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RESEARCH ARTICLE

Screening novel antiviral compounds to treat Clostridioides difficile infections

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Abstract

Clostridioides difficile is a major cause of nosocomial infections, often associated with individuals who have gut dysbiosis from previous antibiotic therapies. C. difficile infections. (CDI) have a high recurrence rate and impose significant financial and mortality burdens on the healthcare system. Therefore, novel anti-C. difficile drugs are urgently needed to treat and reduce the severity and recurrence of infection. In this study, we screened a library of 618 antiviral drugs to identify a potential candidate for repurposing as novel anti-C. difficile therapeutics. Following our preliminary screening, we identified 9 novel compounds that inhibited C: difficile at a concentration of 16 µM or lower. Among these, 4 antiviral compounds demonstrated the most potent anti-C. difficile activity against a panel of 15 C. difficile isolates, with minimum inhibitory concentrations (MICs) comparable to the drug of choice. vancomycin. These include rottlerin (MIC₅₀ = 0.25 µg/mL), a-mangostin (MIC₅₀ = 1 µg/mL), dryocrassin ABBA (MIC_{vn} = 1 µg/mL), and obefazimod (MIC_{vn} = 4 µg/mL). All exhibited minimal to no activity against representative members of the human out microbiota. Interestingly, a-mangostin, a natural xanthone derived from the mangosteen fruit, exhibited strong bactericidal action, clearing a high inoculum of C. difficile in less than an hour. All other drugs exhibited bacteriostatic activity. Given their characteristics, these compounds show great promise as novel treatments for CDI.

Introduction

Healthcare-associated infections are one of the greatest burdens on healthcare systems. Most bacteria responsible for a part of this burden are associated with antimicrobial resistance, with one notable exception: C. difficile. Despite lacking a significant antimicrobial resistance profile. C. difficile is still considered an urgent threat and one of the deadliest bacterial infections responsible for approximately 30,000 deaths annually, primarily affecting older populations [1–5]. The incidence of C. difficile infections (CDI) in the United States has increased over the last two decades and has remained relatively stable from 2021 to 2022, with the COVID-19 pandemic seeing a drop in cases due to improved strategies preventing the spread of

Screening The Novel

Deborah Cartmell

Screening The Novel:

Screening the Novel Robert Giddings, Keith Selby, Chris Wensley, 1990 Screening The Novel Keith Selby, Robert Giddings, 2016-07-27 The book takes as its theme the relationship between literature and the contemporary means of production and distribution collectively termed the media in particular film and television. The intention of the book is to explore and evaluate the mutual opportunities and restrictions in this relationship In the grammar of our culture there seems to be an accepted opinion that print is superior in terms of cultural production to film radio or television that to read a book is somehow a higher cultural activity than seeing a play on television or seeing a film By the same token a novel is a superior work of art to film or television. The longer perspective reveals that traditionally there always is a greater respect paid to the previous mode of literary production poetry was superior to drama poetic drama was superior to the novel and film attained cult and classic status initially over television **Private Screening** Richard North Patterson, 1985 The nation is stunned silent when presidential hopeful James Kilcannon is shot dead point blank while sharing the stage at a benefit concert with his rock star girlfriend Stacy Tarrant Fiercely independent attorney Tony Lord defends the assassin in a sensational trial that erupts into the media event of the decade But the most shocking gambit ever witnessed in the history of television has yet to unfold As America watches a mysterious and ruthless figure known only as Phoenix takes to the airwaves in the ultimate act of high tech terror Holding the wife of a wealthy newspaper mogul and Stacy's manager hostage Phoenix mounts a televised trial of his own in which Stacy Tarrant and Tony Lord are helpless defendants millions of viewers are jurors and unless his chilling demands are met Phoenix is the unstoppable executioner Copyright Libri GmbH All rights reserved **Novel Women** Liora Brosh, 2015-12-04 From Hollywood classics like Jane Eyre or Wuthering Heights to the 1990s wave of Jane Austen films adaptations of the British Nineteenth century novel have been sensationally popular This book examines how British and American filmmakers used the domestic novels of the past to construct stable gender ideals for the present

Screening the Novel Gabriel Miller, 2020 Some of the most memorable movies of Hollywood's Golden Age were based on novels that never received the acclaim they deserved No one who saw Rod Steiger in The Pawnbroker could forget the actor's wrenching performance but does anyone remember the author of the book on which the film was based The same can be said of Jane Fonda in They Shoot Horses Don't They Greta Garbo in Susan Lenox and Humphrey Bogart in The Treasure of the Sierra Madre This book retrieves these novels and re evaluates the careers of the eight neglected novelists whose works inspired eight different directors among them Stanley Kubrick Sidney Lumet John Huston and Sidney Pollack Each chapter offers detailed analysis on both the original text and the resulting movie Taken together the double examination of novel and film raises some important questions about the nature and problems of cinematic adaptation Keighley & Williams' Surgery of the Anus, Rectum and Colon, Fourth Edition Michael R.B. Keighley, Norman S. Williams, 2018-11-06 This fourth edition of Surgery of the Anus Rectum and Colon continues to redefine the field with its comprehensive coverage of

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high specificity and robust reactivity in the complex system such as site specific conjugation by covalent recognition between enzymes and their substrates With the perspective of protein drug interactions this book also reviews the applications of protein immobilization with an emphasis on G protein coupled receptors in drug discovery and protein ligand interaction analysis In addition the merits opportunities and disadvantages are analyzed for different immobilization methods and a perspective for future directions is presented Given its scope this book appeals to a broad readership particularly researchers engaged in the field of analytical chemistry bioconjugate chemistry and chemical biology and other related field as well as Chemical & Metallurgical Engineering Eugene Franz teachers of relevant majors in colleges and universities Roeber, Howard Coon Parmelee, 1924 Advanced Methods in Biomedical Signal Processing and Analysis Kunal Pal, Samit Ari, Arindam Bit, Saugat Bhattacharyya, 2022-09-07 Advanced Methods in Biomedical Signal Processing and Analysis presents state of the art methods in biosignal processing including recurrence quantification analysis heart rate variability analysis of the RRI time series signals joint time frequency analyses wavelet transforms and wavelet packet decomposition empirical mode decomposition modeling of biosignals Gabor Transform empirical mode decomposition The book also gives an understanding of feature extraction feature ranking and feature selection methods while also demonstrating how to apply artificial intelligence and machine learning to biosignal techniques Gives advanced methods in signal processing Includes Screen Adaptations: Jane Austen's Pride and machine and deep learning methods Presents experimental case studies Prejudice Deborah Cartmell, 2010-10-27 The study of literature on screen is a growing area of study in schools and universities Many students have to produce critical essays comparing the novel and film versions of a particular text The Screen Adaptations series offers a wealth of study material from the literary context of the original work through to thought provoking comparisons of the screen versions critical commentary and the afterlife of the films Pride and Prejudice is a classic piece of literature and any new adaptation is a major event With several well known TV and film versions available featuring stars such as Keira Knightley as well as a Bollywood version discussions around how these films interpret the story themes and characters is a popular classroom choice for students and teachers alike This new title in the Screen Adaptations series provides a rich source of material to help students understand and write about the reciprocal relationship between film and literature It offers in depth analysis of the various screen versions and alternative readings as well as critical insight and an interview with writer Andrew Davies best known for his 1995 BBC adaptation Following from Shakespeare Jane Austen on screen is now an established part of literary studies and the volume will conclude with a survey of the growing Microbiology for Cleaner Production and Environmental Sustainability Naga Raju body of literature in the field Maddela, Lizziane Kretli Winkelstroter Eller, Ram Prasad, 2023-08-15 Growth of populations increasing urbanization and rising standards of living due to technological innovations demand not only the meticulous use of shrinking resources but also sustainable ways of producing materials for human welfare Cleaner production involves preventive and protective initiatives

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Tumor-Associated Antigens and Their Autoantibodies: From Discovering to Clinical Utilization Xianggian Guo, Jianying Zhang, Bilian Jin, Qing Zhu, 2022-08-18 **Metagenomics** Munivandi Nagarajan, 2017-10-29 Metagenomics Perspectives Methods and Applications provides thorough coverage of the growing field of metagenomics A diverse range of chapters from international experts offer an introduction to the field and examine methods for metagenomic analysis of microbiota metagenomic computational tools and recent metagenomic studies in various environments The emphasis on application makes this text particularly useful for applied researchers practitioners clinicians and students seeking to employ metagenomic approaches to advance knowledge in the biomedical and life sciences Case study based application chapters examine topics ranging from viral metagenome profiling metagenomics in oral disease and health metagenomic insights into the human gut microbiome and metabolic syndromes and more Additionally perspectives on future potential at the end of each chapter provoke new thought and motivations for continued study in this exciting and fruitful research area Provides thorough coverage of the rapidly growing field of metagenomics with an emphasis on applications of relevance to translational researchers practitioners clinicians and students Features a diverse range of chapters from international experts that offer an introduction to the field and examine methods for metagenomic analysis of microbiota metagenomic computational tools and research pipelines Highlights perspectives on future potential at the end of each chapter to provoke new thought and motivations for continued study in this exciting and fruitful research area Introduction to Pharmacology. Third Edition Mannfred A. Hollinger, 2002-11-28 The first edition of Introduction to Pharmacology has over recent years become a highly influential text among students wishing to acquire a knowledge of pharmacology without having to refer to the larger more detailed traditional pharmacology volumes This revised and updated second edition contains significant new material to bring the reader up to date with the latest practices and principles in pharmacology Exploring the basic principles in both the therapeutic and toxicological aspects of drug use the book employs contemporary examples of medication supplemented with an increased number of accurate and easy to interpret figures and diagrams Additionally Introduction to Pharmacology presents the important concept of understanding the limitations surrounding the drugs that cure replace physiological inadequacies or treat symptoms and which have led to the system of drug classification The broad

scope of the book also encompasses the role of the FDA drugs in sport and the use of animals for drug experimentation A clear and accessible book Introduction to Pharmacology builds on the strengths of the first edition and is an invaluable reference for all students interested in this subject The COVID-19 Aftermath Nima Rezaei, 2024-09-16 This book discusses different challenges imposed to the globe following the COVID 19 pandemic Coronavirus disease 2019 COVID 19 has affected humans individual lives communities and the world It has not only affected human s lives but also environmental and natural systems To better appreciate the pandemic s influence multidisciplinary and interdisciplinary approaches are needed Also lessons learned from facing the pandemic are reviewed to be used for combating the upcoming challenges in healthcare settings mental and psychological health education natural resources energy system environment balance economic stability social relations etc National Theatre Supply Company V. Da Lite Screen Company, Inc ,1936 Impact of AI and Data Science in Response to Coronavirus Pandemic Sushruta Mishra, Pradeep Kumar Mallick, Hrudaya Kumar Tripathy, Gyoo-Soo Chae, Bhabani Shankar Prasad Mishra, 2021-07-22 The book presents advanced AI based technologies in dealing with COVID 19 outbreak and provides an in depth analysis of variety of COVID 19 datasets throughout globe It discusses recent artificial intelligence based algorithms and models for data analysis of COVID 19 symptoms and its possible remedies It provides a unique opportunity to present the work on state of the art of modern artificial intelligence tools and technologies to track and forecast COVID 19 cases It indicates insights and viewpoints from scholars regarding risk and resilience analytics for policy making and operations of large scale systems on this epidemic A snapshot of the latest architectures frameworks in machine learning and data science are also highlighted to gather and aggregate data records related to COVID 19 and to diagnose the virus It delivers significant research outcomes and inspiring new real world applications with respect to feasible AI based solutions in COVID 19 outbreak In addition it discusses strong preventive measures to control such pandemic Nitrogen Compounds: Advances in Research and Application: 2011 Edition ,2012-01-09 Nitrogen Compounds Advances in Research and Application 2011 Edition is a ScholarlyEditions eBook that delivers timely authoritative and comprehensive information about Nitrogen Compounds The editors have built Nitrogen Compounds Advances in Research and Application 2011 Edition on the vast information databases of ScholarlyNews You can expect the information about Nitrogen Compounds in this eBook to be deeper than what you can access anywhere else as well as consistently reliable authoritative informed and relevant The content of Nitrogen Compounds Advances in Research and Application 2011 Edition has been produced by the world's leading scientists engineers analysts research institutions and companies All of the content is from peer reviewed sources and all of it is written assembled and edited by the editors at ScholarlyEditions and available exclusively from us You now have a source you can cite with authority confidence and credibility More information is available at http www ScholarlyEditions com

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