Computational Challenges in Cardiovascular Fluid Mechanics

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History

Thomas Young 1773 – 1829

Both a practicing physician and professor of physics

Other early students of blood circulation: Borelli, Hales, Bernoulli, Euler, Poiseuille, Helmholtz, Fick, ...

As science developed, so did specialization and the study of the cardiovascular system became separated from physical science.\*

\*CG Caro, TJ Pedley, RC Shroter, WA Seed; The Mechanics of Circulation; Oxford University, 1978

## The Cardiovascular System

Three Major Elements – Heart, Blood Vessels, & Blood

The Heart - cardiac muscle tissue

Pumps blood throughout the body

■Four chambers

Right atriumRight ventricleLeft atrium

■Left ventricle



### Circulation

<u>Pulmonary circulation</u>: path of blood from right ventricle through lungs and back to heart

Systemic circulation: path of blood from left ventricle to body and back to heart

<u>Rate</u> of flow through systemic circulation = flow rate through pulmonary circulation



### **Cardiac Cycle**



## The Fluid: BLOOD

#### Whole blood $\sim 7\%$ of human body weight:

#### ~45% Red blood cells ~2% Platelets ~1% White blood cells ~52% Plasma





## **Cardiovascular Flows**

- Pulsatile (periodic) flows: heart rate dependent Womersley #
- Internal (confined; undeveloped) flows
- Moving visco-elastic boundaries (chamber /vessel walls; heart valve leaflets)
- Non- Newtonian (two phase with deformable elements)
- Multi-scale (100 mm to 1 micron)

## **Cardiovascular Flows**

#### Multi-scale

- Laminar flows: Peak Re~6000 (non-diseased resting states – peak vel ~1.2 m/s; dia ~25mm)
- Transition to low Re turbulent (peak ~ 30,000) flows (exercise and/or diseased states - vel ~5m/s)
- Very low Re flows in capillaries: ~ 0.001 (two phase with deformable elements; vel ~1mm/s; dia 10 micron)

## 2D PCMR – Healthy Human LV



## **Identifying Features of BAV**



Trileaflet

Bileaflet

#### Courtesy of Northwestern Radiology

## Heart Valves

#### Aortic valve

Lies between left ventricle and aorta

#### Pulmonic valve

 Lies between the right ventricle and pulmonary artery

#### Mitral valve

Lies between left atrium and ventricle

#### Tricuspid valve

Lies between right atrium and ventricle



## Heart Valves

- Heart valves maintain unidirectional blood flow in the heart
  - Open and close passively due to pressure differential
- Native heart valves may malfunction due to congenital birth defects or disease
  - Valve stenosis (narrowing of valve area) or regurgitation (unwanted backflow through closed valve)



Aortic Valve



C Healthwise, Incorporated

## Heart Valve Disease

- Heart disease is leading cause of death in developed nations
- In US, 2.5% of population has heart valve diseases
- In developing countries, 2.3% of children have Rheumatic Heart Disease
- 1% of population are born w/bicuspid aortic valves
   early calcification of valves

## Six Decades of Progress, but....



## Serious Problems and Complications Associated with Prosthetic Heart Valves

Mechanical Valves

- Thrombosis and thromboembolism
- Anticoagulation related hemorrhage
- Red cell damage and hemolysis
- White cell damage
- Tissue overgrowth
- Damage to endothelial lining of vessel walls in the vicinity of the prosthesis
- Leaks caused by failure of valve to close properly
- Infection
- Tearing of sewing sutures

Bioprosthetic Valves

- Calcification of Valve
- Adverse immune reaction
- Structural failure / Wear and tear
- Perivalvular and paravalvular Leakage
- Tisue overgrowth
- Infection
- Tearing of sewing sutures

## **The Bileaflet Mechanical Valve**

Trend of implantation: 55% bioprosthetic valves vs 45% mechanical valves
The bileaflet mechanical valve design is the valve of choice (>90% of implantation)



## **BMHV** Complications

- Despite evolution and advantages of BMHVs, complications are known to exist
- Major complications are platelet activation and thromboembolic events
  - Clot formation and detachment
- Clot formation in BMHVs have serious consequences
  - Experiments have shown restriction of normal motion of leaflets due to thrombus<sup>1</sup>
  - Detached emboli may lead to vessel occlusion resulting in stroke or death



<sup>1</sup> Baumgartner et al., Circulation, 1993.

## Flow Characteristics through BMHVs



Large scale features

- Pulsatile, turbulent flow
- Complicated geometries and flow fields
- Moving objects / Fluid structure interactions



Small scale features

- Flow through narrow gaps
- Complex 3D vortical structures and regurgitation jets

\* Courtesy: Medtronic Inc.



## **Single-Phase Flow Validation**

- Model single-phase fluid flow through 23mm St. Jude Medical (SJM) valve
  - In aortic position
- Results compared to experimental Particle Image Velocimetry (PIV) data of flow through an *in vitro* flow loop<sup>1,2</sup>
  - Used blood analog fluid, same kinematic viscosity as whole human blood (3.5e<sup>-6</sup> m<sup>2</sup>/s)
- Compare axial velocity for steady and pulsatile flow valve (mean flow)
  - Re = 750 for laminar regime and Re = 5000 for turbulent regime for steady flow
  - Compare mean flow fields and RMS values for pulsatile flow
- Compare instantaneous vorticity contours for pulsatile flow through valve



Computational model of experimental in vitro flow loop setup

<sup>1</sup>Ge et al., Journal of Biomechanical Engineering, **2005**. <sup>2</sup>Dasi et al., Physics of Fluids, **2007**.

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### **SJM Valve Computational Geometry**





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### SJM Valve Computational Geometry



Transparent view of fully open leaflets

Interior view showing fully open leaflets fitting into butterfly hinge geometries Wallace H. Coulter School of Biomedical Engineering Georgia Institute of Technology College of Engineering and Emory University School of Medicine

### **Comparison with Experimental Data: Leaflet Opening**



### **Comparison with Experimental Data: Acceleration**



#### **Comparison with Experimental Data: Peak Flow**



### **Comparison with Experimental Data: Deceleration**



Demonstration of fine, small-scale flow structures

Leaflet wake vortices and sinus recirculation region



Accelerating Flow

Re = 3500



Peak Flow

Re = 5800



Leaflet Closing



### **3D Visualization - Vortical Structures**





### **Blood Damage Quantification**

- Quantifying damage to suspended platelets
- Use of a Blood Damage Index (BDI)
- Linear shear stress exposure time damage accumulation model
- Previously validated against whole human blood damage experiments<sup>1,2</sup>
  - Validated LBM-EBF numerical method for quantifying platelet damage
  - Selected best BDI model in combination with numerical method
- Can track damage of individual platelets or average across platelets

$$BDI = \sum_{t=0}^{t=end} \tau_{\max} \cdot \Delta t$$

 <sup>1</sup> Fallon, A.M. *et al.* "Procoagulant properties..." *Annals of Biomedical Engineering*, 2008.
 <sup>2</sup> J. Wu, B. Min Yun, *et al.*, "Numerical investigation of the effects of channel..." *Annals of Biomedical Engineering*, 2011. Wallace H. Coulter School of Biomedical Engineering *Georgia Institute of Technology College of Engineering and Emory University School of Medicine*

### **Blood Damage Quantification**





B. Min Yun et al., "A numerical investigation of blood damage in the hinge area ...," ABME, 2012.

# Small-Scale Computational Studies



Simon (2010)

Simulation of 3-dimensional hinge flow fields

Pulsatile flow under aortic conditions

# Small-Scale Computational Studies



#### Interfacing Experimental and Computational Approaches to Clinical Translation


#### Modeling in Cardiovascular Devices

#### High level goals:

- Industry for design performance evaluation, regulatory evaluations
- Academia/Research for study of fluid and solid mechanics of devices and blood

#### On a finer scale, what matters?

- Device Performance
  - Durability
  - Fatigue
  - Risk evaluation

#### Hydrodynamics

- Shear stress
- Stasis
- Thromboembolic events
- Solid mechanics
  - Strain
  - Loading profiles



#### **Patient Specific Surgical Planning Models**



#### Patient Specific Surgical Planning Models

#### **Predicting Postoperative Function**



#### Predicting Postoperative Tissue Stress



#### Accuracy is Largely Dependent On

*How close these models mimic native mitral valve geometry, function, and mechanics.* 

#### Numerical Simulations of Native MV

However, without rigorous validation against benchmark data, computational models are of questionable value in their clinical utility

"…three-dimensional in-vitro flow, deformation and strain data with a native mitral valve are sorely needed." – Einstein et. al, 2010

#### Novel Modular Left Heart Simulator



Modular chamber – removable components for various modalities, while maintaining same valve geometry and configuration

## **Characterization Modalities**

MV Geometry	MV Fluid Mechanics	MV Tissue Mechanics					
Micro CT	Hemodynamics	Leaflet Strain					
3D Echo	Stereoscopic PIV	Papillary Muscle Forces					

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Realistic Valvular Anatomy Trough State of the Art Imaging
Micro-CT imaging provides realistic valvular anatomy, including subvalvular apparatus, up to 21 µm resolution, in open and closed configuration



# Valvular Anatomy From Clinical 3D Ultrasound









Schematic of segmentation planes across the mitral valve

RT3DE image of MV with a segmentation plane shown in red

Segmentation plane

with anterior (yellow)

and posterior (green)

leaflets segmented with the user selected

points

Normal Valve



Anterior Leaflet Bellowing



Anterior Leaflet Flail



## **Characterization Modalities**

MV Geometry	MV Fluid Mechanics	MV Tissue Mechanics
Micro CT	Hemodynamics	Leaflet Strain
3D Echo	Stereoscopic PIV	Papillary Muscle Forces

## **Pulsatile Flow Simulator**



# Highly Resolved Flow through the MV



# Out-of-plane velocity & vorticity



Very low magnitude of W in the central plane
Highest vorticity values restricted to edge of diastolic filling jet as expected

## **Characterization Modalities**

MV Geometry	MV Fluid Mechanics	MV Tissue Mechanics
Micro CT 3D Echo	Hemodynamics Stereoscopic	Leaflet Strain Papillary Muscle
		TUICES

## Leaflet Strain



Marker Array 1.5 mm x 1.5 mm Markers ~0.5 mm in diameter

3D marker location accuracy: 65 µm

(Based on camera magnification, pixel size, and the calibration)



Camera 2

Mitral Valve Mounted in the simulator



#### Leaflet Strain







## Subvalvular Force Measurements

In-house fabricated strain gauge transducers measure subvalvular forces throughout the cardiac cycle





(Top) Papillary muscle force transducers measure axial, and two directions of bending force on each papillary muscle

(Left) Miniature chordal force transducers can be applied to the strut, intermediate, and marginal chordae

# Current uses in Computational Modeling





Multi-scale computational model developed at the University of Texas





Fluid-structure interaction model developed at the University of Maine, Pacific Northwes National Labs, and Georgia Tech



Real-time medical image based model developed at Siemens Healthcare

# FSI Analysis - Normal Valve



## Validation Framework for Computational MV Models to Predict MV Closure Geometry



Georgia Institute of Technology College of Engineering and Emory University School of Medicine

# Geometric Model Extraction from microCT

Developed prototype to process microCT volume received from Georgia Tech group and extract geometric models containing:

- Anterior and Posterior leaflet meshes (red)
- Papillary Muscles (green)
- Mitral Valve Chordae Tree (yellow)



# Geometric Model Extraction from 4D Echo

 Adapted Echo prototype to process ex-vivo 4D Echo data received from Georgia Tech group and extract simplified geometric models containing:

Anterior (blue) and Posterior (green) leaflet meshes

Papillary Muscles Tips (green points)



# Bio-mechanical Modeling of MV Physiology



T. Mansi et al., Medical Image Analysis, 2012

Siemens Healthcare

# **Boundary Conditions**



# Clinical Measurements for Quantitative Evaluation

- Selected Clinical measurements relevant for short term post-mv surgery repair:
- Coaptation Area
- Coaptation Length
- As we use a parametric model, additional measurements can be easily added
- Leaflet Tenting Height
   Tenting Area, and Tenting Volume
   Annular-Height-to-Commissural-Width
   Ratio (Saddle Geometry)
   Mitral Annular Area
   Circumference
   Septal-Lateral Diameter
   Transverse Diameter





#### Siemens Healthcare

# **Quantitative evaluation**

- Evaluation of mitral valve closure geometry from TEE, microCT (µCT) and simulated closure from TEE (Sim TEE) on 3 data sets.
- Promising correlation between biomechanically-closed TEE MV and microCT ground truth
- Abstract submitted to BMES (04-23-2014)

	Coaptati	on Leng	th [mm]	Coa	ptation A [mm <sup>2</sup> ]	Area
	TEE	μCΤ	Sim TEE	TEE	μСТ	Sim TEE
Data Set 1	2.62	2.41	2.71	41.2	46.10	45.80
Data Set 2	2.00	1.92	2.05	50.07	48.23	51.42
Data Set 3	2.25	2.12	2.32	56.17	53.17	58.07

# Summary

- Computational MV models need high fidelity boundary conditions & validation:
  - High fidelity MV anatomy obtained through state of the art imaging (µCT, 3D Ultrasound, High speed stereo photogrammetry)
  - Hemodynamic boundary conditions are captured using direct flow and pressure instrumentation
  - Force measurements on MV verify CFD model and adjust material properties models
  - Complex flow fields can be captured through Stereoscopic Particle Image Velocimetry for model validation

#### **ENGINEERING APPROACH TO CLINICAL PROBLEM**

Develop methodology to quantify in vivo patient specific hemodynamic environment from clinical imaging data that achieves the following:

- accurately models physiologic environment
- technically sound (i.e., robust, validated)
- provides clinically useful information
- adaptable to numerous clinical pathologies (disease state, anatomic location)
- *reasonable* time requirements

#### **<u>Clinical Questions</u>**

 Following presentation of coronary disease, can we predict plaque evolution?
 What is the role, if any, of hemodynamics in cardiac allograft vasculopathy?





potential TCFA

Courtesy of P. Stone,, TCT 2011

Lucas H. Timmins, Ph.D., et al.

#### **OVERVIEW OF GT/EMORY COMPUTATIONAL METHODS**



#### **3D GEOMETRY RECONSTRUCTION**

Utilized combination of ANGiography and intravascular UltraSound (ANGUS)

- 3D space of IVUS catheter determined from <u>biplane</u> data
- lumen borders are segments from VH-IVUS images
- images stacked perpendicular to wire, apply global rotation
- branches extended perpendicular from main vessel

Laban et al., *Comp. in Cardiol*, 1995 Krams et al., *ATVB*, 1997



#### **COMPUTATIONAL FLUID DYNAMICS (CFD) TECHNIQUES**

Flow extensions are added

- inlet: smooth transition into domain
- outlet: fully developed flow

Geometry discretized with boundary layer to resolve near-wall flow patterns

**Boundary Conditions** 

- inlet: pulsatile (blunt) velocity waveform
- outlet: pressure-free (traction-free)
- wall: no-slip (i.e.,  $\vec{v} = 0$ ), rigid

Newtonian fluid ( $\rho = 1060 \text{ kg/m}^3$ ,  $\mu = 3.5 \text{ cP}$ )

Post-process (node) data to quantify WSS vectors - temporally average magnitudes; yield timeaveraged WSS





Lucas H. Timmins, Ph.D., et al.

#### **REPRESENTATIVE ASSOCIATION OF BASELINE WSS AND PLAQUE PROGRESSION**



Baseline Pullback

Follow-up Pullback (6 months later)



 developed methods allow for direct and quantitative comparison of hemodynamic environment and clinically measured coronary artery disease progression

Lucas H. Timmins, Ph.D., et al.

#### ASSOCIATION OF WALL SHEAR STRESS AND CHANGE IN PLAQUE COMPOSITION OVER 6 MONTHS

Examined association between wall shear stress and plaque progression in 20 patients

- Low WSS: plaque progression
- Intermediate WSS: plaque regression
- High WSS: progression to vulnerable plaque (significant increase necrotic core tissue)

Utilized computational fluid mechanics as clinical marker to identify variations in coronary artery disease progression



Samady, Eshtehardi, McDaniel, Suo, Dhawan, Timmins, Quyyumi, Giddens, *Circulation*, 2011

Lucas H. Timmins, Ph.D., et al.

# **Translational Impact**



#### **From Bench to Bassinet**

#### **Image-Based Surgical Planning**

A novel means to pre-operatively evaluate blood flow characteristics

 Allow the possibility to tailor the surgery to the patientspecific scenarios

Here, we present our experiences of surgery planning of single ventricle heart defect palliations

## **Total Cavopulmonary Connection**

- The Total Cavopulmonary Connection (TCPC) is a modification of the original Fontan surgical repair
- TCPC results in single ventricle driving flow
- Control of power loss may be critical
- Requires tool for clinical assessment of TCPC performance


# Complications

- Limited Exercise Capacity
- Pulmonary arteriovenous malformations (PAVMs)
- Progressive Cyanosis from shunting
- Thromboembolism
- Arrhythmias
- GI System Disorders
  - Protein Losing Enteropathy
  - Feeding Disorders
  - Liver Failure

## **Complex TCPC Hemodynamics**

- Relevant to some of these long-term complications
- Requires tool for clinical assessment of TCPC performance
- Parameters of interest:
  - central venous pressure
  - power loss across the connection
  - balanced flow to the lungs



### Surgical Planning of TCPC: Methodology

## Methodology Overview





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### In Vivo Flow Fields – PC MRI

- Sequence: BFFE (PC MRI)
- Direction: Coronal
- N = 5-7
- Slice Thickness: ~6 mm
- TR/TE = 42/3.1 69/4.8
- Pixel Size  $\sim = 1 \times 1 \text{ mm}^2$
- Matrix Size: 256 x 256
- Directions: 3 (AP, FH, RL)
- VENC: 80 (FH), 80 (RL), 40 (AP)



#### **3D PC-MRI Velocity Reconstruction Methodology**



# Computational Fluid Dynamics (CFD)



 Method for numerically solving the Navier-Stokes equations of fluid motion for a given model

Widely used in studying cardiovascular flows by using patient-specific vascular models for the fluid domain
 requires input from medical imaging

 CFD yields better spatio-temporal resolution than currently possible with medical imaging

 But patient-specific modeling is dependent on high fidelity imaging to ensure proper *boundary conditions* on the numerical simulations.

# Surgical Planning: Preliminary Designs and What-if Scenarios?

Develop a framework for the surgeon to:

- Envision different scenarios before going into the operating room
- Quantitatively evaluate flow for each of these scenarios
- Estimate the power losses for each scenario
- Optimize the flow distribution to both lungs

## Virtual Surgery Environment



#### **CASE 1:**

# **Correcting Failing Fontans**

2007: Failing Fontan patient presented to CHOP with

- 4 yr. old; Heterotaxy syndrome; Functionally-single ventricle
- Arterio-venous malformations (AVMs) in the left lung.



#### **Patient's Anatomy and Flow**

#### **TCPC anatomy (from MRI)**

**3D PCMRI flow field** 



### Patient's Anatomy and Flow

#### **TCPC anatomy (from MRI)**



**3D PCMRI flow field** 



# Simulations of Pre-Operative Anatomy





#### Objective

**To plan the re-operation, using image-based** computational fluid dynamics (CFD), that will best distribute hepatic factors to the right and left lungs in order to reverse the AVMs

 Assess pre-operative hemodynamics to better understand problem

 Investigate alternate TCPC designs and identify the one that best distributes hepatic blood flow

## **Investigated Options**





Split IVC in 2 branches

Connect AZ to IVC

#### Connect IVC to AZ

# Performance of Investigated Options

**RPA** 

94%

LPA

6%



# Surgery and Clinical Follow-up

- Surgery performed in February 2008
- Five month follow-up:
  - 94% Oxygenation saturation levels (vs. high 60's pre-operatively)
     Significantly better clinically



# Surgical Planning of the Y-graft



## **Translational Test Bed:**

### Y-graft application Comparison to post-operative state

### CASE 2: Planning Initial Fontan Procedure

Kawashima anatomy

2 yr. old patient
HLHS and interrupted IVC
AVMs in right lung



CHOP\_M9

#### **Design Hepatic Baffle to Avoid Poor Flow Distribution Scenarios**

# **Graft Bifurcation to Reduce Sensitivity**

#### **'Traditional Extracardiac'**



**Y-graft** 





#### **Surgical Implementation**





# Comparison to Post-Operative ResultsPre-operative modelPost-operative results



#### **Post-Operative Hemodynamic Outcome**



#### **Post-Operative Hemodynamic Outcome**

#### Post-op Anatomy



#### Virtual model



# Application to Diagnosis and Risk Assessment

Patients with interrupted IVC and Azygous continuation

#### CASE 3: Clinical Case Report

#### Nov. 2009:

- 6 yr. old male with heterotaxy syndrome, unbalanced AV canal to the RV, VSD, hypoplastic LV, and an interrupted IVC with azygous continuation
- Had undergone TCPC completion in Summer 2009, coming for a follow-up scan

Assess risk of PAVM development

TCPC anatomy And time-averaged flow distribution



Cardiac Output: 2.8 L/min

#### In Vivo PC MRI

■ Important flow reversal in the hepatics

Data loss in the SVCs and connection areas due to the limited number of PC MRI slices and high flow disturbances







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#### In Vivo PC MRI

■ Important flow reversal in the hepatics

Data loss in the SVCs and connection areas due to significant de-phasing from flow disturbances and intense mixing





#### Pulsatile CFD Simulations for Detailed In Vivo Hemodynamics

Hepatic streamtraces almost exclusively go to the RPA, apart from a few instances at the peak of the hepatic flow curve



Cardiac Output: 2.8 L/min

#### **Distribution of the Hepatic Nutrients**

- Hepatic nutrients are seeded towards the hepatic inlet and typically take 1.5 cardiac cycle to reach the RPA
- No hepatic particles succeeded to reach the LPA
- This patient might develop left-sided <u>PAVMs</u>

TCPC anatomy And time-averaged flow distribution







#### Summary

- Magnetic Resonance Imaging provides:
  - patient-specific anatomy: computational geometry
  - flow reconstruction: boundary conditions.
- Virtual surgical environment assists in:
  - parametrically investigating various patient-specific surgical options
  - designing pre-operatively the best surgical connection on a patient-specific basis
- Cutting-edge computational fluid dynamic model assesses
  - hepatic flow distribution
  - power loss
  - baffle pressure drop

## Future Challenges











Scan

Image Processing Mesh Numerical Method

Post-process

 Improve image processing to gain more realistic geometries more efficiently

Improve efficiency and quality of mesh generation

Obtain realistic boundary conditions and material properties

 Develop more efficient and more reliable numerical methods (eg: fully coupled FSI with moving boundaries)

Develop more user-friendly and automated post-processing tools

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