

Medical Science

To Cite:

Józefiak A, Szczepanik M, Bochyński C, Hałasiński P, Kropidłowska D, Horbaczewski M, Pielą K, Mazurek J, Mazurek G, Myślička M, Góralski P, Włodarczyk K. Diffuse large B-cell lymphoma (DLBCL) accompanied by a pathologically enlarged pack of lymph nodes: A review. *Medical Science* 2024; 28: e90ms3404
doi: <https://doi.org/10.54905/disssi.v28i150.e90ms3404>

Authors' Affiliation:

- ¹Department of Internal Diseases and Endocrinology, Gabriel Narutowicz Municipal Specialist Hospital in Cracow, Prądnicka 35-37, 31-202 Cracow, Poland
²Department of Pediatrics, District Hospital in Chrzanów, Topolowa 16, 32-500 Chrzanów, Poland
³Department of Internal Medicine, Railway Hospital. Włodzimierz Roefler, MD, PhD in Pruszków, Warszawa 1, 05-800 Pruszków, Poland
⁴Clinical Department of Gastroenterology, Metabolic, Internal Diseases and Dietetics, University Clinical Hospital in Poznań, 49 Przybyszewskiego, 60-355 Poznań, Poland
⁵Polish Red Cross Maritime Hospital in Gdynia, Powstania Styczniowego 1, 81-519 Gdynia, Poland
⁶Hospital Emergency Department, District Health Center Kartuszy, Floriana Ceynowy 7, 83-300 Kartuszy, Poland
⁷Hospital Emergency Department, University Clinical Hospital. Fryderyk Chopin in Rzeszów, Fryderyka Szopena 2, 35-055 Rzeszów, Poland
⁸Primary Health Care, Private Health Care Facility "ALMUS" Jacek Pieniązek, Białoboki 137, 37-207 Gać, Poland
⁹Department of Cardiology and Internal Diseases, Gabriel Narutowicz Municipal Specialist Hospital in Cracow, Prądnicka 35-37, 31-202 Cracow, Poland
¹⁰Wrocław Medical University, wyb. Ludwika Pasteura 1, 50-367 Wrocław, Poland
¹¹University Clinical Center in Gdańsk, Dębinki 7, 80-952 Gdańsk, Poland

*Corresponding Author

Department of Internal Medicine, Railway Hospital. Włodzimierz Roefler, MD, PhD in Pruszków, Warszawa 1, 05-800 Pruszków, Poland
Email: cezary1029@gmail.com

Contact List
 Anna Józefiak jozefiakanna@onet.pl
 Magdalena Szczepanik magdalenaszczepanik97@gmail.com
 Cezary Bochyński cezary1029@gmail.com
 Przemysław Hałasiński przemyslaw.halasiński@tlen.pl
 Dominika Kropidłowska dominika.kropidlowska@tlen.pl
 Maciej Horbaczewski dawidm1990@gmail.com
 Kinga Pielą kinga.pielą@gmail.com
 Jolanta Mazurek mi.jasmin@poczta.onet.pl
 Gabriela Mazurek gabriela.mazurek@gmail.com
 Maria Myślička mariamyslicka3@gmail.com
 Patryk Góralski patryk32150@gmail.com
 Klaudia Włodarczyk włodarczyk.klaudia1@gmail.com

ORCID List
 Anna Józefiak 0000-0003-3968-472X
 Magdalena Szczepanik 0000-0001-5550-0860
 Cezary Bochyński 0009-0001-2672-2445
 Przemysław Hałasiński 0000-0002-7223-3394
 Dominika Kropidłowska 0009-0004-3508-2621
 Maciej Horbaczewski 0009-0002-1842-0991
 Kinga Pielą 0009-0002-1507-8065
 Jolanta Mazurek 0009-0001-4884-4287
 Gabriela Mazurek 0000-0001-7338-9828
 Maria Myślička 0009-0000-6190-9128
 Patryk Góralski 0009-0008-1831-0775
 Klaudia Włodarczyk 0009-0002-0779-8564

Peer-Review History

Received: 12 May 2024
 Reviewed & Revised: 16/May/2024 to 27/July/2024
 Accepted: 31 July 2024
 Published: 07 August 2024

Peer-review Method

External peer-review was done through double-blind method.

Medical Science
 pISSN 2321-7359; eISSN 2321-7367



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Diffuse large B-cell lymphoma (DLBCL) accompanied by a pathologically enlarged pack of lymph nodes: A review

Anna Józefiak¹, Magdalena Szczepanik², Cezary Bochyński^{3*}, Przemysław Hałasiński⁴, Dominika Kropidłowska⁵, Maciej Horbaczewski⁶, Kinga Pielą⁷, Jolanta Mazurek⁸, Gabriela Mazurek⁹, Maria Myślička¹⁰, Patryk Góralski¹¹, Klaudia Włodarczyk¹⁰

ABSTRACT

Introduction: Diffuse large B-cell lymphoma (DLBCL) is a relatively common lymphoma with an aggressive course. A complete cure is possible for this type of lymphoma. The etiology of this cancer is unknown, but a number of factors, including environmental factors, contribute to the onset of this disease. **The aim:** The purpose of this article is to present diffuse large B-cell lymphoma as a curable disease if detected and treated early enough. **Case report:** The described case is a 70-year-old patient who came to the doctor with a neck tumor. After taking an interview, taking a history and physical examination, and making a diagnosis, the patient was diagnosed with Diffuse large B-cell lymphoma. **Results:** DLBCL is the most common lymphoma type in adults, characterized by highly aggressive and dynamic development. Diffuse large B-cell lymphoma most often occurs in older people, around 70 years old. The diagnosis of DLBCL is based on histopathological examination of a sample taken during surgery, i.e., the cancerous lymph node. In this type of lymphoma, complete recovery is possible with early diagnosis and treatment. **Conclusions:** Due to the aggressive development of Diffuse large B-cell lymphoma, only rapid diagnosis and treatment can ensure tumor regression. Diagnosis of cancer is based on histopathological examination, while treatment is based on surgery and subsequent postoperative chemotherapy.

Keywords: Neck tumor, lymphoma, DLBCL, diffuse large B-cell lymphoma

1. INTRODUCTION

Diffuse large B-cell lymphoma (DLBCL) is among the most common groups of lymphomas among lymphoid malignancies. The incidence of DLBCL in Europe is about a dozen cases per 100,000 people in the general population per year. The cancer is more likely to occur in what is considered old age (Zelenetz et al., 2010). The etiology of DLBCL formation is unknown. Scientists believe that a genetic predisposition may be necessary. Immunologic, environmental, infectious, or iatrogenic factors, among others, have a proven association with the occurrence of DLBCL. An increased risk of the disease is found in people with primary and secondary immune disorders, AIDS patients are among a particularly vulnerable group of patients.

Viral infections, e.g., Epstein-Barr virus (EBV), human herpes virus type 8 (HHV-8), human immunodeficiency virus (HIV), and hepatitis C virus (HCV) carry an increased risk of lymphoma. Transplant recipients who are under immunosuppression are at high risk of proliferation of Epstein-Barr virus-infected polyclonal B lymphocytes and subsequent transformation into DLBCL (Blombery et al., 2015). Mechanisms leading to neoplastic transformation of normal lymphocytes in DLBCL are due to the occurrence of genetic instability with subsequent dysregulation of oncogene expression levels or loss of function of tumor suppressor genes (Lenz and Staudt, 2010). In our work, we will present that timely diagnosis and proper treatment of DLBCL lead to full patient cure (Kawano et al., 2021; Eyre et al., 2022).

2. METHODOLOGY

This case study was conducted by searching for current papers on PubMed and Google Scholar using the search phrases (DLBCL) OR (diffuse large B-cell lymphoma) AND (lymphoma). After eliminating duplicates, we appraised all publications using the titles and abstracts. Following an exact revision of complete manuscripts, 20 articles met the inclusion criteria. The research took place in March 2024.

3. CASE STUDY

A seventy-year-old patient presented to the Department of Otolaryngology at the Frederic Chopin Clinical Regional Hospital Number 1 in Rzeszow on the basis of a referral from a family medicine physician. The initial diagnosis, as determined by the admitting physician, was a neck tumor on the left side and status post-treatment for lymphoma. On otolaryngological examination, the patient's general condition was assessed as good: Normal nasal mucosa and nasal patency on the right, dryness after lymphoma treatment, external auditory canals and eardrums of the right and left ears unchanged, oral cavity unchanged, throat unchanged, larynx on indirect laryngoscopy unchanged, on the neck on the left side in the submandibular region palpation of a soft tumor of about 4 cm in diameter (Figure 1).

A blood count results were E $5.06 \times 10^6/\mu\text{L}$, Hb 15.1g/dL, Ht 45.8%, MCV 90.5 fL, Pk $231 \times 10^3/\mu\text{L}$, L $5.67 \times 10^3/\mu\text{L}$, NEUT 69.0%, LYMPH 21.3%, MONO 6.9%, EOS 1.9%, BASO 0.9%, IG% $0.02 \times 10^3/\mu\text{L}$, NRBC% 0.0%, NRBC# $0.0 \times 10^3/\mu\text{L}$, Micro R 1.7%, Macro R 3.9%. On the same day, the patient was qualified for surgery and underwent Exstirpatio Lymphonodulorum colli sin under general anesthesia. During the procedure, a neck skin incision was made on the left side over a packet of group III lymph nodes with a palpable size of 5x3 cm. The surgeons decided to remove the enlarged pathological lymph node, control the bleeding, and apply a seton in the postoperative cavity, sutures, and a dressing. The material was submitted for histopathological verification. The day after the procedure, the patient felt well.

Medical personnel changed her dressing and removed the seton. The wound was healing properly. The patient was discharged home in good condition with recommendations to remove the sutures in 7 days and to use Augmentin 2x1.0. The final clinical diagnosis was locally enlarged lymph nodes. Histopathological examination diagnosed a tumor formed of large atypical lymphocytes, with a diffuse growth type involving adjacent fatty tissue and nerve trunks. IHC profile of tumor cells: CD20+, bcl2+, bcl6+, CD10-, MUM1+, c-myc-(+ <30%), CD5-, CyclinD1-, CD23-. The proliferative activity of Ki-67 is almost 100%. Histopathological picture and IHC profile may correspond to diffuse large B-cell lymphoma: DLBCL NOS, ABC-type. Indication for determination of BCL2, BCL6, and MYC gene rearrangements to exclude double/triple hit lymphoma.



Figure 1 A soft tumor of about 4cm in diameter on the left side of the neck.

4. DISCUSSION

DLBCL is a common type of lymphoma with an aggressive course but with the possibility of a complete cure (Nogai et al., 2011; Yang et al., 2021). It occurs in people of all ages, more commonly in the elderly (Wang, 2023; Kos et al., 2021). Diffuse large B-cell lymphomas (DLBCL, diffuse large B-cell lymphoma) are the most common type of non-Hodgkin's lymphoma in adults and a highly heterogeneous group of diseases in terms of both morphological, biological, and clinical features (Takahara et al., 2023). The basis for diagnosing DLBCL is histopathological examination. The entire lymph node or possibly a fragment of the involved organ should be taken. Histopathologic evaluation needs to be augmented with immunophenotypic studies using monoclonal antibodies (Silkenstedt et al., 2024).

DLBCL lymphoma cells express pan-B antigens (CD19, CD20, CD22, CD79a), with varying percentages of BCL6, BCL2, and CD10, and exceptionally express the CD5 antigen (Chow, 2020; Stachura et al., 2006). In 30-40% of DLBCL cases, there are inappropriate alterations in the BCL6 gene (3q27), which rearrange around gene loci for immunoglobulins in the 14q32, 2p12, or 22q11 region. In 15-30% of patients, t (14; 18) is found, which leads to overexpression of BCL2 (Dargent et al., 2022). The third most common (5-10%) chromosomal aberration is t (8; 14), which runs with increased MYC expression and correlates with extra-nodal localization of DLBCL. In several percent of DLBCL cases, the above abnormalities co-occur. Such lymphomas are characterized by a particularly aggressive

clinical course and usually meet the morphologic criteria of unclassifiable B-cell lymphoma, with traits intermediate between DLBCL and Burkitt's lymphoma (Abrey et al., 2005).

Uncommonly, the diagnosis is supplemented with cytogenetic and molecular studies, which may allow assessment of lymphoid cell clonality or are helpful in identifying characteristic genetic abnormalities for a specific lymphoma subtype (Harkins et al., 2019). Recently, gene expression profiling has become increasingly important to define new molecular subtypes. It is also helpful for confirming the diagnosis of DLBCL that does not meet classic diagnostic criteria (Pasqualucci and Dalla-Favera, 2018). Treatment of a neck tumor depends on the diagnosis of its nature. Histopathological confirmation of either a slice or the entire tumor is necessary. The fate of the patient and the choice of a suitable treatment method depends on the structure and clinical nature of the lesion.

All non-radical surgeries can worsen the prognosis - this applies to tumors but also to congenital lesions. For some inflammatory or traumatic lesions, physicians can use conservative treatment. Surgical treatment often has to be combined with subsequent radiation or chemotherapy. According to statistics, patient survival without therapy for diffuse large B-cell lymphoma is several to several months (Sehn and Salles, 2021). Due to its aggressive nature, if left untreated, it spreads very quickly, mainly through the blood and lymphatic pathways, where it pathologically alters more lymphatic vessels (Messina et al., 2015). An essential step in managing this type of lesion is to exclude its malignancy. This enables the deployment of appropriately selected pharmacological and surgical treatments. (Solimando et al., 2020).

5. CONCLUSIONS

In summary, DLBCL is the most common type of lymphoma in adults characterized by a highly aggressive and dynamic development. The etiology of DLBCL is still not clear. There are quite a few environmental factors that may increase the chance of developing this lymphoma. The diagnosis of DLBCL on histopathological examination of the lymph node collected during surgery. The treatment of choice for this type of tumor is surgery to remove the lesion and subsequent chemotherapy. The patient's health status after treatment depends most on the cancer stage at the time of presentation to the doctor.

Author's Contribution

Anna Józefiak: Conceptualization, methodology, investigation

Magdalena Szczepanik: Conceptualization, methodology, investigation

Cezary Bochyński: Methodology, Review and editing

Przemysław Hałasiński: Conceptualization, writing- rough preparation

Dominika Kropidłowska: Resources, writing- rough preparation

Maciej Horbaczewski: Conceptualization, writing- rough preparation

Kinga Piela: Formal analysis, supervision

Jolanta Mazurek: Review and editing, supervision

Gabriela Mazurek- Visualization, data curation

Maria Myślicka: Formal analysis, supervision

Patryk Góralski: Resources, writing- rough preparation

Klaudia Włodarczyk: Visualization, data curation

Project administration: Cezary Bochyński

Informed consent: Not applicable.

Ethical approval: Not applicable.

Funding

This study has not received any external funding.

Conflict of interest:

The authors declare that there is no conflict of interests.

Data and materials availability

All data sets collected during this study are available upon reasonable request from the corresponding author.

REFERENCES

1. Abrey LE, Batchelor TT, Ferreri AJ, Gospodarowicz M, Pulczynski EJ, Zucca E, Smith JR, Korfel A, Soussain C, DeAngelis LM, Neuwelt EA, O'Neill BP, Thiel E, Shenkier T, Graus F, Van-den-Bent M, Seymour JF, Poortmans P, Armitage JO, Cavalli F; International Primary CNS Lymphoma Collaborative Group. Report of an international workshop to standardize baseline evaluation and response criteria for primary CNS lymphoma. *J Clin Oncol* 2005; 23(22):5034-43. doi: 10.1200/JCO.2005.13.524.
2. Blombery PA, Wall M, Seymour JF. The molecular pathogenesis of B-cell non-Hodgkin lymphoma. *Eur J Haematol* 2015; 95(4):280-93. doi: 10.1111/ejh.12589
3. Chow LQM. Head and Neck Cancer. *N Engl J Med* 2020; 382(1):60-72. doi: 10.1056/NEJMra1715715
4. Dargent JL, Toffoli S, De-Rop C, Hérin M. Fibrin-Associated EBV-Positive Large B-Cell Lymphoma Incidentally Found Within a Multinodular Goiter. *Int J Surg Pathol* 2022; 30(6):658-661. doi: 10.1177/10668969221074604
5. Eyre TA, Savage KJ, Cheah CY, El-Galaly TC, Lewis KL, McKay P, Wilson MR, Evens AM, Bobillo S, Villa D, Maurer MJ, Cwynarski K, Ferreri AJM. CNS prophylaxis for diffuse large B-cell lymphoma. *Lancet Oncol* 2022; 23(9):e416-e426. doi: 10.1016/S1470-2045(22)00371-0
6. Harkins RA, Patel SP, Flowers CR. Cost burden of diffuse large B-cell lymphoma. *Expert Rev Pharmacoecon Outcomes Res* 2019; 19(6):645-661. doi: 10.1080/14737167.2019.1680288
7. Kawano T, Tsuyuki Y, Suzuki Y, Shimada K, Kato S, Takahara T, Mori M, Nakaguro M, Sakakibara A, Nakamura S, Satou A. Clinicopathologic Analysis of Primary Adrenal Diffuse Large B-Cell Lymphoma: A Reappraisal of 23 Japanese Patients Based on EBV Association and PD-L1 Expression in Tumor Cells. *Am J Surg Pathol* 2021; 45(12):1606-1615. doi: 10.1097/PAS.0000000000001809
8. Kos IA, Thurner L, Bittenbring JT, Christofyllakis K, Kaddu-Mulindwa D. Advances in Lymphoma Molecular Diagnostics. *Diagnostics (Basel)* 2021; 11(12):2174. doi: 10.3390/diagnostics11122174
9. Lenz G, Staudt LM. Aggressive lymphomas. *N Engl J Med* 2010; 362(15):1417-29. doi: 10.1056/NEJMra0807082
10. Messina C, Christie D, Zucca E, Gospodarowicz M, Ferreri AJ. Primary and secondary bone lymphomas. *Cancer Treat Rev* 2015; 41(3):235-46. doi: 10.1016/j.ctrv.2015.02.001
11. Nogai H, Dörken B, Lenz G. Pathogenesis of non-Hodgkin's lymphoma. *J Clin Oncol* 2011; 29(14):1803-11. doi: 10.1200/JCO.2010.33.3252
12. Pasqualucci L, Dalla-Favera R. Genetics of diffuse large B-cell lymphoma. *Blood* 2018; 131(21):2307-2319. doi: 10.1182/blood-2017-11-764332
13. Sehn LH, Salles G. Diffuse Large B-Cell Lymphoma. *N Engl J Med* 2021; 384(9):842-858. doi: 10.1056/NEJMra2027612
14. Silkenstedt E, Salles G, Campo E, Dreyling M. B-cell non-Hodgkin lymphomas. *Lancet* 2024; 403(10438):1791-1807. doi: 10.1016/S0140-6736(23)02705-8
15. Solimando AG, Annese T, Tamma R, Ingravallo G, Maiorano E, Vacca A, Specchia G, Ribatti D. New Insights into Diffuse Large B-Cell Lymphoma Pathobiology. *Cancers (Basel)* 2020; 12(7):1869. doi: 10.3390/cancers12071869
16. Stachura J, Rudzki Z., Gałazka K, Jurczak W. Diagnostyka chłoniaków. Zarys standardów diagnostycznych. *Pol J Pathol* 2006; 57:1-5.
17. Takahara T, Nakamura S, Tsuzuki T, Satou A. The Immunology of DLBCL. *Cancers (Basel)* 2023; 15(3):835. doi: 10.3390/cancers15030835
18. Wang SS. Epidemiology and etiology of diffuse large B-cell lymphoma. *Semin Hematol* 2023; 60(5):255-266. doi: 10.1053/j.seminhematol.2023.11.004
19. Yang SH, Park H, Yoo DS, Joo W, Rhoton A. Microsurgical anatomy of the facial nerve. *Clin Anat* 2021; 34(1):90-102. doi: 10.1002/ca.23652
20. Zelenetz AD, Abramson JS, Advani RH, Andreadis CB, Byrd JC, Czuczman MS, Fayad L, Forero A, Glenn MJ, Gockerman JP, Gordon LI, Harris NL, Hoppe RT, Horwitz SM, Kaminski MS, Kim YH, Lacasce AS, Mughal TI, Nademanee A, Porcu P, Press O, Prosnitz L, Reddy N, Smith MR, Sokol L, Swinnen L, Vose JM, Wierda WG, Yahalom J, Yunus F. NCCN Clinical Practice Guidelines in Oncology: non-Hodgkin's lymphomas. *J Natl Compr Canc Netw* 2010; 8(3):288-334. doi: 10.6004/jnccn.2010.0021