

# Hypothermia: what are the trends in recent studies? – a bibliometric analysis with global productivity

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# ABSTRACT

**Aim**: Although the number of global studies on hypothermia, which plays an important role in morbidity and mortality in adults and newborns, has increased, there is still no bibliometric research on this subject in the literature. This study, it was aimed to determine trend topics and global productivity by using various statistical analyzes of scientific articles published on hypothermia.

**Material and Method**: Articles on hypothermia published between 1980 and 2021 were downloaded from the Web of Science (WoS) database and analyzed using various statistical and bibliometric methods. Spearman's correlation coefficient was used for correlation studies. Network visualization maps were used to identify effective studies, global collaborations, and trend topics with citation analyses.

**Results**: Out of a total of 14410 publications, 8157 articles were analyzed. The top 5 contributors to the literature are USA (n=2938, 36%), Japan (737, 9%), UK (641, 7.8%), Germany (576, 7%), and China (544, 6%). was. The first 3 journals that published the most articles were Resuscitation (n=296), Critical Care Medicine (146), Therapeutic Hypothermia, and Temperature Management (135). The top 3 most active institutions were League of European Research Universities (n=448), University of California System (274), and Pennsylvania Commonwealth System of Higher Education (221). The most active author was Marianne Thoresen (n=69).

**Conclusion**: The most studied trend topics in recent years are determined as hypoxic-ischemic encephalopathy, neonatal encephalopathy, out-of-hospital cardiac arrest, neonates, targeted management, therapeutic hypothermia, extracorporeal membrane oxygenation, perioperative hypothermia, emergency medicine, outcome, mortality, and perinatal asphyxia. This study will guide the authors who want to study in this area.

Keywords: Bibliometric analysis, hypothermia, therapeutic, neuroprotection, trends

# INTRODUCTION

Hypothermia, an important issue in the history of medicine, is a decrease in core body temperature below 35°C (Mild hypothermia: 32°C to 35°C (90°F to 95°F), moderate hypothermia: 28°C to 32°C (82°F to 90°F), severe hypothermia: below 28°C (82°F)) and develops when the body's heat loss exceeds heat production (1,2). The temperature regulation center, which makes it possible to keep the core body temperature constant under changing environmental conditions, is located in the hypothalamus. If the body is exposed to cold, various mechanisms come into play to prevent heat loss and increase heat production. Death in hypothermia occurs by the mechanism of heart failure with asystole or ventricular fibrillation, which are factors that contribute to increased catecholamine levels, electrolyte disturbances, and cardiac oxygen depletion (2-4).

Although hypothermia is an emergency requiring immediate treatment, the therapeutic use of hypothermia is an important neuroprotection method. Therapeutic hypothermia is a promising neuroprotective intervention that has been shown to improve outcomes of nerve damage in humans. Until now, it has been proven that many neurological diseases such as stroke, traumatic brain injury, increased intracranial pressure, subarachnoid hemorrhage, spinal cord injury, hepatic encephalopathy, and neonatal peripartum encephalopathy are suppressed by therapeutic hypothermia (5). The neuroprotective role of hypothermia has been well established in cardiac arrest, hypoxic-ischemic encephalopathy, traumatic brain injury, and some other diseases (5-9).

Another special issue of hypothermia is neonatal hypothermia seen in newborns. According to the World

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Health Organization (WHO), neonatal hypothermia is defined as a core body temperature of  $< 36.5^{\circ}$ C or a skin temperature of  $< 36^{\circ}$ C (10).

Bibliometrics examines articles using various statistical methods (11-13). As a result of the analysis of the information obtained from thousands of articles in the literature with various statistical and bibliometric approaches, important information about a subject such as the most active countries, institutions, journals, authors, international collaborations, and past and future trends can be determined (14-16).

Although the number of global studies on hypothermia, which plays an important role in morbidity and mortality in adults and newborns, has increased, there is still no bibliometric research on this subject in the literature. This study, it was aimed to identify trend topics and reveal global productivity by holistically analyzing scientific articles on hypothermia published between 1980 and 2021 using various statistical methods and bibliometric approaches.

# MATERIAL AND METHOD

Since our research article is a bibliometric study, there is no need for an ethics committee approval.

#### Search Strategy

Web of Science Core Collection (WoS by Clarivate Analytics) database was used for the literature review. The search process was determined as 1980 - 2021. All publications with the phrase hypothermia in the title were accessed. So that researchers can access similar documents reproducibility codes: Title "hypothermia", Timespan: 1980-2021 (search findings may vary depending on different access dates, access date: 1 May 2022). VOSviewer (Version 1.6.16, Leiden University's Center for Science and Technology Studies, Netherlands) package program was used to create bibliometric network visualizations as a result of clustering analyses, citation analyses, and trend topic determination analyses (17).

# **Statistical Analysis**

The website 'https://app.datawrapper.de' was used to create the world map showing the distribution of articles by country. The Exponential Smoothing estimator using seasonal smoothing was used in Microsoft Office Excel to estimate the number of articles that could be published in the next 5 years based on past publication trends. Statistical analyzes were performed with SPSS (Version 22.0, SPSS Inc., Chicago, IL, USA) package program. The normal distribution test of the data was analyzed with the Shapiro-Wilks test. Correlation analyzes were performed to determine whether some economic development indicators (Gross Domestic Product (GDP), Gross Domestic Product per capita (GDP per capita), Human Development Index (HDI)) of countries affected hypothermia (data were obtained from the world bank (18)). Correlation analyzes were analyzed using the Spearman correlation coefficient as the data were not normally distributed. p<0.05 was accepted for a statistically significant relationship.

# RESULTS

As a result of the literature review, there were a total of 14410 publications on hypothermia published in all research areas in the WoS database during the 1980-2021 period. Of these publications, 56.6% (n=8157) were Articles, 22.7% (n=3275) Meeting Abstracts, 6.9% (n=1000) Letters, 4.3% (n=615) Proceedings Papers, 4.2% (n=602) were in Review Articles, and the remainder in other publication types (Notes, Corrections, Book Chapters, Early Access, News Items, Book Reviews, Discussions, Poetry, Biographical-Items, Data Papers). Bibliometric analyzes were carried out with 8157 articles published in the Article category out of a total of 14410 publications. 94.2% (n=7686) of these articles were in English and the remainder in other languages (German (156), French (90), Russian (84), Spanish (60), Japanese (23), Turkish (12), Portuguese (9), Czech (8), Polish (7), Italian (6), Korean (3), Serbian (3), Bulgarian (2), Ukrainian (2), Croatian (1), Hungarian (1), Norwegian (1), Serbo Croatian (1), Slovenian (1), Welsh (1)) (Table 1). The h-index of 8157 articles was 158, the average citations per article 25.8, and the sum of times cited 210.431 (without self-citations: 158,258) (Table 1). Most of the articles were scanned in SCI-Expanded (n=7671, 94%) and Emerging Sources Citation Index (ESCI) (n=424, 5.2%).

| Table 1. Language distribution and citations                     |                       |            |                               |        |  |  |
|--|-----------------------|------------|-------------------------------|--------|--|--|
| Language   | Number<br>of articles |            |                               |        |  |  |
| English  | 7686                  |            |                               |        |  |  |
| German   | 156                   |            | h- index                      | 158    |  |  |
| French   | 90                    |            |                               |        |  |  |
| Russian  | 84                    |            | Average                       | 25.8   |  |  |
| Spanish  | 60                    | s          |                               |        |  |  |
| Japanese   | 23                    | articles   | per                           |        |  |  |
| Turkish  | 12                    |            | article                       |        |  |  |
| Portuguese   | 9                     | Fotal 8157 |                               |        |  |  |
| Czech  | 8                     | al 8       | Sum of                        | 210421 |  |  |
| Polish   | 7                     | Iota       | times<br>cited                | 210431 |  |  |
| Italian  | 6                     | -          |                               |        |  |  |
| Korean/Serbian   | 3                     |            |                               | 158258 |  |  |
| Bulgarian/Ukrainian  | 2                     |            | Without<br>self-<br>citations |        |  |  |
| Croatian/Hungarian/ Norwegian/<br>Serbo Croatian/Slovenian/Welsh | 1                     |            |                               | 136238 |  |  |

# Development of publications over the years

The distribution of the number of articles published on hypothermia by years is shown in **Figure 1**. The values

related to the results of the Exponential Smoothing estimation model, which takes into account the seasonal correction used to estimate the number of articles that can be published in the next 5 years, are shown in **Figure 1**. According to the estimation model results, it was predicted that 366 (Confidence Interval 95%: 320-412) articles will be published in 2022 and 394 (CI 95%: 300-489) articles will be published in 2026 (**Figure 1**).



**Figure 1.** A bar chart illustrating the distribution of hypothermia articles published by year, as well as predictions for the number of articles to be published in the next five years.

#### **Research Areas**

The first 15 research areas with the most research on hypothermia are respectively; Neurosciences (1076, 13.2%), Critical Care Medicine (1073, 13.1%), Surgery (1034, 12.6%), Clinical Neurology (777, 9.5%), Pediatrics (755, 9.2%), Cardiac Cardiovascular Systems (726, 8.9 %), Anesthesiology (618, 7.5%), Emergency Medicine (610, 7.4%), Pharmacology Pharmacy (550, 6.7%), Medicine General Internal (494, 6%), Physiology (408, 5%), Peripheral Vascular Disease (339, 4.1%), Respiratory System (328, 4%), Medicine Research Experimental (283, 3.4%), Biochemistry Molecular Biology (206, 2.5%).

#### **Active Countries**

The distribution of the number of articles by country is shown in Figure 2. The first 21 countries that contributed the most to the literature by publishing more than 100 articles are respectively; USA (number of articles, n=2938, 36%), Japan (737, 9%), UK (641, 7.8%), Germany (576, 7%), China (544, 6%), Canada (482, 5.9%), France (304, 3.7%), Sweden (247, 3%), Netherlands (218, 2.6%), South Korea (208, 2.5%), Australia (194, 2.3%), Norway (182, 2.2) %), Italy (181, 2.2%), Austria (174, 2.1%), Spain (152, 1.8%), Turkey (147, 1.8%), Switzerland (144, 1.7%), Brazil (124, 1.5%) , Russia (115, 1.4%), India (103, 1.2%), and Poland (100, 1.2%) (Figure 2). Cluster analysis was performed among 60 countries that have published at least 5 articles from 110 countries that have published articles on hypothermia and whose authors have international cooperation, and it is shown in Figure 3a. According to the results of the clustering analysis, 10 different clusters related to international cooperation were formed (Colors for Cluster 1: red, Cluster 2: green, Cluster 3:

blue, Cluster 4: yellow, Cluster 5: purple, Cluster 6: turquoise, Cluster 7: orange, Cluster 8: brown, Cluster 9: pink, Cluster 10: orange). In addition, the total link strength (international cooperation score) scores showing the cooperation power of 60 countries were calculated and the International cooperation density map created according to these scores was shown in **Figure 3**b (Highest scoring countries: USA (805), England in the UK (423), Germany (345), France (233), Austria (232), Italy (223), Sweden (215), Canada (214), Netherlands (207), Norway (201), Switzerland (201)).



**Figure 2.** Global productivity world map showing the distribution of published articles on hypothermia by country



**Figure 3.** a. Network visualization map of results of cluster analysis showing international cooperation between countries on hypothermia Footnote: Each color denotes a distinct Cluster. The size of the circles representing the countries grows in proportion to the number of articles published by the countries. The lines indicate which countries they collaborate with. b. Density map showing the intensity of international cooperation of countries on hypothermia. Footnote: From blue to red (blue-green-yellow-red), the strength of international cooperation score increases.

#### **Correlation Analysis**

A positive moderate statistically significant correlation was found between the number of articles produced by countries on hypothermia and GDP, GDP per capita, and HDI values (respectively, r=0.709, p<0.001; r=0.702, p<0.001, r=0.666, p<0.001).

# **Active Authors**

The top 10 most active authors on hypothermia are respectively; Thoresen M. (69), Dietrich WD. (45), Kochanek PM. (43), Sterz F. (40), Schwab S. (38), Sessler DI. (37), Shankaran S. (34), Safar P. (32), Yenari MA. (32), Gunn AJ. (31).

# **Active Institutions**

The 15 most active institutions on hypothermia are respectively; League of European Research Universities (448), University of California System (274), Pennsylvania Commonwealth System of Higher Education (221), University of Pittsburgh (178), University of London (142), University of Texas System (139), University of California San Francisco (128), Harvard University (126), Johns Hopkins University (121), Stanford University (111), University College London (109), US Department of Veterans Affairs (106), USA Veterans Health Administration (106), Imperial College London (104), University of Pennsylvania (98).

# **Active Journals**

8157 articles published on hypothermia were published in 1794 different journals. The first 56 journals that contributed the most to the literature by publishing 25 or more articles from these journals, the total number of citations received by the journals and the average number of citations per article are presented in **Table 2**.

# **Citation Analysis**

Among the 8157 articles reviewed, the first 20 articles with the highest number of citations according to the total number of citations are presented in **Table 3**. In the last column of **Table 3**, the average number of citations the articles received per year is given.

| Table 2. The 56 most active journals that have published 25 or more articles on hypothermia |             |             |       |   |    |      |      |  |
|---|-------------|-------------|-------|---|----|------|------|--|
| Journals  | RC          | С           | AC    | Journals  | RC | С    | AC   |  |
| Resuscitation   | 296         | 12056       | 40.7  | Journal of Applied Physiology   | 41 | 1055 | 25.7 |  |
| Critical Care Medicine  | 146         | 8457        | 57.9  | Journal of Maternal-Fetal & Neonatal Medicine                                       | 37 | 306  | 8.3  |  |
| Therapeutic Hypothermia and<br>Temperature Management                                       | 135         | 760         | 5.6   | Life Sciences   | 37 | 774  | 20.9 |  |
| Brain Research  | 117         | 4202        | 35.9  | Acta Paediatrica  | 36 | 599  | 16.6 |  |
| Annals of Thoracic Surgery  | 97          | 2432        | 25.1  | Critical Care   | 36 | 1578 | 43.8 |  |
| Plos One  | 92          | 1444        | 15.7  | Neurological Research   | 36 | 702  | 19.5 |  |
| Journal of Thoracic and<br>Cardiovascular Surgery   | 91          | 3306        | 36.3  | Shock   | 36 | 811  | 22.5 |  |
| Journal of Neurotrauma  | 87          | 3958        | 45.5  | Journal of Neurosurgical Anesthesiology   | 35 | 854  | 24.4 |  |
| Anesthesia and Analgesia  | 86          | 2669        | 31.0  | Journal of Thermal Biology  | 35 | 195  | 5.6  |  |
| Stroke  | 85          | 8541        | 100.5 | American Journal of Physiology-Regulatory<br>Integrative and Comparative Physiology | 34 | 924  | 27.2 |  |
| Anesthesiology  | 81          | 5673        | 70.0  | Anaesthesia   | 34 | 834  | 24.5 |  |
| Journal of Cerebral Blood Flow and Metabolism   | 79          | 5901        | 74.7  | Archives of Disease in Childhood-Fetal and Neonatal Edition                         | 34 | 934  | 27.5 |  |
| Acta Anaesthesiologica Scandinavica   | 76          | 2393        | 31.5  | Neurosurgery  | 34 | 2080 | 61.2 |  |
| American Journal of Emergency<br>Medicine   | 75          | 785         | 10.5  | American Journal of Perinatology  | 33 | 290  | 8.8  |  |
| Journal of Trauma-Injury Infection<br>and Critical Care                                     | 69          | 4465        | 64.7  | Circulation   | 33 | 2621 | 79.4 |  |
| Pediatric Research  | 68          | 2600        | 38.2  | Anaesthesist  | 32 | 194  | 6.1  |  |
| Cryobiology   | 63          | 979         | 15.5  | Pediatric Critical Care Medicine  | 30 | 685  | 22.8 |  |
| Pharmacology Biochemistry and Behavior  | 62          | 1034        | 16.7  | Scandinavian Journal of Trauma Resuscitation & Emergency Medicine                   | 30 | 561  | 18.7 |  |
| European Journal of Pharmacology  | 54          | 1439        | 26.6  | British Journal of Anaesthesia  | 29 | 839  | 28.9 |  |
| Neuroscience Letters  | 49          | 1218        | 24.9  | Experimental Neurology  | 29 | 1116 | 38.5 |  |
| Journal of Perinatology   | 48          | 853         | 17.8  | Intensive Care Medicine   | 29 | 1409 | 48.6 |  |
| Journal of Surgical Research  | 47          | 786         | 16.7  | Neonatology   | 28 | 489  | 17.5 |  |
| Journal of Pediatrics   | 46          | 1920        | 41.7  | Neuroscience  | 28 | 928  | 33.1 |  |
| Bulletin of Experimental Biology and Medicine   | 45          | 26          | 0.6   | Pediatrics  | 28 | 2746 | 98.1 |  |
| Journal of Neurosurgery   | 44          | 3818        | 86.8  | Annals of Emergency Medicine  | 27 | 1134 | 42.0 |  |
| Neurocritical Care  | 42          | 1243        | 29.6  | Neuropharmacology   | 26 | 555  | 21.3 |  |
| Scientific Reports  | 42          | 555         | 13.2  | Psychopharmacology  | 26 | 780  | 30.0 |  |
| European Journal of Cardio-Thoracic<br>Surgery  | 41          | 1072        | 26.1  | Pediatric Neurology   | 25 | 965  | 38.6 |  |
| C: Record count, C: Number of citation, AC: Avera   | ge citation | n per docur | nent  |   |    |      |      |  |

| Tab   | le 3. The top 20 most cited articles on hypothermia  |                                 |   |      |      |       |
|-------|--|---------------------------------|---|------|------|-------|
| No    | Article  | Author                          | Journal   | РҮ   | TC   | AC    |
| 1     | Treatment of comatose survivors of out-of-hospital cardiac arrest with induced hypothermia   | Bernard SA. et al.              | New England<br>Journal of Medicine                          | 2002 | 3687 | 175.5 |
| 2     | Mild therapeutic hypothermia to improve the neurologic outcome after cardiac arrest  | Holzer M. et al.                | New England<br>Journal of Medicine                          | 2002 | 3426 | 163.1 |
| 3     | Whole-body hypothermia for neonates with hypoxic-<br>ischemic encephalopathy   | Shankaran S. et al.             | New England<br>Journal of Medicine                          | 2005 | 1787 | 99.2  |
| 4     | Selective head cooling with mild systemic hypothermia after neonatal encephalopathy: multicentre randomised trial  | Gluckman PD. et al.             | Lancet  | 2005 | 1531 | 85    |
| 5     | Moderate Hypothermia to Treat Perinatal Asphyxial Encephalopathy.  | Azzopardi DV. et al.            | New England<br>Journal of Medicine                          | 2009 | 1117 | 79.7  |
| 6     | Defense strategies against hypoxia and hypothermia   | Hochachka PW.                   | Science   | 1986 | 968  | 26.1  |
| 7     | Treatment of traumatic brain injury with moderate hypothermia  | Marion DW. et al.               | New England<br>Journal of Medicine                          | 1997 | 934  | 35.9  |
| 8     | Effect of mild hypothermia on ischemia-induced release of neurotransmitters and free fatty-acids in rat-brain  | Busto R. et al.                 | Stroke  | 1989 | 925  | 27.2  |
| 9     | Lack of effect of induction of hypothermia after acute brain injury.   | Clifton GL. et al.              | New England<br>Journal of Medicine                          | 2001 | 904  | 41    |
| 10    | Mild hypothermia increases blood loss and transfusion requirements during total hip arthroplasty   | Schmied H. et al.               | Lancet  | 1996 | 613  | 22.7  |
| 11    | Neurological outcomes at 18 months of age after<br>moderate hypothermia for perinatal hypoxic ischaemic<br>encephalopathy: synthesis and meta-analysis of trial data | Edwards AD. et al.              | BMJ-British<br>Medical Journal                              | 2010 | 601  | 46.2  |
| 12    | Hypothermia but not the n-methyl-d-aspartate antagonist,<br>mk-801, attenuates neuronal damage in gerbils subjected to<br>transient global-ischemia                  | Buchan A. and pulsinelli<br>WA. | Journal of<br>Neuroscience                                  | 1990 | 547  | 16.5  |
| 13    | Deep hypothermia with circulatory arrest - determinants of stroke and early mortality in 656 patients  | Svensson LG. et al.             | Journal of Thoracic<br>and Cardiovascular<br>Surgery        | 1993 | 477  | 15.9  |
| 14    | Moderate hypothermia in the treatment of patients with severe middle cerebral artery infarction  | Schwab S. et al.                | Stroke  | 1998 | 473  | 18.9  |
| 15    | Glutamate release and free-radical production following<br>brain injury - effects of posttraumatic hypothermia   | Globus MYT. et al.              | Journal of<br>Neurochemistry                                | 1995 | 453  | 16.1  |
| 16    | Prognostication after cardiac arrest and hypothermia a prospective Study   | Rossetti AO. et al.             | Annals of<br>Neurology                                      | 2010 | 444  | 34.1  |
| 17    | Hypothermia in trauma victims - an ominous predictor of survival   | Jurkovich GJ. et al.            | Journal of Trauma-<br>Injury Infection and<br>Critical Care | 1987 | 444  | 12.3  |
| 18    | Effect of hypothermia on the coagulation cascade   | Rohrer MJ. and natalie AM.      | Critical Care<br>Medicine                                   | 1992 | 438  | 14.1  |
| 19    | Childhood Outcomes after Hypothermia for Neonatal<br>Encephalopathy  | Shankaran S. et al.             | New England<br>Journal of Medicine                          | 2012 | 436  | 39.6  |
| 20    | The effects of mild perioperative hypothermia on blood loss and transfusion requirement  | Rajagopal.an S. et al.          | Anesthesiology  | 2008 | 431  | 28.7  |
| PY: F | Publication year, TC: Total citation, AC: Average citations per year   |                                 |   |      |      |       |

#### **Co-citation Analysis**

There were a total of 116544 studies cited in the references section of all 8157 articles published on hypothermia. Among these studies, the 7 most influential studies with more than 350 citations and the most co-citations are respectively; Bernard et al. (2002) (Number of co-citations: NC=1079), Holzer et al. (2002) (NC=979), Shankaran et al. (2005) (NC=625), Gluckman et al. (2005) (NC=501), Azzopardi et al. (2009) (NC=388), Busto et al. (1989) (NC=386), and Busto et al. (1987) (NC=383) (19-25).

### Keyword Analysis and Trend Topics

In all of the 8157 articles published on hypothermia, 9330

different keywords were used. Among these keywords, 105 different keywords used in at least 20 different articles are shown in **Table 4**. The cluster network visualization map showing the results of the clustering analysis performed between these keywords is shown in **Figure 4**. As a result of the cluster analysis, it was seen that hypothermia subjects were divided into 7 different clusters (Colors for Cluster 1: red, Cluster 2: green, Cluster 3: blue, Cluster 4: yellow, Cluster 5: purple, Cluster 6: turquoise, Cluster 7: orange). The trend network visualization map performed to identify trend topics is shown in **Figure 5**, and the citation network visualization map performed to reveal the most cited topics is shown in **Figure 6**.

| Table 4. The 105 most frequently                  | used keywo        | rds in articles on hypothermia      |                   |                           |                   |
|---|-------------------|-------------------------------------|-------------------|---------------------------|-------------------|
| Keywords  | Number<br>of uses | Keywords                            | Number<br>of uses | Keywords                  | Number<br>of uses |
| hypothermia                                       | 2747              | cooling                             | 56                | core temperature          | 29                |
| therapeutic hypothermia                           | 564               | oxidative stress                    | 54                | hemorrhagic shock         | 29                |
| cardiac arrest                                    | 524               | cerebral blood flow                 | 50                | hibernation               | 29                |
| neuroprotection                                   | 242               | hyperthermia                        | 48                | shivering                 | 29                |
| hypoxic-ischemic (or ischaemic)<br>encephalopathy | 225               | intracranial pressure               | 47                | subarachnoid hemorrhage   | 29                |
| rat (s)   | 215               | magnetic resonance imaging          | 46                | metabolism                | 28                |
| temperature                                       | 181               | nitric oxide                        | 46                | coma                      | 27                |
| cardiopulmonary resuscitation                     | 173               | glutamate                           | 45                | electroencephalography    | 27                |
| mild hypothermia                                  | 172               | hippocampus                         | 45                | microglia                 | 27                |
| resuscitation                                     | 162               | out-of-hospital cardiac arrest      | 45                | critical care             | 26                |
| body temperature                                  | 148               | brain injury                        | 44                | microdialysis             | 26                |
| ischemia  | 143               | encephalopathy                      | 44                | induced                   | 25                |
| traumatic brain injury                            | 143               | moderate hypothermia                | 44                | lactate                   | 25                |
| outcome (s)                                       | 140               | reperfusion                         | 44                | mitochondria              | 25                |
| cerebral ischemia                                 | 135               | sepsis                              | 42                | coagulopathy              | 24                |
| stroke  | 123               | survival                            | 42                | ethanol                   | 24                |
| newborn   | 117               | targeted temperature management     | 42                | head injury               | 24                |
| neonate (s)                                       | 114               | brain edema                         | 41                | reactive oxygen species   | 24                |
| rewarming   | 110               | cytokines                           | 41                | infant                    | 23                |
| apoptosis   | 109               | hypoxia-ischemia                    | 40                | liver                     | 23                |
| inflammation                                      | 97                | myocardial infarction               | 39                | perioperative hypothermia | 23                |
| thermoregulation                                  | 94                | anesthesia                          | 38                | circulatory arrest        | 22                |
| accidental hypothermia                            | 92                | extracorporeal circulation          | 35                | emergency medicine        | 22                |
| induced hypothermia                               | 90                | neurological outcome                | 33                | hypothalamus              | 22                |
| asphyxia  | 88                | shock                               | 33                | lipopolysaccharide        | 22                |
| heart arrest                                      | 88                | children                            | 32                | blood-brain barrier       | 21                |
| cardiopulmonary bypass                            | 80                | mild therapeutic hypothermia        | 32                | brain temperature         | 21                |
| trauma  | 77                | pharmacokinetics                    | 32                | mrı                       | 21                |
| perinatal asphyxia                                | 74                | fever                               | 31                | pediatric                 | 21                |
| brain   | 71                | hemorrhage                          | 31                | brain ischemia            | 20                |
| mortality   | 71                | reperfusion injury                  | 31                | dopamine                  | 20                |
| ventricular fibrillation                          | 67                | spinal cord injury                  | 31                | heart                     | 20                |
| prognosis   | 65                | extracorporeal membrane oxygenation | 30                | kidney                    | 20                |
| neonatal encephalopathy                           | 64                | surgery                             | 30                | microcirculation          | 20                |
| hypoxia   | 59                | birth asphyxia                      | 29                | normothermia              | 20                |



**Figure 4.** Network visualization map for cluster analysis based on keyword analysis performed to identify clustering of hypothermia. Footnote: Each color denotes a distinct cluster. The color of keywords in the same cluster is the same. The circle represents the number of times the keyword is used in articles. The larger the circle represents the number of times the keyword is used in articles.



**Figure 5.** Network visualization map based on keyword analysis to identify past and current trends on hypothermia. Footnote: The article's topicality increases from blue to red as indicated by the indicator in the lower right corner of the figure (blue-green-yellow-red). The circle represents the number of times the keyword is used in articles. The larger the circle represents the number of times the keyword is used in articles.



**Figure 6.** Network visualization map based on keyword analysis performed to identify the most cited topics on hypothermia. Footnote: The number of citations received by the subject increases from blue to red in the indicator in the lower right corner of the figure (blue-green-yellow-red). The circle represents the number of times the keyword is used in articles. The larger the circle represents the number of times the keyword is used in articles.

# DISCUSSION

This study, in which the subject of hypothermia was evaluated, showed that the studies on hypothermia mainly focused on the neuroprotective effect of therapeutic hypothermia. This shows that therapeutic hypothermia is an important research topic. In addition, the increase in the annual average number of publications, especially in the 2008-2021 period, is higher than in previous periods. This may be related to both the development of technologies on hypothermia and the fact that therapeutic hypothermia applications are an important treatment method that keeps up to date.

Heart disease is still one of the main universal reasons for death. Acute myocardial infarction is one of the most common diseases in the world. (26,27). For this reason, millions of people lose their lives every year (28). This situation continues to be a global health problem despite all the precautions and developments in treatment strategies. Management of ischemic brain injury, especially due to late intervention after myocardial infarction, is a very difficult health problem. There are promising results regarding therapeutic hypothermia, which has been increasingly used in recent years in order to eliminate the morbidity and mortality that may occur after this situation (19,29,30). In this bibliometric analysis, although most of the studies involving therapeutic hypothermia applications are associated with ischemic encephalopathy, it is observed that this condition is predominantly associated with a prolonged cardiac arrest that develops after myocardial infarction. Another interesting result is that the broadcasts peaked especially with the COVID-19 pandemic. This situation can be evaluated as a result of the reflection of the therapeutic hypothermia applied to patients due to the increased frequency of severe hypoxia due to COVID-19 pneumonia and subsequent hypoxic encephalopathy in the publications.

When the articles were examined and the number of studies of the countries was evaluated, it was seen that particularly developed countries were at the forefront. This situation does not come as a surprise when we evaluate the economic power of an application that requires advanced technology such as therapeutic hypothermia and the pioneers in the production of these devices. In addition, the cooperation scores of the countries in the studies on these issues were again at the forefront in the developed countries.

When the co-authorship cooperation of countries on hypothermia is evaluated, although some international global collaborations have been observed, it is thought that regional international collaborations based on the geographical neighborhood are common in article production. When the journals that publish the most articles on hypothermia are evaluated, it is seen that therapeutic hypothermia comes to the fore. In addition, it was observed that the studies focused on neurology, anesthesia, and emergency journals. We can recommend that authors who want to publish on hypothermia consider these journals first.

When the analyzed articles were evaluated according to the total number of citations they received and the average number of citations received per year, it was observed that the studies on hypothermia were predominantly post-cardiopulmonary arrest, followed by neonatal encephalopathy. Studies have been published showing that hypothermia treatment applied after postcardiopulmonary arrest and neonatal encephalopathy has positive clinical results. (19,20,23) These results suggest that therapeutic hypothermia applications will intensify in an increasing trend, especially in these subjects. In addition, it is not surprising that similar results are obtained when citation analysis is performed.

Neonatal hypothermia is one of the most important issues in hypothermia. According to the World Health Organization (WHO), neonatal hypothermia is defined as a core body temperature of < 36.5 °C or a skin temperature of < 36 °C (10). Neonatal hypothermia is a global health problem and is still an important risk factor for neonatal morbidity and mortality, especially in lowand middle-income countries (31). In a meta-analysis study conducted for East Africa with 20911 participants (a total of 12 potential studies), the prevalence of neonatal hypothermia was found to be 57.2% (95%CI; 39.5–75.0) (32). Awareness of neonatal hypothermia is still low in some African populations. Risk factors for neonatal hypothermia in the region include poverty, preterm birth, night birth, home birth, low birth weight, early bathing of newborns, late initiation of breastfeeding, and inadequate knowledge among health workers (31,32). Ironically, neonatal hypothermia, which is frequently encountered in underdeveloped regions such as African countries, is reflected in the publications as a condition that needs to be treated, while in developed countries, it gains weight in publications where hypothermia is used for therapeutic purposes (33). In addition, the limited number of studies on neonatal hypothermia may be associated with less sensitivity in underdeveloped populations and ignorance of the diagnosis of neonatal hypothermia.

As a result of our literature review, we did not find any bibliometric study on hypothermia. It can be said as the advantage of our study is that our research is the first bibliometric study on this subject. In addition, the use of many statistical approaches such as international cooperation analysis, citation analysis, trend keyword analysis, and correlation analysis can be said to be the other superior aspects of our study. A limitation of the study is that we used only the WoS database in the literature review. However, citation analyzes cannot be performed in the PubMed database. In addition, the WoS database indexes the articles published in more effective journals (SCI-expanded, ESCI, and SSCI-indexed journals) compared to the Scopus database (34).

# CONCLUSION

In this comprehensive bibliometric research, conducted on hypothermia, it was shared the statistical analysis information of 8157 articles published from the past to the present. The most researched trend topics in recent years were determined as hypoxic-ischemic encephalopathy, neonatal encephalopathy, neonates, targeted management, therapeutic hypothermia, outof-hospital cardiac arrest, extracorporeal membrane oxygenation, perioperative hypothermia, emergency medicine, outcome, mortality, perinatal asphyxia. Developed countries with large economies had a say in global productivity and international cooperation. This study will guide the authors who want to study in this area.

# ETHICAL DECLARATIONS

**Ethics Committee Approval:** Since our research article is a bibliometric study, there is no need for an ethics committee approval.

**Informed Consent:** For this type of study, formal consent is not required.

Referee Evaluation Process: Externally peer-reviewed.

**Conflict of Interest Statement:** The authors have no conflicts of interest to declare.

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