

Review Paper

Medical Causes of Death in Iranian Children Aged 1-59 Months: A Systematic Review and Meta-analysis



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ABSTRACT

Background: Preventing the years of life lost in newborns and children is one of the serious public health problems in underdeveloped countries.

Objectives: This review aimed to highlight the significant determinants related to under-five child mortality in Iran (both in-hospital and out-of-hospital) through a systematic review of the literature.

Methods: EMBASE, PubMed, Scopus, Magiran and Web of Science databases, as well as the Google Scholar search engine, were used for the systematic search of the literature for Iranian studies. Special syntax was used to search the relevant studies for the review. Original research articles in English and Persian until December 31, 2023, were included in the analysis and synthesis of the results. A total of 15 studies were incorporated into this review. Eligible articles were thoroughly appraised and pertinent information was extracted and integrated into the systematic review.

Results: The highest frequency of mortality occurred in boys. The three most significant causes of death among Iranian children aged 1-59 months, respectively, are congenital disorders (28%), respiratory distress syndrome (18%) and non-traffic and traffic accidents (17%).

Conclusions: The most common causes of mortality among Iranian children aged 1-59 include congenital abnormalities, unintentional injuries, mortality due to external factors, and respiratory infections. However, without accurate recording of causes of death using international classification of diseases 10th Revision (ICD-10), as well as conducting autopsy or registering precise age periods for children under five years old, it is not possible to adequately analyze extensive patterns of mortality in the country. Consequently, related public health planning is diminished, which negatively impacts efforts to effectively reduce mortality.

Key Words:

Children, Mortality, Iran

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Introduction

Child mortality rates serve as indicators of a country's development and have been selected as a measure of progress [1]. This is crucial, as it is a significant determinant of health and human advancement, particularly for deaths that are preventable after the first weeks of life [2, 3].

According to the statistics published by the [World Health Organization \(WHO\)](#), there has been a decrease in the under-five mortality rate in recent years. In Iran, the rate was reported as 13.86 deaths per 1000 live births in 2018, which decreased to 13.40 deaths per 1000 live births in 2019. The 2019 disaggregated data showed mortality rates of 14.54 for boys and 13.28 for girls. The latest update from the organization states a global under-five mortality rate of 12.94 deaths per 1000 live births. According to studies, the rate of mortality of babies aged between 1 and 59 months was 80 out of 1000 births in low-income countries and six deaths per 1000 births in high-income countries [4, 5]. Some studies conducted across various regions of Iran have examined factors influencing mortality in children under five years old. The results of these studies indicate that several factors, such as parent's education level and occupation, housing conditions, the distance of children from healthcare facilities, maternal age at marriage and child-birth, developmental indices of provinces, parental literacy rates, maternal employment, average income, life expectancy, birth weight, infant nutrition, and socio-economic status are significant determinants. Additionally, factors, such as birth order, birth spacing, birth complications, like birth injuries and pregnancy disorders, chronic and infectious diseases, and mortality due to accidents have been identified as influential contributors to this rate [6-11].

One of the most important indicators of children's health is their mortality rate [1]. Understanding the causes of mortality is essential for implementing timely and effective preventive and therapeutic measures. Furthermore, through a more practical assessment of the risk factors contributing to children's mortality, the current situation can be improved with appropriate planning and policymaking. Identifying the causes and determinants of mortality is highly important for ensuring the design and appropriate implementation of interventions to improve population health [12].

Therefore, aimed to systematically review the causes of death for children aged 1-59 months in recent years in Iran, in order to identify the factors contributing to

child mortality to reduce mortality and improve children's health in Iran.

Methods

This systematic review and meta-analysis was performed based on the systematic reviews and meta-analysis (PRISMA) guidelines.

Research question

What are the causes of mortality related to children aged 1-59 months in Iran?

Search process

Based on the inclusion criteria, we searched electronic databases, including [EMBASE](#), [PubMed](#), [Scopus](#), [Magiran](#) and [Web of Science](#), as well as the [Google Scholar](#) search engine up to January 2024. Other gray literature (such as conference proceedings and key journals) was also searched. The following operators were used: ([Cause] OR [cause-specific] AND [cause of death] OR [death cause] OR [death causes] OR [mortality] OR [child mortality] Or [infant mortality] OR [child mortalities] OR [mortalities, child] OR [mortality, child] AND [infant] OR [infant, newborn] OR [infant, premature] OR [infant, postmature] OR [newborn infant] OR [newborn infants] OR [newborns] OR (newborn) OR [children] OR [child] OR [child, preschool] OR [childs] AND [1-59-month] OR [2 to 5 years old] OR [1 and 23 months] AND [Iran]).

Target population

The target population included all died Iranian neonates and children aged 1-59 months.

Inclusion and exclusion criteria

The inclusion criteria were all observational studies that reported the causes of death in neonates and children aged 1-59 months, regardless of the language. The exclusion criterion was studies conducted in other countries ([Table 1](#)).

Type of exposure

Death in neonates and children aged 1-59 months.

Data extraction (selection and coding)

Full-text articles meeting the inclusion criteria were evaluated. Information such as the first author's name, year of publication, study location, sample size, and

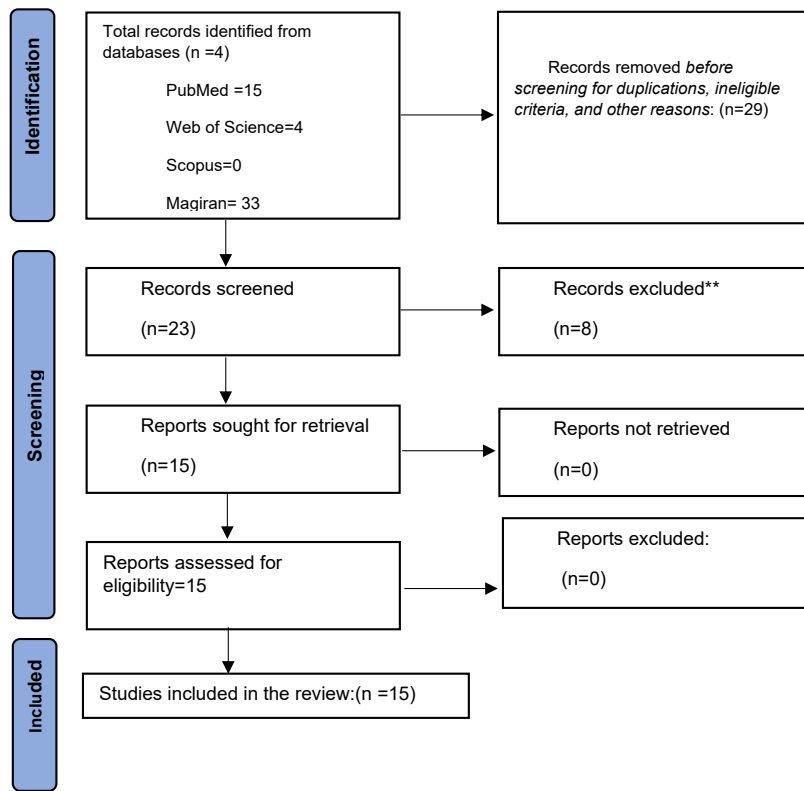


Figure 1. The PRISMA flowchart

causes of death was recorded in an Excel file. If additional information was needed from any of the articles, the corresponding author was contacted. All stages were performed independently by two individuals, and the compiled data were shared and combined.

Risk of bias assessment

The quality of studies was assessed independently by two individuals using the revised CONSORT risk of bias tool for clinical trials [17]. All studies that met the inclusion criteria were evaluated. Any discrepancies in assessments were resolved through discussion between the two individuals.

Analysis methods

Using Stata software, version 17 (Stata Corp LLC, College Station, Texas), the I² statistic was employed as

the criterion for evaluating heterogeneity based on the quality of studies for subgroup analysis. The Begg test was used to assess publication bias. P<0.05 were considered significant in statistical tests.

Results

During the years under study (1999-2020), 15 studies were registered in Iran. Figure 1 shows the number of articles included in the study.

Characteristics of the included studies

After initial evaluations, a total of 15 studies were included in the systematic review [2, 13-26]. The publication period of the articles ranged from 2010 to January 1, 2024 and the sample size of the studies varied from 65 to 3,576 patients (Table 2).

Table 1. The POLIS scale

POLIS Criteria	Patients	Outcome	Location	Indicator	Study Design
Description	Died children aged 1–59 months	Causes of death both in-hospital and out-of-hospital	Iran	Prevalence of causes of death	Observational

POLIS: Patients, outcome, location, indicator, and study design.

Table 2. Characteristics of the included studies

Variables	Chaman et al. 2012 [2]	Rahbar et al. 2013 [13]	Asadza-deh et al. 2022 [14]	Pirzadeh et al. 2022 [15]	Naghbi et al. 2015 [16]	Safari et al. 2014 [17]	Deihim et al. 2014 [18]	Sharaki Vahed et al. 2010 [19]	Nasrabi et al. 2022 [20]	Izadi et al. 2016 [21]	Tajedini et al. 2014 [25]	Ataey et al. 2018 [24]	Mo-menifar et al. 2023 [26]	Khadivi et al. 2020 [22]	Hosseini et al. 2022 [23]
Region	Shahroud, Iran	Iran	Babol	Qazvin	Mazandaran	Semnan	Dezful	Zabol	Neyshabur	Kermanshah	Tehran	Ardabil	Rafsanjan	Isfahan	Babol
Design of study	Nested case-control	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Nested case-control	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional
Study period	June 1999 to March 2009	2007-2008	2018-2020	2017-2020	2012	2002-2012	2010-2014	2007	2015-2019	2011-2014	2012	2011-2015	2011-2020	2012-2018	2009-2020
Sample size	65	5926	123	106	184	65	297	93	112	435	383	553	257	3576	303
Number of males	41	2809	75	41	94	30	171	59		239	201	311	162	1947	179
Number of females	24	3117	46	65	90	35	126	34		196	182	242	95	1627	124
The number of infant deaths (0-28 days)	156		139		115		136	63							
Number of infant deaths (28 days to 1 year)	3674				34		161								
Most significant causes of death in neonates and children aged 1-59 months	Congenital disorders (29.2%) and accidents (27.7%), congenital disorders (29.2%), and accidents (27.7%)	Congenital and chromosomal abnormalities (23.4%) and unintentional accidents (20.5%)	Prematurity (30%), congenital anomalies (25.3%), sepsis (17.1%), and respiratory distress syndrome (13%)	Sepsis (15.85%) and pneumonia (12.19%)	Respiratory diseases=26.1%	Congenital anomalies (n=15), accidents (n=14), and pneumonia (n=11)	Respiratory disorders, Congenital diseases, and asphyxia during birth	Respiratory diseases (n=25), accidents (n=25), and diseases of the digestive system (n=2)	Con-genital anomalies (n=36), respiratory system diseases and chronic diseases (n=19), non-traffic accidents (n=14), and road traffic accidents (n=11)	Accidents (n=81), respiratory system diseases and diseases of the cardiovascular system (n=53)	Accidents (n=59), congenital anomalies (n=67), and cancers (n=43)	Con-genital anomalies (n=276), special conditions originating in the prenatal period (n=119), and accidents (n=81)	Non-con-genital diseases (n=138), congenital diseases (n=48), and accidents (n=24)	Cardio-vascular diseases (n=382), accidents (n=379), diseases of the nervous system (n=253)	Congenital and chromosomal abnormalities (n=100), endocrine, nutritional, and metabolic diseases (n=56), and nervous system diseases (n=23)

Table 3. Quality evaluation of the included studies to check the risk of bias

Author	Was the Research Question or Objective in This Paper Clearly Stated	Was the study Population Clearly Specified and Defined?	Were the Inclusion and Exclusion Criteria for Participation in the Study Prespecified and Applied Uniformly to all Participants?	Were the Outcome Measures (Dependent Variables) Clearly Defined, valid, reliable, and implemented consistently Across all Study Participants?	Was the Timeframe Sufficient so that one Could Reasonably Expect to See an Association Between Exposure and Outcome if it Existed?	For the Analyses in this Paper, Were the Exposure(s) of Interest Measured Before the Outcome(s) Being Assessed?	Quality Assessment Result
Chaman et al. 2012 [2]	Yes	Yes	No	Yes	Yes	Yes	High
Rahbar et al. 2013 [13]	Yes	Yes	No	No	Yes	No	Moderate
Motlagh et al. 2013 [27]	Yes	Yes	No	Yes	No	Yes	Moderate
Asadzadeh et al. 2022 [14]	Yes	Yes	Yes	Yes	No	Yes	High
Pirzadeh et al. 2022 [15]	Yes	Yes	No	No	Yes	No	Moderate
Naghbi et al. 2015 [16]	Yes	Yes	No	Yes	No	Yes	Moderate
Ahmadi et al. 2014 [25]	Yes	Yes	No	Yes	Yes	No	Moderate
Safari et al. (2014) [17]	Yes	Yes	No	Yes	Yes	Yes	High
Deihim et al. 2015 [18]	Yes	Yes	Yes	Yes	Yes	No	High
Sharaki Vahed et al. 2010 [19]	Yes	Yes	No	No	No	No	Low
Nasrabadi et al. 2022 [20]	Yes	Yes	Yes	No	Yes	No	Moderate
Izadi et al. (2016) [21]	Yes	Yes	No	Yes	Yes	No	Moderate
Ataey et al. (2019) [24]	Yes	Yes	No	Yes	No	No	Moderate
Momenifar et al. (2023) [26]	Yes	Yes	No	No	Yes	No	Moderate
Khadivi et al. 2020 [22]	Yes	Yes	No	No	No	No	Low
Hosseini et al. 2020 [23]	Yes	Yes	Yes	Yes	Yes	Yes	High

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Quality assessment

About 31.2% of the studies had a low risk of bias, and 56.2% had a moderate risk. Two articles had a high risk of bias. In most studies, there was no explicit mention of the questions, "Was the timeframe sufficient so that one could reasonably expect to see an association between exposure and outcome if it existed?" and "For the analyses in this paper, were the exposures of interest measured before the outcomes?" This lack of clarity

represented the biggest weakness in the quality of the evaluated studies (Table 3).

The main outcome of interest in this study, the prevalence of congenital disorders, was analyzed in nine studies. The results of the data analysis in this study indicated that the highest frequency of mortality occurs in boys. The three most significant causes of death among Iranian neonates and children aged 1-59 months, respectively, are congenital disorders (28%), respiratory

Table 4. The most common causes of death in neonates and children aged 1-59 months in Iran

Variables	No. of Studies	Chi-squared test of Heterogeneity	P	Overall I-squared (%)	Z	P	Effect Size (95% CI)
Congenital disorders	9	179.24	<0.001	95.54	7.9	<0.001	0.28 (0.21-0.35)
Non-traffic and traffic accidents	9	205.51	<0.001	96.11	7.89	<0.001	0.17 (0.13-0.21)
Respiratory distress syndrome	4	10.94	<0.001	72.58	6.57	<0.001	0.18 (0.12-0.23)

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distress syndrome (18%) and non-traffic and traffic accidents (17%).

In this random-effects model, the mean score of this cause was calculated as effect size (EF)=0.28 [95% CI, 0.21%, 0.35%]. Other important causes of mortality in Iranian children aged 1-59 months include respiratory distress syndrome EF=0.18 [95% CI, 0.12%, 0.23%] and non-traffic and traffic accidents EF=0.17 [95% CI, 0.13%, 0.21%] (Table 4 and Figure 2).

Publication bias

The Begg's test was used to assess publication bias in this outcome. It examines the effect of small studies and shows a P=0.000. Since this value is significant, it indicates that there is no publication bias (adj. Kendall's score (P-Q)=1, SD=1.000, z=0.000, and Pr>|z|=1.000).

Discussion

The results indicated that the highest frequency of mortality occurs in boys. The three most significant causes of death among Iranian neonates and children aged 1-59 months, respectively, are congenital disorders (28%), respiratory distress syndrome (18%) and non-traffic and traffic accidents (17%).

The mortality of infants and children is a public health concern. Congenital abnormalities increase the risk of mortality during infancy and childhood, accounting for 26% of infant deaths and over 10% of deaths in children under ten years old in Europe [27].

Our results also indicated that the primary cause of mortality among children under five years old, with a prevalence of 28%, is congenital disorders. This finding is consistent with the results of other studies conducted in various countries [27-29].

It is crucial to implement primary and secondary preventive strategies, such as early referrals to specialized

centers for high-risk pregnancies and prenatal care. These strategies considerably reduce the death rate from congenital anomalies in children and adolescents [28, 29].

One of the weaknesses identified in the studies was the lack of precise registration of the types of congenital disorders and abnormalities. This oversight could have hindered efforts in primary and secondary prevention over time and impeded more accurate international comparisons. Globally, unintentional injuries and deaths resulting from external factors, such as accidents, being struck by objects at home, swallowing foreign objects, and various poisonings, including methadone poisoning or other toxic substances, are among the leading causes of death in children. These factors account for 30% of deaths in children aged 1 to 3 years, 40% of deaths in 4-year-old children, and 50% to 60% of deaths in children aged 5 to 17 years [30-32].

Since a significant percentage of these external causes of mortality are largely preventable but often overlooked, there is a critical need for effective education on injury prevention. This requires a multi-faceted approach [33]. In addition, conducting education programs on traffic safety is crucially important. In the event of such accidents, it is essential to promptly refer the child to the hospital. Rapid diagnosis and the initiation of treatment significantly increase the chances of a favorable outcome [32].

Our study findings revealed that the third leading cause of death among Iranian children under five years old is related to respiratory infections. Patients succumbed to severe viral or bacterial infections (such as sepsis, pneumonia, and COVID-19) despite receiving treatment, with little chance of survival. The COVID-19 pandemic has also impacted the causes of child mortality and altered their distribution. Most deceased children had an underlying condition, such as congenital abnormalities [32, 34]. The results of this analysis should be discussed, and the conclusions drawn should positively impact the

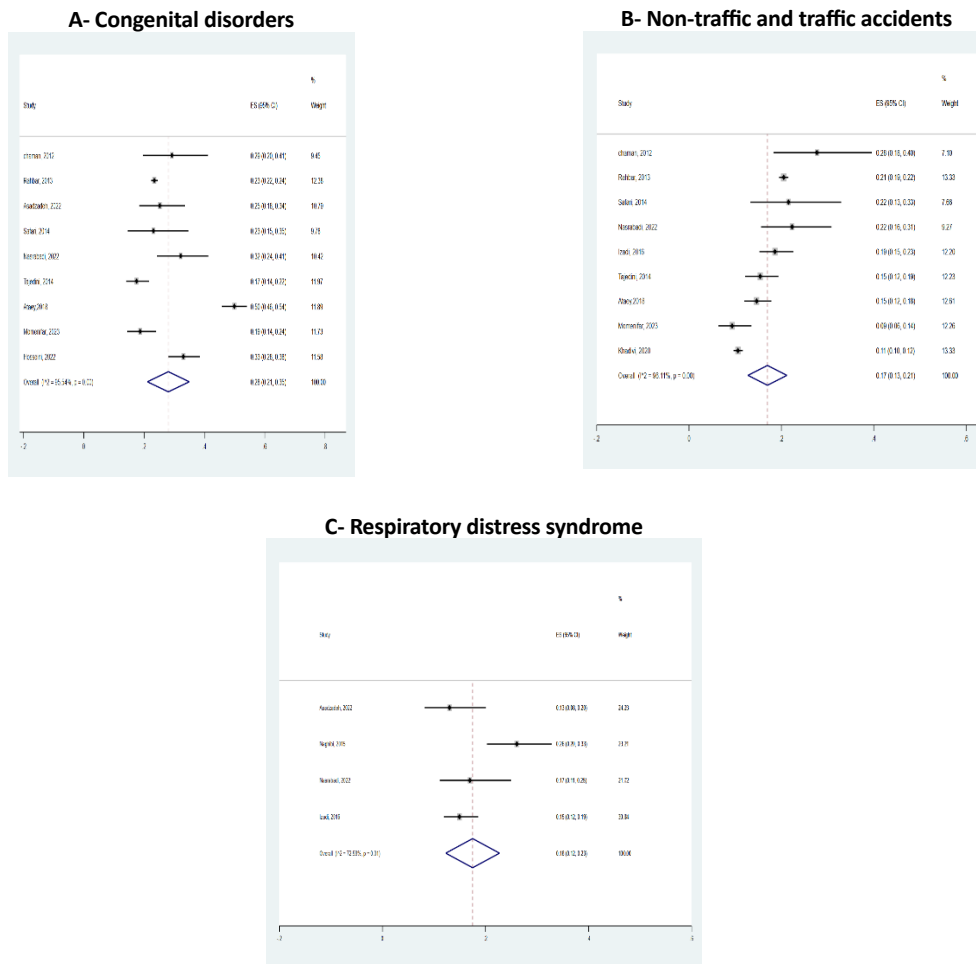


Figure 2. Forest plot of the prevalence of the most significant causes of death in Iranian children (neonates and children aged 1-59 months)

quality of childcare. Similar studies in other countries confirm that in both high and low-income countries, the leading cause of mortality among children under five years old is often respiratory infections and sepsis. This underscores the importance of addressing these issues to improve child health outcomes globally [35, 36]. Low levels of literacy among women may contribute to higher rates of neonatal infections. Consequently, illiterate mothers may struggle to recognize signs of danger and infection-related symptoms in infants, necessitating urgent medical care [36].

Study limitations: If autopsies are not conducted to determine the primary cause of death in deceased individuals, there is essentially no other definitive standard that clearly indicates the actual cause of death in a child. This lack of clarity is evident in studies. Additionally, in most studies, the age of deceased individuals was not categorized into neonatal deaths (0–27 days), post-neo-

natal deaths (28–364 days), or child deaths (over 365 days). Furthermore, none of the causes of death were recorded using international classification of diseases 10th revision (ICD-10) or ICD-9 coding.

Conclusion

The most common reasons for mortality in children aged 1-59 months in Iran are congenital abnormalities, unintentional injuries, and deaths due to external factors and respiratory infections. However, without accurate recording of causes of death using ICD-10 coding, without conducting autopsies or registering more detailed age categories for children under five years old, it is not possible to adequately analyze extensive patterns of mortality in the country. Consequently, related public health planning may suffer, leading to reduced effectiveness in health interventions.

Ethical Considerations

Compliance with ethical guidelines

This study was approved by the Ethics Committee of [Semnan University of Medical Sciences](#) (Code: IR.SEMUMS.REC.1400.10).

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Authors contributions

Conceptualization, project administration and funding acquisition: Gholamreza Mohammadi; Methodology and supervision: Masoudeh Babakhanian and Gholamreza Mohammadi; Software: Masoudeh Babakhanian; Validation: Gholamreza Mohammadi and Raheb Ghorbani; Formal analysis, investigation and writing the original draft: Masoudeh Babakhanian; Resources and data curation: Shaqayeq Khosravifar; Review and editing: Masoudeh Babakhanian, Gholamreza Mohammadi and Raheb Ghorbani; Visualization: Raheb Ghorbani.

Conflicts of interest

The authors declared no conflict of interest.

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