1. Volume V=100000  $\mu^3$ +10000 $\mu^3 \approx 10^5 \mu^3$ . Number of protein molecules is N= 500molecules/minute\*60minutes/hour\*24hours/day\*7days/week/N<sub>a</sub>, N<sub>a</sub> - Avogadro number Concentration C=N/V=8.4\*10<sup>-8</sup>M

2. Dynamic equilibrium: dN<sub>syntetized molecules</sub>/dt=dN<sub>transported to axon</sub>/dt= =1000molecules/minute=K

 $dN_{transported to axon}/dt=v*A*C$ , v-rate of transport (5mm/day), A-cross-sectional area, C stable concentration

 $C=K/(v*A)=2.4*10^{-7}M$ 

3.

- a. 2Dt= $<\Delta X^2$ >, t= $(0.05 \text{ cm}^2)^2/2\text{D}=6.25*10^4\text{ s}$
- b. Presumably the diffusion doesn't have directional preference, so half of synthesized molecules will be in dendrite, and half will be in axon. In dendrite: C=(N/2)/(N<sub>a</sub>\*V<sub>dendrite</sub>)=~21uM, volume V<sub>dendrite</sub>=4 $\mu^{2*}$ 1mm In axon: the first 1 mm (==2\*< $\Delta X^{2}$ >) contains ~95% of molecules diffused in axon (two standard deviations of normal distribution), so average concentration C=0.95\*(N/2)/(N<sub>a</sub>\*V<sub>axon</sub>)=~79uM, volume V<sub>axon</sub>=1 $\mu^{2*}$ 1mm
- 4.  $pH_{medium}=7.4$ , so  $[H^+]=10^{-7.4}M=40nM=C_c$

Surface is negatively charged to V=-50mV, and negative charges are distributed by Boltzmann distribution: surface concentration of positive ions (protons)  $C_s=C_c*exp[zeV/kT]$ , and surface pH=-lgC<sub>s</sub>=6.5

- The surface area of sphere is proportional ("∝") to the r<sup>2</sup>. We have ~25% of radius change ([7.5μ-6μ]/6μ), so we have (1.25)<sup>2</sup>≈1.56 of initial membrane area (~56% stretching). But even 4% stretching leads to lysis, so lymphocytes can swell due to smoothing of folds in membrane.
- 6. At the beginning we have K<sup>+</sup> -defined membrane potential: V<sub>K</sub>=RT/(zF)ln[3mM/140mM]=~-90mV, where is z=1. At the end we can calculate voltage difference again by Nernst equation, but for Ca<sup>2+</sup>: V<sub>Ca</sub>=RT/(zF)ln[0.5mM/2μM]=~+66mV, where is z=2. The change in potential will be ΔV=V<sub>K</sub>-V<sub>Ca</sub>=~-156mV
- 7. Donnan equilibrium (see also Lecture#11):  $C_{K \text{ inside}}/C_{K \text{ outside}} = C_{Cl \text{ outside}}/C_{Cl \text{ inside}}$ , Also we know that  $C_{K \text{ outside}}=1\text{mM} C_{Cl \text{ outside}}=110\text{mM}$ . Another equation – electroneutrality inside the cell (assuming that there are only K & Cl):  $C_{K \text{ inside}}=C_{Cl \text{ inside}}$ . Also x – change in concentrations inside:  $C_{K \text{ inside}}=140+x$ , and

 $C_{Cl inside} = 4 + x$ . By substituting one equation to another :  $x^2+144x+450=0$ , the meaningful solution of this square equation (which will keep concentrations positive) x=-3.2mM, so  $C_{Cl inside}=4+x=0.8$ mM, and  $C_{K inside}=\sim136.8$ mM. Potential difference by Nernst equation (for example for potassium): V=56mV\*lg[1/136.8]=~ -118mV.

- 8.
- a. Bending energy for the membrane with constant curvature E=(B/2)\*(1/R<sub>t</sub>)<sup>2</sup>\*A. Area of cylinder A= $\pi$ \*d\*l=~5 $\mu$ <sup>2</sup> E=7.8\*10<sup>-17</sup> N\*m, KT=~4\*10<sup>-21</sup> N\*m (T=300K) So E= ~KT\*10<sup>4</sup>
- b. Affinity of enhancer (E) to DNA gives us the change in free energy:  $\Delta G_1 = RT^*ln[K_{E-DNA}] = \sim 5.5$  Kcal/mole. For binding E to transcription factor (TF) we know change in entropy and enthalpy of binding, so we also can calculate change in free energy:  $\Delta G_2 = \Delta H T^*\Delta S = -4-1 = -5$  Kcal/mole. For full reaction:  $\Delta G = \Delta G_1 + \Delta G_2 = -10.5$  Kcal/mole, and equilibrium constant:  $K_A = exp[-\Delta G/(RT)] = \sim 4^*10^7$  1/M, and  $K_D = 2.5^*10^{-8}$  M; We also know that we need enhancer concentration equal to dissociation constant for 50% activation of transcription. By knowing the volume of nucleus we can calculate the actual number of molecules:  $N=K_D^*Na^*V = = \sim 1700$  molecules
- 9.  $D=kT/(6\pi\eta r)=2.2*10^{-7} \text{ cm}^2/\text{s}$  (You also can calculate it by using the  $D \propto 1/r$  proportionality from this formula (see comments for (5) of the provided to exam equations):  $D_{1\mu \text{ radius sphere}}=4.4*10^{-9} \text{ cm}^2/\text{s}$ , so  $D_{0.05\mu \text{ radius virus particle}}=2.2*10^{-8} \text{ cm}^2/\text{s}$ )
- 10. The first mechanism linear movement of scanning protein complex with the rate v=30b.p./s=~6x10<sup>-4</sup> cm/s (3Å/b.p.), so scanned distance will be l=t\*v
  For diffusion model scanned distance defined by d=ΔX/2, and <ΔX<sup>2</sup>> = 2D\*t.
  For t=1 min= 60 s: l=~5.4x10<sup>-2</sup> cm=~1800b.p., and d=(1/2)\*(2DT)<sup>1/2</sup> =~1.7um=
  =~5800 b.p.
  For t=1 h = 3600s: l=~3cm=108 Kb.p., d==~13.4um= 45Kb.p.
- 11. L=5000bp, rate of synthesis is v=50bp/s, so time of synthesis is T=L/v=100s. Time between moment of transcription initiations t=300bp/v=6 s.  $100^{th}$  copy of RNA will be synthesized in 99\*6=594<sup>th</sup> second after beginning of synthesis of the first RNA molecule, and will finish synthesis in next T=100s, so cycle for synthesis of 10 molecules is t+T=~700 s. And the gene should be activated

700s/30min=~39% of time.

12. Reynold's number:  $R = \upsilon L\rho/\eta$ , L- here is diameter,  $\upsilon -$  speed,  $\rho -$  density of obnject (particles),  $\eta -$  viscosity of media (we can use viscosity of water here - brain slices are usually covered by thin layer of some liquid). So  $\upsilon = \sim 1.7 m/s$