

Review article

Nipah Virus Outbreaks: A CBRNE Framework for Global Biocontainment

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Abstract

Nipah virus (NiV), a deadly zoonotic pathogen with a fatality rate of 40-75%, continues to pose a significant pandemic threat, as evidenced by recent outbreaks in Kerala, India (2023 and 2024). These events highlight NiV's potential for human-to-human transmission, particularly in healthcare settings, and its ability to cause severe respiratory and neurological disease. Given the absence of approved vaccines or therapeutics, this review explores the implementation of CBRNE (Chemical, Biological, Radiological, Nuclear, and Explosive) strategies for outbreak containment through military-developed protocols including high-level biocontainment units, aerosolized disinfectant systems, and controlled movement zones. The proposed framework addresses NiV's unique challenges by combining rapid deployment of mobile isolation pods, strict corpse management procedures, and specialized healthcare worker protection with broader public health preparedness. Recent responses to epidemics in India demonstrate how the potential integration of CBRNE approaches can reduce transmission risks while preserving essential social functions through preventive training, interagency coordination, and strategic resource allocation. This review provides policymakers with actionable recommendations for mitigating NiV's biological threat through unified military-civilian response architectures.

Key words: Nipah Virus; CBRNE; Health Emergency Management; Global Security.

Introduction

NiV is a recently described paramyxovirus that causes an acute febrile encephalitic, stands as one of the most lethal zoonotic pathogens known to humanity due to its high lethality rate [1]. First identified during the 1998-1999 outbreak in Malaysia and Singapore, where it caused severe encephalitis and respiratory disease among pig farmers, NiV has since emerged periodically in South and Southeast Asia, with Bangladesh and India reporting recurrent cases [2,3]. The virus's natural reservoir is fruit bats the *Pteropus* genus, which shed the virus through urine, saliva, and other excretions, enabling spillover events to intermediate hosts (e.g., pigs) or directly to humans [2,3,4,5]. Recent outbreaks in Kerala, India (2023-2024), with case fatality rates exceeding 70%, have underscored NiV's potential for nosocomial amplification and limited human-to-human transmission-traits that elevate its pandemic risk; however, it should be noted that despite the high case fatality rate,

the number of deaths remained low [5,6,7,8] (Table 1).

Current public health strategies for NiV containment remain reactive, relying heavily on outbreak surveillance, contact tracing, and isolation protocols. However, the virus's high mortality rate, environmental stability, and potential for airborne transmission demand a more robust, preemptive approach [5,9]. This review argues for the integration of CBRNE (Chemical, Biological, Radiological, Nuclear, and Explosive) protocols, traditionally reserved for biowarfare scenarios, into NiV pandemic preparedness plans. By adapting military-grade strategies-such as mobile high-containment units, large-scale decontamination systems, and AI-assisted outbreak modeling-civilian health systems could bridge critical biocontainment, resource allocation, and crisis communication.

The objective of this review is threefold: (1) to synthesize the virological, epidemiological, and clinical features of NiV that necessitate a CBRNE framework; (2) to model a hypothetical pandemic



where scenario NiV acquires enhanced transmissibility; and (3) to propose actionable interventions, from lockdown **CBRNE** enforcement to corpse management, that could mitigate catastrophic outcomes. Climate change amplifies these risks by forcing bat populations to migrate into human-dominated landscapes due to habitat fragmentation and altered fruiting thereby increasing spillover cycles,

opportunities—a pattern documented in South Asian NiV hotspots. As deforestation and anthropogenic pressures further intensify human-bat interfaces, the lessons from NiV preparedness may extend to other high-consequence zoonoses, positioning CBRNE strategies as a cornerstone of 21st-century global health security [10].

Table 1. Chronology of Major Nipah Virus Outbreaks (1998-2024)

Year(s)	Location	Cases (Deaths)	Case Fatality Rate	Transmission Pattern	Key Findings	References
1998-1999	Malaysia/Singap ore	265 (105)	40%	Pig-to-human, limited human- to-human	First identified outbreak; farming nexus	
2001-2023	Bangladesh (Annual)	~300 (~210)	70-90%	Bat-to-human (date palm sap), human- to-human	High CFR; nosocomial superspreading events	[2,8]
2018, 2021, 2023-2024	Kerala, India	23 (17), 1 (1), 6 (4)	74-100%	Bat-to-human, hospital- acquired	Healthcare worker vulnerability	[2,0]
2007	West Bengal, India	5 (5)	100%	Unknown index case, human- to-human suspected	Small cluster with extreme lethality	

CFR = Case Fatality Rate.

NiV: Virology, Epidemiology, and Clinical Features

NiV belong to the genus Henipavirus in the family *Paramyxoviridae* and is relatively large (120–150 nm diameter), enveloped, single-stranded RNA virus [4,10]. Its genome encodes six structural proteins: nucleocapsid (N), phosphoprotein (P), matrix (M), fusion protein (F), glycoprotein (G), and RNA polymerase (L). The F and M proteins play a crucial role in the entry of the virus inside the host cell. In addition, these proteins facilitate viral penetration into endothelial and neuronal cells, explaining NiV's propensity for causing both severe respiratory distress and encephalitis [11].

Molecular studies have identified two primary strains—NiV-Malaysia (NiV-M) and NiV-Bangladesh (NiV-B)—with the latter demonstrating higher mortality and more frequent human-to-human transmission. Viral shedding occurs via respiratory secretions, urine, and saliva, creating multiple routes of exposure in outbreak settings [12,13,14] (Table 2).

Epidemiologically, NiV outbreaks have been largely confined to South and Southeast Asia, where human activities encroach upon the habitats of Pteropus fruit bats, the virus's natural reservoir [13]. Spillover events typically occur through direct contact with infected bats or consumption of contaminated raw date palm sap, followed by secondary transmission among humans via close contact [11,12,13]. The 1998-1999 Malaysian outbreak, linked to pig farming, resulted in 265 cases and 105 deaths, while subsequent outbreaks in Bangladesh and India have shown mortality rates exceeding 70%. Notably, the 2023 Kerala outbreak revealed alarming patterns of nosocomial spread, with healthcare workers accounting for 40% of cases, underscoring the virus's potential to exploit gaps in infection control protocols [14,15].

Clinically, NiV infection manifests in two primary forms: acute encephalitis and severe respiratory syndrome [8,16,17]. Early symptoms—fever, headache, and myalgia—are



nonspecific, often leading to misdiagnosis as influenza or dengue. Within days, neurological signs (disorientation, seizures, coma) or acute respiratory distress emerge, depending on the viral strain and host factors [18]. Magnetic resonance imaging (MRI) of encephalitic cases typically reveals diffuse cortical and brainstem lesions, while pulmonary involvement presents bilateral infiltrates resembling acute (ARDS) syndrome respiratory distress [15,16,17,18]. The absence of licensed vaccines or antivirals forces reliance on supportive care, with

ribavirin and monoclonal antibodies remaining experimental [19]. Survivors frequently exhibit long-term neurological sequelae, including personality changes and residual paralysis, further straining healthcare systems [20].

This triad of virological adaptability, epidemiological volatility, and clinical severity positions NiV as a uniquely challenging pathogen—one that demands innovative containment strategies beyond conventional public health measures [14,21].

Table 2. Comparative Features of Nipah Virus Strains

Characteristic	NiV-Malaysia (NiV-M)	NiV- Bangladesh (NiV-B)	Emerging Variants (Kerala)	References
Primary Reservoir	Pteropus hypomelanus	Pteropus medius	Pteropus giganteus	
Transmission	Swine intermediate host	Direct bat-to- human	Bat/human-to- human	
Human CFR	35-40%	70-90%	70-100%	
Clinical Focus	Encephalitis dominant	Respiratory + neurological	Rapid multi- organ failure	[13,14]
Human-to-Human	Rare	Frequent	Emerging evidence	
Molecular Marker	G protein (E447K mutation)	F protein cleavage efficiency	Enhanced fusion activity	

Containing a Nipah Virus Pandemic: A CBRNE Approach and Global Coordination Framework

The persistent recurrence of NiV outbreaks across South and Southeast Asia necessitates a tiered approach to epidemic preparedness, one that marries conventional public health measures with targeted military-derived containment strategies [14,22,23,24,25]. While communitylevel interventions addressing zoonotic spillover remain foundational, the unique characteristics of NiV - its staggering case fatality rate, propensity for nosocomial amplification, and environmental tenacity - create scenarios where civilian infrastructure becomes rapidly overwhelmed [11,14]. Historical precedents from Ebola and Middle East Respiratory Syndrome (MERS) outbreaks demonstrate that precisely calibrated CBRNE protocols can function as force multipliers when deployed judiciously alongside existing public health frameworks [26,27,28].

The operational superiority of CBRNE strategies manifests most clearly in three critical domains of outbreak response [29]. First, in rapid

case identification and containment, where mobile diagnostic units adapted from biodefense systems achieve laboratory-grade accuracy in field conditions, enabling real-time perimeter control without disrupting ongoing community education initiatives [30,31]. Second, healthcare facility protection, where modular isolation units derived from NATO CBRN standards prevent the hospital-based transmission clusters that accounted for nearly half of cases during recent NiV outbreaks [29,32]. Third, as a bridge to long-term solutions, with military-grade containment buying vital time for vaccine deployment - particularly relevant as several NiV vaccine candidates now progress through clinical trial phases [33].

Economic considerations, while often cited against such high-intensity approaches, must account for both direct costs and catastrophic risk mitigation [28]. Permanent high-containment facilities require capital expenditures orders of



magnitude greater than deployable CBRNE solutions, while the opportunity costs of uncontrolled outbreaks – in lives lost, healthcare systems paralyzed, and economies destabilized – dwarf prevention investments [28,29,32]. This calculus becomes particularly compelling when considering NiV's pandemic potential, a lesson seared into global consciousness by SARS-CoV-2's emergence from another ostensibly "low-mortality" zoonosis [29,32,34].

The ethical implementation of such measures rigorous safeguards. demands Singapore's response blueprint pandemic offers instructive model, combining military-grade outbreak analytics with robust civilian oversight - using anonymized heat mapping rather than individual surveillance, and collocating advanced containment units with community treatment centers to maintain accessibility and public trust. This balanced approach acknowledges that the extraordinary powers invoked during biological crises must be both proportional and transparent

Ultimately, the justification for CBRNE integration lies not in replacing traditional public health, but in providing specialized tools for scenarios where conventional measures falter against particularly virulent pathogens [36,37]. As climate change intensifies human-wildlife interfaces and global connectivity accelerates outbreak potential, such multidimensional preparedness frameworks may well determine whether localized zoonotic events escalate into civilizational threats [10,28].

Conclusion

NiV represents a paradigm-shifting challenge in pandemic preparedness, where conventional public health measures reach their limits against a pathogen combining high mortality, environmental persistence, and nosocomial transmission risks. This review establishes that selectively adapted CBRNE protocols—particularly mobile high-containment units and

References

- Gazal S, Sharma N, Gazal S, Tikoo M, Shikha D, Badroo GA, et al. Nipah and Hendra Viruses: Deadly Zoonotic Paramyxoviruses with the Potential to Cause the Next Pandemic. Pathogens. 2022;11(12):1419.
 - https://doi.org/10.3390/pathogens11121419
- 2. Soman Pillai V, Krishna G, Valiya Veettil M. Nipah Virus: Past Outbreaks and Future Containment. Viruses. 2020;12(4):465. https://doi.org/10.3390/v12040465

precision decontamination systems—can bridge this preparedness gap when integrated with civilian health infrastructure. The success of such hybrid approaches in recent Ebola and MERS outbreaks demonstrates their viability, though full implementation demands three pillars: (1) World Health Organization (WHO)-coordinated military-civilian task forces for cross-border response, (2) regional training hubs for CBRNE-adapted biocontainment, and (3) parallel investment in both vaccine development and outbreak-ready deployment systems.

While ethical governance remains essential—particularly regarding movement restrictions and resource triage—the accelerating frequency of zoonotic spillovers under climate change leaves little margin for delay. The strategies outlined here provide not merely a response framework for NiV, but a scalable prototype for future high-consequence pathogens. Their proactive adoption could redefine global health security from reactive containment to preventable crisis.

Ethics Approval and Consent to Participate

The informed consent was waived because of the retrospective nature of this study.

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Conflict of Interest

The authors have no conflicts of interest to declare.

Authors' Contributions

Conceptualization: GML, PAT; Data curation: GML, AM; Formal analysis: PAT; Investigation: GML, PAT; Methodology: RQ; Project administration: AI, CR, AM; Resources: GG, SR; Supervision: AM; Validation: AI, CR, AM; Visualization: AM, GG, SR; Writing—original draft: GML; Writing—review & editing: all authors.

- 3. Bhowmik A, Hasan M, Redoy MMH, Saha G. Nipah virus outbreak trends in Bangladesh during the period 2001 to 2024: a brief review. Sci One Health. 2024;4:100103.
 - https://doi.org/10.1016/j.soh.2024.100103
- 4. Sabir AJ, Rong L, Broder CC, Amaya M. Cedar virus biology and its applications as a surrogate for highly pathogenic henipaviruses. Cell Insight. 2024;3(4):100181.
 - https://doi.org/10.1016/j.cellin.2024.100181



- 5. Mohapatra P, Nazli Khatib M, Shabil M, Rajput P, Sharma N, Satapathy P, et al. Addressing the Nipah virus threat: A call for global vigilance and coordinated action. Clinical Infection in Practice. 2024;24:100390. https://doi.org/10.1016/j.clinpr.2024.100390
- 6. Sanker V, Vellekkat F, Dave T. Nipah Virus
 Outbreaks in Kerala: An Impending Doom?. Health
 Sci Rep. 2024;7(11):e70195.
 https://doi.org/10.1002/hsr2.70195
- Rahim AA, Chandran P, Bindu V, Radhakrishnan C, Moorkoth AP, Ramakrishnan LV. Recurrent Nipah outbreaks in Kerala: implications for health policy and preparedness. Front Public Health. 2024;12:1356515. https://doi.org/10.3389/fpubh.2024.1356515
- 8. Sahay RR, Patil DY, Chenayil S, Shete AM, Ps KS, Mohandas S, et al. Encephalitis-predominant Nipah virus outbreaks in Kerala, India during 2024. J Infect Public Health. 2025;18(7):102782. https://doi.org/10.1016/j.jiph.2025.102782
- 9. Anish TS, Aravind R, Radhakrishnan C, Gupta N, Yadav PD, Cherian JJ, et al. Pandemic potential of the Nipah virus and public health strategies adopted during outbreaks: Lessons from Kerala, India. PLOS Glob Public Health. 2024;4(12):e0003926. https://doi.org/10.1371/journal.pgph.0003926
- 10. Festa F, Ancillotto L, Santini L, Pacifici M, Rocha R, Toshkova N, et al. Bat responses to climate change: a systematic review. Biol Rev Camb Philos Soc. 2023;98(1):19-33. https://doi.org/10.1111/brv.12893
- 11. Ganguly A, Mahapatra S, Ray S, Chattopadhyay S, Islam MJ, Garai S, et al. The rising threat of Nipah virus: a highly contagious and deadly zoonotic pathogen. Virol J. 2025;22(1):139. https://doi.org/10.1186/s12985-025-02728-4
- 12. Lawrence P, Escudero-Pérez B. Henipavirus Immune Evasion and Pathogenesis Mechanisms: Lessons Learnt from Natural Infection and Animal Models. Viruses. 2022;14(5):936. https://doi.org/10.3390/v14050936
- 13. Chattu VK, Kumar R, Kumary S, Kajal F, David JK. Nipah virus epidemic in southern India and emphasizing "One Health" approach to ensure global health security. J Fam Med Prim Care. 2018;7:275–283. https://doi.org/10.4103/jfmpc.jfmpc 137 18
- 14. Branda F, Ceccarelli G, Giovanetti M, Albanese M, Binetti E, Ciccozzi M, et al. Nipah Virus: A Zoonotic Threat Re-Emerging in the Wake of Global Public Health Challenges. Microorganisms. 2025; 13(1):124.
- https://doi.org/10.3390/microorganisms13010124
- 15. Looi LM, Chua KB. Lessons from the Nipah virus outbreak in Malaysia. Malays J Pathol. 2007;29(2):63-67.
- 16. Lee KE, Umapathi T, Tan CB, Tjia HT, Chua TS, Oh HM, et al. The neurological manifestations of Nipah

- virus encephalitis, a novel paramyxovirus. Ann Neurol. 1999;46(3):428-432.
- 17. Lim CC, Sitoh YY, Hui F, Lee KE, Ang BS, Lim E, et al. Nipah viral encephalitis or Japanese encephalitis? MR findings in a new zoonotic disease. AJNR Am J Neuroradiol. 2000;21(3):455-461.
- 18. Sejvar JJ, Hossain J, Saha SK, Gurley ES, Banu S, Hamadani JD, et al. Long-term neurological and functional outcome in Nipah virus infection. Ann Neurol. 2007;62(3):235-242. doi:10.1002/ana.21178.
 - https://doi.org/10.1002/ana.21178
- 19. Sahay RR, Yadav PD, Gupta N, Shete AM, Radhakrishnan C, Mohan G, et al. Experiential learnings from the Nipah virus outbreaks in Kerala towards containment of infectious public health emergencies in India. Epidemiol Infect. 2020;148:e90.
- https://doi.org/10.1017/S0950268820000825
 20. Chan XHS, Haeusler IL, Choy BJK, Hassan MZ, Takata J, Hurst TP, et al. Therapeutics for Nipah virus disease: a systematic review to support prioritisation of drug candidates for clinical trials. Lancet Microbe. 2025;6(5):101002. https://doi.org/10.1016/j.lanmic.2024.101002
- 21. Sayed A, Bottu A, Qaisar M, Mane MP, Acharya Y. Nipah virus: a narrative review of viral characteristics and epidemiological determinants. Public Health. 2019;173:97-104. https://doi.org/10.1016/j.puhe.2019.05.019
- 22. Khan SU, Gurley ES, Hossain MJ, Nahar N, Sharker MA, Luby SP. A randomized controlled trial of interventions to impede date palm sap contamination by bats to prevent nipah virus transmission in Bangladesh. PLoS One. 2012;7(8):e42689. https://doi.org/10.1371/journal.pone.0042689
- 23. Cappelle J, Hoem T, Hul V, Furey N, Nguon K, Prigent S, et al. Nipah virus circulation at humanbat interfaces, Cambodia. Bull World Health Organ. 2020;98(8):539-547. https://doi.org/10.2471/BLT.20.254227
- 24. Khan S, Akbar SMF, Mahtab MA, Uddin MN, Rashid MM, Yahiro T, et al. Twenty-five years of Nipah outbreaks in Southeast Asia: A persistent threat to global health. IJID Reg. 2024;13:100434. https://doi.org/10.1016/j.ijregi.2024.100434
- 25. Hassan MZ, Rojek A, Olliaro P, Horby P. Improving clinical care of patients in Nipah outbreaks: moving beyond 'compassionate use'. Lancet Reg Health Southeast Asia. 2025;33:100527. https://doi.org/10.1016/j.lansea.2024.100527
- 26. Cenciarelli O, Gabbarini V, Pietropaoli S, Malizia A, Tamburrini A, Ludovici GM, et al. Viral bioterrorism: learning the lesson of Ebola virus in West Africa 2013- 2015. Virus Res 2015;210:318–26. https://doi.org/10.1016/j. virusres.2015.09.002
- 27. Kinsman J, Angrén J, Elgh F, Furberg M, Mosquera PA, Otero-García L, et al. Preparedness and



- response against diseases with epidemic potential in the European Union: a qualitative case study of Middle East Respiratory Syndrome (MERS) and poliomyelitis in five member states. BMC Health Serv Res. 2018;18(1):528. https://doi.org/10.1186/s12913-018-3326-0
- 28. Ludovici GM, Tassi PA, Iannotti A, Russo C, Quaranta R, Manenti G, et al. Bioterrorism and CBRNe threats: The role of Ebola in global security. Ethics Med Public Health. 2025;33:101138. https://doi.org/10.1016/j.jemep.2025.101138
- 29. Malizia A, Filograna L, Sbordone FP, Ciccarese G, Carbone A, Carreri B, et al. Response of a radiology department to the SARS-CoV-2 pandemic: the experience of the hospital "Policlinico Tor Vergata" in Rome. Int J Environ Res Public Health 2022;19:4688. https://doi.org/10.3390/ijerph19084688
- 30. Ludovici GM, Gabbarini V, Cenciarelli O, Malizia A, Tamburrini A, Pietropaoli S, et al. A review of techniques for the detection of biological warfare agents. Def S&T Tech Bull. 2015;8:17–26.
- 31. Mormando G, Paganini M, Alexopoulos C, Savino S, Bortoli N, Pomiato D, et al. Life-Saving Procedures Performed While Wearing CBRNe Personal Protective Equipment: A Mannequin Randomized Trial. Simul Healthc. 2021;16(6):e200-e205. https://doi.org/10.1097/SIH.0000000000000540
- 32. Manenti G, Ludovici GM, D'Amario R, Iannotti A, Russo C, Quaranta R, et al. Implementation and application of a contingency plan in case of an unconventional CBRNe event: A case study in a

- hospital facility. Def S T Tech Bull. 2025;18(1):21-33.
- 33. Mishra G, Prajapat V, Nayak D. Advancements in Nipah virus treatment: Analysis of current progress in vaccines, antivirals, and therapeutics. Immunology. 2024;171(2):155-169. https://doi.org/10.1111/imm.13695
- 34. Klingelhöfer D, Braun M, Naser CA, Brüggmann D, Groneberg DA. Emerging Nipah Virus With Pandemic Potential and High Mortality Rates: Is the Scientific Community Learning From Former Pandemics?. Rev Med Virol. 2025;35(2):e70028. https://doi.org/10.1002/rmv.70028
- 35. Kuguyo O, Kengne AP, Dandara C. Singapore COVID-19 Pandemic Response as a Successful Model Framework for Low-Resource Health Care Settings in Africa?. OMICS. 2020;24(8):470-478. https://doi.org/10.1089/omi.2020.0077
- 36. Petersen L, Havârneanu GM, Arnold A, Carbon D, Görgen T, Gavel A, et al. Applicability of PROACTIVE recommendations on CBRNe risks and threats to passenger rail and metro sectors. J Transp Secur. 2023;16(1):4. https://doi.org/10.1007/s12198-023-00263-3
- 37. Qzih ES, Ahmad MM. Hospital-Based Preparedness Measures for CBRNE Disasters: A Systematic Review. Environ Health Insights. 2024;18:11786302241288859. https://doi.org/10.1177/11786302241288859

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