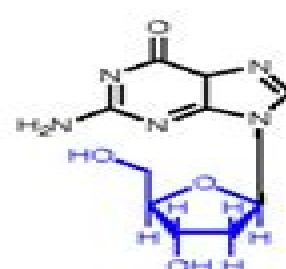


Chapter 6

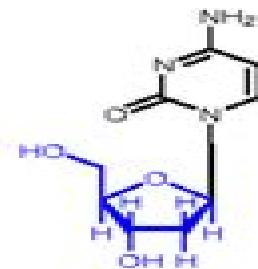
1) Proflavine contains a planar, heteroaromatic, tricyclic ring which could slip between the base pairs of DNA and interact with the base pairs through van der Waals and interactions. The amine groups are likely to be ionised at physiological pH and can form ionic bonds with the charged phosphate groups of the DNA sugar-phosphate backbone (see also section 9.1).

The drug is unlikely to show any selectivity between the DNA of bacteria and the DNA of human cells. Therefore, it cannot be used systemically.

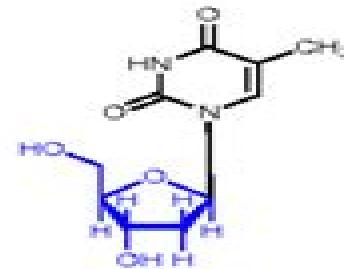
2) The nucleosides which are mimicked are deoxyguanosine, deoxycytidine, and deoxythymidine respectively (see also section 20.6.1)



Deoxyguanosine



Deoxycytidine



Deoxythymidine

3) It is possible to identify five CN fragments within the skeleton of adenine as shown below.



4) The table in appendix 2 can be used to answer this question. A change in Z is least likely to result in a change of amino acid. For example, CUU, CUC, CUA, and CUG all code for leucine.

5)

AGU to ACU

This changes serine to threonine. Both these molecules contain an alcohol functional group in their side chain, and so it is possible that the receptor would still be functional. The increased bulk of threonine might have an effect on the range of ligands that could bind.

Answers Patrick Medicinal Chemistry

AW Rasmussen

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Table of Contents Answers Patrick Medicinal Chemistry

1. Understanding the eBook Answers Patrick Medicinal Chemistry
 - The Rise of Digital Reading Answers Patrick Medicinal Chemistry
 - Advantages of eBooks Over Traditional Books
2. Identifying Answers Patrick Medicinal Chemistry
 - 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 Answers Patrick Medicinal Chemistry
 - User-Friendly Interface
4. Exploring eBook Recommendations from Answers Patrick Medicinal Chemistry
 - Personalized Recommendations
 - Answers Patrick Medicinal Chemistry User Reviews and Ratings
 - Answers Patrick Medicinal Chemistry and Bestseller Lists
5. Accessing Answers Patrick Medicinal Chemistry Free and Paid eBooks
 - Answers Patrick Medicinal Chemistry Public Domain eBooks
 - Answers Patrick Medicinal Chemistry eBook Subscription Services
 - Answers Patrick Medicinal Chemistry Budget-Friendly Options
6. Navigating Answers Patrick Medicinal Chemistry eBook Formats

- ePub, PDF, MOBI, and More
 - Answers Patrick Medicinal Chemistry Compatibility with Devices
 - Answers Patrick Medicinal Chemistry Enhanced eBook Features
7. Enhancing Your Reading Experience
- Adjustable Fonts and Text Sizes of Answers Patrick Medicinal Chemistry
 - Highlighting and Note-Taking Answers Patrick Medicinal Chemistry
 - Interactive Elements Answers Patrick Medicinal Chemistry
8. Staying Engaged with Answers Patrick Medicinal Chemistry
- Joining Online Reading Communities
 - Participating in Virtual Book Clubs
 - Following Authors and Publishers Answers Patrick Medicinal Chemistry
9. Balancing eBooks and Physical Books Answers Patrick Medicinal Chemistry
- Benefits of a Digital Library
 - Creating a Diverse Reading Collection Answers Patrick Medicinal Chemistry
10. Overcoming Reading Challenges
- Dealing with Digital Eye Strain
 - Minimizing Distractions
 - Managing Screen Time
11. Cultivating a Reading Routine Answers Patrick Medicinal Chemistry
- Setting Reading Goals Answers Patrick Medicinal Chemistry
 - Carving Out Dedicated Reading Time
12. Sourcing Reliable Information of Answers Patrick Medicinal Chemistry
- Fact-Checking eBook Content of Answers Patrick Medicinal Chemistry
 - Distinguishing Credible Sources
13. Promoting Lifelong Learning
- Utilizing eBooks for Skill Development
 - Exploring Educational eBooks
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 - Interactive and Gamified eBooks

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