

Response to Rituximab Induction Is a Predictive Marker in B-Cell Post-Transplant Lymphoproliferative Disorder and Allows Successful Stratification Into Rituximab or R-CHOP Consolidation in an International, Prospective, Multicenter Phase II Trial

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A B S T R A C T

Purpose

The Sequential Treatment of CD20-Positive Posttransplant Lymphoproliferative Disorder (PTLD-1) trial (ClinicalTrials.gov identifier, NCT01458548) established sequential treatment with four cycles of rituximab followed by four cycles of cyclophosphamide, doxorubicin, vincristine, and prednisone (CHOP) chemotherapy as a standard in the management of post-transplant lymphoproliferative disorder (PTLD) and identified response to rituximab induction as a prognostic factor for overall survival. We hypothesized that rituximab consolidation might be sufficient treatment for patients with a complete response after rituximab induction.

Patients and Methods

In this prospective, international, multicenter phase II trial, 152 treatment-naïve adult solid organ transplant recipients, with CD20⁺ PTLD unresponsive to immunosuppression reduction, were treated with four weekly doses of rituximab induction. After restaging, complete responders continued with four courses of rituximab consolidation every 21 days; all others received four courses of rituximab plus CHOP chemotherapy every 21 days. The primary end point was treatment efficacy measured as the response rate in patients who completed therapy and the response duration in those who completed therapy and responded. Secondary end points were frequency of infections, treatment-related mortality, and overall survival in the intention-to-treat population.

Results

One hundred eleven of 126 patients had a complete or partial response (88%; 95% CI, 81% to 93%), of whom 88 had a complete response (70%; 95% CI, 61% to 77%). Median response duration was not reached. The 3-year estimate was 82% (95% CI, 74% to 90%). Median overall survival was 6.6 years (95% CI, 5.5 to 7.6 years). The frequency of grade 3 or 4 infections and of treatment-related mortality was 34% (95% CI, 27% to 42%) and 8% (95% CI, 5% to 14%), respectively. Response to rituximab induction remained a prognostic factor for overall survival despite treatment stratification.

Conclusion

In B-cell PTLD, treatment stratification into rituximab or rituximab plus CHOP consolidation on the basis of response to rituximab induction is feasible, safe, and effective.

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INTRODUCTION

Post-transplant lymphoproliferative disorders (PTLDs) are a serious but rare consequence of immunosuppression after solid organ transplantation (SOT). Their rarity, variety of histologic manifestations, and the complex medical

history of patients with PTLD have slowed the development of evidence-based therapies. For all of the rarer subtypes and in the relapsed or refractory setting, case reports and small case series remain the only source of evidence.^{1,2}

Although the histologic range stretches from polymorphic PTLD to monomorphic lymphoma-type PTLD, the majority of cases are of CD20⁺

ASSOCIATED CONTENT

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B-cell lineage.³ In pediatric CD20⁺ PTLD, favorable results have been reported in a phase II trial of rituximab, cyclophosphamide, and corticosteroids.⁴ Through international cooperation, we have been able to assemble adult patient cohorts large enough for meaningful first-line therapy trials. The phase II Sequential Treatment of CD20-Positive Post-Transplant Lymphoproliferative Disorder (PTLD-1) trial recruited 70 patients from 2003 to 2007 and established sequential treatment (ST) with four cycles of weekly rituximab followed by four cycles of chemotherapy with cyclophosphamide, doxorubicin, vincristine, and prednisone every 21 days (CHOP-21) as a standard in CD20⁺ PTLD after SOT.⁵ Median overall survival (OS) was 6.6 years, a clear improvement over the preceding smaller rituximab monotherapy trials (1.2 to 3.5 years).⁶⁻⁸ Toxicity, particularly treatment-related mortality (TRM), was 13%, thus lower than in the preceding retrospective case series of first-line chemotherapy in PTLD (up to 31%).^{5,9-15}

We observed that response to four cycles of rituximab induction was a prognostic factor for OS after completion of ST.⁵ On this basis, we hypothesized that rituximab consolidation might be sufficient treatment for patients with a complete response (CR) after rituximab induction. The PTLD-1 protocol was therefore amended in 2006 to introduce risk-stratified sequential treatment (RSST) with rituximab consolidation for patients in CR after rituximab induction. Treatment of patients not in CR after four weekly cycles of rituximab was changed from CHOP-21 to rituximab plus cyclophosphamide, doxorubicin, vincristine, and prednisone every 21 days (R-CHOP-21). The rationale for the latter had several components; large trials in immunocompetent patients with diffuse large B-cell lymphoma (DLBCL) demonstrated a higher efficacy of R-CHOP than CHOP.^{16,17} Moreover, safety concerns with regard to the use of R-CHOP in immunosuppressed patients at the time the protocol for PTLD-1 was developed in 2002 started to be allayed by 2006.¹⁸ The goal of this trial was to demonstrate the feasibility, safety, and efficacy of RSST on the basis of patient response to rituximab induction.

PATIENTS AND METHODS

In 2006, after inclusion of 70 patients, the second planned interim analysis of the PTLD-1 trial (ClinicalTrials.gov identifier, NCT01458548) was performed, and response to four courses of rituximab was identified as a prognostic factor for OS.⁵ The protocol was amended to introduce RSST, the results of which are reported in this article. The trial design outside the treatment schedule remained unchanged. The trial was stopped after it had reached its target recruitment (225 patients in total, 150 treated with RSST).

Study Design and Patients

An international, prospective, multicenter, open-label, phase II trial was performed at 32 centers in Germany, Belgium, France, Australia, Poland, and Italy. Treatment-naïve adult SOT recipients diagnosed with CD20⁺ PTLD were enrolled after activation of the amendment in their participating country from October 24, 2006, until October 3, 2014. Inclusion and exclusion criteria remained unchanged from the original PTLD-1 trial⁵ and also included response failure to upfront immunosuppression reduction (with or without antiviral therapy), measurable disease > 2 cm in diameter (and/or bone marrow involvement), and an Eastern Cooperative Oncology Group performance status ≤ 2. The extent and duration of upfront immunosuppression reduction were at the

discretion of the treating physician, but usually calcineurin inhibitors were reduced by 30% to 50%, and azathioprine or mofetil mycophenolate were stopped. Response failure to immunosuppression reduction was defined as stable disease at 2 to 4 weeks after immunosuppression reduction or as progressive disease at any time. The main exclusion criteria were CNS involvement, a history of HIV infection, and the presence of severe organ dysfunction not related to PTLD.

Diagnostic tissue samples were reviewed by an expert hematopathologist and classified according to 2004 WHO criteria. Epstein-Barr virus (EBV) association was confirmed by in situ hybridization for EBV-encoded small RNA transcripts. Disease stage at enrollment was determined through a complete patient history; physical examination; laboratory investigations (including full blood count, lactate dehydrogenase [LDH] activity and renal and liver function tests); bone marrow biopsy findings; cerebrospinal fluid analysis; and computed tomography (CT) scans of the head, chest, and abdomen. The responsible local ethics committees approved the trial, and all patients gave written informed consent according to the Declaration of Helsinki.

Treatment Plan

Treatment consisted of rituximab (375 mg/m² intravenously [IV]) on days 1, 8, 15, and 22 followed by interim staging by CT scan (days 40 to 50; Fig 1). Starting on day 50, patients with CR at interim staging (low-risk group) continued with four courses of rituximab monotherapy (375 mg/m² IV) every 21 days, whereas all others (high-risk group) received four cycles of R-CHOP-21 (rituximab 375 mg/m² IV on day 1, cyclophosphamide 750 mg/m² IV on day 1, doxorubicin 50 mg/m² IV on day 1, vincristine 1.4 mg/m² [maximum, 2 mg] IV on day 1, and prednisone 50 mg/m² orally on days 1 through 5, every 21 days). In case of clinical signs of disease progression at any time during rituximab monotherapy or before interim staging, restaging was performed prematurely, and R-CHOP-21 was commenced immediately if disease progression was confirmed. Supportive treatment with granulocyte colony-stimulating factor after R-CHOP-21 chemotherapy was obligatory. *Pneumocystis jirovecii* chemoprophylaxis was recommended. The final

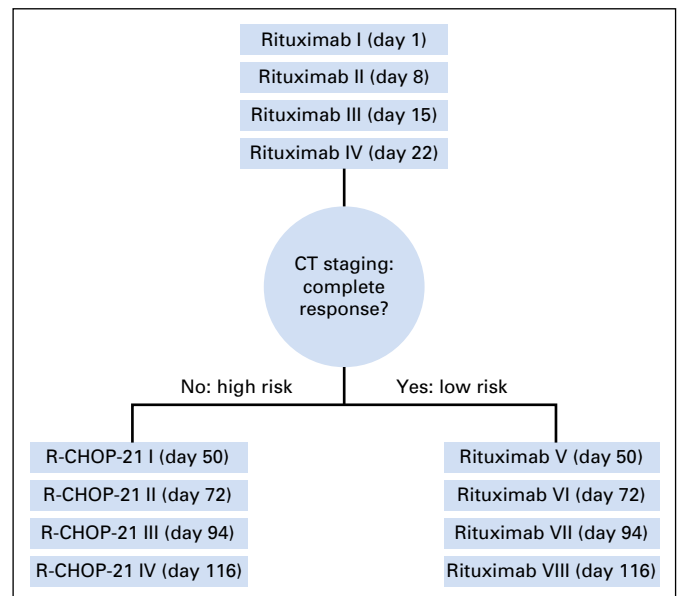


Fig 1. Risk-stratified sequential treatment schedule. Rituximab signifies rituximab 375 mg/m² intravenously (IV), R-CHOP-21 signifies rituximab 375 mg/m² IV on day 1 plus cyclophosphamide 750 mg/m² IV on day 1, doxorubicin 50 mg/m² IV on day 1, vincristine 1.4 mg/m² (maximum, 2 mg) IV on day 1, and prednisone 50 mg/m² orally on days 1 through 5, every 21 days. In case of progressive disease from day 1 through day 50, patients proceeded to R-CHOP-21 immediately. CT, computed tomography.

response assessment was performed 1 month (± 7 days) after the last cycle of therapy. Subsequently, patients underwent follow-up examinations every 3 months for 2 years, every 6 months for years 3 through 5, and annually thereafter. Interim, final response, and follow-up assessments included a complete patient history, physical examination, laboratory investigations, and CT scans of the chest and abdomen. Further investigations, such as bone marrow biopsy, CT scans of the head, or endoscopy, were performed if clinically indicated to determine remission status. Follow-up data were evaluated up to July 2015, with a median follow-up of 4.5 years.

Statistical Analysis

The primary end point was treatment efficacy measured as response rate in patients who completed therapy and response duration (RD) in those who completed therapy and responded. Secondary end points were frequency of infections, TRM, OS, and time to progression (TTP) in the intention-to-treat (ITT) population. Response to treatment and disease progression were classified according to WHO criteria using CT imaging. RD was defined from the date of best response (CR or partial response) to disease progression, whereas TTP was defined from start of treatment to disease progression (all patients). OS was defined from start of treatment to death attributable to any cause. Adverse events and serious adverse events were documented according to the WHO toxicity grading scale. Analysis was by ITT.

CI and best point estimates for observed response rates were calculated using the adjusted Wald method. Time-to-event outcomes were described using Kaplan-Meier statistics. Exploratory analyses were performed using two-sided stratified log-rank tests as well as χ^2 tests for categorical variables, and the independent samples Kruskal-Wallis test was used for continuous variables. Multivariable analyses were performed with Cox regression models (log-rank ratio test, backward elimination). The two-sided significance level was set at .05, and SPSS 22.0.0.0 statistical software (IBM Corporation, Chicago, IL) was used for all analyses. The results of the 70 patients treated with the original PTLD-1 trial protocol (the ST cohort)⁵ and its subgroups (on the basis of rituximab response) were used for post hoc comparisons of efficacy, survival, and toxicity.

RESULTS

Patients

One hundred fifty-two patients were enrolled at centers in Germany (72 patients), Belgium (36 patients), France (24 patients), Australia (seven patients), Poland (seven patients), and Italy (six patients). Their baseline characteristics are listed in Table 1. Median age was 56.4 years (range, 18 to 82 years). Sixty-nine patients had undergone kidney transplantation, 40 patients had undergone liver transplantation, 18 patients had undergone lung transplantation, 15 patients had undergone heart transplantation, five patients had undergone heart and kidney transplantation, three patients had undergone kidney and pancreas transplantation, and two patients had undergone heart and lung transplantation. Median time from transplantation to PTLD was 9.0 years. Most patients (112 of 152 [74%]) were diagnosed with monomorphic DLBCL-type PTLD, 67 of 144 (47%) had EBV-associated tumors and 101 of 151 (67%) had Ann Arbor Conference classification of disease stage III or IV. Ninety-seven (65%) of 150 patients had an elevated serum LDH activity at diagnosis, and 55 (38%) of 143 had an international prognostic index (IPI) score of ≥ 3 (risk factors are age > 60 years, Ann Arbor stage \geq III, Eastern Cooperative Oncology Group performance status ≥ 2 , elevated LDH, and more than one extranodal disease manifestation).¹⁹ Four patients were

Table 1. Baseline Characteristics of Patients Enrolled (intention-to-treat population [n = 152])

Characteristic	No. (%)
Median age (range), years	56.4 (18-82)
≥ 60 years of age	60 (40)
Male	115 (76)
Transplant type	
Kidney	69 (45)
Liver	40 (26)
Lung	18 (12)
Heart	15 (10)
Heart and kidney	5 (3)
Kidney and pancreas	3 (2)
Heart and lung	2 (1)
Median time from transplantation to PTLD (range), years	9.0 (0.2-27.9)
< 1 year	32 (21)
≥ 1 year	120 (79)
Histology	
Early lesion	2 (1)
Polymorphic	20 (15)
Monomorphic	129 (85)
Burkitt	6 (4)
DLBCL	112 (74)
Other B-cell, CD20 ⁺	8 (5)
Other B-cell, CD20 ⁻ *	3 (2)
Multicentric Castleman disease*	1 (1)
EBV association (n = 144)	
EBV associated	67 (47)
Non-EBV associated	77 (53)
Ann Arbor Conference classification of disease stage (n = 151)	
I	30 (20) [†]
II	20 (13)
III	22 (15)
IV	79 (52)
Lactate dehydrogenase (n = 150)	
Within normal range	53 (35)
Elevated	97 (65)
Nodal disease (n = 151)	110 (73)
Extranodal disease (n = 151)	108 (72)
GI	43 (28)
Liver	34 (23)
Lung	26 (17)
Kidney	4 (3)
Bone marrow	12 (8)
Graft	13 (9) [‡]
International prognostic index (n = 143)	
< 3	88 (62)
≥ 3	55 (38)
ECOG performance status (n = 144)	
0	40 (28)
1	66 (46)
2	32 (22)
3	6 (4)

Abbreviations: DLBCL, diffuse large B-cell lymphoma; EBV, Epstein-Barr virus; ECOG, Eastern Cooperative Oncology Group; PTLD, post-transplant lymphoproliferative disorder.

*Diagnosis changed upon pathology review.

[†]This includes 21 patients in stage IE.

[‡]Eight of 13 were patients who had undergone lung transplantation.

reclassified with a diagnosis other than CD20⁺ PTLD on pathology review.

Treatment

Of the 152 patients enrolled, one died before the start of treatment. One hundred forty-eight patients could be evaluated for

response to rituximab induction, 134 of whom had received all four scheduled applications (Fig 2). Thirty-seven (25%) of 148 patients achieved CR at interim staging and were allocated to rituximab monotherapy consolidation in the low-risk group. Three of these patients did not receive further treatment—one patient choose to withdraw from further treatment, one was withdrawn after GI perforation, and one died as a result of pulmonary hemorrhage. Thus, 34 patients received rituximab monotherapy consolidation. Of the 111 patients who were not in CR after rituximab induction (high-risk group), 100 went on to receive treatment with R-CHOP-21. Two patients died before treatment continuation (carotid perforation and liver abscesses). Nine patients were withdrawn from treatment because of: progressive disease that involved the CNS (two patients); renal failure (two patients); physician choice in favor of radiotherapy (two patients, both of whom in partial response); and GI perforation, hepatitis B viral infection, and hypokinetic cardiomyopathy (one patient each). Ninety-two patients could be evaluated for response to R-CHOP-21. Four patients died during therapy. Three patients were withdrawn from therapy as a result of infectious complications, and one patient was lost to follow-up.

Although early PTLD, EBV association, and low baseline IPI were significantly more common in the low-risk group than in the high-risk group (Data Supplement), 31 of 37 patients in the low-risk group had monomorphic PTLD, 21 of 37 had late PTLD, 13 of 36 had EBV-negative tumors, and eight of 34 had an IPI ≥ 3. Of note, six of 18 patients with PTLD who had undergone lung transplantation, a subgroup with historically poor OS,²⁰ were allocated to the low-risk group.

Outcome

The overall response rate (ORR) of RSST was 88% (111 of 126 patients; 95% CI, 81% to 93%) and the CR rate was 70% (88 of 126; 95% CI, 61% to 77%). Median RD (Fig 3A) was not reached;

the 3-year Kaplan-Meier estimate was 82% (95% CI, 74% to 90%). In the ITT population (152 patients), median TTP (Fig 3B) was not reached. The 3-year Kaplan-Meier estimate was 75% (95% CI, 67% to 82%). Median OS (Fig 3C) was 6.6 years (95% CI, 5.5 to 7.6 years) with a 3-year estimate of 70% (95% CI, 62% to 77%). These results were confirmed by a per-protocol analysis (Data Supplement).

Toxicity

Fifty-seven (63%) of 91 patients experienced grade 3 or 4 leukopenia (95% CI, 52% to 72%; no repeat blood counts in 60 patients), whereas 52 (34%) of 151 patients experienced grade 3 or 4 infections (95% CI, 27% to 42%). The most common infection experienced by patients was febrile neutropenia (24 patients), whereas *Clostridium difficile* colitis, *P jirovecii* pneumonia (PcP), and invasive aspergillosis were experienced by three patients each. At least two of the patients with PcP did not receive prophylaxis, and two of those who experienced PcP were low-risk patients. Twelve (8%) of 151 patients experienced treatment-related mortality (95% CI, 5% to 14%). Five patients died as a result of infections, two each from hemorrhage and the sequelae of GI perforation and one as a result of an unknown cause. During the follow-up period, one patient experienced fatal progressive multifocal leukoencephalopathy and one patient experienced secondary acute myeloid leukemia. Only five of 52 patients who experienced grade 3 or 4 infections were in the low-risk group, and all but one treatment-related death occurred in the high-risk group.

Prognostic Factors

Response to four applications of rituximab was a highly significant predictor of TTP and OS despite treatment stratification (n = 148; both P < .001; Data Supplement). We can confirm the significance of the baseline IPI (< 3 or ≥ 3) previously reported as

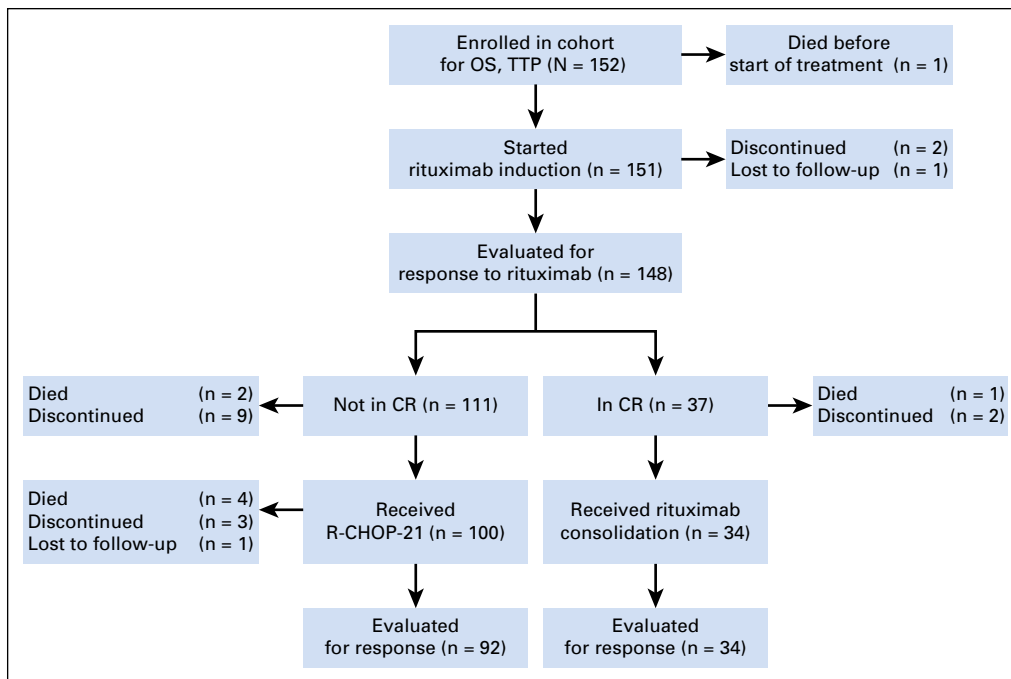


Fig 2. Diagram of number of patients enrolled, treated, and evaluated for response. CR, complete response; ITT, intention to treat; OS, overall survival; R-CHOP-21, rituximab plus cyclophosphamide, doxorubicin, vincristine, and prednisone; TTP, time to progression.

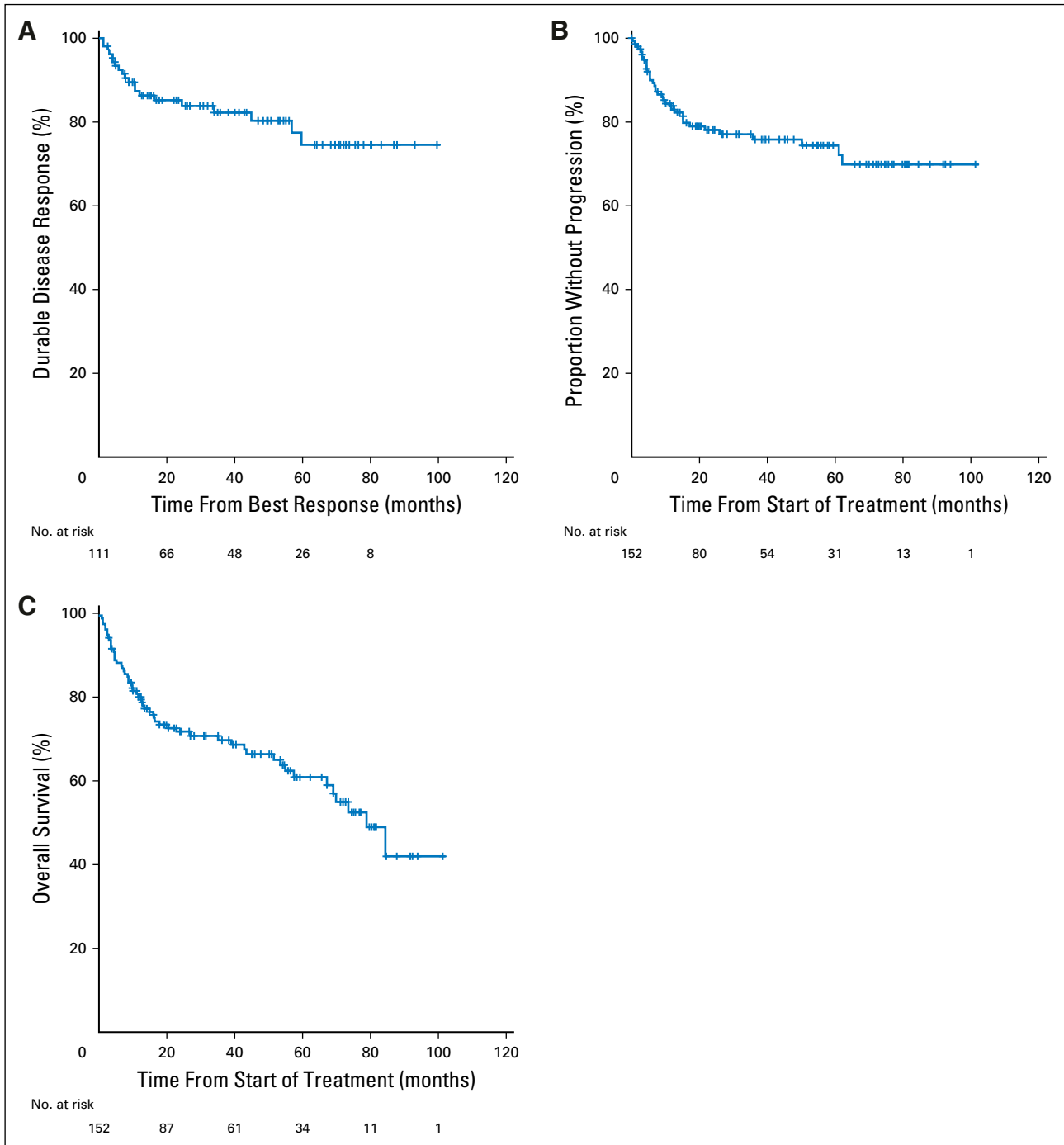


Fig 3. Response duration, time to progression, and overall survival. Median time of follow-up was 4.5 years. (A) Response duration (patients in complete response or partial response). (B) Time to progression (all patients). (C) Overall survival (all patients).

a significant prognostic factor for OS in PTLD-1 ST²¹ in the RSST cohort for TTP and OS (complete IPI data available in 143 patients; $P = .001$; Data Supplement). On the other hand, there was no significant difference in ORR between EBV-positive and EBV-negative PTLD (48 [86%] of 56 patients and 59 [92%] of 64 patients; $P = .255$). No significant differences in TTP ($P = .908$) or OS ($P = .793$) were found (Data Supplement). In a multivariable analysis (Data Supplement), both response to four applications of

rituximab and the baseline IPI (< 3 or ≥ 3) were highly significant independent prognostic factors for TTP and OS.

Comparison With PTLD-1 ST

Baseline characteristics of both trial cohorts were similar, and the only significant difference was time from transplant to PTLD (Data Supplement). The overall response rate of RSST was 111

(88%) of 126 patients compared with 53 (90%) of 59 patients in the PTLD-1 ST cohort.⁵ Median OS was identical (6.6 years), and 3-year Kaplan-Meier estimate was 70% (95% CI, 62% to 77%) compared with 61% (95% CI, 49% to 72%) in PTLD-1 ST. The comparisons for RD (3-year estimates, 82% [95% CI, 74% to 90%] ν 74% [95% CI, 62% to 86%]) and TTP (3-year estimates, 75% [95% CI, 67% to 82%] ν 69% [95% CI, 57% to 80%]) were favorable. The frequency of both grade 3 or 4 infections (34% ν 41%) and TRM (8% ν 13%) were lower in RSST.

Low-Risk Group and Comparison With PTLD-1 ST

The TTP estimate in the low-risk rituximab consolidation group was 89% (95% CI, 76% to 100%) at 3 years compared with 69% (95% CI, 44% to 95%) in the 14 patients in PTLD-1 ST who had reached CR with rituximab induction and continued ST with CHOP chemotherapy (Fig 4A). OS in these two cohorts was similar, with 3-year Kaplan-Meier estimates of 91% (95% CI, 82% to 100%) and 86% (95% CI, 67% to 100%), respectively (Fig 4B). An analogous comparison of the high-risk R-CHOP consolidation group with patients in the PTLD-1 ST group who had not reached CR after rituximab induction can be found in the Data Supplement.

DISCUSSION

When published in 2012, the PTLD-1 ST trial with 70 patients had been the largest prospective trial in PTLD and demonstrated an

unprecedented median OS of 6.6 years.⁵ We present the results of a prospective trial with more than twice as many patients recruited in six countries from a wide range of clinical settings.

The results of the 70 patients treated with ST in the PTLD-1 trial from 2003 to 2007 provide a suitable benchmark. Despite the limiting of chemotherapy to the high-risk group, the ORR of 88% and median OS of 6.6 years of RSST closely match the results of ST, where all patients received CHOP chemotherapy. Furthermore, the Kaplan-Meier estimates of RD, TTP, and OS compare favorably, and the infection and mortality safety parameters were lower in the RSST cohort.

The 3-year TTP of 89% (95% CI, 76% to 100%) in the low-risk rituximab consolidation group confirmed the key hypothesis of this protocol—A CR to rituximab induction identifies a group of patients with B-cell PTLD who do not need chemotherapy. This is further supported by our observation that response to rituximab monotherapy is a predictive marker for OS and TTP.

The safety profile of RSST was favorable. TRM was 8% and thus comparable to that reported in immunocompetent patients with DLBCL older than 60 years of age (7% with six cycles of R-CHOP every 14 days in RICOVER-60 [Six Versus Eight Cycles of Biweekly CHOP-14 With or Without Rituximab in Elderly Patients With Aggressive CD20+ B-Cell Lymphomas] and 6% with eight cycles of CHOP-21 with or without rituximab in LNH98.5 [CHOP Chemotherapy Plus Rituximab Compared With CHOP Alone in Elderly Patients With DLBCL]).^{16,22} R-CHOP immunochemotherapy in the high-risk patients did not result in excess toxicity or mortality. We conclude that R-CHOP, the proven standard in

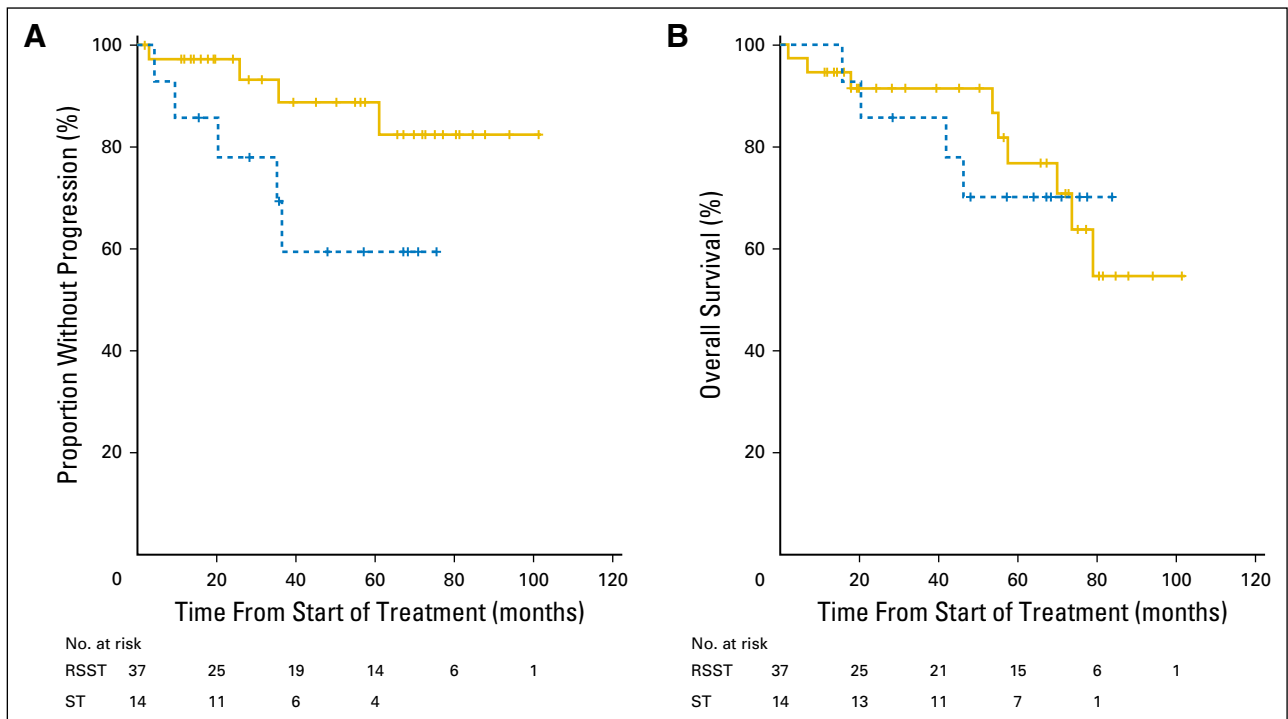


Fig 4. Patients in complete response after rituximab induction (low-risk group). Time to progression and overall survival in the risk-stratified sequential treatment (RSST) cohort (n = 37; solid line) and the sequential treatment (ST) cohort (n = 14; dashed line). (A) Time to progression. The 3-year Kaplan-Meier estimate was 89% (95% CI, 76% to 100%) compared with 69% (95% CI, 44% to 95%) in the 14 patients in the PTLD-1 (Sequential Treatment of CD20-Positive Post-Transplant Lymphoproliferative Disorder trial) ST cohort. (B) Overall survival. The 3-year Kaplan-Meier estimate was 70% (95% CI, 62% to 77%) compared with 61% (95% CI, 49% to 72%) in PTLD-1 ST cohort. Of the six late deaths that occurred in the RSST low-risk cohort, two were attributable to progressive PTLD (after first and second relapse) whereas four were not (one death as a result of unknown causes and three as a result of infections).

immunocompetent patients with CD20⁺ DLBCL^{16,17,22} can be safely used in PTLD as part of ST. However, the spectrum of infections observed included entities typically associated with longstanding immunosuppression (PcP, aspergillosis, progressive multifocal leukoencephalopathy).²³⁻²⁵

The optimal treatment of PTLD has long been a source of controversy.²⁶ This study lend further support to the argument that B-cell PTLD should not be treated with upfront R-CHOP immunochemotherapy in analogy with immunocompetent patients with DLBCL. Upfront chemotherapy in PTLD, to our knowledge, has never been tested in a prospective setting. In retrospective case series of CHOP or CHOP-like protocols, TRM has been reported to be as high as 26% and 31%.^{12,15} We observed a more acceptable rate of TRM (13%) in our previous prospective trial of ST, where CHOP was administered after rituximab induction, possibly as a result of reduced tumor burden and a delay of 50 days between reduction of immunosuppression and start of chemotherapy.⁵ The current trial demonstrates that approximately 25% of patients with PTLD do not need chemotherapy.

Furthermore, the results with continued rituximab strongly suggest that rituximab consolidation is superior to no consolidation (ie, that eight, not four, courses of rituximab are the best available therapy for patients in CR after rituximab induction). Although we have not formally tested this hypothesis, the TTP with RSST in the low-risk group (97% at 24 months; Fig 4A) compares favorably with previous trials where only four cycles of rituximab were administered. In the German and French rituximab monotherapy trials, four of 25 patients in CR experienced a relapse within 12 months, and Blaes et al reported a median duration of CR of 8 months.^{27,28}

In summary, this study establishes the feasibility, efficacy, and safety of RSST in CD20⁺ PTLD. In the absence of any randomized trial data, the results define RSST as a new therapeutic standard in adult CD20⁺ PTLD after SOT and demonstrate that PTLD is a successfully treatable lymphoma.

AUTHORS' DISCLOSURES OF POTENTIAL CONFLICTS OF INTEREST

Disclosures provided by the authors are available with this article at ascopubs.org/journal/jco.

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AUTHORS' DISCLOSURES OF POTENTIAL CONFLICTS OF INTEREST

Response to Rituximab Induction Is a Predictive Marker in B-Cell Post-Transplant Lymphoproliferative Disorder and Allows Successful Stratification Into Rituximab or R-CHOP Consolidation in an International, Prospective, Multicenter Phase II Trial

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