

EE 6301
Week1 Introduction
1.

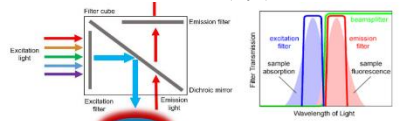
Week2 Optical Biosensor

- 1. Structure of Biosensor:
Light source -> sensor -> light analyzer
- 2. Key Parameters of Biosensors:
Specificity: what is the analyte being measured
Sensitivity: the quantity (or concentration) of the analyte (LOD) (=slope)
Detection limit / Limit of detection (LOD)
LOD = 3 x s / m
- 3. Light sources: (Lasers, LED, lamps)
s: standard deviation;
m: slope;
ceil
- 4. General requirements on sources:
Spatial coherence
Low noise (RIN)
Adequate power output (SNR)
- 5. Light as Waves - The EM Wave Model:
E(z,t) = E0 cos(omega t - kz)
angular frequency: omega = 2pi nu
wavelength: lambda = c/nu
wave vector: k = 2pi/lambda
speed of light: c = 3 x 10^8 m/s
- 6. Light as Particles - The Particle Model
Energy is delivered in photons (光子).
Energy of one photon: E = h*nu = 1239.8/lambda[nm]
Planck's constant: h = 6.63 x 10^-34 Joule.s
1W = 1J/s 1eV = 1.602 x 10^-19J
Photon momentum (动量): p = h/lambda
- 7. Refractive Index
n = Speed in Vacuum / Speed in medium
Snell's Law: n1 sin(theta1) = n2 sin(theta2)
Total internal Reflection: theta_c = sin^-1(n2/n1)
- 8. Evanescent Wave Field
Penetration depth: d = lambda_0 / 4pi * sqrt(n1^2 sin^2 theta_1 - n2^2) (assuming incident angle = 90°)
High intensity: I(d) = exp(-d/L_0)
- 9. Light-matter Interactions
Incident = Reflected + Transmitted + Absorbed
Absorption / A+Luminescence / Scattering
- 10. Absorption spectroscopy
Absorbance: A = epsilon * L = -log10(I/I0)
I(lambda) = I0(lambda) * e^-epsilon(lambda) * CL, epsilon = epsilon / 2.303
epsilon: molar absorption coefficient (M^-1 cm^-1)
c: molar concentration (M)
L: optical path length (cm)
- 11. Fluorescence Lifetime & Quantum Yield:
Phi = N_emit / N_absorb = k_f / (k_f + k_nr) = k_f / tau_s
k_f: fluorescence rate constants
- 12. Scattering (no absorption):
Elastic scattering: no energy change
Inelastic scattering: energy change
Frequency change, Vibration, Raman
- 13. Raman scattering
stokes: E down, f down; anti-stokes: E up, f up
wavenumber: nu_bar = 1/lambda (cm^-1)
- 14. Grating
Dispersion: d sin theta = m * lambda
Resolving Power: R = lambda/dlambda = mN
Delta lambda: resolution m: diffraction order Ceil(N)

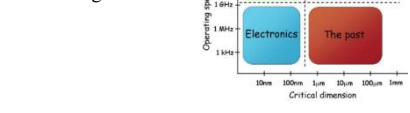
CW: continuous wave
CW: continuous wave

Week3 Optical Biosensor (2)

- 1. Fluorescence Biosensors-(Dye) Labeled Sensors:

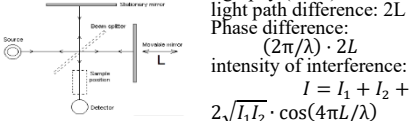


- 1) Excitation of a fluorophore through the absorption of light energy. 2) A transient excited lifetime with some loss of energy. 3) Return of the fluorophore to its ground state, accompanied by the emission of light.
- 2. Selecting Filters:
- 3. Label-Free Sensor Working Principle
Sensitivity: S_lambda[nm]/RIU = (lambda_2 - lambda_1)/(n_2 - n_1)
Resonant wavelength: lambda = 2*pi*r*n_eff/m
r: fiber radius n_eff: effective refractive index
Full Width at Half Maximum:
FWHM = x_2 - x_1
f(x_1) = f(x_2) = 0.5*f_max
Q-factor: Q = f_0/Delta f Delta f由0.5I_max决定
f_0: Resonant/central frequency
Figure of Merit: FOM = Sensitivity/Delta f
Surface Plasmon Biosensor:
Materials:
Al/Cu/Ag/Au



Week4 Biomedical Imaging (1)

- 1. Light propagation in tissues:
attenuation coefficient: mu_t[cm^-1] = mu_a + mu_s
mu_a: absorption coefficient mu_s: scattering
mu_a = probability of absorption / unit pathlength = inverse of the mean-free-path for absorption
light attenuation:
I(x) = I_0 * e^-mu_t * L = I_0 * e^-(mu_a + mu_s)L
loss of incident light: dI = absorption + scattering
- 2. Fresnel reflection coefficients (normal incident):
R = (n_1 - n_2)^2 / (n_1 + n_2)^2, n_1 incident
- 3. Property of thin lens:
thin lens equation: (1/S_1) + (1/S_2) = (1/f)
Magnification: M = S_2/S_1
S_1: object distance, f: focal length
- 4. Diffraction Limit:
Resolution:
Delta d = 0.61 lambda_excited / NA
Numerical aperture:
NA = n sin theta
Width of the central band of the Airy disk:
D = 1.21 lambda / (n sin theta)
In practice: M Delta d / Delta x = 2.3 ~ 3
Delta x: the pixel size of the detector
- 5. Two-photon excited fluorescence
Excited energy = h*nu_0.5
- 6. Optical Coherence Tomography (OCT)
light path difference: 2L
Phase difference: (2*pi/lambda) * 2L
intensity of interference:
I = I_1 + I_2 + 2*sqrt(I_1*I_2) * cos(4*pi*L/lambda)



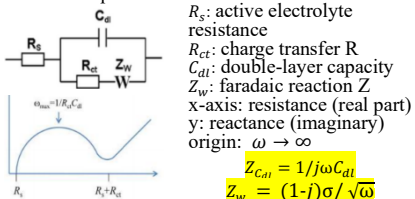
Week5 Optical Bioimaging and Devices

- 1. Physics of Ultrasound:
p = p_0 * e^-alpha x cos(omega t - kx)
angular frequency: omega = 2pi f
propagation constant: k = 2pi/lambda
alpha: attenuation coefficient
Attenuation = alpha (dB/(MHz cm)) * [cm] * f [MHz]

- 1 是往返总路程
intensity of the acoustic wave: I(t) = p^2(t)/Z
[W/m^2] p(t): pressure, Z: impedance
- 2. Confusion Matrix
Sensitivity = TP / (TP + FN)
Specificity = TN / (FP + TN)
Data on slanted line are large -> good model
Receiver Operator Characteristic (ROC)
Area Under Curve = 1 -> perfect

Week6 Electrochemical Biosensor

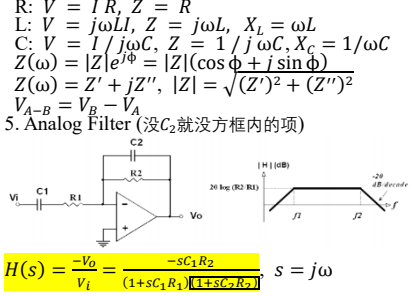
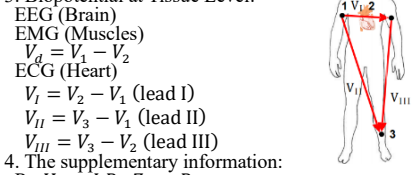
- 1. Electrochemical Calculation
Coulomb Calculation Formula: Q = I * t
Q: Coulombs(C); I: Current (A, amps); t: time(s)
Faraday Constant Formula:
1 mol of electrons = 1F = 96485 C/mol
Avogadro's Num: N_A approx 6.02 x 10^23 particles/mol
Moles of electrons: n_e[mol] = Q/F
Moles of element: n_element[mol] = n_e/n
n: electrochemical equivalent
Mass: m_element = n_element * M_element = rho * V
M_element: Molar mass of silver
- 2. Randle equivalent circuit:
R_s: active electrolyte resistance
R_ct: charge transfer R
C_dl: double-layer capacity
Z_w: faradaic reaction Z
x-axis: resistance (real part)
y: reactance (imaginary)
origin: omega -> infinity
Z_Cdl = 1/j*omega*C_dl
Z_w = (1-j)*sigma/sqrt(omega)



- 3. Bode plot:
x-axis: logarithmic scale of the frequency
y-axis: logarithm of the impedance Z & phase shift Phi
omega = 2pi f

Week12 Bioelectricity and Biopotential

- 1. Ionic concentrations and channels
Current: I_K = g_K (V_m - E_K)
V_m = V_in - V_out < 0
E_K = ((RT)/(zF)) * ln([K+]_out/[K+]_in)
g_K: conductance, 电阻的倒数
E_K: Equilibrium potential for potassium
R: Universal gas constant (J/(mol.K));
T: Temperature (K) z: 离子电荷数(K+ -> +1)
F: Faraday's constant
[K+]_out: Concentration of K+ outside the cell
- 2. five steps on the cell membrane:
Time(ms)
Membrane potential (mV)
Resting potential
Depolarization
Action potential
Repolarization
Hyperpolarization
Threshold potential
- 3. Biopotential at Tissue Level:
EEG (Brain)
EMG (Muscles)
V_d = V_1 - V_2
ECG (Heart)
V_I = V_2 - V_1 (lead I)
V_II = V_3 - V_1 (lead II)
V_III = V_3 - V_2 (lead III)
The supplementary information:
R: V = I R, Z = R
L: V = j*omega*L, Z = j*omega*L, X_L = omega*L
C: V = 1/j*omega*C, Z = 1/j*omega*C, X_C = 1/omega*C
Z(omega) = |Z|e^j*phi = |Z|(cos phi + j sin phi)
Z(omega) = Z' + jZ'', |Z| = sqrt(Z'^2 + Z''^2)
V_A-B = V_B - V_A
- 5. Analog Filter (没C_2就没方框内的项)

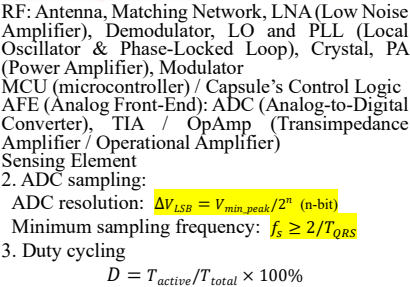


H(s) = -V0 / (Vi * (1 + sC1R1)(1 + sC2R2)) s = j*omega

H(jf) = H0 * (j(f/f1)) / (1 + j(f/f1)(1 + R1/R2)) H0 = -R2/R1
f1 = 1 / (2*pi*R1*C1) f2 = 1 / (2*pi*R2*C2)

Week8

- 1. Wireless Sensing System

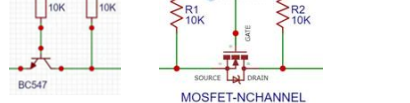


RF: Antenna, Matching Network, LNA (Low Noise Amplifier), Demodulator, LO and PLL (Local Oscillator & Phase-Locked Loop), Crystal, PA (Power Amplifier), Modulator
MCU (microcontroller) / Capsule's Control Logic
AFE (Analog Front-End): ADC (Analog-to-Digital Converter), TIA / OpAmp (Transimpedance Amplifier / Operational Amplifier) Sensing Element

- 2. ADC sampling:
ADC resolution: Delta V_LSB = V_min_peak/2^n (n-bit)
Minimum sampling frequency: fs approx 2/T_QRS
- 3. Duty cycling
D = T_active / T_total approx 100%

- 4. Baud rate
R_baud = R_bit / k, k = log2 M
k: Bits per Symbol; M: Modulation Order

- 5. Logic level shifter
bidirectional
unidirectional



- input-BJT-output 5V-close-3.3V
input-MOSFET-output 5V-close-3.3V
0V-open-0V 0V-open-0V

- 6. In vivo wireless communication
L_antenna approx lambda_c/4

lambda_c = v_0 / f_c * sqrt(epsilon_r) * lambda_c = v_0 / f_c * sqrt(epsilon_r)

lambda_c: Wavelength of carrier; v_0: Wave speed in vacuum; epsilon_r: Relative permittivity (相对介电常数); epsilon_r: Relative permittivity

Week9

- 1. Amplifier Switch
NPN: V_B 高开
N-channel: V_GS > V_th 开

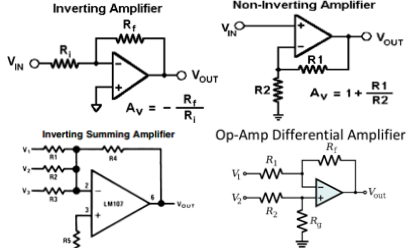


Transistor: Input and output are not insulated



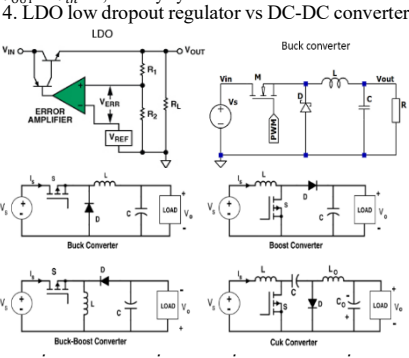
Relay: Input and output are insulated

2. Operational amplifier



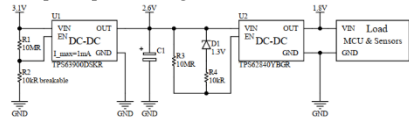
V_OUT = -R4 * (V1/R1 + V2/R2 + V3/R3)
V_OUT = ((Rf+R1)/R2) * V2 - (Rf/R1) * V1

- 3. Pulse width modulation (PWM)
V_OUT approx V_in * D, D: duty cycle
- 4. LDO low dropout regulator vs DC-DC converter

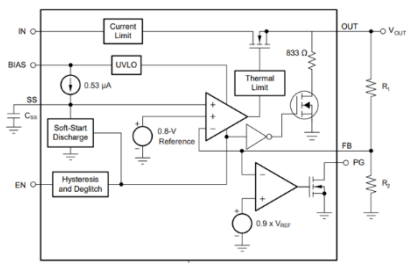


v_in * i_in = eta * v_out * i_in
v_in * i_in = eta * v_out * i_out
v_out = v_ref * (1 + R1/R2)
D = v_out / (v_out + v_in)
V_o = D * V_s
V_o = V_s / (1 - D)
V_o = D * V_s

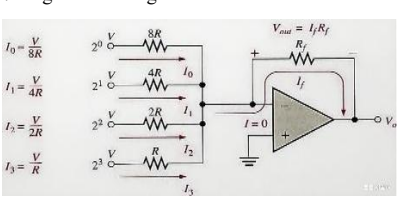
- 5. Super caps buffering circuit



6. TPS74201 (Soft Starter)

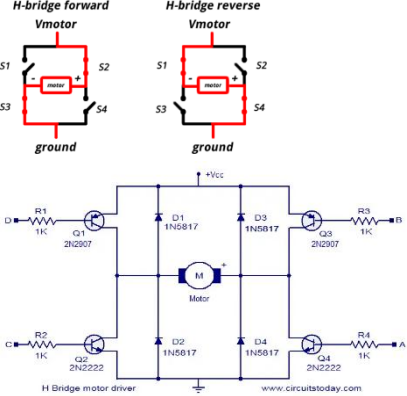


C_SS从0.53uA 处储能后缓慢放电, C_SS up T_start down
7. Digital to analog converter



Week10

1. H-bridge driver



Week1

Sensor 定义

A sensor is a device that detects and responds to input (stimulus) from its physical environment. Physical Changes -> Sensor -> Output Signals

Week 2 光学传感器 Optical Biosensor

. Fluorescence intensity is defined as the amount of photons emitted per unit time and per unit volume of solution

. Brightness

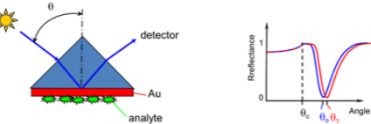
$$\text{Brightness} = \epsilon \times \Phi$$

光栅 (grating) 角度分光能力 单位角度内有多少波长成分 d 角度/d 波长 分光能力越大越好，在一个角度内要包含更少的波长成分

Week 3 Resonator 共振光学传感器

Surface Plasmon Resonance

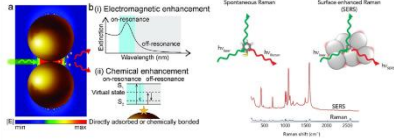
纳米结构下的缝，会让光密度变大，用 Al, Ag, Cu, Au 生成 plasmonic，当激光照射金属表面，会吸收特定波长的光发热，形成一个波谷 dip dark signal 暗信号 改变外部浓度，比如特定病毒附着，dip 会发生偏移，可能是激光角度也可能是波长 Localized Surface Plasmons 把金属 film 变成细小颗粒 拉曼散射 SERS surface enhanced Raman Scattering



金属纳米颗粒靠近或化学键有自己的 bend 会发生共振，在某些波长下会出现 peak 或 dip，增强拉曼散射，使信号放大

Week4 成像技术

The enhancement factor can be as much as $\sim 10^6$ to $\sim 10^{12}$



Penetration depth 光信号深入人体组织深度后，信噪比为 1 的深度；红外对人体组织的穿透更深，用于计算心跳，血氧饱和度，利用血红蛋白携带氧气后的吸收差异

Optical Coherence Tomography OCT 主要用于眼科检测，发射后，距离的细微差异带来相位差，图像主要与反射率有关。是眼睛的纵切面成像

Hyperspectral Imaging 包含更丰富的光谱，全波长的信息 (传统的只包含 RGB 三个波长的信息)，扫描成像

Week 5 非光学成像技术

Ultrasound Imaging 基于反射，声波频率只有百 Khz，因此频率比光低，波长更大，身体不同组织对超声波的衰减是不同的，且与频率有关，深度定义也是信噪比为 1 的地方。但是超声波是反射的，要计算来回的距离。因此实际深度是一半。

声光成像 Photoacoustic Imaging 声和光的优势，声更深入，光分辨率更高

laser pulse -> absorption -> thermal expansion -> acoustic waves -> ultrasound detection -> image formation

Week6 电化学传感器

化学氧化还原反应 Redox reaction 产生电子流，体液 blood, saliva 唾液, tears, urine, sweat

电压 Potentiometric，电流 Amperometric，阻抗 Conductometric 三种传感原理

电压是参考值，要有确定的 GND

Week7 人体电信号

对单个的 cell，可以用 patch electrode 直接插入细胞测试；也可以用微电极阵列 Multielectrode Array 测试细胞表面电压神经细胞的内外压差是 -70mv，由于钠钾泵，静息状态，内部 K 多，外部 Na 多

肌肉 EMG 电位用两个电极，测量两电极的差，有效降

噪

心脏 ECG 的用三个电极，两臂和腿，由于心脏有吸气和泵血

脑电波 EEG 用 20 个电极在头的各个部位

LDO 用于低压降，纹波小，电阻分压原理

DCDC 用于高压降，纹波大，但是效率很高，PWM 分压

计算反馈电阻 $V_{out} = V_{ref} \cdot (1 + R1/R2)$ ，计算软启动时间 $Q = CV = It$ 恒流源给外部电容充电到 V_{ref} 所耗费的时间即软启动时间，10ms 左右；超级电容放电， $E = 0.5 \cdot C \cdot V^2$

H 桥驱动

二极管作用续流，电极可视为一个大电感，停转时候会有反向的巨大电压，防止烧毁