# Determining Parathyroid Vascularity in a Clinical Setting

Vanderbilt University

# **Department of Biomedical Engineering**

Gabriella Caires de Jesus Tianhang Lu Itamar Shapira James Tatum Yu Zhou

## Abstract

Endocrine surgeons at the Vanderbilt University Medical Center have requested a device that will aid in the identification of perfused parathyroid tissue over poorly perfused parathyroid tissue. If a parathyroid gland's perfusion source is damaged during surgical operation, it may become unperfused and lose parathyroid function, specifically bodily calcium homeostasis; moreover, in such a situation the parathyroid is minced and reimplanted in the sternocleidomastoid muscle. Consequently, physicians have a need for quantification of the perfusion of parathyroid gland. To address this clinical need, a team of senior undergraduate biomedical engineering students will design and build a device to quantify the perfusion of parathyroid tissue in order to replace the subjective visual method currently employed. The team members are Gabrielle Caires de Jesus, Tianhang Lu, Itamar Shapira, James Tatum, and Yu Zhou. The advisors for this project are Dr. Matthew Walker III, Dr. Anita Mahadevan-Jansen, and PhD student Melanie McWade. The overall objectives for this project are as follows:

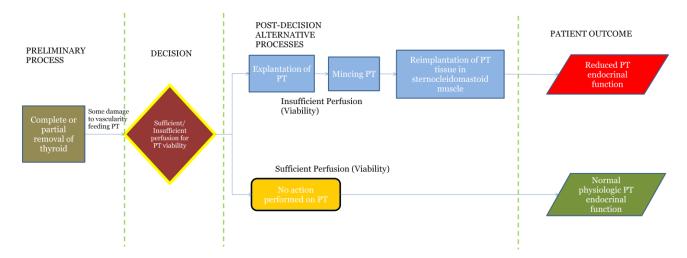
- Consider and assess several imaging techniques to determine parathyroid tissue perfusion and to select an optimal technique for use in a clinically useful device.
- Implement optimal imaging technique in a sterilizable, clinically safe, and rapidly analyzing device adapted to use by surgeons *in situ*. Ideal implementation of the technique will result in a modular device easily integrated into the existing fluoroscopy instrumentation.
- Establishing proof of concept (POC) with *ex vivo* animal testing and preliminary clinical trials and creating documentation to encourage further licensure agreement with Anasys Instruments.

The success of the project will be based off our device's ability to determine parathyroid perfusion in a clinical setting.

## Introduction

#### **Problem Statement:**

Endocrine surgery often requires thyroidectomy or parathyroidectomy. Following total or incomplete thyroidectomy, parathyroid tissues are often exposed. During such procedures, vascular tissues connecting the parathyroid gland to circulation may be damaged. Doctors must decide whether to reimplant the parathyroid tissue or leave it in the thyroid bed; however, surgeons lack an objective measure to make such a determination.



## Needs statement:

A means of assessing tissue viability is necessary, whereas perfusion of tissue is a chief indicator of such viability; thus, the device should give a quantitative flow measure. The device should allow extremely rapid collection of data such that it may be used in surgical settings; consequently, an optical method will be required. Any optical technology should take care not to injure (ablate) sensitive parathyroid tissue. The device should build off of the success of previous devices intended to assist in thyroidectomies and parathyroidectomies: this eases obstacles that practitioner dogma poses and may encourage late adopters to consider the device. A modular device with high compatibility and easy integration with existing thyroidectomy and parathyroidectomy accessories is highly desirable. Regardless, such equipment must be compact and portable, such that it will not obstruct bench space or reduce ergonomics in a surgical environment. Lastly, producing a modular device will be cost effective as well.

## **Team Composition**

The team working on this project is composed of senior level biomedical engineering students. Each member has completed the same core coursework, and the only differences are in the electives and extracurricular research or employment positions. Itamar Shapira is the team's communication and organizational lead. He outlines the goals for each meeting to ensure the goals are completed in a timely manner. He also handles the communication between the advisors and keeps the rest of the team informed. James Tatum is the optics lead for the group. He has taken the biomedical engineering optics course and has the most experience regarding the fundamentals of lasers within the group. He ensures that the team receives the necessary education in the background material and suggests avenues of further scientific inquiry. Tianhang Lu is a double major in both biomedical engineering and economics. He has previous experience in product design and development. His role will primarily focus on the financial and marketing aspect of the project. Gabriela Caires de Jesus maintains documentation for the group and has experience with a diversity of imaging techniques. Yu Zhou has researched in the biophotonics lab and has worked with several methods of optical spectroscopy. He has an interest in the medical field and will be the medical lead for the group, maintaining contact with the medical personnel associated with the project.

The DISC profiles of the team members match up well with a balanced group. We have one D dominant profile and the rest are SC dominant or C dominant. This is a good balance mostly due to the single D (dominant) profile. Multiple Ds can negatively impact group

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productivity due to the controlling nature of that profile. The S profiles are perfect for pushing through the workload and the Cs are great for making sure everything is done correctly. Though we are missing the I personality, the group has a good level of cohesion which offsets the need for an I dominant person.

## **Project Advisors**

There are three advisors who will be providing guidance on this project.

*Dr. Anita Mahadevan-Jansen* is Assistant Professor of Biomedical Engineering and is a leading expert on the use of laser spectroscopy to identify brain tumors.

*Dr. Matthew Walker III* holds the full time faculty Position of Associate Professor of the Practice of Biotechnology/Bioengineering at Vanderbilt University. His research includes unveiling systems level orthogonal biomarkers at the structure/function interface of disease. *Melanie McWade* - Is a Biomedical Engineering PhD candidate in Dr. Mahadevan-Jansen's laboratory currently studying the fluorophore that gives thyroid tissue its high fluorescence.

## **History and Context**

We initially conducted research on current vascularity identification techniques. A chart of potential imaging solutions has been iteratively revised, initially growing before being narrowed to several leading candidates. Following a meeting with our clinician, Dr. James Broome, the focus of our progress has pivoted to an examination of parathyroid gland perfusion as a main criterion of tissue viability in the thyroid bed. At the end of the fall semester, several viable techniques are being considered for a best candidate for device implementation in the spring semester. Presently, the leading candidates are laser speckle imaging and laser Doppler flowmetry. Dr. Anita Mahadevan-Jansen's biophotonics laboratory is currently testing a fluoroscopy device that can distinguish parathyroid tissue from thyroid tissue using near infrared spectroscopy (NIRS). Our first intended users will be practitioners at Vanderbilt University Medical Center (VUMC) currently using the fluoroscopy device to distinguish between thyroid and parathyroid tissue. These practitioners have a high patient volume for such procedures and their high levels of experience makes them optimal early adopters.

### **Work Process and Outcomes**

The primary outcome of this project is to create a device that can determine the perfusion state of the parathyroid gland remaining in the thyroid bed, improving patient outcomes following endocrine surgery. This outcome must be met for the project to be considered successful. Ideally, a threshold value for perfusion can be determined and the flow can be quantified and compared to this threshold value.

In order to meet this succeed in device development, several objectives must be completed in order to have a functioning device. We must assess multiple imaging techniques and select an optimal method. POC must be verified in *ex vivo* experiments. Several optical methods of are currently under consideration, including spectroscopic techniques.

Demonstration of POC may be aided by the creation of a phantom. Creating such a phantom will allow repeatable demonstration of the diagnostic capabilities of the device.

We are pursuing several secondary outcomes, including modular design and rapid data acquisition. Ideally, we intend to construct a modular component easily integratable with the current fluoroscopy device that is being tested by the Biophotonics department. A modular device will lower costs and increase the ease of acceptance by physicians already familiar with the fluoroscopy device. The device should allow for rapid data acquisition given its use in a surgical setting. The current fluoroscopy device requires three seconds to produce each of five or six measurements.

## **Evaluation and sustainability plan**

The success of our project depends on the ability of our imaging device to (quantitatively) determine perfusion state of parathyroid tissue in a clinical setting.

With respect to distinguishing perfused from unperfused tissue, our device must acquire a signal representing perfusion while filtering out distortion noise. Our method of detection may have a defined threshold in order to aid surgeons considering explantation and reimplantation. Also in evaluating our level of success, it is vital that our optical technology does not injure tissues. There are several existing candidate imaging solutions that might meet our criteria. Given this, the achievability creating a device capable making such determination of perfusion is feasible within the Senior Design project timeline.

Evaluation of the success comes in the form of POC documentation. Data created using a good phantom model for the parathyroid will demonstrate diagnostic success.

Maintaining similar data acquisition times to the preceding fluoroscopy device is a measure of high success. Mechanical and data analysis integration with the preexisting fluoroscopy device will be a measure of high success.

## Appendix

## Budget

Our budget greatly depends on the method for determining perfusion. The optical probe price varies depending on the spectroscopy method used. The current fluoroscopy device uses a \$5,000 probe while simple fibers (SiO<sub>2</sub>) costs less than a dollar. In vivo specimens would be necessary for POC. We have permission to use all of the spectroscopy devices owned by

Vanderbilt Biophotonics. If we determine that a device not owned by Vanderbilt is needed, then we would need to add that to the expenditures. Also, costs of possible future integration with current techniques could be part of the expenditure, but that is very far in the future.

# Tianhang (Kevin) Lu

Permanent Address: 6799 Parklake Dr. Mason, OH 45040

Tianhang.lu@vanderbilt.edu (513) 295-8220 Current Address: 1215 15<sup>th</sup> Ave. South Nashville, TN 37212

## **EDUCATION**

Vanderbilt University, Nashville, TN Double Major: Bachelor of Engineering in Biomedical Engineering Bachelor of Arts in Economics Expected Graduation Date: May 2014

## WORK EXPERIANCE

WORK EAPERIANCE	
Osto360, Nashville TN	Summer 2013
Biomedical Engineering Intern	
<ul> <li>Integral part of the Research and Development team</li> </ul>	
• Identified materials and mechanisms that led to the successful proof	of concept for a new product
• Developed first prototype for a new planned service	
• Compiled and maintained a directory containing over 2000 relevant	medical professionals
• Part of multiple investment pitches including the company's bid for	Healthbox
<ul> <li>Maintained relationships with partners</li> </ul>	
Vanderbilt Dining	2011 to 2012
• Food Server at CT West	
Lumen Learning Center	2009 to 2010
• Math tutor for students grade 1 through 12	
EXTRACURRICULAR EXPERIENCE	
Vanderbilt University Crew Team	2010 to 2011
• Rower for the Vanderbilt Crew Team	
• Dedicated 22 hours a week and over 800 total hours	
Vanderbilt Students Volunteers for Science	2012 to 2013
• Taught students at local schools	
• Lead a group of 5 every week	
Greater Cincinnati Chinese School	2007 to 2010
Volunteer Director	
<ul> <li>Founded the volunteering section</li> </ul>	
<ul> <li>Distributed responsibilities to other volunteers</li> </ul>	
<ul> <li>Dedicated 5 hours a week, 750 hours total</li> </ul>	

# LANGUAGE SKILLS

- Fluent in Mandarin
- Studied Spanish for 4 years

## **COMPUTER SKILLS**

• VB, C++, JAVA, MATLAB, Microsoft Office, LabVIEW, R Commander, Computer Hardware Installation

# James Tatum

james.m.tatum@vanderbilt.edu (901)497-8920

Current Address: Vanderbilt University PMB 356703 2301 Vanderbilt Place Nashville TN 37235 – 6307 Permanent Address: 6735 Spencer Forrest Cv E Memphis, TN 38141

## Education

**Vanderbilt University**, Nashville, TN Bachelor of Engineering, Biomedical Engineering (August 2010 to May 2014) Pre-medical

• Engineering GPA: 3.1/4.0

## Coursework

Circuits I • Circuits II • Biomaterials • Linear Algebra and Differential Equations • Matlab Systems Physiology • Fundamentals of Medical Imaging • Biological Basis of Imaging

## Skills

- MatLab
- Mathematica
- Photoshop CS5
- Photography

## Experience

**Kelly Services at New Breed Logistics**, Memphis, TN *Materials Handler* (June 2012 to July 2012)

- Picked orders for New Breed Logistics
  - Worked with a team of 10-14 pickers to complete 4000 to 8000 orders per day on schedule

## Leadership

## Vanderbilt Running Club

Practice Captain (January 2010 to Present)

- Designed workouts and led groups of 5-10 runners through Nashville
- Ensured all runners in my group completed the workout safely

Apparel Coordinator (January 2012 to Present)

- Designed t-shirts and running singlets for use by the club
- Worked with suppliers and other club board members to ensure everyone was satisfied with the product

# Gabriela Caires de Jesus

Current Address Vanderbilt University, PMB 350012 Nashville, TN gabriela.caires.de.jesus@vanderbilt.edu (615) 602-6317 Permanent Address: Rua Praia de Touros , 9135 Natal, RN, Brazil, 59092-110

EDUCATION	ſ
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#### VANDERBILT UNIVERSITY, Nashville, TN **Exchange Student** September, 2013 to May, 2014 GPA: -/4.0 FEDERAL UNIVERISTY OF RIO GRANDE DO NORTE, Natal, RN, Brazil Bachelor of Engineering, Biomedical Engineering February, 2010 to December, 2015 GPA: 3.5 / 4.0 **RELEVANT PROJECTS** FEDERAL UNIVERISTY OF RIO GRANDE DO NORTE, Natal, RN, Brazil Development and construction of a device to measure the conductivity of a solution The circuit was set up on a BreadBoard using basic electrical components • February, 2013 to June, 2013 Assembly of an ECG/EMG/EEG using basic electrical components February, 2012 to July, 2012 The device was assembled at a pre-printed circuit with basic electric components and the achieved signal was then filtered using software Pure Data Article: "Atuação do Engenheiro Biomédico no Hospital Universitário Onofre Lopes" October, 2012 "Performance of the Biomedical Engineer at the University Hospital Onofre Lopes" Article presented at the 18th CIENTEC - Week of Science, Technology and Culture Development of a Matlab script to evaluate the gait February, 2012 to July, 2012

- The script was generate using the Robotics Toolbox
- Data from joints was used to simulate a robot in order to achieve quantitative information about the gait

#### WORK EXPERIENCE

DISTANCE EDUCATION SECRETARIAT OF THE FEDERAL UNIVERSITY	OF RIO GRANDE DO NORTE
Test analyzer	November, 2012 to July, 2013
<ul><li>Worked testing the software that was developed to coordinate all the courses</li><li>Was part of the customer support team</li></ul>	of the Distance Education at the University
FEDERAL UNIVERSITY OF RIO GRANDE DO NORTE	
<ul> <li>Assistant of the Brazilian Robotics Olympiad</li> <li>Collaborated with the organization of the Brazilian Robotics Olympiad</li> </ul>	May, 2010 to December 2010

#### CCAA – Anglo-American Cultural Centre

**English Teacher** 

• Worked as an English teacher mostly for kids and teenagers

#### SOCIAL EXPERIENCE AND SPORT

Kung Fu Classes

May, 2012 to February, 2013

September, 2009 to July, 2010

#### COURSEWORK

#### VANDERBILT UNIVERSITY

Biomedical Instrumentation • Design of Biomedical Devices and Systems I • Senior Engineering Design Seminar • Image Processing • Introduction to Tissue Engineering

#### **TECHNICAL SKILLS**

Matlab • MS Office Suite

#### LANGUAGES

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**Itamar Shapira** 

11006 Stone Mountain Dr. Huntsville, Alabama 35803 (256) 658-2318 itamar.shapira@vanderbilt.edu DOB: July 2, 1993

Educational History: Virgil I. Grissom High School 7901 Bailey Cove Rd SE Huntsville, AL 35802 (256) 428-8000 Attended: Freshman - Senior

University of Alabama at Birmingham (UAB) Birmingham, Alabama, 35294 Attended: First year Undergraduate

Vanderbilt University Nashville, Tennessee, 37240 (615) 322-7311 Attended: Second year Undergraduate-Present

#### Major: Biomedical Engineering

#### Awards/Honors/Achievements:

First Year:

Science and Technology Honors Program Alpha Lambda Delta Honor Society (Top 5% Freshman Class) Freshman Forum Member (UAB USGA) UAB Golden Excellence Scholarship UAB Presidential Honors (both semesters)

#### Second Year:

Fueling Innovation Forum Summer 2013 (UAB Science and Technology Honors Program)

#### Third Year:

[On track to become] Fastest graduate of Vanderbilt Undergraduate Biomedical Engineering Department

#### **Research Experience:**

First Year:

Student Researcher in lab of Dr. Ho-Wook Jun, Fall 2011/Spring 2012 Biotechnology and Research Applications Course, Spring 2012

#### Second Year:

Student Researcher in lab of Dr. Elizabeth Sztul Summer 2013

#### Skill Set Specifics

Familiarity or competence with the following biological laboratory techniques:

- Basic biotechnology: PCR, Agarose Gel Electrophoresis, Polyacrylamide Gel Electrophoresis, Bacterial Plating, Ligation, Transformation, Plasmid DNA Isolation, Spectroscopy, Restriction Enzyme Digest, Protein Isolation, Cell Culture

#### Competent:

-Statistics, Word, Excel, Powerpoint, MATLAB Scripting, Creo Parametric Solid Modeling, Engineering design, Scientific/Technical writing

#### Extra-curricular Activities:

#### First Year:

Freshman Forum Member (UAB USGA)

- Student Issues Board

Biomedical Engineering Society (BMES)

Second and Third Years:

Vanderbilt Hillel

-Religious Life Committee member

-Hillel Israel Committee member

-Founder of Tiyul (Hillel's first Jewish outdoors group)

Dores for Israel (pro-Israel political advocacy organization)

-Attendee to AIPAC Saban conference in Washington, D. C., January 2013, July 2013

-Attendee to AIPAC Policy Conference in Washington, D. C., March 2013

- Chabad Jewish Student Organization
- -Chabad Student Board

#### Miscellaneous:

Multilingual (fluent in Hebrew and English; some French and German) Avid runner

Mobile 513.259.8365 Station & 6915 Nashville TN 37235 Yu Zhou yu.zhou.1@vanderbilt.edu

## Academics

Vanderbilt University Current major: Biomedical Engineering GPA: 3.636 Year of graduation: 2014 Hours earned: 116.0 Quality hours: 88.0

#### Extracurricular Activities

Aug 10	Vanderbilt Rowing Team (18 Hrs/Week + Competition on weekends)
to Present	<u>Team President</u> : liaison to the University, approves all team funds and monitor team finances, leads recruitment and fundraisers, organize and run board meetings
	<u>Varsity Rower</u> : Rowed in the top varsity boat for the past 2 years, rowed in the top freshmen boat freshmen year. Competed in nationwide regattas including Head of the Charles (world's largest two-day regatta), championship events in the Head of the Hooch (second largest regatta in U.S: champ 4+ and champ 8+).
	ACRA: American Collegiate Rowing Association (national club rowing competition)
	15th (novice 8+, 2011) 16th (varsity 8+, 2012) 18th (varsity 8+, 2013)
	2-time ACRA Second Team Academic All-American
Research	

#### Research

Jan 11 to May12	The SyBBURE Searle Undergraduate Research Program (10Hrs/Wk) SyBBURE stands for Systems Biology and Bioengineering Undergraduate Research Experience Two paid research projects in the SyBBURE microfluidics lab: <u>Independent project</u> : Finding a cheaper and more hydrophobic alternative to dialysis membrane using block co-polymers and infused porogens. The filter is designed for improved reading of mass spectrometry of biological samples.
May 12	Team project: Building a microfluidic device that tracks over 40 generations of yeast cells (10 <sup>12</sup> number of cells) to determine the mechanisms of cell aging. My part in the project is in the design and fabrication of the yeast tracking device <b>Research Assistant in the Department of Biophotonics</b> (40Hrs/Wk)
to Aug' 12	The focus of the research is the development of optical spectroscopy for diagnosis of disease and guidance surgery. My involvement is the characterization of bone matrix properties of breast and prostate bone metastasis in mice. I identified the characteristic patterns of cancerous bone metastasis in mice.
Volunteer Service	

Jan '12 Monroe Carell Jr. Children's Hospital Volunteer (2Hrs/Wk) Part of a national reading program. Job includes reading books in the waiting room to children demonstrating to parents and caregivers how to read to children. Also encouraged visiting families to sign up for Tennessee reading programs, that sends books to children to encourage reading and improve literacy. (2 Hours/Week) to May 12

## **Additional Supporting Documents**



Figure: Near-infrared fluorescence spectroscopy system showing a (1) spectrometer, (2) 785-nm diode laser, (3)

laptop computer, and (4) hand-held fiber optic probe. (Courtesty of Mahadevan-Jansen laboratory)