# New England Journal of Medicine, 2010, in press. CONFIDENTIAL DOCUMENT—DO NOT COPY OR DISTRIBUTE Supplementary Table 1. Grading of peak acute GVHD severity. 1

GVHD GRADE	Skin Stage <sup>a</sup>		LIVER STAGE <sup>b</sup>		GUT STAGE <sup>c</sup>
1	1 or 2	and	0	and	0
2	3	or	1	or	1
3	4	or	2 - 3	or	2 - 3
4	_		4	or	4

<sup>&</sup>lt;sup>a</sup> Skin stage 1, rash involving <25% of body surface; stage 2, 25-50%; stage 3, >50%; stage 4, generalized erythroderma with bullae

<sup>&</sup>lt;sup>b</sup> Liver stage 1, total serum bilirubin 2-2.9 mg/dL; stage 2, 3-5.9 mg/dL; stage 3, 6-14.9 mg/dL; stage 4,  $\geq$ 15 mg/dL (1 mg/dL=17.1  $\mu$ mol/L).

<sup>&</sup>lt;sup>c</sup> Gut stage 1, diarrhea 500-999 mL/day or biopsy-proven upper gut involvement; stage 2, diarrhea 1000-1499 mL/day; stage 3, diarrhea 1500-1999 mL/day; stage 4, diarrhea > 2000 mL or severe abdominal pain with or without ileus.

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Supplementary Table 2. The Pretransplant Assessment of Mortality (PAM) index and the

score associated with each component, not including the conditioning regimen component (see Methods).<sup>25</sup>

COMPONENT	Score
Age	
< 20 years	1
20-30 years	1
30-40 years	1
40-50 years	1
50-60 years	3
> 60 years	5
Donor Type	
Matched sibling	1
Unrelated	3
Non-sibling or mismatched sibling	4
Disease Risk <sup>a</sup>	
Low	1
Intermediate	8
High	12
Serum Creatinine Level	
$\leq 1.2 \text{ mg/dL}$	1
> 1.2  mg/dL	8
Serum alanine aminotransferase level	
≤ 49 U/L	1
> 49 U/L	2
$\overline{\text{FEV}_1}$	
> 80%	1
70-80%	3
< 70%	6

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Carbon monoxide diffusing capacity	
> 80%	1
70-80%	1
< 70%	4

<sup>a</sup> Low-risk disease includes chronic myeloid leukemia in chronic phase, refractory anemia, aplastic anemia, and inborn errors of the immune system. Intermediate-risk disease includes chronic myeloid leukemia in accelerated phase, chronic myeloid leukemia in remission following treatment of blast crisis, acute leukemia or lymphoma in remission, refractory anemia with excess blasts, chronic lymphcytic leukemia, paroxysmal nocturnal hemoglobinuria. High-risk disease includes chronic myeloid leukemia in blast crisis, juvenile chronic myeloid leukemia,, acute leukemia or lymphoma in relapse, refractory anemia with excess blasts in transition, solid tumors, autoimmune diseases.

#### CONFIDENTIAL DOCUMENT—DO NOT COPY OR DISTRIBUTE Supplementary Table 3. The Hematopoietic Cell Transplant-specific Comorbidity Index

## (HCT-CI) and the score associated with each comorbidity.<sup>26</sup>

Comorbidity	<b>DEFINITION</b>	SCORE
Arrhythmia	Atrial fibrillation or flutter, sick sinus syndrome,	1
	or ventricular arrhythmias	
Cardiac	Coronary artery disease (one or more vessel-	1
	coronary artery stenosis requiring medical	
	treatment, stent, or bypass graft), congestive heart	
	failure, myocardial infarction, or ejection fraction	
	≤ 50%	
Inflammatory bowel disease	Crohn's disease or ulcerative colitis	1
Diabetes	Requiring treatment with insulin or oral	1
	hypoglycemic medication but not controlled with	
	diet alone	
Cerebrovascular disease	Transient ischemic attack or cerebro-vascular	1
	accident	
Psychiatric disturbance	Depression or anxiety requiring psychiatric	1
	consult and/or treatment at the time of HCT	
Hepatic, mild	Chronic hepatitis, total serum bilirubin greater	1
	than the upper limit of normal to 1.5 times the	
	upper limit of normal, or AST/ALT greater than	
	the upper limit of normal to 2.5 times the upper	
	limit of normal.	
Obesity	Patients with a body mass index > 35 kg/m <sup>2</sup> for	1
	adults or with body mass index-for-age percentile	
	of≥95 <sup>th</sup> percentile for children	
Infection	Documented infection or fever of unknown	1
	etiology requiring antimicrobial treatment before,	
	during, and after the start of conditioning	
	regimen	

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Rheumatologic	Systemic lupus erythematosis, rheumatoid	2
	arthritis, polymyositis, mixed connective tissue	
	disease, or polymyalgia rheumatica	
Peptic ulcer	Requiring treatment	2
Moderate/severe renal	Serum creatinine > 2 mg/dL, on dialysis, or prior	2
	renal transplantation	
Moderate pulmonary	DL <sub>CO</sub> and/or FEV <sub>1</sub> 66%-80% or dyspnea on	2
	slight activity	
Prior solid tumor	Treated at any time point in the patient's past	3
	history, excluding non-melanoma skin cancer	
Heart valve disease	Except asymptomatic mitral valve prolapse	3
Severe pulmonary	Carbon monoxide diffusion capacity and/or FEV <sub>1</sub>	3
	≤ 65% or dyspnea at rest or requiring oxygen	
Moderate/severe hepatic	Liver cirrhosis, total serum bilirubin greater than	3
	1.5 times the upper limit of normal, or AST/ALT	
	greater than 2.5 times the upper limit of normal.	

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**Supplementary Table 4. Additional Patient Characteristics** 

CHARACTERISTIC	1993-1997	2003-2007	p-VALUE
	(N=1418)	(N=1148)	
Patient/Donor Gender			0.30
F/F	256 (18%)	234 (20%)	
F/M	307 (22%)	247 (22%)	
M/F	344 (24%)	288 (25%)	
M/M	511 (36%)	379 (33%)	
Patient/Donor CMV Serostatus			< 0.001
~ <del>-</del> -/- <del>-</del> -	417 (29%)	311 (27%)	
+/-	326 (23%)	349 (30%)	
-/+	210 (15%)	118 (10%)	
_/_	465 (33%)	370 (32%)	
Ethnicity			< 0.001
Caucasian	1221 (86%)	899 (78%)	
Hispanic	75 (5%)	52 (5%)	
Native American	17 (1%)	22 (2%)	
African American	33 (2%)	23 (2%)	
Asian	30 (2%)	50 (4%)	
Other	33 (2%)	75 (7%)	
Unknown	9 (1%)	26 (2%)	
Pretransplant serum creatinine <sup>25</sup>			0.001
≤1.2 mg/dL	1355 (96%)	1063 (93%)	
>1.2 mg/dL	63 (4%)	85 (7%)	
Pretransplant serum alanine			<0.001
aminotransferase <sup>25</sup>	1060 (75%)	1020 (89%)	
≤49 U/L	312 (22%)	128 (11%)	
>49 U/L	46 (3%)		
Unknown			

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Pretransplant FEV <sub>1</sub> <sup>25</sup>			0.20
>80%	922 (65%)	750 (65%)	
70-80%	287 (20%)	256 (22%)	
<70%	70 (5%)	54 (5%)	
Unknown	139 (10%)	88 (8%)	
Pretransplant carbon monoxide			< 0.001
diffusing capacity (DL <sub>CO</sub> ) <sup>25</sup>			
>80%	811 (57%)	497 (43%)	
70-80%	365 (26%)	422 (37%)	
<70%	91 (6%)	123 (11%)	
Unknown	151 (11%)	106 (9%)	