

Nontyphoidal Salmonella Outbreaks Associated With Chocolate Consumption

A Systematic Review

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Background: A large, cross-border outbreak of nontyphoidal salmonellosis connected to chocolate product consumption was recently reported. This occurrence motivated us to conduct a comprehensive review of existing literature concerning outbreaks of nontyphoidal salmonellosis associated with chocolate consumption.

Methods: We performed a systematic review following the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines (PROSPERO CRD42022369023) in 3 databases: U.S. National Library of Medicine, Web of Science and Excerpta Medica. Google Scholar and the bibliography of each identified report were also screened. Eligible were articles published after 1970, describing outbreaks of more than 10 patients with a nontyphoidal salmonellosis associated with chocolate consumption.

Results: Twenty-three articles were included, which described 12 outbreaks involving a total of 3266 patients. All outbreaks occurred in high-income countries: 1 was limited to 1 city, 6 involved 1 country and the remaining 5 involved 2 or more countries. Six outbreaks peaked in winter, 3 in autumn, 2 in spring and 1 in summer. Children were mainly affected. No predominant serotype was identified.

Conclusions: Our data documents that chocolate is an optimal medium for the transmission of nontyphoidal salmonellosis. A connected worldwide reporting system including high-income, middle-income and low-income countries is crucial to detect infectious diseases outbreaks in an early phase and avoid their spread.

Key Words: child, chocolate, nontyphoidal salmonella, outbreak

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Domestic and wild animals, including mammals, poultry and reptiles, are the natural hosts of nontyphoidal Salmonella (hereafter referred to as Salmonella, for simplicity), which is one of the most frequently reported causative agents for foodborne outbreaks worldwide.^{1–3} Traditionally, foodborne Salmonella infections have been associated with eggs and poultry.^{1–3} Nowadays many other foods are implicated, such as dried fruits and vegetables, nuts, spices, cereal-based and confectionery products.^{1–4}

The link between chocolate consumption and nontyphoidal salmonellosis has been established since the 1950s, with several documented outbreaks in the literature.¹

Over the past 5 decades, considerable progress has been made in optimizing the food supply chain, including improvements in production and distribution processes. These advances have been mainly developed in high-income countries (HICs). Surveillance systems at national and supranational levels, such as Pulse NET (in the US) and the European Centre for Disease Prevention and Control in Europe, play a crucial role in monitoring the epidemiology of nontyphoidal Salmonella infections. Rapid disease cluster detection, sharing of molecular subtyping data via surveillance reports or online platforms like Enterobase and close collaboration with the Food and Drug Administration in the US and the European Food Safety Authority in Europe are essential in reducing the likelihood of widespread outbreaks. Such structured networks are less common in middle-income countries and low-income countries, although efforts to improve the situation, such as the PulseNet International network, are underway in a subset of countries.

The increasing globalization has amplified the potential for a single outbreak to spread rapidly, thus affecting a very large number of subjects.^{2–4,5}

In spring 2022, a large outbreak of nontyphoidal salmonellosis linked to chocolate products from a factory in Belgium affected more than 450 patients in 16 countries.⁶

The existing literature lacks a comprehensive epidemiologic analysis of the characteristics, patterns and outcomes of outbreaks of nontyphoidal Salmonella infections linked to the consumption of chocolate.

METHODS

Data Search

The protocol for this systematic review has been registered on PROSPERO (CRD42022369023). The principles underlying the UK Economic and Social Research Council guidance on the conduct of narrative synthesis and the 2020 version of the Preferred

Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines were employed.⁷

A literature search with no language restrictions using the terms (“salmonella” OR “salmonellosis”) AND “chocolate” was carried out using the US National Library of Medicine, Excerpta Medica and Web of Science databases in November 2022 and repeated before submission (August 10, 2023). Google Scholar and the bibliography of each identified report were also screened.⁸ Because reports published until 1970 were poorly documented, only reports published thereafter were considered.⁹

Eligibility Criteria, Data Extraction and Quality Assessment

For the final analysis, we retained exclusively outbreaks of more than 10 patients diagnosed with an intestinal Salmonella infection following the consumption of any sort of food containing chocolate (such as confectionery products, cakes, pastries and ice cream). Chocolate-containing food had to have been considered as the source of infection according to one of the following 3 conditions: (1) microbiologic studies revealing the presence of the same Salmonella strain in both fecal specimens and food samples; (2) epidemiologic evidences indicating that all infected individuals consumed the same food; (3) a combination of both microbiologic and epidemiologic evidences.

The following data were extracted from the included papers: timeframe and geographic area of the outbreak, chocolate vehicle, microbiologic studies in fecal samples and/or in the vehicle of infection, number of cases, demographics, hospitalization rate, complications and outcome. If 2 or more reports provided information on the same outbreak, the corresponding data were cumulated. The Newcastle-Ottawa Quality Assessment Scale was employed to assess the risk of bias in the included studies.¹⁰

Two authors (Z.P. and P.B.F.) performed in duplicate the literature search, the selection of eligible reports, data extraction and quality assessment. Disagreements between individual judgements were solved by consensus meetings with a third author (L.K.). The data were input into a predefined worksheet by one author (Z.P.) and the accuracy of the data entry was verified by the second author (P.B.F.).

RESULTS

Search Results and Quality of Studies

The literature search process is summarized in Figure, Supplemental Digital Content 1, <http://links.lww.com/INF/F396>. Twenty-three articles published between 1972 and 2022 were retained for the final analysis: 20 were written in English, 1 in Italian, 1 in Norwegian and 1 in Swedish.^{6,11–32} The mentioned articles describe 12 outbreaks of nontyphoidal salmonellosis linked with chocolate consumption. The risk of bias assessment score could only be calculated for 7 studies (see Table, Supplemental Digital Content 2, <http://links.lww.com/INF/F397>) that were classified as case-control studies. Only 2 studies were of good quality.^{18,25} The remaining studies could not be analyzed because they did not meet the criteria for classification as either case-control or cohort studies.

Findings

The characteristics of the 12 outbreaks, involving a total of 3266 patients, are presented in Table 1. Two outbreaks took place in the 1970s, 3 in the 1980s, 1 in the 1990s, 3 in the decade 2000–2009 and 3 thereafter. Six outbreaks peaked in winter, 3 in autumn, 2 in spring and 1 in summer.

One outbreak was limited to 1 city. Six outbreaks involved 1 country and the remaining 5 outbreaks involved 2 or more countries. On 3 occasions, the outbreak spread across 2 continents.

No predominant Salmonella serotype was identified. Between 1970 and 1989, conventional microbiologic laboratory techniques were applied to identify the responsible strain.^{11–19} Subsequently, pulsed-field gel electrophoresis,^{20,24,27,28} multilocus variable-number tandem repeat analysis²⁷ and whole genome sequencing^{6,30,31} were also employed. The number of cases in each outbreak ranged from 29 to 1111, median 115 cases (interquartile range: 77–385 cases). In 4 studies, the median age of the patients was not provided. Among the remaining studies, the median age ranged from 3 to 15 years. The hospitalization rate ranged between 3% and 41%, median 16% (interquartile range: 10%–29%). No infections involving the bloodstream or other normally sterile sites and fatalities were reported as a direct consequence of the Salmonella infection.

DISCUSSION

Chocolate has peculiar characteristics that make it an optimal medium for the spread of Salmonella. Three factors account for that. First, the low water content and the high fat level of chocolate increase the thermal resistance of Salmonella. Second, higher temperatures during chocolate production, despite eliminating Salmonella, would worsen its taste. Third, Salmonella may persist for more than 1 year in chocolate. These factors, combined with the very small amount of Salmonella required to initiate an infection, illustrate the challenge of preventing outbreaks of this infection.^{33–37}

This systematic literature review describes the characteristics of 12 human nontyphoidal salmonellosis outbreaks associated with chocolate consumption in the past 50 years. Between 1 and 3 outbreaks occurred during each decade. All the reported outbreaks were observed in HICs. This finding may be related, on one hand, to the large availability of industrially produced food and, on the other hand, to the presence of effective outbreak detection and control networks in these countries. It is also conceivable that outbreaks detected in middle-income and low-income countries have not been reported.

Conventional culture-based diagnostic techniques are time-consuming and labor intensive. Today's molecular methods offer rapid and precise detection, enabling early identification and management of foodborne illnesses.³⁸

The results of this review confirm that nontyphoidal salmonellosis occurs most often in children. However, the factors underlying this tendency are elusive. First, it has been suggested but never proven that children might be more susceptible to intestinal infections because the immune system is still developing. Second, person-to-person transmission of Salmonella might likely be more common in children due to behaviors that increase germ exposure (such as putting hands contaminated with bacteria into mouth and having less developed hygiene practices). Additionally, the allure of chocolate in children could also play a contributing factor.^{1–3,32–34,39}

Nontyphoidal salmonellosis outbreaks typically occur during warm season. The majority of Salmonella outbreaks temporally associated with chocolate consumption were observed in the cold season. We speculate that very popular seasonal products such as chocolate Santa Clauses and chocolate Easter bunnies might account for this observation.

No cases of death were reported, likely because nontyphoidal Salmonella disease is self-limiting and does not require treatment other than hydration. Furthermore, extraintestinal complications are uncommon in immunocompetent individuals living in HICs.⁴⁰

This study has limitations. An outbreak is a sudden increase of infections linked to a common source. It is important to note that the definition of an outbreak can vary depending on specific jurisdictions and circumstances. Local authorities often have own guidelines and protocols for identifying an outbreak.

These assumptions likely underlie the high heterogeneity and variability observed in the outbreaks included in this analysis. It is tempting to speculate that some reports may have predominantly focused on documenting severe cases rather than mild ones. On the other hand, some studies may have included only primary cases while other studies did not differentiate between primary and secondary cases. Since the serotypes identified in sporadic cases do not differ from those identified in outbreaks, certain small outbreaks may go undiagnosed. The age of patients was not homogeneously provided in all included studies. A further limitation is that only a minority of reports were of good quality.

CONCLUSIONS

This systematic review of outbreaks of nontyphoidal salmonellosis induced by chocolate consumption highlights a recurring pattern of outbreaks occurring every decade over the past 50 years. The outbreaks primarily occurred in HICs. Children were most affected. Despite the high number of cases, no fatalities were reported, indicating a mild disease severity.

Surveillance reporting systems have a pivotal role in detecting foodborne outbreaks. In particular, prompt timing and coordination with stakeholders at an international level is crucial. The example from the 2022 nontyphoidal Salmonella outbreak is impressive: on March 27, 2022 the United Kingdom notified a cluster of cases with Salmonella typhimurium infection with an unidentified source. The outbreak was quickly linked by investigators to a chocolate firm in Belgium, and a global product recall was initiated on April 10, 2022.

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