

# First-Line Therapy of Primary Central Nervous System Lymphoma: A Russian Multicenter Study

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## Introduction

Primary central nervous system lymphoma (PCNSL) is a rare subtype of extranodal lymphomas with an aggressive clinical course. Currently, there are no systematic data available in the Russian Federation regarding treatment outcomes and prognosis in patients with PCNSL.

The aim of this study was to evaluate the clinical and epidemiological characteristics of patients with PCNSL, the current first-line therapeutic landscape, and treatment outcomes in real-world clinical practice.

## Methods

From 2010 to 2025, a total of 205 adult patients with histologically confirmed PCNSL from 23 centers across Russia were included in the study.

Key patient and disease characteristics are presented in Table 1.

## Results

As first-line therapy, 96% of patients (n=196) received immuno- and/or chemotherapy (ICT) regimens, 3% (n=6) underwent radiotherapy, and 1% (n=3) received glucocorticoids alone.

Within the ICT group, rituximab was administered to 83% of patients (n=162) with CD20-positive PCNSL. High-dose methotrexate (HD-MTX)-based regimens accounted for 71% of ICT cases (n=139). The most frequently used HD-MTX-based regimens were R-HDMTX-AraC (35%, n=49), R-MP(D)V ± lenalidomide (17%, n=23), R-HDMTX (14%, n=19), and R-HDMTX-temozolomide (10%, n=14). The MATRix protocol was administered in 5% of cases (n=7). The remaining patients (29%, n=57) received other intensive regimens (R-DeVIC – 5%, n=3) or non-intensive regimens (68%, n=39), most commonly temozolomide and/or MTX combinations ± other agents (49%, n=28). In 26% (n=15) of these patients, intra-arterial chemotherapy with MTX was performed using temporary opening of the blood–brain barrier.

In the ICT group, treatment response was assessed in 166 patients (85%). An objective response (OR) was achieved in 71% (n=118), including complete response (CR) in 42% (n=69) and partial response (PR) in 29% (n=49). Stable disease (SD) was observed in 11% (n=19), while 17% (n=29) experienced progressive disease (PD). Treatment-related mortality prior to restaging occurred in 8% (n=15). In the radiotherapy group, 83% (n=5) achieved CR, while 1 patient had PD.

At a median follow-up of 12 months (range 0.5–151), 2-year overall survival (OS) was 49%, and 2-year progression-free survival (PFS) was 33% (Figure 1).

In the ICT group, 35% of patients with OR (n=41) received consolidation therapy: 26 with CR and 15 with PR. High-dose chemotherapy followed by autologous hematopoietic stem cell transplantation (auto-HSCT) was performed in 23 patients, while 17 underwent whole-brain radiotherapy (WBRT), and 1 received a combination of both. Among patients who underwent consolidation, 2-year OS was 84.7% and 2-year PFS was 68.4%. Outcomes did not differ between patients in CR and PR at the time of consolidation: OS 88.6% vs. 80% (p=0.73), PFS 72.2% vs. 63.4% (p=0.86). Consolidation method did not impact outcomes: OS after auto-HSCT vs. WBRT was 79.4% vs. 90% (p=0.39), and PFS 72.9% vs. 66.2% (p=0.86), respectively.

## Conclusions

The clinical profile of patients was generally consistent with previously published data, although some selection factors related to sex, age, and HIV status were noted. Most patients were treated with regimens aligned with international guidelines, though a significant proportion received suboptimal approaches. The best survival outcomes were observed in patients who achieved an objective response and subsequently underwent consolidation therapy. The choice of consolidation modality did not affect prognosis.

**Table 1. Key patient and disease characteristics**

Characteristics	n=205 (100%)
<b>Sex</b>	
Male	83 (40.5%)
Female	122 (59.5%)
<b>Age at diagnosis, median (range)</b>	58 (27–82)
<b>ECOG performance status</b>	
ECOG 1	53 (26%)
ECOG ≥2	152 (74%)
<b>Immunocompromised status at diagnosis</b>	
HIV-positive	10 (5%)
Kidney transplant	1 (0.5%)
History of malignancy	4 (2%)

<b>MSKCC risk score</b>	
Low risk	27 (13%)
Intermediate risk	77 (38%)
High risk	101 (49%)
<b>Diagnostic procedure</b>	
Surgical resection	113 (55%)
Stereotactic biopsy	92 (45%)
<b>Histologic subtype</b>	
Diffuse large B-cell lymphoma	197 (96%)
High-grade B-cell lymphoma	4 (2%)
Marginal zone lymphoma	2 (1%)
Burkitt lymphoma	1 (0.5%)
Peripheral T-cell lymphoma	1 (0.5%)

Figure 1. Overall survival and progression-free survival of patients with PCNSL

