

### 1. Bio-Molecule Detector

One application for electronics that has gained a lot of attention over the past several years is in so-called “bio-molecule” detection. The idea is to build a system that detects the presence of specific molecules and/or cells (e.g., specific viruses, proteins, etc.) in a biological sample; if this detection can be performed automatically and using relatively low-cost components, it can have a dramatic impact on a number of areas such as medical diagnosis, drug development, DNA sequencing, etc.

In this problem we’ll look at how some of the techniques we learned about in the touchscreen module can be applied to realize a hypothetical bio-molecule detector. (Real bio-molecule detection systems involve quite a bit more complexity than what we’ll include here, but in many designs the same basic principles apply.)

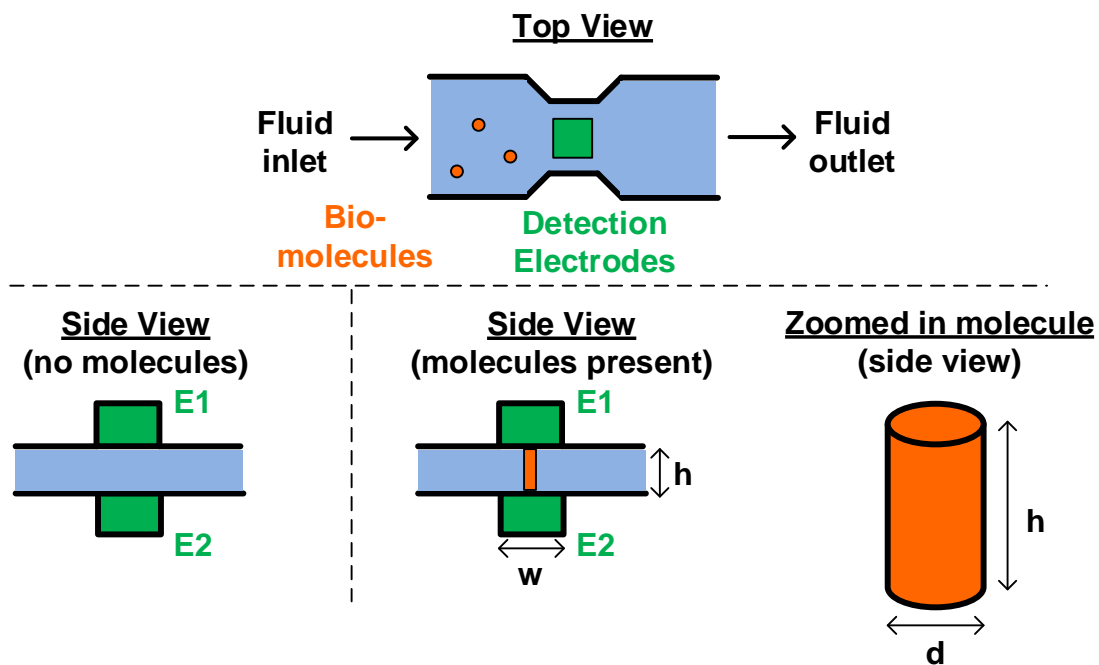


Fig. 1: Cartoon of our simplified bio-molecule detector system.

As shown in Fig. 1 above, the detector works by flowing a liquid that may or may not contain the biomolecules through a region in the device that has electrodes on the top and bottom of the liquid channel. The electrodes (E1/E2 in Fig. 1) are chemically “functionalized” (using e.g. some appropriately designed antibodies) so that if the specific bio-molecule of interest is present in the fluid sample, one or more of the molecules will get physically trapped between the two electrodes

(bottom right of Fig. 1). After all of the fluid has been cleared out of the device (i.e., so that if there are bio-molecules present, there is only air in between the two electrodes E1/E2), we can then figure out whether or not one or more bio-molecules were trapped by measuring the resistance between the two electrodes, the capacitance between the two electrodes, or both.

- (a) Let's first assume that we want to detect the presence of a bio-molecule by measuring resistance. If no bio-molecule is present, what should be the resistance between E1/E2? As shown in Fig. 1, if each bio-molecule is a cylinder with diameter  $d = 10\text{nm}$ , height  $h = 100\text{nm}$ , and has a resistivity  $\rho = 100 \Omega\cdot\text{m}$ , what would be the resistance between E1 and E2 if only a single bio-molecule has been trapped? Note that you can assume that the trapped molecule is exactly vertically oriented when it is trapped – i.e., the top and bottom faces of the molecule are both aligned with surfaces of the electrodes.
- (b) Using the same numbers for  $d$ ,  $h$ , and  $\rho$  as part a), as a function of the number of trapped bio-molecules  $N_{\text{molecules}}$ , what is the resistance between E1 and E2? (Note that you can assume that  $N_{\text{molecules}}$  is small enough that all of the molecules fit within the electrode area, and that all of the molecules are still trapped in an exactly vertical orientation.)
- (c) Given your answers to parts (a) and (b), design a circuit that will output a +5V voltage if more than 5 molecules are trapped, and 0V if 4 or fewer molecules are trapped.
- (d) Now let's assume that the bio-molecules aren't conductive at all (i.e.,  $\rho = \infty \Omega\cdot\text{m}$ ), and so we will instead try and detect the change in capacitance caused by the presence of trapped bio-molecules. Assuming that the electrodes are square (from the top view) and have a size length  $w = 10\mu\text{m}$ , that  $h$  is still 100nm, and that the permittivity of the bio-molecule is  $\epsilon = \epsilon_r * \epsilon_0 = 10 * 8.85\text{e-}12 \text{ F/m}$ , what is the capacitance between E1 and E2 if no bio-molecules are present?
- (e) Using the same parameters as part (c), what is the capacitance between E1 and E2 if a single bio-molecule is trapped? How about if  $N_{\text{molecules}}$  are trapped?
- (f) Given your answers to parts (d) and (e), design a circuit that will output a +5V voltage if more than 5 molecules are trapped, and 0V if 4 or fewer molecules are trapped.
- (g) We may not know in advance whether the bio-molecule will be conductive, and so we might want to build our detector circuit so that it is capable of measuring either the resistance or the capacitance between E1/E2. Design a circuit that will output a voltage that is proportional to the resistance between E1/E2 (if measuring resistance), or output a voltage that is inversely proportional to the capacitance between E1/E2 (if measuring capacitance). Note that you can assume that if your circuit is configured to measure capacitance, the resistivity of the bio-molecule is infinite (i.e., you will always be measuring either purely resistance or purely capacitance).

- (h) Because the bio-molecules are small and the binding process that traps them isn't perfect, the measurements we get from any real detector can be quite noisy. Because of this, one of your colleagues suggests to use the circuit you designed in part (g) to measure both the capacitance and the resistance of a sample, and to then somehow use that information to get a more noise tolerant estimate of  $N_{\text{molecules}}$ . Calling the measured resistance  $R_{\text{meas}}$  and the measured capacitance  $C_{\text{meas}}$ , using the tools we've learned in EE16A, formulate and describe a strategy that may achieve this goal.

## 2. Digital-to-Analog Converter

As we saw in homework 6, one device that finds a lot of usage is a “digital-to-analog converter” (DAC) that allows us to translate signals from the digital representation we use in e.g. our computers to an analog quantity we can use in the “real” world. In this problem we'll look at one implementation of such a DAC that converts the digital codes into an analog voltage. The DAC design we'll be working with is shown below in Fig. 2, and is known as an “R-2R” DAC.

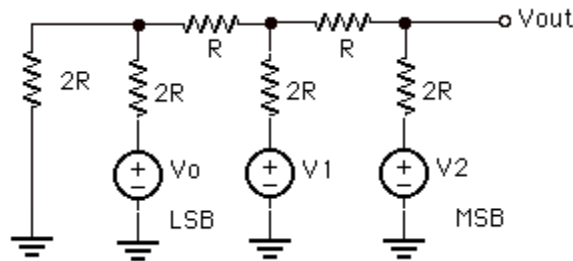


Fig. 2: Digital-to-analog converter circuit.

Note that throughout this problem, we will assume that a digital “1” translates in to a voltage of 1V, and a digital “0” translates in to a voltage of 0V.

- For a digital input of 100 (i.e.,  $V_2 = 1.2\text{V}$ ,  $V_1 = 0\text{V}$ , and  $V_0 = 0\text{V}$ ), what is the output voltage  $V_{\text{out}}$ ?
- For a digital input of 001 (i.e.,  $V_2 = 0\text{V}$ ,  $V_1 = 1.2\text{V}$ , and  $V_0 = 0\text{V}$ ), what is the output voltage  $V_{\text{out}}$ ? (Hint: you may find it easier to solve this problem by remembering that you can “transform” between voltage and current sources.)
- For a digital input of 010, what is the output voltage  $V_{\text{out}}$ ?
- It turns out that by combining the results of parts (a), (b), and (c), it can be shown that the circuit in Fig. 2 can be modeled as the circuit shown below in Fig. 3, where  $R_T = R$ , and  $V_T = V_2/2 + V_1/4 + V_0/8$ .

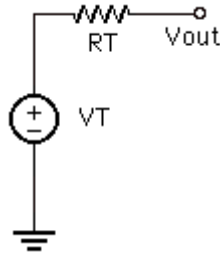


Fig. 3: R-2R DAC equivalent circuit.

Using this model for the R-2R DAC circuit and assuming that  $R_T = 1\text{k}\Omega$ , design a circuit that would provide an output voltage that swings from  $-1.75\text{V}$  (corresponding to 000) to  $1.75\text{V}$  (corresponding to 111) while driving a  $100\Omega$  load resistance.

### 3. “Timer” Circuit

As we saw in the locationing module, keeping track of the amount of that has elapsed between the occurrence of two events (e.g., receiving a signal from one satellite vs. another) can be extremely useful. Therefore, in this problem we will explore the design of a circuit that can produce a periodic voltage waveform, where the period of that waveform will be set by the values we choose for our components. In particular, we want to design a circuit that will output  $+5\text{V}$  for half of the period, and  $-5\text{V}$  for the other half of the period – i.e., your circuit should output a square wave with a 50% duty cycle.

In order to realize this circuit, you are allowed to use any combination of the following components:

- Ideal op-amps
- Resistors
- Capacitors
- Switches
- Batteries (i.e., voltage sources)

If you need some control signals (like those we used in the touchscreen module from the Arduino) that drive some switches in order to reset and/or initialize some voltages within your circuit, please feel free to utilize those as well.

- (a) Sketch a design for a circuit that achieves the timer functionality described above. Don’t worry about setting the value of the period yet or the values of the any of the components yet – just show a schematic for the circuit. (Hint: If driven by a fixed current, how does the voltage across a capacitor change over time?)

- (b) Now select component values for your design such that the period of your timer circuit is  $100\mu\text{s}$ .

#### 4. Estimating Friends

You've recently been hired at a new and growing social-network startup, and you want to develop a model that can predict the number of friends new users on the network will have.

- (a) You collect a bunch of data regarding the age of the current users and the number of friends they have. Using this data, you'd like to develop a predictive model for the number of friends a new user will have. The data you have collected looks like the data in Fig. 4.

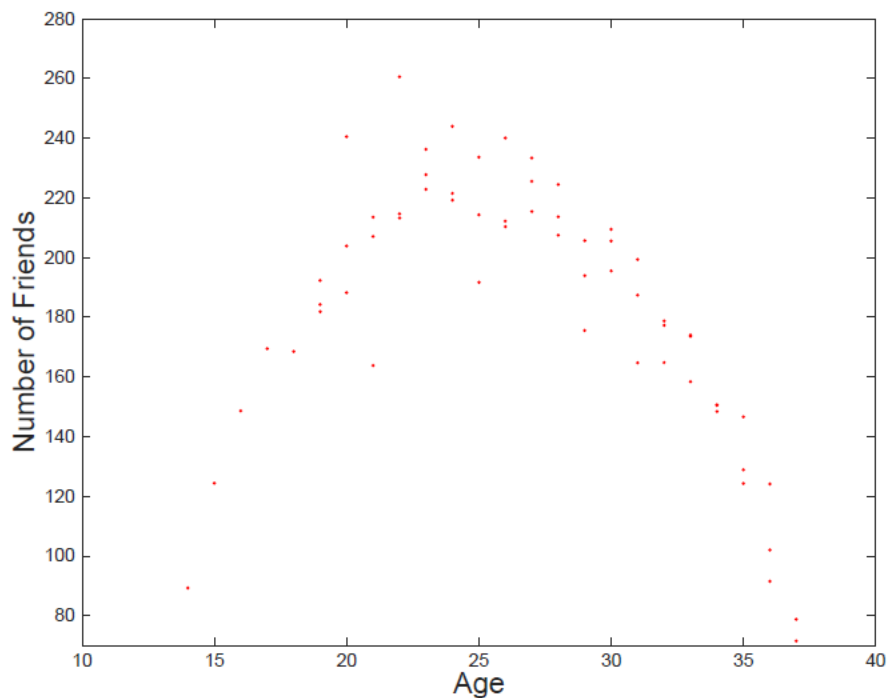


Fig. 4: Number of friends vs. age

If the points you have are of the form  $(a_i, n_i)$ , where  $a_i$  is the age of the  $i$ th user and  $n_i$  is the number of friends, what kind of model would you use to capture the relationship between age and number of friends? If a new user comes in with age  $\alpha$ , how would you predict the number of friends they have? (You don't need to provide any numerical answers – just explain the procedure you would use.)

- (b) You realize that just age is not a perfect predictor of the number of friends – clearly there are other factors that influence this. In fact, you realize people who live in urban areas tend to have more friends than people who live in rural areas. So you develop a metric called “urbanity” ( $u$ ) that maps every location to a number from 1 to 10 where 10 is most urban and 1 is the most rural setting. You expect to have a linear relationship between the number

of friends of a user and the degree of urbanity  $u_i$  of the  $i$ th user's location. How would you augment your model above to account for this? Now, if you have a new user come in with age  $\alpha$  and urbanity  $u$ , how would you predict their number of friends?

## 5. Medical Imaging

It turns out that solving underdetermined systems of equations can be very useful in medical imaging applications. Specifically, we are often in a situation where we want to image parts of the body in order to find abnormalities in the tissue. One of the techniques to do this is called tomography, and we described the basics behind in lecture. Tomography uses the fact that different materials in the body absorb different amounts of light, and thus readings of the absorption of beams of light from various directions onto the tissue can be used to reconstruct an image of the tissue.

We won't be developing a true tomography system here, but let's consider a simplified model that utilizes the same basic principles. Consider an unknown vector  $x = [x_1 \ x_2 \ \dots \ x_n]^T$  that represent the "image" of the tissue (i.e., samples of light absorption of different parts of the body) and where  $n = 10^8$ . Let's assume that if the tissue is normal and healthy – which should be the case for the large majority of the pixels in the image – the absorption of light is essentially equal to 0. Let's further assume that if there is e.g. a blood clot (or some other abnormality) in the tissue, the absorption value is very high.

Because each absorption measurement may take anywhere from several ms to even several seconds in order to achieve reasonable accuracy, it is typically infeasible to get  $10^8$  or more measurements in order to estimate the value of every pixel in the image using a technique like least squares. Because of this, we would like to use the fact that number of blood clots is very low, and thus the number of non-zero entries within the vector  $x$  should also be very low. This situation can be approximated by thinking about the vector  $x$  as having a small norm.

Although we can't measure the entire tissue image directly, let's say that we are able to get  $10^3$  measurements of the total absorption for a volume of tissue that spans multiple pixels in the original image. In other words, the result of each absorption measurement  $b_i$  is equal to the dot product of a "sampling vector"  $a_i$  (note that  $a_i$  is a column vector) and the image vector  $x$ .

- (a) Set up a matrix-vector equation using the  $a_i$ 's,  $b_i$ 's and  $x$  that captures the measurements that were taken.
- (b) Recalling that a vector with a small number of non-zero entries should have a small norm, how could you "solve" the system of equations you set up in part (a) to find a good estimate for the image vector  $x$ ?

## 6. Practice with Underdetermined Systems

Note that you can find the numerical solutions for the problems below using a computer; on the actual exam you will not be asked to manually invert anything larger than a  $2 \times 2$  matrix. We would however encourage you to compute by hand the intermediate terms (e.g.,  $AA^T$ ) of the minimum norm solution for the problems below.

(a) Find the minimum norm  $x$  for the system

$$\begin{bmatrix} 1 & 1 & 1 & 1 & 1 \\ 1 & 1 & 0 & 1 & 1 \\ 1 & 1 & 1 & 1 & 0 \end{bmatrix} \begin{bmatrix} x_1 \\ x_2 \\ x_3 \\ x_4 \\ x_5 \end{bmatrix} = \begin{bmatrix} 10 \\ 8 \\ 8 \end{bmatrix}$$

(b) Find the minimum norm  $x$  for the system

$$\begin{bmatrix} 1 & 1 & 1 & 0 & 1 \\ 1 & 1 & 0 & 1 & 1 \\ 1 & 1 & 1 & 1 & 0 \end{bmatrix} \begin{bmatrix} x_1 \\ x_2 \\ x_3 \\ x_4 \\ x_5 \end{bmatrix} = \begin{bmatrix} 10 \\ 8 \\ 8 \end{bmatrix}$$