#### REVIEW

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# Current Perspectives and Trend of Acupuncture in Breast Cancer-Related Symptoms: A Bibliometric Study

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**Purpose:** This bibliometric research aims to delineate global publication trends and emerging research interests in the use of acupuncture for breast cancer (BC)-related symptoms treatment over the past three decades. Furthermore, it identifies influential institutions, potential collaborative partners, and future research trends, thereby providing guidance for relevant, novel research directions.

**Methods:** Scientific publications related to acupuncture for BC-related symptoms were gathered from the Web of Science Core Collection (WoSCC) from 1993 to 2023. Four software applications were principally used to analyze the resulting data: the "bibliometrix" package in the R environment (version 4.2.3), VOSviewer, CiteSpace6.1.R6, and the bibliometrics website. These applications were employed to evaluate different parameters.

**Results:** A total of 621 papers on acupuncture in BC-related symptoms treatment were analyzed. The United States, China, and South Korea contributed the most, with Memorial Sloan Kettering Cancer Center, and Columbia University leading institutions. It is interesting to mention that Mao, Jun J. and Molassiotis, A. feature among the top 10 authors and co-cited authors. JAMA is the leading journal, with an ongoing focus on acupuncture's effectiveness. Keywords show that the initial research focus was mainly on "vasomotor symptoms", but in recent years there has been a gradual shift towards "pain", "chemotherapy-induced peripheral neuropathy (CIPN)", "electroacupuncture", and "non-specific effects".

**Conclusion:** Acupuncture has demonstrated a unique value in the process of adjuvant treatment of BC-related symptoms, and has been shown to be effective in reducing pain, eliminating fatigue, and improving quality of life. The study of the mechanisms of acupuncture and the application of electroacupuncture are possible future research priorities in this field. This study offers a deep perspective on acupuncture for BC research, highlighting key points and future trends.

Keywords: breast cancer-related symptoms, acupuncture, bibliometric study, cancer pain, CiteSpace, VOSviewer

## Introduction

Breast cancer (BC) is the most common female malignancy, accounting for 30% of all female malignancies.<sup>1</sup> Globally, BC affects about 2.1 million people a year.<sup>2</sup> Due to early diagnosis and better treatments, the mortality rates in BC have steadily fallen in recent years.<sup>3</sup> The most commonly used treatment approaches for BC include surgery, endocrine therapy, and chemotherapy.<sup>4</sup> However, these treatments are accompanied by significant adverse effects.<sup>5</sup> BC

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Finding a safe and effective strategy to enhance the QOL of BC survivors is critical. Based on suggestions from international clinical practice guideline development groups, acupuncture is a safe and ancient medical therapy for a number of cancer-related symptoms.<sup>10</sup> Numerous clinical studies on the efficacy and safety of acupuncture for treating BC-related symptoms have been published over the past 30 years.<sup>11</sup> For example, acupuncture not only could significantly relieve joint pain associated with aromatase inhibitors among BC survivors,<sup>12</sup> but it could also minimize the opioid dose and treat opioid-related adverse effects.<sup>13</sup> Meanwhile, Chien reported that acupuncture dramatically attenuated BC-related menopausal symptoms, and treatment gains were maintained up to 3 months after the completion of treatment.<sup>14</sup> Therefore, analyzing the application trends, research status, and hotspots of acupuncture for BC-related symptoms is critical for use with acupuncture in clinical practice and deeper fundamental research.

Bibliometrics has been extensively used in diverse clinical fields as an emerging method in recent years.<sup>15</sup> It is the application of statistical and mathematical techniques to scholarly publications, revealing multiple facets and study trends in that area.<sup>16</sup> In addition, the quantity of bibliometrics literature has exponentially increased over the past few years. Citespace is the most commonly used information visualization software based on citation analysis theory.<sup>17</sup> VOSviewer visualizes literature data in a multivariate, time-phased, and dynamic citation analysis language.<sup>16,18</sup> Using the Bibliometric software to examine articles in the R language environment is simple and quick.<sup>19</sup> In recent years, acupuncture-related bibliometrics have grown exponentially, including acupuncture for pain,<sup>20</sup> depression,<sup>21</sup> and inflammation.<sup>17</sup> To the best of our knowledge, acupuncture therapy's effects on BC-related symptoms have not yet been the subject of a thorough bibliometric analysis.

In this study, we use the bibliometric approach and scientific knowledge map to graphically analyze the research progress, current status, and hotspots of acupuncture for BC-related symptoms throughout the previous 30 years from multiple perspectives. We also predict future research trends in this area.

## **Material and Methods**

#### Source of Data and Search Strategy

The data was gathered from the Web of Science Core Collection (WoSCC) database. The WoSCC was selected because it is the most extensive and frequently utilized database, embodying the most significant and appropriate journals.<sup>22</sup> The search method included the topics "breast neoplasm" and "acupuncture therapy", with the limited time set from 1993 to 2023. Literature that was an article or review was included, and the language was set to English. Table 1 displays the precise search techniques and outcomes.

## Data Collection and Analysis

The data were retrieved by two independent reviewers from the WoSCC database. Any differences were settled by the decision of a third senior reviewer. With the help of the intrinsic function of WoSCC, the publication features were analyzed to provide a number of bibliometric parameters, primarily the Hirsch index, yearly publications, the output of countries, journals and citations. Furthermore, the impact factor of every journal was calculated using the Journal Citation Reports 2022 edition.

Data were transformed to "txt" format, designated "download\_\*.txt", and then loaded into the "bibliometrix" package in the R environment (version 4.2.3), VOSviewer, bibliometrics (<u>http://bibliometric.com/</u>), and CiteSpace V6.1.6 for analysis (Figure 1). The "bibliometrix" package in the R environment (version 4.2.3) function was used to automatically transform and analyze the bibliographic data of the selected articles. We analyzed information related to countries, the author's countries and keywords. The collaborative networks between institutions and authors were examined using

Set	Results	Search Query
#1	439,175	((TS=("breast neoplasm*" OR "breast neoplas*" OR "breast cancer*" OR "breast carcinoma*" OR "breast carcin*" OR "breast tumor*" OR "breast tumour*" OR "breast oncolog*" OR "mammary carcinoma*" OR "mammary cancer*" OR "mammary neoplasm*" OR "mammary tumor*" OR "mammary tumour*" OR "malignant neoplasm of breast" OR "malignant tumor of breast" OR "breast malignant tumor*" OR "cancer mammary" OR "carcinoma mammae" OR "cancer of breast")) AND DT=(Article OR Review)) AND LA=(English) Indexes=WoSCC, Timespan=1993-01-01 to 2023-03-21
#2	28,988	((TS=(acupunct* OR eletro-acupunct* OR electroacupunct* OR "acupuncture therapy" OR "acupuncture treatment*" OR "body acupuncture" OR "needle acupuncture" OR "manual acupuncture" OR "acupuncture points" OR electroacupuncture OR "warm acupuncture" OR moxibust* OR "acupuncture point*" OR "Auricular acupuncture" OR acupress* OR acupoint* OR auriculotherap* OR auriculoacupunct* OR meridian*)) AND DT=(Article OR Review)) AND LA=(English) Indexes=WoSCC, Timespan=1993-01-01 to 2023-03-21
#3	621	#1 AND #2

Table I The Topic Search Query

VOSviewer. The collaborative analysis of countries was specifically plotted using the bibliometrics website. CiteSpace's parameters were set as follows: (1) time slice: 1993–2023, 1 year per slice; (2) node type was chosen at a time; (3) 50 best items were used as selection criteria; (4) pruning: pathfinder. In addition, the timeline view of co-cited references and all items' keywords were captured, along with powerful citation burst maps.

# Results

## Analysis of Annual Publications

The volume of literature in a field and its changing trend can reflect the development stage. A total of 621 documents about acupuncture in BC research were included, including 442 "Articles" and 179 "Review Articles". Although the time frame of our search was nearly 30 years, Figure 2 demonstrates that the first study in this field was published in 1995.



Figure I Flow-chart of the study.



Figure 2 Annual trends of global scientific publications from 1995 to 2023.

From 1995 to the present, the number of publications shows a gradual upward trend over time. It is important to mention that there has been a substantial increase in 2018. Based on the time of publication and quantitative characteristics of the literature, research on acupuncture BC may be separated into two phases, namely the basic phase (1995–2017) and the significant growth phase (2018–2022). 272 documents were published, accounting for 51.04% of the total in Period 2. We anticipated that 85 papers would be written on acupuncture for BC in 2023 based on the fitting curve.

## Analysis of Countries

Forty-nine countries published more than one paper on acupuncture for the BC study between 1995 and 2023. Table 2 depicts the ten most productive nations and regions. The USA published the most papers (n = 226, 36.39%), followed by China (n = 159, 25.60%) and South Korea (n = 51, 8.21%). USA had the most citations (8936 times), the highest H-index (49) and centrality (0.33). The map of cross-country collaborations shows the density of collaboration between countries

Rank	Countries	Count	Centrality	H-Index	Citations
I	USA	226	0.33	49	8936
2	China	159	0.21	24	2014
3	The United Kingdom	51	0.19	24	2060
4	South Korea	51	0.16	16	883
5	Australia	41	0.04	14	907
6	Canada	34	0.03	14	1628
7	Germany	25	0.05	13	468
8	Israel	20	0	9	259
9	Italy	17	0.07	7	437
10	Sweden	14	0	9	433

 Table 2 Top 10 Productive Countries Related to Acupuncture for BC

(Figure 3A). The USA has research collaborations with many countries, with China Canada, and UK collaborating most closely. However, the level of collaboration between other countries is lower. In Figure 3B, we find that the USA has long been in first place in terms of annual postings until 2015. China's annual publication volume jumped in 2015 and was roughly equal to the USA in the following years, surpassing it in 2019. As shown in Figure 3C and D, papers in acupuncture for BC were mostly published in North America, Asia, and European nations.

## Analysis of Institutions

These papers were contributed by 1011 institutions, with a total of 161 papers being provided by the top 10 institutions, accounting for 25.93% of all the publications (Table 3). Memorial Sloan Kettering Cancer Center ranked first (n = 37, 5.96%), followed by Columbia University (n = 17, 2.74%). Memorial Sloan Kettering Cancer Center is ranked first in terms of intermediary centrality, indicating that it is very influential in the field of acupuncture for BC. Meanwhile, we noticed that although Harvard University is only 7<sup>th</sup> in relation to the number of articles published, it is second in terms of intermediary centrality at 0.13, indicating that this institution is also making an important contribution to the field.

## Authors and Co-Cited Authors

Over 3212 authors and 14,391 co-cited authors are included in this area of study. Among these authors, the top three were Mao, Jun J. (n = 27, 4.35%), Ben-arye, Eran (n = 15, 2.41%), and Bao, Ting (n = 12, 1.93%). The term "co-cited authors" refers to authors who have two (or more) writings referencing them at the same time. Among the leading 5 co-citation authors, Molassiotis, A. (n = 167, 26.89%) ranked first, followed by Mao, Jun J. (n = 144, 23.19%), and Deng G. (n = 110, 17.71%). Author/co-cited author collaboration network visualization is depicted in Figures 4A and B. There is close cooperation between authors, and the centrality of each author is much less than 0.1.



Figure 3 (A) Country Collaboration Map. (B) The changing trend of the annual publication quantity in the top ten countries from 1995 to 2023. (C) The international collaborations' visualization map of countries/regions. (D) Corresponding Author's Countries.

Rank	Institutions	Count	Centrality	H-Index	Citations
I	Mem Sloan Kettering Canc Ctr	37	0.15	18	1404
2	Columbia Univ	17	0.13	12	1731
3	Beijing Univ Chinese Med	16	0.02	8	127
4	Shanghai Univ Tradit Chinese Med	15	0.03	5	149
5	Clalit Hlth Serv	14	0.04	8	119
6	China Acad Chinese Med Sci	13	0.05	8	202
7	Harvard Univ	13	0.13	17	1406
8	Univ Michigan	12	0.03	9	575
9	Korea Inst Oriental Med	12	0.08	8	209
10	Technion Israel Inst Technol	12	0.02	8	139

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## Analysis of Journals and Cited Journals

The 621 articles collected were allocated to 217 publications. The top 10 journals published roughly 32.37% of all publications (Table 4). Among them, Integrative Cancer Therapies ranked first (47 articles, 7.57%), followed by Supportive Care in Cancer (36 articles, 5.79%) and Journal of Alternative and Complementary Medicine (19 articles, 3.06%).

The top three co-cited journals were the Journal of Clinical Oncology, Support Care Cancer, and Breast Cancer Research and Treatment (Table 5). This suggests that these journals hold significance in acupuncture research for the treatment of BC. The Jama-Journal of the American Medical Association, with the highest impact factor (IF) of 157.335, was also quite active.

Figure 5 demonstrates that the majority of the articles were published in the field of "Medicine, Medical, and Clinical" on the left, with three important outward citation paths visualized as green curves, and they were significantly influenced by the fields of "Molecular, Biology, Genetics", "Health, Nursing, Medicine", and "Psychology Education Social".

## Analysis of References

Table 6 presents the top ten papers ranked by co-citation frequency in descending order. The ten most co-cited references include one clinical practice guideline, two systematic reviews, and seven clinical studies. Excitingly, one clinical study



Figure 4 A network map showing authors (A) or co-cited authors (B) involved in acupuncture in research in relation to BC.

Rank	Journal	Count	Citations	IF	JCR
I	Integrative Cancer Therapies	47	719	3.077	Q2
2	Supportive Care in Cancer	36	687	3.359	Q
3	Journal of Alternative and Complementary Medicine	19	492	2.381	Q3
4	Medicine	18	50	1.817	Q3
5	Breast Cancer Research and Treatment	17	773	4.624	Q2
6	Acupuncture in Medicine	16	277	1.976	Q3
7	Journal of Clinical Oncology	15	2181	50.717	QI
8	Cancer	12	466	6.921	QI
9	BMC Complementary and Alternative Medicine	11	170	4.782	QI
10	Complementary Therapies in Clinical Practice	10	71	3.577	Q2

Table 4 The Top 10 Journals Involved in Research on Acupuncture in Relation to BC

Table 5 The Top 10 Co-Cited Journals Involved in Research on Acupuncture in Relation to BC

Rank	Co-Cited Journals	Count	Centrality	IF	JCR
I	Journal of Clinical Oncology	478	0.01	50.717	QI
2	Support Care Cancer	323	0.01	3.359	Q2
3	Cancer	308	0.01	6.921	QI
4	Breast Cancer Research and Treatment	284	0.05	4.624	Q2
5	Journal of Alternative and Complementary Medicine	276	0.01	2.381	Q3
6	Integrative Cancer Therapies	261	0.02	3.077	Q2
7	Jama-Journal of the American Medical Association	246	0.02	157.335	Q
8	Journal of Pain and Symptom Management	240	0.01	5.576	QI
9	Acupuncture in Medicine	234	0.03	1.976	Q3
10	Complementary Therapies in Medicine	224	0.01	3.335	Q2

was the most cited paper in the area from 1993 to 2023. These references were published in important and renowned journals. The review that had the greatest centrality of co-cited sources was in the journal of BREAST.

According to the dynamics of the research area, Figure 6 displays the top 25 references in the citation boom that occurred throughout the growth of acupuncture for BC research from 1993 to 2023. References with high strength levels are usually significant turning moments in the scientific mapping area. The milestone paper is a clinical trial of acupuncture on aromatase inhibitor-related joint pain.<sup>12</sup> This literature can be broadly categorized into two phases: the first phase of research focuses on clinical studies related to acupuncture for the treatment of vasodilatory symptoms of BC, especially hot flashes; the outbreak of literature in the last decade can be categorized as the second phase, where research is being transformed into clinical studies related to acupuncture for the treatment of pain and peripheral neuropathies of BC and related guidelines.

The cluster analysis of reference allowed us to observe those 10 clusters. Four of these clusters correlated with disease names, including #3 menopausal problem, #4 inhibitor-induced arthralgia, #6 reducing hot flash, and #7 night sweat. Another quartet focused on treatment techniques such as #0 cancer care, #1 acupuncture therapy, #2 alternative



Figure 5 A network map showing Journals and Co-cited Journals involved in acupuncture in research in relation to BC.

therapeutic approaches, and #9 Chinese medicine. The top ten clusters that were most recently active included #1 acupuncture therapy, #4 inhibitor-induced arthralgia, and #5 nonspecific effects (Figure 7).

## Analysis of Keywords

Publications yielded a total of 2171 keywords related to acupuncture for BC. Since 1993, a total of 25 keywords have experienced significant bursts. Figure 8 shows that keywords such as inhibitor-induced validation (2018–2023), inhibitor induced arthralgia (2019–2023), cancer pain (2019–2023), fatigue (2019–2023), therapy (2020–2023), induced peripheral neuropathy (2020–2023), safety (2020–2023), electroacupuncture (2021–2023), pain (2021–2023), and risk factor (2021–2023) have gained prominence in recent years. This indicates that research concerning these topics is currently garnering attention and signifies potential developmental trends in the field.

A cluster analysis of the keywords revealed 10 clusters. These clusters pertained mostly to disease name (#1, #3, #4), treatment (#0, #2, #5, #6, #8), and mechanism (#7). For a deeper exploration of characteristics of the temporal evolution

Rank	Co-Cited Reference	Count	Centrality	IF
I	Hershman DL, 2018, J AM MED ASSOC, V320, P167	53	0.02	157.335
2	Greenlee H, 2017, CANCER J CLIN, V67, P195	45	0.03	286.13
3	Molassiotis A, 2012, J CLIN ONCOL, V30, P4470	34	0.01	50.717
4	Lesi G, 2016, J CLIN ONCOL, V34, P1795	31	0.08	50.717
5	Walker EM, 2010, J CLIN ONCOL, V28, P634	29	0.07	50.717
6	Mao JJ, 2014, EUR J CANCER, V50, P267	28	0.1	10.002
7	He YH, 2020, JAMA ONCOL, V6, P271	27	0.02	33.006
8	Hervik J, 2009, BREAST CANCER RES TR, VII6, P311	26	0.07	4.624
9	Garcia MK, 2013, J CLIN ONCOL, V31, P952	26	0.05	50.717
10	Bokmand S, 2013, BREAST, V22, P320	25	0.15	4.254

Table 6 Top 10 Co-Cited References with the Highest Frequency

#### **Top 25 References with the Strongest Citation Bursts**

References	Year	Strength	Begin	End	1993 - 2023
Hervik J, 2009, BREAST CANCER RES TR, V116, P311, DOI 10.1007/s10549-008-0210-3, DOI	2009	11.79	2009	2014	
Frisk J, 2008, CLIMACTERIC, V11, P166, DOI 10.1080/13697130801958709, DOI	2008	8.74	2009	2013	
Deng G, 2007, J CLIN ONCOL, V25, P5584, DOI 10.1200/JCO.2007.12.0774, DOI	2007	7.34	2009	2012	
Nir Y, 2007, MATURITAS, V56, P383, DOI 10.1016/j.maturitas.2006.11.001, DOI	2007	6.77	2009	2012	
Walker EM, 2010, J CLIN ONCOL, V28, P634, DOI 10.1200/JCO.2009.23.5150, DOI	2010	13.43	2011	2015	
Crew KD, 2010, J CLIN ONCOL, V28, P1154, DOI 10.1200/JCO.2009.23.4708, DOI	2010	10.46	2012	2015	
Molassiotis A, 2012, J CLIN ONCOL, V30, P4470, DOI 10.1200/JCO.2012.41.6222, DOI	2012	14.13	2013	2017	
Bokmand S, 2013, BREAST, V22, P320, DOI 10.1016/j.breast.2012.07.015, DOI	2013	8.94	2013	2017	
Liljegren A, 2012, BREAST CANCER RES TR, V135, P791, DOI 10.1007/s10549-010-1283-3, DOI	2012	6.59	2013	2017	
Bao T, 2013, BREAST CANCER RES TR, V138, P167, DOI 10.1007/s10549-013-2427-z, DOI	2013	6.34	2013	2018	
Garcia MK, 2013, J CLIN ONCOL, V31, P952, DOI 10.1200/JCO.2012.43.5818, DOI	2013	10.54	2014	2018	
Bao T, 2014, CANCER-AM CANCER SOC, V120, P381, DOI 10.1002/cncr.28352, DOI	2014	7.41	2014	2017	
Mao JJ, 2014, EUR J CANCER, V50, P267, DOI 10.1016/j.ejca.2013.09.022, DOI	2014	9.74	2015	2019	
Frisk J, 2012, SUPPORT CARE CANCER, V20, P715, DOI 10.1007/s00520-011-1134-8, DOI	2012	6.49	2015	2017	
Mao JJ, 2015, J CLIN ONCOL, V33, P3615, DOI 10.1200/JCO.2015.60.9412, DOI	2015	9.05	2016	2020	
Mao JJ, 2014, CANCER-AM CANCER SOC, V120, P3744, DOI 10.1002/cncr.28917, DOI	2014	8.34	2016	2018	
Lesi G, 2016, J CLIN ONCOL, V34, P1795, DOI 10.1200/JCO.2015.63.2893, DOI	2016	10.29	2017	2021	
Chien TJ, 2015, J ALTERN COMPLEM MED, V21, P251, DOI 10.1089/acm.2014.0083, DOI	2015	8.17	2017	2020	
Greenlee H, 2017, CA-CANCER J CLIN, V67, P195, DOI 10.3322/caac.21397, DOI	2017	13.71	2018	2023	
Hershman DL, 2018, JAMA-J AM MED ASSOC, V320, P167, DOI 10.1001/jama.2018.8907, DOI	2018	18.28	2019	2023	
Chiu HY, 2017, EUR J CANCER CARE, V26, P0, DOI 10.1111/ecc.12457, DOI	2017	7.23	2019	2023	
Lyman GH, 2018, J CLIN ONCOL, V36, P2647, DOI 10.1200/JCO.2018.79.2721, DOI	2018	7.79	2020	2023	
Bao T, 2018, EUR J CANCER, V101, P12, DOI 10.1016/j.ejca.2018.06.008, DOI	2018	6.55	2020	2023	
He YH, 2020, JAMA ONCOL, V6, P271, DOI 10.1001/jamaoncol.2019.5233, DOI	2020	11.74	2021	2023	_
Lu WD, 2020, ONCOLOGIST, V25, P310, DOI 10.1634/theoncologist.2019-0489, DOI	2020	6.57	2021	2023	

Figure 6 Top 25 references with the strongest citation bursts from 1995 to 2023.



Figure 7 References cluster time-line map related to acupuncture on BC from 1995 to 2023.

Keywords	Year	Strength	Begin	End	1995 - 2023
united states	1998	7.18	1998	2009	
complementary therapy	1998	3.39	1998	2007	
prostate cancer	2001	3.22	2001	2014	
acupuncture point	2003	3.08	2003	2011	
applied relaxation	2005	4.21	2005	2013	
flashe	2008	3.36	2008	2013	
postmenopausal women	2002	3.29	2010	2011	
intervention	2009	4.33	2014	2018	
integrative medicine	2006	4.76	2015	2019	
prevention	2007	4.14	2017	2018	
traditional chinese medicine	2004	3.99	2017	2019	
validation	2018	4.53	2018	2023	
inhibitor induced arthralgia	2019	5.1	2019	2023	
cancer pain	2019	4.37	2019	2023	
fatigue	2006	3.1	2019	2023	
cancer-related fatigue	2009	2.91	2019	2021	
therapy	2000	6.52	2020	2023	
induced peripheral neuropathy	2014	4.79	2020	2023	
sham acupuncture	2014	4.12	2020	2021	
physical activity	2000	3.92	2020	2021	
american society	2015	2.94	2020	2021	
safety	2014	2.94	2020	2023	
electroacupuncture	2003	5.27	2021	2023	
pain	2002	3.9	2021	2023	
risk factor	2015	3.65	2021	2023	

#### **Top 25 Keywords with the Strongest Citation Bursts**

Figure 8 Top 25 keywords with the strongest citation bursts from 1995 to 2023.

of the knowledge base in acupuncture for BC, cluster analysis was performed using the timeline view tool of CiteSpace (Figure 9). The clusters that were most recently active included #0 complementary therapies, #1 hot flashes, #3 symptom cluster, #4 nausea, #5 trial, and #8 traditional Chinese medicine.

Thematic evolution maps offer an alternative perspective on identifying research hotspots and trends. From 1995 to 2012, the primary research focus was on "prevalence". The mainstream research topic shifted to "Quality of life" during the years 2013–2016, while "fatigue" emerged as a novel keyword from 2017–2019. In particular, the research on acupuncture for BC from 2020 to 2023 showed a notable increase in the appearance of "mechanisms" and "expression" (Figure 10A).

Moreover, we constructed a thematic map of hotspots, as shown in Figure 10B. The upper right quadrant, exemplifying strong centrality and high density, indicates that the research themes symbolized by the keywords in this region are highly popular. Consequently, prevalent themes during the period 1993–2003 include"hypnosis""clinical oncology program""risk factors" validation", and "functional assessment.

## Discussion

## General Information

For this particular investigation, we conducted a thorough exploration of the WoSCC databases to locate articles related to acupuncture for BC that were published between 1995 and 2023. This scientometric study comprised 621 articles published in 217 journals by 1011 institutions in 49 countries.



Figure 9 Keyword cluster time-line map related to acupuncture on BC from 1995 to 2023.



Figure 10 (A) Thematic map of keywords. (B) Keyword evolution analysis.

We can classify three-decades of development into two periods as the basic phase (1995–2017) and the significant growth phase (2018–2022). There was an increased number of patients developing complications such as pain, acupuncture has been widely used with its unique advantages in analgesia.<sup>23</sup> Aromatase inhibitors (AIs) have been included in the 2018 treatment guidelines for BC. About half of women who take AIs develop arthritis-like joint stiffness and pain, some clinical evidence suggests that acupuncture can successfully manage joint stiffness and pain.<sup>24</sup> Consequently, research on BC acupuncture therapy-related research has developed to a new level.

The number of articles published in the USA was far in advance and had taken the lead. One in eight women is likely to have BC, indicating that the USA is a country with a high rate of BC.<sup>25</sup> That could be hundreds of millions of dollars a year in research funding in the United States.<sup>26</sup> China was the only developing nation among these ten most published countries, which is closely linked with the increased provision in the medical field.<sup>27</sup> Memorial Sloan Kettering Cancer

Center, established in 1884, is consistently ranked in the top two of the best hospitals in the United States for cancer specialties. Mao Jun J., Bao Ting and Deng Gary are leading members of the institution researching the field.

The top four active authors were from the United States. This finding may mean that US researchers have made a significant contribution to this study. In terms of co-cited authors, the top ranking was Molassiotis Alex, followed by Mao Jun J. and Deng Gary. Molassiotis Alex (The Hong Kong Polytechnic University) is widely recognized for his substantial contributions to the development of clinical practice guidelines for evidence-based combination therapy for BC.<sup>28</sup> Mao Jun J.'s research is important for reducing peripheral neuropathy induced by neoadjuvant paclitaxel in BC patients.<sup>29</sup> Deng Gary focused on the study of acupuncture for dyspnea in BC patients and chronic fatigue after chemotherapy.<sup>30</sup>

#### **Knowledge Base**

The top 10 co-cited journals are mostly top-tier publications and consist of seven clinical trials, two reviews, and one clinical practice recommendation. Seven clinical studies outline the process of discovery and development of acupuncture for BC-related symptoms. The seven clinical studies included acupuncture for vasodilatory symptoms in women with BC-related symptoms,<sup>31–34</sup> aromatase inhibitor-related joint pain,<sup>12,35</sup> and cancer-caused fatigue.<sup>36</sup> Research shows acupuncture's increasing popularity as an alternative medicine worldwide due to its effectiveness. A high-IF study by Hershman included 226 early-stage BC patients who were taking AIs through a multicenter collaboration. They concluded that the true acupuncture group significantly reduced joint pain at 6 weeks.<sup>12</sup> However, accuracy issues were found in their test results.

Three articles, consisting of one clinical practice guideline and two systematic reviews, were examined. Metaanalyses and systematic reviews of *RCT* showed that acupuncture is effective in reducing cancer pain, nausea and can reduce the use of painkillers.<sup>37,38</sup> Additionally, in 2017, the Society for Integrative Oncology issued a clinical practice guideline that highlighted the role of acupuncture in symptom relief and acupuncture for the treatment of chemotherapyinduced nausea and vomiting (CINV) with a "B" grade of recommendation.<sup>28</sup>

#### **Research Hotspots**

The research hotspots, in addition to the symptoms such as pain and vasodilatory symptoms, which are also focused on QOL and electroacupuncture, are major highlights of the treatment.

Over 70% of advanced cancer patients experience moderate to severe pain,<sup>39</sup> adding to the current opioid crisis and highlighting the need for nonpharmacological treatments.<sup>40</sup> An evidence-based guideline strongly recommends acupuncture for arthralgia-induced pain,<sup>13</sup> especially in patients receiving aromatase inhibitors. A network meta-analysis of 17 RCTs involving 1516 participants ranked acupuncture as the top intervention for improving pain intensity,<sup>41</sup> showing acupuncture plays a great role in pain management. Apart from this, multiple RCTs suggested acupuncture can reduce CINV,<sup>42</sup> despite its limited role in prevention.<sup>43</sup> Importantly, acupuncture can also alleviate vasomotor symptoms like hot flushes and night sweats, which are prevalent in 65% of postmenopausal BC patients.<sup>44</sup> As the number of survivors increases, addressing these symptoms becomes crucial.<sup>45</sup> Hormone replacement therapy (HRT), often used for hot flashes, is controversial in BC patients.<sup>46</sup> Clinical studies have shown that acupuncture is effective in relieving hot flashes without elevating estradiol levels or inducing cancer recurrence.<sup>33</sup> The underlying mechanism of acupuncture, possibly by increasing in endorphin levels and reducing the level of the potent vasodilator calcitonin gene-related peptide continues to be a focus.<sup>47</sup> Moreover, acupuncture seems to significantly improve night sweats in BC survivors.<sup>48</sup> Measuring treatment effects on QOL is crucial, particularly for BC patients undergoing therapies such as anthracyclines (ANT), which often have significant side effects.<sup>49,50</sup> Acupuncture is effective in the secondary prevention of myelo-suppression during chemotherapy, and QOL improves in women during treatment.<sup>51</sup>

Electroacupuncture, which combines electrostimulation and traditional acupuncture needling, cannot only improve treatment efficiency but also be a targeted route of drug delivery. Preliminary findings suggest electroacupuncture can promote paclitaxel accumulation by altering the microvasculature and microenvironment of mouse BC tumors.<sup>52</sup> In addition, electroacupuncture significantly induced apoptosis in the tumors.<sup>53</sup> Mechanistic analyses have shown that electroacupuncture has been shown to directly decrease triple-negative breast cancer (TNBC) tumor growth by inhibiting

proteins involved in tumor angiogenesis and extracellular matrix, suppressing triple-negative BC-induced inflammation, and upregulating nerve growth factor receptors.<sup>54</sup> Therefore, future studies should focus on how acupuncture plays various roles in a more targeted way during each stage of BC.

## **Research Frontiers**

Pain is not only a current research hotspot; we predict that it is also a future research trend. In order to better investigate and characterize the cutting-edge field, we conducted a keyword and co-citation analysis of the keywords and co-cited literature for emergence and clustering. Current studies focused on "chemotherapy-induced peripheral neuropathy (CIPN)", "cancer-related fatigue (CRF)" and "nonspecific effect".

CIPN poses a significant clinical issue for BC patients,<sup>55</sup> with a prevalence rate ranging between 30% and 97%.<sup>56</sup> Mostly, it results in paresthesia and pain, which, according to the current interventions, are not sufficiently addressed.<sup>55</sup> However, promising evidence from acupuncture studies suggests pain reduction and enhanced nerve conduction in CIPN patients.<sup>57</sup> While findings from a phase IIA trial indicate acupuncture's potential in reducing CIPN's severity among BC patients undergoing neoadjuvant therapy,<sup>29</sup> it is essential to note the necessity for larger-scale studies to solidify these conclusions.

CRF has received inadequate attention, diagnosis, and care. A Study tracking fatigue trajectories in BC patients found that high fatigue is often accompanied by depressive tendencies and higher rates of sleep disorders.<sup>58</sup> A randomized controlled study found that acupuncture relieved physical and psychological fatigue and reduced the incidence of anxiety and depression.<sup>36</sup> Acupuncture should be advocated as an effective alternative therapy for CRF patients, particularly those with BC.<sup>59</sup> According to research, acupuncture works to reduce fatigue by influencing the variety and amount of intestinal bacteria, enhancing intestinal barrier performance, and modifying brain gut peptides.<sup>60</sup> With the advancement of genome and transcriptome sequencing, they showed that acupuncture can improve CRF in BC by blocking the Leptin/AMPK signaling pathway and thereby minimizing mitochondrial functional impairment.<sup>61</sup> While acupuncture presents several potential benefits in BC treatment, particularly in managing CIPN and CRF, future studies need to provide larger samples and maintain stringent controls to ensure the results' validity and reliability.

Increasing evidence shows that acupuncture therapy has nonspecific effects in addition to specific therapeutic effects.<sup>62</sup> Meta-analysis has shown that the clinical efficacy of "comfort needling" is also influenced by nonspecific effects and that comfort needling has some specific potency due to the contact of the treatment modality with the skin, resulting in the projection of somatosensory fibers to various brainstem nuclear area and hypothalamus.<sup>63</sup> Acupuncture does not only provide analgesia by suppressing the sensation of pain, but it can also provide analgesia by intervening on emotions and cognitions (expectations) to relieve the sensation of pain.<sup>64</sup> To further measure the specificity impact of acupuncture and to encourage the growth of acupuncture on a global scale, sham acupuncture control should be implemented in a way that minimizes the influence of factors such as patient-doctor communication, patients' expectations, and the attitude of the practitioner.

## Strengths and Limitations

Our study provides a systematic analysis of acupuncture treatment for BC-related symptoms, which can provide some guidance. We also used multiple bibliometric programs to analyze the research hotspots and trends in multiple dimensions. However, this study has several drawbacks. First, we just included articles from WoSCC. Again, the retrieved articles were limited to those published in English. Finally, complete annual data for 2023 is not yet available, new hotspot areas may have been missed, and further updates are needed for future studies.

# Conclusion

In summary, acupuncture treatment for BC-related symptoms is a challenging subject in clinical and basic research. Acupuncture treatment for BC-related symptoms is a relatively new field, receiving increased attention since 2018. The use of acupuncture in symptom management of BC has been broadening and the scope of research has gradually overstepped from single symptom management to an integrated oncology treatment model. A number of clinical trials have been conducted to validate the efficacy and safety of acupuncture therapy for the treatment of BC-related symptoms, with studies focusing on acupuncture's improvement of BC-related

symptoms and adverse effects after BC treatment. There are increasing studies on acupuncture in the treatment of pain, vasodilatory symptoms, CIPN, and CRF. These are important for promoting the use of acupuncture therapy and improving the quality of life of BC patients. Improving these symptoms and studying their mechanisms will be the focus of future research. In the future, basic and translational research should be carried out to reveal the mechanism of the effect of acupuncture and moxibustion, and at the same time, multi-center and large sample RCTs should be conducted to provide more credible data, and multi-methods combination therapy research should be carried out to enrich and standardize the treatment of breast cancer, so as to bring about a new development and change in the field.

## **Data Sharing Statement**

The data can be acquired directly from the WoSCC.

## **Ethics Approval**

This article does not contain any studies involving either human or animal subjects, thus, institutional review board ethical approval was not required.

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## **Author Contributions**

All authors agreed to submit the work to the current journal, gave final approval of the version to be published, and agreed to be responsible for all aspects of the work. They also all actively participated in the conception and design, acquisition, analysis, and interpretation of the data, contributed significantly to the article's writing, or critically revised it for important intellectual content.

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## Disclosure

The authors declare that they have no conflicts of interest in this work.

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