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Running title: Factors affecting survival after alloSCT

Factors influencing survival after allogeneic stem cell transplantation for hematologic malignancies in adult patients: A retrospective cohort study

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Allogeneic stem cell transplantation (alloSCT) remains the established main treatment option with 22 23 curative potential for many hematologic malignancies. We conducted a retrospective analysis of 104 adult patients who underwent allogeneic stem cell transplantation between March 2013 and November 24 25 2023. Kaplan–Meier survival analysis, the chi-square test, and Cox regression models were used to identify risk factors and outcomes. The median follow-up of the cohort was 19 (0.3-128.1) months. The 26 median age of the recipients was 49 (19-65) years, and 57 (54.8%) recipients were males. Ninety 27 (86.5%) patients had a matched sibling, and 14 (13.5%) had a haploidentical donor. According to the 28 multivariable analysis, a body mass index (BMI) ≥ 30 kg/m2 (p = 0.02) and status without chronic 29 graft-versus-host disease (cGVHD) (p = 0.04) were significantly associated with worse overall survival. 30 A BMI > 30 kg/m2 was also predictive of worse relapse-free survival (p = 0.01). The cumulative 31 incidence rates of nonrelapse mortality (NRM) and relapse mortality (RM) at 1 year were 8.5% (95%) 32 CI; 4.3-16.5%) and 26.7% (95% CI; 19.1-37.4%), respectively. Patients without cGVHD had 33 significantly higher RM than patients with cGVHD (p < 0.001), whereas patients with cGVHD had 34 significantly higher NRM (p = 0.01). Patients with a BMI ≥ 30 kg/m² had significantly more 35 posttransplant fatal events (p < 0.001). Our analysis revealed that a BMI > 30 kg/m² and a status 36 without cGVHD were significantly associated with worse OS. NRM was higher in patients with 37 cGVHD, whereas patients without cGVHD died mostly from relapses. 38

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Key words: graft versus host disease; cGVHD; allogeneic stem cell transplantation; relapse;
 nonrelapse mortality

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Hematopoietic cell transplantation (HCT) is an established curative procedure for various
hematological diseases. Since the first published report from the European Society for Bone Marrow
Transplantation (EBMT) describing activity in hematopoietic stem cell transplant centers in Europe in

1991, this survey has been published annually by the EBMT and currently includes data from more 47 than 800,000 transplants [1, 2]. The number of HCTs in European centers and collaborating countries is 48 49 increasing, with approximately 20,000 (41%) allogeneic transplantations (alloSCTs), although we can 50 observe the development of novel therapies, especially targeted immunotherapies and chimeric antigen 51 receptor T-cell therapies [2]. Long-term survival and outcomes are affected by considerable early and late transplantation-related mortality (TRM) and morbidity, which are caused mainly by infections, 52 acute and chronic graft-versus-host disease (aGVHD, cGVHD) and toxicity related to conditioning 53 54 regimens. Especially cGVHD, is a particularly devastating syndrome, typically requiring prolonged use of immunosuppressive agents, with a median duration of 2 to 3.5 years [3], cGVHD is therefore 55 56 associated not only with increased late mortality but also with a significant impact on long-term quality of life and functional status among hematopoietic cell transplantation (HCT) survivors [4, 5]. 57 58 Improvements in HLA typing, earlier patient referrals, supportive care, and the development of nonmyeloablative and reduced-intensity conditioning regimens have been achieved over the years, 59 increasing the age limit of alloSCT to above 70 years [6-10]. 60

Our transplantation center was established in 1992 and started with autologous and later alloSCTs with matched related donors, and as of 2021, we have started to perform haploidentical stem cell transplantations (haploSCTs). Since 2021, our institution has reported annual activity to the EBMT Activity Survey. Here, we present our transplantation experiences between 2013 and 2023 concerning alloSCTs for various onco-hematological diseases. We focused on biological and transplantation variables in the cohort, analyzed their impact on survival in recipients after alloSCT, and identified the primary causes of posttransplant mortality.

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69 **Patients and methods**

Study design and data collection. This was a retrospective single-center analysis of all adult patients (aged > 18) who received their first alloSCT for various onco-hematological diseases at the National Cancer Institute in Slovakia from March 2013 until November 2023. This study exclusively included alloSCT recipients from fully matched sibling and haploidentical family donors. All data collection was performed via the National Cancer Institute and University Hospital in Bratislava medical records database of transplanted patients. The study was approved by the Institutional Review Board (Approval No. aloHTC01/2022/a), and a waiver of consent was granted.

The data collected included biological and transplantation recipient and donor characteristics (age, sex,
cytomegalovirus serostatus, body mass index (BMI) at the time of diagnosis, Karnofsky performance

status, HCT-CI, diagnosis and status at transplantation), and transplantation-related factors, including baseline cytomegalovirus (CMV) serostatus of the donor and recipient, intensity of the conditioning regimen, graft source, dose of CD34+ cells, graft versus host disease (GVHD) prophylaxis, incidence and severity of acute and chronic GVHD, length of hospitalization, relapses and mortality.

Definitions. Grading of aGVHD (classic, persistent, recurrent, late-onset) and cGVHD (classic and overlap) was performed according to criteria established at a given time (Przepiorka 1995; Harris 2016, Mount Sinai Acute GVHD International Consortium = MAGIC; National Institutes of Health (NIH) Consensus Development Projects on Criteria for Clinical Trials in Chronic GVHD 2005 and 2014) [11-14]. Patients had to survive more than 100 days to be included in the assessment of cGVHD (87 patients).

In the case of relapses and mortality, we monitored the time from alloSCT to relapse (TTR) and the time from relapse to death (TTD), relapse mortality (RM), and nonrelapse mortality (NRM). NRM was defined as any cause of death without evidence of relapse/persistence/progression, and RM was defined as death from relapse/persistence/progression as the primary or secondary cause of death. OS was defined as the time from alloSCT until death from any cause, and RFS was defined as the time from alloSCT until relapse/progression or death from any cause. Patients alive without evidence of disease relapse, persistence or progression were censored on November 30, 2023.

The myeloablative conditioning regimen consisted of busulfan+cyclophosphamide, 96 cyclophosphamide+total body irradiation (TBI) or fludarabine+TBI. The following reduced-intensity 97 nonmyeloablative regimens were used: fludarabine+melphalan, fludarabine+busulfan or 98 and combinations of cyclophosphamide or fludarabine with a lower dose of TBI. The standard GVHD 99 prophylaxis was cyclosporin (CyA) in combination with methotrexate or mycophenolate mofetil, and 100 five patients were administered rabbit antithymocyte-globulin (ATG). GVHD prophylaxis for 101 102 haploidentical transplant consisted of tacrolimus, MMF and posttransplantation cyclophosphamide.

According to the World Health Organization (WHO) international BMI classification, patients were classified as obese when their BMI was $\geq 30 \text{ kg/m}^2$ [15]. We divided patients into 2 groups: BMI < 30 kg/m² and BMI $\geq 30 \text{ kg/m}^2$.

106 **Statistical methods.** Biological characteristics were described by using an editor, MS Excel v365. We 107 described continuous variables as medians (ranges) and categorical variables as numbers and 108 percentages. The Kaplan– Meier method was used to estimate OS and RFS, and the results are reported 109 as medians with 95% confidence intervals (CIs) and survival percentages.

110 Cumulative incidence functions were used to estimate the incidence of relapse, NRM, and acute and 111 chronic GVHD. The results are reported as hazard ratios (HRs) and 95% confidence intervals (CIs).

For univariable analysis, categorical variables were compared via the chi-square test or Fisher's exact test (as appropriate), and time-to-event variables were compared via the log-rank test. Multivariable analyses were performed via the Cox regression test for all factors with p-values < 0.2 from univariable analysis with potential risks for OS and RFS, and the logistic regression test was used for categorical variables with potential risks of cGVHD and a BMI \geq 30 kg/m². A p-value \leq 0.05 was considered statistically significant. Statistical analyses were performed with Statistical and Power Analysis Software, NCSS 2025.

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120 **Results**

Patient and transplant characteristics. At a median follow-up of 19 (0.3-128.1) months, all 104 transplanted patients were included in the analysis. The median age of the recipients at alloSCT was 49 (19-65) years, and 57 (54.8%) recipients were male. The stem cell source was exclusively peripheral blood in 100% of the patients. The median dose of CD34+ cells used for alloSCT was 4.9×10^6 /kg. The distributions of patients according to diagnosis and other cohort demographic and transplant characteristics are described in Table 1.

Graft-versus-host disease. We recorded aGVHD of any grade in 52 (50%) patients, and the most common grade was Grade II in 19 (18.3%) patients. The distribution of patients according to grade is reported in Table 2, and Figure 1 shows the percentage of patients according to organ involvement for aGVHD. The onset of aGVHD occurred at a median of 54.5 (15-241) days. The cumulative incidence of aGVHD on day +100, grades II-IV, was 25.3% (95% CI: 17.8-35.8%), and that of grades III-IV was 12.8% (95% CI: 7.2-23.0%).

133 Forty-four (50.6%) patients developed cGVHD, with a median time from alloSCT to the onset of cGVHD of 8.6 months (3.6-44.3). At the time of cGVHD onset, 3 (6.8%) patients had overlapping 134 135 cGVHD, and 41 (93.2%) had classic cGVHD. Severe cGVHD appeared in 26 (59.1%) patients. The severity of cGVHD, organ manifestations and NIH scores are shown in Table 2 and Figure 2. The 136 cumulative incidences of cGVHD 1, 2 and 3 years after alloSCT were 46.3% (95% CI; 35.9-59.6%), 137 62.4% (95% CI: 51.0-76.4%) and 65.6% (95% CI: 53.8-79.9%), respectively. Due to the high 138 139 cumulative incidence of cGVHD, we investigated potential correlations between cGVHD and monitored clinical variables (donor type, donor sex, conditioning regimen, complete remission status 140

- before alloSCT, age $< 40 \text{ vs} \ge 40$, age $< 50 \text{ vs} \ge 50$, Karnofsky performance status, BMI, and prior aGVHD) using the chi-square test. However, no statistically significant associations were found.
- 143 Survival. The median OS and RFS after transplantation were 28.6 months (95% CI 17.0-51.6) and 15.1 months (95% CI 10.1-26.2), respectively. The 3- and 5-year OS rates for the entire patient group were 144 145 44.7% (CI 95%; 34.3-55.1%) and 38.6% (CI 95%; 28.0-49.1%), respectively, and the RFS rates were 36.6% (CI 95%; 26.6-46.6%) and 30.5% (CI 95%; 20.5-40.5%), respectively (Figure 3). Univariable 146 analysis revealed several risk factors for worse OS and RFS. Karnofsky score < 90% (p=0.02; p=0.01), 147 status without complete remission before alloSCT (p=0.01; p < 0.001), BMI > 30 kg/m² (p < 0.001; p < 0.001; p < 0.001; p < 0.001), BMI > 30 kg/m² (p < 0.001; p < 0.001; p < 0.001; p < 0.001, BMI > 30 kg/m² (p < 0.001), p < 0.001; p < 0.001; p < 0.001, p <148 (0.001) and status without cGVHD (p=0.01; p=0.03) were associated with worse OS and RFS (Table 3, 149 150 Figure 4). The results of the multivariable analysis for OS and RFS are shown in Table 4. BMI \geq 30 kg/m^2 (p=0.02) and status without cGVHD (p=0.04) were significantly associated with worse OS. A 151 BMI \ge 30 kg/m² (p=0.01) also significantly worsened RFS, whereas it was not confirmed for patients 152 without cGVHD (0.08). A Karnofsky score < 90% and status without complete remission were not 153 predictors of OS or RFS in the multivariable analysis. 154
- Relapse analysis, relapse mortality and nonrelapse mortality. At a median of 5.1 months (1.4-61.6), 35 (33.7%) patients experienced relapse after alloSCT, 18 (17.3%) patients experienced persistence/progression of the disease, and evaluations were not available for 4 (3.9%) patients. The median time to death of patients who experienced relapse after alloSCT was 4.3 months (0.2-103.7). The cumulative incidence rates of relapse were 29.5% (95% CI: 21.3-40.8%) and 35.3% (95% CI: 26.3-47.4%) at 1 and 2 years, respectively.
- At the end of follow-up, 44 (42.3%) patients were alive, and 60 (57.7%) patients had died. Diseaserelated causes of death (relapse/progression/persistence) and nonrelapse mortality are shown in Table 2. In terms of nonrelapse mortality, in addition to infection and GVHD, 4 (6.7%) other causes of death were noted: hepatorenal failure due to drug-induced liver injury, diffuse alveolar damage due to hemorrhage, thrombotic microangiopathy and secondary graft failure. The cumulative incidence rates of 1- year NRM and RM were 8.5% (95% CI; 4.3-16.5%) and 26.7% (95% CI; 19.1-37.4%), respectively.
- Since a BMI \ge 30 kg/m² and status without cGVHD adversely affected patient survival, we analyzed mortality in these patient groups. According to the univariable analysis, obese patients did not have significantly more cases of relapse (p=0.25) or nonrelapse mortality (p=0.08; (infection, GVHD, other)) than patients with a BMI < 30 kg/m², but as a whole group, they had significantly more posttransplant events (p < 0.001).

173 According to the univariable analysis, patients without cGVHD had significantly higher RM than patients with cGVHD did (p < 0.001). We found significantly higher NRM in patients with cGVHD 174 (p=0.01), especially mortality due to infections which was close to statistical significance (p=0.06). The 175 3-year OS in the groups of patients without cGVHD and with mild, moderate, and severe cGVHD was 176 177 38.7% (95% CI: 22.9-54.4%), 100%, 66.2% (95% CI: 41.6-90.8%), and 62.3% (95% CI: 42.6-81.9%), respectively, with a median OS of 18.3 months, not reached, 96.3 months, and 56.5 months. 178 respectively, with evidence of statistical significance (log-rank test p=0.05). The difference in 5-year 179 RFS between the groups of patients was not significant (p=0.11). 180

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182 Discussion

183 This analysis summarizes the real-life data of allogeneic stem cell transplantation from a single center184 in Slovakia.

A recent survey by the European Society for Blood and Marrow Transplantation (EBMT) describing 185 activity in hematopoietic cell transplantation (HCT) centers in Europe reported that the main diagnoses 186 with indications for alloSCT were acute myeloid leukemia (39%), acute lymphoblastic leukemia (17%) 187 and myelodysplastic syndrome or myelodysplastic/myeloproliferative neoplasm overlap (13%) [16]. In 188 our cohort, the most frequently represented diagnoses were AML (33.7%), NHL (16.3%) and ALL 189 (12.5%). The higher number of NHLs in our study reflects the 10-year follow-up, while the EBMT 190 reported annually data. Notably, the number of NHLs indicated for alloSCT in our center decreased 191 192 with the start of CAR-T-cell therapy in 2023.

The cumulative incidence rates of aGVHD at days +100, grades II-IV (25.3%) and grades III-IV 193 (12.8%) correspond to the results of Greinix et al. [17], with reported incidences of 28% and 11%, 194 respectively. Acute GVHD of the skin is reported to be the most affected organ, followed by the GIT 195 196 and liver [18]. The same order of organ involvement in aGVHD was confirmed in our analysis (skin, lower GIT, liver, upper GIT). cGVHD is associated with declines in all aspects of life across all age 197 198 groups. It also leads to increased late mortality [19, 20]. The cumulative incidence of cGVHD at 1 year 199 after alloSCT was 46.3%, which was similar to the 42.3% reported in the study of Langer et al. [18] The 2-year cumulative incidence of cGVHD was 62.4% higher than that reported by Arora et al. (47%) 200 [21]. In that study, correlations between the incidence of cGVHD and the type of donor, CMV 201 202 serostatus, disease diagnosis, and race/ethnicity were found. In our study, chi-square analysis revealed that none of the monitored variables (donor type, donor/recipient sex, conditioning regimen, CR before 203 alloSCT, age, Karnofsky index, BMI, previous aGVHD) were predictive of cGVHD. Potential 204

205 explanations for the higher cumulative incidence of cGVHD could be the 100% use of peripheral blood (PB) as a stem cell source and the limited use of ATG - administered in only 5 patients. The majority of 206 207 sibling transplants (within the total cohort of 104 patients) were performed without ATG, a wellestablished agent for reducing the risk of cGVHD. Following the European Society for Blood and 208 209 Marrow Transplantation (EBMT) guidelines introduced in 2020, we have progressively adopted ATG 210 as a standard component of GVHD prophylaxis in matched related and unrelated donor transplants. 211 We also refer to a multicenter prospective study evaluating the 2-year cumulative incidence of cGVHD, where in the whole cohort, it was 35%, with the highest incidence in PB recipients at 43.7%, compared 212 with 34.4% for bone marrow and 31.6% for cord blood, although the difference was not statistically 213 significant (p=0.26) [22]. In our cohort, patients with cGVHD of any severity had longer 5-year OS 214 than patients without cGVHD. A possible explanation, consistent with previously published data, is 215 that patients with cGVHD had protective graft-versus-tumor effect, as they had significantly lower 216 number of relapses and lower relapse mortality compared to patients without cGVHD [18, 23]. Patients 217 with cGVHD primarily died from nonrelapse causes, especially infections, supporting the notion that 218 cGVHD contributes to transplantation related mortality. This observation is in correlation with the 219 findings of Bhatt et al. who reported that cGVHD was significantly associated with a higher risk of 220 NRM and lower risk of relapse, regardless of age [24]. Furthermore they also reported that severe 221 cGVHD was associated with shorter OS, whereas mild and moderate cGVHD were associated with 222 longer OS. These results are consistent with our findings, where patients with mild and moderate 223 cGVHD had longer 3-year OS compared to those without cGVHD or with severe cGVHD. 224

With respect to 3-year and 5-year OS and RFS, our analysis demonstrated that a BMI $> 30 \text{ kg/m}^2$ was 225 226 significantly associated with worse OS and RFS. We confirmed that this group of patients had more posttransplant events of any type in relation to higher relapse and nonrelapse mortality than did patients 227 with a BMI < 30 kg/m². Doney et al. reported that very obese patients (BMI \ge 35 kg/m²) had increased 228 NRM, which was associated with the intensity of the conditioning regimen, and with long-term follow-229 230 up, they reported increased NRM in obese patients (BMI 30.0-34.9) [25]. In our study, we did not find any correlation between BMI or NRM and the intensity of the conditioning regimen. We agree with 231 Doney et al. that patients need to be informed of their increased risk of NRM based on their BMI and 232 233 need to be controlled in their comorbidities, such as hypertension, diabetes, and hyperlipidemia.

The lower OS and RFS in our cohort may have been affected by the greater proportion of patients without CR before alloSCT and therefore the greater number of patients who died from relapse and persistence/progression in the first year after transplantation. The reported data summarize more than 10 years of activity and outcomes of HSCT within an unselected patient cohort from a single transplantation center in Slovakia. Our analysis has several limitations; thus, the results must be interpreted with caution. These include the retrospective design of the study, the small number of patients in a single institution, slow patient recruitment in a low population country, a nonhomogeneous cohort: various diagnoses, the use of different treatment protocols that change over time according to the availability of innovative drugs, and new treatment possibilities for GVHD.

This study identified important factors that negatively affect overall and relapse-free survival, namely, status without cGVHD and BMI \ge 30 kg/m², and revealed that obese patients had more posttransplant complications associated with mortality. Patients with cGVHD died mostly from nonrelapse causes, especially infections, whereas patients without cGVHD had significantly higher relapse mortality.

The factors affecting OS and RFS are potentially modifiable, but further improvements in transplant practices and supportive care are needed to decrease relapses and other transplant complications, especially cGVHD. Some patients, however, benefit from transplantation. An evaluation of quality of life before and after alloSCT would also be meaningful.

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- 349350 Figure Legends
- 351
- 352 Figure 1. Organ involvement in acute graft-versus-host disease.
- 353
- **Figure 2.** The organ manifestation and NIH scoring of chronic graft-versus-host disease.
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- **Figure 3.** Overall and relapse-free survival of the cohort.

- 358 Figure 4. OS and RFS according to Karnofsky score, status of complete remission before alloSCT,
- BMI and presence of cGVHD.
- 360

Table 1. Patient and transplant character	
Patients' characteristics	N (%)
umber of transplanted patients, N (%)	104 (100%)
ge of recipient/donor, years	
nedian (range)	49 (19-65)/46 (18-69)
< 40, N (%)	28 (26.9%)/32 (30.8%)
2 40, N (%)	76 (73.1%)/72 (69.2%)
ex, recipient/donor, N (%)	
nale	57 (54.8%)/56 (53.8%)
emale	47 (45.2%)/48 (46.2%)
liagnosis, N (%)	
ML	35 (33.7%)
	13 (12.5%)
MDS	6 (5.8%)
NHL	17 (16.3%)
HL	12 (11.6%)
CML	
	4 (3.8%)
other MPN	4 (3.8%)
	5 (4.8%)
MPAL, BPDCN	3 (2.9%)
T-PLL	3 (2.9%)
MM	2 (1.9%)
Karnofsky performance status at the	
ime of alloSCT, N (%)	
00-100%	60 (57.7%)
< 90%	44 (42.3%)
nedian (range)	90 (60-100%)
3MI at alloSCT, N (%)	
$< 18.5 \text{ kg/m}^2$	6 (5.8%)
$18.5-24.9 \text{ kg/m}^2$	50 (48.1%)
$25-29.9 \text{ kg/m}^2$	30 (28.8%)
$\geq 30 \text{ kg/m}^2$	18 17.3%)
baseline CMV serostatus, N (%)	
recipient+ donor+	84 (80.8%)
recipient- donor+	3 (2.9%)
recipient+ donor-	10 (9.6%)
recipient-donor-	7 (6.7%)
Transplant characteristics	<u>(0.776)</u> N (%)
Donor	11 (70)
	00(8650)
MSD	90 (86,5%)
aplo	14 (13,5%)
CT-CI score, N (%)	
)	33 (31.7)
/2	37 (35.6%)
≥ 3	34 (32.7)
conditioning, N(%)	

Table 1. Patient and transplant characteristics of the cohort.

RIC+NMA	63 (60.6%)
GVHD prophylaxis	
MTX based	64 (61.6%)
MMF based	28 (26.9%)
other	12 (11.5%)
disease status before alloSCT, N (%)	
CR	54 (51.9%)
no CR	50 (48.1%)
disease status after alloSCT, N (%)	
CR	80 (76.9%)
no CR	20 (19.2%)
NA	4 (3.9%)

362 Abbreviations: MPAL-mixed-phenotype acute leukemia; BPDCN-blastic plasmacytoid dendritic cell 363 neoplasm; MM-multiple myeloma; T-PLL-T-prolymphocytic leukemia; aGVHD-acute graft-versus-

364 host disease; cGVHD-chronic graft-versus-host disease; CMV-cytomegalovirus; MTX-methotrexate;

365 MMF-mycophenolate mofetil; CR-complete remission

COX

867	Table 2. Characteristics of patients with aGVHD and cGVHD	and causes of death.
	parameter	N (%)
	aGVHD, N (%) according to MAGIC	52 (50%)
	Gr I (% of patients with aGVHD/% of all patients)	12 (23.1%/11.5%)
	Gr II	19 (36.5%/18.3%)
	Gr III	18 (34.6%/17.3%)
	Gr IV	3 (5.8%/2.9%)
	NIH severity of cGvHD at maximum severity,	
	only patients alive after 3 months after alloSCT; (87	
	patients)	44 (50.6%)
	any severity	2 (4.5%)
	mild	16 (36.4%)
	moderate	26 (59.1%)
	severe	
	causes of death	
	relapse and persistence/progression related	42 (70%)
	relapse	28 (46.7%)
	persistence/progression	14 (23.3%)
	non-relapse-related	18 (30%)
	aGVHD	2 (3.3%)
	cGVHD	1 (1.7%)
	infections	11(18.3%)
	others Abbreviations: aGVHD-acute graft-versus-host disease; cGV	4 (6.7%)
869		

characteristic	number of patients	HR (CI 95%)	p-value OS	HR (CI 95%)	p-value RFS
age of recipient					
< 40, N (%)	28	0.76 (95% CI 0.44-1.31)	0.35	0.88 (95% CI 0.53-1.48)	0.64
\geq 40, N (%)	76				
sex, recipient/donor, N (%)					
male	57	1.04 (95% CI 0.62-1.72)	0.89	0.84 (95% CI 0.59-1.54)	0.84
female	47		C		
Karnofsky index (%)		A			
90-100%	60	0.56 (95% CI 0.33-0.94)	0,02	0.55 (95% CI 0.33-0.91)	0.01
< 90%	44				
HCT-CI score, N (%)					
0-2	69	0.83 (95% CI 0.48-1.43)	0.48	1.05 (95% CI 0.63-1.75)	0.85
\geq 3	34				
BMI at alloSCT, N (%)					
$< 30 \text{ kg/m}^2$	86	0.31 (95% CI 0.13-0.71)	< 0.001	0.34 (95% CI 0.15-0.77)	< 0.00
$\geq 30 \text{ kg/m}^2$	18				
conditioning, N (%)					
MAC	41	1.13 (95% CI 0.67-1.9)	0.65	1.10 (95% CI 0.67-1.81)	0.69
RIC+NMA	63				
donor type, N (%)					
match sibling	90	1.07 (95% CI 0.44-2.61)	0,88	1.53 (95% CI 0.71-3.29)	0.35
haploidentical	14				
disease status before alloSCT, N (%)					
CR		0.54 (95% CI 0.33-0.90)	0.02	0.44 (95% CI 0.27-0.72)	< 0.00
non CR	77	0.34 (93% CI 0.33-0.90)	0.02	0.44 (93% CI 0.27-0.72)	< 0.00
	23				
ABO matching					
matched	61				
major mismatched	17		0.70		0.27
minor mismatched	23				
bidirectional	3				
aGVHD, N(%) according to MAGIC		1.23 (95% CI 0.74-2.05)	0.52	1.2 (95% CI 0.74-1.94)	0.45

370 Table 3. Univariate analysis of biological and transplant characteristics and correlations with OS and RFS (log-rank test).

yes					
no	53				
	51				
cGVHD according to NIH, N (%)					
yes	44	0.46 (95% CI 0.26-0.84)	0.01	0.55 (95% CI 0.32-0.94)	0.02
no	44				

Abbreviations: MAC-myeloablative; RIC+NMA-reduced intensity and nonmyeloablative; CR-complete remission; aGVHD-acute graft-371 372 versus-host disease; cGVHD-chronic graft-versus-host disease; OS-overall survival; RFS-relapse-free survival

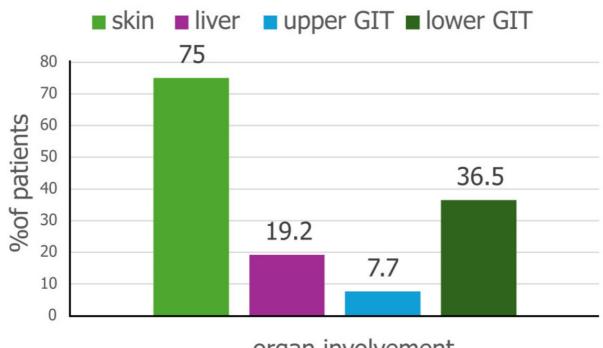
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	373	Table 4. Multivariable	analysis	of transplant	outcomes	(Cox regression test)).
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	OS		RFS	
variable	RR (95% CI)	p-value	RR (95% CI)	p-value
Karnofsky index (%)				
90-100%	1.3 (0.7-2.6)	0.38	1.3 (0.7-2.4)	0.41
< 90%				
BMI at alloSCT, N (%)				
$< 30 \text{ kg/m}^2$	2.6 (1.2-5.6)	0.02	2.6 (1.2-5.5)	0.01
$\geq 30 \text{ kg/m}^2$				
lisease status before alloSCT, N (%)				
CR		0.67	14(0.9.2.0)	0.27
no CR	1.2 (0.7-2.2)	0.67	1.4 (0.8-2.6)	0.27
cGVHD according to NIH, N (%)				
yes	0.5 (0.2, 1, 0)	0.04		0.00
10	0.5 (0.3-1.0)	0.04	0.6 (0.4-1.1)	0.08
Abbreviations: OS-overall survival;	RFS-relapse-free	survival: R	R-risk ratio: BMI-	body mass i
CR-complete remission	~ rempse nee	II , II , II		
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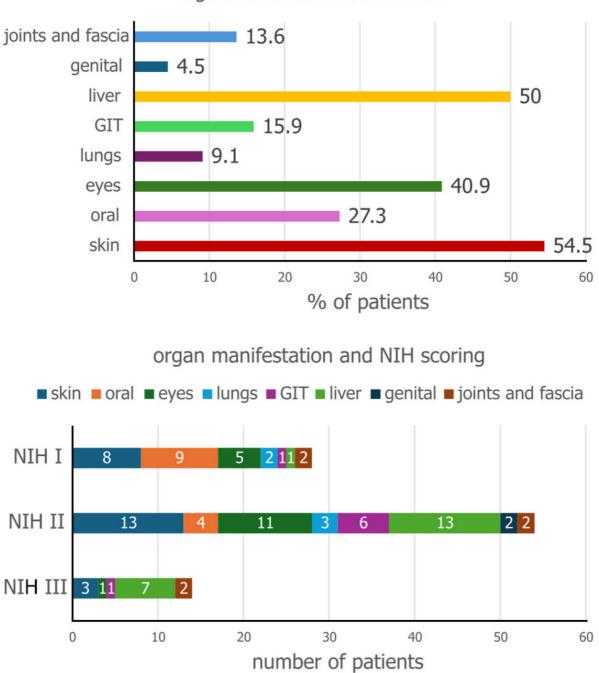
Abbreviations: OS-overall survival; RFS-relapse-free survival; RR-risk ratio; BMI-body mass index;





organ involvement

Fig. 2 Download full resolution image



organ manifestation of cGVHD



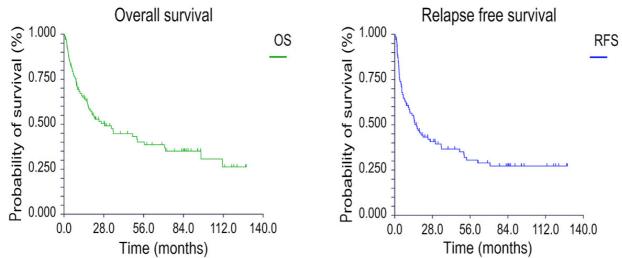


Fig. 4 Download full resolution image

