

Test 4 Review

Anthony Yu

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1 Cancer

1.1 9 Hallmarks

Making more cells isn't enough for cancer to proliferate. These cells start by overdividing; but they will quickly get killed by the various regulatory checkpoints in our body, such as immune system or apoptosis. Thus they need to satisfy all the hallmarks.

1. Proto-oncogens mutate and become hyperactive: proto-oncogenes are genes that can turn into a cancer gene (AKA an oncogene). They usually help the cell go through cell cycle. An example is rtk, a protein associated with a proto-oncogene. (recall that it is the stick thing that has 3 branches, which combine to form a dimer when they receive a growth signal). Once the proto-oncogene becomes mutated, rtk will keep telling the cell to divide.

See the sheet for more details on this process.

2. Anti-oncogenes mutate and become inhibited: For example, p53 and pRB genes, which are master breaks of the cell cycle, stop working.
3. Apoptotic genes mutate. This blocks apoptosis and fail to trigger it. Example would be bcl-2 proteins block apoptosis, when they activate the cells no longer suicide.
4. Telomerase genes become active and extend telomeres to help cells bypass hayflick limit and make them immortal
5. Cells must acquire the ability to produce VEGF and other growth factors that stimulate growth of blood vessels to nourish tumor (a process sustained angiogenesis)
6. Metastasis: cancer cells separate from their intercellular junctions and invade other tissues.
7. Changes to metabolism. Some cells do anaerobic glycolysis even in presence of oxygen.

8. Avoiding immune destruction. Straightforward.
9. Phenotypic plasticity: Cancer cells de-differentiate and return to a more stem-cell like state.

1.2 Causes of mutation

1. Random mutations due to replication errors
2. Ionizing radiation
3. Mutagenic materials, such as arsenic in water
4. Viral infections promote proto-oncogenes or deactivate tumor suppressor genes. HPV.

1.3 Cancer in old age

most people have cancer in old age because the longer you live, the more likely there is for a mutation to occur. It's a chance game.

1.4 Early cancer

1. inheritance of mutated proto-oncogenes (thus present in all cells!)
2. Inheritance of defective DNA repair genes
3. different versions of detoxifying enzymes that can make more/less harmful molecules from original molecules, or work faster/slower
4. Impaired immune system
5. Exposure to carcinogens
6. disease

Definition 1 (Carcinogens). *Covalent bonds between carcinogens and DNA cause mutations*