

**a** The string looped around a pencil and twisted represents a double helix without supercoils.

#### Demonstration of Supercoiling



**b** As the string unwinds, supercoils form ahead of the unwinding if the helix is not allowed to rotate.



c Rotation relieves the tension from supercoiling.

#### Semi discontinuous DNA replication





#### **Restrictions on DNA polymerases**



**b** DNA polymerase cannot extend the nucleotide chain in the  $3' \rightarrow 5'$  direction.



c DNA polymerase cannot initiate synthesis on a template strand without paired nucleotides.

Figure 2.18 Restrictions on DNA polymerases.

## An RNA primer for initiation of DNA synthesis



## Primer removal by DNA polymerase I



**a** DNA polymerase III leaves a nick between the 3' end of the newly sythesized fragment and the 5' end of the RNA primer.



**b** DNA polymerase I binds at the nicked site.



#### Molecular structure of DNA Polymerase III holoenzyme in E.coli



#### Model of DNA replication in Prokaryotes with simultaneous synthesis of leading and lagging strands



leading and lagging strands.

## **Prokaryotic DNA Polymerases**

Table 2.1 Prokaryotic DNA Polymerases		
Polymerase	Functions	
DNA polymerase I	Removal of nucleotides during DNA repair $(5' \rightarrow 3' \text{ exonuclease});$	
	synthesis of DNA during repair;	
	synthesis of short gaps in DNA;	
	primer removal $(5' \rightarrow 3')$ exonuclease);	
	proofreading $(3' \rightarrow 5' \text{ exonuclease})$	
DNA polymerase II	Synthesis of DNA during repair;	
	proofreading $(3' \rightarrow 5' \text{ exonuclease})$	
DNA polymerase III	DNA synthesis; proofreading $(5' \rightarrow 3' \text{ exonuclease})$	

### **Proofreading newly synthesized DNA**



**b** DNA polymerase adds a mispaired nucleotide.



**c** DNA polymerase reverses direction and acts as a  $3' \rightarrow 5'$  exonuclease to remove the mispaired nucleotide.



## **Eukaryotic DNA Polymerases**

Table 2.2 Eukaryotic DNA Polymerases		
Mammalian Polymerase	Corresponding Polymerase in Yeast	Functions
α	pol I	Synthesis of lagging strand; primer synthesis
β	none	Synthesis of DNA during repair
δ	pol III	Synthesis of leading strand; proofreading $(3' \rightarrow 5' \text{ exonuclease})$
3	pol II	Synthesis of DNA during repair; proofreading $(3' \rightarrow 5' \text{ exonuclease})$
γ	mitochondrial DNA polymerase	Mitochondrial DNA synthesis; proofreading $(3' \rightarrow 5' \text{ exonuclease})$

### **Eukaryotic DNA Polymerases**



a The 48 and 58 kD subunits of DNA polymerase α may function on their own as a primase. **b** They may also function as a primase when part of the entire enzyme.

**Figure 2.24** DNA polymerase  $\alpha$  in mammals.

#### Model of DNA replication in Eukaryotes with simultaneous synthesis of leading and lagging strands



# **Origin of replication**



## **Bidirectional replication**



**a** Replication "bubbles" forming from two origins of replication.



**b** Replication bubbles (indicated by arrows) in Drosophila melanogaster DNA.

Figure 2.27 Bidirectional replication. (Photo courtesy of D. S. Hogness.)

Strategy for replicating circular DNA (Theta mode replication in *E. coli*)



#### Electron micrograph of theta mode replication in E. coli





### **Rolling circle mode replication**





# **Linear DNA Replication**



**Figure 2.32** The linear DNA replication paradox. How can the gap at the end of the linear molecule be filled?

#### **D** loop mode replication



#### A model for replicating the ends of linear chromosomes in eukaryotes

