

# BioGears: An OpenSource Human Physiology Engine

## Agenda (1:30pm – 5:00 pm)

1. Engine Overview [~50min]
  - *Bio Break [10min]*
2. Using the Graphical User Interface (GUI) [~50min]
  - *Bio Break [10min]*
3. Interacting with the Application Program Interface (API) [~50min]
4. Questions and Help [until 5pm]

## Recommended Preparation

- If you would like to follow along using the GUI (item 2), please download and unzip the Toolkit from:  
<https://biogearsengine.com/download>



# BioGears Overview

Presentation for MMVR

Presenters: Jeff Webb, Rachel Clipp, PhD, Aaron Bray  
Applied Research Associates, Inc. (ARA)

7 April 2016

# BioGears Overview Agenda

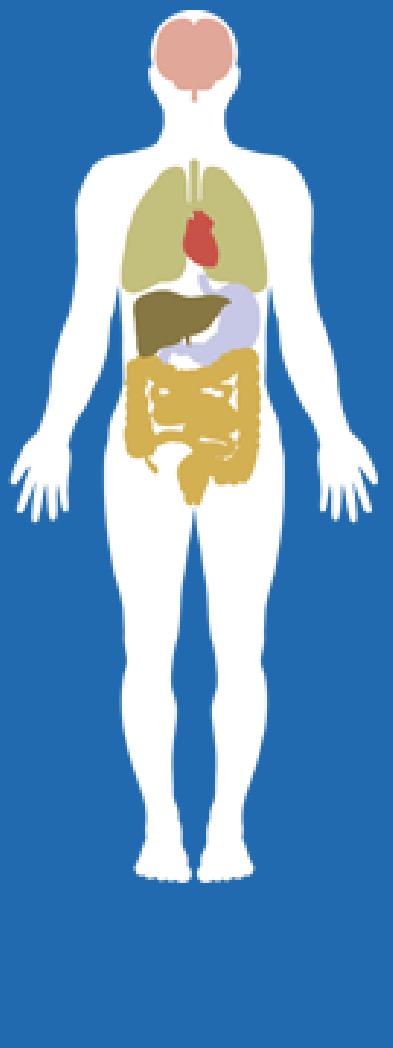
- Program Overview
- Modeling Approach
- Verification and Validation
- Systems Review
- Architecture and Tools
- Coming Soon

# PROGRAM OVERVIEW

# Project Information

- **Organization:** Applied Research Associates, Inc. (ARA)
- **Telemedicine & Advanced Technology Research Center (TATRC)**  
**Award #:** W81XWH-13-2-0068
- **Principal Investigator:** Mr. Jeff Webb
- **Amount:** \$6,959,593
- **Period of Performance:** Sept 2013 – Sept 2018
- **License:** Apache 2.0 permissive free software
- **Disclaimer:** *This work is supported by the US Army Medical Research and Materiel Command. The views, opinions and/or findings contained in this report are those of the author(s) and should not be construed as an official Department of the Army position, policy, or decision unless so designated by other documentation.*

# High Level Objectives



- Create a **publicly available** physiology research platform that enables accurate and consistent simulated **physiology** across training applications
- **Lower the barrier** to create medical training content
- **Engage the community** to develop and extend physiology models
- Meet the **training needs** of the military
- **Expand the body of knowledge** regarding the use of simulated physiology for medical education



## Patients

 Anesthesia Machine

 Inhaler

| External Systems  |
|---|
|  <b>System Interactions</b>                    |
|  <b>Environment</b><br>Substances & Thermal    |
| Internal Systems  |
|  <b>Cardiovascular</b><br>Hemodynamics & ECG   |
|  <b>Respiratory</b><br>Gas Exchange            |
|  <b>Blood Chemistry</b><br>Quantities & Panels |
|  <b>Renal</b><br>Blood Filtering               |
|  <b>Nervous</b><br>Sympathetic & Parasymp.     |
|  <b>Endocrine</b><br>Hormones                  |
|  <b>Gastrointestinal</b><br>Ingestion        |
|  <b>Energy</b><br>Thermal & Metabolic        |
|  <b>Drugs</b><br>PK/PD                       |

| Features   |                                   |  |         |            |              |             |        |         |
|--|-----------------------------------|--|---------|------------|--------------|-------------|--------|---------|
|  <b>Organs &amp; Tissues</b><br>Extravascular & Intravascular   |                                   |  |         |            |              |             |        |         |
|  <b>Homeostatic Feedback</b> <ul style="list-style-type: none"> <li>• Baroreceptors &amp; Osmoreceptors</li> <li>• Chemoreceptors</li> <li>• Local Autoregulation</li> </ul>  |                                   |  |         |            |              |             |        |         |
|  <b>Physics Based Approach</b> <ul style="list-style-type: none"> <li>• Fluid Mechanics – Gases &amp; Fluids</li> <li>• Thermodynamics – Heat Transfer</li> </ul>   |                                   |  |         |            |              |             |        |         |
|  <b>Substances</b> <table border="0"> <tr> <td>Transport, Diffusion, &amp; Clearance</td> <td></td> </tr> <tr> <td>• Gases</td> <td>• Hormones</td> </tr> <tr> <td>• Hemoglobin</td> <td>• Nutrients</td> </tr> <tr> <td>• Ions</td> <td>• Drugs</td> </tr> </table> | Transport, Diffusion, & Clearance |  | • Gases | • Hormones | • Hemoglobin | • Nutrients | • Ions | • Drugs |
| Transport, Diffusion, & Clearance  |                                   |  |         |            |              |             |        |         |
| • Gases  | • Hormones                        |  |         |            |              |             |        |         |
| • Hemoglobin   | • Nutrients                       |  |         |            |              |             |        |         |
| • Ions   | • Drugs                           |  |         |            |              |             |        |         |

| Interfaces   |
|--|
|  <b>Inputs</b> <ul style="list-style-type: none"> <li>• Parameter Setting</li> <li>• Chronic Conditions</li> <li>• Insults</li> <li>• Interventions</li> </ul>        |
|  <b>Outputs</b> <ul style="list-style-type: none"> <li>• Acute Events</li> <li>• Clinical Assessments</li> <li>• System Vitals</li> <li>• Compartment Data</li> </ul> |
|  <b>Scenarios</b> <ul style="list-style-type: none"> <li>• Static and Repeatable and/or</li> <li>• Real-Time Dynamic</li> </ul>                                      |

# Milestones

[Link to Version History](#)

|      |   |
|------|---|
| FY13 | September: project kick-off   |
| FY14 |   |
| FY15 | October: <b>Alpha Build</b> 1.0.0 release and website launch<br>March: 2.0.0 release<br>July: 3.0.0 release   |
| FY16 | October: <b>Beta Build</b> 4.0.0 release and users conference<br>December: 5.0.0 release<br>March: 5.1.0 release<br><br>Summer: planned <b>Release Candidate</b> 6.0.0 <ul style="list-style-type: none"><li>• System updates</li><li>• Serialization, modularity, and optimization</li></ul> |
| FY17 | Several releases throughout <ul style="list-style-type: none"><li>• New systems/features</li></ul>  |
| FY18 | October: <b>Final Contractual</b> release<br>Maintenance only   |



# Current Collaborators and Integrators

- 2,700+ downloads of the engine and related files since the Alpha Build Release (October 2014)
- 1,400+ pages of physiology modeling methodology and software documentation, and validation data available to our user community
- Used by the government/military, academic institutions, and commercial businesses
- Let us know if/when you use BioGears and we can add you to the home page of our website!



**SynDaver™ Labs**

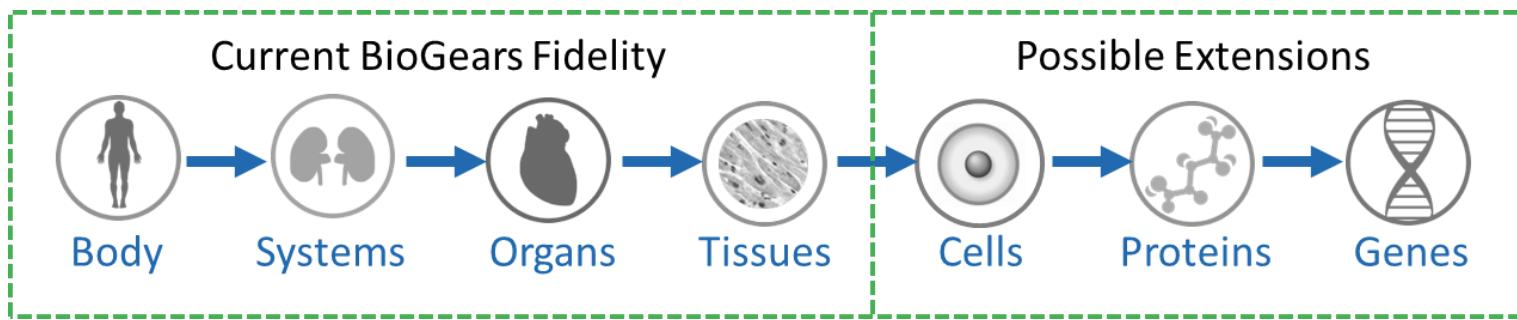


# MODELING APPROACH

# Engine Overview

## Modeling Approach:

- Top-down approach to model development with bottom-up hooks for engine expansion
- Multi-scale for varying fidelity, allowing integration of models from any level



## Computation Approach:

- Time-stepping transient analysis for linearization of differential equations
  - Currently 90Hz for 2x real-time simulation on standard laptops
- Dynamically change/add/remove elements to represent physiological mechanisms
- Stabilization analysis for initialization and implementation of conditions
- Designed with low computational overhead
  - Faster than real-time on typical PC, multiple instances on single or multicore processors
  - Build Targets include Windows, Mac, Linux, and Raspberry Pi

# Background: Lumped Parameter Modeling

- Discrete entities that approximate the behavior of a distributed system
- Electronic-Hydraulic/Thermal Analogy: body system fluid dynamics and thermodynamics modeled using electrical circuit math
- Generalized definitions of Nodes, Paths, and Elements for simple understanding, implementation, and modification

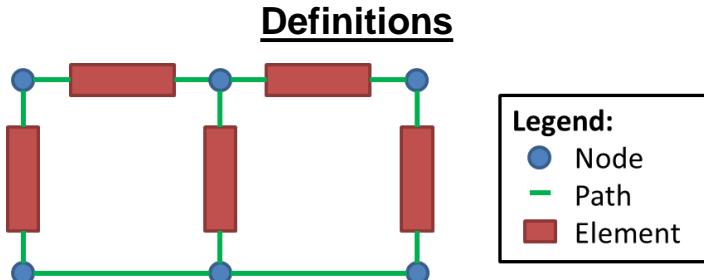
## Lumped Parameter Modeling of Fluids

$P$ =Pressure,  $F$ =Flow,  $R$ =Resistance,  $C$ =Compliance,  $I$ =Inertance

| Element         | Circuit Analogy |       | Pipe Analogy |   | Flow Equation                                |
|-----------------|-----------------|-------|--------------|---|--|
| Node            | Junction        | ●     | Junction     |    | Total Flow = 0                               |
| Path            | Wire            | —     | Rigid Pipe   |    | Solve Directly                               |
| Resistance      | Resistor        | —w—   | Aperture     |    | $F = P/R$                                    |
| Compliance      | Capacitor       | —     | Diaphram     |    | $F(t) = C \frac{dP(t)}{dt}$                  |
| Inertance       | Inductor        | —m—   | Heavy Paddle |    | $F = \frac{1}{L} \int_{t_0}^t P dt + F(t_0)$ |
| Switch          | Switch          | —\—   | Gate Valve   |    | Solve Directly                               |
| Valve           | Diode           | —>—   | Check Valve  |   | Solve Directly                               |
| Pressure Source | Voltage Source  | +—    | Pump         |  | Solve Directly                               |
| Flow Source     | Current Source  | —(+)— |              |  | $F = F$                                      |

# Lumped Parameter Approach

- Data model
  - Generic and reusable node and path definitions
  - Uses the same equations/code with native units
- Other element types not in table:
  - Switch
  - Valve/diode
  - Polarized elements

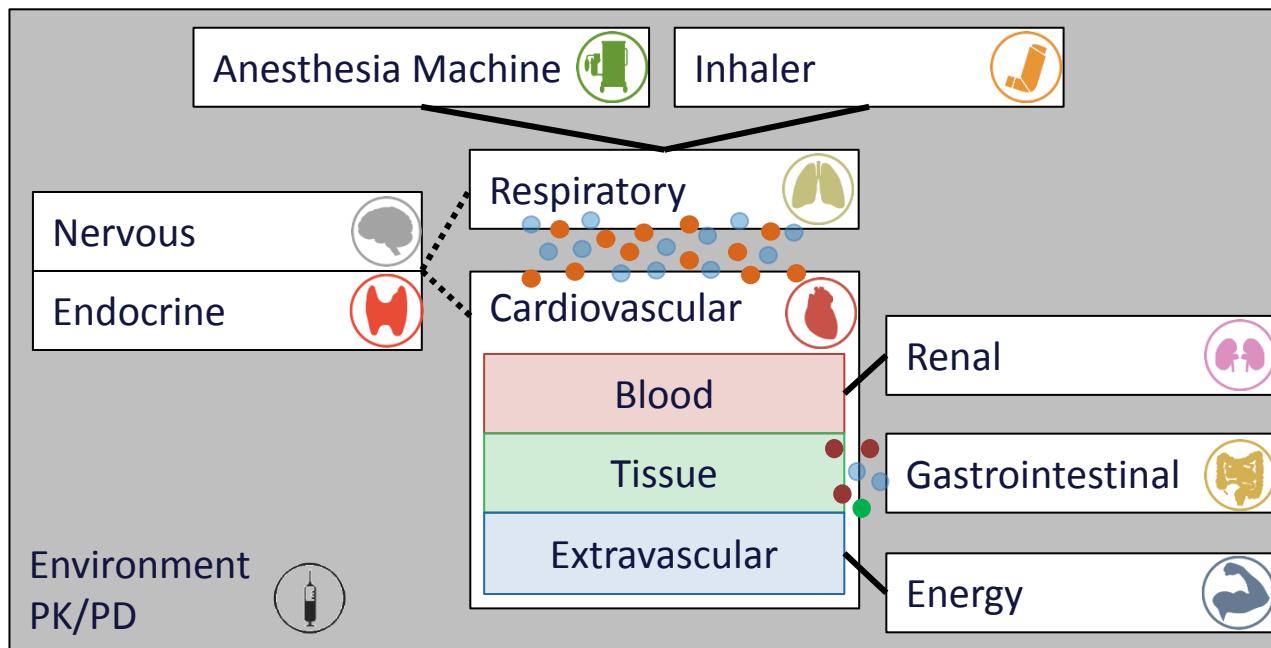


## Circuit Types and Elements

| Description |            | BioGears Included        |                  |                |                   | Other Possible Extensions |                 |                         |                      |
|-------------|------------|--------------------------|------------------|----------------|-------------------|---------------------------|-----------------|-------------------------|----------------------|
|             |            | Generic                  | Electrical       | Fluid          | Thermal           | Translational             | Rotational      | Concentrational         |                      |
| Variables   | Primary    | Across                   | Potential        | Voltage        | Pressure          | Temperature               | Velocity        | Angular Velocity        | Concentration        |
|             | Through    | Flux                     | Current          | Flow           | Heat Flow         | Force                     | Torque          | Flow                    |                      |
|             | Integrated | $\int(Across) \cdot dt$  | -----            | Magnetic Flux  | Pressure Momentum | -----                     | Displacement    | Angular Displacement    | -----                |
|             | Integrated | $\int(Through) \cdot dt$ | Quantity         | Charge         | Volume            | Heat                      | Momentum        | Angular Momentum        | Quantity             |
| Elements    | Passive    | Resistor                 | Resistance       | Resistance     | Resistance        | Resistance                | Damping         | Damping                 | Clearance            |
|             | Passive    | Capacitor                | Capacitance      | Capacitance    | Compliance        | Capacitance               | Mass            | Inertia                 | Capacitance          |
|             | Passive    | Inductor                 | Inductance       | Inductance     | Inertance         | Inductance                | Stiffness       | Torsional Stiffness     | -----                |
|             | Source     | Across Source            | Potential Source | Voltage Source | Pressure Source   | Temperature Source        | Velocity Source | Angular Velocity Source | Concentration Source |
|             | Source     | Through Source           | Flux Source      | Current Source | Flow Source       | Heat Flow Source          | Force Source    | Torque Source           | Flow source          |

Key: In the CDM

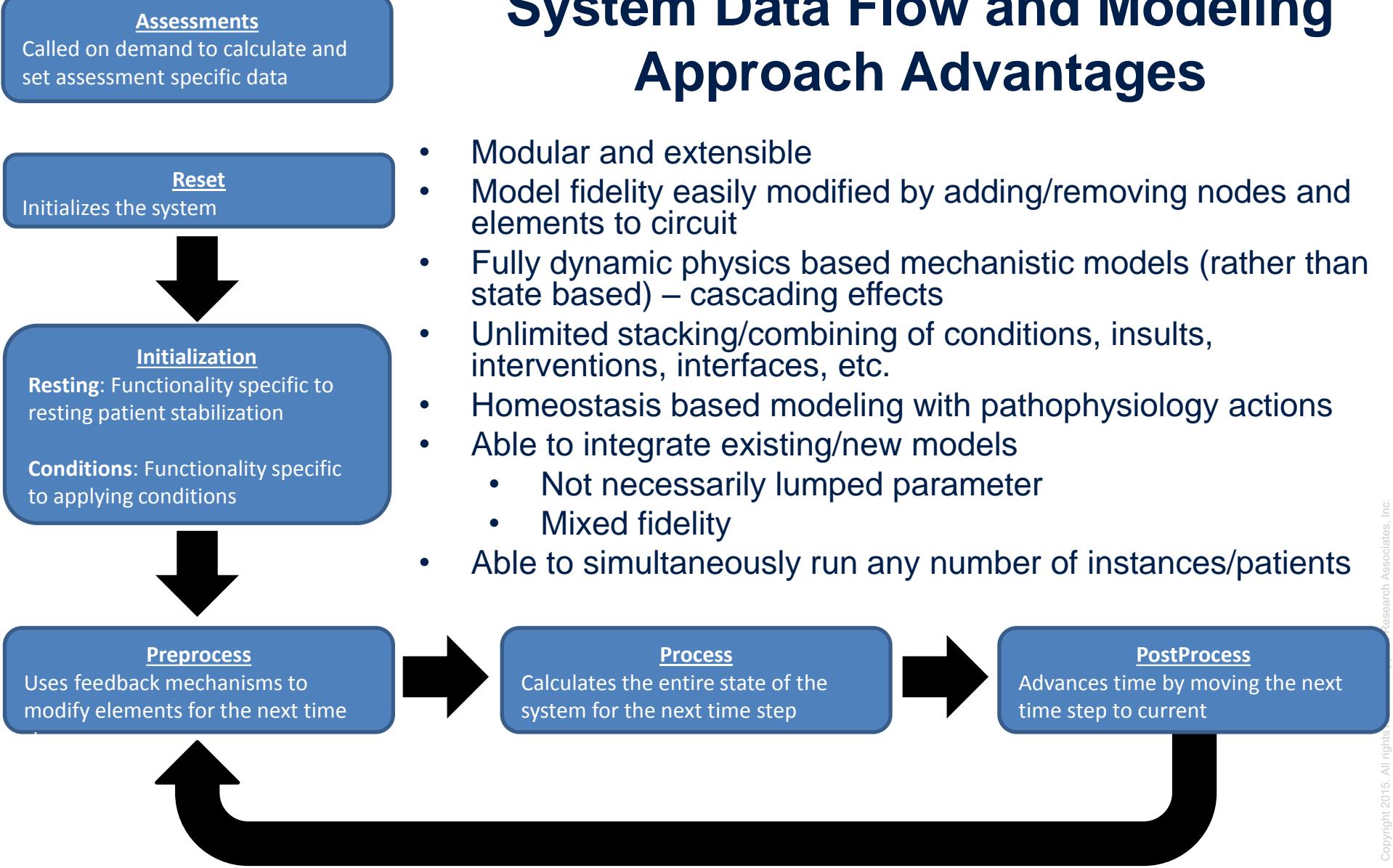
# Physiology System Interaction



## Connection types:

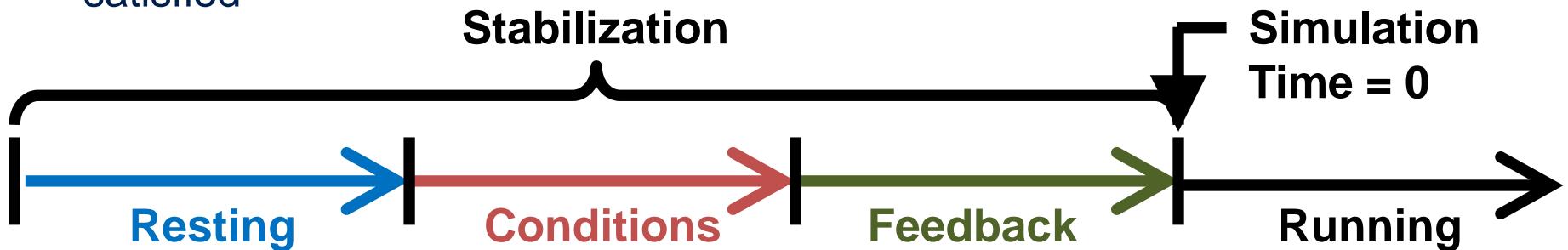
- Direct circuit connection – e.g. Anesthesia Machine and Respiratory
- Feedback – e.g. Nervous and Endocrine
- Substance exchange – e.g. Respiratory and Cardiovascular gas exchange

# System Data Flow and Modeling Approach Advantages



# Engine Initialization

- Dynamic stabilization drives towards patient homeostasis
- Each step the engine executes until all specified stabilization criteria are satisfied



## Step 1: Patient initialization

- Systems feedback modifies values to achieve specified patient parameters (e.g. baseline mean arterial pressure)
- Standard environment used

## Step 2: Condition initialization

- Conditions applied to represent new patient homeostatic state
- Environment changes applied to simulate long term acclimation

## Step 3: Feedback Mechanisms

- Feedback mechanisms that would interfere with chronic conditions (e.g. baroreceptors) are activated

## Simulation begins

- Acute insults, interventions, and parameter modification applied instantaneously through actions

## Patients

- Properties modify system setup, circuit values, and feedback parameters

## Substances

- Physical and transport related properties
- E.g. *MolarMass*, *IonicState*, *Sedation*

## Compounds

- Concentrations of multiple substances
- E.g. *Saline*, *Blood*

## Environments

- Surrounding properties
- E.g. *AmbientTemperature*, *ClothingResistance*

## Nutrition

- Meal composition (masses)
- E.g. *Protein*, *Calcium*

## Stabilization

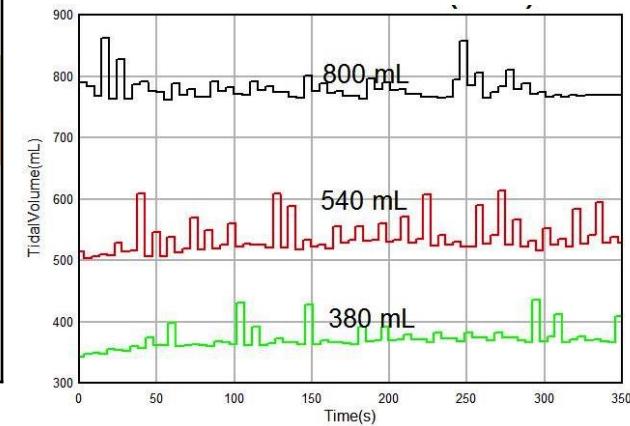
- Percent difference criteria for stabilization – resting and conditions

# Engine Data Library

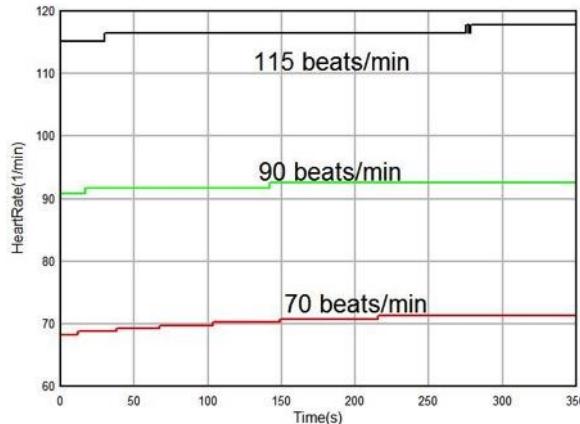
## Patient Modification Example

| Patient File Parameters |                                 |  |
|-------------------------|---------------------------------|--|
| Gender                  | FunctionalResidualCapacity      |  |
| Age                     | 2: HeartRateBaseline            |  |
| 1: Weight               | 3: MeanArterialPressureBaseline |  |
| Height                  | RespirationRateBaseline         |  |
| BodyFatFraction         | RightLungRatio                  |  |
| CarinaToTeethDistance   | TotalBloodVolumeBaseline        |  |
| Contractility           |                                 |  |

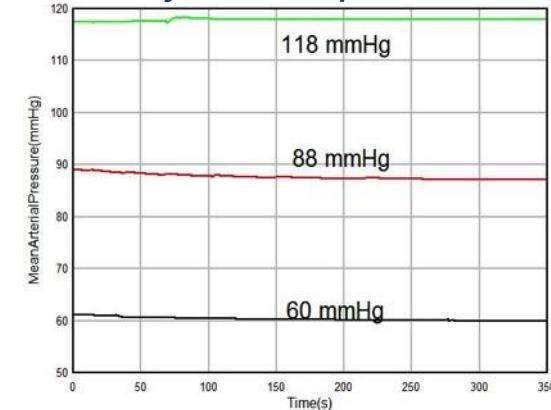
1 Tidal Volume varies with weight



2 Driver is modified to vary heart rate



3 Circuit is modified to vary arterial pressure



# Compartment Example: Kidney Definition

Left Kidney

Left Renal Vein

Left Renal Artery

Left Nephron

Left Afferent Arteriole

Left Glomerular Capillaries

Left Efferent Arteriole

Left Peritubular Capillaries

Left Bowmans Capsules

Left Tubules

Left Ureter

Key:

Vascular

Urine

- Anatomical Compartments defined by sub-circuits and allow access via an anatomy tree
- Several overlapping compartment types
  - Fluid
  - Liquid
    - Vascular
    - Urine
    - Chyme
  - Gas
    - Pulmonary
  - Tissue
  - Thermal
  - Electrical
- Compartment **properties** are combined from children (liquid example)
  - **Volume** is a sum
  - **InFlow** is a sum
  - **OutFlow** is a sum
  - **Pressure** comes from an assigned child node (sum does not make sense)
- **Substance quantities** (mass, concentration, etc.) are calculated on demand for any level in hierarchy
- There are thousands of compartment data values that are updated each time-step

# VERIFICATION AND VALIDATION

# Verification

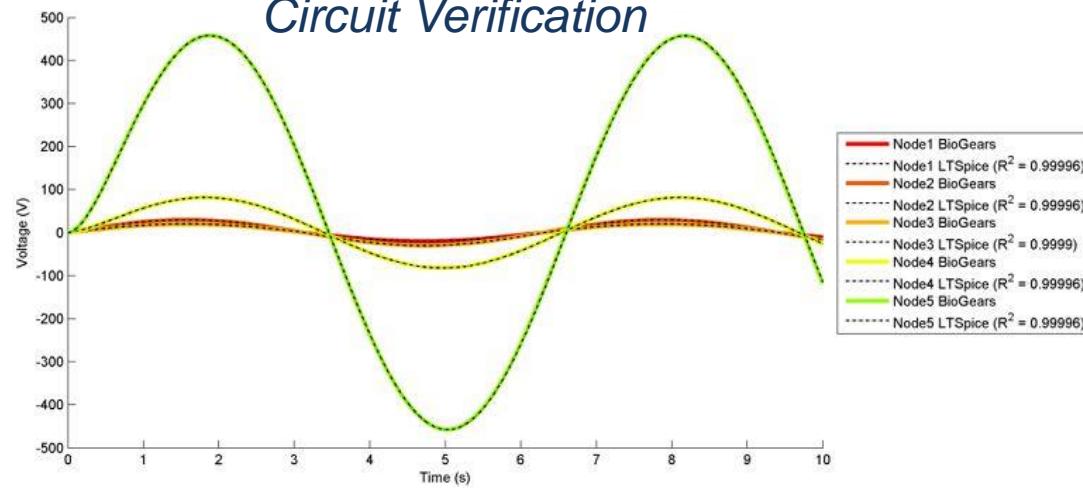
1. **Full Scenario Suite:** BioGears test suite includes scenarios to test all patient files, patient actions, substance effects, and equipment performance. Completed for 3 circuit unit tests and 80+ scenarios.
2. **Circuit Verification:** Verify that the BioGears circuit solver is producing comparable results to established circuit solvers. Completed for 100+ circuit elements/combinations.
3. **Timing:** Suite is executed by team members throughout development. It is also automatically executed when the code base is altered in the repository.
4. **Output:** Each scenario indicates the physiologic outputs for validation. Each output is tested to ensure it is within 2% of the results in the code repository. Comparison and error plots (shown) are generated.
5. **Full Results:** An email is sent to all team members when verification is complete. Each scenario either Passes (green) or Fails (red). Failures must be addressed by the team.

|   |   |   |                    |
|---|---|---|--------------------|
| VentilatorPressureLossScenarioResultsReport | 1 | 1 | 21.258999824523926 |
| YpieceDisconnectScenarioResults Report      | 1 | 1 | 23.98900008201599  |
| AirwayInsultObstructionResults Report       | 0 | 1 | 14.930999994277954 |
| AtropineScenarioResultsReport               | 0 | 1 | 13.121000051498413 |
| BasicScenario1ResultsReport                 | 0 | 1 | 18.195000171661377 |

## Scenario Verification



## Circuit Verification



## Source of Failure



# Calibration and Validation

1. **Verification:** Unit tests ensure correct implementation and sound physics principles for all tools
2. **System Level Validation:** All major systems (cardiovascular, respiratory, blood chemistry, etc.) are validated for clinical output level data
3. **Compartment Level Calibration:** Individual organs (kidney, liver, etc.) or functional units (trachea, alveoli, etc.) are validated wherever possible
4. **Scenario Calibration & Validation:** Every insult, intervention, and assessment includes a matrix with validation data for whole body combined effects from multiple systems
5. **Combined Scenario Validation:** All four showcases and several other scenarios validated for combined effects – heavily leveraged SME consultants Bryan Bergeron MD and Nicholas Moss PhD

## Showcase Scenario Combined Effects Validation

| Scenario           | Number (%) of Validation Measures in Deviation Category |            |            | Total |
|--------------------|---|------------|------------|-------|
|                    | < 10%   | 10 – 30%   | > 30%      |       |
| Combat Multitrauma | 59 (64.8%)  | 10 (11.0%) | 22 (24.2%) | 91    |
| Asthma Attack      | 26 (65.0%)  | 7 (17.5%)  | 7 (17.5%)  | 40    |
| Heat Stroke        | 52 (76.5%)  | 10 (14.7%) | 6 (8.8%)   | 68    |
| Exposure           | 26 (86.7%)  | 2 (6.7%)   | 2 (6.7%)   | 30    |

# SYSTEMS REVIEW

# Cardiovascular

## Feedback

- Heart driver – variable compliance

## Actions

- Cardiac Arrest
- CPR
- Hemorrhage
- Pericardial Effusion

## Conditions

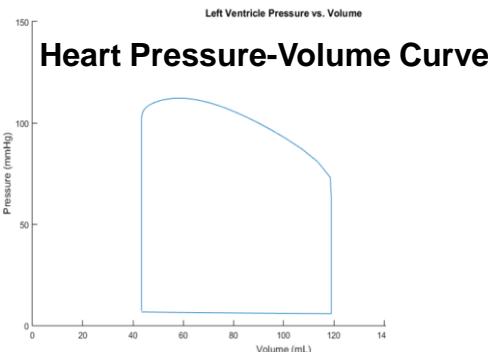
- Anemia
- Arrhythmia
- Bradycardia
- Tachycardia
- Heart Failure
- Pericardial Effusion

## Events

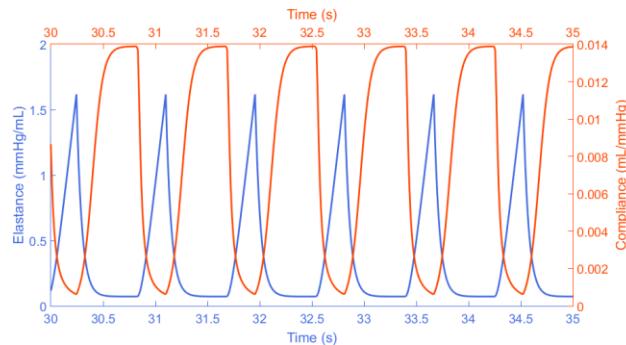
- Asystole
- Bradycardia & Tachycardia
- Cardiac Arrest
- Hypovolemic Shock

## Assessments

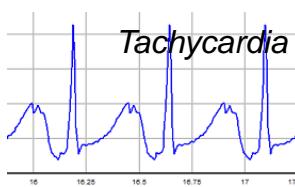
- None



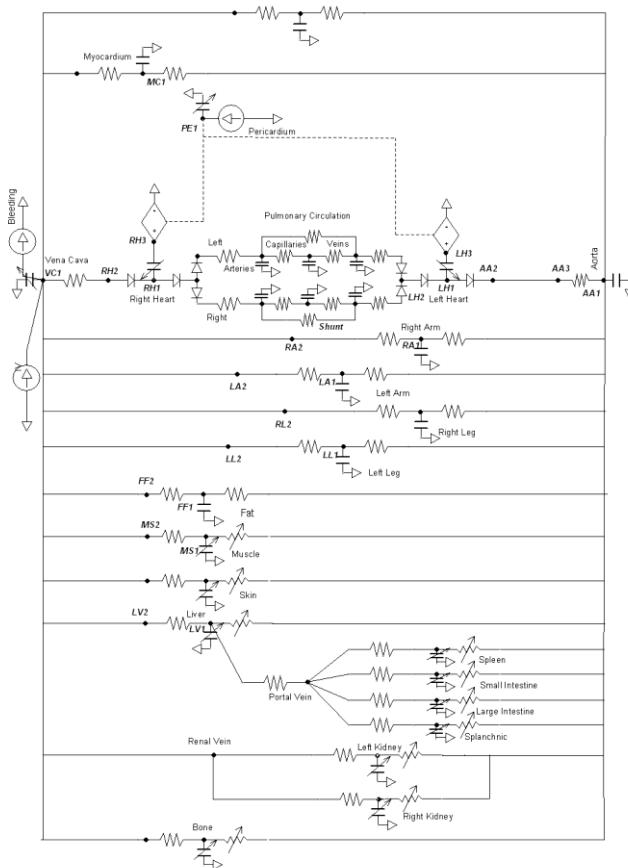
**Heart Elastance and Compliance**



**ECG Outputs**



**Circuit Diagram**



**Resting Validation**

| Data Type   | Number (%) of Validation Measures in Deviation Category |           |          | Total |
|-------------|---|-----------|----------|-------|
|             | < 10%   | 10 – 30%  | > 30%    |       |
| System      | 18 (75.0%)  | 4 (16.7%) | 2 (8.3%) | 24    |
| Compartment | 54 (84.4%)  | 4 (6.3%)  | 6 (9.4%) | 64    |

# Respiratory

## Feedback

- Respiratory driver – from aorta O<sub>2</sub> and CO<sub>2</sub>
- Pleural compliance – from lung volume

## Actions

- Airway Obstruction
- Bronchoconstriction
- Asthma Attack
- COPD Bronchitis
- Intubation
- Pneumothorax
- Conscious Respiration
- Occlusive Dressing
- Needle Decompression

## Conditions

- COPD (Bronchitis and Emphysema)
- Lobar Pneumonia
- Respiratory Acidosis & Alkalosis
- Max Pulmonary Ventilation Rate
- Right Mainstem Intubation

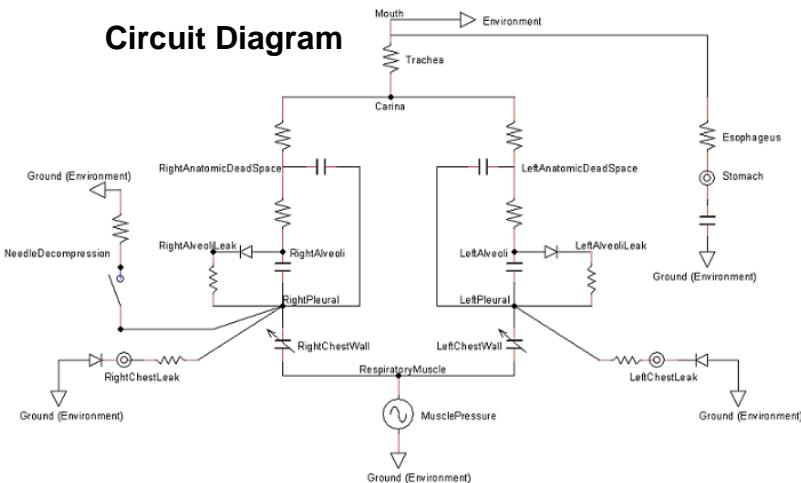
## Events

- Bradypnea & Tachypnea

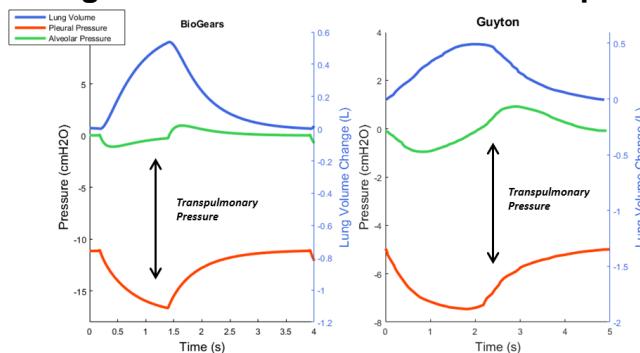
## Assessments

- Pulmonary Function Test

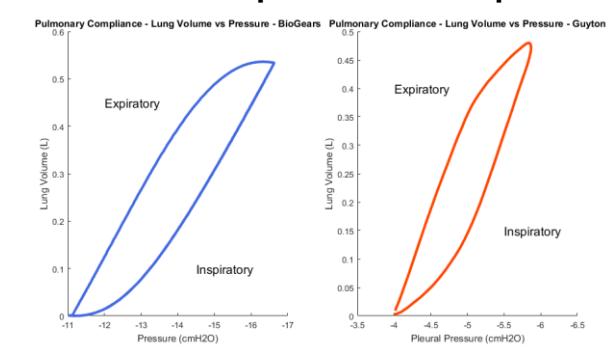
## Circuit Diagram



## Lung Volume and Pressures vs Accepted



## Variable Compliance vs Accepted



## Resting Validation

| Data Type   | Number (%) of Validation Measures in Deviation Category |          |           | Total |
|-------------|---|----------|-----------|-------|
|             | < 10%   | 10 – 30% | > 30%     |       |
| System      | 10 (83.3%)  | 0 (0.0%) | 2 (16.7%) | 12    |
| Compartment | 31 (86.1%)  | 2 (5.6%) | 3 (8.3%)  | 36    |

# Renal

## Feedback

- Ultrafiltration – from substance molecular weight/radius and charge
- Colloid osmotic pressure – from local Albumin concentration
- Tubuloglomerular – from tubules sodium delivery
- Osmoreceptor – from blood osmolarity
- Active and Passive Reabsorption – from ratio to fluid flow
- Gluconeogenesis
- Drug clearance

## Actions

- Urinate

## Conditions

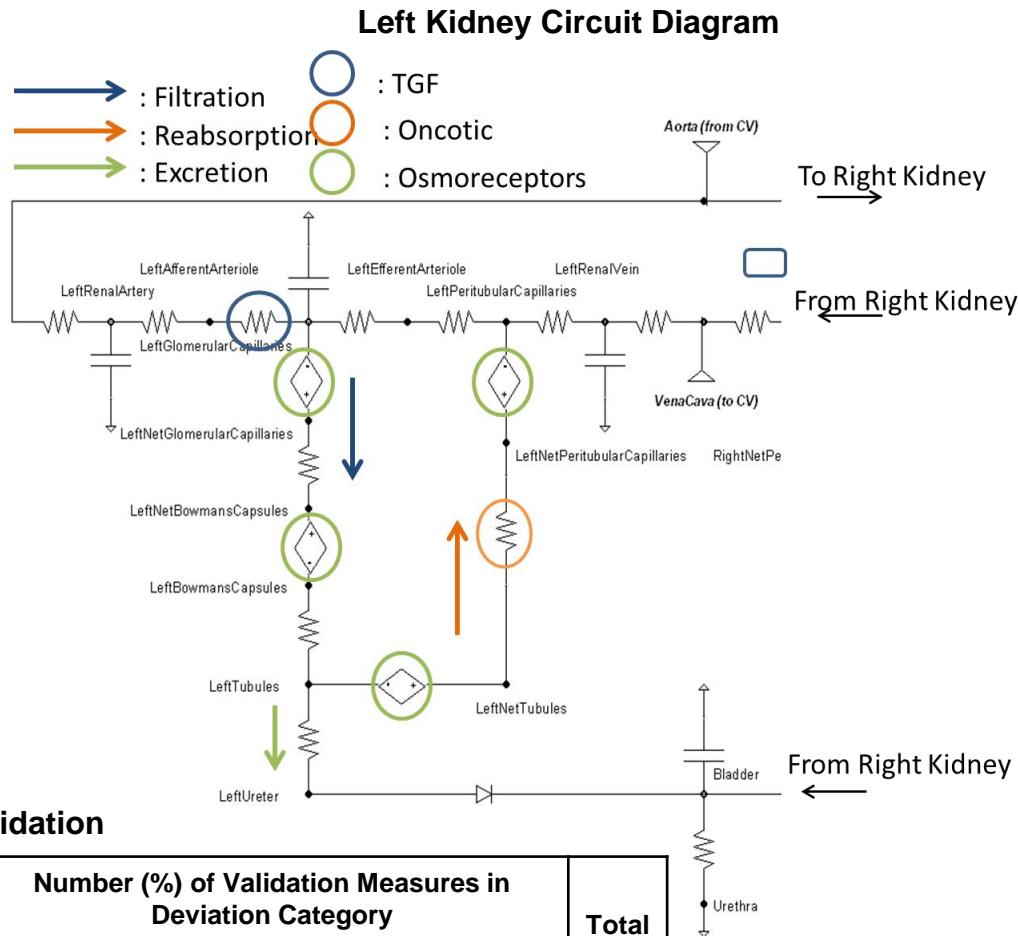
- Renal Stenosis

## Events

- Diuresis & Antidiuresis
- Natriuresis
- Dehydration
- Functional Incontinence

## Assessments

- Urinalysis



## Resting Validation

| Data Type        | Number (%) of Validation Measures in Deviation Category |            |            | Total |
|------------------|---|------------|------------|-------|
|                  | < 10%   | 10 – 30%   | > 30%      |       |
| System           | 28 (56.0%)  | 7 (14.0%)  | 15 (30.0%) | 50    |
| Compartment      | 25 (55.6%)  | 12 (26.7%) | 8 (17.8%)  | 45    |
| Substance Params | 15 (50.0%)  | 6 (20.0%)  | 9 (30.0%)  | 30    |
| Assessment       | 6 (100.0%)  | 0 (0.0%)   | 0 (0.0%)   | 6     |

# Environment

## Settings

- Air Density
- Air Velocity
- Ambient Temperature
- Atmospheric Pressure
- Clothing Resistance
- Emissivity
- Mean Radiant Temperature
- Relative Humidity
- Respiration Ambient Temperature
- Ambient Substance

## Actions

- Environment Change
- Thermal Application
  - Active Heating
  - Active Cooling
  - Applied Temperature

## Conditions

- Environment Change

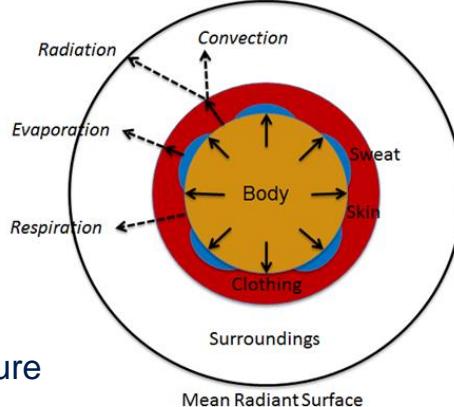
## Events

- None

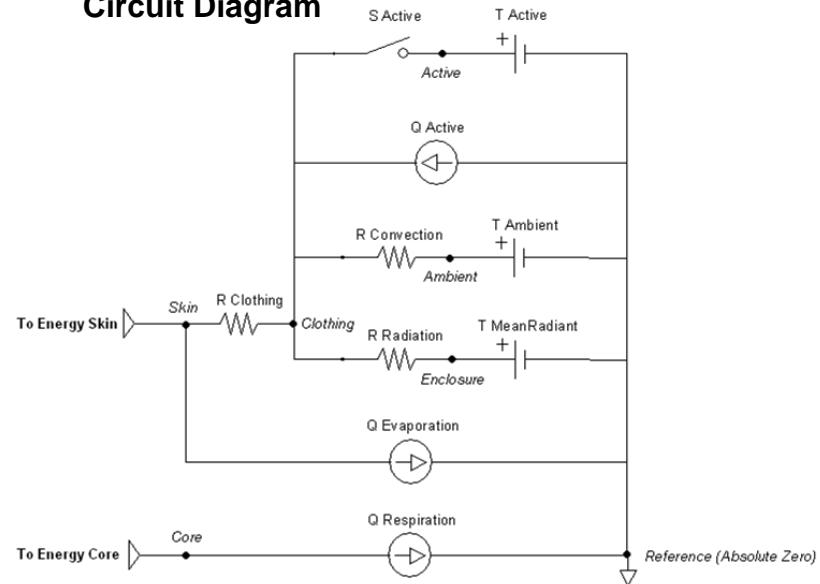
## Assessments

- None

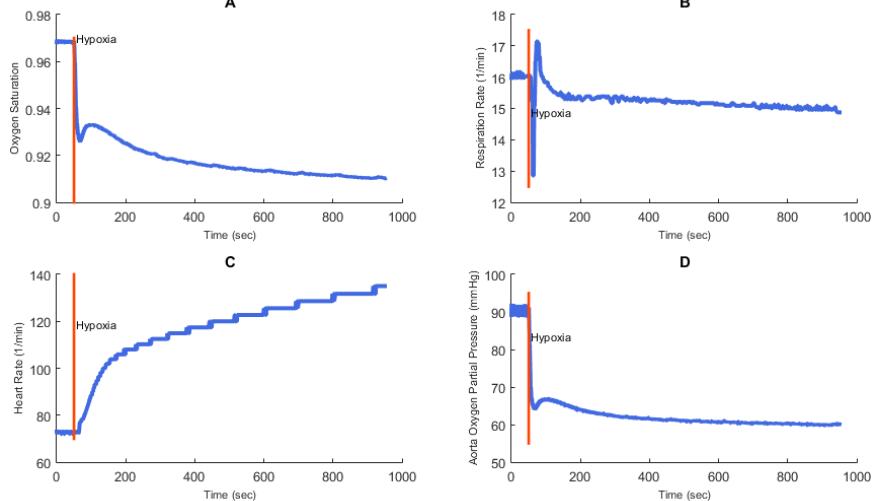
## Thermoregulation Model



## Circuit Diagram



## High Altitude Change Scenario Results



# Energy

## Feedback

- Thermal feedback on vascular resistance
- Sweating response due to increase core temperature
- Shivering response due to decreased core temperature
- Metabolic effects on Cardiovascular and Respiratory systems during physical activity

## Actions

- Exercise

## Conditions

- Dehydration
- Starvation

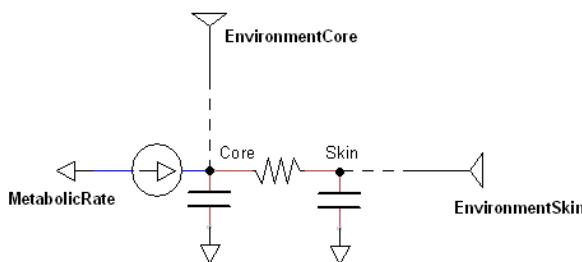
## Events

- Fasciculation
- Fatigue
- Hyperthermia
- Heat Stroke
- Metabolic/Respiratory Acidosis & Alkalosis

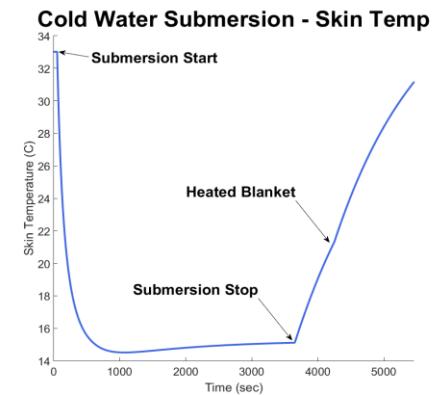
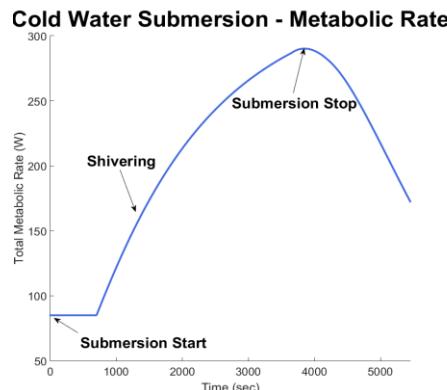
## Assessments

- None

Circuit Diagram

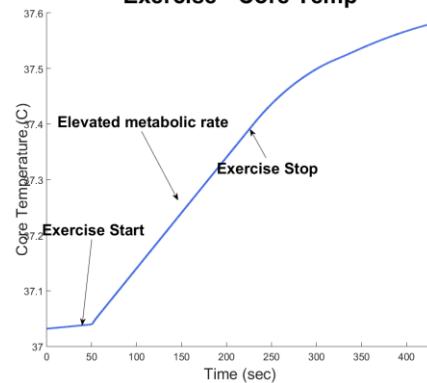


Cold Water Submersion Scenario Results

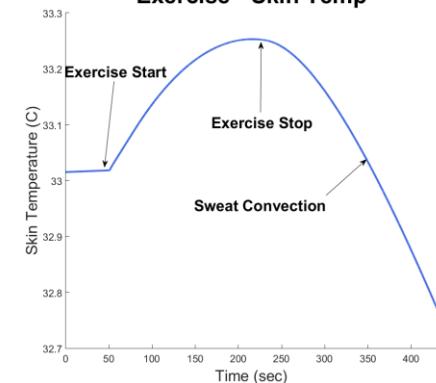


Exercise Submersion Scenario Results

Exercise - Core Temp



Exercise - Skin Temp



Resting Validation

| Data Type | Number (%) of Validation Measures in Deviation Category |          |          | Total |
|-----------|---|----------|----------|-------|
|           | < 10%   | 10 – 30% | > 30%    |       |
| System    | 9 (100.0%)  | 0 (0.0%) | 0 (0.0%) | 9     |

# Drugs

## Feedback

- PK
  - Partition Coefficients
  - Clearance
- PD
  - Drug Effects

## Actions

- IV Fluid Administration
- IV Drug Administration
- IM Drug Administration

## Conditions

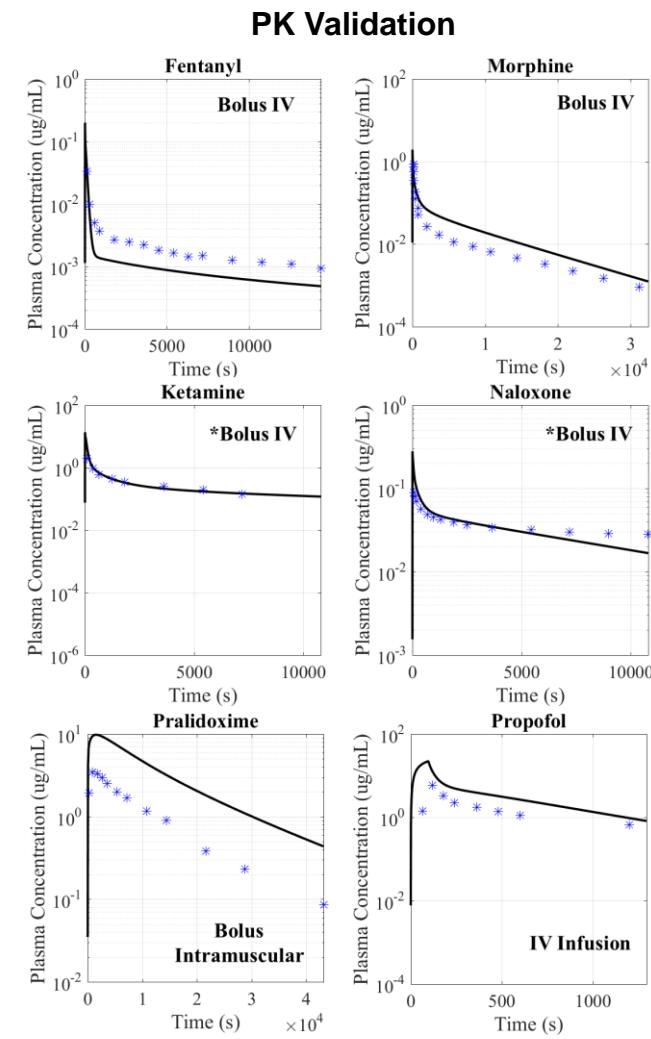
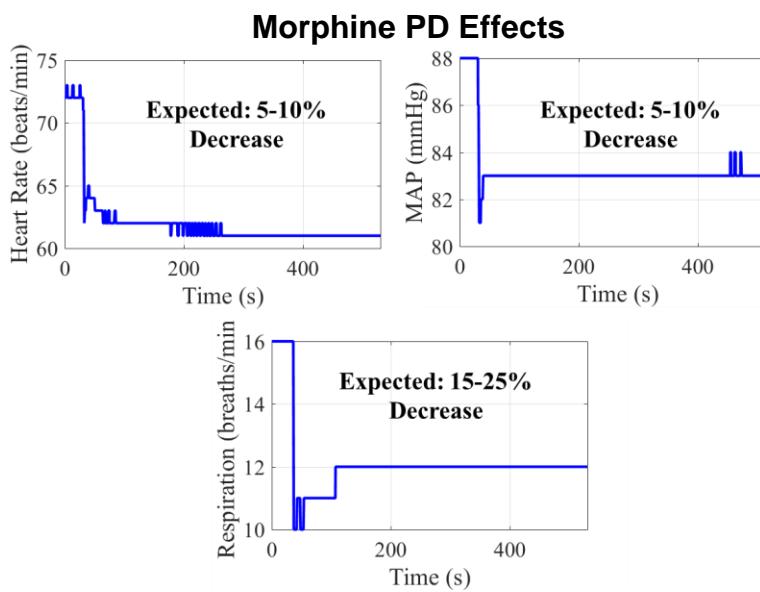
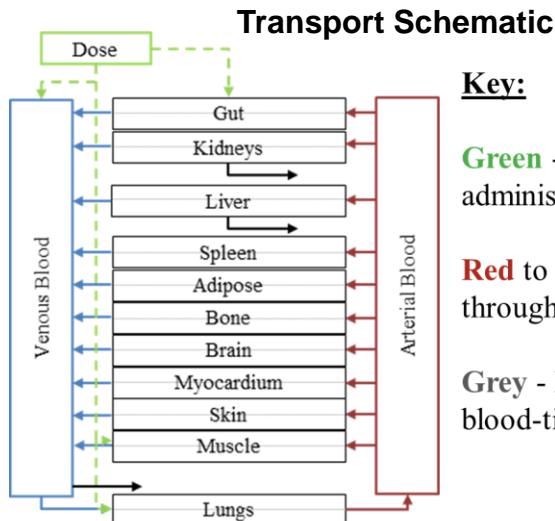
- None

## Events

- None

## Assessments

- None



## Settings

- General
  - State (on/off)
  - Ventilator Pressure
  - Respiratory Rate
  - Positive End Expired Pressure
  - Inspiratory Expiratory Ratio
  - Relief Valve Pressure
  - Ventilator Pressure
  - Oxygen Bottle One Volume
  - Oxygen Bottle Two Volume
- Gas Inlet Settings
  - Inlet Flow
  - Oxygen Fraction
  - Primary Gas (Nitrogen/Air)
  - Oxygen Source (Bottle 1/Bottle 2/Wall)
- Related Settings
  - Airway Mode (Free/Mask/Tube)
  - Substance Administration

## Actions

- Expiratory/Inspiratory Valve Leaks/Obstructions
- Soda Lime Failure
- Mask/Tube Leak
- Vaporizer Failure
- Ventilator Pressure Loss
- Oxygen Port/Tank Pressure Loss
- Endotracheal Intubation
- Esophageal Intubation

## Conditions

- Environment Change

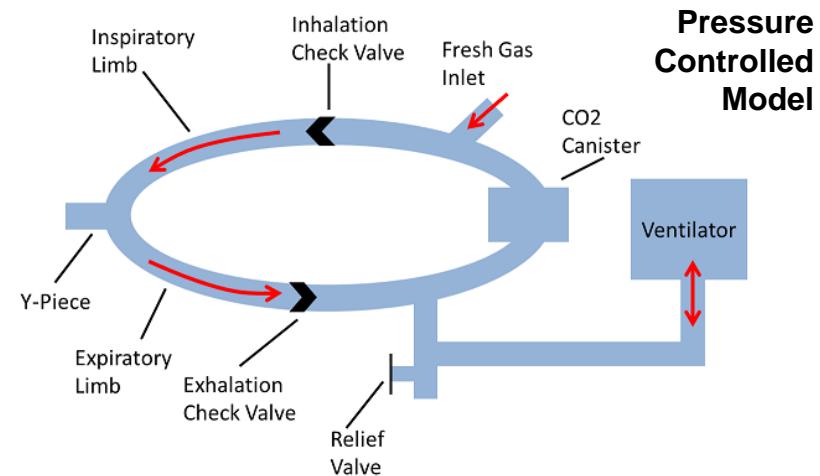
## Events

- Oxygen Bottle Exhausted
- Relief Valve Active

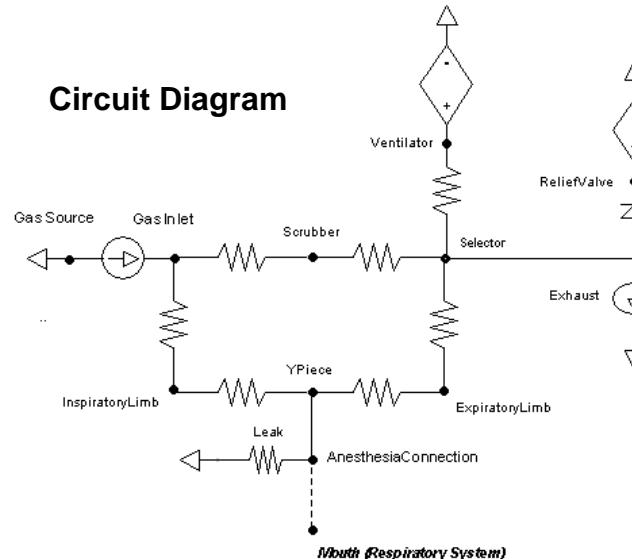
## Assessments

- None

# Anesthesia Machine



**Circuit Diagram**



# Inhaler

## Settings

- Substance
- Metered Dose
- Nozzle Loss
- Spacer Volume

## Actions

- Actuation

## Conditions

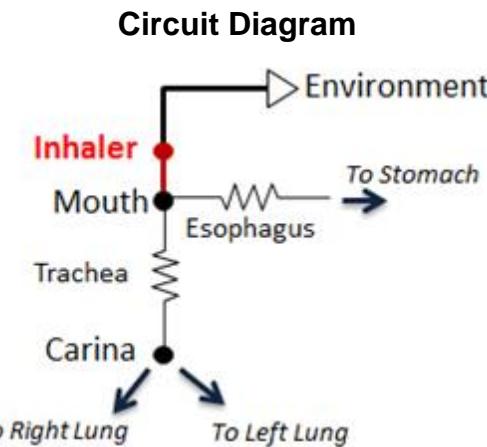
- None

## Events

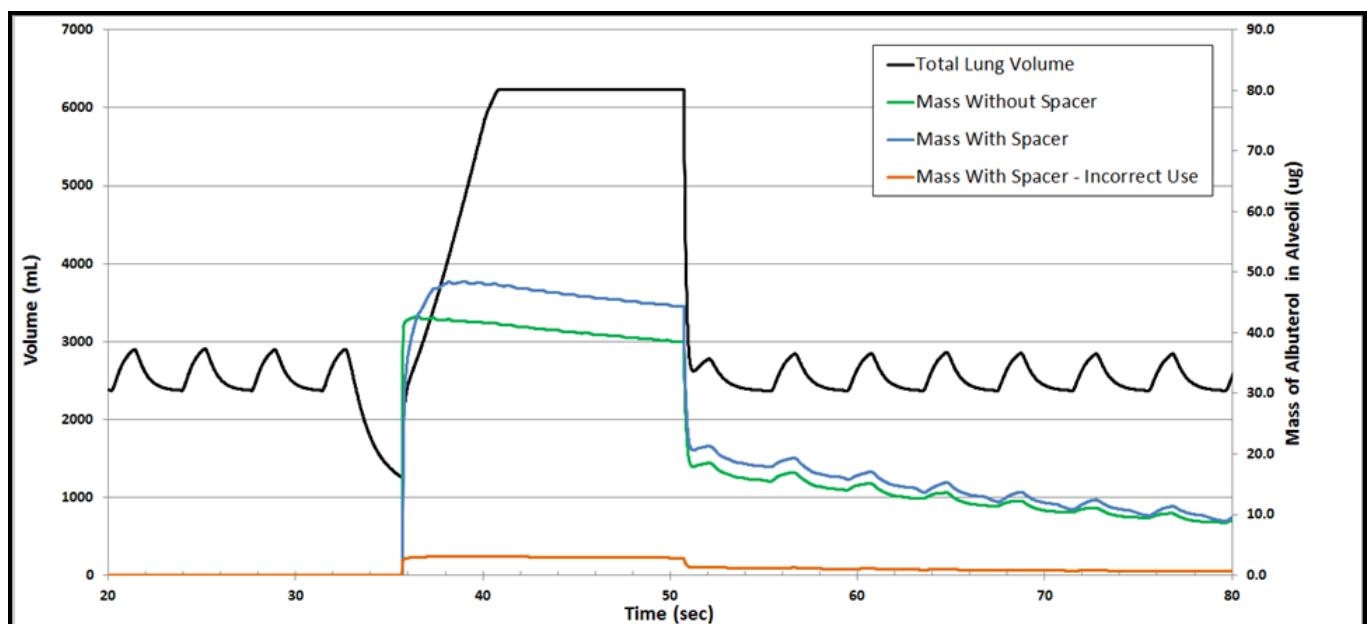
- None

## Assessments

- None



Albuterol Administration Results



# Remaining Systems

## Feedback

- Baroreceptors [Nervous]
- Epinephrine and Norepinephrine release [Endocrine]
- Produce Albumin [System Interactions / Hepatic]
- Acid-Base balance (including O<sub>2</sub> & CO<sub>2</sub> saturation & pH) [System Interactions]
- Gas exchange (Alveolar transfer) [System Interactions]

## Actions

- Consume Meal [Gastrointestinal]

## Conditions

- None

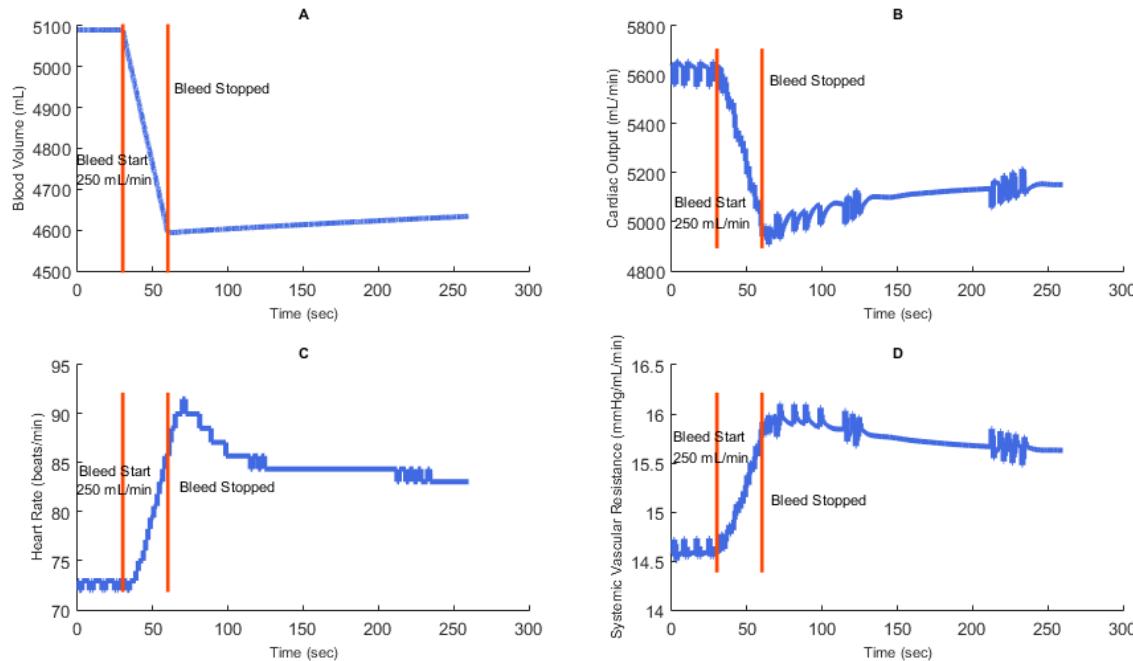
## Events

- Irreversible state
- Hypercapnia & Hypoxia [Blood Chemistry]
- Brain & Myocardium Oxygen Deficit [Blood Chemistry]

## Assessments

- Complete Blood Count [Blood Chemistry]
- Comprehensive Metabolic Panel [Blood Chemistry]

## Hemorrhage Scenario Results



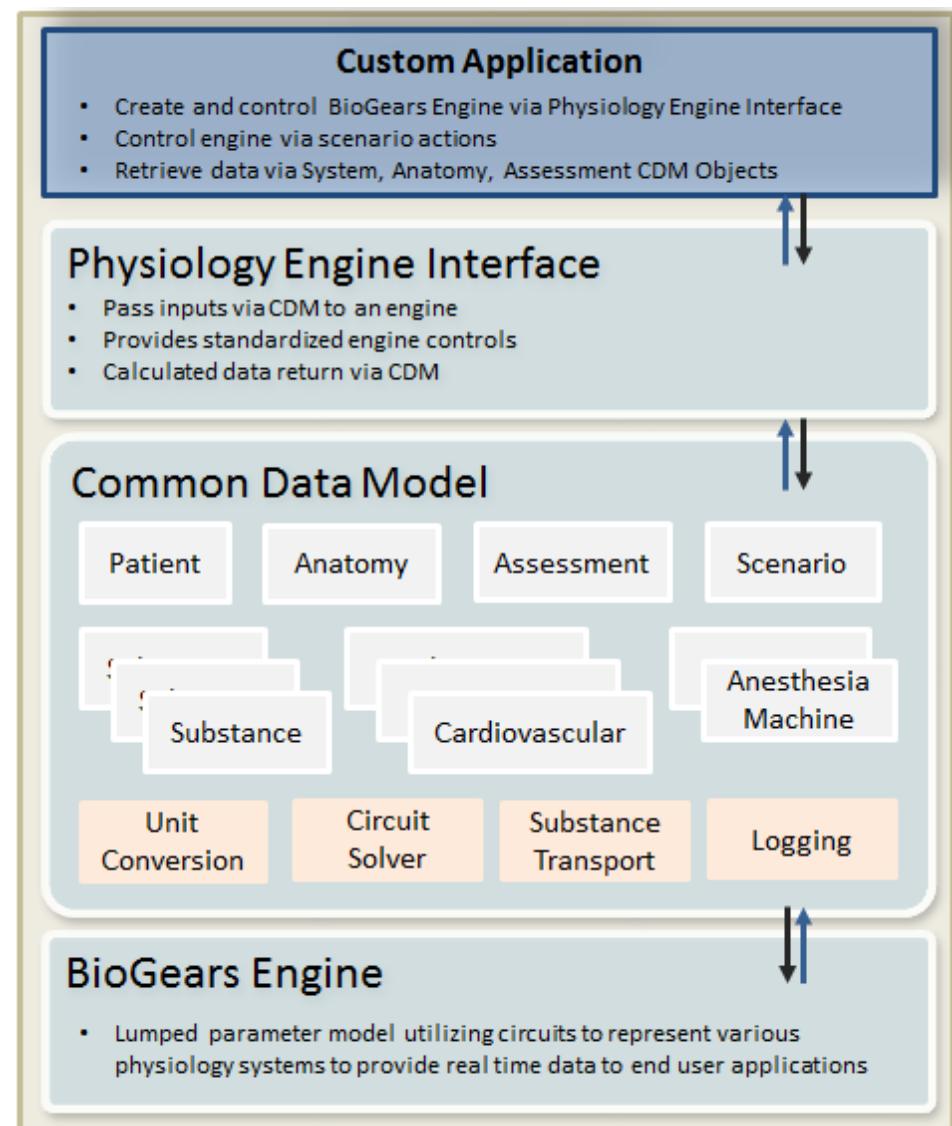
## Resting Validation

| Data Type   | Number (%) of Validation Measures in Deviation Category |          |           | Total |
|-------------|---|----------|-----------|-------|
|             | < 10%   | 10 – 30% | > 30%     |       |
| System      | 37 (84.1%)  | 2 (4.5%) | 5 (11.4%) | 44    |
| Patients    | 85 (100.0%)   | 0 (0.0%) | 0 (0.0%)  | 85    |
| Assessments | 16 (100.0%)   | 0 (0.0%) | 0 (0.0%)  | 16    |

# ARCHITECTURE AND TOOLS

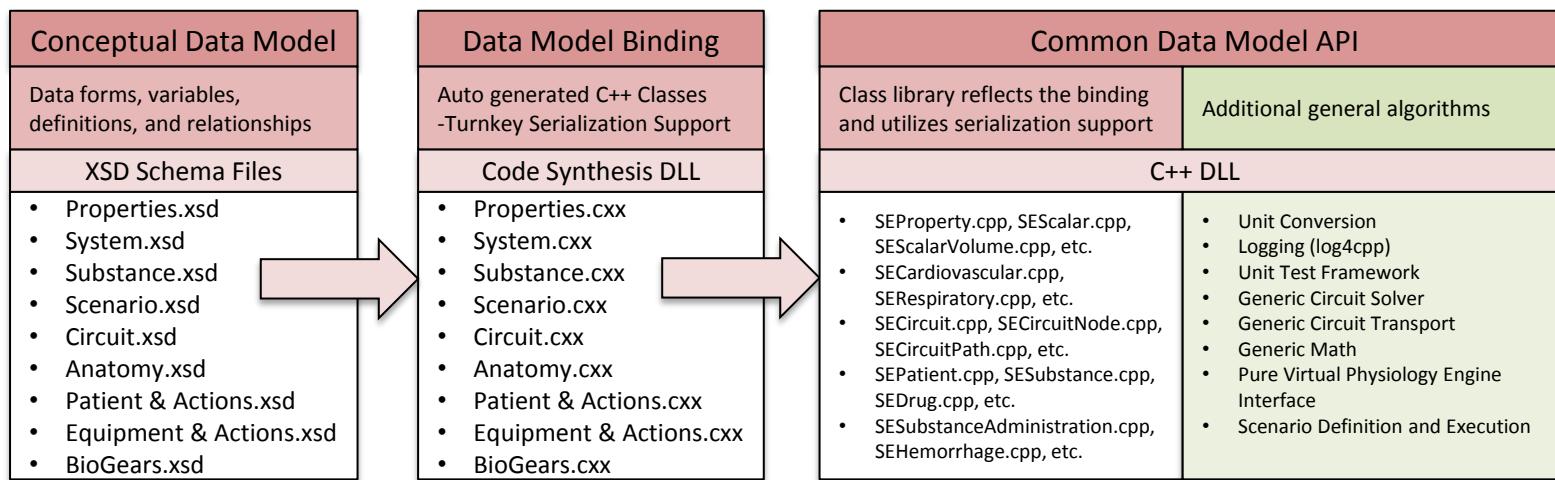
# Software Architecture

- **Common Data Model (CDM):** Well-defined, intuitive, interchangeable format to standardize interfaces
  - Standardized inputs, outputs, units, and naming conventions to aid model additions and external model integrators
- **Application Programming Interface (API):** Easy integration and interaction in any programming language
  - Data organized logically by Anatomy so that users are able to easily find and pull relevant data
- **Software Development Kit (SDK):** Application examples and stand-alone execution
  - Tutorials, How-to's, scenario examples



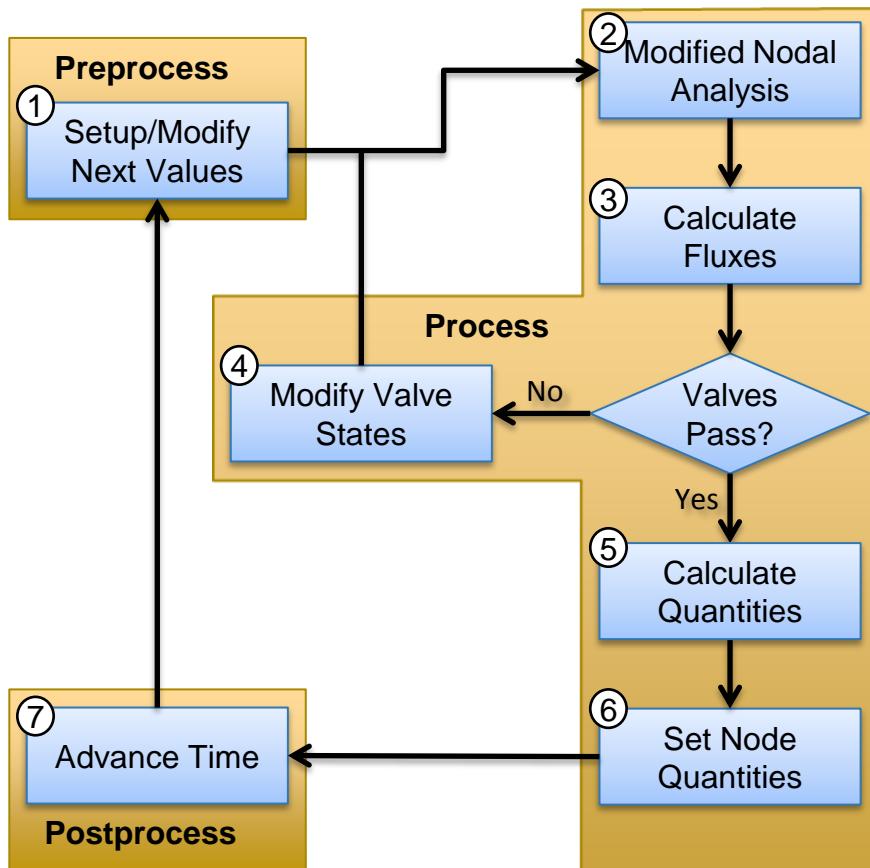
# Common Data Model Overview

- Common data structures for modeling and simulation of the human body
  - Not specific to any methodology, including BioGears
- Separates the physiological data from the physiological modeling methodology
- Object Oriented Design of class structures providing a unified set of tools that promotes fast development, compatible data sets, and well-defined interfaces
- Provides a well-defined data interchange format that disparate models can use for standardizing inputs and outputs between each other
  - Allows for specific extensions, but interfaces are defined by the CDM



# Circuit Solver

- Fully dynamic Modified Nodal Analysis solver for any valid closed-loop circuit
- Solves circuit types with any units: Electrical, Fluid, Thermal



## Preprocess:

1. Systems use “Current” values to setup/modify “Next” values via feedback mechanisms (outside of the solver)

## Process:

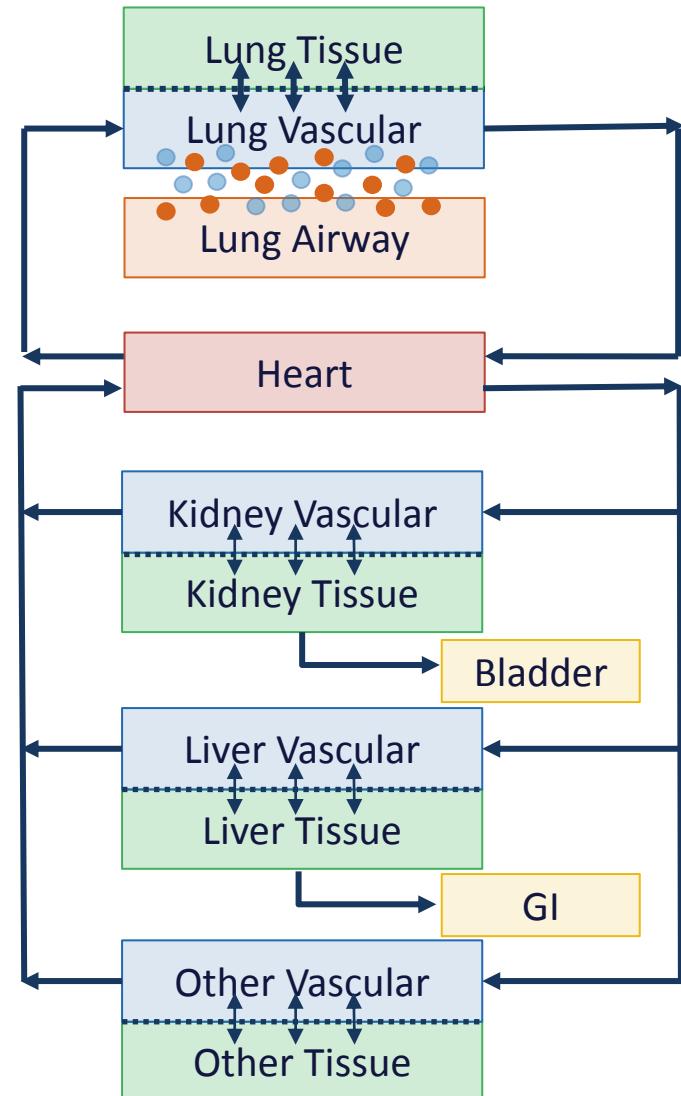
2. Perform numerical integration by using linearization (first order approximations) through Modified Nodal Analysis ( $Ax=b$ )
  - a. Use KCL (total Flux at each node = 0) to Calculate the Jacobian matrix ( $A$ ) and right-hand side vector ( $b$ ) for each Node Potential and Potential Source Flux ( $x$ )
  - b. Use the Eigen templated library linear solver (FullPivLU) to solve for  $x$  vector
3. Calculate unknown Fluxes – using Trapezoid Rule where applicable
4. Calculate Valves using assumed diode states (cannot be solved directly) – iterate as necessary
5. Calculate and increment Compliance Path Quantities – No other elements have dynamic Quantities (rigid pipes)
6. Set Node Quantities (based on Path Quantities)
  - Note: Transporter is called here

## Postprocess:

7. Advance time by moving “Next” to “Current” values

# Transporter

- Common Data Model:
  - Substances move with the fluid to each node in the circuit
  - No particle deposition
  - Mass is updated as at each time step based on flow into and out of nodes
  - Concentration and partial pressure are updated after the mass
- Systems Interactions:
  - Partial pressure driven diffusion moves substances between two nodes ( $O_2$ ,  $CO_2$ ,  $N_2$ )
  - Perfusion limited diffusion moves substances across the blood/tissue barrier based on flow and partition coefficients (Drugs)
  - Systemic clearance removes substances from the vena cava to represent metabolic or other clearance mechanisms
  - Hepatic clearance removes substances from the liver to represent the ability of the liver to metabolize or remove substances



# Developer GUI

## Motivation

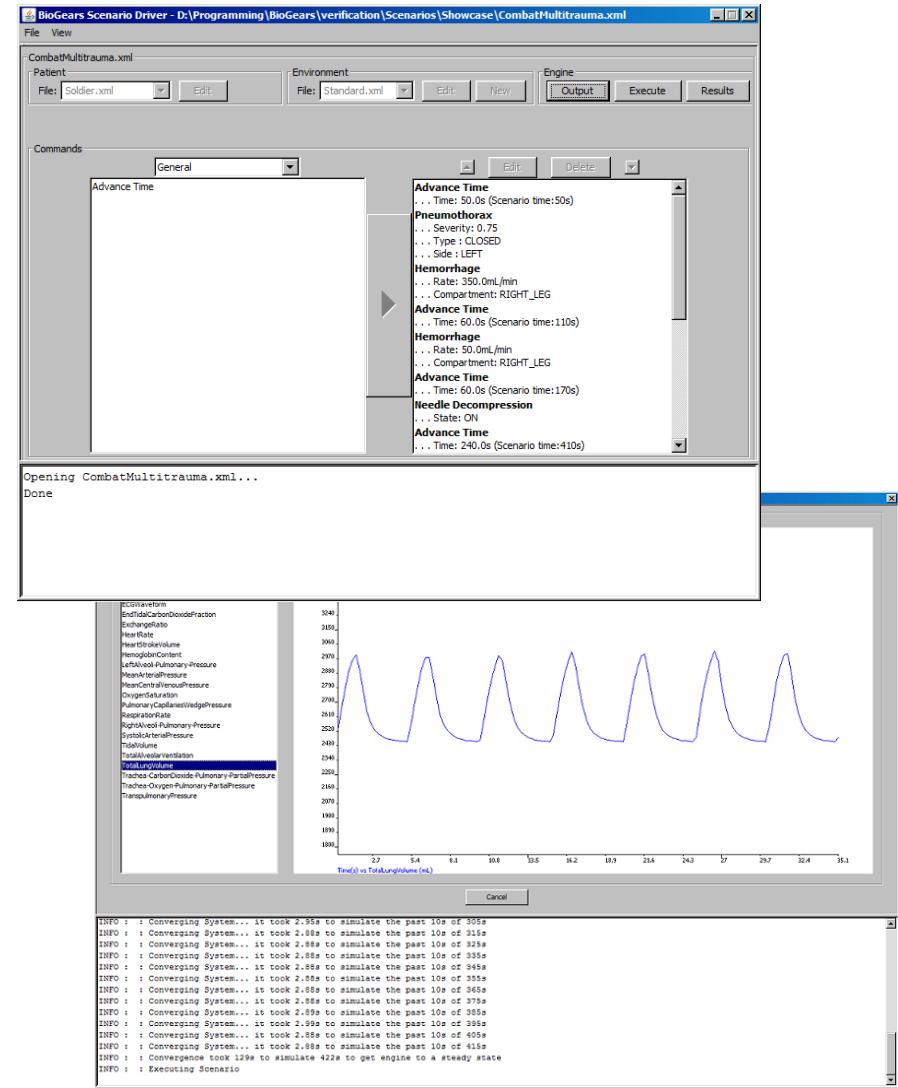
- Data driven developers' tool to demonstrate basic functionality
- Test Engine without command line
- Ready for testing out-of-the-box (no compiling)
- Simple interface for creating, editing, and executing scenario file, and creating resulting plots of requested data

## Expectations

- Only requirements are to allow editing and execution of scenarios – not a major focus
- Not heavily QA'ed – some known bugs being continuously fixed

## Limitations

- Clunky Java Swing for rapid prototyping via the API



# COMING SOON

# High Level Near Term Tasks

## Current (hopefully deployed this summer with version 6.0.0):

- Bug fixing and system refinement
- Optimization and increased simulation speed
- State serialization – saving and loading simulations
- Modularity – more easily replace entire systems
- Renal feedback updates
- Acid-base balance – O<sub>2</sub> & CO<sub>2</sub> saturation modifications
- Total body substance balance and new substances

## Near-Term (FY16):

- Nervous system additions
- Exocrine additions
- Endocrine additions
- Vascular fluid exchange
- Pneumothorax updates
- Gastrointestinal updates

## Long-Term (FY17):

- Patient modifications (gender, body mass, etc)
- Intoxications – Ketamine proof of concept
- Airborne agents (Nerve/Pulmonary/Smoke/CO) and vaporization
- Diuretics
- Additions to blood assessments and pulmonary function test improvement

## How to Contribute

- Use the software for any and all applications (please let us know)
- Report problems
- Submit code
  - Currently just email us  
(<https://www.biogearsengine.com/workwithus>)
  - Moving to a public repository – GitHub/BitBucket hosted
- Post and respond to Forums  
(<https://www.biogearsengine.com/forums>)



# Contact

[www.biogearsengine.com/workwithus](http://www.biogearsengine.com/workwithus)

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