



## Adult T cell Leukemia and Lymphoma Patients Undergoing Allogeneic Hematopoietic Stem Cell Transplant Risk Evaluation

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### Abstract

Disease status at allogeneic hematopoietic cell transplantation (HCT) is an important pretransplant prognostic factor of HCT in adult T cell leukemia/ carcinoma (ATL); still, other prognostic factors, including comorbidities, weren't prophetic in small cohort analyses. Several scoring systems (HCT-specific comorbidity indicator (HCT- CI)/ modified European Group for Blood and Gist Transplantation threat score (mEBMT)) have been espoused to prognosticate HCT issues in other hematologic malice. We retrospectively estimated HCT- CI and mEBMT to prognosticate nonrelapse mortality (NRM) in 824 ATL cases registered in the Japan Society for Hematopoietic Cell Transplantation TRUMP database, from 2008 until 2013. A advanced HCT- CI was associated with lesser NRM when comparing HCT- CI 0 versus HCT- CI 1 to 3 and HCT- CI 0 versus HCT- CI  $\geq 4$ . An advanced mEBMT score wasn't associated with advanced NRM when comparing mEBMT 0 to 3 with 4 to 6. Because ATL cases are aged and accordingly at threat of fresh complications, we developed an optimized prognostic indicator for ATL (ATL- HCT- PI) using known threat factors age, HCT- CI, and patron – philanthropist coitus combination. The ATL- HCT- PI scores effectively prognosticated the 2- time NRM (22.0, 27.7, and 44.4, independently). Thus, the recently developed ATL- HCT- PI, in combination with other threat factors, is more useful for prognosticating NRM in HCT for ATL cases.

**Keywords:** Allogeneic hematopoietic cell transplantation; Comorbidities; Adult T cell leukemia/ carcinoma; Hematologic malice; Mortality

### Introduction

Adult T cell leukemia/ carcinoma (ATL) is an intractable hematologic malice caused by the idle infection of mortal T cell leukemia contagion type I. The 3- time overall survival (zilches) of ATL cases entering optimal chemotherapy is roughly 24 (2). Allogeneic hematopoietic stem cell transplantation (HCT) is a seductive volition modality that offers long- term absolution in some cases with ATL [1]. A civil retrospective analysis showed that the 3- time OS for the entire cohort was 33(5). Specially, the accretive prevalence of treatment-related mortality (TRM) at 3 times in this cohort was veritably high (roughly). Thus, there's a critical need for clinical results to drop TRM. One possible result involves a more accurate vaticination of the threat of TRM before allo- HCT. Several prognostic indicators for ATL have been reported but were either determined in ATL cases without allo- HCT or have no way been validated in terms of zilches or TRM for cases with ATL entering allo- HCT 6, 7, 8. Hence, it's unclear whether these indicators can be applied to cases with ATL witnessing allo- HCT. The threat factors associated with zilches or TRM include the use of cord blood as a patron source, development of grades III to IV acute graft- versus- host complaint, presence of expansive habitual graft- versus- host complaint, manly coitus, performance status (PS) 2 to 4, late transplantation from opinion, and old age 5, 9, 10, 11, 12. Still, other factors related to pretransplantation conditions, including comorbidities, haven't been completely estimated in cases with ATL witnessing allo- HCT [2].

The HCT-specific comorbidity indicator (HCT- CI) is extensively used to prognosticate TRM. The HCT- CI was established using a large data set and was set up to be reproducible in numerous transplantation settings 14, 15, 16, 17. Still, limited data are available on whether HCT- CI is suitable for use among carcinoma cases entering allo- HCT. By assaying a limited number of non-Hodgkin carcinoma cases (n = 63), Pollack et al. (19) reported that the use of HCT- CI isn't as salutary in non-Hodgkin carcinoma cases as in leukemia cases. In addition, Mori. Reported that the Pre-transplantation Assessment of Mortality

score isn't as useful in cases with ATL entering allo- HCT compared with that in cases with other hematologic malice [3].

In the present study we tested the capability of HCT- CI to prognosticate nonrelapse mortality (NRM) in cases with ATL entering allo- HCT by using the largest ATL allo- HCT registry available worldwide. Also, we tested the mileage of the modified European Group for Blood and Gist Transplantation (mEBMT) threat score and other reported pretransplant threat factors. In addition, we propose a new optimized pretransplant prognostic indicator for cases with ATL entering allo- HCT, called the ATL HCT prognostic indicator (ATL- HCT- PI). This new prognostic indicator more predicts NRM and OS than either the HCT- CI or mEBMT score. To our knowledge this is the largest conformational study of HCT- CI and mEBMT scores with a single complaint reality [4].

### Materials and Method

#### Data collection

ATL cases who passed their first allo- HCT between January 1, 2008 and December 31, 2013 (n = 830), when HCT- CI was introduced into registry data parameters, were considered for this study. Data were collected from the civil check database of the Japan Society for Hematopoietic Cell Transplantation (JSHCT). The data were streamlined as of December 2014. Five cases with missing HCT- CI data

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and 1 case entering autologous HCT were barred. Thus, an aggregate of 824 cases were eventually included in this analysis [5].

Data collected for analysis included the cases' clinical characteristics, similar as HCT- CI parameters ATL clinical subtype age at HCT, coitus, patron coitus, survival data at the last follow- up, PS at HCT, complaint status at HCT, stem cell sources, HLA codifying in serologic resolution, ABO blood group, conditioning authority, date at HCT, patron relationship with the philanthropist (related or unconnected), date of death, and primary cause of death. The study was approved by the data operation panels of the JSHCT and by the institutional ethical commission of Kagoshima University Hospital [6].

## Delineations

Zilches were calculated from the date of HCT until the date of death or last follow- up. The registered causes of death were distributed as either ATL- related or NRM. ATL- related mortality was defined as death with relapse or progression of ATL in cases, grounded on the judgment of each registered croaker. NRM was defined as any death other than ATL- related mortality. However, for illustration, we classified such a case as ATL- related mortality, If the primary cause of death was graft- versus- host complaint or infection after the relapse or progression of ATL. The need for a myeloablative exertion authority and reduced- intensity exertion authority was registered grounded on the judgment of each croaker. An HLA match was defined when the serologic resolution codifying for HLA- A, HLA- B, and HLA- DR was identical [7].

## Statistical analysis

The probability of OS was estimated according to the Kaplan- Meier system. The prevalence of NRM was calculated using accretive prevalence estimates with the contending threat model, while considering ATL- related mortality as a competitive event. Univariate Cox commensurable retrogression analysis was used to estimate the threat factors for zilches. The variables tested in the univariate analysis for transplant issues were PS, coitus, patron – philanthropist coitus combination, ABO mismatch, serologic HLA status, and age. Multivariate analysis was performed while counting for the contending threat structure. Was detected on univariate Cox retrogression analysis and Gray's test. A significance position of was needed in the multivariate analysis. Fine and Gray commensurable hazard modelling was used to estimate the effect of the same variables used in the univariate analysis of NRM and ATL- related mortality on the accretive prevalence of NRM and ATL- related mortality. A backward selection algorithm was used to elect the final model [8].

## Discussion

In the present study we assessed whether the case-, complaint-, and transplantation- related variables and pretransplant comorbidities, represented by HCT- CI and the mEBMT threat score, enhanced the prognostication of NRM, ATL- related mortality, and OS in a cohort of 824 ATL cases treated in Japan between 2008 and 2013. To our knowledge this score has not been anatomized in a large cohort of ATL cases. Our analysis showed that HCT- CI could identify comorbid conditions in only 36.9 of cases, which is fairly lower than that reported

preliminarily (52 to 80). The thickness in comorbidity rendering across different observers remains a concern, because a brief training program is generally proposed for easing dependable assessment of comorbidities. still, a large prospective confirmation study that examined 8115 cases treated with allogeneic HCT reported that 48 of the registered cases had an HCT- CI score of 0 (28). In that cohort cases were aged and further refractory to chemotherapy before allo- HCT. Thus, it's possible that cases with further comorbidity were ineligible to suffer allo- HCT [9].

Pulmonary complaint, hepatic complaint, diabetes, and infection had the loftiest frequency in our cohort, whereas other comorbidities were infrequently observed. Among the comorbidities included in the HCT- CI, only severe pulmonary comorbidities were set up to be prophetic of increased NRM; the remaining comorbidities had no significant prognostic value on NRM by multivariate analysis. When assaying the compound score, we observed a trend for inferior zilches and advanced NRM in cases with advanced HCT- CI. still, prognostic significance wasn't achieved when threat was stratified according to the original report( score 0, low; score 1 to 2, intermediate; score  $\geq$  3, high. nonetheless, when the threat was stratified as score 0, low; score 1 to 3, intermediate; and score  $\geq$  4, high we observed a significantly inferior zilches and advanced NRM in cases with advanced threat. This difference in our population could be attributed to the specific characteristics of our patient cohort [10].

## Conflict of Interest

None

## Acknowledgement

None

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