The Cell Cycle

Chapter 12

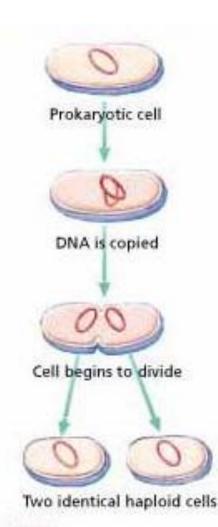
Key Roles of Cell Division

- 1. Continuity of life is based on reproduction of cells, or cell division
- 2. Reproduction of an entire organism; ex: an amoeba is a one celled organism.
- 3. Production of progeny from multicellular organisms. Ex: plant cuttings
- Sexually reproducing organisms from single cell (fertilized egg → fetus → infant)
- 5. Renewal & repair of damaged or worn out cells

Cell Division Roles

- Prokaryotic cells
 - Binary fission = reproduction
 - Origin of Replication = point where replication begins in bacteria DNA

- Eukaryotic cells
 - Development, growth, and repair



Cell Cycle

- Cell cycle life of a cell from the time it is formed until its own division into two daughter cells
 - Passes identical genetic material to cellular offspring.

Cellular Organization Genetic Material

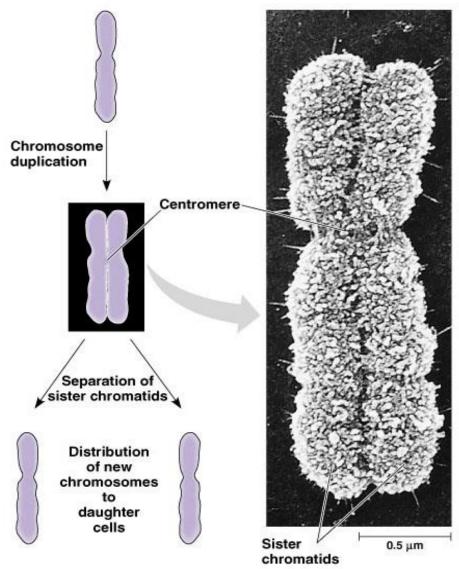
- **<u>DNA</u>** cell's genome, genetic material
 - Typical human has ~ 2n of DNA
 - All new cells will have an EXACT copy of the DNA
- <u>Chromosomes</u> coiled DNA
 - Structure that contains all the cell's packaged DNA
 - Eukaryotic chromosomes made of <u>chromatin</u> complex of DNA and associated proteins that helps maintain the structure of the chromosome
 - Each chromosome carries 100s-1000s of genes

Chromosome Numbers

- Each organism has a characteristic number of chromosomes.
- Human somatic cells (body cells) have 46 chromosomes
- Gametes (reproductive cells sperm / egg) have half the # of chromosomes (human eggs and sperm have 23 chromosomes).

Chromosome Structure

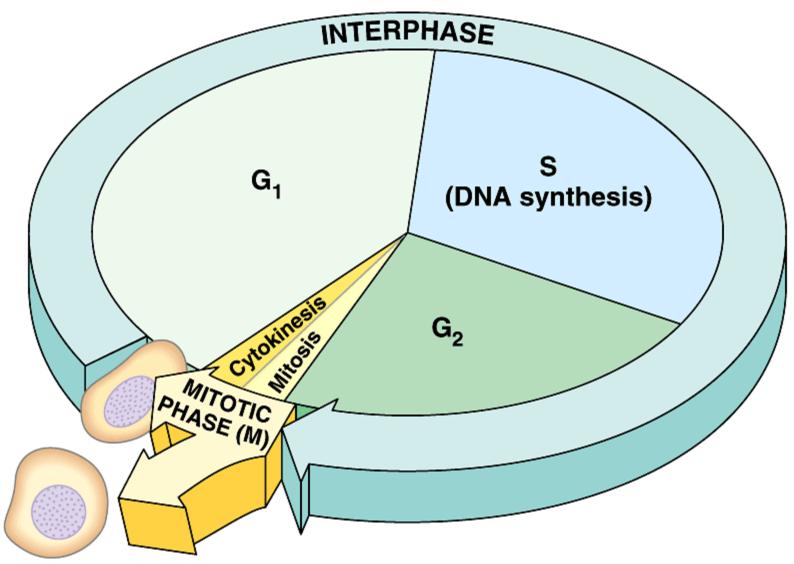
- Non-dividing cells' chromosomes are in the form of <u>CHROMATIN</u>
- Following DNA replication chromosomes coil & condense
- Duplicated chromosomes have 2 halves = <u>SISTER</u> <u>CHROMATIDS</u>
- Chromatids are connected by a <u>CENTROMERE</u>



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Phases of the cell cycle

- Interphase (90% of cell's life)
 - 1. G₁ Phase
 - 2. S Phase
 - 3. G₂ Phase
- M phase mitotic phase (10% of cell's life)
 - 1. Prophase
 - 2. Metaphase
 - 3. Anaphase
 - 4. Telophase
- Cytokinesis



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Interphase

- Accounts for ~ 90% of cell's cycle
- Cell grows and copies chromosomes
- Divided into 3 subphases:
 - 1. G_1 (first gap)
 - 2. S phase (synthesis)
 - 3. G₂ (second gap)
 - During all 3 subphases cell grows by producing proteins & cytoplasmic organelles

- Human cell takes ~ 24 hours to divide
 - Chromosomes duplicated during sphase (10-12hrs)
 - Cell grows in G1 (5-6hrs)
 - Cell continues to grow in G2 (4-6hrs)
 - M phase (< 1 hr)

MITOSIS

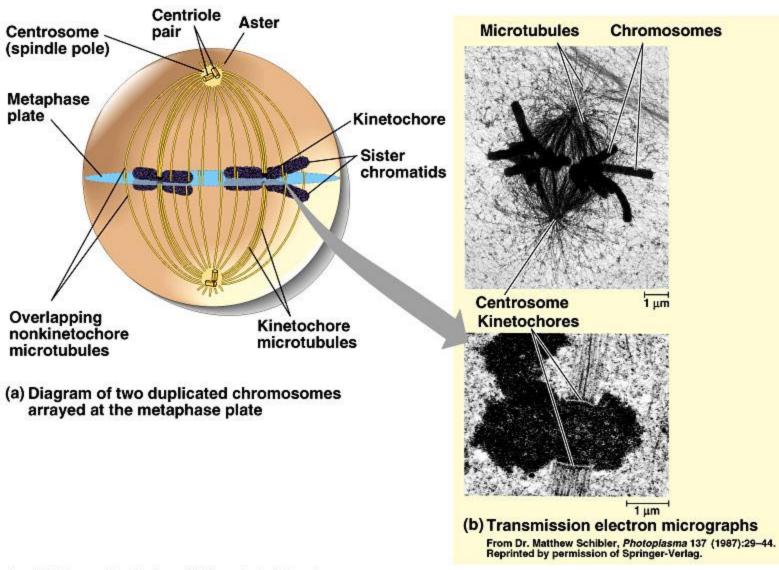
- **MITOSIS** Division of the **NUCLEUS**
- **<u>CYTOKINESIS</u>** = division of the cytoplasm
 - Human body ~ 200 trillion somatic cells (we all started as one)
 - Mitosis **MAINTAINS** the chromosome number
 - If a cell begins w/ 46 chromosomes, the new cell will have 46 chromosomes.

Cytoskeleton – Role in Cell Cycle

- Microtubules
 - Hollow tubes that move chromosomes
 - Centrioles 9 sets of triplet microtubules
 - Centrosome microtubule-organizing center near the nucleus
- Microfilaments
 - Actin 2 strands intertwined
 - Helps with cleavage furrow formation

Structures involved in cell division

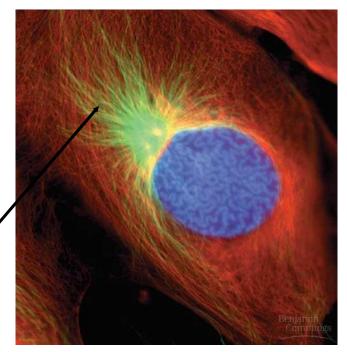
- 1. <u>Spindle fibers</u> (mitotic spindle) fibers made of microtubules and associated proteins that move the chromosomes during division
- 2. <u>Centrosome</u> nonmembranous organelle that organize and produce the spindle
 - Single centrosome replicates during interphase
 - <u>Asters</u> radial arrays of microtubules forming from the centrosome
- **3.** <u>Kinetochore</u> group of proteins associated with sections of chromsomal DNA at the centromere
 - Place of microtubule attachment



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M phase – Mitosis - Prophase

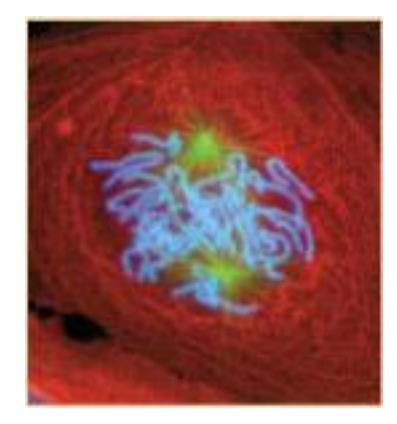
- Chromatin condenses into chromosomes becoming visible under light microscope
- Nucleoli disappear
- Duplicated chromosomes w/ 2 sister chromatids
- Mitotic spindles form
- Lengthening of spindles pushes centrosomes away from each other.



- •Lung cells from newt
- •22 chromosomes
- •Chromosomes- blue
- •Microtubules- green
- •Intermediate filaments- red

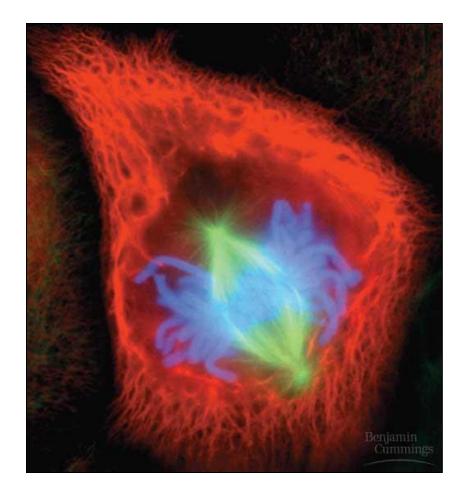
M phase – Mitosis - Prometaphase

- Nuclear envelop starts to breakup
- Microtubules can now interact with chromosomes and attach to centromere
- Each chromatid now has a kinetochore
- Nonkinetochore mircotubules interact with those on the opposite pole



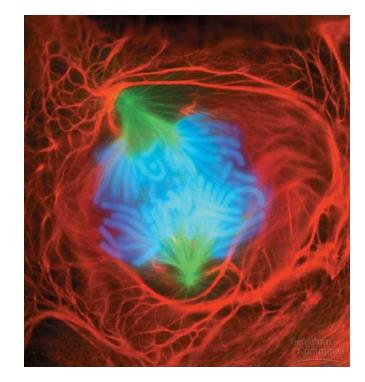
M phase – Mitosis – Metaphase

- Longest stage of mitosis ~ 20 minutes
- Centrosomes to opposite ends
- Chromosomes line up at equator = <u>metaphase plate</u>; middle of the cell because of tugging from kinetochore microtubules
- Microtubules that originate from the centrosomes are attached to each side of the sister chromatid's kinetochore
- Microtubule = spindle b/c of shape.



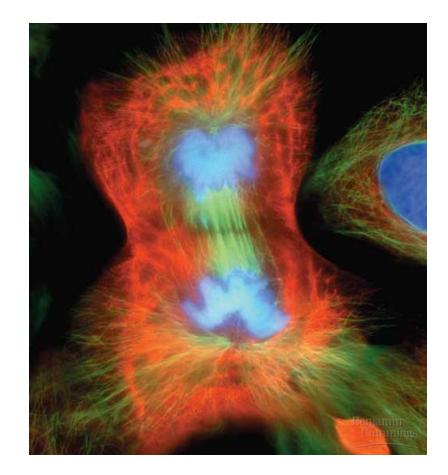
M phase – Mitosis – Anaphase

- Shortest stage
- Sister chromatids are pulled apart by microtubules (spindle fibers)
 - Caused by action of motor proteins as they depolymerize the kinetochore microtubules at the kinetochore end
 - This action shortens the fibers
- Chromosomes move toward opposite ends of cell
- Cell elongates due to nonkinetochore microtubules moving past one another also using motor proteins
- End of anaphase, two ends of cell have EQUIVALENT & complete new set of chromosomes



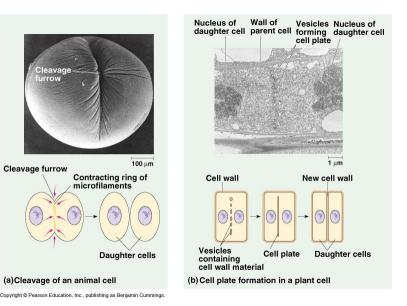
M phase – Mitosis – Telophase

- Daughter cell nuclei and nucleoli begin to form
- Nuclear envelop forms around each set of chromosomes
- Chromosomes uncoil to chromatin state
- Nuclear division is complete



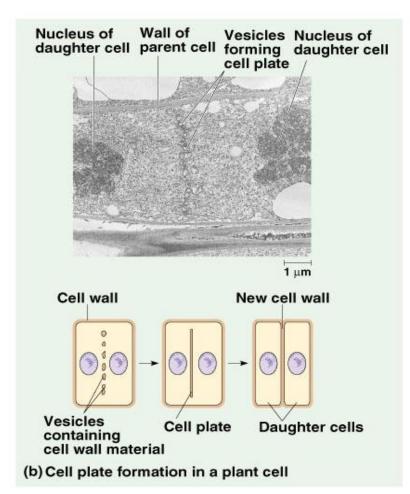
M phase – Mitosis – Cytokinesis

- Animal cells
 - Cleavage process that separates the two daughter cells
 - Cleavage furrow when a rin of actin forms on the cell surface and then interacts with myosin protein
 - It begins to contract until the cell is divided



M phase – Mitosis – Cytokinesis

- Plant Cells
 - Cell plate forms from the fusion of membrane vesicles made from the golgi apparatus
 - Deposits of cell wall material are collected in cell plate
 - Plasma membrane forms followed by cell wall from cell plate contents.



Cell Cycle Control

- 3 checkpoints throughout the cell cycle
 G1, G2, and M
- Cyclin dependent kinases
- Cyclin increases in number in the S phase and then breaks down after Mitosis
- Cyclin binds to Cdks to activate them
 - Becomes Mitosis promoting factor MPF

Cell Cycle regulation

- Growth factors = proteins released by certain cells that stimulates other cells to divide
 - Platelet derived growth factor PDGF
 - Released from platelets
 - Bind to tyrosine kinase receptors on cells called fibroblasts (cells that aid in wound healing)
 - Triggers the fibroblasts to move past the G1 checkpoint

Cell Cycle regulation

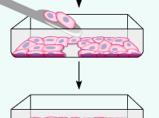
- Density-dependent inhibition = phenomenon where crowded cells stop dividing
 - In the lab, cells will fill a space. When some cells are removed, mitosis is triggered and the cells divide to fill the space again



Cells anchor to dish surface and divide (anchorage dependence).

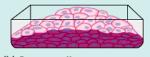


When cells have formed a complete single layer, they stop dividing (densitydependent inhibition).



If some cells are scraped away, the remaining cells divide to fill the gap and then stop (densitydependent inhibition).

(a) Normal mammalian cells



Cancer cells do not exhibit anchorage dependence or density-dependent inhibition.

(b) Cancer cells

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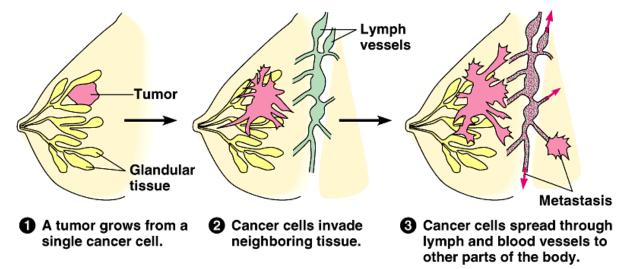
Cell Cycle regulation

 Anchorage dependence = to divide, cells must be attached to a substratum

- Cancer cells do not exhibit density-dependent inhibition or anchorage dependence
 - Normally cells undergo apoptosis when an irreparable mistake occurs in DNA replication, but cancer cells bypass that normal control

Cancer

- Transformation normal cell to cancer cell
- Tumor: benign or malignant
- Metastasis



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• HeLa Cell video