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Asian Journal of Oncology



## **Original** Article

# A bibliometric analysis of CiteSpace software-based immunotherapy for the treatment of diffuse large B-cell lymphoma

## Weichen Si, BACHELOR<sup>1</sup>

<sup>1</sup>Department of Acupuncture and Moxibustion, Beijing Direction Community Health Service Station, Beijing, China

# ABSTRACT

**Objectives:** To analyze the literature data and research status of immunotherapy for the treatment of diffuse large B-cell lymphoma since the establishment of the Web of Science (WOS) core database.

Material and Methods: The WOS core database was searched for literature related to immunotherapy for diffuse large B-cell lymphoma, and the included literature was formatted, cleaned, node merged, and analyzed using CiteSpace software. Based on the parameters set, the included literature was analyzed for trends in publications, author publications and inter-author collaborations, national publications, global institutional publications and inter-institutional collaborations, citations, keyword co-occurrence, keyword emergence, and keyword clustering. The final visual knowledge map was created.

**Results:** A total of 370 articles were selected for inclusion. The highest number of annual publications was in 2021. Four individuals, Marconato Laura; Ansell, Stephen M; Xiao Min; and Aresu Luca, were the most published scholars. The United States with 152 publications was the country with the highest number of publications. Benjamin J is the most cited scholar in this field. The top three most cited keywords were "expression," "diffuse large b-cell lymphoma," and "rituximab." "Bone marrow transplantation" was the first and longest-running keyword. "Cancer immunotherapy," "resistance," and "cytokine release syndrome" are still hot topics. The keyword clusters "pd-l1," "antibody-based," "immunotherapy," and "cd19" were the main clusters studied.

**Conclusion:** After visualization and analysis, the recent research and hot trends in the field of immunotherapy for diffuse large B-cell lymphoma were reviewed using knowledge mapping and further presented in a visualized form, providing a reference for further development of related research in the future.

Keywords: Immunotherapy, Diffuse large B-cell lymphoma, CiteSpace, Metrology

# INTRODUCTION

The application of immunotherapy in the treatment of diffuse large B-cell lymphoma is one of the hot topics in the field of tumor therapy, and it is of great significance to the bibliometrics research in this field. Specifically, this study can help readers understand the research progress and trend of immunotherapy in the treatment of diffuse large B-cell lymphoma, and provide scientific basis and reference for clinical practice. Analyzing and evaluating the quality and quantity of existing research helps us understand the overall situation of research in this field, helps to find the defects and deficiencies in research, and provides the direction of improvement and perfection for future research. It will provide important reference information for policy makers, managers, researchers, and investors in relevant research fields to help

them formulate more scientific and rational development strategies and plans and promote further development and research in this field. To provide the academia and the public with the latest information on research progress in this field, the public's attention and awareness of this field should be improved, and the achievements and technologies in this field should be popularized.

# MATERIAL AND METHODS

## Etiology of diffuse large B-cell lymphoma

Diffuse large B-cell lymphoma (DLBCL) is a type of non-Hodgkin's lymphoma (NHL) that originates from B cells in the lymphatic system.<sup>[1]</sup> The causes of DLBCL are not fully understood, but several factors have been identified that

Corresponding author: Dr. Weichen Si, Department of Acupuncture and Moxibustion, Beijing Direction Community Health Service Station, Beijing, China. weichensil4@outlook.com

Received: 24 April 2023 Accepted: 15 May 2023 Published: 21 June 2023 DOI 10.25259/ASJO\_1\_2023

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may increase the risk of developing this condition: Genetic mutations: Certain genetic mutations have been associated with an increased risk of developing DLBCL, including mutations in genes that regulate cell growth and division.<sup>[2]</sup> Immune system dysfunction: DLBCL may occur when the immune system is weakened, such as in people with HIV/ AIDS or those who have undergone an organ transplant and are taking immunosuppressive drugs.<sup>[3]</sup> Viral infections: Some viral infections, such as Epstein-Barr virus (EBV) and human herpesvirus 8 (HHV-8), have been linked to an increased risk of developing DLBCL. Environmental factors: Exposure to certain environmental toxins, such as pesticides, may increase the risk of developing DLBCL.<sup>[4]</sup> Age: DLBCL is more common in older adults, with the average age at diagnosis being around 65 years. It is important to note that while these factors may increase the risk of developing DLBCL, not everyone who is exposed to these risk factors will develop the disease.<sup>[5]</sup> In addition, many people with DLBCL have no identifiable risk factors, and the exact cause of their lymphoma is unknown.

## Pathogenesis of diffuse large B-cell lymphoma

DLBCL involves the transformation of B cells in the lymphatic system into malignant cells that grow and divide uncontrollably.<sup>[6]</sup> This can result in the formation of tumors and the spread of cancer to other parts of the body.<sup>[6]</sup> DLBCL arises from a particular type of B cell known as the germinal center B cell, which normally plays a role in the immune response to infection. However, in DLBCL, these cells acquire mutations that disrupt their normal function, causing them to proliferate abnormally.<sup>[7]</sup> There are several genetic alterations that have been associated with DLBCL pathogenesis.<sup>[5]</sup> The most common genetic alteration is a translocation between the BCL2 and IGH genes, which results in overexpression of the BCL2 protein<sup>2</sup>. This protein helps to prevent apoptosis (programmed cell death) and can promote the survival of cancer cells.<sup>[8]</sup> Other genetic alterations that can contribute to DLBCL pathogenesis include mutations in genes involved in B cell signaling pathways, as well as alterations in genes that regulate cell growth and division.<sup>[4]</sup> In addition to genetic alterations, DLBCL pathogenesis can also be influenced by epigenetic changes, such as alterations in DNA methylation or histone modifications, which can affect gene expression<sup>[1]</sup>. The precise sequence of events that lead to DLBCL pathogenesis can be complex and may involve interactions between multiple genetic and environmental factors.<sup>[9]</sup> However, a better understanding of the underlying molecular mechanisms involved in DLBCL pathogenesis may lead to the development of more effective treatments for this disease.[2,3]

# The treatment of diffuse large B-cell lymphoma

DLBCL typically depends on several factors, including the patient's age and overall health, the stage and aggressiveness of thelymphoma, and the presence of any genetic abnormalities.<sup>[10]</sup> First-line treatment for DLBCL typically involves combination chemotherapy, which is often administered with the monoclonal antibody rituximab.<sup>[10,11]</sup> The most commonly used chemotherapy regimen for DLBCL is called R-CHOP, which includes rituximab, cyclophosphamide, doxorubicin, vincristine, and prednisone.<sup>[12]</sup> This regimen is typically administered every 21 days for six cycles. For patients who are not candidates for R-CHOP, alternative chemotherapy regimens, such as EPOCH or dose-adjusted R-EPOCH, may be used. In some cases, radiation therapy may be added to the treatment regimen. Second-line treatment options for DLBCL include high-dose chemotherapy followed by autologous stem cell transplantation (ASCT), which involves the use of the patient's own stem cells to replace the damaged bone marrow.<sup>[13]</sup> Other options may include targeted therapies, such as the monoclonal antibody drug brentuximab vedotin or the immunomodulatory drug lenalidomide. Adjuvant therapy, which is given after the initial treatment, may include radiation therapy or maintenance therapy with rituximab.<sup>[14]</sup> These therapies are designed to help prevent the recurrence of the lymphoma. For patients with relapsed or refractory DLBCL, salvage chemotherapy followed by ASCT may be considered.<sup>[15]</sup> Other treatment options for relapsed or refractory DLBCL may include targeted therapies, such as the BTK inhibitor ibrutinib or the PI3K inhibitor idelalisib, as well as immunotherapy, such as CAR T-cell therapy.<sup>[16]</sup>

# Overview of immunotherapy

Immunotherapy is a type of cancer treatment that harnesses the body's own immune system to fight cancer.<sup>[17]</sup> It is designed to stimulate the immune system to recognize and attack cancer cells, thereby slowing or stopping the growth and spread of cancer. One of the major advantages of immunotherapy is its specificity.<sup>[18]</sup> Unlike traditional cancer treatments, such as chemotherapy, which can kill both cancerous and healthy cells, immunotherapy drugs are designed to target cancer cells specifically.<sup>[19]</sup> This means that they can be more effective in treating cancer while minimizing damage to healthy cells. Another advantage of immunotherapy is its potential to provide long-lasting responses.<sup>[20]</sup> In some cases, immunotherapy has been shown to cause complete remission, meaning that the cancer is no longer detectable in the body, and patients can remain cancerfree for years after treatment. Moreover, immunotherapy is a relatively new field, and researchers continue to develop and refine new immunotherapy treatments.<sup>[21]</sup> This means that there is potential for even greater advancements in the future, with the development of more effective and targeted immunotherapies.

## **Research Methodology**

Bibliometric analysis involves the use of quantitative methods to analyze scientific publications in order to evaluate the impact of a particular field or subject area.<sup>[22]</sup> In the case of DLBCL, a type of blood cancer that affects B cells, bibliometric analysis could provide insights into the research trends, collaborations, and impact of scientific publications related to this disease.<sup>[23]</sup> One potential implication of bibliometric analysis of DLBCL is that it could help identify the most influential authors, institutions, and countries in the field. This could inform decisions about where to allocate funding and resources for research, as well as identify potential collaborations and partnerships.<sup>[24]</sup> Another implication is that bibliometric analysis could provide insights into the research trends and gaps in DLBCL.<sup>[25]</sup> For example, by analyzing the keywords used in DLBCL publications, it may be possible to identify emerging areas of research or areas where there is a lack of research. In addition, bibliometric analysis could help evaluate the impact of DLBCL research on clinical practice and patient outcomes.<sup>[22,23]</sup> By analyzing citations of DLBCL publications in clinical guidelines and other publications, it may be possible to assess the extent to which research has influenced clinical decision-making and patient care. Overall, bibliometric analysis of DLBCL could provide valuable insights into the research trends, collaborations, and impact of scientific publications related to this disease, which could inform future research and clinical practice.<sup>[25]</sup>

This research uses Web of Science (WOS) as the data source. Search subject terms: TS = ("Diffuse Large B-cell Lymphoma" OR "ALL") AND TS=("immunotherapy"). The search time was set from library creation to December 2022. The retrieved literature was manually screened to exclude literature with titles, abstracts, content not clearly relevant to DLBCL, conference papers, scientific and technical results, and newspapers. Finally, the relevant documents that met the criteria were exported in plain text format, named "download\_\*\*.txt" and imported into CiteSpace (6.1.R3) for format conversion, data cleaning, node merging, and data analysis. The time slice was 1 year, and the source of the subject terms was selected by default. According to each setting parameter, the included related literature was analyzed for publication trend, author publication and inter-author collaboration, publication volume by country, publication volume by global institutions and inter-institutional collaboration, literature cited, keyword co-occurrence, keyword emergence, keyword clustering, and the network knowledge graph of DLBCL was drawn. The software prompts are combined with manual literature reading and information integration for in-depth analysis of the mapping information.

CiteSpace is a bibliometric analysis software tool that enables the visualization and analysis of citation networks in scientific literature.<sup>[26]</sup> It is widely used in various fields, including science, technology, and medicine, to identify research trends, key contributors, and emerging areas of research. CiteSpace uses a variety of metrics and algorithms to identify important research themes and patterns of collaboration among researchers.<sup>[27]</sup> The software can be used to identify the most frequently cited papers in a particular field, the most influential authors, and the most productive institutions. In addition, it can analyze co-citation networks to identify clusters of related research topics, and it can identify bursty keywords that are becoming increasingly popular in a given field. The research methodology employed by CiteSpace is important because it provides a systematic, objective, and quantitative approach to bibliometric analysis. By analyzing citation networks, CiteSpace can reveal patterns and trends in scientific research that may not be readily apparent through other methods.<sup>[28]</sup> This information can be used to inform research policy, identify promising areas for future research, and evaluate the impact of scientific publications. The use of CiteSpace can also help researchers to identify potential collaborators and to stay up-to-date with the latest research in their field. By analyzing the citation patterns of researchers and institutions, CiteSpace can help to identify important research trends and collaborations that might not be immediately apparent.<sup>[29]</sup> Overall, the research methodology employed by CiteSpace is an important tool for bibliometric analysis and can provide valuable insights into the structure and evolution of scientific research.

# RESULTS

# Results of the trend analysis of publications

A total of 370 articles related to immunotherapy for DLBCL were obtained from the WOS database of 2022. The trend and distribution of the literature on DLBCL obtained from the CiteSpace analysis is shown in Figure 1, where the first appearance of DLBCL in the last 20 years was in 2003. The years 2003–2010 were the initial stage of DLBCL research, with less than 10 articles published per year and few relevant research results; 2011–2018 saw a slight increase in research literature (11–32 articles) and a significant upward trend; 2019–2022: After a small decrease, the annual number of articles peaks at 62 in 2021 and then decreases slightly in 2022. The total number of papers on immunotherapy for DLBCL from the web of science core build to 2022 is 370. New high-level research on immunotherapy for DLBCL is

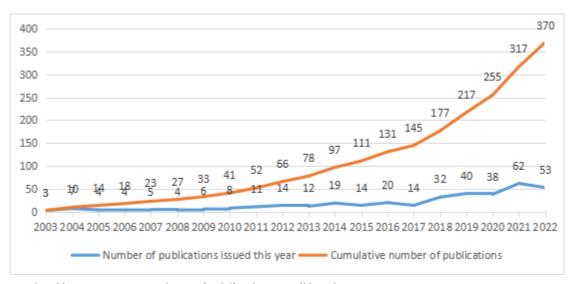


Figure 1: Research publications on immunotherapy for diffuse large B-cell lymphoma.

emerging, and there is still much room for future research development.

# Results of the analysis of authorship and collaboration between authors

A total of 496 authors were included in the resulting map, with the highest number of publications being six. The four authors with the highest number of publications were Marconato Laura; Ansell, Stephen M; Xiao Min; and Aresu Luca. The number of articles published by Stefano was five. The number of articles published by Dybkaer Karen, Tzankov Alexandar, Xu-monette, and Zijun Y was four. The number of authors with three articles is 22, the number of authors with two articles is 56, and the number of authors with one article is 405. Marconato Laura; Ansell, Stephen M; Xiao Min; and Aresu Luca are the most influential authors in the field of immunotherapy for DLBCL. Each node in Figure 2 represents an author. The larger the node, the greater the number of publications, and the thicker the line, the closer



**Figure 2:** Co-presentation of core authors in the research literature on immunotherapy for diffuse large B-cell lymphoma.

Author	Year	Number of publications
Marconato Laura	2014	6
Ansell Stephen M	2009	6
Xiao Min	2019	6
Aresu Luca	2014	6
Young Ken H	2018	5
Comazzi Stefano	2014	5
Dybkaer Karen	2017	4
Tzankov Alexandar	2019	4
Xu-monette Zijun Y	2018	4
Li Yong	2019	3

the connection between the authors. The 10 scholars with the largest number of publications are shown in Table 1.

#### Analysis of the volume of articles published by country

As shown in Table 2 and Figure 3, there is a wide disparity in the number of articles published in different countries, with the United States ranking first with the highest number of articles at 152, creating a fault line between the second place. The second highest number of articles was published by PEOPLES R CHINA with 71 articles, FRANCE with 40 articles, and GERMANY with 37 articles. In summary, there is a wide disparity in research on diffuse large B-cell lymphoma across countries, with the United States having the most research, more than twice as many as the second most published country, suggesting that the United States is significantly ahead of the world in research in this area. In developing countries other than China, there is less research on immunotherapy for DLBCL and further improvement is needed. Si: Bibliometric analysis of immunotherapy for diffuse large B-cell lymphoma

Table 2: Top 10 countries in terms of number of articles published.								
CountryYearNumber of articlesCountryYearNumber of a								
USA	2003	152	England	2004	26			
Peoples R China	2011	71	Spain	2004	22			
France	2006	40	Switzerland	2012	21			
Germany	2004	37	Canada	2003	19			
Italy	2006	35	Netherlands	2003	15			

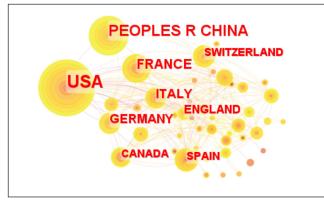


Figure 3: Volume of relevant literature by country.

# Results of the analysis of the number of articles published by institutions and inter-institutional collaboration

A total of 328 institutions were included in the 370 included papers, and the top 10 institutions in terms of number of publications are shown in Table 3. A visual analysis of the collaboration network was conducted [Figure 4]. A total of 51 nodes and 276 links were obtained from the analysis,

**Table 3:** Top 10 research institutions in terms of number of articlespublished.

Institution name	Year	Centrality	Number of articles
UTMD Anderson Cancer Center	2007	0.06	24
Harvard University	2011	0.08	20
Institut National de la Sante	2012	0.07	18
et de la Recherche Medicale (Inserm)			
UDICE-French Research	2012	0	14
Universities			
Memorial Sloan Kettering	2010	0.05	14
Cancer Center Assistance Publique Hopitaux	2012	0.04	13
Paris (APHP)	••••		
Fred Hutchinson Cancer Center	2004	0.07	11
Mayo Clinic	2005	0.05	11
Dana-Farber Cancer Institute	2013	0.03	11
Cornell University	2010	0.01	10

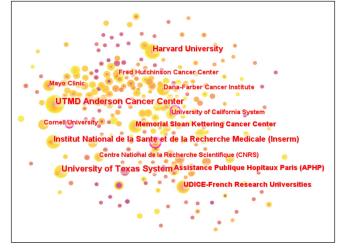


Figure 4: Display of collaborative network of relevant literature research institutions.

indicating that there are a large number of institutions studying immunotherapy for DLBCL and that the institutions are closely linked to each other. For DLBCL research, interinstitutional coordination and collaboration leads to more impactful research results. UTMD Anderson Cancer Center with 24 articles has published the most number of articles. Harvard University has published 20 articles, and Institut National de la Sante et de la Recherche Medicale (Inserm) has published 18 articles. Nine institutions have published more than 10 articles.

## Analysis of citations to the literature

As shown in Table 4 and Figure 5, Benjamin J. Chen; Bjoern Chapuy; Jing Ouyang, *et al*, published in *CLIN CANCER RES/CLINICAL CANCER RESEARCH*, PD-L1 Expression Is Characteristic of a Subset of PD-L1 Expression Is Characteristic of a Subset of Aggressive B-cell Lymphomas and Virus-Associated Malignancies is one of the most influential studies in the field of immunotherapy for DLBCL. There are two papers with more than 10 citations in the field of immunotherapy for DLBCL. Thirteen papers were cited more than five times. Seven papers were cited five times and ten papers were cited four times. Fourteen papers were cited three times. See Table 3 for information on authors,

Table 4: Information or	Table 4: Information on authors, journals, and literature with more than six citations.								
Highly cited scholar	Name of journal	DOI number	Number of citations	Centrality	Year				
Chen BJ <sup>[30]</sup>	CLIN CANCER RES	10.1158/1078-0432.CCR-13-0855	12	0.26	2013				
Ansell SM <sup>[31]</sup>	NEW ENGL J MED	10.1056/NEJMoa1411087	11	0.05	2015				
Coiffier B <sup>[32]</sup>	NEW ENGL J MED	10.1056/NEJMoa011795	10	0.19	2002				
Armand P <sup>[33]</sup>	J CLIN ONCOL	10.1200/JCO.2012.48.3685	10	0.3	2013				
Cheson BD <sup>[34]</sup>	J CLIN ONCOL	10.1200/JCO.2006.09.2403	9	0.18	2007				
Pfreundschuh M <sup>[35]</sup>	LANCET ONCOL	10.1016/S1470-2045(06)70664-7	8	0.09	2006				
Kiyasu J <sup>[36]</sup>	BLOOD	10.1182/blood-2015-02-629600	7	0.02	2015				
Herbst RS <sup>[37]</sup>	NATURE	10.1038/nature14011	7	0	2014				
Swerdlow SH <sup>[38]</sup>	BLOOD	10.1182/blood-2016-01-643569	7	0.02	2016				
Feugier P <sup>[39]</sup>	CLIN ONCOL	10.1200/JCO.2005.09.131	7	0.26	2005				
Coiffier B <sup>[40]</sup>	BLOOD	10.1182/blood-2010-03-276246	7	0.4	2010				
Andorsky DJ <sup>[41]</sup>	CLIN CANCER RES	10.1158/1078-0432.CCR-10-2660	7	0.01	2011				

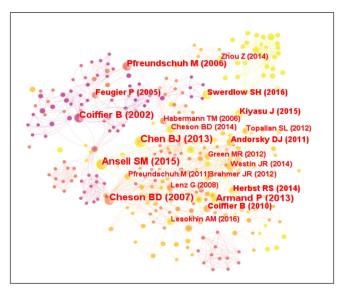


Figure 5: Network diagram of citation information.

journals, and literature with more than six citations. Among the journals with more than six citations BLOOD appeared three times, the highest number. CLIN CANCER RES, NEW ENGL J MED, and J CLIN ONCOL appeared two times each. LANCET ONCOL, NATURE, and CLIN ONCOL appeared one time each.

# **Keyword analysis**

## Keyword co-occurrence analysis

Keywords are a high level summary of the main content of a piece of literature and can therefore be analyzed to examine topical issues in a particular field. Figure 6 shows the keyword co-occurrence mapping of the literature related to immunotherapy for DLBCL, obtained by co-occurrence analysis of keywords in the literature through CiteSpace. There are a total of 422 nodes and 2447 linked lines in the graph. Each

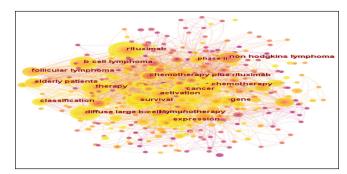


Figure 6: Keyword co-linear mapping of related literature.

node represents a keyword, with larger nodes representing more frequent occurrences of the keyword, and lines between nodes representing the degree of interconnectedness. The top three keywords were "expression," "diffuse large b-cell lymphoma," and "rituximab." The top 20 keywords were disease names: "diffuse large B-cell lymphoma," "non-Hodgkin's lymphoma," "B-cell lymphoma," "follicular lymphoma," and four others. The keywords related to treatment methods were: "rituximab," "chemotherapy," "immunotherapy," "cancer immunotherapy," and "transplantation" [Table 5].

# Keyword emergence analysis

A keyword emergent analysis can reflect the hot topics of research in a particular field at a particular time. A keyword emergent analysis yielded a graph of emergent terms in the field of immunotherapy for DLBCL [Figure 7]. "Bone marrow transplantation" was the earliest and longest-running term, appearing from 2004 to 2015." "Immune response" was also the earliest, appearing from 2004 to 2010. The three keywords that appeared until 2022, the last year of inclusion, were "cancer immunotherapy," "resistance," and "cytokine release syndrome," suggesting that this is a hot area of current research.

Table 5: Top 20 keyword information.								
Keyword	Frequency	Year	Centrality	Keyword	Frequency	Year	Centrality	
expression	71	2004	0.17	follicular lymphoma	29	2004	0.13	
diffuse large B-cell lymphoma	68	2005	0.09	elderly patients	27	2007	0.07	
rituximab	57	2009	0.11	classification	21	2004	0.09	
therapy	47	2003	0.07	chop	21	2008	0.03	
survival	46	2009	0.11	phase ii	19	2003	0.06	
cancer	46	2012	0.09	outcome	18	2017	0.03	
chemotherapy	43	2003	0.18	cancer immunotherapy	16	2011	0.05	
non-Hodgkin's lymphoma	43	2003	0.13	activation	16	2009	0.05	
B-cell lymphoma	42	2006	0.12	transplantation	16	2017	0.03	
immunotherapy	32	2004	0.14	gene	15	2004	0.07	

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Keywords	Year	Strength	Begin	End	2003 - 2022	
one marrow transplantation	2004	3.04	2004	2015		

bone marrow transplantation	2004	3.04	2004	2015
immune responses	2004	2.17	2004	2010
high dose therapy	2005	2.53	2005	2015
epstein barr virus	2006	2.39	2006	2013
non hodgkins lymphoma	2003	10.42	2007	2015
chemotherapy plus rituximab	2007	3.38	2007	2014
chop	2008	2.86	2008	2016
autologous transplantation	2008	2.22	2008	2017
chop chemotherapy	2004	3.01	2010	2014
trial	2003	3.15	2011	2012
prognosis	2011	2.88	2011	2014
follicular lymphoma	2004	4.56	2015	2017
t cells	2015	4.37	2015	2019
brentuximab vedotin	2015	2.54	2015	2019
hodgkin lymphoma	2016	2.66	2016	2019
death ligand 1	2018	6.14	2018	2020
cancer immunotherapy	2011	3.1	2018	2022
classical hodgkin lymphoma	2019	2.49	2019	2020
resistance	2019	2.23	2019	2022
cytokine release syndrome	2020	2.4	2020	2022

Figure 7. Anal	weie of konword	prominence in	related literature.
rigure /: Anai	lysis of Keyword	prominence m	related interature.

## Keyword clustering analysis

The keywords were analyzed by Log-Likelihood Ratio (LLR) algorithm, and the keywords in the field of immunotherapy for diffuse large B-cell lymphoma were divided into nine clusters [Figure 8 and Table 6]. The keyword clustering map is shown below. Several types of clusters were formed, with #0 being the largest, #1 the second largest, and so on. Among them "pd-l1," "antibody-based," "immunotherapy," and "cd19" clusters are predominantly studied. As can be seen from the keyword clustering map in Figure 6, the clusters cover each other, indicating that the clusters are closely linked and the research topics have a concentrated character.

# DISCUSSION

As can be seen from the study results, the research on DLBCL immunotherapy has been on the rise over the past 20 years. Although the initial research literature is small and the related research results are limited, the research in this field has shown

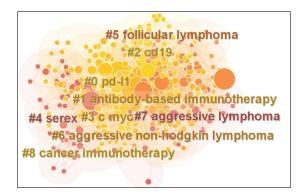


Figure 8: Cluster analysis of keywords in related literature.

a clear upward trend in recent years. This indicates that the research in the field of DLBCL immunotherapy is gradually receiving more attention, and there is great potential for future research development.

International Research Gaps: The study found that the United States leads the way in DLBCL immunotherapy research,

Table 6: Keyword clustering.			
Top terms	Size	Silhouette	Mean
pd-l1 (18.94, 1.0E-4); immune checkpoint (12.6, 0.001); expression (11.27, 0.001); survival (9.48, 0.005); pd 1 blockade (9.44, 0.005)	77	0.694	2014
antibody-based immunotherapy (17.51, 1.0E-4); diffuse large B-cell lymphoma (15.59, 1.0E-4); lymphoma and Hodgkin's disease (13.99, 0.001); chop (10.48, 0.005); chemotherapeutic approaches (10.48, 0.005)	64	0.74	2013
cd19 (13.88, 0.001); B cell (11.35, 0.001); cellular (7.56, 0.01); chimeric antigen (7.56, 0.01); loncastuximab tesirine (7.56, 0.01)	53	0.713	2015
c myc (13.68, 0.001); ct (9.1, 0.005); phase i (9.1, 0.005); gene expression (9.1, 0.005); flow cytometry (9.1, 0.005)	46	0.768	2014
serex (15.18, 1.0E-4); cancer testis antigen (10.11, 0.005); gene (7.36, 0.01); dendritic cells (6.45, 0.05); b-cell lymphoma (5.42, 0.05)	44	0.897	2007
follicular lymphoma (10.46, 0.005); bispecific antibody (7.9, 0.005); radioimmunotherapy (7.9, 0.005); low grade (7.9, 0.005); detude des lymphm (7.9, 0.005)	44	0.843	2008
aggressive non-Hodgkin's lymphoma (10.49, 0.005); immunohistochemistry (5.25, 0.05); alk plus large B cell lymphoma (5.23, 0.05); treatments (5.23, 0.05); bcl 6 protein expression (5.23, 0.05)	37	0.702	2011
aggressive lymphoma (10.46, 0.005); autologous transplantation (7.66, 0.01); hematopoietic stem cell transplantation (7.08, 0.01); ctl019 chimeric antigen receptor (7.08, 0.01); autologous stem cell transplantation (7.08, 0.01)	24	0.872	2008
cancer immunotherapy (20.69, 1.0E-4); antibody (15.12, 0.001); oncogenic stat3 signaling (12.99, 0.001); lymphoma microenvironment (12.99, 0.001); dlbcl mouse models (12.99, 0.001)	23	0.863	2016

with more than twice as many articles as other countries. In contrast, other developing countries except China have less research on immunotherapy for DLBCL. This means that other countries' research in this field needs to be further upgraded and improved to narrow the gap with leading countries.

Collaboration and interinstitutional collaboration: The findings show that the field of DLBCL immunotherapy involves multiple research institutions, and there is a strong collaboration between these institutions. In particular, UTMD Anderson Cancer Center, Harvard University, and French National Institute of Health and Medical Research. Many research articles have been published by institutions such as Inserm. This suggests that cross-institutional collaboration and collaboration in DLBCL's research contributes to more impactful research outcomes. Therefore, future research can continue to encourage collaboration between institutions to advance the field.

Hot research areas: Keyword analysis showed that the hot research areas mainly focused on the expression of DLBCL, Rituximab and other treatment methods. In addition, treatment methods such as immunotherapy, chemotherapy, cancer immunotherapy and transplantation are also receiving attention. Future research could further explore these hot areas and further study them.

# CONCLUSIONS

DLBCL is a malignant B-cell lymphoma and is one of the most common types of NHL.<sup>[1]</sup> Traditional treatments

include radiotherapy and chemotherapy, but these treatments can cause many side effects and are not effective for some patients.<sup>[12,13]</sup> Immunotherapy is a treatment that targets the tumor's immune system and can kill cancer cells by activating or boosting the body's immune system. For patients with DLBCL, immunotherapy may involve using specific antibodies to recognize and attack cancer cells, or by transferring immune cells to attack cancer cells.<sup>[17,18]</sup>

Conducting a bibliometric study can help us better understand the literature on immunotherapy in the treatment of DLBCL in order to understand current research trends and hot spots in the field. This information can provide an important reference for clinicians or researchers to help them produce more impactful research in this area and provide valuable references and insights for future relevant research. In addition, bibliometric studies can help us understand the research hotspots and knowledge gaps in the field, and provide directions and recommendations for future relevant research.

## Declaration of patients consent

Patient's consent not required as there are no patients in this study.

## Financial support and sponsorship

Nil.

## **Conflicts of interest**

There are no conflicts of interest.

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**How to cite this article:** Si W. A bibliometric analysis of CiteSpace software-based immunotherapy for the treatment of diffuse large B-cell lymphoma. Asian J Oncol, 2023;9:1.