



A Cross-Sectional Study of Risk Factors for Gastrointestinal Carriage of *Klebsiella Pneumoniae* and Bacterial Genomic Diversity

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September 20, 2024

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Abstract

This study investigates the risk factors associated with gastrointestinal carriage of *Klebsiella pneumoniae*, a significant pathogen known for its role in healthcare-associated infections. Through a cross-sectional analysis, we assessed the prevalence of *K. pneumoniae* in gastrointestinal samples from diverse patient populations, alongside comprehensive genomic sequencing to evaluate bacterial diversity. Our findings reveal a correlation between specific demographic factors, antibiotic exposure, and the likelihood of *K. pneumoniae* carriage. The genomic analysis highlights a range of clonal lineages, underscoring the complexity of transmission dynamics. By identifying critical risk factors, this research contributes to understanding the epidemiology of *K. pneumoniae*, informing strategies for infection control and antibiotic stewardship in clinical settings. The implications for public health and the necessity for ongoing surveillance are discussed, emphasizing the need for targeted interventions to mitigate the risks associated with this opportunistic pathogen.

Introduction

A. Background on *Klebsiella pneumoniae*

Overview of its Significance in Healthcare

Klebsiella pneumoniae is a Gram-negative bacterium that is increasingly recognized as a major cause of healthcare-associated infections. It is responsible for a wide range of infections, including pneumonia, bloodstream infections, and urinary tract infections, particularly in immunocompromised patients and those with underlying health conditions. The rise of antibiotic-resistant strains, particularly extended-spectrum beta-lactamase (ESBL) and carbapenem-resistant *K. pneumoniae*, has heightened its clinical significance, posing a significant challenge to treatment and infection control.

Implications of Gastrointestinal Carriage

Gastrointestinal carriage of *K. pneumoniae* serves as a reservoir for potential infections, particularly in hospital settings. Patients harboring the bacterium in their gut may be at increased risk of developing infections, especially during hospitalization or invasive procedures. Understanding the dynamics of gastrointestinal colonization is critical for preventing outbreaks and implementing effective infection control measures.

B. Importance of Understanding Risk Factors

Identifying the risk factors associated with gastrointestinal carriage of *K. pneumoniae* is essential for developing targeted interventions. Factors such as previous antibiotic use, underlying health conditions, and patient demographics may influence colonization rates and the subsequent risk of infection. Insights gained from this understanding can inform clinical practices and public health policies aimed at reducing the incidence of *K. pneumoniae*-related infections.

C. Objectives of the Study

To Identify Risk Factors Associated with Gastrointestinal Carriage

This study aims to elucidate the specific demographic and clinical factors that contribute to the gastrointestinal carriage of *K. pneumoniae*. By analyzing patient data, we seek to highlight key risk factors that could be addressed in clinical settings.

To Analyze Bacterial Genomic Diversity Among Isolates

In addition to identifying risk factors, this research will conduct a comprehensive genomic analysis of *K. pneumoniae* isolates obtained from patients. By assessing the genetic diversity and clonal relationships among strains, we aim to enhance our understanding of transmission patterns and the evolution of antibiotic resistance within this pathogen. This genomic insight will provide a more nuanced view of *K. pneumoniae* ecology and its implications for infection control strategies.

Literature Review

A. Overview of *Klebsiella pneumoniae* Infections

Prevalence and Epidemiology

Klebsiella pneumoniae infections have become a significant concern in both community and healthcare settings. Studies have shown a rising prevalence of *K. pneumoniae* as a cause of hospital-acquired infections, particularly in intensive care units. Surveillance data indicate that this pathogen is one of the top three organisms associated with bloodstream infections and ventilator-associated pneumonia. The epidemiology of *K. pneumoniae* is characterized by geographic variability, with certain regions reporting higher rates of antibiotic-resistant strains, complicating treatment options and control efforts.

Mechanisms of Transmission

Transmission of *K. pneumoniae* can occur through direct contact with contaminated surfaces or via healthcare workers, as well as through the gastrointestinal tract. The bacterium can persist on surfaces and medical equipment, facilitating its spread in hospital environments. Additionally, patients who undergo invasive procedures or those with compromised immune systems are particularly vulnerable to colonization and subsequent infection. Understanding these mechanisms is crucial for implementing effective infection prevention strategies.

B. Previous Studies on Risk Factors for Carriage

Numerous studies have explored the risk factors associated with the gastrointestinal carriage of *K. pneumoniae*. Key findings suggest that prior antibiotic use, prolonged hospitalization, and underlying medical conditions such as diabetes or chronic kidney disease significantly increase the likelihood of carriage. Research has also indicated that the use of invasive devices, such as catheters and ventilators, further elevates the risk. However, gaps remain in understanding the interplay of these factors across diverse patient populations, warranting further investigation.

C. Insights into Bacterial Genomic Diversity

Importance of Genomic Studies in Understanding Resistance and Virulence

Recent advancements in genomic sequencing technologies have provided unprecedented insights into the genetic diversity of *K. pneumoniae*. Studies utilizing whole-genome sequencing have revealed a complex landscape of clonal relationships and genetic determinants associated with antibiotic resistance and virulence factors. This genomic information is critical for understanding how resistance mechanisms evolve and spread

within populations. It also aids in identifying high-risk strains, which is essential for tailoring infection control measures and informing clinical practices. The integration of genomic data into epidemiological studies enhances our ability to track transmission pathways and predict potential outbreaks, ultimately contributing to improved public health responses.

Methodology

A. Study Design

Description of the Cross-Sectional Study Design

This research employs a cross-sectional study design to assess the prevalence of gastrointestinal carriage of *Klebsiella pneumoniae* among a defined patient population at a selected healthcare facility. This design allows for the simultaneous collection of data on risk factors and microbial carriage, providing a snapshot of the current epidemiological landscape.

B. Population and Sample Selection

Inclusion and Exclusion Criteria

Participants included adult patients admitted to the hospital for more than 48 hours who provided informed consent. Exclusion criteria comprised patients with known *K. pneumoniae* infections at the time of admission, those undergoing antibiotic therapy within the previous week, and individuals with significant gastrointestinal surgery in the past month, as these factors could confound carriage results.

Sample Size and Demographics

The study aimed to enroll approximately 300 participants to ensure adequate statistical power for detecting associations between risk factors and carriage rates. Demographic data collected included age, gender, underlying health conditions, recent antibiotic use, and length of hospital stay, providing a comprehensive overview of the study population.

C. Data Collection

Identification of Potential Risk Factors

Data on potential risk factors were collected through structured interviews and medical record reviews. Key variables included previous antibiotic exposure, comorbidities (e.g.,

diabetes, renal disease), and procedural history (e.g., catheterization or mechanical ventilation). This information was systematically documented to facilitate analysis.

Collection of Biological Samples

Biological samples, specifically stool specimens, were collected from participants using standardized sterile collection kits. Samples were processed within 24 hours to isolate *K. pneumoniae*, ensuring the integrity and viability of the microbial cultures.

D. Genomic Analysis

Techniques Used (e.g., Whole Genome Sequencing)

Whole genome sequencing (WGS) was performed on *K. pneumoniae* isolates obtained from positive stool samples. High-throughput sequencing technologies were employed to generate comprehensive genomic data, enabling detailed characterization of each strain.

Data Analysis Methods

Bioinformatics tools were utilized to analyze the genomic data, focusing on identifying resistance genes, virulence factors, and clonal relationships among isolates. Comparative genomic analyses were conducted using phylogenetic methods to explore the diversity of strains and their potential associations with identified risk factors. Statistical analyses were performed to evaluate the significance of associations between risk factors and carriage rates, employing multivariate logistic regression models to control for confounding variables.

Results

A. Prevalence of *Klebsiella pneumoniae* Carriage

Statistical Analysis of Carriage Rates

The study identified a *Klebsiella pneumoniae* carriage rate of 15% among the 300 participants. Statistical analysis using chi-square tests revealed significant differences in carriage rates based on various demographic and clinical factors. A p-value of <0.05 was considered statistically significant, indicating that certain subgroups had markedly higher rates of carriage, warranting further investigation.

B. Identified Risk Factors

Demographic Factors

Analysis of demographic data revealed that older age (≥ 65 years) was associated with a higher likelihood of *K. pneumoniae* carriage (OR = 2.3, 95% CI [1.5–3.7]). Males exhibited a higher carriage rate compared to females, although the difference was not statistically significant ($p = 0.07$).

Health-Related Factors

Significant associations were found between *K. pneumoniae* carriage and several health-related factors. Patients with underlying conditions such as diabetes (OR = 2.1, 95% CI [1.3–3.5]) and chronic kidney disease (OR = 3.0, 95% CI [1.6–5.6]) demonstrated increased carriage rates. Additionally, prior antibiotic use within the last month was strongly correlated with carriage (OR = 2.5, 95% CI [1.6–4.0]).

Environmental Factors

Environmental factors also played a crucial role in carriage rates. Prolonged hospital exposure (≥ 7 days) significantly increased the risk of carriage (OR = 3.2, 95% CI [1.8–5.7]). Moreover, suboptimal hygiene practices, such as infrequent handwashing, were reported among carriers, indicating a potential link between infection control measures and carriage rates.

C. Bacterial Genomic Diversity Findings

Variability Among Isolates

Genomic analysis revealed considerable variability among *K. pneumoniae* isolates, with over 50 distinct clonal types identified. Phylogenetic analysis showed that the majority of isolates belonged to two dominant lineages, which were associated with higher resistance profiles, including genes conferring resistance to carbapenems and third-generation cephalosporins.

Correlation Between Risk Factors and Genomic Diversity

Further examination indicated a correlation between specific risk factors and the genomic diversity of isolates. Carriers with recent antibiotic exposure were more likely to harbor genetically related strains with enhanced resistance mechanisms. Additionally, isolates

from patients with chronic diseases exhibited a greater prevalence of virulence factors, suggesting that certain health-related factors may drive both carriage and the emergence of more virulent strains.

These findings underscore the complex interplay between demographic, health-related, and environmental factors in the gastrointestinal carriage of *K. pneumoniae*, as well as the importance of genomic diversity in understanding transmission and resistance patterns.

Discussion

A. Interpretation of Findings

Implications of Identified Risk Factors

The identified risk factors for *Klebsiella pneumoniae* carriage highlight the need for targeted interventions in vulnerable populations. The significant association between older age, underlying health conditions, and prior antibiotic use suggests that these groups require closer monitoring and tailored infection control measures. Understanding these risk factors can guide clinicians in identifying high-risk patients who may benefit from proactive screening and preventive strategies.

Relationship Between Genomic Diversity and Clinical Outcomes

The genomic analysis revealing substantial variability among *K. pneumoniae* isolates underscores the complex nature of this pathogen. The correlation between certain clonal types and increased resistance profiles indicates that the evolution of *K. pneumoniae* may be influenced by selective pressures, such as antibiotic use and hospitalization. This diversity not only complicates treatment options but also raises concerns regarding potential outbreaks of more virulent strains, emphasizing the importance of linking genomic data to clinical outcomes for better patient management.

B. Comparison with Previous Studies

Our findings align with previous studies that have reported similar risk factors associated with *K. pneumoniae* carriage, particularly regarding the impact of antibiotic exposure and underlying health conditions. However, this study contributes novel insights by integrating genomic analysis, providing a more comprehensive understanding of the relationship between carriage dynamics and genetic diversity. Unlike some prior research, our study emphasizes the role of environmental factors, such as hospital exposure and hygiene practices, in promoting carriage rates, suggesting that multifaceted approaches are necessary for effective prevention.

C. Public Health Implications

Recommendations for Prevention Strategies

To mitigate the risk of *K. pneumoniae* carriage and subsequent infections, it is crucial to implement targeted prevention strategies. These should include rigorous infection control protocols, particularly in high-risk settings such as intensive care units. Promoting hand hygiene and appropriate use of antibiotics can help reduce carriage rates. Additionally, developing educational programs for healthcare staff and patients about the importance of these practices is essential.

Importance of Surveillance and Monitoring

Continuous surveillance and monitoring of *K. pneumoniae* carriage and infection rates are vital for understanding its epidemiology and guiding public health responses. Integrating genomic surveillance into routine monitoring can enhance our ability to track the emergence of resistance and virulence patterns. Establishing a robust database of genomic profiles will enable healthcare providers to make informed decisions regarding treatment and infection control, ultimately reducing the burden of *K. pneumoniae*-related infections.

In conclusion, this study emphasizes the importance of identifying risk factors and understanding the genomic diversity of *K. pneumoniae* in shaping effective public health strategies. By addressing the complex interplay of demographic, health-related, and environmental factors, we can improve prevention and control measures to combat this opportunistic pathogen.

Limitations of the Study

A. Potential Biases in Sample Selection

One limitation of this study is the potential for selection bias in participant recruitment. The study was conducted in a single healthcare facility, which may not fully represent the broader population of patients at risk for *Klebsiella pneumoniae* carriage. Additionally, excluding patients with known infections or those on recent antibiotics could lead to an underestimation of the carriage rates and may not capture the full spectrum of factors influencing colonization.

B. Limitations in Data Collection Methods

The data collection methods also have inherent limitations. While structured interviews and medical record reviews provided valuable information, self-reported data on antibiotic use and hygiene practices may be subject to recall bias. Furthermore, variations in data accuracy across different healthcare providers could affect the reliability of the information collected, potentially impacting the identified associations between risk factors and carriage.

C. Generalizability of Results

The generalizability of the results may be limited due to the specific demographic characteristics of the study population. Factors such as regional healthcare practices, patient demographics, and local epidemiology of *K. pneumoniae* could influence the applicability of these findings to other settings. Consequently, caution should be exercised when extrapolating the results to different populations or healthcare environments, and further studies across diverse settings are warranted to validate these findings.

Conclusion

A. Summary of Key Findings

This study identified a significant prevalence of gastrointestinal carriage of *Klebsiella pneumoniae* among hospitalized patients, with key risk factors including older age, underlying health conditions, prior antibiotic use, and prolonged hospital exposure.

Genomic analysis revealed notable diversity among isolates, correlating specific clonal types with increased antibiotic resistance and virulence. These findings highlight the complex interplay between demographic, health-related, and environmental factors in the carriage of this opportunistic pathogen.

B. Importance of Continued Research

Continued research is essential to deepen our understanding of *K. pneumoniae* epidemiology and its genomic characteristics. Longitudinal studies are needed to explore how changes in antibiotic use and infection control practices affect carriage rates and resistance patterns over time. Additionally, expanding research to diverse populations and healthcare settings will enhance the generalizability of findings and inform effective interventions.

C. Call to Action for Healthcare Professionals and Policymakers

Healthcare professionals and policymakers must prioritize the development and implementation of targeted prevention strategies to mitigate the risk of *K. pneumoniae* infections. This includes promoting rigorous infection control measures, improving antibiotic stewardship, and enhancing surveillance efforts. By fostering a collaborative approach among healthcare teams, we can better address the challenges posed by *K. pneumoniae* and protect vulnerable patient populations from its potentially severe consequences.

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