CHAPTER 19

THE ORGANIZATION AND CONTROL OF EUKARYOTIC GENOMES

I. Introduction

- A. Gene expression in eukaryotes has two main differences from the same process in prokaryotes.
 - 1. First, the typical multicellular eukaryotic genome is much larger than that of a bacterium.
 - Second, cell specialization limits the expression of many genes to specific cells.
- B. The estimated 35,000 genes in the human genome includes an enormous amount of DNA that does not program the synthesis of RNA or protein.
- C. This DNA is elaborately organized.
 - Not only is the DNA associated with protein to form chromatin, but the chromatin is organized into higher organizational levels.
- D. Level of packing is one way that gene expression is regulated.
 - Densely packed areas are inactivated.
 - Loosely packed areas are being actively transcribed.

II. Eukaryotic Chromatin Structure

- A. Chromatin structure is based on successive levels of DNA packing
 - While the single circular chromosome of bacteria is coiled and looped in a complex, but orderly
 manner, eukaryotic chromatin is far more complex.
 - Eukaryotic DNA is precisely combined with large amounts of protein.
 - 3. During interphase of the cell cycle, chromatin fibers are usually highly extended within the nucleus.
 - Eukaryotic chromosomes contain an enormous amount of DNA relative to their condensed length.
 - Each human chromosome averages about 2 x 10 nucleotide pairs.
 - b) If extended, each DNA molecule would be about 6 cm long, thousands of times longer than the cell cliameter.
 - c) This chromosome and 45 other human chromosomes fit into the nucleus.
 - d) This occurs through an elaborate, multilevel system of DNA packing.
 - Histone proteins are responsible for the first level of DNA packaging.
 - a.) Their positively charged amino acids bind tightly to negatively charged DNA.
 - b) The five types of histones are very similar from one eukaryote to another and are even present in bacteria.
 - Unfolded chromatin has the appearance of beads on a string, a nucleosome, in which DNA winds around a core of histone proteins.
 - The beaded string seems to remain essentially intact throughout the cell cycle.
 - 8. Histones leave the DNA only transiently during DNA replication.
 - They stay with the DNA during transcription.
 - a) By changing shape and position, nucleosomes allow RNA-synthesizing polymerases to move along the DNA.
 - As chromosomes enter mitosis the beaded string undergoes higher-order packing.
 - 11. The beaded string coils to form the 30-nm chromatin fiber.
 - This fiber forms looped domains attached to a scaffold of nonhistone proteins.
 - In a mitotic chromosome, the looped domains coil and fold to produce the characteristic metaphase chromosome.
 - 14. These packing steps are highly specific and precise with particular genes located in the same places.

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Frank R. Spellman, Joanne E. Drinan

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Table of Contents Prentice Hall Biology Chapter 10

- 1. Understanding the eBook Prentice Hall Biology Chapter 10
 - o The Rise of Digital Reading Prentice Hall Biology Chapter 10
 - Advantages of eBooks Over Traditional Books
- 2. Identifying Prentice Hall Biology Chapter 10
 - Exploring Different Genres
 - $\circ\,$ Considering Fiction vs. Non-Fiction
 - Determining Your Reading Goals
- 3. Choosing the Right eBook Platform
 - Popular eBook Platforms
 - Features to Look for in an Prentice Hall Biology Chapter 10
 - User-Friendly Interface
- 4. Exploring eBook Recommendations from Prentice Hall Biology Chapter 10
 - Personalized Recommendations

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

- 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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