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## Review Article

### **A COMPREHENSIVE REVIEW OF COVID-19: EPIDEMIOLOGY, CLINICAL FEATURES, AND MANAGEMENT STRATEGIES**

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#### **ABSTRACT**

Coronavirus disease 2019 (COVID-19) is a highly transmissible and pathogenic viral infection caused by the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2). Since its emergence in December 2019, the disease has rapidly evolved into a global health crisis, leading to significant morbidity, mortality, and socioeconomic disruptions worldwide. The virus primarily spreads through respiratory droplets, aerosols, and close human contact, contributing to its widespread transmission. COVID-19 presents with a diverse spectrum of clinical manifestations, ranging from asymptomatic or mild cases to severe pneumonia, acute respiratory distress syndrome (ARDS), multi-organ failure, and death, particularly in vulnerable populations such as the elderly and individuals with underlying comorbidities.

This review provides a comprehensive analysis of the epidemiology, clinical features, diagnostic approaches, and management strategies for COVID-19. A systematic assessment of published literature highlights the critical role of vaccination in reducing disease severity and mortality. Various vaccine platforms, including mRNA, viral vector, protein subunit, and inactivated virus vaccines, have significantly altered the course of the pandemic, although concerns regarding waning immunity and emerging variants persist. In addition to vaccines, antiviral therapies such as remdesivir, molnupiravir, and nirmatrelvir-ritonavir, along with

immunomodulatory treatments including corticosteroids and monoclonal antibodies, have been instrumental in improving patient outcomes.

Non-pharmaceutical interventions (NPIs), such as social distancing, mask mandates, and lockdown measures, have played a crucial role in mitigating viral transmission, especially in the early phases of the pandemic. However, the socioeconomic and psychological consequences of these measures highlight the need for balanced public health strategies. The emergence of SARS-CoV-2 variants with enhanced transmissibility and immune escape potential continues to pose challenges for disease control, necessitating continuous genomic surveillance and adaptive vaccine strategies.

This review aims to provide a holistic perspective on the lessons learned from the COVID-19 pandemic, emphasizing the importance of global collaboration, evidence-based policymaking, and sustained investment in medical research. Strengthening healthcare infrastructure, enhancing pandemic preparedness, and fostering international cooperation are essential to mitigating the impact of COVID-19 and addressing future pandemics effectively.

## **KEYWORDS**

Antiviral therapies, COVID-19, Diagnostic modalities, Epidemiology, Management strategies, Pandemic, SARS-CoV-2, Vaccine efficacy, Variants, Viral transmission

## **INTRODUCTION**

### **HISTORICAL CONTEXT OF PANDEMICS**

Throughout history, pandemics have had a profound impact on human civilization, shaping societies, economies, and healthcare systems. Notable pandemics include the Spanish flu of 1918, which resulted in an estimated 50 million deaths worldwide, demonstrating the devastating potential of viral outbreaks. More recently, the H1N1 influenza pandemic in 2009, although less severe, highlighted the importance of vaccine development and global coordination in managing infectious diseases. Additionally, the emergence of coronaviruses, including the Severe Acute Respiratory Syndrome (SARS) outbreak in 2002 and the Middle East Respiratory Syndrome (MERS) in 2012, underscored the zoonotic origins of novel pathogens and the challenges in containing respiratory viruses. These historical pandemics provided valuable lessons in surveillance, vaccine development, and public health preparedness, many of which were revisited during the COVID-19 pandemic. However, despite past experiences, the unprecedented scale and rapid global spread of COVID-19 presented unique challenges, requiring swift adaptations in healthcare and policy responses.<sup>1,2</sup>

### **Early Response and Initial Challenges**

The initial global response to COVID-19 varied significantly across nations, with many governments struggling to assess the pandemic's severity and implement appropriate containment measures. Delays in recognizing its pandemic potential, compounded by limited testing capacity and insufficient data, hindered early mitigation efforts. Healthcare systems worldwide faced shortages of personal protective equipment (PPE), hospital beds, ventilators, and critical medical supplies, placing immense strain on frontline workers. Furthermore, the absence of clear treatment guidelines and the reliance on experimental therapies created uncertainty in clinical management. In an attempt to control the virus, governments implemented lockdowns, travel restrictions, and social distancing measures, but these efforts were often inconsistent due to political, economic, and societal pressures. Countries with strong public health infrastructure and proactive containment measures were more successful in controlling early outbreaks, while others experienced uncontrolled viral spread, overwhelming healthcare facilities and leading to high mortality rates.<sup>3,4</sup>

### **Socioeconomic Impact of COVID-19**

The COVID-19 pandemic triggered an unprecedented global economic downturn, disrupting industries, labor markets, and international trade. Business closures, reduced consumer spending, and strict lockdowns led to widespread job losses, with unemployment rates soaring in many countries. Small and medium-sized enterprises (SMEs), which form the backbone of most economies, were particularly vulnerable, struggling to survive prolonged shutdowns and supply chain disruptions. The crisis also exposed weaknesses in global healthcare systems, particularly in low- and middle-income countries, where inadequate resources and underfunded medical infrastructure resulted in disproportionately high case fatalities. Education systems faced disruptions as schools and universities transitioned to remote learning, exacerbating educational inequalities and mental health concerns among students. Social isolation, economic hardship, and uncertainty about the future led to a surge in mental health disorders, including anxiety, depression, and post-traumatic stress disorder (PTSD). Additionally, the pandemic disproportionately affected marginalized communities, widening existing socioeconomic disparities and emphasizing the need for more resilient and equitable public health systems.<sup>5</sup>

### **Political and Healthcare System Responses**

Governments worldwide adopted varying approaches to managing COVID-19, ranging from strict lockdowns and aggressive testing strategies to more relaxed policies aimed at achieving herd immunity. Countries such as New Zealand and South Korea successfully implemented early containment strategies, including mass testing, contact tracing, and quarantine measures, which significantly reduced transmission rates. In contrast, other nations struggled with inconsistent policies, delayed responses, and public resistance to health mandates. The effectiveness of each country's response depended on factors such as healthcare infrastructure, political leadership, public trust in health authorities, and vaccine availability. The rapid development and deployment

of COVID-19 vaccines represented a turning point in the fight against the pandemic, but vaccine distribution disparities between high-income and low-income countries highlighted global health inequities. Misinformation and vaccine hesitancy further complicated efforts to achieve widespread immunization. Despite these challenges, the pandemic accelerated advancements in telemedicine, digital health technologies, and global health cooperation, reinforcing the importance of evidence-based policymaking and international collaboration in preparing for future pandemics.<sup>6</sup>

## **Epidemiology**

### **Origin and Evolution of SARS-CoV-2**

SARS-CoV-2, the causative agent of COVID-19, is a zoonotic virus with a genetic structure closely related to bat coronaviruses, particularly those found in *Rhinolophus* bat species. The virus shares significant homology with SARS-CoV, the pathogen responsible for the 2002 SARS outbreak, suggesting a similar evolutionary trajectory. Although the precise origin of SARS-CoV-2 remains a topic of scientific investigation, evidence indicates that it likely emerged from an animal reservoir, with bats serving as the primary source. The possibility of transmission through an intermediate host, such as pangolins or other wildlife species, has been suggested, but conclusive proof is lacking. The zoonotic spillover event that enabled the virus to cross the species barrier and infect humans highlights the complex interplay between wildlife ecosystems, human activities, and emerging infectious diseases. Understanding the origins of SARS-CoV-2 is critical for preventing future zoonotic outbreaks and improving surveillance at the human-animal interface.<sup>7,8</sup>

### **Transmission Dynamics**

SARS-CoV-2 is primarily transmitted through respiratory droplets released during coughing, sneezing, speaking, or breathing. Aerosolized particles, particularly in poorly ventilated or enclosed spaces, further enhance the risk of transmission, making super-spreader events a significant driver of the pandemic. Contaminated surfaces, although less common as a primary mode of transmission, also played a role in viral spread, emphasizing the importance of hand hygiene and environmental disinfection. The high transmissibility of SARS-CoV-2 is attributed to its ability to bind efficiently to the angiotensin-converting enzyme 2 (ACE2) receptor, which is abundantly expressed in the human respiratory tract. Closed environments, such as healthcare settings, public transport, and crowded indoor venues, amplified transmission rates, particularly during the early phases of the pandemic when public health measures were inconsistently implemented.<sup>9,10</sup>

### **Role of Asymptomatic Carriers in Disease Spread**

One of the most challenging aspects of containing SARS-CoV-2 has been its ability to spread through asymptomatic and pre-symptomatic individuals. Asymptomatic carriers, who show no clinical signs of

infection, and pre-symptomatic individuals, who are infectious before the onset of symptoms, contributed significantly to undetected transmission chains. Studies estimate that a substantial proportion of infections were transmitted by individuals who were unaware of their infectious status, complicating contact tracing and quarantine efforts. This silent transmission underscored the importance of widespread testing, universal masking, and proactive public health strategies to identify and isolate carriers, regardless of symptom presentation.<sup>11</sup>

### **Seasonal and Geographical Variations**

The transmission of COVID-19 exhibited seasonal and geographical trends influenced by various factors. Case surges were often observed during colder months, particularly in regions with temperate climates. Lower temperatures and reduced humidity are believed to facilitate viral stability and transmission, while increased indoor gatherings during winter further exacerbate spread. Geographical variations in case severity and outcomes were shaped by factors such as population density, urbanization, and access to healthcare services. Resource-limited settings faced disproportionately higher mortality rates due to inadequate medical infrastructure and delayed access to diagnostics and therapeutics. Additionally, differences in cultural practices, public health policies, and levels of community compliance with interventions contributed to regional disparities in case dynamics. Understanding these variations is crucial for tailoring context-specific responses to future outbreaks and pandemics.<sup>12</sup>

### **Clinical Features**

#### **Symptomatology and Disease Progression**

COVID-19 presents with a wide spectrum of clinical manifestations, ranging from asymptomatic or mild cases to severe respiratory distress and multi-organ failure. The initial symptoms typically appear within 2 to 14 days post-exposure and often include fever, dry cough, sore throat, and fatigue. Many patients also experience anosmia (loss of smell) and ageusia (loss of taste), which have emerged as distinguishing features of COVID-19. In some cases, gastrointestinal disturbances such as nausea, diarrhea, and abdominal discomfort may occur, particularly in pediatric and immunocompromised populations. While the majority of individuals recover with mild to moderate symptoms, severe cases can progress to pneumonia, acute respiratory distress syndrome (ARDS), cytokine storm, and multi-organ dysfunction, necessitating intensive care and ventilatory support. The unpredictable nature of disease progression underscores the importance of early diagnosis and timely medical intervention.<sup>13</sup>

#### **Complications and Long-Term Effects (Long COVID)**

A significant proportion of COVID-19 survivors experience lingering symptoms, a condition commonly referred to as “Long COVID” or post-acute sequelae of SARS-CoV-2 infection (PASC). These persistent

symptoms can last for weeks or months beyond the acute phase of infection and may include chronic fatigue, brain fog, difficulty concentrating, shortness of breath, muscle weakness, and sleep disturbances. Additionally, some individuals report cardiovascular complications such as myocarditis, arrhythmias, and increased risk of thrombotic events, raising concerns about long-term health consequences. Neurological symptoms, including headaches, dizziness, and mood disorders, have also been documented, suggesting potential neuroinflammatory and autoimmune mechanisms triggered by viral infection. The burden of Long COVID highlights the need for continued research, rehabilitation programs, and long-term healthcare strategies to support affected individuals.<sup>14</sup>

### **Risk Factors and Vulnerable Populations**

While COVID-19 can affect individuals of all ages, certain populations are at a significantly higher risk of developing severe disease and complications. Elderly individuals, particularly those above the age of 65, face the highest mortality rates due to age-related immune decline and the presence of multiple comorbidities. Immunocompromised individuals, including those undergoing chemotherapy, organ transplant recipients, and patients with autoimmune diseases, are more susceptible to severe infections due to weakened immune defenses. Pre-existing conditions such as diabetes, hypertension, chronic kidney disease, and obesity have been strongly associated with increased hospitalization and mortality rates. Lifestyle factors, including smoking, poor nutrition, and lack of physical activity, further exacerbate the risk of severe outcomes, reinforcing the importance of targeted prevention strategies for high-risk groups.<sup>15,16</sup>

### **Impact on Different Age Groups and Comorbidities**

The clinical presentation and severity of COVID-19 vary significantly across different age groups. Children generally experience milder symptoms, with many remaining asymptomatic or presenting with only mild fever and respiratory symptoms. However, rare but severe cases of multisystem inflammatory syndrome in children (MIS-C) have been reported, characterized by systemic inflammation, fever, rash, and organ dysfunction, often requiring intensive care. Young adults and middle-aged individuals typically recover with mild to moderate symptoms, but those with underlying health conditions face a heightened risk of severe illness. In contrast, older adults, especially those with pre-existing cardiovascular, metabolic, or pulmonary diseases, experience higher morbidity and mortality rates due to increased susceptibility to severe complications. The disproportionate impact of COVID-19 on individuals with comorbidities highlights the critical role of vaccination, early detection, and proactive management in reducing disease severity and mortality.<sup>16,17</sup>

### **Diagnosis**

#### **Overview of Diagnostic Tools**

The diagnosis of COVID-19 relies on a combination of molecular, antigen-based, and imaging techniques to confirm infection and assess disease severity. Nucleic acid amplification tests (NAATs), primarily reverse transcription-polymerase chain reaction (RT-PCR), serve as the gold standard for detecting SARS-CoV-2 RNA. Antigen tests provide rapid detection by identifying viral proteins, making them useful for mass screening, although they have lower sensitivity compared to RT-PCR. In cases where molecular tests yield inconclusive results or symptoms persist despite negative test results, imaging modalities such as chest X-rays and computed tomography (CT) scans assist in detecting characteristic lung abnormalities, including ground-glass opacities and bilateral lung involvement.<sup>18,19</sup>

### **Sensitivity and Specificity of Tests**

RT-PCR remains the most sensitive and specific diagnostic method, with accuracy exceeding 95% in ideal conditions. However, test performance is influenced by factors such as sample collection timing, viral load, and specimen quality. Rapid antigen tests, while providing quicker results (within 15–30 minutes), demonstrate reduced sensitivity, particularly in asymptomatic individuals and early-stage infections. Serological tests detecting antibodies against SARS-CoV-2 are valuable for epidemiological studies and assessing post-infection or post-vaccination immunity but are not reliable for acute diagnosis.<sup>20</sup>

### **Advances in Point-of-Care Testing**

The development of portable diagnostic kits and home-based rapid tests has significantly improved accessibility to COVID-19 testing. These point-of-care tests, including lateral flow assays and CRISPR-based diagnostic tools, have enabled early detection and faster isolation of infected individuals, particularly in remote or resource-limited settings. Advances in microfluidic technology and biosensor-based diagnostics continue to enhance the accuracy and affordability of rapid tests, paving the way for more decentralized testing strategies.<sup>21</sup>

### **Role of AI and Machine Learning in Diagnostics**

Artificial intelligence (AI) and machine learning have played a transformative role in COVID-19 diagnostics. AI-enhanced imaging techniques, including deep-learning algorithms applied to CT scans and chest X-rays, have improved early detection and differentiation from other respiratory diseases. Predictive modeling using AI has also been instrumental in forecasting disease progression, identifying high-risk patients, and optimizing healthcare resource allocation. AI-powered chatbots and virtual health assistants have further facilitated remote symptom assessment and triaging, reducing the burden on healthcare systems.<sup>22,23</sup>

## **Treatment and Management**

### **Antiviral Therapies and Their Mechanisms**

The treatment landscape for COVID-19 has evolved with the development of antiviral drugs aimed at reducing viral replication and disease severity. Remdesivir, an RNA polymerase inhibitor, has been approved for hospitalized patients with moderate to severe disease. Nirmatrelvir-ritonavir (Paxlovid) and molnupiravir, oral antiviral agents, have shown efficacy in reducing hospitalization and mortality in high-risk outpatients. These antivirals work by targeting viral replication mechanisms, preventing disease progression when administered early in the infection.<sup>24</sup>

### **Immunomodulatory Treatments**

Given the role of hyperinflammation in severe COVID-19 cases, immunomodulatory therapies have been crucial in disease management. Corticosteroids such as dexamethasone have been widely used to reduce systemic inflammation and lower mortality in critically ill patients. Monoclonal antibodies, including tocilizumab and baricitinib, target inflammatory cytokines such as interleukin-6 (IL-6) and Janus kinase (JAK) pathways, respectively, mitigating the cytokine storm that contributes to severe complications.<sup>25</sup>

### **Supportive Care and Hospital Management**

For severe COVID-19 cases, supportive care remains the cornerstone of treatment. Oxygen therapy, ranging from nasal cannula to high-flow oxygen and non-invasive ventilation, is essential for managing hypoxia. In critical cases, mechanical ventilation and extracorporeal membrane oxygenation (ECMO) provide life-saving respiratory support. Anticoagulation therapy is often administered to prevent thromboembolic complications associated with COVID-19. Intensive monitoring of organ function and timely interventions are crucial in managing critically ill patients.<sup>26</sup>

### **Role of Integrative Medicine and Alternative Therapies**

Ayurvedic, traditional Chinese medicine, and other complementary therapies have been explored as adjunct treatments for COVID-19. Herbal formulations, including those with immunomodulatory and antiviral properties, have been investigated for their potential to alleviate symptoms and enhance recovery. However, scientific validation through rigorous clinical trials remains necessary to establish their efficacy and safety. Integrative medicine approaches, including yoga, meditation, and nutritional interventions, have also gained attention for their role in supporting overall well-being and immune resilience.<sup>27</sup>

### **Public Health Measures**

#### **Role of Non-Pharmaceutical Interventions**

Non-pharmaceutical interventions (NPIs) have been fundamental in mitigating the spread of COVID-19. Social distancing measures, mask mandates, frequent hand hygiene, and improved ventilation in indoor spaces



have significantly reduced transmission rates. Public compliance with these interventions has varied across regions, affecting overall effectiveness.

### **Effectiveness of Lockdowns and Social Distancing**

Lockdowns and movement restrictions, implemented globally during the early phases of the pandemic, successfully slowed viral transmission and prevented healthcare system overloads. However, prolonged lockdowns resulted in severe economic downturns, increased unemployment, and exacerbated mental health issues. The trade-off between public health protection and economic stability remains a crucial consideration for future pandemic responses.

### **Global Strategies for Pandemic Containment**

Countries that adopted aggressive testing, contact tracing, and isolation strategies witnessed better outbreak control. Nations such as South Korea, New Zealand, and Taiwan successfully managed early outbreaks through rapid containment measures and widespread public cooperation. The integration of digital health tools, including contact-tracing apps and surveillance systems, further enhanced pandemic response capabilities.<sup>27,18</sup>

### **Economic and Mental Health Impacts**

The COVID-19 pandemic has significantly impacted global economies, leading to business closures, supply chain disruptions, and financial crises. Unemployment and economic instability have exacerbated mental health conditions, including anxiety, depression, and post-traumatic stress disorder (PTSD). The psychological burden of social isolation, grief, and uncertainty has highlighted the need for robust mental health support systems and policy interventions.

### **Vaccination Strategies**

#### **Development and Approval of Vaccines**

The rapid development of COVID-19 vaccines marked a significant milestone in global health. mRNA-based vaccines (Pfizer-BioNTech and Moderna) and vector-based vaccines (AstraZeneca, Sputnik V, and Johnson & Johnson) emerged as leading immunization strategies. These vaccines underwent accelerated clinical trials, demonstrating high efficacy in preventing severe disease and hospitalization.

#### **Efficacy and Safety Concerns**

While vaccine trials confirmed strong protective efficacy, concerns over rare adverse events, such as myocarditis (particularly in younger populations receiving mRNA vaccines) and vaccine-induced thrombotic thrombocytopenia (VITT) associated with adenoviral vector vaccines, necessitated continuous safety

monitoring. Despite these concerns, the benefits of vaccination far outweighed the risks, significantly reducing morbidity and mortality rates.

### **Breakthrough Infections and Booster Shots**

The emergence of SARS-CoV-2 variants, particularly Delta and Omicron, led to breakthrough infections in vaccinated individuals. While vaccines remained effective in preventing severe outcomes, waning immunity necessitated the introduction of booster doses. Booster campaigns aimed to enhance immunity, especially in vulnerable populations, ensuring prolonged protection against evolving variants.

### **Global Vaccination Disparities**

Despite advancements in vaccine development, disparities in vaccine access persisted across nations. High-income countries achieved high vaccination coverage early on, whereas low- and middle-income countries faced challenges in procurement, distribution, and public acceptance. Global initiatives such as COVAX aimed to bridge these gaps, yet vaccine inequity remained a critical barrier to achieving worldwide herd immunity. Addressing these disparities through improved supply chains, knowledge-sharing, and equitable distribution policies remains essential for future pandemic preparedness.

### **Future Preparedness and Conclusion**

#### **Strengthening Global Health Policies**

The COVID-19 pandemic has highlighted the critical need for a unified and coordinated global approach to pandemic preparedness and response. Strengthening international health policies requires sustained investments in public health infrastructure, robust healthcare systems, and efficient global supply chains. Governments and international organizations, including the World Health Organization (WHO) and the United Nations (UN), must collaborate to establish standardized pandemic response protocols, ensuring that resources such as vaccines, therapeutics, and medical supplies are rapidly deployed where they are needed most. Additionally, creating financial mechanisms to support low- and middle-income countries in pandemic response efforts will enhance global resilience against future infectious disease threats.

#### **Surveillance and Early Warning Systems**

Investing in real-time surveillance systems is essential for early detection and containment of emerging infectious diseases. Genomic surveillance, which involves continuous monitoring of viral mutations and pathogen evolution, plays a crucial role in identifying potential threats before they escalate into global pandemics. Advances in artificial intelligence (AI) and big data analytics can enhance the predictive capabilities of disease surveillance systems by analyzing patterns in healthcare records, social behavior, and environmental changes. Strengthening the capacity of international networks such as the Global Influenza

Surveillance and Response System (GISRS) and integrating AI-driven predictive modeling will allow for more effective outbreak forecasting and rapid response strategies.

### **Research Priorities for Pandemic Prevention**

To mitigate future pandemics, research must prioritize the development of broad-spectrum antivirals and universal vaccines capable of targeting multiple virus strains. Unlike strain-specific vaccines, universal vaccines would provide long-lasting immunity against various coronaviruses and influenza viruses, reducing the need for frequent booster shots. In addition, research on innovative therapeutic approaches, such as CRISPR-based antiviral treatments and monoclonal antibody therapies, could offer new strategies for combating viral infections. AI-driven diagnostics and wearable biosensors can revolutionize disease detection, enabling early identification of infections and personalized treatment plans. Moreover, global collaborations between academic institutions, pharmaceutical companies, and public health organizations will be key in accelerating scientific breakthroughs and ensuring equitable access to medical innovations.

### **Lessons from COVID-19 for Future Outbreaks**

The COVID-19 pandemic has provided invaluable lessons for future pandemic preparedness. First, global collaboration and information sharing must be prioritized to facilitate coordinated responses. The rapid development of COVID-19 vaccines demonstrated the potential of scientific innovation when researchers, governments, and private companies work together. Second, public trust in healthcare systems and science-based policies is essential for the successful implementation of health measures such as vaccination campaigns and non-pharmaceutical interventions. Transparent communication, combatting misinformation, and community engagement will be fundamental in addressing vaccine hesitancy and ensuring compliance with public health guidelines. Third, strengthening healthcare infrastructure, particularly in low-resource settings, will be necessary to enhance the capacity to handle future outbreaks. Investment in training healthcare workers, improving hospital preparedness, and expanding access to essential medical supplies will contribute to a more resilient global health system.<sup>29,30</sup>

### **CONCLUSION**

The COVID-19 pandemic has underscored the devastating impact of infectious diseases on global health, economies, and societies. However, it has also demonstrated the remarkable power of scientific progress, international cooperation, and adaptive public health strategies. By strengthening global health policies, investing in surveillance systems, prioritizing pandemic-related research, and applying lessons learned from COVID-19, the world can be better prepared to face future pandemics. A proactive, science-driven approach will be essential in mitigating the risks of emerging infectious diseases, ensuring global health security, and safeguarding the well-being of future generations.

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