

**1. pH, pK<sub>a</sub>, and all that (25 pts):**

(a; 9 pts) The imidazole side chain of histidine has a pK<sub>a</sub> of 6.04 (referring to the protonated side chain). Draw the dominant prototropic forms of histidine at (1) pH 4, (2) pH 8, and (3) pH 12. Don't worry about C<sub>α</sub> stereochemistry.

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(b; 6 pts) Calculate the ratio of neutral to protonated histidine at pH 6.5.

(c; 10 pts) We have mentioned that the  $pK_a$  of an amino acid side chain can change substantially in different protein contexts. If a histidine side chain were found buried among isoleucines in the center of a protein, what effect would this have on the actual  $pK_a$  of that particular side chain, and why? What if there were a buried aspartate as well?

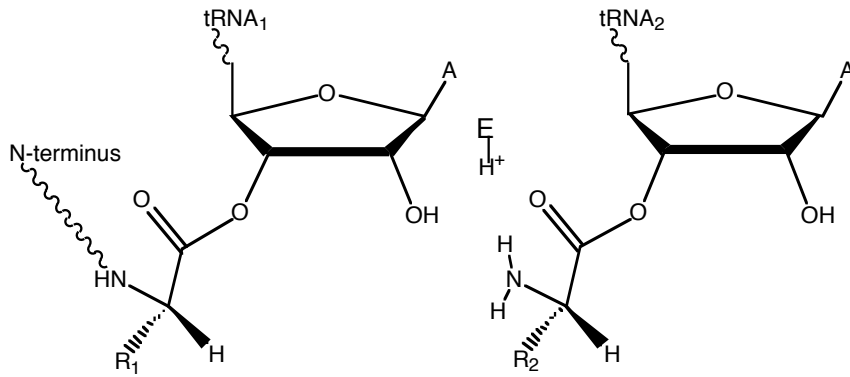
## **2. Amino acid and peptide bond structure and chemistry (26 pts):**

(a; 6 pts) Name the two sulfur-containing amino acids, and very briefly list one unique functions for each.

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(b; 15 pts) Draw the structure of Valine-Proline-Glutamate at pH 7. Make the valine-proline peptide bond *cis* and the proline-glutamate peptide bond *trans*. Give the 3- and 1-letter codes for each amino acid below your structure.

(b; 5 pts) The structure below shows two charged tRNA molecules aligned for the process of peptide bond synthesis. Draw the first step of the reaction mechanism, leading to the key intermediate.



### 3. Thermodynamics (25 pts):

(a; 6 pts) What are the two most important functions for lipids?

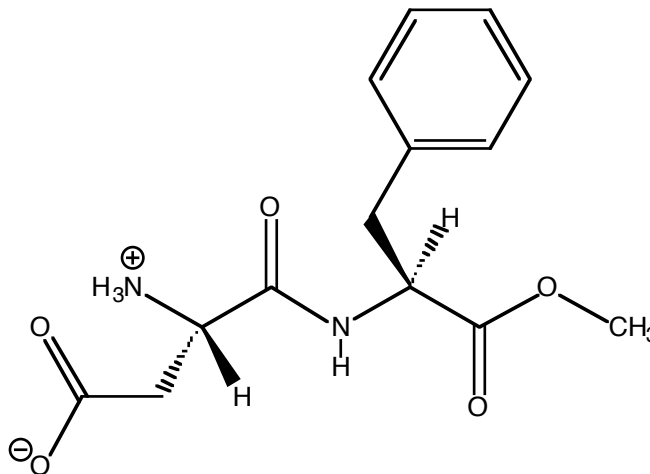
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(b; 9 pts) Briefly describe how typical aerobic cells maintain order in the face of the universal tendency toward increasing entropy.

(c; 10 pts) You have discovered an enzyme that converts substance A to substance B without requiring any input of free energy (i.e. the reaction proceeds without high-energy cosubstrates like ATP). The equilibrium lies far to the side of B. What then must be true about any process that carries out net conversion of B to A under the same conditions? Also, explain why your discovery either does or does not preclude the possibility that there may be a separate energy-consuming path for going from A to B, and a biological rationale for your answer.

**4. Intermolecular interactions(22 pts):**

(a; 12 pts) The structure of the artificial sweetener Aspartame is shown below. Identify potential hydrogen bond donors and acceptors, and electrophilic carbons. Why must phenylketonurics avoid Aspartame? (Memory jogger: would Aspartame absorb UV light?)



(b; 10 pts) Briefly describe the origin of the hydrophobic effect. Give the signs of  $\Delta H^\circ$ ,  $\Delta S^\circ$ , and  $\Delta G^\circ$  for the process of dissolving a long alkyl chain in water.

Score: 1. pH, pK<sub>a</sub>, and all that (25 pts): \_\_\_\_\_

2. Amino acid and peptide bond structure and chemistry (26 pts): \_\_\_\_\_

3. Thermodynamics (25 pts): \_\_\_\_\_

4. Intermolecular interactions(22 pts): \_\_\_\_\_

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**Total: out of 100 (2 pts for Honor Pledge)** \_\_\_\_\_