- 1. The chemical structure of a new degradable polymer, poly(ethylene glycol)-b-polylysine-b-poly(D,L-lactide), is shown below.
  - a. Based on chemical structure alone, discuss your prediction of whether this material will degrade more quickly or more slowly than poly(L-lactide) *in vivo*.
  - b. Cite 2 physical data you would want to know to make a better prediction in part (a), and explain why these would aid in this estimate.
  - c. What will the end-point degradation products of this polymer be if it is degraded by incubation *in vitro* in phosphate buffered saline, pH 7.4 at 37°C? Would these products be different if the material were degraded *in vivo*?

$$\overset{a}{\operatorname{CH}_{3}} \circ - \left( \overset{b}{\operatorname{CH}_{2}} \overset{b}{\operatorname{CH}_{2}} \circ \right)_{X} \left( \overset{0}{\operatorname{C-CH}_{1}} \overset{c}{\operatorname{CH}_{1}} \overset{e}{\operatorname{NH}_{2}} \right)_{Y} \left( \overset{0}{\operatorname{C-CH}_{1}} \overset{f}{\operatorname{C-CH}_{1}} \circ \right)_{Z}$$

 The generation of acidic or basic degradation products is a serious issue complicating the design of devices for *in vivo* application, even for well-established materials like poly(lactideco-glycolide). Propose an approach to limit or eliminate the generation of a low pH within and near a PLGA device implant. Be creative! (limit ~1/2 page explanation). 3. Propose a route to synthesize the hydrolyzable polymer shown below. Show the structure of the monomers you would use and any initiators/catalysts.

$$\begin{array}{cccc} \mathsf{CH}_3 & \mathsf{O} & \mathsf{O} & \mathsf{H} \\ \mathsf{CH}_3 \text{-} \mathsf{C}\text{-} \mathsf{O}\text{-} \mathsf{C}\text{-} \mathsf{CH}_2\text{-} \mathsf{CH}_2\text{-} \mathsf{O}\text{-} (\mathsf{C}\text{-} \mathsf{C}\text{H}\text{-} \mathsf{O}\text{-} \mathsf{C}\text{-} \mathsf{C}\text{H}\text{-} \mathsf{N})_{\mathsf{n}}\text{-} \mathsf{H} \\ \mathsf{CH}_3 & \mathsf{O} & \mathsf{CH}_3 & (\mathsf{C}\text{H}_2)_4 \\ \mathsf{NH}_2 \end{array}$$

4. In order to carry out studies of polymer hydrolysis on reasonable experimental timescales, a number of researchers have used elevated temperature to speed up degradation. Using the data given below, determine the degradation mechanism (surface or bulk erosion) for a sample of polycaprolactone ( $\rho = 1.146 \text{ g/cm}^3$ ) 0.02m thick degraded in water at 37°C and whether the mechanism is changed by degrading the sample at 85°. (Hint: assume both the hydrolysis rate constant and the diffusivity of water in the polymer have an Arrhenius form.)

At 37°C:  $k = 9.7 \times 10^{-8} \text{ s}^{-1}$   $D_{H2O} = 10^{-8} \text{ cm}^2 \text{s}^{-1}$ 

At 50°C:  $k = 5.4 \times 10^{-7} \text{ s}^{-1}$   $D_{H2O} = 3.5 \times 10^{-8} \text{ cm}^2 \text{s}^{-1}$ 

k = rate constant of PCL bond hydrolysis

 $D_{H2O}$  = diffusivity of water in PCL