# <<基础分子生物学>>

#### 图书基本信息

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### <<基础分子生物学>>

#### 前言

The fast pace of modern molecular biology research is driven by intellectual curiosity and major challenges in medicine, agriculture, and industry. No discipline in biology has ever experienced the explosion in growth and popularity that molecular biology is now undergoing. There is intense public interest in the Human Genome Project and genetic engineering, due in part to fascination with how our own genes influence our lives. With this fast pace of discovery, it has been difficult to find a suitable, up-to-date textbook for a course in molecular biology. Other textbooks in the field flail into two categories: they are either too advanced, comprehensive, and overwhehmingly detailed, with enough material to fill an entire year or more of lectures, or they are too basic, superficial, and less experhmental in their approach. It is possible to piece together literature for a molecular biology course by assigning readings from a variety of sources. However, some students are poorly prepared to learn material strictly from lectures and selected readings in texts and the primary literature that do not match exactly the content of the course. At the other end, instructors may find it difficult to decide what topics are the most important to include in a course and what to exclude when presented with an extensive array of choices. This textbook aims to fill this perceived gap in the market. The intent is to keep the text to a manageable size while covering the essentials of molecular biology. Selection of topics to include or omit reflects my view of molecular biology and it is possible that some particular favorite topic may not be covered to the desired extent. Students often complain when an instructor teaches "straight from the textbook," so adding favorite examples is encouraged to allow instructors to enrich their course by bringing to it their own enthusiasm and insight. **Approach** central theme of the textbook is the continuum of biological understanding, starting with basic properties of genes and genomes, RNA and protein structure and function, and extending to the complex, hierarchical interactions fundamental to living organisms. A comprehensive picture of the many ways molecular biology is being applied to the analysis of complex systems is developed, including advances that reveal fundamental features of gene regulation during cell growth and differentiation, and in response to a changing nvironment, as well as developments that are more related to commercial and medical applications. Recent advances in technology, the process and thrill of discovery, and ethical considerations in molecular biology research are emphasized. text highlights the process of discovery - the observations, the questions, the experimental designs totest models, the results and conclusions - not just presenting the "facts." At the same time the language of molecular biology is emphasized, and a foundation is built that is based in fact. It is not feasible to examine every brick in the foundation and still have time to view the entire structure. However, as often as possible real examples of data are shown, e.g. actual results of an EMSA, Western blot, or RNA splicing assay. Experiments are selected either because they are classics in the field or because they illustrate a particular approach frequently used by molecular biologists to answer a diversity of questions.

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#### 内容概要

The beginnings of molecular biology, The structure of DNA, Genome organization: from nucleotides to chromatin, The versatility of RNA, From gene to protein, DNA replication and telomere maintenance, DNA repair and recombination, Recombinant DNA technology and molecular cloning.

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#### 书籍目录

- 1 The beginnings of molecular biology
- 1.1 Introduction
- 1.2 Historical perspective

Insights into heredity from round and wrinkled peas: Mendelian genetics

Insights into the nature of hereditary material: the transforming principle is DNA

Creativity in approach leads to the one gene-one enzyme hypothesis

The importance of technological advances: the Hershey-Chase experiment

A model for the structure of DNA: the DNA double helix

Chapter summary

Analytical questions

Suggestions for further reading

- 2 The structure of DNA
- 2.1 Introduction
- 2.2 Primary structure: the components of nucleic acids

Five-carbon sugars

Nitrogenous bases

The phosphate functional group

Nucleosides and nucteotides

- 2.3 Significance of 5 and 3
- 2.4 Nomenclature of nucleotides
- 2.5 The length of RNA and DNA
- 2.6 Secondary structure of DNA

Hydrogen bonds form between the bases

Base stacking provides chemical stability to the DNA double helix

Structure of the Watson-Crick DNA double helix

Distinguishing between features of alternative double-helical structures

DNA can undergo reversible strand separation

2.7 Unusual DNA secondary structures

Slipped structures

Cruciform structures

Triple helix DNA

Disease box 2.1 Friedreichs ataxia and triple helix DNA

2.8 Tertiary structure of DNA

Supercoiling of DNA

Topoisomerases relax supercoiled DNA

What is the significance of supercoiting in vivo?

Disease box 2.2 Topoisomerase-targeted anticancer drugs

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3.1 Introduction

3.2 Eukaryotic genome

Chromatin structure: historical perspective

Histones Nucleosomes

Beads-on-a-string: the 10 nm fiber

The 30 nm fiber Loop domains

Metaphase chromosomes

Alternative chromatin structures

3.3 Bacterial genome

3.4 Plasmids

3.5 Bacteriophages and mammalian DNA viruses

Bacteriophages

Mammalian DNA Viruses

3.6 Organelle genomes: chloroplasts and mitochondria

Chloroplast DNA(cpDNA)

Mitochondrial DNA (mtDNA)

Disease box 3.1 Mitochondrial DNA and disease

3.7 RNA-based genomes

Eukaryotic RNA viruses

Retroviruses

Viroids

Other Subviral pathogens

Disease box 3.2 Avian flu

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4 The versatility of RNA

4.1 Introduction

4.2 Secondary structure of RNA

Secondary structure motifs in RNA

Base-paired RNA adopts an A-type double helix

RNA helices often contain noncanonical base pairs

4.3 Tertiary structure of RNA

tRNA structure: important insights into RNA structural motifs

Common tertiary structure motifs in RNA

4.4 Kinetics of RNA folding

4.5 RNA is involved in a wide range of cellular processes

4.6 Historical perspective: the discovery of RNA catalysis

Tetrahymena qroUP I intron ribozyme

RNase P ribozyme

Focus box 4.1: The RNA World

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Mode of ribozyme action

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Large ribozymes

Small ribozymes

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5 From gene to protein

5.1 Introduction

5.2 The central dogma

5.3 The genetic code

Translating the genetic code

The 21st and 22nd genetically encoded amino acids

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5.4 Protein structure

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Secondary structure

Tertiary structure

Quaternary structure

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5.5 Protein function

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modifications

Allosteric regulation of protein activity

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Macromolecular assemblages

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Disease box 5.1 Prions

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6 DNA replication and telomere maintenance

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6.2 Historical perspective

Insight into the mode of DNA replication: the Meselson-Stahl

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Insight into the mode of DNA replication: visualization of

replicating bacterial DNA

6.3 DNA synthesis occurs from 5 3

6.4 DNA polymerases are the enzymes that catalyze DNA

synthesis

Focus box 6.1 Bacterial DNA polymerases

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6.5 Semidiscontinuous DNA replication

Leading strand synthesis is continuous

Lagging strand synthesis is discontinuous

6.6 Nuclear DNA replication in eukaryotic cells

Replication factories

Histone removal at the origins of replication

Prereplication complex formation at the origins of

replication

Replication Licensing: DNA only replicates once per cell

cycle

Duplex unwinding at replication forks

RNA priming of Leading strand and Lagging strand DNA

synthesis

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Maturation of nascent DNA strands

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Focus box 6.2 The naming of genes involved in DNA replication

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6.7 Replication of organelle DNA

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Replication of cpDNA

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6.8 Rolling circle replication

6.9 Tetomere maintenance: the role of tetomerase in DNA

replication, aging, and cancer

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Solution to the end replication problem

Maintenance of telomeres by telomerase

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Telomerase, aging, and cancer

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function

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7 DNA repair and recombination

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7.2 Types of mutations and their phenotypic consequences

Transitions and transversions can lead to silent, missense, or

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Expansion of trinucleotide repeats leads to genetic

instability

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7.3 General classes of DNA damage

Single base changes

Structural distortion

DNA backbone damage

Cellular response to DNA damage

7.4 Lesion bypass

7.5 Direct reversal of DNA damage

7.6 Repair of single base changes and structural distortions by

removal of DNA damage

Base excision repair

Mismatch repair

Nucleotide excision repair

Disease box 7.1 Hereditary nonpolyposis colorectal cancer: a defect

in mismatch repair

7.7 Double-strand break repair by removal of DNA damage

Homologous recombination

Nonhomologous end-joining

Disease box 7.2 Xeroderma pigmentosum and related disorders:

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Disease box 7.3 Hereditary breast cancer syndromes: mutations in

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8 Recombinant DNA technology and molecular cloning

9 Tools for analyzing gene expression

10 Transcription in prokaryotes

11 Transcription in eukaryotes

12 Epigenetic and monoallelic gene expression

13 RNA processing and post-transcriptional gene regulation

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16 Genome analysis: DNA typing, genomics and beyond

17 Medical molecular biology

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#### 章节摘录

Other subviral pathogens Other subviral pathogens include satellite RNAs and virusoids. Viroids replicate autonomously by using host-encoded RNA polymerase. In contrast, satellite RNAs multiply only in the presence of a helper virus that provides the appropriate RNA-dependent RNA polymerase. Some of the larger satellite RNAs may encode a protein. Satellite RNAs are found in plants (e.g. satellite tobacco necrosis virus) and animals. A well known human satellite RNA is hepatitis delta virus (HDV). HDV is a small single-stranded A virusoid is an RNA molecule that does not encode any proteins and RNA satellite of hepatitis B virus. depends on a helper virus for replication and capsid formation. Virusoids occur in association with viruses causing plant diseases such as velvet tobacco mottle and subterranean clover mottle. They are sometimes regarded as a subtype of satellite RNA. The virusoid genome resembles a viroid and consists of circular, single-stranded RNA with self-cleaving activity (see Section 4.7). Chapter summary The genomes of most organisms are made of DNA; certain viruses and subviral pathogens have RNA genomes. Eukaryotic DNA combines with basic protein molecules called histones to form structures known as nucleosomes. Each nucleosome contains four pairs of core histones (H2A, H2B, H3, and H4) in a wedge-shaped disk, around which is wrapped 146 bp of DNA. The linker histone H1 is bound to DNA between the core histone octamers, where the DNA enters and exits the nucleosome. The first order of chromatin folding is represented by a string of nucleosomes. This 10 mn nucleosome fiber is further folded into a 30 nm fiber in a zig-zag ribbon structure, which is then folded into loop domains, and finally the metaphase chromosome. Each chromosome is composed of one linear, double-stranded Bacterial chromosomal DNA exists as one double-stranded, circular DNA molecule DNA molecule. organized into a condensed structure called a nucleoid. Plasmids are self-rephcating small, double-stranded, circular or linear DNA molecules carried by bacteria, some fungi, and some higher plants. Plasmids are important tools for recombinant DNA technology. Bacteriophages and mammalian DNA viruses have DNA genomes that occur in a variety of forms, ranging from double-stranded to single-stranded DNA and linear to circular forms. Viruses either package their genomes with their own basic proteins, or use host cell histones. **Both** mitochondria and chloroplasts contain their own genetic information. The small, double-stranded DNA genomes are usually, but not always, circular and there are multiple copies per organdie. Organelle genomes are maternally inherited.

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