811)	0.128	0.122	1.55 (0.88)	1.06 (0.811)	0.685	0.611
FSID	7.75 (1.72)	7 (1.84)	0.768	0.845	3.49 (2.57)	10.71 (1.84)	0.028	0.046	9.27 (1.81)	7.99 (1.95)	0.632	0.712	7.34 (2.07)	7.34 (2.07)	0.82	0.792
BPII	3.19 (0.717)	1.64 (0.766)	0.154	0.029	1.30 (1.01)	2.10 (0.766)	0.536	0.804	2.96 (0.748)	1.04 (0.804)	0.098	0.018	3.69 (0.837)	0.876 (0.804)	0.021	0.004
BPIF	3.29 (0.824)	1.08 (0.881)	0.0802	0.0384	1.61 (1.17)	1.12 (0.881)	0.742	0.402	2.75 (0.862)	0.779 (0.926)	0.131	0.063	1.86 (0.966)	0.3 (0.926)	0.253	0.132

SES, Socioeconomic Status; SE, Standard Error; PSQI, Pittsburgh Sleep Quality Index; FSII, Fatigue Symptom Index Intensity; FSIF, Fatigue Symptom Index Frequency; FSID, Fatigue Symptom Index Duration; BPII, Brief Pain Index Intensity; BPIF, Brief Pain Index Frequency.