#### SYLLABUS IS TENTATIVE IT WILL BE UPDATED on a WEEKLY BASIS ACCORDING to PROGRESS in CLASS

Course Code Course Title Instructor	BME3142 Biomedical Modeling and Simulation Kamuran A. Kadıpaşaoğlu <u>kamuran@yildiz.edu.tr</u> <u>http://avesis.yildiz.edu.tr/kamuran/</u>
Graduate Assistant	Ümmühan Zengin Özer
Textbooks	Guyton AC and Hall JE <u>Textbook of Medical Physiology, 11<sup>th</sup> edition</u> Elsevier, Philadelphia, 2006 <u>Michael CK Khoo</u> <u>Physiologic Control Systems</u> IEEE Press Series in BME, 2000 <u>Neumann DA</u> <u>Kinesiology of the Musculoskeletal System, 3<sup>rd</sup> Ed</u> Elsevier, St. Louis, MO, 2017
Recommended Reading	<ul> <li>J.T. Ottesen, M.S. Olufsen, and J.K. Larsen</li> <li>Applied Mathematical Models in Human Physiology</li> <li>BioMath-Group, Depts Mathematics &amp; Physics</li> <li>Roskilde University, Denmark, 2003</li> <li>Batzel JJ, Kappel F, Schnedits D, Tran HT</li> <li>Cardiovascular and Respiratory Systems: Modeling,</li> <li>Analysis and Control</li> <li>SIAM (Society for Industrial and Applied Mathematics),</li> <li>Philadelphia, 2007</li> </ul>

#### **Course Objectives**

The objective of the course is to introduce and apply general theories for modelling and simulation of systems relevant within biomedical engineering. This includes both physical and physiological models.

#### Course Content

Introduction to concepts of system modeling, model formalism and its relationship to different simulation strategies. Application of general and specific methods to analyze and model systems. Implementation and simulation of models in a computing environment (Matlab/Simulink). Discrete-time and stochastic simulation methods. Evaluation of model applicability, accuracy, and robustness.

#### **Course Learning Outcomes**

- **1.** Identify and describe general principles for modeling and simulating a system.
- 2. Apply these principles when designing mathematical models for a number of realistic systems.
- **3.** Implement and use computer-based modeling and simulation for studying research relevant problems within the field of biomedical engineering.
- **4.** Analyze simulation results obtained from produced models using visualizations, graphics, and other tools.
- 5. Evaluate the applicability and usability for different models and simulation techniques.

#### Program Outcomes (PO)

- **1.** Understanding of the ethics and principles of engineering.
- **2.** Understanding the concept of biomedical engineering and its main fields of study.
- 3. Learning the basic terminology and the analogies between systems used in biomedical engineering.
- **4.** To gain awareness about entrepreneurship, project management, sustainability in engineering.
- **5.** Establishing effective oral and written communication

#### Program Outcomes (PO) Relevant to the Course

- 4. PO-2.1) Ability to define, formulate and solve complex engineering problems.
- 5. PO-2.2) Ability to choose and apply appropriate analysis and modeling methods for this purpose.
- 8. PO-4.1) Ability to develop, select and use modern techniques and tools necessary for the analysis and solution of complex problems encountered in engineering applications.
- 9. PO-4.2) Ability to use information technologies effectively.

# **Course Learning Outcomes & Program Outcomes Matrix**

<u>CLO</u>	1	2	3	4	5
PO-4	5	5	4	5	3
PO-5	5	5	5	5	5
PO-8	3	3	5	5	5
PO-9	3	3	5	5	4

Welcome to Class everyone.

Before starting this semester, it may be wise to be advised about the course organization so as to maximize your classroom performance and get the most fun out of the experience.

To begin, please follow the steps below.

- 1. Go to <u>http://avesis.yildiz.edu.tr/kamuran/</u>
- 2. Click on My Shares (Duyuru ve Dokümanlar)
- 3. Find the file "**Rules and Guidelines**"
  - a. Download & save and print it
  - b. Read, understand, and even memorize its contents.
  - c. Pay particular attention to Homework Policy and Grading Policy
- On the same page, find the file
   "YourCourseCode\_Syllabus\_Semester\_Year\_#".
  - a. The file will be updated on a weekly basis.
  - b. Make sure you read the assignment(s) **BEFORE** coming to each class, **STARTING THIS WEEK**!
  - c. Download and save any **Textbook**(s) that is (are) posted.
  - 5. It is highly recommended that you
    - a. Read the assigned chapter
    - b. Solve the example and homework problems
    - c. Prepare your questions for me

# **BEFORE COMING TO CLASS.**

6. There will be a quiz assignment every week.

I am looking forward to an educational and productive semester.

Good luck to all.

KAK

# HOMEWORK and GRADING POLICY

- 1. There will be a HW assignment every week, except the first week, the weeks before the exams, and the holiday weeks (if any) for a maximum total of 11 HWs.
- 2. Overall HW grade counts for 35% of final grade.
- 3. You can follow up weekly HW grades from **BME3142 GRADES**, check out the solution sheet from the updated syllabus, and see the highest and lowest scoring submissions when the Homework link on the relevant week becomes active (past the HW submission deadline).
- 4. Midterm and Final exam grades count for 30% and 40% of the final grade, respectively.

Overall Homework Grade	%40
Midterm	%20
Final	%40

- I. Classroom Rules
- II. Recap:
  - 1. Differential Equations and Laplace Transform Lecture Link 1
  - 2. System Analogies Lecture Link 2

# III. Cell Membrane: Introduction Lecture Link 3 Physiology Web

- IV. Homework
  - 1. Read <u>Academic Dishonesty</u> Krishna Bista (2011)
  - 2. Study Lecture Links above
  - 3. Study Class Notes Chapter 1 Cell Membrane
  - **4.** Prepare for quiz next week

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#### I. TRANS-MEMBRANE POTENTIAL

#### MUST STUDY: Class Notes Chapter 1 Cell Membrane

Chemical and Electrical Gradient Chemical and Electrical Flux, Fick's Law Gibb's Free Energy, Nernst Equation Gibson-Hodgkin-Katz Equation (<u>Link</u> from LibreTexts)

#### II. PERSONA

Alan Hudgkin Andrew Huxley Kenneth Cole Hugh Huxley Jean Hansen

#### III. Action Potential Propagation

Neuron Anatomy: Axon, Dendrite, Synapse Polarization, Depolarization, Repolarization External Input:

- · Depolarization: Action Potantial
- Repolarization: Na-K Pump (ATP) Rigor Mortis

#### IV. Modeling the Membrane: Hudgkin-Huxley Model

$C_{\rm m}(dV/a$	$I_{t} = I - I_{m} = I - g_{Na}m^{3}h(V - V_{Na}) - g_{K}n^{4}(V - V_{K}) - g_{L}(V - V_{L})$	V <sub>res</sub>	t = -60  mV
d <b>m</b> /dt =	$\alpha_m(1-m) - \beta_m m$	$V_{Na}$ $V_{V}$	= 50  mV = - 77 mV
$d\mathbf{n}/dt =$	$\alpha_n(1-n) - \beta_n n$	$V_{\rm L}$	= - 54.402 mV
d <b>h</b> /dt =	$\alpha_h(1-h) - \beta_h h$	$g_{\rm Na}$	$= 120 \text{ mmho/cm}^2$ $= 36 \text{ mmho/cm}^2$
where,		$g_{\rm L}$	$= 0.3 \text{ mmho/cm}^2$
V	= Membrane potential	E	$= V - V_{\text{rest}}$
1	= Sum of external and synaptic currents entering the cell	$\alpha_m$	$= 0.1(25-E)/(e^{(25-E)/10} - 1)$
$I_m$	= Membrane current	$\beta_m$	$= 4 e^{-E/18}$
<i>m</i> , <i>n</i> , <i>h</i>	= State variables	$\alpha_n$	$= 0.01(10 - E)/(e^{(10 - E)/10} - 1)$
V Na	= Equilibrium (or reversal) potential at which the net flow of Na ions is zero	<i>B</i> .,	$= 0.125e^{-E/80}$
<sup>V</sup> к	= Equilibrium (or reversal) potential at which the net flow of K ions is zero	pn a	$= 0.07e^{-E/20}$
V <sub>L</sub>	= Equilibrium (or reversal) potential at which leakage is zero	R.	$= 1/(1 + e^{(30-E)/10})$
$C_{\rm m}$	= Membrane capacitance	$p_h$	= 1/(1 + e)
$g_{_{ m Na}}$	= Sodium channel conductivity		
$g_{\rm K}$	= Potassium channel conductivity		
$g_{\rm L}$	= Leakage channel conductivity		
<i>а.<sub>т. п. h</sub></i>	= Suitable rate coefficients		
$\beta_{m, n, h}$	= Suitable rate coefficients		

$$\begin{aligned} & \alpha_n = \frac{0.01(v+50)}{1-\exp\left(\frac{-(v+50)}{10}\right)} & \beta_m = 4.0 \exp\left(-0.0556(v+60)\right) \\ & \alpha_h = 0.07\exp\left(-0.05(v+60)\right) \\ & \beta_n = 0.125 \exp\left(\frac{-(v+60)}{80}\right) & \beta_h = \frac{1}{1+\exp\left(-0.1(v+30)\right)} \\ & \alpha_m = \frac{0.1(v+35)}{1-\exp\left(\frac{-(v+35)}{10}\right)} \\ & \alpha_m = \frac{0.1(v+35)}{1-\exp\left(\frac{-(v+35)}{10}\right)} & g_{Na} = 1.2 \text{ mS/cm}^2 \\ & E_{Na} = 55.17 \text{ mV} & g_K = 0.36 \text{ mS/cm}^2 \\ & E_K = -72.14 \text{ mV} & g_l = 0.003 \text{ mS/cm}^2 \\ & E_l = -49.42 \text{ mV} \end{aligned}$$





# V. HOMEWORK

- 1. Read plagiarism article
- 2. Study:
  - Solution of 1<sup>st</sup> order diff. equation
  - <u>System Analogies</u>
  - <u>Class Notes Chapters 3 and 4</u>
- 3. Study
  - Equilibrium Potential
  - Nernst Equation
  - Gibson-Hodgkin-Katz Equation
  - Give Examples
- 4. Study:
  - Proteins and Hudgkin-Huxley Model
  - <u>Spiking Neuron Models: Single Neurons, Populations, Plasticity</u> (Chapter I-2.2: Hudgkin-Huxley Model, In Gerstner and Kistler, Cambridge University Press, 2002)
  - <u>Neocleous and Schizas</u>: Paper format, Formulas available, Simulink complicated
  - <u>Murat Sağlam</u>: MathWorks example, No formulas, Simulink very clear.
  - Various Membrane Models
- 5. Create and simulate HH model in Simulink Bring laptop for Quiz 2

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I. Nervous Tissue/System Overview			
	<u>Nerve and Nervous System</u>		
	Action Potential Depolarization R		

Action Potential, Depolarization, Repolarization Nervous System Neuron, Axon, Dendrite, Synapse (Ramon y Cajal) Sensory (afferent) and Motor (efferent) nerve fibers Reptilian/Limbic/Cortical Brain Voluntary and Involuntary Systems Sympathetic and Parasympathetic Systems Acetylcholine (Loewi) /Atropine (Linneaus) Neurotransmitters

#### II. Introduction to Muscle

Rhoades Ch 8

Muscle Slides 1

Skeletal, Smooth, and Cardiac Muscle

Skeletal Muscle Anatomy and Physiology

Fascia, Fiber, Fibril, Filament

Sarcomere Anatomy

Actin/Myosin Bridge and Sliding Filament Theory

Active Force Generation

Force vs. Time: Twitch, Treppe (Summation), Tetanus, Parallel and Series Arrangement of Sarcomeres Active State

# III. Modeling the Muscle: Active Force-Length Curve

#### Muscle Slides 2

Contractile Element (CE, Active State) Sarcomere Length-Tension Relationship Resting vs Stretched Muscle Titin Elasticity (PE)-Nonlinear Passive Force (Preload): Initial stretch Total Force= Active Force+ Passive Force

# IV. HOMEWORK: Study

Khan Academy, <u>Biology Library</u>, Human Biology (GO to Unit 33): **The neuron** and the nervous system (all 10 entries) AND Muscles (all 4 entries) <u>Muscle Slides 3</u>

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# I. Modeling the Muscle-IV: Force-Length Relationship <u>Inbar & Adam (1976)</u> (Original Paper) <u>Simple Muscle Model</u> (Derivation of Model Equations)

- 1. Most General Muscle Equation:  $\dot{T} = -a.T + b.\dot{x} + c.x + d.AS$
- 2. Sudden Stretch
- 3. Special Cases: No Parallel Elastic Element ( $k_{PE}=0$ ) Isotonic Shortening ( $\dot{F}_{ex} = 0$ ) and linear F-v Relationship Isometric Tetanus ( $\dot{F}_{CE} = \dot{x}_m = 0$ )

# II. Modeling the Muscle-V: Force-Velocity Relationship

- 1. <u>AV Hill (1938) (Original Paper)</u>
- 2. <u>Teaching from classic papers</u> This paper provides an explanation of model's logic for
  - a. Creating the Isometric L-T Curve
  - b. Sudden Release and Isotonic Shortening
  - c. External Mechanical Work and Heat of Shortening

# III. <u>Consequences of the Hill Model</u>

# Study Well:

- 1. Non-dimensional Form of the Hill Equation
- 2. Modeling the nonlinear series elasticity
- 3. Modeling the time course of a twitch



Hyperbolic Force-Velocity Relationship

180

160

#### **HOMEWORK 4:**

# 1. STUDY and CREATE all MODELS in Matlab

a. Modeling the Active State (Tension vs. Length Relationship)

Inbar & Adam (1976) Theory Guidelines Model\_Guidelines

# b. Hill Model (Muscle Power: Force-Velocity Relationship)

Teaching from classic papers

- Derive the non-dimensional version of Hill Equation
- Starting from Hill equation, derive an expression for the velocitydependent damping coefficient of the muscle.
- Create an m.file to reproduce the F-v relationship of skeletal muscle, for varying preloads and afterloads.
- Get inspired by the Matlab model in it but do not copy it.
- Add Figure Legends and Axis Labels
- Produce **constant-afterload** gridlines.
- c. Modeling the Passive Force

# **Consequences of the Hill Model**

**Problem 1** (McMahon TA, Chapter 1, Solved Problems):

- Model active force (CE) and passive force (SE)
- Model the exponential decay of isometric force
- Modeling the Tetanus

Duration of stimulation is  $t_1$  (C in Problem) Duration of rest between stimulations is  $t_2$  (A in Problem) Time between two consecutive stimulations is  $T=t_1+t_2$ Maximum force that can be reached is AS (active state at tetanus) AS will be achieved at  $t=nT+t_1$ 

Model as much as you can, from this point on, about the contractile properties of the muscle (characterized by b and k) in terms of n and T.

2. Bring laptop for quiz

I. REFLEX MODEL (<u>Lecture 2<sup>nd</sup> hour</u>):

Feedback Control of Reflex (Modeling the Muscle-Spindle System) Soechting et al 1971 (Original Paper) Khoo NM Reflex: Textbook version using Torsional Mechanical System (Moments and Angles as variables)

<u>Class Notes: Spinal Reflex Mechanism</u>: Derivation of Textbook version equations using Translational Mechanical System (Forces and Distances)

- a. Derive the Khoo Model Equations
- b. Create Reflex Model in Simulink for different versions of  $\beta$
- c. Add transport Delay,  $T_d$

#### II. CARDIOVASCULAR SYSTEM-I: INTRODUCTION (<u>Lecture 3<sup>rd</sup> hour</u>) <u>Guyton Ch.9</u> and/or <u>Meiss Ch 10</u>

- 1. Anatomy
  - a. Nouns and Adjectives
  - **b.** Cardiac Chambers
  - c. Cardiac Valves
  - d. Major Arteries and Veins
  - e. Systemic and Pulmonary Circulations



<u>Ashkan Jamali</u>, Study of the influence of different inflow configurations on Computational Fluid Dynamics in mechanical heart valve prostheses, MS Thesis, February 2017

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# HOMEWORK 5 for Quiz 4

# 1. Spinal Reflex Model: Prepare Working Simulink Model

- a. Derive the Khoo Model Equations. Use Translational (Force and Displacement) instead of Rotational (Moment and Angle) system
- b. Create the Model in Simulink for different versions of  $\beta$
- c. Add transport Delay,  $T_d$
- d. Discuss
- 2. Cardiovascular System: Read and Learn

Guyton Ch.9 and/or Meiss Ch 10

#### MIDTERM PREPARATION

- 1. Study <u>Problem 1</u> for tetanus. Assume
- 2. Study the modeling of the time course of a twitch based on <u>Consequences</u> <u>of the Hill Model</u>
- 3. Study:
  - a. <u>Guyton Ch.9</u> and/or <u>Meiss Ch 10</u>
  - b. https://folk.ntnu.no/stoylen/strainrate/What does.html
  - c. <u>ClassLecture-Introduction</u>
  - d. <u>Class Notes: Intro to CVS</u>

- 6 No Quiz INTRO to HEART <u>(WATCH ppt)</u> (Lecture Notes 2)
  - I. CARDIOVASCULAR SYSTEM-I: INTRODUCTION Guyton Ch.9 and/or Meiss Ch 10
    - 1. Anatomy
      - a. Nouns and Adjectives
      - b. Cardiac Chambers
      - c. Cardiac Valves
      - **d.** Major Arteries and Veins
      - e. Systemic and Pulmonary Circulations

<u>Ashkan Jamali</u>, Study of the influence of different inflow configurations on Computational Fluid Dynamics in mechanical heart valve prostheses, MS Thesis, February 2017



- 2. Wiggers Diagram (<u>Watch</u>)
  - a. Pressures (LV, Ao, LA) and LV Volume
  - **b.** Exponential (1<sup>st</sup> order) Response
  - c. Cardiac Phases
    - i. Diastole (IVR and Filling)
    - ii. Systole (IVC and Ejection: Auxotonic Contraction)

#### 3. PV Loop

- a. Opening and Closure of the Valves
- b. Cardiac Phases
- c. Ventricular Stiffness (P-V Relationship)



II. Modeling Cardiac Physiology (RC Circuits)

# 1. Hydraulic Analogue



Read: Westerhof, N., Lankhaar, JW. & Westerhof, B.E. The arterial Windkessel. *Med Biol Eng Comput* **47**, 131–141 (2009). https://doi.org/10.1007/s11517-008-0359-2

# 2. Electrical Analogue: The Windkessel Simplification

This is a subject that we have already covered in relation to 1<sup>st</sup> order system response. Here, the exponential response of the arterial system is calculated and the shape of the aortic pressure during systole and diastole are explained. Make sure that you can understand the derivation and interpretation of EACH equation.



Garrett AS, Pham T, Loiselle D, Han JC, Taberner A. Mechanical loading of isolated cardiac muscle with a real-time computed Windkessel model of the vasculature impedance. Physiol Rep. 2019 Sep;7(17):e14184. doi: 10.14814/phy2.14184.

# III. HOMEWORK

 Solve <u>Dynamic Model</u> using ALL concepts you studied, particularly those relevant to modeling, system analogies, isometric and isotonic contraction Windkessel (4-Node) Modeling

windkessel (4-Node) Modeling

- a. Electrical analogue circuit
- b. System dynamics
- c. State-space representation
- d. Matlab

# IV. Start working on your Midterm Exam

- a. H&H Membrane Model
  - <u>Neocleous and Schizas</u>: Paper format, Formulas available, Simulink complicated
  - <u>Murat Sağlam</u>: MathWorks example, No formulas, Simulink very clear.
- b. Inbar and Adam Muscle Model
  - <u>Theory Guidelines (Explanation of Model Logic)</u>
  - <u>Model Guidelines</u> (Explanation of Simulink Model)

- c. Hill Model for F-v Relationship
  - AV Hill (1938) (Original Paper)
  - <u>Teaching from classic papers</u>
- d. Tetanus Model <u>MacMahon's Twitch Model-Extended</u>
- e. Khoo Reflex Model
  - <u>Soechting et al 1971</u> (Original Paper)
  - <u>Khoo NM Reflex</u>: Textbook version using Torsional Mechanical System (Moments and Angles as variables)
  - <u>Class Notes: Spinal Reflex Mechanism</u>: Derivation of Textbook version equations using Translational Mechanical System (Forces and Distances)
- f. Study and Explain the Cardiovascular System
  - <u>Guyton Ch.9</u>
  - <u>Lecture Slides 6</u>
- g. Static Model (Khoo from Guyton)

Understand and explain

- Starling's Law of the Heart
- The concept of Mean Systemic Filling Pressure;
- **Build** the Simulink model

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# I. STATIC MODEL of the PERIPHERAL CIRCULATION: Cardiac Output-Venous Return Balance (Khoo)

Physiologic Control Systems

This section covers the Guyton model as described in Khoo 3.5. You should know/learn the following concepts

- Length-Tension Relationship (Skeletal Muscle)
- Pressure-Volume Relationship (Cardiac Muscle)
- Cardiac P-V Loops
- Cardiac Cycle
- Wiggers Diagram
- Stroke Volume, Heart Rate and Cardiac Output
- Stroke Work
- Contractility
- Elastance vs Compliance of LV
- Electrical/Hydraulic Modeling

# **II. MODELING IN LIFE SCIENCES**

- 1. Uses
  - a. Educational
  - b. Performance Testing of Novel Devices
- 2. Types
  - a. in silico (Computational/Numeric/Mathematical)
  - *b. in vitro* (Physical/Benchtop)
  - c. in vivo (Experimental/animal)
  - d. Clinical (Human trials in phases)
- 3. Model Accuracy vs. Model Complexity
  - a. *in silico* model can be made as complex as desired for high accuracy
  - b. in silico model must be duplicated in vitro for
    - i. reproducibility of *in silico* environment
    - ii. duplicability of *in silico* findings
  - c. Therefore, complexity of *in silico* model is limited for practical considerations

#### III. From *in silico* to *in vitro*

- 1. Modeling Ventricles
  - a. Piston-in-cylinder
  - b. Fluid-filled sac
  - c. Flexible membrane
- 2. Modeling Capacitances
  - a. Constant Capacitance (see below, Section II:2-a.ii)
  - b. Variable Capacitance (next week)
- 3. Modeling Pressures
  - a. Low Pressure ( $P_{LA}, P_V \le 20 \ mmHg$ ) Hydraulic pressure only
  - b. High Pressure ( $P_{LA}, P_V \ge 80 \ mmHg$ ) Hydraulic plus Pneumatic Pressure

# WK HR SUBJECT

8 Holiday

# WKHRSUBJECT9MIDTERM EXAM

Open Notes Laptop with MATLAB No Internet

# 10 PNEUMATIC SYSTEM

1. Analogy

Independent Variable	Mass	m
Dependent Variables		
Effort	Pressure	Р
Motion	Mass Flow Rate	'n
System Constants		
PE Storage	Variable Capacitance	RT/V
HE Dissipation	Valve Resistance	1/g

# 2. Governing Equations



- a. Case 1: Single Tank with Constant Volume
  - i. Ideal Gas Law

$$P_2 = P_{atm} = 0$$
 (gauge pressure)

ii. Capacitance Equation

$$PV = mRT = \rho VRT$$

$$P = \rho RT = \frac{m}{V}RT = \frac{RT}{V}m = e_pm$$

Pneumatic Elastance 
$$\triangleq e_p = \frac{1}{C_p} = \frac{RT}{V}$$

#### iii. Kirchhoff's Current Law

$$\dot{m}_1 - \dot{m}_2 = \dot{m}$$

iv. Ohm's Law

$$P_C - P_1 = R_1 \dot{m}_1$$
$$g_1 (P_C - P_1) = \dot{m}_1$$

3. Putting it all Together

$$\dot{P}_1 = ?$$

# IV. HOMEWORK (Use this format)

- 1. Guyton Circulatory Equilibrium Model
  - a. Matlab
  - b. Discussion
- 2. Pneumatic Modeling
  - a. Single Constant-Volume Reservoir with inlet and outlet valves

$$P_{2} = P_{compressor}$$
$$P_{2} = 0 < P_{vacuum} < P_{atm}$$

- 11 Lecture Notes <u>1<sup>st</sup> hr</u>, <u>2<sup>nd</sup> hr</u>
  - I. Left Ventricle as Pneumo-Hydraulic System



Figure 2. Detail from the MCL schematic, showing the gas pressure regulation for the preload and afterload reservoirs. Symbols are P: Pressure, Q: Flow, g: Viscous Admittance, *m*: Gas Mass Flow Rate. Subscripts are Com: Compressor, vac:Vacuum, h: Hydraulic, p: Pneumatic.

# II. System Dynamics and Lyapunov Controller

Read and Learn, but DO NOT Copy

If you cannot write **ALL** equations from memory, **DO NOT** write anything at all! Because if you do, it will be tantamount to academic suicide!

# III. HOMEWORK (<u>Use this Format</u>)

Simulink model of 1-tank system using vacuum tank to suck air out

- 1. Set reference pressure to
  - a. Constant
  - b. Square wave
  - c. Sinusoidal wave
- 2. Show reference and control pressures and valve action for each case.

# WKHRSUBJECT11Lecture 1st hr, 2nd hrModeling the LV

#### HOMEWORK

For a pneumo-hydraulic bench-top system constructed with an air compressor ( $P_c = 10 \text{ atm}$ ), a pressure reduction tank ( $P_1$ ), a tank partially containing water as shown below ( $P_T$ ), and a vacuum pump ( $P_V = 0.5 \text{ atm}$ ), all placed in series,

- 1. Calculate the dynamics of  $P_1$  and  $P_{LV}$
- 2. The control law for the valves so that the two error terms below are driven to zero

a. 
$$\epsilon_1 = 1 atm - P_1$$
  
b.  $\epsilon_2 = P_{LV} - P_T$ 

Obtain *P*<sub>LV</sub> from your electrical analogue *in silico* system

3. Operate the system in Simulink



12 Congestive Heart Failure Mechanical Circulatory Assistance Left Ventricular Assist Devices (LVAD) Preclinical Performance Testing LVAD Characteristic H-Q curves Cardiovascular Mock Circuits Hybrid Circuits

> Class Lectures  $1^{\text{st}}$  hr,  $2^{\text{nd}}$  hr,  $3^{\text{rd}}$  hr. <u>PowerPoint Presentation</u>

13 OPTIMAL CONTROL THEORY: Introduction MUST READ: <u>Class Notes (Chapter X-1): Optimal Control-Historical Background</u> Lecture 11\_Class Notes <u>1</u>, <u>2</u>, <u>3</u> Lecture 11\_<u>Recording</u>

#### I. What is Optimal in Nature

- 1. Catenary: Minimize potential energy
- 2. Stone throw: Minimize Lagrangian
- 3. Light: Minimum time or trajectory?
- 4. Plants: Minimize volume, maximize exposure to light
- 5. Ants, Spiders and Dogs: Minimize time to target
- 6. Heart: Maximize Stroke Efficiency?

#### II. Calculus of Variations:

#### <u>Historical Overview</u>: Review Article (in Turkish) <u>Lagrangian and Action</u>

**1.** Definition of the Lagrangian, L

$$\mathcal{L} = \mathcal{L} [\text{KE}(\dot{x}) - \text{PE}(x)] = \mathcal{L}(\dot{x}, x, t)$$

2. Cost Function (Action): Definite Integral of Lagrangian

$$\mathcal{A}[y(x)] = \int_{x_1}^{x_2} \mathcal{L}\left[y(x), \frac{d}{dx}y(x), x\right] dx$$

- III. Euler-Lagrange Equations Derivation of E L Equations
- IV. Tautochrone and Brachistochrone Problems Cycloid, Tauto- and Brachistochrone Equations

#### V. HOMEWORK 8:

- 1. SYLLABUS
- 2. WEEKLY SPECIAL
  - **a.** Derive the E-L Equation
  - **b.** Derive the Cartesian coordinates of a **Cycloid**
  - c. Solve 3 problems using the E-L Equation
    - i. Dog catches bone in sea
    - ii. Pendulum
    - iii. Your problem

- 14 (May 23)OPTIMAL CONTROL THEORY<br/>Lecture 12\_Class Notes 1, 2, 3<br/>Lecture Recording
  - I. Constrained Optimization Read and Study Example Problems:
    - a. Lagrange Multipliers Lagrange Multipliers

#### II. State-Space Analysis Configuration Space, Phase Space, State Space Expression of System Dynamics as All-Integral Block Diagram State Variables= Output of Integrator Blocks State Matrix, Input Matrix, Output Matrix,

- III. Beyond Euler-Lagrange Read and Study Examples: <u>Hamiltonian and PMP</u>
  - **a.** Legendre Transformation
  - **b.** Hamiltonian
  - c. Introduction to Pontryagin Maximum Principle (PMP)
    - i. Setting up the Problem in State-Space
    - **ii.** Defining the Hamiltonian
    - iii. Solving the Adjoint Dynamics

# IV. Examples

Theory and Examples

- a. Minimum Energy (Linear Pendulum, Polar Coordinates <u>Pendulum</u>
- **b.** Minimum Path (2 points, Cartesian Coordinates) <u>Channel Crossing:</u> Ferry Problem (Example 3)
- **c.** Minimum time <u>Moon Landing</u> (Example 1)
- d. Catenary, Rocket Car, Life Guard (Dog)

# V. HOMEWORK 9:

Read: <u>Class Notes 10</u>

Read: <u>Optimal Control of CV System (Article in Turkish)</u> Hamiltonian and PMP

Study: Moon Landing

#### 15 (Make up)

# OPTIMAL CONTROL THEORY Lecture 12\_Class Notes <u>1</u>, <u>2</u>, <u>3</u> Lecture Recording

(Unfortunately, I forgot to share the screen for the first hour. So, you might want to just listen for the first hour and start viewing after 59:30 (unless you want to see nothing but my face close up on the screen for a full hour)

#### I. PMP Applications to CVS

- 1. <u>Yamashiro</u>
- 2. Litvak and Yamashiro
- 3. Noldus

#### II. FINAL EXAM:

- **1.** Syllabus:
  - **a.** Correct/Update all weeks
  - **b.** Improve Examples
  - c. Solve ALL Weekly Special Problems
- **2.** Article 1 or 2 above:
  - a. Explain
  - **b.** Discuss
  - **c.** Criticize
  - d. Solve and Plot