Association of severity of organ involvement with mortality and recurrent malignancy in patients with chronic graft-versus-host disease

Yoshihiro Inamoto,¹ Paul J. Martin,¹ Barry E. Storer,¹ Jeanne Palmer,² Daniel J. Weisdorf,³ Joseph Pidala,⁴ Mary E. D. Flowers,¹ Mukta Arora,³ Madan Jagasia,⁵ Sally Arai,⁶ Xiaoyu Chai,¹ Steven Z. Pavletic,⁷ Georgia B. Vogelsang,⁸ and Stephanie J. Lee¹ on behalf of the Chronic GVHD Consortium

¹Clinical Research Division, Fred Hutchinson Cancer Research Center, Seattle, WA; ²Division of Hematology and Oncology, Mayo Clinic Arizona, Scottsdale, AZ, ³Blood and Marrow Transplant Program, University of Minnesota, Minneapolis, MN; ⁴Blood and Marrow Transplantation, Moffitt Cancer Center, Tampa, FL; ⁵Hematology and Stem Cell Transplant Program, Vanderbilt University Medical Center, Nashville, TN; ⁶Division of Blood and Marrow Transplantation, Stanford University Medical Center, Stanford, CA; ⁷National Cancer Institute, Bethesda, MD; and ⁸Oncology, Johns Hopkins Hospital, Baltimore, MD, USA

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Supplementary Table S1. Comparison of risk categories in the validation set

	New mortality model, no. of visits		
	Low	Intermediate	High
NIH global score, no. of visits			
Mild	106	10*	0
Moderate	123*	219	0
Severe	13*	73*	78

^{*}Visits with different risk categories according to the two models.

Supplementary Table S2. Causes of differences in risk category

NIH global score	New mortality model	Cause of difference	No. of visits	% in total 622 visits
Moderate	Low	Score 2 in skin ± other organs	61	9.8
		Score 2 in liver ± organs other than skin	22	3.5
		Score 1 in more than 2 organs	21	3.4
		Score 2 in eye	12	1.9
		Score 2 in joint/fascia or genital tract	7	1.1
Severe	Intermediate	Lung score 2	27	4.3
		Skin score 3	22	3.5
		Liver score 3	8	1.3
		Lung score 2 + joint/fascia score 3	1	0.2
		Score 3 in other organs	15	2.4
Severe	Low	Eye score 3	10	1.6
		Joint/fascia score 3	2	0.3
		Genital tract score 3	1	0.2
Mild	Intermediate	Gastrointestinal tract score 1	10	1.6

Supplementary Table S3. Associations of organ scores with risk of overall mortality

Skin 0 388 1.00 (reference) 1 246 1.57 (0.96-2.57) 0 2 252 1.93 (1.17-3.18) 0	P HR [†] (95% CI) P 1.00 (reference) .07 .07 1.44 (0.88-2.37) 0.15 .01 2.11 (1.28-3.50) 0.004 .001 3.40 (2.07-5.61) <0.001
0 388 1.00 (reference) 1 246 1.57 (0.96-2.57) 0 2 252 1.93 (1.17-3.18) 0 3 148 3.64 (2.21-6.01) <0	.07 1.44 (0.88-2.37) 0.15 .01 2.11 (1.28-3.50) 0.004
1 246 1.57 (0.96-2.57) 0 2 252 1.93 (1.17-3.18) 0 3 148 3.64 (2.21-6.01) <0	.07 1.44 (0.88-2.37) 0.15 .01 2.11 (1.28-3.50) 0.004
2 252 1.93 (1.17-3.18) 0 3 148 3.64 (2.21-6.01) <0	.01 2.11 (1.28-3.50) 0.004
3 148 3.64 (2.21-6.01) <0	,
	.001 3.40 (2.07-5.61) <0.001
Mouth	
0 407 1.00 (reference)	
1 398 1.32 (0.90-1.95) 0	.16
2 114 1.80 (0.94-3.45) 0	.078
3 17 0.51 (0.07-3.86) 0	.51
Eye	
0 370 1.00 (reference)	
	.77
2 188 1.08 (0.63-1.83) 0	.79
3 49 1.02 (0.39-2.65) 0	.97
GI tract	
0 530 1.00 (reference)	1.00 (reference)
	.006 1.78 (1.13-2.80) 0.013
2-3 [‡] 69 2.01 (0.98-4.12) 0	.058 1.54 (0.75-3.17) 0.24
Liver	<u> </u>
0 441 1.00 (reference)	1.00 (reference)
1 303 1.31 (0.85-2.00) 0	.22 1.31 (0.85-2.02) 0.22
2 144 1.66 (0.94-2.93) 0	.082 1.49 (0.84-2.65) 0.17
3 54 4.71 (2.30-9.66) <0	.001 4.45 (2.14-9.24) < 0.001
Lung	
0 418 1.00 (reference)	1.00 (reference)
1 354 2.18 (1.37-3.48) 0	.001 2.10 (1.31-3.37) 0.002
2 150 4.14 (2.54-6.77) <0	.001 4.41 (2.68-7.27) < 0.001
3 24 6.24 (2.85-13.7) <0	.001 6.24 (2.84-13.7) < 0.001
Joint/fascia	
0 486 1.00 (reference)	
	.93
2 110 1.59 (0.81-3.11) 0	.18
3 14 1.63 (0.22-11.9) 0	.63
Genital tract§	
0 213 1.00 (reference)	
1 83 0.76 (0.26-2.23) 0	.61
2-3 [‡] 53 1.55 (0.68-3.56) 0	.30

^{*}Total number of patients contributing to the score at least at one visit.

[†]Models were adjusted for time after transplantation, transplant center, patient age, stem cell source, disease risk, cytomegalovirus status, HLA and donor type, gender mismatch, conditioning intensity, prior acute GVHD and thrombocytopenia at the visit.

[‡]Scores 2 and 3 were combined since score 3 was documented in small numbers of patients.

[§]Female patients.

Supplementary Table S4. Associations of organ scores with risk of recurrent malignancy

	_	Univariate	
Organ score	No. of patients*	HR [†] (95% CI)	\boldsymbol{P}
Skin			
0	383	1.00 (reference)	
1	244	1.42 (0.83-2.43)	0.21
2	247	1.19 (0.67-2.11)	0.56
3	146	1.20 (0.56-2.59)	0.64
Mouth			
0	401	1.00 (reference)	
1	391	1.53 (0.97-2.40)	0.065
2	111	0.59 (0.18-1.92)	0.38
3	17	1.41 (0.19-10.5)	0.74
Eye			
0	362	1.00 (reference)	
1	340	0.71 (0.44-1.16)	0.17
2-3 [‡]	200	0.38 (0.18-0.77)	0.007
GI tract			
0	522	1.00 (reference)	
1	234	1.25 (0.72-2.17)	0.43
2-3 [‡]	68	1.05 (0.33-3.40)	0.93
Liver			
0	434	1.00 (reference)	
1	297	1.49 (0.93-2.41)	0.10
2	142	1.30 (0.63-2.68)	0.48
3	53	0.64 (0.09-4.72)	0.66
Lung			
0	411	1.00 (reference)	
1	348	1.05 (0.63-1.75)	0.85
2	147	1.45 (0.77-2.71)	0.25
3	23	2.58 (0.79-8.45)	0.12
Joint/fascia			
0	477	1.00 (reference)	
1	250	1.34 (0.82-2.20)	0.24
2-3 [‡]	110	0.68 (0.24-1.91)	0.46
Genital tract§			
0	208	1.00 (reference)	
1	82	0.44 (0.10-1.89)	0.27
2-3 [‡]	52	1.85 (0.68-4.99)	0.23

^{*}Total number of patients contributing to the score at one or more visits.

[†]Models were adjusted for time after transplantation, transplant center and disease risk.

[‡]Scores 2 and 3 were combined since score 3 was documented in small numbers of patients.

[§]Female patients.

Supplementary Table S5. Associations of organ involvement with risk of recurrent malignancy

Involved organ site	No. of patients*	HR [†] (95% CI)	P
Skin	424	1.28 (0.83-1.98)	0.26
Mouth	420	1.39 (0.89-2.18)	0.14
Eye	405	0.58 (0.37-0.90)	0.015
Gastrointestinal tract	269	1.21 (0.72-2.04)	0.47
Liver	363	1.40 (0.90-2.18)	0.13
Lung	402	1.22 (0.77-1.91)	0.39
Joint/fascia	291	1.19 (0.74-1.90)	0.48
Genital tract [‡]	104	0.97 (0.41-2.29)	0.94
Mucosa§	508	1.12 (0.69-1.84)	0.64

^{*}A total number of patients contributing to involvement at least at one visit.

[†]Models were adjusted for time after transplantation, transplant center and disease risk.

[‡]Female patients.

[§]Involvement of either mouth, eyes or genital tract.