PROJECT TITLE: Improving skeletal muscle function in distal arthrogryposis

Sakthivel Sadayappan, PhD, MBA
Professor of Internal Medicine
Associate Chairman for Basic Research,
Internal Medicine
Director of Heart Branch of the Heart,
Lung and Vascular Institute
Division of Cardiovascular Health and
Disease, College of Medicine
University of Cincinnati

Cardiovascular Center, Rm 4935
231 Albert Sabin Way
Cincinnati, OH 45267-0575, USA
Phone: +1 513-558-7498
FAX: +1 513-558-2884
Email: sadayasl@ucmail.uc.edu

Project Description

1. Area of the research
Distal arthrogryposis is an autosomal dominant congenital anomaly characterized by contractures of the distal regions of the hands and feet with no facial involvement or any additional anomalies prior to birth. Distal arthrogryposis type 1 affects an estimated 1 in 10,000 people worldwide. Distal arthrogryposis is related to decreased fetal movement during development which can be caused by environmental factors, single gene changes, chromosomal abnormalities and various syndromes. Treatment depends on the signs and symptoms found in each person, but it may include physical therapy, removable splints, exercise, and/or surgery. Research studies involve characterizing skeletal muscle function by determining muscle-specific gene and protein expression and measuring skeletal muscle function in vitro.

2. Research tasks the student will be performing
Undergraduate students will perform DNA and RNA extraction, real-time polymerase chain reaction, agarose gel electrophoresis, protein isolation, quantitation, standard graph creation and Western blot analysis. Furthermore, students will use skinned muscle strips from mouse model of distal arthrogryposis to measure skeletal muscle function in vitro using the Aurora
Scientific 1300A 3-in-1 System. Briefly, the muscle of interest is surgically
removed and placed in an oxygenated and temperature-controlled bath to
maintain viability. One tendon is anchored to a fixation clamp, while the
other tendon is sutured onto a dual-mode lever. The muscle is then
stimulated by field electrodes fixed inside the bath to measure muscle
contractility. This technique requires two weeks of training prior to
performing the experiments.

3. Training that the mentor will provide to the WISE student
The objective of the twelve-week Women in Science and
Engineering (WISE) Program is to expose undergraduate women to skeletal
muscle research. The skeletal muscle system is responsible for movement of
the human body, posture, movement of substances inside the body and
generation of body heat. Therefore, studying skeletal muscle physiology in
health and disease is essential for those contemplating careers in skeletal
muscle biology. We hypothesize that undergraduate students fill these
positions, as the upcoming innovators who will bring new ideas and
technologies to improve skeletal muscle function in distal arthrogryposis. As
proposed in this study, students will be exposed to systematic and critical
thinking, planning, teamwork, experimental design and documentation thereof.
They will learn how to review the pertinent literature, ask relevant
questions, and develop an appropriate hypothesis. They will learn how to
perform experiments in a rigorous and reproducible manner. In addition,
students will be asked to attend research seminars and present the results of
their studies at lab meetings. Overall, the WISE training plans will include
orientation, hands-on research experience, weekly discussions of projects,
weekly journal club, workshops on pertinent research tools, and poster
presentations at the one-day local College of Medicine symposium.

4. Specific requirements, if any, that the mentor expects the student to meet
We prefer students who are highly motivated and interested in skeletal muscle
research which is another focus of our research group.