



# From empirical to precision therapy in ICUs: Rethinking antibiotic use after COVID-19

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The coronavirus disease 2019 (COVID-19) pandemic caused a unique surge in empiric broad-spectrum antibiotic use in intensive care units (ICUs), even when bacterial coinfection was unlikely [1-3]. Such widespread empirical prescribing, while understandable under crisis conditions, has likely accelerated antimicrobial resistance (AMR) in critical care settings, exacerbating challenges for post-pandemic infection management. In this regard, we believe the time is suitable to advocate a transition from empirical toward precision-based antimicrobial therapy in ICUs, integrating diagnostics, stewardship, and clinical decision support to better shape antibiotic use. During the early waves of COVID-19, studies documented that antibiotic prescribing rates were high, reaching nearly 70% in ICU and hospitalized patients, despite bacterial coinfection rates typically being low [1,4,5]. For example, an observational analysis revealed that empirical antibiotic overuse in a COVID-19 ICU correlated with increased spread of carbapenem-resistant Gram-negative (CR-GNB) pathogens [6]. These observations underscore a critical gap between prevailing clinical practices and the true microbiological risk profile in critically ill patients, a gap largely driven by diagnostic uncertainty and the fear of overlooking bacterial superinfection. Transitioning to precision therapy requires several integrated strategies. First, rapid diagnostics (e.g. multiplex polymerase chain reaction [PCR] panels, molecular pathogen detection, antimicrobial resistance gene assays) used within an antimicrobial stewardship framework have already demonstrated reduced time to optimal therapy, shorter hospital stays, and mortality benefits in bloodstream infections [7,8]. A network meta-analysis of 88 studies found that combining rapid diagnostic tests (RDTs) with stewardship programs reduced mortality compared to standard blood culture alone (odds ratio [OR] 0.72; 95% CI 0.59–0.87) [9]. However, RDTs alone offered no benefit, underlining the importance of the stewardship component. Second, diagnostic stewardship ordering the right test for the right patient at the right time is essential to avoid overinterpretation or misapplication of molecular results [10]. Third, artificial intelligence-driven decision support, local antibiogram integration, risk stratification tools, and biomarker-guided algorithms (e.g., procalcitonin, CRP kinetics) can help clinicians calibrate antibiotic initiation and adjustment more precisely [11,12].

Realizing this paradigm shift requires investment in infrastructure, 24/7 molecular testing capacity, and integration of infectious diseases, pharmacy, and microbiology teams into ICU workflows.

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Institutional protocols should adopt rapid diagnostic-guided pathways with continuous monitoring, while clinical and observational studies are needed to validate precision algorithms, particularly in post-COVID populations. Building acceptance among intensivists depends on outcome evidence confirming the safety of de-escalation in high-risk patients, consistent with the stewardship principle of delivering the right drug at the right time and for the right duration [13].

In conclusion, the empirical antibiotic paradigm that dominated ICU practice during COVID-19 was a necessary short-term response to uncertainty, but it is unsustainable in a post-pandemic era faced with accelerating AMR. ICUs must evolve toward precision antimicrobial prescribing, leveraging diagnostics, stewardship, and smart decision support to preserve therapeutic options, reduce collateral harm, and optimize patient outcomes. We call upon critical-care leaders, journal editors, and funding agencies to prioritize and support research, infrastructure, and guideline updates that embed precision therapy into standard ICU practice.

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## Authors' contributions

HA, NK, KK: conceived the idea and drafted the initial version of the manuscript. NK, KK: contributed to the literature review and critical discussion. HA, NK: provided revisions for intellectual content and supervised the overall development of the work. All authors approved the final version of the manuscript.

## Conflict of interest

No potential conflict of interest was reported by the authors.

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Not applicable.

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