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Titolo	Polymyxin Antibiotics: From Laboratory Bench to Bedside [[electronic resource] /] / edited by Jian Li, Roger L. Nation, Keith S. Kaye
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Nota di contenuto	Overview Multidrug resistant Gram-negative pathogens: the urgent need for 'old' polymyxins Discovery, antibacterial spectrum and chemistry Mechanisms of antibacterial activity Mechanisms of resistance Bioanalysis and stability of polymyxins In vitro pharmacodynamics and PK/PD in animals Antibacterial susceptibility breakpoints Labeling of parenteral products of colistin methanesulfonate Meta-analysis of polymyxin use in patients Clinical use of colistin Clinical use of polymyxin B Pharmacokinetics, pharmacodynamics, toxicodynamics and dosing recommendations Rational combinations with other antibiotics Toxicities in patients: management and mechanisms Antiendotoxin effect Discovery of novel polymyxin-like antibiotics Conclusion.
Sommario/riassunto	This volume is the first-ever complete treatise on polymyxins and presents the most comprehensive and up-to-date reviews of all major research and clinical topics from chemistry, microbiology, pharmacology, clinical use, to drug discovery. All chapters were written by internationally leading researchers and clinicians in the field. It is our wish that readers discover the importance of polymyxin structure in relation to the mechanisms of activity, resistance and toxicity. We emphasized that reliable analytic methods for polymyxins are critical

when investigating their pharmacokinetics (PK) and pharmacodynamics (PD). The complicated dose definitions and different pharmacopoeial standards have already compromised the safe use of polymyxins in patients. Therefore, informed by the latest pharmacological information, scientifically-based dosing recommendations have been proposed for intravenous polymyxins. Considering the PK/PD limitations and potential development of resistance, polymyxin combinations are encouraged; however, the current literature has not shown definite microbiological benefits, possibly because most clinical studies to date overlooked key PK/PD principles. Nephrotoxicity is the major dose-limiting factor and it is imperative to elucidate the mechanisms and develop novel approaches to minimize polymyxinassociated toxicities. In addition, the anti-endotoxin effect of polymyxins supports their clinical use to treat Gram-negative sepsis. Fortunately, the discovery of new-generation polymyxins with wider therapeutic windows has benefited from the latest achievements in polymyxin research. This book provides extensive pharmacological information on polymyxins to infectious diseases clinicians, pharmacists, clinical microbiologists, antimicrobial pharmacologists, and pharmaceutical scientists, and is an essential read for those who aim to develop novel polymyxins and improve their clinical use as a last-line defense against Gram-negative 'superbugs'.