



بسم الله الرحمن الرحيم  
 مادة البيومن أسهل وامتع المواد في هذا العالم  
 لذلك قررنا ان نجعلها أسهل للجميع



شعارنا : دعونا سويا نفك السوبر كويل

\* تعريف بالمذكرة :

- شاملة لجميع المواضيع . ( كاملة )
- تركيز على النقاط الهامة .
- جداول تسهل عملية الحفظ .
- أسئلة عامة .
- ( الحلم أصبح حقيقة )

Team leader : **مجهول**

Team group :

أبويسرا

Blue eye

Dr.noop

Ocean

والشكر للجنود المجهولين الذين فضلوا عدم ذكر أسمائهم



**\* Prokaryotic DNA synthesis (REPLICATION) :**

**\* Semiconservative Replication :**

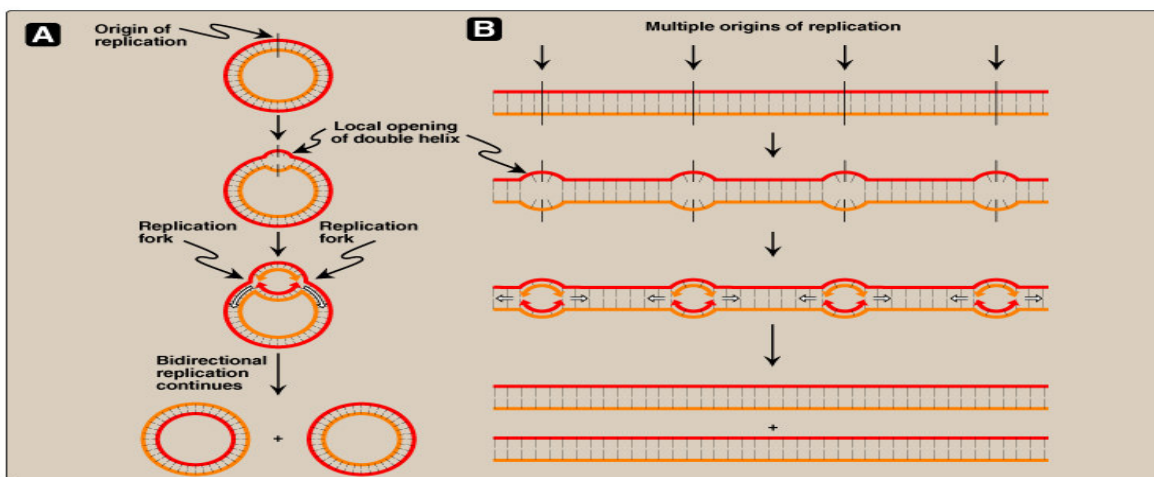
- When the two strands of the DNA double helix are separated, each can serve as a template for the replication of a new complementary strand.
- Each of the individual parental strands remains intact in one of the two new Duplexes . ( T ) . ( important )
- (i.e. one of the parental strands is conserved in each of the two new duplexes) .

**A. Separation of the two complementary DNA strands:**

☐ Before the replication , separation of the two parental strands must occur first why ?  
 Because polymerases use only single – stranded DNA .

You will have a question in this schedule studied hard

<i>Prokaryotes</i>	<i>Eukaryotes</i>
DNA replication begins at <b>single</b> , unique nucleotide sequences (this site is called <b><u>origin of replication</u></b> )	-In eukaryotes , replication begins at <b>multiple</b> sites along the DNA helix this is referred to as <b>short consensus sequence</b> (exclusively composed of <b>AT base pairs</b> ) <b>- multiple origins of replication .</b>
<b>Slower replicating than eukaryotes</b>	<b>Rapidly replicating</b>



**Figure 29.9** Replication of DNA: origins and replication forks. A. Small prokaryotic circular DNA. B. Very long eukaryotic DNA.

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**C. Formation of the replication fork :**

- As the two strands unwind & separate they form a V where active synthesis occurs , this region is called replication fork
- replication fork moves along the DNA molecule as synthesis occurs in both direction (bidirectional)

**1. proteins required for DNA strand separation :**

✓ called **prepriming** complex & include :

( AT least two question from this schedule )

DNA protein	Single-stranded DNA-binding(SSB) proteins	DNA helicases
<ul style="list-style-type: none"> <li>• 20 to 50 monomers bind to specific nucleotide sequences at origin of replication <b>(which rich in AT base)</b></li> <li>• <b>ATP is required</b></li> <li>• Cause <b>double stranded</b> DNA to melt (separate) &amp; , forming <b>localized</b> regions of <b>single</b> stranded DNA .</li> </ul>	<ul style="list-style-type: none"> <li>• Also called <b>helix-destabilizing proteins</b></li> <li>• Bind only to <b>single</b> (هام) stranded DNA</li> <li>• Bind <b>cooperatively</b> (هام) (binding of one molecule make it easier for binding of additional molecules)</li> <li>• Are <b>not enzyme</b> (هام) (but shift the equilibrium between single &amp; double stranded DNA in the direction of single stranded form)</li> <li>• Functions of SSB proteins:                             <ol style="list-style-type: none"> <li>I. Keep the two strands of DNA separated in the area of replication origin ( prevent reformation of double-helix</li> <li>II. Protect the DNA from nucleases (that cleave single stranded DNA)</li> </ol> </li> </ul>	<ul style="list-style-type: none"> <li>• Bind to <b>single stranded</b> DNA near the replication fork , then move to neighboring double stranded region , &amp; unwinding the strands</li> <li>• <b>ATP is required</b></li> <li><b>( be careful DNA helicase Bind to single strand )</b></li> <li>( I never seen an exam without this question )</li> </ul>

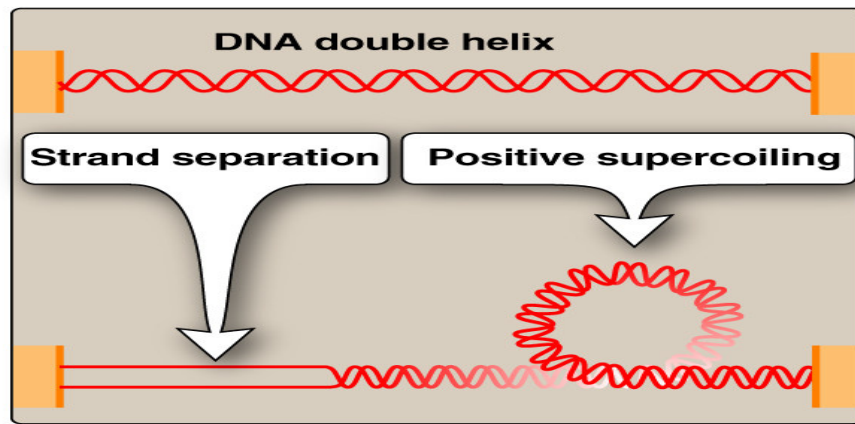


## \* Solving the problem of supercoils 😊 :

☐ **Positive** supercoils (supertwist) in the region of DNA **ahead** ( be careful not before ) of the replication fork interfere with further unwinding of double helix .

### DNA topoisomerases

are responsible for removing these supercoils



**Figure 29.11**  
Positive supercoiling resulting from DNA strand separation.

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DNA topoisomerases I	DNA topoisomerases II
<ul style="list-style-type: none"> <li>Reversibly cut <b>a single strand</b> of double helix . ( MCQ )( NOT double)</li> <li>- <b>(by nuclease then ligase)</b> (هام جدا)</li> <li>* nuclease ( strand – cutting )</li> <li>* ligase ( strand resealing )</li> </ul>	<ul style="list-style-type: none"> <li>-cuts <b>double</b> strands</li> <li>- causes a second stretch of the DNA double helix to pass through the break &amp; finally , reseals the break .</li> </ul>
<ul style="list-style-type: none"> <li><b>They do not require ATP</b> (but store the energy from cleaving of phosphodiester bond)</li> </ul>	USE ATP
<ul style="list-style-type: none"> <li>Reliving (relaxing) of:                             <ul style="list-style-type: none"> <li>✓ Negative supercoils in <b>prokaryotic</b> DNA (e.g. in <i>E. coli</i>)</li> <li>✓ Negative &amp; positive supercoils in <b>eukaryotic</b> DNA</li> </ul> </li> </ul>	<ul style="list-style-type: none"> <li>- Relax both <b>positive &amp; negative supercoils</b></li> <li>-Required for separation of interlocked molecules of DNA following a chromosomal replication (<b>in prokaryotes &amp; eukaryotes</b>)</li> </ul>
<p>-Transient "<b>nick</b>" in one DNA Strand ( NOT Both strand ) (MCQ)</p> <p>- the intact DNA –strand is passed through the break before resealed .</p>	<ul style="list-style-type: none"> <li>- <b>DNA gyrase :</b> (م يخلو منه اختيار )                             <ul style="list-style-type: none"> <li>a) A type II topoisomerase</li> <li>b)found in <i>E.coli</i></li> <li>c) Introduce negative supercoils</li> <li>d)into relaxed circular DNA</li> <li>e)(to facilitates future replication of DNA)</li> <li>f) Use ATP .</li> </ul> </li> <li>- <b>Antimicrobial agents</b> ( bacteria DNA gyrase ), <u>quinolones</u> (as ciprofloxacin)</li> <li>- <b>Anticancer agents : etoposide ( target human topoisomerase II )</b></li> </ul>

● **Negative supercoils :** contain fewer turns of helix than relaxed DNA.

● **Positive supercoils :** contain more turns of helix than relaxed DNA.

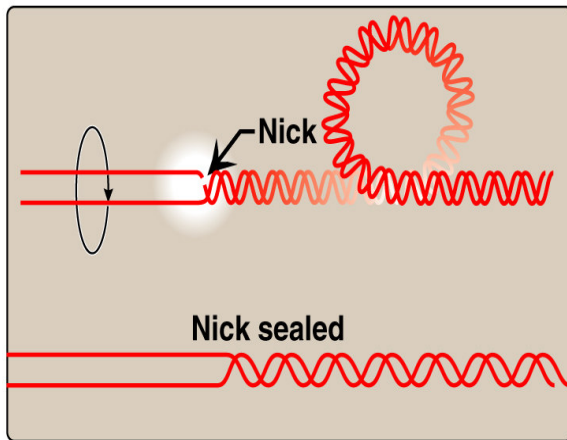


Figure 29.12  
Action of type I DNA topoisomerases.

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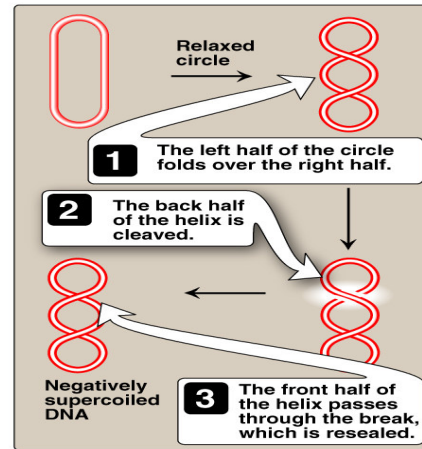


Figure 29.13  
Action of type II DNA topoisomerase.

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## D. Direction of DNA replication:

- DNA polymerases (that responsible for of DNA template are):
  - ✓ Only read parental nucleotide sequences in  $3' \rightarrow 5'$  direction
  - ✓ They synthesized the new DNA in  $5' \rightarrow 3'$  antiparallel direction

■ So, beginning with one prenatal double helix, the two newly synthesized DNA must grow in opposite directions:

- 1- One in  $5' - 3'$  direction towards the replication fork
- 2- One in  $5' - 3'$  direction away from the replication fork.

### 1. Leading strand:

- copied in  $5' \rightarrow 3'$  direction , toward the replication fork
- synthesized almost continuously ( MCQ )

### 2. Lagging strand:

- copied in  $5' \rightarrow 3'$  direction , away from the replication fork
- synthesized discontinuously with small fragments of DNA near the replication fork , is called okazaki fragments (MCQ )
- note : okazaki fragments are eventually joined to become a single continuous strand