Introduction to Biomedical Engineering

Biomedical Instrumentation

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Outline

- Chapter 8 and chapter 5 of 1st edition: Bioinstrumentation
 - Bridge circuit
 - Operational amplifiers, instrumentation amplifiers
 - Frequency response of analog circuits, transfer function
 - Filters
 - Non-ideal characteristics of op-amps
 - Noise and interference
 - Electrical safety
 - Data acquisition (sampling, digitization)

Overview of biomedical instrumentation

Basic instrumentation system



Emphasis of this module will be on instruments that measure or monitor physiological activities/functions

Types of medical instrumentation

- Biopotential
- Blood (pressure, flow, volume, etc)
- Respiratory (pressure, flow rate, lung volume, gas concentration)
- Chemical (gas, electrolytes, metabolites)
- Therapeutic and prosthetic devices
- Imaging (X-ray, CT, ultrasound, MRI, PET, etc.)
- Others

Characteristics of some bio-signals

Parameter	Voltage	Frequency (Hz)
ECG (skin)	0.5-4 mV	0.01-250
EEG (scalp)	5-200 μV	DC-150
EGG (skin)	10-1000 μV	DC-1
EGG (stomach)	0.5-80 mV	DC-1
EMG (needle)	0.1 - 5 mV	DC-10,000
EOG (contact)	50-3500 μV	DC-50
ERG (contact)	0-900 μV	DC-50
Nerve	0.01-3 mV	DC-10,000

EGG (electrogastrogram): measures muscular activity of the stomach EOG (ElectroOculoGram): measures the resting potential of retina ERG (ElectroRetinoGram): measures the electrical response of retina to light stimuli

Signal amplification

- Gain up to 10⁷
- Cascade (series) of amplifiers, with gain of 10-10000 each
- DC offset must be removed (ex. by HPF with a cutoff frequency of 1Hz)
- Further reduction of the common-mode signal

Analog circuits

Wheatstone bridge circuit



The measured V_{ab} can be used to obtain R which represents the unknown resistance of devices such a strain gauge and a thermistor

Operational amplifier (op-amp)



$$V_{out} = A(v_p - v_n)$$

Open-loop voltage gain

 $A \sim 10^6$

For ideal op-amps:

• No current flows into or out of the input terminals (input impedance)

- $v_p = v_n$ since A ~ 10⁶
- Output impedance
 0

Cautions for op-amp circuits

Op-amps are used with (negative) feedback loops for stability Must be in the active region (input and output not saturated)

Voltage follower or unity buffer



Advantage: input current is ~0, high input impedance. Output current drawn from the op-amp can drive a load (Z_L) or next stage of circuit; particularly useful as the first stage for physiological measurements



Summing amplifier



You can add more input signals...

 $V_{out} = -(V_1 + V_2 + V_3)$



Subtractor



If a differential signal (ex. ECG leads, bipolar EMG) is measured across the input terminals

Differential gain
$$G_d = \frac{V_{out}}{V_2 - V_1} = \frac{R_2}{R_1}$$

Common-mode rejection ratio CMRR of the differential amplifier



If a common-mode voltage at both inputs is $V_{cm}=(V_1+V_2)/2$

Then the common-mode gain =

$$G_{cm} = \frac{V_{out}}{V_{cm}} = \frac{R_1 R_4 - R_2 R_3}{R_1 (R_3 + R_4)}$$

CMRR is defined as:

$$CMRR = 20\log \left| \frac{G_d}{G_{cm}} \right|$$

Homework:

- Derive the expression for $G_d=V_{out}/V_d$ (in terms of $R_1 \sim R_4$) with a differential input $V_d=V_2-V_1$
- Suppose you use 4 resistors 100KΩ±1%, calculate the CMRR 10 times using random numbers for errors in resistance

More on differential amplifier



- For measuring biopotentials, voltage gain can be obtained by subsequent amplifier stages
- Input impedance is small ~R

• In ECG, the impedance of skin is $\sim M\Omega$ (can be lowered to 15-100K Ω by applying electrolyte gel)

• Mismatches in R reduce the CMRR

Instrumentation amplifier



Instrumentation amplifier v_1 R R 3 R_{gain} V_{out} R R R ٧, 2R $G_d = 1$ $G_{cm} \approx 0$ $G_d = 1 +$ $R_{_{gain}}$

Provides good CMRR without the need for precisely matching resistors

 $G_{cm} = 1$

Example of common-mode voltage

Interference from power line (60Hz) can induce current i_{db}



Driven-right-leg circuit

Output is connected to the right leg through a surface electrode, which provides negative feedback



Driven-right-leg circuit



Time-varying signals

 Any signal can be decomposed into a series of sinusoidal waveforms with various frequencies (Fourier transform)

• In other words, we only need to describe/model a single sinusoidal waveform and the results can be generalized to any waveform that might occur in the real world

Sinusoidal signals have amplitude, frequency and phase

$$v_1(t) = V_1 \cos(\mathbf{w}t + \mathbf{q}) = V_1 \cos(2\mathbf{p}ft + \mathbf{q})$$

<u>Phasors</u>: complex numbers (magnitude and phase angle) representing the sinusoidal signal (without the frequency)

$$\hat{V}_1 = V_1 e^{j\boldsymbol{q}} = V_1 \angle \boldsymbol{q} \qquad e^{j\boldsymbol{q}} = \cos \boldsymbol{q} + j\sin \boldsymbol{q}$$

Time-varying signals and circuits

Since capacitors and inductors introduce phase shift to the signal, their impedances Z can be expressed in phasors as following

$$\hat{V}_{1} = Z\hat{I} \qquad Z_{R} = R$$

$$Z_{L} = j\mathbf{w}L = \mathbf{w}Le^{j\mathbf{p}/2}$$

$$Z_{C} = \frac{1}{j\mathbf{w}C} = \frac{1}{\mathbf{w}C}e^{-j\mathbf{p}/2}$$

For example, the voltage across a capacitor is generated by electric charges accumulated in the capacitor current leads voltage

Laplace domain analysis

Use <u>Laplace transform</u> (time-domain s-domain) to describe timevarying signals \Rightarrow Differential equations become algebraic equations



Inverse Laplace transform is used when we want to obtain the time-domain signals (ex. transient response)

Laplace transform

Definition

$$L\{f(t)\} = F(s) = \int_0^\infty f(t)e^{-st}dt$$

Some properties of Laplace transform

Operation	Time Function	Laplace Transform
Linear combination	Af(t) + Bg(t)	AF(s) + BG(s)
Multiplication by e^{-at}	$e^{-at}f(t)$	F(s + a)
Multiplication by t	tf(t)	-dF(s)/ds
Time delay	$f(t-t_0)u(t-t_0)$	$e^{-it_0}F(s)$
Differentiation	f'(t)	$sF(s) = f(0^-)$
	f''(t)	$s^2 F(s) - sf(0^-) - f'(0^-)$
Integration	$\int_{0^{-}}^{t} f(\lambda) \ d\lambda$	$\frac{1}{s} F(s)$

Laplace transform pairs

f(t)	F(s)
А	$\frac{A}{s}$
u(t) - u(t - D)	$\frac{1 - e^{-sD}}{s}$
t	$\frac{1}{s^2}$
t	$\frac{r!}{s^{r+1}}$
e^{-at}	$\frac{1}{s+a}$
te ^{-at}	$\frac{1}{(s + a)^2}$
$t^{r}e^{-at}$	$\frac{r!}{(s+a)^{r+1}}$
$\sin \beta t$	$\frac{\beta}{s^2 + \beta^2}$
$\cos (\beta t + \phi)$	$\frac{s\cos\phi-\beta\sin\phi}{s^2+\beta^2}$
$e^{-at}\cos(\beta t+\phi)$	$\frac{(s+a)\cos\phi-\beta\sin\phi}{(s+a)^2+\beta^2}$

Transfer function

Relationship between the input and output

$$T(s) = \frac{V_{out}(s)}{V_{in}(s)}$$

• Since s = jw = j2pf

T(s) also provides information on the frequency and phase of the circuit – frequency response



Transfer function – example



$$Z_{i}(s) = R_{1} + \frac{1}{sC_{1}} = \frac{sR_{1}C_{1} + 1}{sC_{1}} \qquad Z_{f}(s) = \frac{1}{\frac{1}{R_{2}} + sC_{2}} = \frac{R_{2}}{1 + sR_{2}C_{2}}$$

$$T(s) = -\frac{Z_f(s)}{Z_i(s)} = \frac{sR_2C_1}{(1+sR_1C_1)(1+sR_2C_2)}$$

26

Transfer function – example2



Frequency response

 The transfer function can be factored into poles and zeros

$$T(s) = K \frac{(s+z_1)(s+z_2)\cdots}{(s+p_1)(s+p_2)\cdots}$$

Alternatively

$$T(s) = K' \frac{(1+s/z_1)(1+s/z_2)\cdots}{(1+s/p_1)(1+s/p_2)\cdots} = K' \frac{(1+jw/z_1)(1+jw/z_2)\cdots}{(1+jw/p_1)(1+jw/p_2)\cdots}$$

$$T(jw) = |T(jw)|e^{jq(w)}$$
Magnitude response
Phase response

Frequency response – LPF



Magnitude response

$$|T(j\mathbf{w})| = |K'| \frac{1}{\sqrt{1 + \frac{\mathbf{w}^2}{p_1^2}}} \qquad \text{Gain} \quad \begin{cases} |K'| = R_2/R_1 & \text{DC} (\omega=0) \\ \frac{K'}{\sqrt{2}} & \text{When } \omega = p_1 \end{cases}$$

Frequency response– LPF

At the cut-off frequency f_c : the magnitude response is

$$\left|T(j2\mathbf{p}f_c)\right| = \frac{\left|T(j\mathbf{w})\right|_{\max}}{\sqrt{2}}$$

(-3dB power attenuation)

In this example

$$\boldsymbol{w}_c = 2\boldsymbol{p}f_c = p_1$$
$$\therefore f_c = \frac{1}{2\boldsymbol{p}R_2C}$$



Frequency response – HPF



$$|T(jw)| = \frac{wR_2C}{\sqrt{1+w^2R_1^2C^2}} \qquad \text{Gain} \quad \begin{cases} 0 & \text{DC }(\omega=0) \\ -\frac{R_2}{R_1} & \text{When } \omega \end{cases}$$
$$w_c = 2pf_c = p_1 \qquad \therefore f_c = \frac{1}{2pR_1C}$$

Frequency response – HPF, BPF



Homework:

• For the HPF shown in slide 31, show that the magnitude response is $\frac{1}{\sqrt{2}}$ of the maximum at the cut-off frequency ω_c

Active filters

Frequency characteristics of analog filters



The most important is the amplitude response which represents how the amplitudes of different frequency components are modified by the filter

Active filters

Response of low-pass Butterworth filters with different orders (-3dB frequency is normalized at 1)



Sharper knee with higher orders

Butterworth filter of order n

$$\frac{V_{out}}{V_{in}} = \frac{1}{\sqrt{1 + (f / f_c)^{2n}}}$$

Chebyshev filter of order n

$$\frac{V_{out}}{V_{in}} = \frac{1}{\sqrt{1 + e^2 C_n^2 (f / f_c)}}$$

 C_n is the Chebyshev polynomial of the first kind of degree n, is a constant that sets the passband ripple

Active filters

Comparison of several 6-pole low-pass filters



Transfer Function	Frequency-Domain Cha	Time-Domain Characteristics		
	Ripple	Stopband	Phase	Group Delay
Chebyshev	Equal ripple flat	Steep	Poor	Poor
Butterworth	Smooth	Moderate	Moderate	Moderate
Bessel	Maximum smoothness	Weak	Very flat	Very flat

Active filter circuits – VCVS



es	Butter- worth K	Bessel		Chebyshev (0.5dB)		Chebyshev (2.0dB)	
Pol		fn	к	f _n	к	fn	к
2	1.586	1.272	1.268	1.231	1.842	0.907	2.114
4	1.152	1.432 1.606	1.084 1.759	0.597 1.031	1.582 2.660	0.471 0.964	1.924 2.782
6	1.068 1.586	1.607 1.692	1.040 1.364	0.396 0.768	1.537 2.448	0.316 0.730	1.891 2.648
	2.483	1.908	2.023	1.011	2.846	0.983	2.904
8	1.038 1.337 1.889	1.781 1.835 1.956	1.024 1.213 1.593	0.297 0.599 0.861	1.522 2.379 2.711	0.238 0.572 0.842	1.879 2.605 2.821
	2.610	2.192	2.184	1.006	2.913	0.990	2.946



VCVS filter design

- Each circuit is a 2-pole filter; i.e. for an n-pole filter, you need to cascade n/2 VCVS sections
- Within each section, set $R_1 = R_2 = R$ and $C_1 = C_2 = C$
- Set the gain K according to the table
- For Butterworth filters

$$RC = \frac{1}{2pf_c}$$
 f_c is the -3dB frequency

- For Bessel and Chebyshew low-pass filters

$$RC = \frac{1}{2\mathbf{p}f_n f_c}$$

- For Bessel and Chebyshew high-pass filters

$$RC = \frac{1}{2\mathbf{p}f_c / f_n}$$

Non-ideal op-amp

<u>Input bias current</u> I_B : simply the base or gate currents of the input transistors (could be either current source or sink) – the effect of I_B can be reduced by selecting resistors to equalize the effective impedance to ground from the two inputs



Non-ideal op-amp

Input offset current I_{OS} : difference in input currents between two inputs; typically 0.1~0.5 I_B

<u>Input offset voltage</u>: the difference in input voltages necessary to bring the output to zero (due to imperfectly balanced input stages)



The offset voltage can be eliminated by adjusting null offset pots on some op-amps

Non-ideal op-amp

<u>Voltage gain</u>: typically 10^5 - 10^6 at DC and drops to 1 at some f_T (~ 1-10 MHz); when used with feedback (closed-loop gain = G), the bandwidth of the circuit will be f_T/G





<u>Output current</u>: due to limited output current capability, the maximum output voltage range (swing) of an op-amp is reduced at small load resistances

Practical considerations

- Negative feedback (resistor between the output and the inverted input terminal) provides a linear input/output response and in general stability of the circuit
- Choose resistor values $1k\Omega$ - $1M\Omega$ (best $10k\Omega$ -100k Ω)
- Match input impedances of the two inputs to improve CMRR
- Equalize the effective resistance to ground at the two input terminals to minimize the effects of I_B

Matching effective impedance to ground

The voltage gain is 5 for both circuits



40KΩ 10KΩ = 8KΩ

So the effective impedance to ground from both input terminals is the same

Noise

- Interference from outside sources
 - Power lines, radio/TV/RF signals
 - Can be reduced by filtering, careful wiring and shielding
- Noise inherent to the circuit
 - Random processes
 - Can be reduced by good circuit design practice, but not completely eliminated

Signal-to-noise ratio

$$SNR = 20 \log \left| \frac{V_{s(rms)}}{V_{n(rms)}} \right| \quad dB \qquad V_{rms} = \left[\frac{1}{T} \int_0^T v^2(t) dt \right]^{1/2}$$

Noise

- Types of fundamental (inherent) noise:
 - Thermal noise (Johnson noise or white noise)
 - Shot noise
 - Flicker (1/f) noise

Noise

Thermal noise: generated in a resistor due to thermal motion of atoms/molecules

$$V_{noise}(rms) = \sqrt{4kTRB}$$

k: Boltzmann's constant

- T: absolute temperature (°K)
- R: resistance (Ω)
- B: bandwidth f_{max}-f_{min}

Thermal noise contains superposition of all frequencies \Rightarrow white noise

Shot noise: arises from the statistical uncertainty of counting discrete events

Shot noise =
$$\sqrt{\frac{dn}{dt}}\Delta t \approx \sqrt{n}$$

 $S / N = \frac{n}{\sqrt{n}} = \sqrt{n}$

dn/dt is the count rate Λt is the time interval for the measurement

Flicker (1/f) noise: power spectrum is $\sim 1/f$; somewhat mysterious; found related to resistive materials of resistors and their connections.

Interference

Electric fields existing in power lines can couple into instruments and even the human body (act as capacitors)



Electromagnetic interference

Magnetic fields in the environment can be picked up by a conductor and results in an induced current



Electromagnetic interference

Time-varying magnetic field induces a current in a closed loop



Reduce induced current by minimizing the area formed by the closed loop (twisting the lead wires and locating close to the body)

Electrical safety

Physiological effects of electricity (for a 70kg human)



- Threshold of perception: >0.5mA at 60Hz and >2-10mA at dc
- Let-go current: the <u>maximal</u> current at which the subject can withdraw voluntarily (>6mA)

• Respiratory paralysis: involuntary contraction of respiratory muscles (>18-22mA)

• Ventricular fibrillation: the current excites part of the heart muscle (>75-400mA)

Electrical safety

The effects of electricity depend on many conditions such as sex, frequency, duration, body weight and points of entry



- The mean value for threshold of perception is 0.7mA for women and 1.1mA for men
- The mean let-go current is 10.5mA for women and 16mA for men

Macroshock vs. microshock



The risk of fibrillation is small due to wide distribution of current through the body (only a small fraction flows through the heart)

Fibrillation can be caused by microshock currents $80-600\mu A$ For safety, the limit to prevent microshocks is $10\mu A$

Let's start with the power line...

Simplified electric power distribution circuits



It's the "hot" lines that are at high voltages to ground

Macroshock hazards



Electric faults

 happen when the hot conductor (high voltage) gets in contact with metal chassis or cabinet that is not grounded properly

 can be caused by failures of insulation, shorted components (e.g. due to mechanical failure), strain and abuse of power cords, plugs and receptacles

Microshock hazards

Generally result from leakage currents

- \bullet small currents (~ μA) flow between two adjacent conductors that are insulated from each other
- mostly flow through capacitance between the two conductors
- some are resistive through insulation, dust, or moisture



Microshock hazards

Another example: ground potential differences (when "ground" is no longer at ground) \Rightarrow current flows from one "ground" to another <u>through the patient</u>



Solution – isolated power distribution

<u>Ground fault</u>: a short circuit between the hot conductor and ground injects large currents into the grounding system

• the hot conductors can be isolated from ground using an isolation transformer



Power-isolation transformer system

If there is only one ground fault between one of the conductors and ground, there will be no surge current. This fault can be detected by the monitor system (and removed to prevent real hazard to the patients).

Solution – grounding system

- All the receptacle grounds and conductive surfaces in the vicinity of the patient are connected to the <u>patient-</u> <u>equipment grounding point</u> (with resistance = 0.15Ω)
- The difference in potential between the conductive surfaces must be = 40mV
- Each <u>patient-equipment grounding</u> <u>point</u> is connected individually to a <u>reference grounding point</u> that is in turn connected to the building ground



Solution – electrical isolation

To prevent leakage currents going through the patient's heart directly (microshocks), all patient leads need to be isolated electrically from the AC power lines \Rightarrow isolation amplifiers

- break the ohmic continuity of electric signals between the input and output \Rightarrow impedance across the barrier > 10M Ω
- include different supply-voltage sources and different grounds on each side of the isolation barrier



Isolation amplifier example

Optically coupled signal transmission via LED and 2 matched photodiodes



Other examples of isolation amplifiers include those using <u>transformers</u> and <u>capacitors</u> (signal is usually frequency-modulated)

A/D conversion

Conversion of Analog signal to Digital (integer) numbers



Continuous time \rightarrow discrete time interval ΔT

A/D conversion is a process to

- "Sample" a real world signal at finite time intervals
- Represent the sampled signal with finite number of values

Sampling rate (frequency)

How fast do we need to sample? First define the sampling frequency:

$$f_{sampling} = \frac{1}{\Delta T}$$
 (sample/s)

Intuitively, we must sample <u>fast enough</u> to avoid distortion of the signal or loss of information \Rightarrow easier to explain in the frequency domain

 $f_{sampling} > 2f_{max}$ (sampling theorem)

where f_{max} is the highest frequency present in the analog signal

What happens if the above criterion is not met?

- Loss of high frequency information in the signal
- Even worse, the data after sampling may contain false information about the original signal \Rightarrow frequency aliasing

Sampling

In the frequency domain, sampling of the signal at f_{sampling} results in duplicates of the spectrum that are shifted by $m_{f_{\text{sampling}}}$ (m is an integer)



The sampling theorem essentially requires the spectrum of signal not overlapping with its duplicates

Frequency aliasing

When the sampling theorem condition is not satisfied



The high-frequency region overlaps and shape of spectrum is changed (summed). The process is <u>not reversible</u> \Rightarrow information is lost

Anti-aliasing

In the real world, no signal is strictly band-limited. But an effective bandwidth can be defined and used to find the sampling frequency
To avoid frequency aliasing, a low-pass filter is applied to the signal prior to sampling



Data acquisition hardware

Lots of commercial products to choose from. <u>National Instruments</u>, for example, has families of products with a variety of features



Data acquisition hardware

Examples from National Instruments			Input	Aggregate		
	Product	Bus	Analog Inputs ¹	Resolution (bits)	Sampling Rate (kS/s) ²	Input Range (V)
	PCI-6014	PCI	16 SE/8 DI	16	200	±0.05 to ±10
	PCI-6013	PCI	16 SE/8 DI	16	200	±0.05 to ±10
	PCI-6010	PCI	16 SE/8 DI	16	200	±0.2 to ±5
¹ SE–Single-ended, DI–differential ² All channels share one analog-to-digital converter.						

Input resolution: for 16 bits $\Rightarrow 2^{16}$ digital levels

If the input range is $\pm 5 V$, the minimum detectable signal level is

$$\frac{10V}{2^{16}} = \frac{10V}{65535} = 0.15mV$$

In practice, it is desirable to match the range of analog signal to the input range of the data acquisition hardware to increase the overall resolution of amplitude sampling

References

- The Art of Electronics (2nd ed.), by Paul Horowitz and Winfield Hill
 - Ch5: Active filters
 - Ch7: Precision circuits and low-noise techniques
- Medical Instrumentation: application and design, 3rd ed., edited by John G. Webster
 - Ch3: Amplifiers and Signal Processing
 - Ch14: Physiological effects of electricity