## The Cytoskeleton

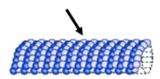
The cytoskeleton provides mechanical support and shape to cells.

It consists of different  $\underline{protein}$  structures including  $\underline{microtubules}$ , which are found in all eukaryotic cells.

Tubulin protein

Microtubules Structure

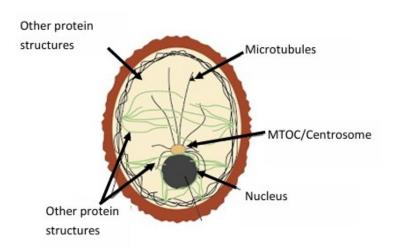
Hollow cylinders composed of the protein tubulin.

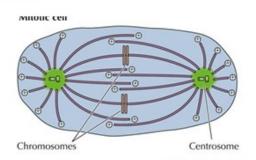


#### Microtubules Function

- 1. Control the movement of membrane-bound organelles.
- 2. Control the movement of <u>chromosomes</u> though formation of <u>spindle fibres</u> that are active during cell division.

Microtubules radiate from the microtubule organising centre (MTOC)/ centrosome near the nucleus.





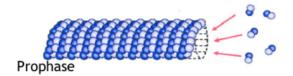
# Cell Division & cytoskeleton

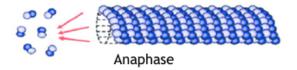
Cell Division requires <u>remodelling</u> of the cytoskeleton through <u>polymerisation</u>.

and <u>depolymerisation</u> of <u>tubulin</u> in the formation/ break down of <u>spindle fibres</u>.

Formation of Spindle fibres (polymerisation)

Breakdown of Spindle fibres (depolymerisation)





### \*\*\*Likely applied knowledge Question\*\*\*

Chemotherapy cancer drugs such as <u>colchicine</u> and <u>paclitaxel</u> are designed to destroy microtubule formation in tumour cells that show an increase rate of cell division.

#### Disadvantage

Cell division in normal cells also inhibited OR cannot move organelles within cell.

Other drugs interfere with another stage of the cell cycle such as <u>cisplatin</u> which prevents successful DNA replication.

# The Cell Cycle

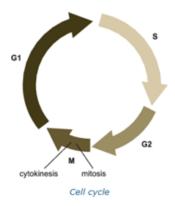
The Cell Cycle

Consists of interphase and mitotic phase.

### a) Interphase

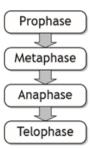
This Phase involves growth of the cell and DNA synthesis.

- G1 Growth phase
- S DNA replication
- G2 Second growth phase

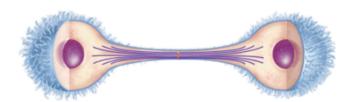


### b) Mitotic (M) Phase

1. Mitosis The chromosomal material is separated by the spindle fibres in a <u>4 stage</u> process.

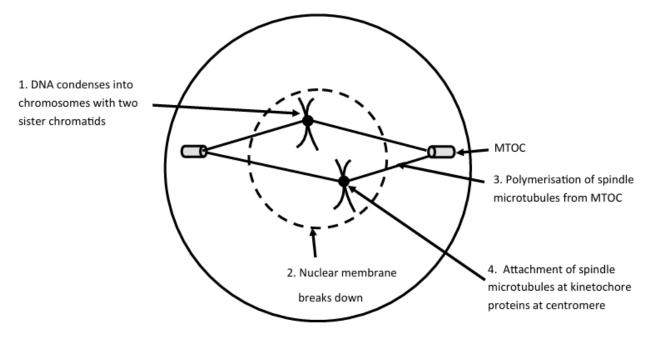


2. Cytokinesis The <u>cytoplasm</u> is <u>separated</u> into two daughter cells.



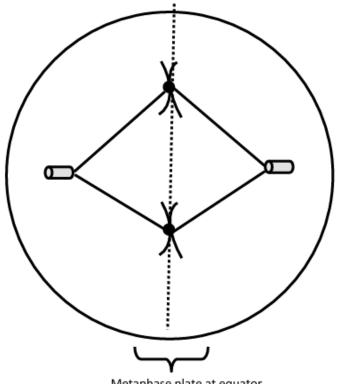
# **Stages of Mitosis**

- 1.Prophase
- a. DNA condenses into chromosomes each consisting of two sister chromatids.
- b. The <u>nuclear membrane</u> breaks down.
- <u>Spindle microtubules</u> extend from the MTOC by <u>polymerisation</u> attach to chromosomes via their kinetochores in the centromere region.



#### 2. Metaphase

Chromosomes are aligned at the metaphase plate (equator of the spindle).

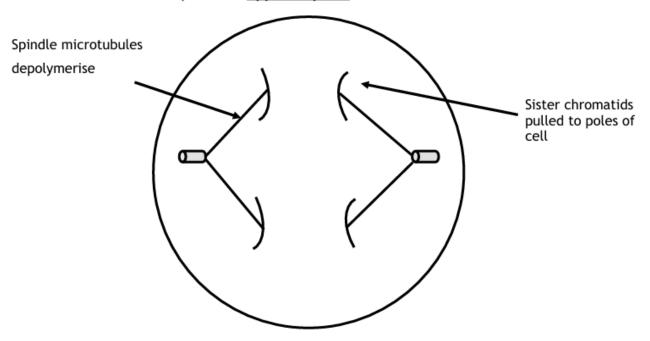


Metaphase plate at equator

# **Stages of Mitosis**

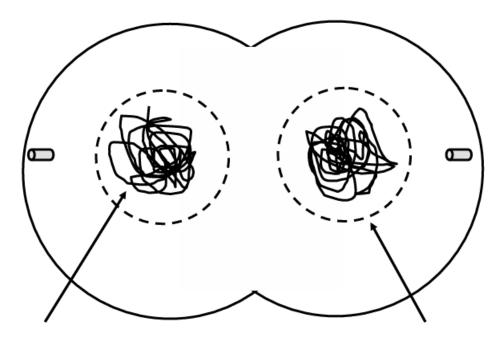
# 3. Anaphase

Spindle microtubules shorten by <u>depolymerisation</u>, separating <u>sister chromatids</u> which are pulled to <u>opposite poles</u>.



### 4. Telophase

The chromosomes <u>decondense</u> and <u>nuclear membranes</u> are formed around them.



Chromosomes decondense

Nuclear membrane reformed

# **Cell Cycle Checkpoints**

Three Cell Cycle Checkpoints

Mechanisms within the cell that assess the condition of the cell during the cell cycle and halt progression to the next phase until certain requirements are met.

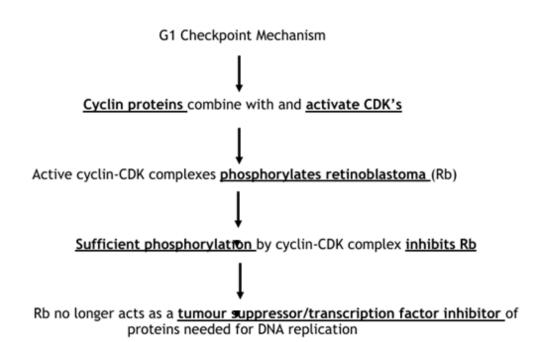
Name of Checkpoint	Cell Checkpoint	Halts entry before
G1 checkpoint	Assesses cell size through sufficient cyclin proteins that accumulate during <u>cell growth</u>	S phase
G2 Checkpoint	Success of <u>DNA replication</u> & any <u>DNA damage</u> is assessed.	M phase
M Checkpoint	Checks whether chromosomes have aligned correctly on the <u>metaphase plate</u> and attached to the spindle microtubules	Anaphase

#### Cyclin proteins

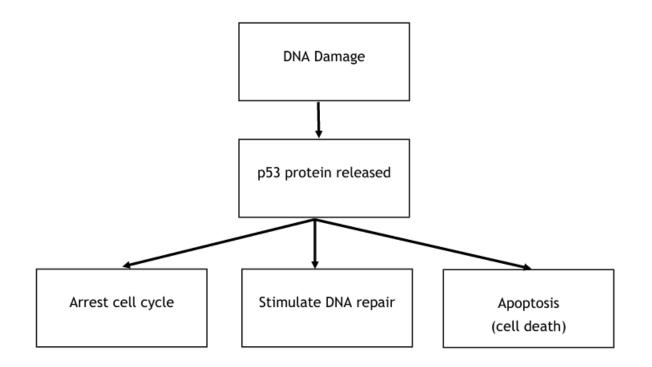
Accumulate during cell growth and are involved in regulating the cell cycle.

Cyclins combine with and activate CDK's.

Active cyclin-CDK complexes **phosphorylate specific proteins** that regulate progression through the cycle. If **sufficient phosphorylation** is reached, progression occurs.

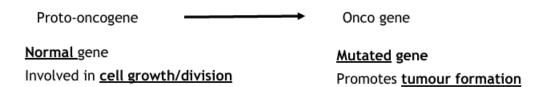


# DNA Damage & p53



# Cell Cycle Rate

- 1. <u>Uncontrolled reduction</u> in the rate of the cell cycle
  - may result in degenerative disease.
- 2. <u>Uncontrolled increase</u> in the rate of the cell cycle
  - may result in tumour formation.



## **Apoptosis**

Apoptosis

- 1. External Cell death signal Lymphocytes
- Internal Cell death signal <u>DNA damage/absence of growth factors (GF)</u>

External death signal molecules

- 1. Lymphocyte releases cell death signal
- 2. This signal molecule bind to a <u>surface receptor</u> protein
- 3. A protein cascade is triggered within the cytoplasm
- 4. Resulting in activation of <u>caspases (proteases)</u> that cause destruction of the cell

Internal death signal

- 1. DNA damage/absence of GF activates p53 tumour suppressor protein.
- 2. Resulting in activation of <u>caspases</u> that cause destruction of the cell

Importance of Apoptosis

Remove cells no longer required during the process of <u>development/metamorphosis</u>

