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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

Keith Selby, Robert Giddings

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 Screening 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 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 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 common and rare colorectal conditions advances in the molecular biology and genetics of colorectal diseases and new laparoscopic techniques Contributions from international experts on specialized topics and various new illustrations ensure that the extensive text is not only current and authoritative but easy to understand No other book provides the expertise of a world class editorial team with the cutting edge knowledge you need to master colorectal surgery Photosynthesis Research for Food, Fuel and Future Tingyun Kuang, Congming Lu, Lixin Zhang, 2013-08-31 Photosynthesis is the process by which plants algae and certain species of bacteria transform solar energy into chemical energy in the form of organic molecules In fact all life on the planet ultimately depends on photosynthetic energy conversion The book provides a compressive and state of the art of very recent progress on photosynthesis research

The topics span from atom to intact plants from femtosecond reactions to season long production from physics to agronomy The book is to offer advanced undergraduate students graduate students and research specialists the most recent advances in the all aspects of photosynthesis research The book is intended to offer researchers detailed information on the most recent advances in all aspects of photosynthesis research Tingyun Kuang is a professor at Institute of Botany the Chinese Academy of Sciences CAS and the Academician of CAS Congming Lu is a professor at Institute of Botany CAS Lixin Zhang is a professor at Institute of Botany CAS and the Chief Scientist in the National Basic Research Program of China on G Protein-Coupled Receptors Xinfeng Zhao, Qian Li, Jing Wang, Qi Liang, Jia Quan, 2023-02-20 This book summarizes the progress made to functional immobilize G protein coupled receptors GPCRs through site specific or orientated recognition in both non covalent and covalent manners. The last decade is the dawn of the post structural biology era for G protein coupled receptor research As an emerging approach for state of the art immobilization this book discusses efforts to explore the elegance of naturally occurring biochemical reactions by using their 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 teachers of relevant majors in colleges and universities Chemical & Metallurgical Engineering Eugene Franz 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 machine and deep learning methods Presents experimental case studies **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

This book delves into Screening The Novel. Screening The Novel is a vital topic that must be grasped by everyone, from students and scholars to the general public. This book will furnish comprehensive and in-depth insights into Screening The Novel, encompassing both the fundamentals and more intricate discussions.

- 1. This book is structured into several chapters, namely:
 - Chapter 1: Introduction to Screening The Novel
 - Chapter 2: Essential Elements of Screening The Novel
 - Chapter 3: Screening The Novel in Everyday Life
 - Chapter 4: Screening The Novel in Specific Contexts
 - ∘ Chapter 5: Conclusion
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- 4. In chapter 3, the author will examine the practical applications of Screening The Novel in daily life. The third chapter will showcase real-world examples of how Screening The Novel can be effectively utilized in everyday scenarios.
- 5. In chapter 4, the author will scrutinize the relevance of Screening The Novel in specific contexts. The fourth chapter will explore how Screening The Novel is applied in specialized fields, such as education, business, and technology.
- 6. In chapter 5, this book will draw a conclusion about Screening The Novel. The final chapter will summarize the key points that have been discussed throughout the book.
 - The book is crafted in an easy-to-understand language and is complemented by engaging illustrations. This book is highly recommended for anyone seeking to gain a comprehensive understanding of Screening The Novel.

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Table of Contents Screening The Novel

1. Understanding the eBook Screening The Novel

- The Rise of Digital Reading Screening The Novel
- Advantages of eBooks Over Traditional Books
- 2. Identifying Screening The Novel
 - Exploring Different Genres
 - Considering Fiction vs. Non-Fiction
 - Determining Your Reading Goals
- 3. Choosing the Right eBook Platform
 - Popular eBook Platforms
 - Features to Look for in an Screening The Novel
 - User-Friendly Interface
- 4. Exploring eBook Recommendations from Screening The Novel
 - Personalized Recommendations
 - Screening The Novel User Reviews and Ratings
 - Screening The Novel and Bestseller Lists
- 5. Accessing Screening The Novel Free and Paid eBooks
 - Screening The Novel Public Domain eBooks
 - Screening The Novel eBook Subscription Services
 - Screening The Novel Budget-Friendly Options
- 6. Navigating Screening The Novel eBook Formats
 - ePub, PDF, MOBI, and More
 - Screening The Novel Compatibility with Devices
 - Screening The Novel Enhanced eBook Features
- 7. Enhancing Your Reading Experience
 - Adjustable Fonts and Text Sizes of Screening The Novel
 - Highlighting and Note-Taking Screening The Novel
 - Interactive Elements Screening The Novel
- 8. Staying Engaged with Screening The Novel
 - Joining Online Reading Communities
 - Participating in Virtual Book Clubs
 - Following Authors and Publishers Screening The Novel

- 9. Balancing eBooks and Physical Books Screening The Novel
 - Benefits of a Digital Library
 - Creating a Diverse Reading Collection Screening The Novel
- 10. Overcoming Reading Challenges
 - Dealing with Digital Eye Strain
 - Minimizing Distractions
 - Managing Screen Time
- 11. Cultivating a Reading Routine Screening The Novel
 - Setting Reading Goals Screening The Novel
 - Carving Out Dedicated Reading Time
- 12. Sourcing Reliable Information of Screening The Novel
 - Fact-Checking eBook Content of Screening The Novel
 - Distinguishing Credible Sources
- 13. Promoting Lifelong Learning
 - Utilizing eBooks for Skill Development
 - Exploring Educational eBooks
- 14. Embracing eBook Trends
 - Integration of Multimedia Elements
 - Interactive and Gamified eBooks

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 $reclaiming\ the\ spirit\ gay\ men\ and\ women\ come\ to\ terms\ with\ religion$

reconstructing proto-afroasiatic proto-afrasian

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