



Bovine Brucellosis: A Zoonotic Disease of Public Health and Economic Concern in Ethiopia

Bekiyad Shasho Daro^{*1}, Yared Adisu Deneke², and Jiregna Gari Negasa³

^{*1}Department of Veterinary Epidemiology, College of Veterinary Medicine, Haramaya University, P.O. Box-138, Dire Dawa, Ethiopia

²Department of Veterinary Public Health, College of Veterinary Medicine, Haramaya University, P.O. Box-138, Dire Dawa, Ethiopia

³Department of Veterinary Microbiology, College of Veterinary Medicine, Ambo University, P.O. Box-19, Ambo, Ethiopia

***Corresponding Author:** Bekiyad Shasho Daro, Department of Veterinary Epidemiology, College of Veterinary Medicine, Haramaya University, P.O. Box-138, Dire Dawa, Ethiopia. ORCID iD: <http://orcid.org/0000-0001-5060-2657>

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Abstract

Bovine brucellosis is a bacterial disease that affects cattle and other animals; the main species of bacteria causing this disease are *Brucella abortus* and *Brucella melitensis*. The objective of this appraisal is to articulate the epidemiology, molecular characteristics, and socio-economic importance of bovine brucellosis. It is one of the most widespread zoonoses transmitted by animals which can bring about a serious public health and economic consequences acting as a constraint to livestock production in low-income countries like Ethiopia. The global and regional distribution of bovine brucellosis varies depending on the prevalence of infection in animals, the control measures implemented, the surveillance systems, the environmental factors and the human behavior. Consumption of raw dairy products is the most common way of the disease transmission. Direct contact, venereal transmission, breast-feeding, and sexual contact are other means of transmission. Previous studies have suggested that the prevalence of the disease in exotic and cross-bred, older animals, female cattle, and cattle that have aborted previously, as compared to indigenous animals, male cattle, and cows that have no abortion history respectively. Signs of the disease vary depending on stage of infection, age and sex of the animal, and the presence of other diseases. Diagnosis can be done by different methods like bacteriological culture, molecular techniques, and serological tests. There are many different species and strains, which can be distinguished by their genetic characteristics. Even if diseased, the treatment is not likely to be recommended; because, it is ineffective, costly, and has a risk of creating antibiotic-resistant strains of *Brucella abortus*. That is why the prevention and control are based on a combination of measures, such as vaccination, identification, test-and-slaughter, quarantine, biosecurity, and public education. Since there is lack of policies and programs for bovine brucellosis control and eradication in Ethiopia, further researches have to be conducted in order to address these challenges.

Keywords: Bovine Brucellosis, Economic Impact, Public Health Importance, Risk Factors

Introduction

Bovine brucellosis is a bacterial zoonotic disease that affects cattle and humans, causing abortion, infertility, reduced milk production, and other economic losses in livestock, and fever, headache, joint pain, and other chronic symptoms in humans [1]. The disease is caused by *Brucella abortus*, and occasionally by *B. melitensis*, which are transmitted through contact with infected animals or their products, such as milk, meat, or placenta. Bovine brucellosis is considered the world's most common bacterial zoonosis and one of the top five priority zoonotic diseases in Ethiopia [2].

Ethiopia has one of the largest cattle populations in Africa, with an estimated 60.39 million heads in 2018 [3]. However, the productivity and profitability of the livestock sector are hampered by various factors, including infectious diseases such as bovine brucellosis. Previous studies have reported variable prevalence

rates of bovine brucellosis in different regions of Ethiopia, ranging from 0.06% to 18.9%. The disease is endemic and widely distributed in the country, especially in pastoral and agro-pastoral areas where cattle are kept together with small ruminants or camels that may harbor *B. melitensis*. Moreover, the lack of adequate diagnostic facilities, vaccination programs, public awareness, and control measures contribute to the persistence and spread of the disease [4].

Livestock production is one of the main sources of income and livelihood for the people in the zone [5]. However, there is limited information on the status of bovine brucellosis and its public health significance in Ethiopia. The findings of previous studies may provide baseline data for designing and implementing appropriate interventions to prevent and control bovine brucellosis and protect animal and human health [6,7].

A bacterial disease caused by *Brucella abortus* that mainly affects cattle, water buffalo, and bison. It can cause abortion, birth of stillborn or weak calves, retained placenta, and reduced milk production in infected animals. It can also be transmitted to humans through direct contact with infected animals or consumption of contaminated animal products. The frequency of individuals in a population who test positive for a specific disease based on serology (blood serum) specimens [8]. It often reflects the level of exposure or immunity to a pathogen in a population. The determination of the genetic characteristics of an organism or a pathogen using molecular techniques such as DNA sequencing, PCR, or hybridization. It can help identify the species, strain, or genotype of the organism or pathogen and reveal its phylogenetic relationships, genetic diversity, or evolutionary history. Something that increases the risk or susceptibility of an individual or a population to a disease or an adverse outcome. Risk factors can be biological, environmental, behavioral, or social. For example, age, breed, sex, herd size, management practices, and vaccination status are some of the risk factors for bovine brucellosis [9].

The bacteria can be transmitted to humans through contact with infected animals or their secretions, or through consumption of contaminated, unpasteurized dairy products. Human brucellosis can cause fever, fatigue, joint pain, sweating, enlarged liver and spleen, and complications in pregnant women [2]. According to a study by, globally, 500000 cases of human brucellosis are reported annually [1]. The prevalence of human brucellosis differs between areas and has been reported to vary with standards of personal and environmental hygiene, animal husbandry practices, and species of the causative agent and local methods of food processing. In Ethiopia, human brucellosis is considered an underreported and neglected disease due to lack of awareness, diagnostic facilities, and effective control measures [10].

Brucellosis is a bacterial disease that affects various animal species, especially cattle, and can cause serious economic losses and public health risks. In Ethiopia, bovine brucellosis is endemic and poses a threat to the livestock sector and the rural livelihoods. However, there is limited information on the Seroprevalence, molecular characterization and risk factors for bovine brucellosis in different regions of the country. Therefore, the objective of this paper is:

- To review on the epidemiology, molecular characteristics and socio-economic impact of bovine brucellosis and
- To assess the existing policies and programs for control and eradication of bovine brucellosis.

Epidemiology of Bovine Brucellosis

Bovine brucellosis is a bacterial disease that affects cattle and other animals. It can also be transmitted to humans through direct contact with infected animals or their products, such as milk and cheese. Brucellosis is found globally and is a public health concern in many countries. The main species of bacteria causing bovine brucellosis are *Brucella abortus* and *Brucella melitensis* [11].

The Global and Regional Distribution and its Prevalence

Bovine brucellosis is a bacterial disease caused by *Brucella abortus*, which mainly infects cattle and causes abortions or reproductive failure. The disease is one of the most widespread zoonoses transmitted by animals and has serious public health and economic consequences. The global and regional distribution of bovine brucellosis varies depending on the prevalence of infection in animals, the control measures implemented, the surveillance systems, the environmental factors and the human behavior [12]. According to the World Organization for Animal Health (WOAH), bovine brucellosis is found in all continents except Antarctica, but it is more prevalent in some regions than others. The highest incidence is observed in the Middle East, the Mediterranean region, sub-Saharan Africa, China, India, Peru, and Mexico. Currently, countries in central and southwest Asia are seeing the greatest increase in cases [13].

In Africa, bovine brucellosis is endemic in most countries and poses a major threat to livestock production and public health [14]. The disease is often underreported due to the lack of diagnostic facilities, awareness and political will. In Ethiopia, bovine brucellosis has been reported in different regions and agro-ecological zones, with Seroprevalence ranging from 0.8% to 8.8%. The risk factors for bovine brucellosis in Ethiopia include herd size, breed, age, sex, abortion history, grazing system, contact with wildlife and consumption of raw milk [15].

The Main Transmission Routes and Risk Factors

The most common way to be infected is by eating or drinking unpasteurized/raw dairy products. When sheep, goats, cows, or camels are infected, their milk becomes contaminated with the bacteria. If the milk from infected animals is not pasteurized, the infection will be transmitted to people who consume the milk and/or cheese products [16]. Breathing in the bacteria that causes brucellosis may also lead to infection. This risk is generally greater for people in laboratories that work with the bacteria. In addition, slaughterhouse and meat-packing employees have also been known to be exposed to the bacteria and ultimately become infected [17].

Bacteria can also enter wounds in the skin/mucous membranes through contact with infected animals. This poses a problem for workers who have close contact with animals or animal excretions (newborn animals, fetuses, and excretions that may result from birth). Such workers may include, slaughterhouse workers, meat-packing plant employees, veterinarians [12]. People who hunt animals may also be at risk. When they are in contact with infected animals, exposure to the bacteria may occur through skin wounds, accidentally ingesting undercooked meat, or inhaling the bacteria while dressing their game. Commonly infected animals include: bison, elk, caribou, moose and wild hogs [18].

Venereal transmission by infected bulls to susceptible cows appears to be rare. Transmission may occur via artificial insemination when *Brucella*-contaminated semen is deposited in the uterus; however, apparently, not when deposited in the midcervix. Person-to-person spread of brucellosis is extremely rare. Infect-

ed mothers who are breast-feeding may transmit the infection to their infants. Sexual transmission has been rarely reported. While uncommon, transmission may also occur via tissue transplantation or blood transfusions [19].

The prevalence of the disease in exotic and cross-bred animals is higher than in indigenous cattle. The prevalence of the disease increases with age as older animals have more exposure to the infection. Female cattle are more susceptible to the infection than the male ones due to reproductive hormones and abortion events. Cattle that have aborted previously are more likely to be infected than those that have not [20]. Larger farms with poor hygiene and biosecurity practices are more prone to the infection than smaller farms with better management. Cattle that are frequently moved or mixed with other herds are more likely to be exposed to the infection than those that are isolated [21].

Clinical Signs and Diagnosis

The clinical signs of bovine brucellosis vary depending on the stage of infection, the age and sex of the animal, and the presence of other diseases. The most common signs are abortion (usually in the last trimester of pregnancy), retained placenta, metritis, orchitis, epididymitis, and arthritis. However, some infected animals may show no signs or only mild signs of infection, making the diagnosis difficult [22].

The diagnosis of bovine brucellosis can be done by using different methods, such as bacteriological culture, molecular techniques, and serological tests. Bacteriological culture is the gold standard method for confirming the presence of *Brucella abortus* in samples such as milk, blood, aborted fetuses, or tissues. However, this method is time-consuming, labor-intensive, and requires biosafety level-3 facilities [23]. Molecular techniques, such as polymerase chain reaction (PCR), can detect the DNA of *Brucella abortus* in samples with high sensitivity and specificity. However, these techniques are also expensive and require specialized equipment and trained personnel. Serological tests are the most widely used methods for screening and surveillance of bovine brucellosis. These tests detect antibodies against *Brucella abortus* in serum samples using different antigens and formats [9]. Some of the commonly used serological tests are the Rose Bengal Plate Test (RBPT), the complement fixation test (CFT), the enzyme-linked immunosorbent assay (ELISA), and the fluorescence polarization assay (FPA). However, these tests have some limitations, such as cross-reactivity with other bacteria, lack of standardization, and interference by vaccination [24].

Treatment and Prevention

The treatment of bovine brucellosis is not recommended because it is ineffective, costly, and poses a risk of creating antibiotic-resistant strains of *Brucella abortus*. Moreover, treated animals may still remain as carriers and shedders of the bacteria [25]. Therefore, the prevention and control of bovine brucellosis are based on a combination of measures, such as vaccination, identification, testing, culling, quarantine, biosecurity, and public education. Vaccination is one of the most important tools for reducing the prevalence and incidence of bovine brucellosis in endemic areas. The most commonly used vaccines are live at-

tenuated strains of *Brucella abortus*, such as strain 19 and RB51. These vaccines induce protective immunity in cattle by stimulating both humoral and cellular responses. However, these vaccines also have some drawbacks, such as causing abortion in pregnant animals, inducing antibodies that interfere with serological diagnosis, and being potentially infectious to humans [6].

Identification and testing are essential for detecting infected animals and tracing their contacts. Culling is the removal of infected animals from the herd to prevent further transmission and contamination. Quarantine is the isolation of new or suspect animals until they are proven to be free of infection [26]. Biosecurity is the implementation of hygienic practices to prevent the introduction or spread of infection within or between herds. Public education is the dissemination of information and awareness about bovine brucellosis to farmers, veterinarians, consumers, and other stakeholders to promote compliance with prevention and control measures [27].

Molecular Characterization of Bovine Brucellosis

Genetic Diversity and Evolution of *Brucella* Species and Strains
Brucella has many different species and strains, which can be distinguished by their genetic characteristics. Some of these genetic characteristics are single nucleotide polymorphisms (SNPs), which are variations in the DNA sequence of a gene. SNPs can be used to trace the evolutionary history and geographic origin of *Brucella* species and strains [28]. One of the main methods to study the genetic diversity and evolution of *Brucella* is whole genome sequencing (WGS), which is the process of determining the complete DNA sequence of an organism's genome. WGS can reveal the SNPs and other genetic features of *Brucella*, such as genes, plasmids, insertion sequences, and genomic islands. WGS can also help to identify new or atypical *Brucella* strains that do not fit into the typical classification based on phenotypic or serological tests [29].

Some studies have used WGS to analyze the global evolution and phylogeography of *Brucella melitensis*, the most common cause of human brucellosis. For example, performed an analysis of the SNP of the complete genomes of 98 *B. melitensis* strains isolated in different regions of the world [30]. They identified five main genotypes and 13 subgenotypes of *B. melitensis*, and determined their regions of origin and pathways of distribution. They also found that some subgenotypes were associated with specific hosts, such as sheep, goats, or camels [31].

Another example is, who focused on the Brucellaceae family, which includes *Brucella* and other related genera, such as *Ochrobactrum* and *Pseudochrobactrum* [32]. They compared 145 Brucellaceae genomes with over 40 others from the wider order Rhizobiales to resolve phylogenetic ambiguities. They showed that *Brucella* is a monophyletic group within the *Ochrobactrum* diversity, and that *Ochrobactrum* itself is polyphyletic, splitting into two clades. They also recognized substantial undiagnosed diversity in *Ochrobactrum* spp. and *Pseudochrobactrum* spp. These studies demonstrate how WGS can provide valuable insights into the genetic diversity and evolution of *Brucella* species and strains, as well as their relationships with other mem-

bers of the Brucellaceae family. This knowledge can help to improve the diagnosis, prevention, and control of brucellosis in animals and humans [29].

Advantages and Limitations of Different Molecular Techniques for Typing and Identifying Brucella Isolates

16S rRNA gene sequencing is based on the amplification and sequencing of a highly conserved region of the bacterial 16S rRNA gene. It can provide accurate identification of Brucella species and subspecies, as well as phylogenetic analysis. However, it is relatively expensive, time-consuming, and requires specialized equipment and expertise [33]. Real-time PCR can provide rapid and sensitive identification of Brucella species and subspecies, as well as quantification of bacterial load. However, it is also costly, requires specific primers and probes, and may be affected by PCR inhibitors or cross-contamination [34].

Enterobacterial repetitive intergenic consensus (ERIC) PCR can provide high-resolution typing of Brucella isolates, as well as epidemiological analysis and outbreak investigation. However, it is technically demanding, prone to variability and reproducibility issues, and may not discriminate between closely related strains [35]. The amplification and analysis of multiple loci containing variable-number tandem-repeats (VNTRs), which are short DNA sequences repeated in tandem, can provide high discriminatory power, stability, and evolutionary insight for Brucella typing. However, it is labor-intensive, requires standardized protocols and databases, and may not reflect the true phylogeny of Brucella [36].

The Current Knowledge and Challenges of Molecular Epidemiology of Bovine Brucellosis

Bovine brucellosis is a zoonotic infection caused by bacteria of the genus Brucella, mainly Brucella abortus, that affects cattle and other domestic and wild animals. It is a major public health problem and a cause of economic losses in many countries, especially in developing regions [20]. Molecular epidemiology of bovine brucellosis is the study of the genetic diversity, evolution, transmission, and population structure of Brucella strains isolated from cattle and other hosts. It can provide insights into the origin, spread, and control of the disease [37].

Molecular typing methods, such as multilocus variable-number tandem-repeat analysis (MLVA), multilocus sequence analysis (MLSA), single nucleotide polymorphism (SNP) typing, and whole genome sequencing (WGS), have been used to characterize Brucella strains from different geographic regions, animal species, and sources of infection [30]. These methods have revealed a high genetic diversity and variability of Brucella strains within and between countries, reflecting the complex epidemiology and ecology of the disease [38].

Molecular epidemiology has also identified the emergence and dissemination of new Brucella variants, such as biovars 5b and 6 in Europe, biovar 3b in Africa, and biovar 1 in Asia, that pose new challenges for diagnosis and control; and has also contributed to the understanding of the transmission dynamics and risk

factors of bovine brucellosis, such as animal movement, trade, husbandry practices, wildlife reservoirs, and human activities [39,18].

However, molecular epidemiology of bovine brucellosis also faces some limitations and difficulties, such as the lack of standardized and harmonized typing methods and databases, the limited availability and accessibility of molecular data from endemic areas, the need for more representative and comprehensive sampling strategies, and the integration of molecular data with epidemiological, clinical, and environmental information [40]. Therefore, molecular epidemiology of bovine brucellosis is a valuable tool for improving the surveillance, prevention, and control of the disease, but it requires further development and application in different settings and contexts.

Bovine Brucellosis in Ethiopian Perspective

Bovine brucellosis a major public health problem and a constraint to livestock production in Ethiopia. The epidemiological situation and trends of bovine brucellosis in Ethiopia are influenced by various factors, such as the management system, the breed of cattle, the diagnostic methods, the geographical location, and the environmental conditions. According to the prevalence of bovine brucellosis in Ethiopia ranges from 1.2% to 22.5% at the individual level, and from 3.3% to 68.6% at the herd level [6]. The disease is more common in pastoral and mixed farming systems, where cattle are kept in close contact with humans and other animals. The exotic and cross-bred cattle are more susceptible to brucellosis than the local breeds, because they have lower immunity and higher reproductive performance [27]. The disease causes abortion, infertility, reduced milk production, and decreased weight gain in cattle, resulting in significant economic losses [41].

The diagnosis of bovine brucellosis is challenging, because the clinical signs are not specific and the laboratory tests are not standardized or widely available. The most commonly used diagnostic methods in Ethiopia are the Rose Bengal Plate Test (RBPT) and the Complement Fixation Test (CFT), which have different sensitivities and specificities [42]. Molecular techniques, such as polymerase chain reaction (PCR), are more accurate and reliable, but they require sophisticated equipment and trained personnel [24]. Therefore, there is a need for rapid, simple, and cost-effective diagnostic tools that can be applied in field settings.

The distribution of bovine brucellosis in Ethiopia varies according to the geographical regions and climatic conditions. The disease is more prevalent in lowland areas than in highland areas, because of the higher temperature and humidity that favor the survival of Brucella organisms [43]. The disease is also influenced by seasonal factors, such as rainfall, drought, and migration of animals [44]. The risk factors for bovine brucellosis include the herd size, the sex and age of cattle, the history of abortion, the contact with wildlife, and the consumption of raw milk or milk products [45].

The public health implications of bovine brucellosis in Ethiopia

are serious, because humans can acquire the infection through direct or indirect contact with infected cattle or their products. The symptoms of human brucellosis include fever, headache, joint pain, fatigue, and enlarged spleen or liver [18]. The disease can also cause chronic complications, such as endocarditis, arthritis, osteomyelitis, meningitis, or orchitis [46]. The treatment of human brucellosis requires prolonged antibiotic therapy, which can be expensive and have adverse effects. The prevention and control of human brucellosis depends on the control of bovine brucellosis [47]. Therefore, based on this analysis, it can be concluded that there is a need for further research on the seroprevalence, molecular characterization, and risk factors for bovine brucellosis in different regions of Ethiopia.

The Socio-Economic Impact and Public Health Implications

The socio-economic impact and public health implications of bovine brucellosis in Ethiopia can be evaluated based on the prevalence and distribution of the disease in different regions, agro-ecological zones, and livestock production systems [12]. This can help to estimate the burden of the disease on the livestock sector and identify the high-risk areas for transmission and control interventions. The molecular characterization and genetic diversity of *Brucella* strains can help to understand the epidemiology, evolution, and transmission dynamics of the disease, as well as to develop appropriate diagnostic tools and vaccines. The risk factors associated with the disease in cattle and humans can help to identify the main sources and routes of exposure, as well as the potential risk groups and behaviors that facilitate the spread of the disease [48].

The clinical manifestations, diagnosis, treatment, and prevention of bovine brucellosis in cattle and humans can help to assess the health impact of the disease on both animal and human populations, as well as to evaluate the effectiveness and feasibility of existing or novel control strategies [49]. The economic losses and social consequences can also help to quantify the direct and indirect costs of the disease on the livestock industry, human health, food security, and livelihoods of affected communities [34].

The Existing Policies and Programs for Control and Eradication
The goal of bovine brucellosis control and eradication is to prevent or eliminate the disease from domestic cattle and other susceptible animals. Different countries have different policies and programs for achieving this goal, depending on the prevalence, distribution, and impact of the disease in their regions. Some of the common strategies include, surveillance and reporting, vaccination, diagnostic testing and laboratory services, test-and-slaughter policy, education and awareness, and the socio-cultural and economic factors [41].

Surveillance and reporting involve testing animals for brucellosis infection, tracing the source and contacts of infected animals, reporting the results to the authorities, and maintaining a database of the disease situation [50]. Surveillance and reporting are essential tools for monitoring the progress of control and eradication efforts, identifying high-risk areas, and implementing appropriate interventions in *Brucella* infection [51].

Vaccination involves immunizing animals with a vaccine that protects them from brucellosis infection or reduces the severity of the disease. Vaccination can be compulsory or voluntary, depending on the level of risk and the availability of resources which can reduce the incidence and transmission of bovine brucellosis, but it cannot eliminate the disease completely [52]. Vaccinated animals should be identified with a tag or a mark to distinguish them from unvaccinated ones.

Diagnostic testing is necessary for confirming the diagnosis of suspected cases, screening animals for trade or movement, evaluating the effectiveness of vaccination programs, and verifying the absence of infection in eradication areas [53]. Test-and-slaughter policy involves culling animals that test positive for brucellosis infection, as well as their contacts or offspring. It is aimed at removing the source of infection from the population and preventing further spread of the disease [54]. Test-and-slaughter policy should be accompanied by compensation for the owners of culled animals, as well as disinfection of contaminated premises.

Education and awareness involve informing and educating farmers, consumers, health workers, and other stakeholders about bovine brucellosis, its causes, symptoms, transmission, prevention, control, and eradication. This can increase the knowledge and understanding of the disease, improve the compliance and cooperation with control and eradication measures, reduce the risk behaviors and practices that facilitate the spread of the disease, and promote the adoption of good animal husbandry and hygiene practices [55].

Socio-cultural and economic factors involve considering the social, cultural, economic, and environmental factors that influence the behavior and practices of livestock owners and consumers regarding bovine brucellosis. These factors can affect the acceptance, feasibility, sustainability, and effectiveness of control and eradication programs [2]. Low awareness and knowledge of the disease among farmers and consumers, lack of access to diagnostic tests and laboratories, high cost and low availability of vaccines; inadequate compensation for culled animals, poor enforcement of quarantine and movement restrictions; traditional customs such as raw milk consumption or animal sharing, poverty and food insecurity that limit the adoption of preventive measures, plus weak coordination and collaboration among different sectors involved in animal health are among some of the factors that may hinder bovine brucellosis control and eradication in Ethiopia [4].

Conclusion and Recommendations

Bovine brucellosis is a serious zoonotic disease that affects cattle and other animals, as well as humans. It has a global and regional distribution, and it can cause significant economic losses and public health risks in Ethiopia. The disease is transmitted by various routes, such as consumption of unpasteurized dairy products, direct contact, venereal transmission, breast-feeding, and sexual contact. The diagnosis can be done by different methods, such as bacteriological culture, molecular techniques, and serological tests. The treatment is not recommended due to its ineff-

fectiveness, costliness, and risk of creating antibiotic-resistant strains of *Brucella abortus*. Therefore, prevention and control measures are essential to eradicate the disease from Ethiopia. Based on the above conclusion, we can recommend the following actions to address the problem:

- Strengthening the surveillance systems for bovine brucellosis at national and local levels.
- Implementing effective vaccination programs for cattle and other animals at risk.
- Improving the identification and testing of infected animals through test-and-slaughter schemes.
- Enhancing the quarantine and biosecurity measures to prevent the spread of the disease within and between farms.
- Raising public awareness about the signs, symptoms, transmission modes, prevention strategies, and treatment options of bovine brucellosis.
- Developing policies and programs for bovine brucellosis control and eradication in Ethiopia.

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