

Chapter 3: Modeling with First-Order Differential Equations Project for Section 3.1
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Project for Section 3.1

Is AIDS an Invariably Fatal Disease?

by Ivan Kramer

This essay will address and answer the question: Is the acquired immunodeficiency syndrome (AIDS), which is the end stage of the human immunodeficiency virus (HIV) infection, an invariably fatal disease?

Like other viruses, HIV has no metabolism and cannot reproduce itself outside of a living cell. The genetic information of the virus is contained in two identical strands of RNA. To reproduce, HIV must use the reproductive apparatus of the cell it invades and infects to produce exact copies of the viral RNA. Once it penetrates a cell, HIV transcribes its RNA into DNA using an enzyme (reverse transcriptase) contained in the virus. The double-stranded viral DNA migrates into the nucleus of the invaded cell and is inserted into the cell's genome with the aid of another viral enzyme (integrase). The viral DNA and the invaded cell's DNA are then integrated, and the cell is infected. When the infected cell is stimulated to reproduce, the proviral DNA is transcribed into viral DNA, and new viral particles are synthesized. Since anti-retroviral drugs like zidovudine inhibit the HIV enzyme reverse transcriptase and stop proviral DNA chain synthesis in the laboratory, these drugs, usually administered in combination, slow down the progression to AIDS in those that are infected with HIV (hosts).

What makes HIV infection so dangerous is the fact that it fatally weakens a host's immune system by binding to the CD4 molecule on the surface of cells vital for defense against disease, including T-helper cells and a subpopulation of natural killer cells. T-helper cells (CD4 T-cells, or T4 cells) are arguably the most important cells of the immune system since they organize the body's defense against antigens. Modeling suggests that HIV infection of natural killer cells *makes it impossible for even modern antiretroviral therapy to clear the virus [1]*. In addition to the CD4 molecule, a virion needs at least one of a handful of coreceptor molecules (e.g., CCR5 and CXCR4) on the surface of the target cell in order to be able to bind to it, penetrate its membrane, and infect it. Indeed, about 1% of Caucasians lack coreceptor molecules, and, therefore, are completely *immune* to becoming HIV infected.

Once infection is established, the disease enters the acute infection stage, lasting a matter of weeks, followed by an *incubation period*, which can last two decades or more! Although

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Table of Contents Cengage Access Project 3 Answers

1. Understanding the eBook Cengage Access Project 3 Answers
 - The Rise of Digital Reading Cengage Access Project 3 Answers
 - Advantages of eBooks Over Traditional Books
2. Identifying Cengage Access Project 3 Answers
 - 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 Cengage Access Project 3 Answers
 - User-Friendly Interface
4. Exploring eBook Recommendations from Cengage Access Project 3 Answers

- Personalized Recommendations
 - Cengage Access Project 3 Answers User Reviews and Ratings
 - Cengage Access Project 3 Answers and Bestseller Lists
5. Accessing Cengage Access Project 3 Answers Free and Paid eBooks
 - Cengage Access Project 3 Answers Public Domain eBooks
 - Cengage Access Project 3 Answers eBook Subscription Services
 - Cengage Access Project 3 Answers Budget-Friendly Options
 6. Navigating Cengage Access Project 3 Answers eBook Formats
 - ePub, PDF, MOBI, and More
 - Cengage Access Project 3 Answers Compatibility with Devices
 - Cengage Access Project 3 Answers Enhanced eBook Features
 7. Enhancing Your Reading Experience
 - Adjustable Fonts and Text Sizes of Cengage Access Project 3 Answers
 - Highlighting and Note-Taking Cengage Access Project 3 Answers
 - Interactive Elements Cengage Access Project 3 Answers
 8. Staying Engaged with Cengage Access Project 3 Answers
 - Joining Online Reading Communities
 - Participating in Virtual Book Clubs
 - Following Authors and Publishers Cengage Access Project 3 Answers
 9. Balancing eBooks and Physical Books Cengage Access Project 3 Answers
 - Benefits of a Digital Library
 - Creating a Diverse Reading Collection Cengage Access Project 3 Answers
 10. Overcoming Reading Challenges
 - Dealing with Digital Eye Strain
 - Minimizing Distractions
 - Managing Screen Time
 11. Cultivating a Reading Routine Cengage Access Project 3 Answers
 - Setting Reading Goals Cengage Access Project 3 Answers
 - Carving Out Dedicated Reading Time
 12. Sourcing Reliable Information of Cengage Access Project 3 Answers

- Fact-Checking eBook Content of Cengage Access Project 3 Answers
 - 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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