

Introduction & Aim

Chimeric antigen receptor T-cell (CAR-T) therapy has improved outcomes in relapsed or refractory non-Hodgkin lymphoma, but treatment failure, prolonged cytopenias, infection, and mortality remain difficult to predict. Preinfusion inflammatory and hematologic biomarkers may capture host immune state, marrow reserve, and end-organ vulnerability before cellular therapy. This synthesis evaluates translational evidence linking preinfusion inflammatory and hematologic biomarker signatures with CAR-T efficacy, hematologic toxicity, infection risk, and survival in non-Hodgkin lymphoma.

Design & Methods

A focused translational evidence synthesis included three primary clinical biomarker studies evaluating pre-CAR-T inflammatory or hematologic risk models in non-Hodgkin lymphoma. Studies were identified through a literature search for preinfusion biomarker models in CAR-T recipients, with inclusion restricted to models reporting both development and external validation cohorts. Extracted data included biomarker panels, model performance, and clinical outcomes, including treatment failure, survival, and hematologic toxicity.

3 translational studies | 2 validated biomarker models



InflaMix: inflammatory and end-organ vulnerability



CAR-HEMATOTOX: hematologic and inflammatory vulnerability

SHARED: Hemoglobin + CRP

INFLAMIX



Albumin • AST • ALP • LDH



Inflammatory +
end-organ vulnerability



Death or relapse



HR 2.98



Development: n=149 NHL CAR-T recipients
External validation: 3 cohorts; n=688

CAR-HEMATOTOX



Platelets • ANC • Ferritin



Marrow +
inflammatory vulnerability



Severe neutropenia



AUC 0.89



Development: n=258 intent-to-treat LBCL patients
External validation: cohorts n=91 and n=109

Results

- InflaMix identifies inflammatory and end-organ vulnerability associated with death or relapse (HR 2.98; 95% CI 1.60–4.91; p<0.001).
- CAR-HEMATOTOX identifies hematologic and inflammatory vulnerability associated with prolonged severe neutropenia (AUC 0.89; sensitivity 89%; specificity 68%).

Limitations

- Evidence was limited to three studies with heterogeneous patient populations and clinical endpoints.
- Differences in biomarker definitions and model application limited direct comparison.
- Further prospective, multicenter validation is needed before routine clinical use.

Conclusion

Preinfusion biomarker models may support individualized risk assessment for treatment failure and hematologic toxicity before CAR-T therapy.

References

1. Raj SS, Fei T, Fried S, et al. Nat Med. 2025;31:1183–1194.
2. Rejeski K, Perez A, Sesques P, et al. Blood. 2021;138:2499–2513.
3. Rejeski K, Perez A, Iacoboni G, et al. J Immunother Cancer. 2022;10:e004475.

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