Question 1
You are studying a human membrane protein that acts as an enzyme and has Guanosine triphosphate (GTP) as its substrate. Binding of GTP to the active site of this protein is shown in the following schematic. For simplicity only the side-chains of the important amino acids in the active site are shown. The alpha carbon atom of each amino acids is indicated with an *.

![Diagram of GTP binding and amino acids]

a) Circle the strongest interaction that exists between……..

- Side-chain of Lys<sub>100</sub> and the phosphate group of GTP.

  **Hydrogen Ionic Hydrophobic Interaction /Van der Waals forces Covalent**

- Side-chain of Glu<sub>150</sub> and the ribose sugar of GTP.

  **Hydrogen Ionic Hydrophobic Interaction /Van der Waals forces Covalent**

- Side-chain of Tyr<sup>201</sup> and the guanine base of GTP.

  **Hydrogen Ionic Hydrophobic Interaction /Van der Waals forces Covalent**

b) You make mutations in the GTP binding pocket of this enzyme and examine the effect of each mutation on the binding of GTP. Give the most likely reason why each mutation has the stated effect. **Note:** Consider each mutation independently.

- Lys<sub>100</sub> mutated to Arg results in an enzyme that still binds to GTP.

- Lys<sub>100</sub> mutated to Glu results in an enzyme that cannot bind to GTP.
**Question 1 continued**

c) This enzyme is a glycoprotein, and requires addition of the carbohydrate fructose for its function. Fructose (shown below) forms a glycosidic bond with the side-chain of the amino acid serine of this enzyme. Draw the R group of serine and circle its reactive atoms that will form a covalent bond with fructose. Also circle only one of the reactive groups on fructose that could participate in the formation of this bond.

![Fructose structure](image)

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d) This enzyme is a membrane protein and has a linear stretch of amino acids that spans the lipid bilayer of the membrane. Below are three different options for the amino acid sequence of this linear stretch of amino acids.

- Option 1: lys-cys-ser-trp-tyr-asp-leu-his-gly-arg-leu
- Option 2: leu-ala-gly-cys-ala-val-ile-leu-ala-phe-trp
- Option 3: gly-thr-tyr-ser-ala-gly-glu-glu-lys-thr-ser

Circle the option that most likely forms a stretch of protein that spans the lipid bilayer? **Explain** briefly why you selected this option.

e) This enzyme is a glycoprotein, and requires addition of the carbohydrate fructose for its function. Full activity of this enzyme requires both the carbohydrate (fructose) and protein components. You isolate the enzyme and subject it to the following treatments in a test tube and measure its activity. You then return the samples in each tube to the pre-treatment conditions and measure the enzyme activity again. Complete the following table for each treatment. **(Note: Consider each treatment independently).**

<table>
<thead>
<tr>
<th>Tube</th>
<th>Treatment</th>
<th>Bonds disrupted by treatment</th>
<th>Protein active/inactive when returned to pre-treatment conditions? Explain.</th>
</tr>
</thead>
<tbody>
<tr>
<td>#1</td>
<td>Lactase (hydrolyzes the sugar lactose to glucose and galactose)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>#2</td>
<td>Trypsin (cleaves the protein at Lys)</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

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**Question 2**

DNA polymerase is an enzyme that catalyzes the **polymerization** that involves the addition of the incoming nucleotides to a growing strand of DNA. Each polymerization may be regarded as a pair of **coupled reactions** as follows.

- **Reaction 1:** Binding of the incoming nucleotide triphosphate to the active site of the enzyme followed by its **hydrolysis** to nucleotide monophosphate.

- **Reaction 2:** Formation of a **phosphodiester bond** between the nucleotide monophosphate and the last base of the growing strand of DNA.

a) Based on the information provided, which of these two reactions (reaction 1/reaction 2) is most likely to have a higher free energy change (+ΔG)? **Explain** why you selected this option.

b) Briefly explain why it is important to couple reaction 1 with reaction 2 during DNA polymerization.

c) From the choices below, circle the reaction parameter(s) that is reduced by DNA polymerase and briefly explain why this enhances the rate of reaction.

<table>
<thead>
<tr>
<th>Reaction equilibrium</th>
<th>ΔG</th>
<th>Activation energy</th>
</tr>
</thead>
</table>

**Question 3**

You are studying two traits using a mouse model. The mutant mice are **small** and **lethargic** whereas the normal mice are **large** and **active**. You cross a true breeding **large** and **lethargic** mouse with true breeding **small** and **active** mouse. All of the resulting F1 mice are **small** and **lethargic**.

a) What are the genotypes of the true breeding parental mice? Use the nomenclature outlined below.

- In each case, use the uppercase letter for the allele associated with the dominant phenotype and the lowercase letter for the allele associated with the recessive phenotype.
- For the size (i.e. large or small) use D or d to designate the alleles.
- For the activity (i.e. active or lethargic) use G or g to designate the alleles.

<table>
<thead>
<tr>
<th>Parent</th>
<th>Genotype</th>
</tr>
</thead>
<tbody>
<tr>
<td>Large and lethargic</td>
<td></td>
</tr>
<tr>
<td>Small and active</td>
<td></td>
</tr>
</tbody>
</table>
Question 3 continued
b) You then cross two of the F1 mice. If these two genes were unlinked, based on Mendel’s law, about how many large and active mice do you expect out of a total of 320?

c) You find that the two genes are linked. If the map distance between the two genes is 20 cM, out of a total of 400 offspring, how many will show the non-recombinant/parental phenotypes?

Question 4
Your next experiment involves the fruit fly. In the parental (P) generation, you mate a true-breeding female fly that has short antennae and red eyes with a true-breeding male fly that has long antennae and white eyes. All of the flies in the F1 generation have long antennae and red eyes.

Note: Assume the genes for these two traits are located on autosomes. For the alleles that regulate antennae length, use the letters “A” and “a” and for the alleles that regulate eye color, use the letters “B” and “b.” Use the uppercase letters to represent the alleles associated with dominant phenotypes and lowercase letters to represent the alleles associated with recessive phenotypes.

a) Give the genotypes of the flies in the P generation.

Genotype of the female Fly:  
Genotype of the male Fly:  

b) Give the genotypes of the gametes produced by the flies in the P generation.

i. Genotype of the gametes produced by the female Fly:

ii. Genotype of the gametes produced by the male Fly:

c) Give the genotypes of the flies that are produced in the F1 generation.

d) Assuming that the genes that regulated antennae length and eye color assort independently, give ALL the possible genotypes of the gametes that are produced by the F1 flies.

e) You mate a female F1 fly with a true-breeding male fly that has short antennae and white eyes.

i. What is the genotype of the male fly in this mating experiment?

ii. What is the genotype of the gametes produced by the male fly in this experiment?
**Question 4 continued**

f) You **mate two F1 flies** and obtain 1600 offspring in the F2 generation. If the genes that regulate antennae length and eye color **assort independently**, complete the table below for each type of progeny in the F2 generation.

<table>
<thead>
<tr>
<th>Phenotypes of the flies in F2 generation</th>
<th>Corresponding genotypes</th>
<th>Approximate number</th>
</tr>
</thead>
</table>

**Question 5**

a) An error occurs during division of cells that make up the lining of the intestines such that daughter cells inherit an abnormal number of parental chromosomes. Is this more likely to be an error in mitosis or in meiosis? **Explain** briefly.

b) A child is born with three copies of chromosome #21 in nearly every cell in its body. This is a disorder commonly called Down Syndrome. Does this disorder most likely reflect an error that occurred during a mitotic cell division or during a meiotic cell division? **Explain** briefly.

c) During which type of cell division would you expect to see chiasmata? **Explain** briefly.

**Question 6**

The following “line-angle” drawings represent three chemical structures. On each drawing, the hydrogen atoms that should be bonded to the NON-carbon atoms are missing.

![Chemical Structures](image_url)
a) For each structure, show the position of All carbon (C) and all hydrogen (H) atoms. **Note:** If there is a charge present make sure you take it into account.

b) Give the chemical formula of each of the structures shown by the line angle drawings.

A:  
B:  
C:  

**Question 7**
The following human pedigree shows the inheritance pattern of a specific disease within a family. Assume that the individuals marrying into the family for all generations (except the parental generation) do not have the allele associated with the disease phenotype and that no other mutation arises spontaneously. Also assume complete penetrance.

a) **Circle** the most likely mode of inheritance for this disease. Choose from: autosomal dominant, autosomal recessive, X-linked dominant, X-linked recessive.

b) Write **all possible** genotypes of the following individuals in the pedigree. Use the uppercase “A” or “X^A” for the allele associated with the dominant phenotype and lowercase “a” or “X^a” for the allele associated with the recessive phenotype.

Genotype(s) of Individual 2:  
Genotype(s) of Individual 4:  

The following human pedigree shows the inheritance pattern of a specific disease within a family. Assume that the individuals marrying into the family for all generations (except the parental generation) do not have the allele associated with the disease phenotype and that no other mutation arises spontaneously. Also assume complete penetrance.

1st Pedigree

1 2 3 4 5

- Affected female
- Affected male
- Unaffected female
- Unaffected male

a) **Circle** the most likely mode of inheritance for this disease. Choose from: autosomal dominant, autosomal recessive, X-linked dominant, X-linked recessive.

b) Write **all possible** genotypes of the following individuals in the pedigree. Use the uppercase “A” or “X^A” for the allele associated with the dominant phenotype and lowercase “a” or “X^a” for the allele associated with the recessive phenotype.

Genotype(s) of Individual 2:  
Genotype(s) of Individual 4:  

c) What is the probability that Individual 5 will be a carrier?