Better transplant outcome with pre-transplant marrow response after hypomethylating treatment in higher-risk MDS with excess blasts

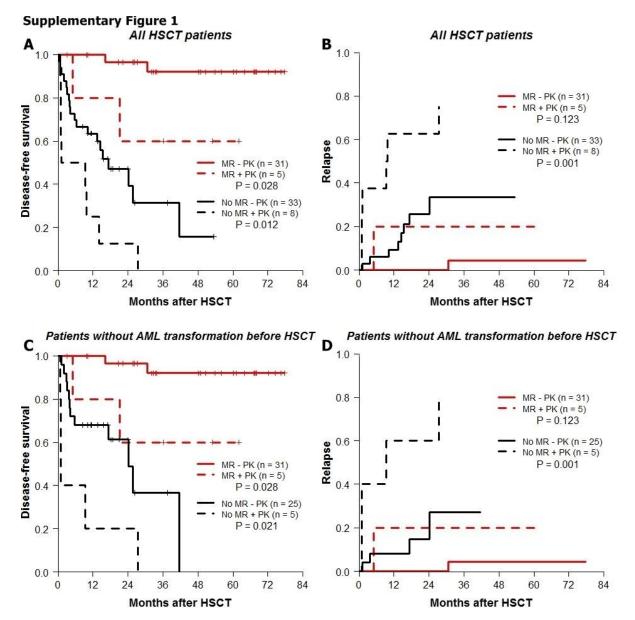
Supplementary Material

Supplementary Table 1 Prognostic factors of marrow response achievement and HMT-related mortality

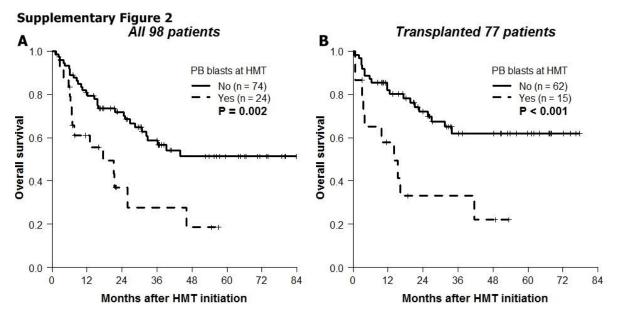
| _ | Achievement of marrow response | | | | HMT-related mortality | | | | | |
|---------------------------------|--------------------------------|------|-----------------------|------------------|-----------------------|------|-----------------------|------|-------------------|------|
| | Univariate analysis | | Multivariate analysis | | Univariate analysis | | Multivariate analysis | | | |
| | E/N | % | P | OR (95% CI) | P | E/N | % | P | OR (95% CI) | P |
| Age | | | | | | | | | | |
| <53 | 29/49 | 59.2 | | | | 2/49 | 4.1 | | | |
| >53 | 21/49 | 42.9 | 0.16 | | | 6/49 | 12.2 | 0.27 | | |
| Sex | | | | | | | | | | |
| Male | 26/61 | 42.6 | | 1 | | 4/37 | 10.8 | | | |
| Female | 24/37 | 64.9 | 0.04 | 0.53 (0.22-1.29) | 0.17 | 4/61 | 6.6 | 0.47 | | |
| Drug | | | | | | | | | | |
| Decitabine | 19/32 | 59.4 | | | | 3/32 | 9.4 | | | |
| Azacitidine | 31/66 | 47.0 | 0.18 | | | 5/66 | 7.6 | 0.71 | | |
| MDS type | | | | | | | | | | |
| De novo | 48/90 | 53.3 | | | | 7/90 | 7.8 | | | |
| Secondary | 2/8 | 25.0 | 0.12 | | | 1/8 | 12.5 | 0.51 | | |
| WHO diagnosis | | | | | | | | | | |
| RAEB-1 | 14/20 | 70.0 | | | | 3/20 | 15.0 | | | |
| RAEB-2 | 33/73 | 45.2 | | | | 5/73 | 6.8 | | | |
| CMMoL-1/2 | 3/5 | 60.0 | 0.12 | | | 0/5 | 0 | 0.59 | | |
| IPSS | | | | | | | | | | |
| Intermidiate-2 | 37/70 | 52.9 | | | | 5/70 | 7.1 | | | |
| High | 13/28 | 46.4 | 0.36 | | | 3/28 | 10.7 | 0.69 | | |
| Cytogenetic risk | | | | | | | | | | |
| Good/Intermediate | 41/75 | 54.7 | | | | 3/75 | 4.0 | | 1 | |
| Poor | 9/23 | 39.1 | 0.24 | | | 5/23 | 21.7 | 0.02 | 4.12 (0.78–21.78) | 0.10 |
| Normal karyotype | | | | | | | | | | |
| Yes | 18/38 | 44.7 | | | | 2/38 | 5.3 | | | |
| No | 33/60 | 55.0 | 0.41 | | | 6/60 | 10.0 | 0.49 | | |
| WBC ($x10^{6}/L$) | | | | | | | | | | |
| < 3000 | 30/54 | 55.6 | | | | 2/54 | 3.7 | | | |
| ≥ 3000/ | 20/44 | 45.5 | 0.42 | | | 6/44 | 13.6 | 0.14 | | |
| ANC (x10 ⁶ /L) | | | | | | | | | | |
| < 1000 | 30/52 | 57.7 | | | | 2/52 | 3.8 | | | |
| ≥ 1000 | 20/46 | 43.5 | 0.22 | | | 6/46 | 13.0 | 0.14 | | |
| Hemoglobin (g/dL) | | | | | | | | | | |
| < 10 | 44/80 | 55.0 | | | | 7/80 | 8.8 | | | |
| ≥ 10 | 6/18 | 33.3 | 0.12 | | | 1/18 | 5.6 | 1.0 | | |
| Platelets (x10 ⁹ /L) | | | | | | | | | | |
| < 100 | 35/73 | 47.9 | | | | 7/73 | 9.6 | | | |
| ≥ 100 | 15/25 | 60.0 | 0.36 | | | 1/25 | 4.0 | 0.46 | | |
| PB blast | | | | | | | | | | |
| No | 43/73 | 58.9 | | 1 | | 4/74 | 5.4 | | 1 | |
| Yes | 7/25 | 28.0 | 0.01 | 0.33 (0.12–0.93) | 0.04 | 4/24 | 16.7 | 0.1 | 3.36 (0.64–17.80) | 0.15 |

| < 15% | 32/57 | 56.1 | 4/57 | 7.0 |
|-------|-------|----------|------|----------|
| ≥ 15% | 18/41 | 439 0.31 | 4/41 | 9.8 0.72 |

HMT, hypomethylating treatment; OR, odds ratio; CI, confidence interval; MDS, myelodysplastic syndrome; WHO, World Health Organization; RAEB-1, refractory anemia of excess blast -1; RAEB-2, refractory anemia of excess blast -2; CMMoL -1/2, chronic myelomonocytic leukemia -1 & -2; IPSS, International Prognostic Scoring System; WBC, white blood cell; ANC, absolute neutrophil count; PB, peripheral blast; BM, bone marrow



Supplementary Figure 1. Posttransplantation outcome according to combined risk groups of marrow response (MR) and poor karyotype (PK). Probability of (A) DFS and (B) cumulative incidence of relapse in all HSCT patients (n = 77); probability of (C) DFS and (D) cumulative incidence of relapse in 66 HSCT patients who did not experience AML transformation before HSCT.



Supplementary Figure 2. Kaplan-Meier analyses for survival from HMT according to the presence of PB blasts at HMT. The influences of PB blasts on HMT response and leukemic transformation were further translated into significant differences in 4-year OS from HMT according to the presence of PB blasts among (A) all higher-risk MDS patients who received HMT (n = 98; 20.5% \pm 11.0% vs. 52.0% \pm 6.9%) and among (B) those who received HSCT following HMT (n = 77; 22.0% \pm 12.6% vs. 61.7% \pm 7.1%).