Biology

Concepts and Applications | 9e Starr | Evers | Starr

Chapter 8

DNA Structure and Function

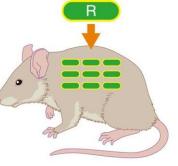
8.1 How Was DNA's Function Discovered?

- The substance we now call DNA was first described in 1869 by Johannes Miescher
- Miescher determined that DNA is not a protein, and that it is rich in nitrogen and phosphorus
 - He never learned of its function

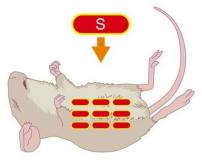


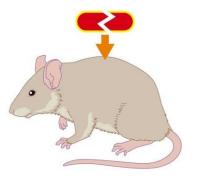
Patrick Landmann/Science Source

- Sixty years after Miescher's work, Frederick Griffith unexpectedly uncovered a clue about DNA's function
 - Heat destroyed the ability of lethal S bacteria to cause pneumonia, but it did not destroy their hereditary material
 - The hereditary material could be transferred from dead S cells to live R cells



A Griffith's first experiment showed that R cells were harmless. When injected into mice, the bacteria multiplied, but the mice remained healthy.





B The second experiment showed that an injection of S cells caused mice to develop fatal pneumonia. Their blood contained live S cells.

C For a third experiment, Griffith killed S cells with heat before injecting them into mice. The mice remained healthy, indicating that the heat-killed S cells were harmless.

D In his fourth experiment, Griffith injected a mixture of heat-killed S cells and live R cells. To his surprise, the mice became fatally ill, and their blood contained live S cells.

- In 1940, Oswald Avery and Maclyn McCarty identified that the "transforming principle" was a nucleic acid
 - Lipid- and protein-destroying enzymes did not block the S cell's transformation of R cells
 - DNA-degrading enzymes, but not RNAdegrading enzymes, prevented transformation
 - They concluded that DNA must be the transforming principle

- Essential properties of hereditary material:
 - A full complement of hereditary information must be transmitted along with the molecule
 - An equal amount of hereditary material must be found in each cell of a given species
 - The hereditary material must not change
 - The hereditary material must be capable of encoding the enormous amount of information required to build a new individual

- In the late 1940s, Alfred Hershey and Martha Chase established that DNA transmits a full complement of hereditary information
 - They established that the material bacteriophage (a virus that infects bacteria) injects into bacteria is DNA, not protein

ANIMATION: The Hershey and Chase Experiments

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- In 1948, André Boivin and Roger Vendrely established that body cells of any individual of a species contain precisely the same amount of DNA
- Daniel Mazia's laboratory discovered that DNA content does not change over time
 - Established that DNA is not involved in metabolism

8.2 How Was DNA's Structure Discovered?

- Building blocks of DNA
 - DNA is a polymer of nucleotides, each with a five-carbon sugar, three phosphate groups, and one of four nitrogen-containing bases

Animation: Subunits of DNA

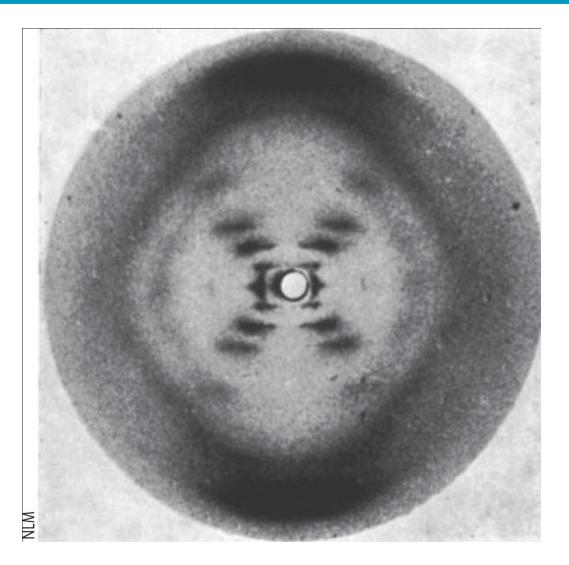
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- 1950: Erwin Chargaff made two important discoveries about DNA
 - Chargaff 's first rule: the amounts of thymine and adenine are identical, as are the amounts of cytosine and guanine (A = T and G = C)
 - Chargaff 's second rule: DNA of different species differs in its proportions of adenine and guanine

- 1950s: James Watson and Francis Crick suspect that DNA is a helix
 - Made models from scraps of metal connected by suitably angled "bonds" of wire
- Rosalind Franklin made the first clear xray diffraction image of DNA as it occurs in cells
 - She calculated that DNA is very long and identified a repeating pattern





- Structure of DNA helix:
 - Two sugar-phosphate chains running in opposite directions, and paired bases inside
 - Bonds between the sugar of one nucleotide and the phosphate of the next form the backbone of each chain (or strand)

- Structure of DNA helix (cont'd.):
 - Hydrogen bonds between the internally positioned bases hold the two strands together
 - Only two kinds of base pairings form (supports Chargaff 's first rule):
 - A to T
 - G to C

Animation: DNA close up

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DNA's Base Sequence

- The two strands of DNA match
 - The strands are complementary: the base of each nucleotide on one strand pairs with a suitable partner base on the other
 - The base-pairing patterns A to T and G to C
 is the same in all molecules of DNA

DNA's Base Sequence (cont'd.)



DNA's Base Sequence (cont'd.)

- How can just two kinds of base pairings give rise to the incredible diversity of traits we see among living things?
 - The order of nucleotides in a strand of DNA (DNA sequence) varies tremendously among species
 - Explains Chargaff 's second rule

DNA's Base Sequence (cont'd.)

- DNA molecules can be hundreds of millions of nucleotides long
 - So their sequence can encode a massive amount of information
- DNA is the basis of life's unity
 - Variations in its nucleotide sequence are the foundation of life's diversity; defines species and distinguishes individuals

8.3 What Is a Chromosome?

- DNA in a single human cell is about 2 meters (6.5 feet) long
- How can that much DNA pack into a nucleus that is less than 10 micrometers in diameter?
 - Proteins associate with the DNA and help keep it organized

What Is a Chromosome? (cont'd.)

- Chromosome: structure that consists of DNA and associated proteins
 - Carries part or all of a cell's genetic information
- Histone: type of protein that structurally organizes eukaryotic chromosomes
- Nucleosome: a length of DNA wound twice around a spool of histone proteins

ANIMATION: Chromosome structural organization

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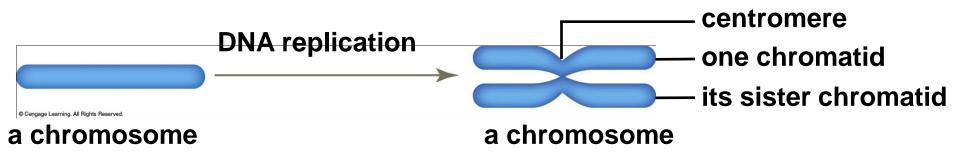
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What Is a Chromosome? (cont'd.)

- During most of a cell's life, each chromosome consists of one DNA molecule
- When the cell prepares to divide, it duplicates its chromosomes by DNA replication
 - After replication, each chromosome consists of two DNA molecules (*sister chromatids*) that attach at a *centromere* region

What Is a Chromosome? (cont'd.)



Chromosome Number and Type

- Each species has a characteristic chromosome number (number of chromosomes in its cells)
 - Examples:
 - The chromosome number of oak trees is 12
 - The chromosome number of humans is 46

- Human body cells have two sets of 23
 chromosomes—two of each type
 - Having two sets of chromosomes means these cells are *diploid*
- Karyotype: an image of an individual's diploid set of chromosomes

ANIMATION: Karyotype preparation

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- Autosome: a chromosome that is the same in males and females
 - Two autosomes of a pair have the same length, shape, and centromere location
 - They hold information about the same trait

- Members of a pair of sex chromosomes differ between females and males
 - The body cells of typical human females have two X chromosomes (XX)
 - The body cells of typical human males have one X and one Y chromosome (XY)
- Environmental factors (not sex chromosomes) determine sex in some invertebrates and reptiles



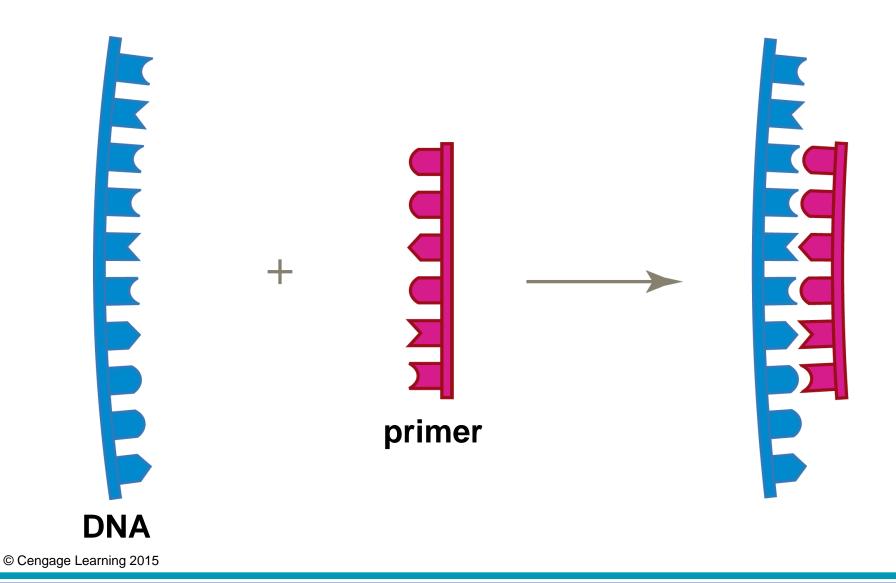
8.4 How Does a Cell Copy Its DNA?

- In preparation for division, a cell copies its chromosomes so that it contains two sets
 - The process by which a cell copies its DNA is called DNA replication

Semiconservative Replication

- Before DNA replication, a chromosome consists of one molecule of DNA (one double helix)
- As replication begins, enzymes break the hydrogen bonds that hold the double helix together
 - The two DNA strands unwind and separate

- Another enzyme constructs *primers*: short, single strands of nucleotides
 - Primers serve as attachment points for DNA polymerase, the enzyme that assembles new strands of DNA
 - A primer base-pairs with a complementary strand of DNA



- The establishment of base-pairing between two strands of DNA is called *nucleic acid hybridization*
 - Hybridization is spontaneous, driven by hydrogen bonding between bases of complementary strands
- DNA polymerases attach to the hybridized primers and begin DNA synthesis

- Each nucleotide provides energy for its own attachment to the end of a growing strand of DNA
 - Two of the three phosphate groups are removed when a nucleotide is added to a DNA strand
- The enzyme *DNA ligase* seals any gaps, so the new DNA strands are continuous

- Both of the two strands of the parent molecule are copied at the same time
- As each new DNA strand lengthens, it winds up with its template strand into a double helix
- Semiconservative replication produces two copies of a DNA molecule: one strand of each copy is new, and the other is parental

ANIMATION: DNA replication in detail

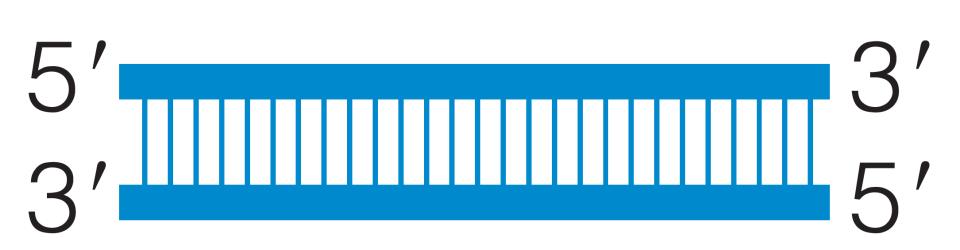
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Directional Synthesis

- Each strand of DNA has two ends
 - The last carbon atom on one end of the strand is a 5' (5 prime) carbon of a sugar
 - The last carbon atom on the other end is a 3' (three prime) carbon of a sugar

Directional Synthesis (cont'd.)

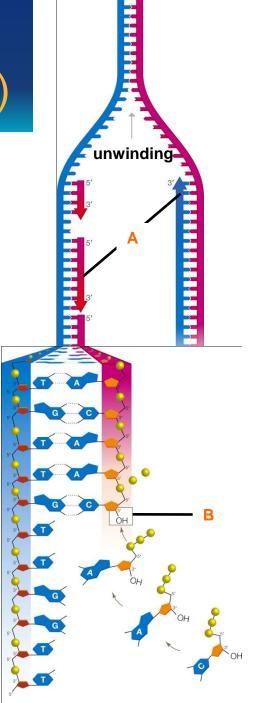


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Directional Synthesis (cont'd.)

- DNA polymerase can attach a nucleotide only to a 3' end
 - DNA synthesis proceeds only in the 5' to 3' direction
- One new strand of DNA is constructed in a single piece during replication
 - Synthesis of the other strand occurs in segments that must be joined by DNA ligase

Directional Synthesis (cont'd.)



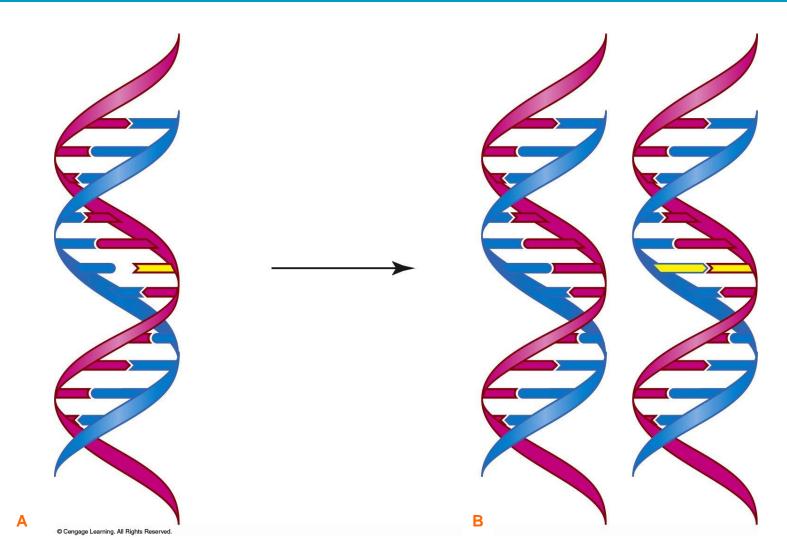
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8.5 What Causes Mutations?

- Mistakes can and do occur during DNA replication
- Examples:
 - The wrong base is added to a growing DNA strand
 - A nucleotide gets lost, or an extra one slips in

- Most replication errors occur because DNA polymerases work very fast
- Luckily, most DNA polymerases also proofread their work
 - They can correct a mismatch by reversing the synthesis reaction to remove the mispaired nucleotide

- Replication errors may occur after a cell's DNA gets broken or damaged
 - DNA polymerases do not copy damaged DNA very well
- When proofreading and repair mechanisms fail, an error becomes a *mutation*
 - A permanent change in the DNA sequence of a cell's chromosome



• Mutations can form in any type of cell

 Those that occur during egg or sperm formation can be passed to offspring

 Mutations that alter DNA's instructions may have a harmful or lethal outcome

- Most cancers begin with a mutation

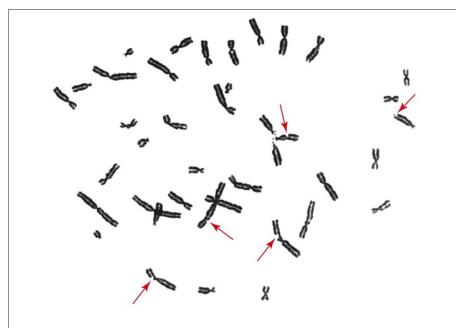
• Not all mutations are dangerous

Some give rise to variation in traits; basis for evolution

Agents of DNA Damage

- Ionizing radiation from x-rays, most UV light, and gamma rays may cause DNA damage:
 - Breaks DNA
 - Causes covalent bonds to form between bases on opposite strands
 - Fatally alters nucleotide bases
 - Causes adjacent nucleotide dimers to form

Agents of DNA Damage (cont'd.)

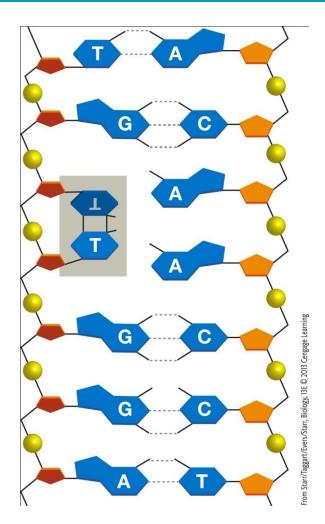


Olga Shovman, Andrew C. Riches, Douglas Adamson, and Peter E. Bryant. An improved assay for radiation-induced chromatid breaks using a colcemid block and calyculin-induced PCC combination. Mutagenesis (2008) 23(4): 267-270 first published online March 6, 2008 doi:10.1093/mutage/gen009, by permission of Oxford University Press



main, Courtesy of Janis Ruksans; inset, Frank Sommariva/image/imagebroker.net/SuperStock.

Agents of DNA Damage (cont'd.)



a thymine dimer

8.6 How Does Cloning Work?

- Cloning: making an identical copy of something
- Reproductive cloning: technology that produces genetically identical individuals
 - Example: artificial embryo splitting

How Does Cloning Work? (cont'd.)

- Animal breeders sometimes want an exact copy of a specific individual
 - Use a cloning method where a somatic cell is taken from an adult organism (contains master blueprint for new individual)
 - An adult somatic cell will not start dividing to produce an embryo because the cell has already *differentiated* (obtained specialized characteristics)

How Does Cloning Work? (cont'd.)

- Somatic cell nuclear transfer (SCNT) can undifferentiate a somatic cell by turning its unused DNA back on
 - An unfertilized egg's nucleus is replaced with the nucleus of a donor's somatic cell
 - The egg's cytoplasm reprograms the transplanted DNA to direct the development of an embryo, which is then implanted into a surrogate mother

ANIMATION: How Dolly was created

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How Does Cloning Work? (cont'd.)



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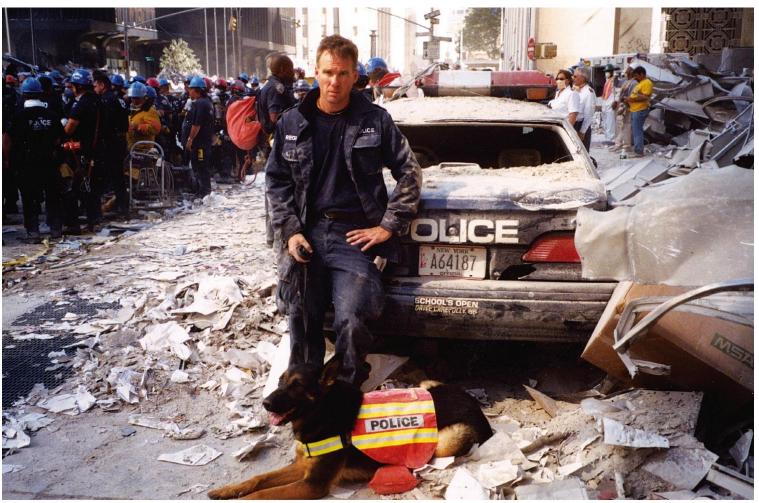
How Does Cloning Work? (cont'd.)

- As techniques become routine, cloning humans is no longer only within the realm of science fiction
- SCNT is already being used to produce human embryos for medical purposes, called *therapeutic cloning*
 - Example: researchers are using SCNT to study how heart defects cause developing heart cells to malfunction

8.7 A Hero Dog's Golden Clones

- Trakr, who died in 2009, was a hero dog who helped rescuers at the World Trade Center on 9/11
 - Through the Golden Clone Giveaway Trakr's DNA was used to make clones
- Cloning animals raises ethical questions about cloning humans
 - Is it acceptable to clone a lost child for a grieving parent?

A Hero Dog's Golden Clones (cont'd.)



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