

#### Dear Friends and Colleagues,

This year has seen great progress in our research programs targeting paralysis. Five FDA-approved clinical trials targeting spinal cord injury (SCI) and peripheral nerve injury continue to provide exciting findings. The Phase 1 safety trial evaluating autologous human Schwann cell transplantation in subacute injury subjects is nearing completion with promising results. Millions of Schwann cells have been successfully transplanted into four subjects with no adverse effects. Based on these promising

results, we have now obtained permission form the FDA to test this therapy chronically injured individuals for the first time. This chronic SCI trial will be the first to combine cell transplantation with extensive rehabilitation. Our FDA approved trial for repairing a severe peripheral nerve injury has also shown promise in promoting return of motor and sensory function in the leg of one subject. To treat neuropathic pain after SCI, we are testing the beneficial effects of Deep Brain Stimulation. This FDA approved program has already resulted in encouraging findings that suggest a new strategy

of State School 74 Ch School Physics Par Wheel Physics

Barth Green, M.D.,F.A.C.S. W. Dalton Dietrich, Ph.D.

for addressing this difficult to treat condition experienced by many individuals living with SCI. Our Brain-Machine Interface FDA program is merging biological and biomedical disciplines to show proof-of-concept of operating an upper extremity neural prosthetic thereby enhancing independence. Finally, another FDA approved trial testing the safety and efficacy of adult mesenchymal stem cells has also shown encouraging results

in one subject. Together these reparative trials represent the most comprehensive program in the world testing experimental therapies for spinal cord injury and related disorders.

The benefits of therapeutic hypothermia and targeted temperature management in acute brain and spinal cord injured patients continue to be recognized. Over 40 acutely injured SCI patients have undergone this experimental therapy that is

showing long term benefits in neurologic function. In the area of traumatic brain injury, the most severely injured individuals are also being treated with cooling strategies in an international multicenter trial. clinical These neuroprotective programs are additional examples of how Miami Project-led basic and translational studies have been successfully moved into the clinic.

Multiple clinical programs including our Miami Project "Boot Camp" are actively investigating other aspects of SCI that may also improve function. Our ultimate goal is to

combine the state-of-the-art rehabilitation and conditioning strategies with cell therapies and other regenerative approaches to target functional recovery, neuropathic pain, male fertility, spasticity, and bladder function. The Miami Project is committed to developing whole life strategies that can maximize quality of life and good health as our scientists continue to strive to develop new therapeutic interventions.

Discovery research, which fuels our translational and clinical programs, is clarifying molecular and cellular mechanisms underlying cell death, axonal regeneration, and circuit repair. Ultimately, this knowledge will be combined with our current therapeutic interventions to maximize functional recovery. The clarification of critical gaps in our knowledge regarding axonal regeneration and circuit plasticity will improve our chances of developing successful cures for paralysis.

The Miami Project to Cure Paralysis was established in 1985 to develop novel therapies to improve function in paralyzed individuals. Today, our discoveries are being successfully translated to people and are changing the way we provide clinical care. Our program is unique in that it touches upon multiple areas of medical research including education, discovery, translational and clinical trials. These are very exciting times within The Miami Project to Cure Paralysis and we thank our friends, colleagues, and research participants for their long-term support and commitment to our research.

Sincerely,

W. Dalton Dietrich, Ph.D.

Scientific Director, The Miami Project to Cure Paralysis Kinetic Concepts Distinguished Chair in Neurosurgery Senior Associate Dean for Discovery Science Professor of Neurological Surgery, Neurology and Cell Biology

Barth A. Green, MD Co-Founder and Chairman, The Miami Project to Cure Paralysis Professor and Chairman, Department of Neurological Surgery Professor, Departments of Orthopaedics and Rehabilitation Medicine University of Miami Miller School of Medicine



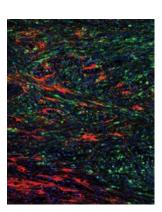
#### Research Review 2014

2	Message from the Chairman, Dr. Barth A. Green
	and Scientific Director, Dr. W. Dalton Dietrich

5 Fundraising, Administrative, and Scientific Support Staff

#### RESEARCH HIGHLIGHTS

- 6 Phase I Chronic Schwann Cell Trial
- 9 Monica Perez, PT, PhD, Martin Oudega, PhD, Join The Miami Project
- 10 Hyperthermia in Neurotrauma
- 11 Faculty Highlight Mousumi Ghosh, PhD
- 12 Whole Body Vibration and Bones
- 13 Engineered Schwann Cell Grafts
- 14 Active Clinical Studies and Trials
- 16 Educational Outreach
- 18 Research Funding in 2014
- 22 Faculty Profiles
- 33 NIH Summer Student Research Highlights
- 34 Scientific Research Publications 2014



On the Cover Schwann cell transplant: host cord interface in chronic spinal cord injury model

**Editor & Contributor,** Kim Anderson-Erisman, Ph.D.

Managing Editor, Scott Roy
Graphic Editor, Robert Camarena
Contributors, Diana Berning, Dalton
Dietrich

**Photographs & Graphics,** Robert Camarena, Mousumi Ghosh, Damien Pearse, Haruo Kanno, Coleen Atkins, Helen Bramlett

Website: www.themiamiproject.org

#### **Physical Address:**

1095 NW 14th Terrace, Miami, FL 33136 Phone: 305-243-6001

#### **Mailing Address:**

Post Office Box 016960, R-48 Miami, FL 33101

The Project is published biannually by The Miami Project to Cure Paralysis, a Center of Excellence at the University of Miami Leonard M. Miller School of Medicine.
All contents ©2015. Reproduction in whole or in part without written permission is prohibited. The University of Miami is an Equal Opportunity/Affirmative Action Employer.

Privacy Policy: If you do not wish to receive *The Project* in the future, please send your request to the address above or call 305-243-7147.

The Miami Project scientific team is grateful for the dedication and hard work of the fundraising, administrative, and scientific support staff. This incredible group of people spend countless hours providing direct clerical and administrative support to the research staff, and raising the precious private funds to support Miami Project research endeavors.



## Fundraising, Administrative, and Scientific Support Staff

#### Management

Marc A. Buoniconti President

Stephanie Sayfie Aagaard Director, Major Gifts, Corporate Relations, Marketing and Events

> Randy Carlson Director, Finance

Francine Y. Schoen, MLS Development Director, Foundation Relations

> Marcela Ward Manager, Human Resources

Suzanne M. Sayfie *Executive Director* 

Kim Anderson-Erisman, Ph.D. Director, Education

Randy L. Medenwald Director, Government Relations

Kristin Wherry Director, Buoniconti Fund National Chapters

Cynthia Jones Manager, Facilities and **Operations** 

Diana C. Berning Center Administrator

Teri I. Bendell Director, Web Services, Auction, and Database

> Scott P. Roy Director, Public Relations and **Communications**

Jacqueline Manzano Manager, Marketing

Jon-Roi Vega Senior Manager, Sponsored Programs

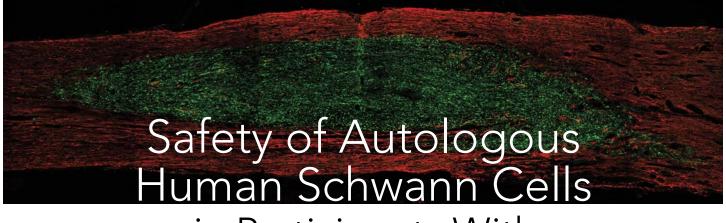
#### **Support Staff**

Teresa Arcay Angel Loor Rebecca Avshalom Maria-Jose Luna Javier Burgos Maria Muniz Robert Camarena Nathalie Perez Maria Chagoyen Carole Robbins Danielle Cilien Pablo Sanchez Georgina Escobar Maria Serna Letitia Fisher Joel Sola **Evelyn Gomez** Erika Suazo Cynthia Jones Jacqueline Tommasino

#### Scientific Staff & Trainees

Michele Kenny

Alexander Marcillo Post-doctoral Fellows **Graduate Students** Medical/Residents/Observorships **Undergraduate Students** Volunteers Other students Research Staff



# in Participants With Chronic Spinal Cord Injury Receiving Rehabilitation

A t long last, we have begun our first clinical trial testing autologous human Schwann cell (ahSC) transplantation in people living with chronic spinal cord injury (SCI). This is also a Phase I trial focused on safety and feasibility outcomes. Since October 2012, we have been conducting a separate Phase I trial targeting sub-acute SCI.

As we announced in October 2014, we obtained FDA approval after having submitted for review additional data regarding transplantation of Schwann cells into rodents and some pigs with chronic SCI. We now have ethics approval from the University of Miami Institutional Review Board and are open for enrollment.

This new trial will be primarily focused on safety, but in addition it will involve a preliminary evaluation of the efficacy of combining Schwann cells with exercise and rehabilitation. For humans with chronic SCI, we hypothesize that axons might show improved function if myelin repair is induced with the implantation of ahSC. In addition, spinal cord cavitation may be reduced and neural sprouting and plasticity may be enhanced via neurotrophic effects. In this trial, participants will receive fitness conditioning and locomotor rehabilitation prior to transplantation in order to validate the stability of their neurological baseline and enhance their ability to undergo surgery with few complications. They will also receive fitness conditioning and rehabilitation post-transplantation to maintain health and promote neuronal activity and potential neuroplasticity.

We only have FDA approval to transplant a maximum of 10 people – remember that it is a Phase I safety trial. In order to reduce risk, there are a number of inclusion and exclusion criteria. Some of which are listed here.

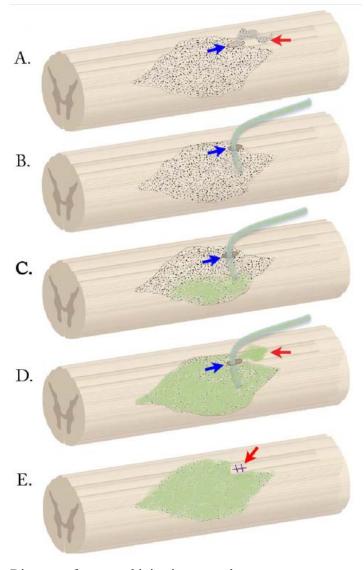


Diagram of proposed injection procedure

#### Participant inclusion criteria:

- Persons with traumatic SCI that occurred a minimum of 12 months prior to enrollment;
- Between the ages of 18 and 65 years at last birthday;
- SCI between cervical level C5 and thoracic level T12 as defined by the most caudal level of intact motor and sensory function on the International Standards for Neurological Classification of Spinal Cord Injury (ISNCSCI):
- ASIA Impairment Scale (AIS) grade A, B, or C at time of enrollment;
- Spinal cord lesion size that is less than or equal to 3 cm in length and less than or equal to 2 cc in volume when viewed on magnetic resonance imaging (MRI)

#### Participant exclusion criteria (not all are listed here):

- Persons with penetrating injury of the spinal cord or complete transection of the cord, as identified by MRI;
- Persons with a cavity structure that would preclude successful transplantation, as identified by MRI;
- Intolerance to functional electrical stimulation of muscles:
- Presence of disease that might interfere with participant safety, compliance, or evaluation of the condition under study;
- Body Mass Index (BMI) > 35;
- Persons with pre-existing conditions that would preclude satisfactory sural nerve harvest;

We will be staggering enrollment in 2 cohorts. Progression through the cohorts is based on safety, going from low risk to higher risk, and there are certain lock-out periods between participants.

#### Cohort 1: Thoracic Injuries with Grades A, B, C

We will enroll up to 4 participants who have thoracic spinal cord injuries between the levels of T2 to T12 with Grades A, B, or C in cohort 1.

We will enroll 2 participants with Grade A injuries before we enroll any participants with Grades B or C injuries.

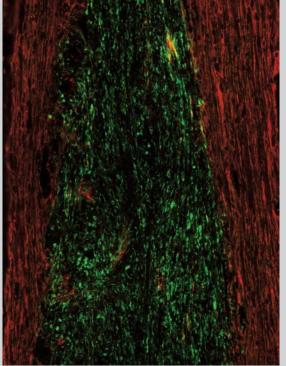
#### Cohort 2: Cervical Injuries with Grades A, B, C

We will enroll up to 6 participants with cervical spinal cord injuries between the levels of C5 to T1 with Grades A, B, or C in cohort 2.

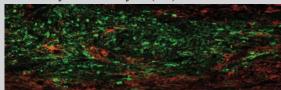
We will enroll 2 participants with Grade A injuries before we enroll any participants with Grades B or C injuries.

There are 5 stages of Screening to progress through before the actual Schwann cell transplantation occurs and each participant's time commitment to the trial will be approximately 10 months. They will be followed in a separate long-term monitoring protocol annually for a total of 5 years post-transplantation.

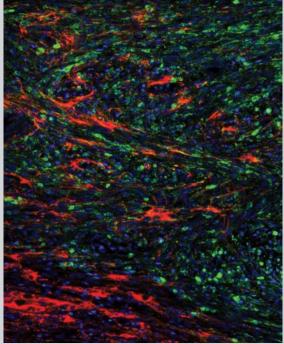
One of the stages involves harvesting a segment of each participant's own Schwann cells from one of their sural nerves, a sensory nerve above the ankle. The Schwann cells will then be derived from the nerve and processed



Rostral end of Schwann cell graft (green) surrounded by host astrocytes (red)



Entire transplant (green) surrounded by astrocytes



Caudal end of SC transplant (green) with host cord astrocyte interface (red)

in a culturing facility to generate the number of cells necessary for transplantation, and to undergo a purification process. All procedures will be conducted in Miami at the University of Miami Miller School of Medicine. Because the Schwann cells are obtained from each individual participant, there is reduced concern of immune rejection and immunosuppressant medication is not required. Once the cells are removed from the participant, they have to be handled in accordance to current Good Manufacturing Practices (GMP). Because the cells will eventually be injected into the spinal cord, this process is required to ensure that the cells are prepared consistently and without contaminants.

Another stage before the transplantation, which will resume after the transplantation, involves the fitness and locomotor rehabilitation training. This involves at-home and onsite components that must be completed every week. So, the individuals that end up qualifying and agreeing to participate have a very large commitment to think about.

#### At home, 3 days every week:

Upper Extremity Circuit Resistance Training

#### On-site, 2 days every week:

**Body-weight Supported Locomotor Training** 

Lower Extremity Functional Electrical Stimulation cycling

All procedures for this research study are for research purposes only, and, therefore, are free of charge. However, participants not local to Miami will need to relocate here for at least 10 months. For those non-local participants, we have a housing program available free of charge once they start the pre-transplantation fitness and locomotor rehabilitation training program. No travel or food expenses will be covered. There are strict regulations on the amount of money people can receive to participate in clinical trials; this is to prevent individuals from participating in risky interventions just for the money.

To find out more information about the trial and prescreening, contact The Miami Project Education Department at 305-243-7108 or MPinfo@med.miami.edu. More information about all of our clinical trials and studies is available at http://www.themiamiproject.org/trials



#### Monica Perez, PT, Ph.D.

The Miami Project is very happy to announce that Dr. Monica Perez will be joining us as a new clinical research faculty! Dr. Perez is joining us from the University of Pittsburg. She actually earned her PhD degree in the Department of Physical Therapy at the University of Miami in 2003. From there she did a post-doctoral fellowship at the University of Copenhagen followed by a fellowship at the National Institute of Neurological Disorders and Stroke (NINDS). Her research focuses on neural control of movement, particularly the corticospinal tract. She is currently funded to investigate Neural control of bilateral hand and arm movements after SCI (NINDS), Corticospinal function after human SCI (NINDS), Enhancement of hand motor function after cervical SCI (Veterans Affairs), and Role of the motor cortex in recovery of hand function after SCI (Craig H. Nielsen Foundation, CHNF). "I am extremely happy to have the opportunity to join The Miami Project to Cure Paralysis. The goal of my research is to understand how the brain and spinal cord contribute to control residual voluntary movement after SCI. Our mechanistic approach is used to develop strategies to facilitate voluntary motor output. For the next years our research will focus on two main areas. First, we will work on protocols to maximize plasticity at different sites within the central nervous system to enhance voluntary motor output using noninvasive stimulation techniques and

motor training. Second, we will combine methodologies to better characterize SCI and their impact on the neural control of movement. These methodologies include mapping of multiple motor cortical regions, estimations of activity in the corticospinal pathway and their input to spinal motoneurones, and coupling between motor cortical areas and different muscle groups."

#### Martin Oudega, Ph.D.

Dr. Martin Oudega will be joining us as a new basic science research faculty! Dr. Oudega earned his PhD at the University of Leiden in the Netherlands. He did a postdoctoral fellowship at the University of California, San Diego and then at the University of Miami as well. He became faculty at Johns Hopkins University and then the University of Pittsburgh. "I am thrilled to join The Miami Project and to start working with its great scientists and wonderful administrative and technical supporting staff. My main research focus is on developing cell-based strategies for repair of anatomical damage and improving functional impairments after SCI. I am particularly interested in the molecular mechanisms underlying cell transplant-mediated repair and in how natural and synthetic biomaterials can augment the repair potential of intraspinal cell transplants." He is currently funded to investigate ESHU for optimizing BMSC transplant survival and their spinal cord repair efficacy (Wings for Life), Role of Ang-1 in vascular stabilization in BMSC-mediated tissue spinal cord repair (CHNF), and Fibronectin for enhancing BMSC-mediated spinal cord repair (Morton Foundation).

Congratulations to Drs. Perez and Oudega and welcome to the family!

# Heat Can Worsen the Effect of Mild Traumatic Brain Injury

In the clinical setting, fever can significantly worsen morbidity and mortality after severe TBI. But what about all the mild TBIs and concussions that occur?

Our researchers have demonstrated over several years the importance of temperature in neurotrauma; particularly, how lowering the temperature (hypothermia) can be therapeutic. This research directly led to the ongoing trial at the University of Miami and Jackson Memorial Hospital evaluating the safety and efficacy of mild therapeutic hypothermia in acute cervical spinal cord injury (SCI). It also led to the current HOPES trial, *HypOthermia for Patients* requiring Evacuation of Sudural Hematoma, a randomized-controlled trial for individuals that sustained a severe traumatic brain injury (TBI) plus developed an acute subdural hematoma that requires surgical removal. Hypothermia is used during the surgical removal of the hematoma. What about elevated temperature, hyperthermia? Dr. Dietrich and others have shown that hyperthermia after moderate to severe TBI can worsen tissue damage and outcome in experimental models. In the clinical

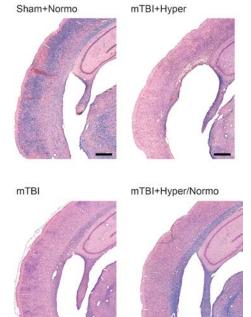


setting, fever can significantly worsen morbidity and mortality after severe TBI. But what about all the mild TBIs and concussions that occur? Is temperature management important there?

Drs. Coleen Atkins and Dalton Dietrich recently investigated whether hyperthermia at the time of a mild TBI worsened/aggravated associated cognitive deficits. In addition to a couple control groups, they tested the following injury/temperature conditions:

- 1. Normothermic mTBI a mild TBI was administered while the brain and body were at 98.6°F, then that temperature was maintained for 4 hours:
- 2. Hyperthermic mTBI a mild TBI was administered while the brain and body were at 102.2°F, then that temperature was maintained for 4 hours;
- 3. Hyperthermic/normothermic mTBI (cooled hyperthermic) - a mild TBI was administered while the body and brain were at 102.2°F; that temperature was maintained for 15 minutes, then cooled to 98.6°F and maintained for 4 hours.

Sustaining a mild TBI while hyperthermic caused greater brain tissue damage compared to a mild TBI



Examples of brain tissue damage under different temperature conditions.

while normothermic. Interestingly, bringing the hyperthermic brain temperature down to normal within 15 minutes post-injury significantly reduced the amount of tissue damage.

This also had an effect on memory. Sustaining a mild TBI while normothermic had no effect on longterm memory, but sustaining a mild TBI while hyperthermic caused persistent deficits in long-term memory. These long-term memory deficits were prevented by cooling hyperthermic mild TBI animals to normothermia within 15 min after the mild TBI. Sustaining a mild TBI while normothermic or hyperthermic caused persistent deficits in working memory, which were not improved by cooling, however. Overall, these results suggest that temperature management strategies immediately after mild TBI may be promising to pursue clinically.

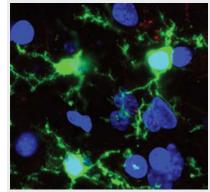


## Mousumi Ghosh, PhD

r. Mousumi Ghosh is a new Research Assistant Professor in the Department of Neurological Surgery and The Miami Project. Dr. Ghosh, a native of India, holds her Bachelors and Masters degrees in Chemistry and Microbiology from the University of Poona, India. While an undergraduate, she became interested in academic research and earned her PhD in Biochemistry in 1997 from the University of Calcutta, India. Her postdoctoral training focused on studying cell-signaling mechanisms and protein-protein interactions between the heterotrimeric G proteins and their downstream signaling targets at the University of Rochester, NY, under Dr. Alan Smrcka. She then joined The Miami Project in 2007 as a postdoctoral associate with Dr. Damien Pearse, where she began developing methods for altering glial cell reactivity and cell-to-cell interactions in response to central nervous system (CNS) injury or disease to promote neuroprotection and repair. During this time Dr. Ghosh investigated 1) the surface modification of Schwann cells to enhance their capacity for migration and axon regeneration following transplantation within the injured spinal cord, 2) microglia phenotypic conversion to alter their properties from inflammatory to reparative, and 3) the manipulation of intracellular signaling pathways in astrocytes to abolish their reactivity and production of axon growth inhibitory matrix molecules. Dr. Ghosh's research in spinal cord injury repair was recognized by The Sam Schmidt Paralysis Foundation and The American Spinal Injury Association through The Outstanding Young Investigator Award given to her in 2009.

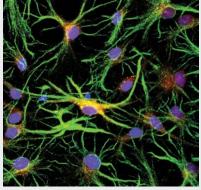
The present focus of Dr. Ghosh's laboratory is on altering the hostile environment of the injured or diseased CNS to one that is conducive to repair through conversion of microglia and macrophages from a cytotoxic M1 ("bad") to a reparative M2 ("good") state. By understanding the signals that govern macrophage-microglia phenotype, she aims to develop novel, molecular, and immuno-pharmacological therapeutic strategies to promote M1 to M2 conversion of these cells to induce neuroprotection, neuroplasticity, and/or disease remission. Her laboratory is also interested in understanding how altering the immunophenotypical profile of macrophages and microglia can affect host glial responses, including the formation of the glial scar and oligodendrocyte migration and myelination as well as influence the ability of transplanted cells, such as Schwann cells and stem cells, to mediate neurorepair.

#### **M1-M2 MACROPHAGES** AND MICROGLIA



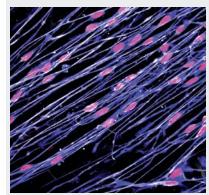
Understanding the signaling pathways involved in classical and alternative activation of microglia/ macrophages

#### **HOST GLIAL CELL RESPONSES**



Inducing an environment for repair in the injured/diseased CNS by immunomodulation

#### **CELL THERAPIES**

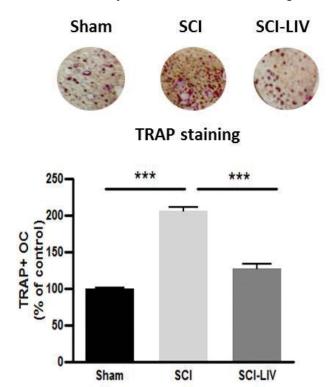


Altering host immune responses to improve the neurorepair provided by cell therapies

## Can Whole Body Vibration Influence Bone Health After (Spinal Cord Injury) SCI?

As many of our readers know, SCI causes many more problems aside from not being able to move. One big problem is bone density loss, which can lead to osteoporosis. Putting weight on bones (weight bearing) and having muscle contractions are two important components of maintaining healthy bones. Healthy bone metabolism is a balance between old bone degradation (resorption) and new bone formation. SCI alters bone metabolism; there is an increase in bone resorption, but a decrease in new bone formation. This leads to an overall loss of bone density.

Vibration is a component of weight bearing and whole body vibration has been investigated by many research groups as a therapeutic intervention for osteoporosis, especially in post-menopausal women. The potential benefits of whole body vibration have been investigated



Examples of osteoclasts (top) with LIV reducing the osteoclastogenesis (part of the bone resportion process) triggered by SCI (graph)

for other body systems as well. In fact, in our 2010 winter edition of the *Research Review* we wrote an article about the effect of whole body vibration on muscle spasticity in people with motor incomplete SCI. Seeing those beneficial results promoted an interest in going "back to the bench" to evaluate how/if vibration alters bone metabolism after SCI.

Drs. Helen Bramlett and Dalton Dietrich, along with colleagues in New York, tested the effect of low-intensity, high-frequency mechanical vibration (LIV) in rodents that had a moderate mid-thoracic spinal contusion (Bramlett et al., 2014, *Osteoporos. Int.*). They waited for 28 days before starting LIV for a couple reasons: 1) to mimic the clinical situation of several weeks passing after SCI before rehabilitation begins and 2) animals with this type of injury begin to regain weight-bearing ability by that time and it was hypothesized that weight bearing might enhance any benefits due to LIV. LIV was given 2x/day (15 minutes/ session), 5 days/week for 35 days. The vibration frequency delivered was in the 40 Hz range, which was similar to what Ness and Field-Fote (2009, *Gait & Posture*) did in humans with SCI.

They found that LIV administered at that time point, dose, and duration did not alter gross bone mineral density or trabecular bone architecture in that experimental model of SCI. However, they did identify some significant changes in bone metabolism. Osteoblasts are the cells associated with bone formation and the chemical osteocalcin is secreted only by osteoblasts, so osteocalcin is a biomarker of bone formation. The SCI itself resulted in decreased osteocalcin biomarker levels in the blood, which was increased back to normal levels by LIV. This suggests that LIV has a positive effect on new bone formation.

Additionally, there appeared to be an effect on certain aspects of bone resorption. Osteoclastogenesis is the development of osteoclasts and osteoclasts are the cells associated with bone resorption. In this experiment, the SCI resulted in increased osteoclastogenic potential of bone marrow precursor cells and LIV reduced this potential by 70%. This suggests that LIV may be reducing some components of the increased bone resorption induced by SCI.

Though bone metabolism is a complicated process, just like everything else with SCI, it appears that LIV is worth pursuing more as a therapeutic intervention for bone health after SCI!

...we support the idea that a combination of interventions is required to promote the most repair and functional improvement in people.

# **Engineered** Schwann Cell Grafts

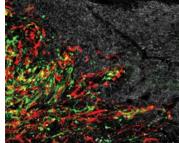
Tany years of animal research and many scientists have Leader demonstrated that Schwann cells transplanted into the spinal cord injury (SCI) site can create an environment that promotes various activities important for repair. Indeed we now have two Phase I clinical trials testing the safety of transplanting Schwann cells by themselves. However, we also know that Schwann cells in combination with other treatments can promote more repair – remember, the spinal cord is very complex and we support the idea that a combination of interventions is required to promote the most repair and functional improvement in people.

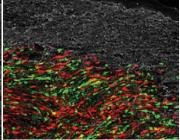
Drs. Mary Bunge, Damien Pearse, and colleagues recently published new results from a very exciting combination strategy (Journal of Neuroscience, 2014, 34(5):1838 –1855). The "players" in the combination strategy were:

- 1. rat Schwann cells,
- 2. D15A this is a growth factor that has been modified to act as two growth factors (Neurotrophin-3 and Brain-derived neurotrophic factor), and
- 3. Chondroitinase ABC (ChABC) an enzyme that can break apart many of the proteins that are found in scar tissue surrounding the SCI site.

In addition to multiple experimental controls, they transplanted the following combinations one week after a mid-thoracic contusion injury in rodents:

- 1. Schwann cells that secrete D15A only
- 2. Schwann cells that secrete ChABC only
- 3. Schwann cells that secrete BOTH D15A and **ChABC**





Graft:host interface. Schwann cells engineered to secrete D15A and ChABC (left) migrate into the caudal host spinal cord tissue better than control Schwann cells (right).

They followed the animals for 3 months and then analyzed many different aspects of repair and function. They found that the transplanted Schwann cells that had been engineered to secrete D15A and ChABC at the same time had the best effect. The main results from that combination showed:

- 1. the greatest number of Schwann cells surviving in the graft site as well as the greatest number of axons that were myelinated (insulated) by Schwann cells,
- 2. more nerve fibers from the brainstem grew into the graft as well as the host tissue around and below the graft,
- 3. more nerve fibers from the cortex of the brain remained closer to the graft (did not die back) and grew/sprouted around and below the graft,
- 4. a significant improvement in aspects of locomotion, and
- a reduction in pain-like sensations.

These results are very exciting because they show that Schwann cell grafts engineered to secrete a multifunctioning growth factor in addition to altering the scar composition lead to more axonal regeneration and functional improvement than either strategy administered alone. These are the kind of animal experiments that will help guide us as we move forward into testing combination treatments in future clinical trials.



If you would like to be considered for these or future Miami Project trials or studies, please see call The Miami Project Education Office at 305-243-7108.

## **Biological Trials**

Hypothermia

Schwann Cells

Riluzole

**HuCNS-SC** Stem Cells

Phase I Phase I Chronic Acute

Phase I Acute

Phase II Acute

Phase II Chronic

T3-T11 AIS A

C5-T12 AIS A, B, C

C1-C8 AIS A, B

C4-C8 AIS A, B, C

C5-C7 AIS A, B, C



**Online Surveys** 

- Pain After SCI
  - www.surveymonkey.com/s/SCIpain
- Basic Pain Dataset Self-Report www.surveymonkey.com/s/paindatast

**Activity and Nutrient Modifying** 

- Cardiometabolic Risk, Obesity, and Cardiovascular Disease in SCI
- Effect of an Omega-3 Supplement Intervention on Cardiometabolic Health
- Obesity/Overweight in Early and Chronic SCI: Lifestyle Intervention Program
- Clinically Meaningful Changes in Wheelchair Propulsion Stress
- Training Programs to Improve Outcomes for Individuals with SCI

**Male Fertility** 

- Fertility Evaluation
- Treatment for Infertility

**Spasticity** 

Quantifying Spasticity in Activities of Daily Living

**Aging** 

- Telomeres as an Aging and Health Biomarker in Persons with SCI
- Muscle Weakness and Fatigue with Age After SCI
- Impact of Time Post-Injury on the Bladder Inflammatory Profile

**Brain Machine** Interface

- Treatment for Pain and Autonomic Dysreflexia in SCI with Deep Brain
- Assessment of Candidates and Design Considerations for Neuroprosthetic Devices for Chronic SCI











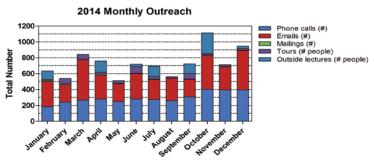
The Education Department: (clockwise from upper right) Kathleen Nagle, Kim Anderson-Erisman, PhD (Director), Letitia Fisher, Maria Chagoyen, and Danielle Cilien.

### The Miami Project believes that an important component of developing treatments for paralysis involves communication with the community.

The Education department, directed by Kim Anderson-L Erisman, Ph.D., is responsible for helping thousands of our community members each year. The other valuable members of the department are Maria Chagoyen, Danielle Cilien, Letitia Fisher, and Kathleen Gilsenan. Each year the department answers thousands of phone calls and emails to provide people with information about all of our research programs and clinical studies as well as provide information about rehabilitation, clinical care referral, resources for living with paralysis, and advice about research from around the world. We also conduct numerous tours and



Images from the 2014 Community Open House and 2014 Brain Fair "Build-a-Spinal Cord" exhibit.



lecturers about our research. The graph shows the total number of people reached each month during 2014 outreach activities. The Education department also assists all of The Miami Project clinical research faculty with recruitment for their clinical studies. To participate in research studies individuals must first complete an Intake form, which provides us with preliminary injury characteristics. Then you receive a phone call from us to discuss the studies that you pre-qualify for and determine whether you are interested in proceeding with any studies. If so, we set up an appointment for you to come to our research center for a neurologic exam ("ASIA") and introduction to the laboratories.

On April 5, 2014 the Education department hosted the 4th Annual Miami Project Community Open House. We enjoy this opportunity to open up our doors to the public to answer questions and share information, as well as to hear direct input from our community. The afternoon began with an interactive session about *Health & Wellness* related to spinal cord injury (featuring Drs. Mark Nash, Rachel Cowan, Christine Thomas, Eva Widerström-Noga, and Diana Cardenas). This included cardiovascular issues in relation to diet and exercise, in addition to chronic inflammation and sleep disruptions. Mobility was also discussed, particularly how it is impacted by strength, the environment, and wheelchair



configuration, with suggestions on how each of these factors might be manipulated to enhance mobility. Chronic pain is a significant problem to many people living with SCI and input was provided about the bio-psycho-social aspects of pain and their interactions.

This was followed by a *Meet the Scientists* session in which the audience met 5 of our faculty (Drs. Damien Pearse, Paula Monje, Pantelis Tsoulfas, Vance Lemmon, and Brian Noga), heard brief explanations of their current research, and had the opportunity to ask questions. The final component of the day was Behind-the-Scenes tours of four laboratories, including a cell culture lab, regeneration and imaging lab, human locomotor and functional electrical stimulation lab, and human sensory lab. The 5th Annual Community Open House is scheduled for April of 2015.

The education department also participated in the 5<sup>th</sup> Annual Brain Fair with a Build-a-Spinal Cord exhibit/ activity and gave invited lectures to multiple SCI consumer groups. If you have questions, don't hesitate to email us at mpinfo@med.miami.edu or call us at 305-243-7108.

## Gail F. Beach Memorial Lecture Series

The Miami Project has brought many renowned neuroscientists from around the world to our campus as part of The Gail F. Beach Memorial Visiting Lectureship Series. The lectureship series is dedicated to Gail F. Beach, a schoolteacher and person with SCI,

whose generosity and foresight provides outstanding educational opportunities for The Miami Project researchers and our neuroscience colleagues at the University of Miami.



Gail F. Beach Memorial Lecturer Vittorio Gallo, PhD, with Faculty member Roberta Brambilla, PhD

November 5, 2014 Vittorio Gallo, PhD

Children's National Medical Center Washington, DC

December 3, 2014

#### Pramod Dash, PhD

University of Texas Medical School at Houston Houston, TX

January 7, 2015

#### Susan Harkema, PhD

University of Louisville - Louisville, KY

February 4, 2015

#### **Armin Curt, MD, FRCPC**

University of Zurich - Zurich, Switzerland

March 4, 2015

#### V. Reggie Edgerton, PhD

University of California, Los Angeles Los Angeles, CA

April 1, 2015

#### Larry M. Jordan, PhD

University of Manitoba - Winnipeg, Canada

May 6, 2015

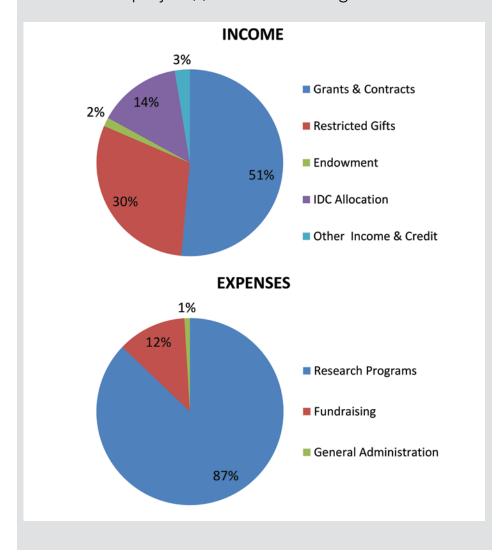
#### T. George Hornby, PhD, PT

University of Illinois at Chicago – Chicago, IL

Each year, Miami Project scientists seek funding for their research by submitting proposals to the National Institutes of Health, the premier scientific body in the United States, as well as other funding agencies and foundations.



Their scientific peers rate the merits of these proposed experiments in a highly competitive process and only the best are funded. The agencies and organizations listed below supported the Principal Investigator(s) and the scientific project(s) indicated during 2014.



#### **American Heart Association Scientific Development Grant**

Dr. Juan Pablo De Rivero Vaccari (P.I.)

-Activation of Rig-like Receptor Signaling after Focal Cerebral Ischemia

#### **Bryon Reisch Paralysis Foundation**

Dr. Kim Anderson-Erisman (P.I.) -The Safety of Autologous Human Schwann Cells – Screening

#### **Children's Tumor Foundation**

Dr. Cristina Fernandez-Valle (P.I.), Dr. Paula Monje (Co-I.)

-Creation of Human Merlin-Null Schwann cells for NF2 Studies

#### **Christopher & Dana Reeve Foundation**

Dr. James Guest (Center P.I.) -North American Clinical Trials Network

-CTN7-2014: Safety and Pharmacokinetics of Riluzole in **Patients** 

#### **Craig H. Neilsen Foundation**

Dr. Kim Anderson-Erisman (P.I.) -Miami Project Education Program

Dr. Nancy Brackett (P.I.), Dr. Juan Pablo De Rivero Vaccari (Co-I.) -Improving Reproductive Function in Men with Spinal Cord Injury

Dr. Robert Keane (P.I.), Dr. Juan Pablo De Rivero Vaccari (Co-I.) -Inflammasome Regulation Following Spinal Cord Injury

Dr. Daniel Liebl (P.I.) -A Novel Eph Receptor-Mediated Mechanism of Cell Death Following Spinal Cord Injury

Dr. Brian Noga (P.I.), Dr. Ian Hentall

-Acute Facilitation of Walking After SCI Using Deep Brain Stimulation

Dr. Kevin Park (P.I.), Dr. Jae Lee (Co-I.) -Novel Combinatorial Approaches to Promote Axon Regrowth After Chronic SCI

Dr. Urs Rutishauser (P.I.), Dr. Damien Pearse (Co-I.) -Optimization of PST-Engineered Schwann Cells for SCI Repair

Dr. Jacqueline Sagen (P.I.) -Utilizing Designer Genes to Alleviate Chronic SCI Pain

Dr. Christine Thomas (P.I.) -Weakness and Fatigue with Age after Spinal Cord Injury

#### Department of Defense (DoD) **Spinal Cord Injury Research Program of the Office of the Congressionally Directed Medical Research Programs**

Dr. Rachel Cowan (P.I.) -Fitness and Independence After SCI: Defining meaningful Change and Thresholds

Dr. W. Dalton Dietrich (P.I.), Dr. Michael Wang (Partner P.I.) -Biomarkers for Spinal Cord Injury-**Related Medical Complications** 

Dr. W. Dalton Dietrich (P.I.), Dr. John Bixby (Co-I.), Dr. Helen Bramlett (Co-I.), Dr. Jae Lee (Co-I.), Dr. Vance Lemmon (Co-I.), Dr. Daniel Liebl (Co-I.), Dr. Kevin Park (Co-I.), Dr. Pantelis Tsoulfas (Co-I.) -Battlefield Exercise and Combat Related Spinal Cord Injury

Dr. Jonathan Jagid, (P.I.), Dr. Ian Hentall (Co-I.), Dr. Alberto Martinez-Arizala (Co-I.), Dr. Eva Widerström-Noga (Co-I.)

-Treatment of Pain and Autonomic Dysreflexia in Spinal Cord Injury with Deep Brain Stimulation

Dr. Mark Nash (P.I.)

-Obesity/Overweight in Persons with Early and Chronic SCI: A Randomized, Multicenter, Controlled Lifestyle Intervention

Dr. Kevin K. Park (P.I.) -Novel Combinatory Approaches to Repair Visual System After Optic Nerve Damage

Dr. Damien Pearse (P.I.), Dr. Mary Bartlett Bunge (Partner P.I.), Dr. James Guest (Partner P.I.), Dr. Dalton Dietrich (Co-I.) -Schwann Cell (SC) Implantation for SCI Repair: Optimization of Dosing, Long-Term Cell Persistence, and the Evaluation of Toxicity and Tumorigenicity

Dr. Damien Pearse (P.I.), Dr. Howard Levene (Partner P.I.) -Translation of Novel PDE4 Inhibitors for the Treatment of Acute Spinal Cord Injury

Dr. Shirin Shafazand (P.I.), Dr. Mark S. Nash (Co-P.I.) -Neuro-cognitive Decline and Sleep-Disordered Breathing After SCI

Dr. Eva Widerström-Noga (P.I.), Dr. Kim Anderson-Erisman (Co-I.), Dr. Alberto Martinez-Arizala (Co-I.) -Experiences of Living with Persistent Pain After a Spinal Cord Injury

#### Department of Defense (DoD) **Psychological Health and Traumatic Brain Injury Program of the Office** of Congressionally Directed Medical **Research Programs**

Dr. W. Dalton Dietrich (P.I.), Dr. Helen Bramlett (Co-I.) -Operation Brain Trauma Therapy

-The Importance of Temperature in the Pathophysiology of Mild Repetitive **Brain Injury** 

-The Use of Pro-Neurogenic

Molecules to Promote Recovery of Function Following Acute and Chronic Traumatic Brain Injury

Dr. Jed Hartings (P.I.), Dr. M. Ross Bullock (Co-I.) -Spreading Depressions as Secondary Insults after Traumatic Injury to the Human Brain

#### **Department of Defense US Army Medical Research and Materiel Command (DAMD)**

Dr. M. Ross Bullock (P.I.) -Clinical Phase IIB Trial of Oxycyte Perfluorocarbon in Severe Human Traumatic Brain Injury -Laboratory studies evaluating PFC in models of Penetrating and Closed TBI

#### E. Matilda Ziegler Foundation for the Blind

Dr. Kevin Park (P.I.) -Novel Combinatorial Approaches to Enhance Retinal Ganglion Cell Survival and Axon Regeneration after Optic Nerve Injury

#### FISM Fondazione Italiana Sclerosi Multipla (Italian Multiple Sclerosis Foundation)

Dr. Roberta Brambilla (P.I.) -The Pro-Remyelination Effect of Transmembrane Tumor Necrosis Factor: Investigation Into the Role of Tumor Necrosis Factor Receptor 2

#### Florida Department of **Transportation**

Dr. Gillian Hotz (P.I.) -WalkSafe/Bikesafe Project, Social Marketing Campaign, and School-Specific Action Plan

#### **International Spinal Research Trust**

Dr. James Guest (Center P.I.) -Cellular Trials to Support Corticospinal Axon Regeneration in Non-Human Primates

#### **Medtronic Spinal and Biologics**

Dr. Allan Levi (Site-P.I.), Dr. Barth Green (Co-I.)

-Study of the Bryan™ Cervical Disc Prosthesis in the Treatment of Degenerative Disc Disease – Pivotal IDE Study

#### **National Eye Institute**

Dr. Kevin K. Park (P.I.)
-Regeneration and Reconnection of Damaged Optic Nerve

## **National Football League Charities Medical Research**

Dr. Coleen Atkins (P.I.)
-The Effects of Mild Hyperthermia on
Outcome after Mild Traumatic Brain
Injury

## National Heart, Lung, and Blood Institute

Dr. Stephan Schürer (P.I.), Dr. Vance Lemmon (Co-I.)

-LINCS Information FramEwork (LIFE) to integrate and Analyze Diverse Data Sets

#### National Human Genome Research Institute

Dr. Vance Lemmon (P.I.)
-Bioassay Ontology and Software
Tools to Integrate and Analyze Diverse
Data Sets

## National Institute of Child Health and Human Development

Dr. Vance Lemmon (Co-P.I.), Dr. John Bixby (Co-P.I.)

-Novel Gene Targets for CNS Axonal Regeneration

## National Institute of Disability & Rehabilitation Research

Dr. Diana Cardenas (P.I.), Dr. Rachel Cowan (Co-I.), Dr. Mark Nash (Co-I.) -South Florida Spinal Cord Injury Model Systems

Dr. Michael Boninger (P.I.), Dr. Kevin Dalal (Site P.I., Miami), Dr. Rachel

Cowan (Site Co-P.I., Miami)
-Collaboration on Mobility Training

Dr. Mark Nash (P.I.), Dr. Kim Anderson-Erisman (Co-I.) -Sleep Disordered Breathing in Persons with Chronic Tetraplegia: Characterization and Intervention

Dr. Suzanne Groah (P.I), Dr. Mark Nash (Co-I.)

-Rehabilitation Research and Training Center on Secondary Conditions in SCI

## National Institute of Neurological Disorders & Stroke

Dr. Kim Anderson-Erisman (Co-P.I.), Dr. W. Dalton Dietrich (Co-P.I.) -NIH Neurotrauma Summer Research Experience Program

Dr. Coleen Atkins (P.I.)

-Rehabilitation Strategies for Memory Dysfunction after Traumatic Brain Injury

-The Role of Phosphodiesterase 4B in Inflammation after Trauma (Fellowship)

Dr. Mary Bartlett Bunge (Co-P.I.), Dr. John Bethea (Co-P.I.), Dr. Ian Hentall (Co-I.), Dr. Paula Monje (Co-I.), Dr. Kevin Park (Co-I.), Dr. Patrick Wood (Co-I.)

-Cytological Studies of Developing and Mature Neurons

Dr. John Bixby (P.I.), Dr. Vance Lemmon (Co-I.)

-Novel Compounds that Overcome Glial Inhibition of Axonal Regeneration

Dr. W. Dalton Dietrich (P.I.), Dr. Helen Bramlett (Co-I.), Dr. Daniel Liebl (Co-I.), Dr. Pantelis Tsoulfas (Co-I.) -Mechanisms of Recovery Following Traumatic Brain Injury Dr. W. Dalton Dietrich (P.I.), Dr. Helen Bramlett (Co-I.), Dr. Coleen Atkins (Co-I.)

-The Importance of Temperature on Inflammation after TBI -Cyclic Nucleotide Regulation in

Traumatic Brain Injury

Dr. W. Dalton Dietrich (P.I.), Dr. Robert Keane (Co-P.I.)

-Therapeutic Neutralization of the Inflammasome after Spinal Cord Injury

Dr. Gillian Hotz (Site-P.I.), Dr. M. Ross Bullock (Site Co-P.I)

-Transforming Research and Clinical Knowledge in Traumatic Brain Injury

Dr. Jae Lee (P.I.)

-Role of Fibroblasts in Axon Regeneration After SCI -Translational Profile of Perivascular Fibroblasts After Spinal Cord Injury -RAP as a Therapeutic Compound for Neuronal Regeneration After Spinal Cord Injury (Sub-contract)

Dr. Vance Lemmon (P.I.), Dr. John Bixby (P.I.), Dr. Stephan Schürer (P.I.) -Regenbase: A Searchable Database to Organize Regeneration Knowledge via Ontologies

Dr. Daniel Liebl (P.I.)

-Ephrins Regulate Stem Cell Proliferation following Traumatic Brain Injury

-Eph Receptors Regulate Vascular Growth Following Traumatic Brain Injury (Fellowship)

-Modulating Post-injury Gliotransmitter Levels leads to improved Synaptic Function (Fellowship)

Dr. Paula Monje (P.I.)

-Phenotypic and Functional Analysis of Human Schwann Cells for Potency Assay Development Dr. Damien Pearse (P.I.)

-Therapeutic Targeting of Intracellular Mechanisms Involved in Glial Scar Formation

Dr. Grace Zhai (P.I.), Dr. Pantelis Tsoulfas (Co-I.) -Mechanisms of Neuronal

Maintenance and Protection

#### Robert J. Kleberg, Jr., and Helen C. **Kleberg Foundation**

Dr. Kim Anderson-Erisman (P.I.), Dr. Rachel Cowan (Co-I.), Dr. Edelle Field-Fote (Co-I.), Dr. Mark Nash (Co-I.), Dr. Christine Thomas (Co-I.), Dr. Eva Widerström-Noga (Co-I.) -Exercise and Locomotor Training Required for Clinical Trials Targeting Chronic Spinal Cord Injury

#### **Paralyzed Veterans of America Foundation**

Dr. Mousumi Ghosh (P.I.), Dr. Damien Pearse (Co-I.) -Down-Regulating PDE4A in Astrocytes to Promote Axon Regeneration After SCI

#### **Sheila and David Fuente Neuropathic Pain Program**

Dr. Jacqueline Sagen (P.I.) -Development of a Phantom Limb Pain Model for Novel Therapeutic Interventions

#### State of Florida Brain and Spinal Cord Injury Program, Department of Health, and Red Light Camera Fund

-These three state funds contribute to several research programs within The Miami Project to Cure Paralysis

#### StemCells Inc.

Dr. Allan Levi (Site-P.I.)

-A Single-Blind, Randomized, Parallel Arm, Phase II Proof-of-Concept Study of the Safety and Efficacy of HUCNS-SC Transplantation in Cervical Spinal Cord Injury

#### **The Pew Charitable Trusts**

Dr. Kevin Park (P.I.)

-Visual System Repair Following Optic Nerve Damage

#### **University of Miami SAC Awards**

Dr. Robert Keane (P.I.), Dr. Juan Pablo De Rivero Vaccari (Co-I.)

-Therapeutic Neutralization of the Inflammasome after Spinal Cord Iniurv

#### **US Army Medical Research and Material Command**

Dr. Treena Arinzeh (P.I.), Dr. Mary Bartlett Bunge (Sub-site P.I.)

-A Combination Tissue Engineering Strategy for Schwann Cell-Induced Spinal Cord Repair

#### **Veterans Administration Biological Laboratory Research and Development**

Dr. Helen Bramlett (P.I.)

-Novel Treatment Strategies for Targeting Post-Traumatic Epilepsy

#### **Veterans Administration** Rehabilitation Research and **Development**

Dr. Victor Arvanian (P.I.), Dr. Damien Pearse (Co-I.)

-Enhancing Plasticity in a Damaged Spinal Cord to Repair Transmission and Function

Dr. Robert Jackson (P.I.), Dr. Diana Cardenas (Co-I.)

-Rehabilitation of IPF Patients: Effects of Exercise and Oxidant Stress

Dr. Damien Pearse (P.I.)

-Enhancing the Reparative Efficacy of Schwann Cells Following Chronic SCI









# The Miami Project To Cure Paralysis **Faculty Profiles**

The faculty of The Miami Project are a talented multidisciplinary team. In the following Profiles, each faculty member describes their specific research focus and highlights of recent progress.



W. DALTON DIETRICH, III, PH.D. Scientific Director, The Miami Project **Kinetic Concepts Distinguished Chair in Neurosurgery** Senior Associate Dean for Discovery Science Professor, Departments of Neurological Surgery, Neurology, and Cell Biology & Anatomy

Neuroprotection and Improved Recovery of Function following CNS Trauma My research interest is the pathobiology and treatment of CNS injury in both the acute and chronic setting. Animal models of cerebral ischemia, and brain and spinal cord trauma are utilized to investigate the mechanisms of tissue injury. The ultimate goal is to target specific injury processes for pharmacological intervention, including the addition of growth factors, to promote circuit plasticity, regeneration and recovery of function.



BARTH A. GREEN, M.D., F.A.C.S. Co-Founder, The Miami Project Professor and Chairman, Department of Neurological Surgery Translational Interventions

Over the recent years my research efforts have mainly involved taking the cutting edge basic neuroscience work product and data created by our Miami Project team from the bench to our UM affiliated clinics and hospitals. A good example of such translational research efforts has included the use of modest hypothermia for neuroprotection both in cases of acute spinal cord injury and for use in the operating room for patients undergoing high risk spinal cord surgery. I am also privileged to be able to collaborate with The Miami Project cellular transplantation programs and have been working on projects involving adult mesenchymal stem cells as well as being part of the major effort transforming our successful Schwann cell laboratory model into clinical trials. Another area of clinical interest and research includes the diagnosis and treatment of Chiari I malformation with and without syringomyelia.



MARY BARTLETT BUNGE, PH.D. Christine E. Lynn Distinguished Professor in Neuroscience Professor, Departments of Cell Biology, Neurological Surgery, and Neurology Development of Combination Strategies with Schwann Cells to Repair the Injured Spinal Cord

The goal in my laboratory is to foster regeneration of axons across and beyond a spinal cord injury (SCI). To improve regeneration of axons, we are investigating reducing the accumulation of proteoglycans (molecules that inhibit axonal growth), improving survival of transplanted Schwann cells (SCs), and genetically engineering SCs before transplantation to improve their growth factor-secretion capability or neurons to enhance their ability to regrow axons, and testing matrices for efficacy after injury. We pay particular attention to the interface between the SC implant and the host spinal cord.



JOHN BIXBY, PH.D.

Professor, Departments of Molecular & Cellular Pharmacology and Neurological Surgery, Center for Computational Sciences, Institute for Human Genomics, Sylvester Cancer **Center, Vice Provost for Research** 





High Content Screening and Functional Genomics of the Nervous System

Our laboratory has developed methods to test thousands of genes or chemicals in hundreds of thousands of neurons each week and obtain quantitative information about cell morphology and gene expression. This "high throughput" capability allows us to tackle questions about development and regeneration using systems biology approaches. The Lemmon-Bixby lab has several ongoing projects related to axon regeneration. One project is to test the roles of known signaling proteins called protein kinases. In this screen we have tested hundreds of kinases by overexpression and have also tested >1500 kinase inhibitors, many of which strongly promote neurite growth in vitro. Using machine learning and cheminformatics (collaboration with Dr. Stephan Schürer) we can identify critical kinases and their signaling networks as well as potential lead therapeutic compounds. A second project is based on the observation that peripheral sensory neurons initiate a genetic program appropriate for axonal regeneration after injury. Our laboratory is combining next-generation sequencing with cell-based phenotypic screening to identify genes and microRNAs that are part of this genetic program. Finally, in collaboration with Dr. S. Schürer and Dr. Ubbo Visser, we are developing RegenBase, an information system that integrates diverse data on nerve regeneration after spinal cord injury with data from other information resources.



M. ROSS BULLOCK, M.D., PH.D. **Professor, Department of Neurological Surgery Director, Clinical Neurotrauma** 

#### Preclinical Mechanistic and Neuroprotection Research in Traumatic Brain Injury and Clinical Trials, and Neuromonitoring Techniques in the Injured Brain

We are near completion of a two year grant from the Department of Defense to evaluate the neuroprotective effect of Perfluorocarbons in four rodent models of traumatic brain injury (penetrating brain injury, closed traumatic brain injury with secondary hypoxia, tissue culture with stretch injury, and mechanistic and safety studies). These oxygen carriers have shown benefit in previous studies involving fluid percussion injury and subdural hematoma models. We are also evaluating hypothermia neuroprotection, in humans and animals, using novel biomarkers. We are also obtaining pilot data with FDA approved Human stem cells, transplanted into the rat brain, as therapy for Penetrating TBI.

#### DIANA CARDENAS, M.D., M.H.A. Professor and Chair, Department of Rehabilitation Medicine Chief of Service & Medical Director, Department of Rehabilitation Medicine

**Pain Interventions and Prevention of Urinary Tract Infections** 

The goals of my research are to help find therapeutic interventions that improve recovery, reduce secondary conditions, and create a better life for persons with SCI and other conditions that impair physical or cognitive function. Currently, my research focus is in the areas of neuropathic pain and neurogenic bladder management.





ROBERT W. KEANE, PH.D. Professor, Departments of Physiology & Biophysics, and Neurological Surgery **Regulation of Innate Immunity after CNS Trauma** 

Innate immunity is the first line of defense against pathogens and host-derived signals of cellular stress. My research focuses on investigating mechanisms that direct normal innate immunity and its dysregulation in central nervous system injury and disease, including (1) agonists and activation mechanisms of inflammasomes, (2) regulatory mechanisms that potentiate or limit inflammasome activation after injury, and (3) emerging data linking inflammasome proteins as biomarkers for CNS injury.

#### ALLAN D. LEVI, M.D., PH.D., F.A.C.S.

Professor, Departments of Neurological Surgery, Orthopedics, and Rehabilitation Medicine Chief of Neurospine Service, Jackson Memorial Hospital/Chief of Neurosurgery, University of Miami Hospital

Cellular Transplantation Strategies after SCI/Systemic Hypothermia after Acute SCI

My clinical research interests currently focus on developing cellular transplantation strategies to repair injuries within both the human central and peripheral nervous system. I am currently Co-PI on our clinical trial "Transplantation of Autologous Human Schwann Cells (SCs) to Repair the Injured Spinal Cord - Phase I - safety study". This represents a first-in-man dose escalation study of autologous human SCs for patients with sub-acute thoracic SCI (T3 to T11). We are also very interested in the use of SCs for peripheral nerve injuries with long segmental defects and have



performed such a transplantation in a patient with a significant acute sciatic nerve injury. Hypothermia continues to show promise in a variety of acute central nervous system injuries. There are various factors that need to be considered with systemic cooling of the SCI patient, including methods of cooling, window from injury to initiation, duration and depth of hypothermia, rate of re-warming, etc. While profound levels of hypothermia (T <32°C) can be difficult to administer and are subject to increased complication rates, mild (modest) levels of hypothermia (T 32-34°C) have been shown to provide significant protection against traumatic and ischemic neuronal cell death. I am currently the PI of our institutional protocol studying systemic hypothermia induced via an intravascular catheter and continued for 48 hours after acute cervical SCI.



DANIEL J. LIEBL, PH.D. **Professor, Department of Neurological Surgery** 

Molecular Mechanisms that Regulate Cellular Dysfunction and Death Following CNS Injury, and Mechanisms to Promote Regeneration and Recovery

The goal of my laboratory is to identify the mechanisms that lead to CNS pathophysiology and its regenerative potential. We focus on a family of molecules, called eprhins and Eph receptors, which play important roles in the developing, regenerating, and injured nervous systems. Specifically, we are currently interested in areas of adult neurogenesis, neuroprotection, apoptotic cell death, synaptic plasticity, regeneration, and therapeutic strategies. Overall, our approach is to develop novel strategies to minimize CNS damage and maximize regeneration/tissue repair, which can be best achieved through a comprehensive mechanistic approach.

#### MARK S. NASH, PH.D., F.A.C.S.M. Professor, Departments of Neurological Surgery, Rehabilitation Medicine, and Kinesiology & Sports Sciences

Physiological Assessment of Secondary Complications following SCI: Electrical Stimulation, Cardiometabolic and Vascular Pathophysiology, Cardioendocrine Pathology, and Exercise and Dietary Biochemistry

One of the enduring goals of The Miami Project has been to test and then translate strategies that optimize health of persons with SCI. A significant target for this strategy has focused on physical activity to lessen secondary risks of SCI associated with physical deconditioning and an accelerated trajectory of cardioendocrine disease. We also examine complementary themes to validate exercise prescription after SCI, identify optimal dietary composition and causes for overeating, and use of prescription and non-prescription agents that reduce hazards of fasting and postprandial lipid disorders, dysglycemia, and vascular inflammatory stress.





DAMIEN D. PEARSE, PH.D. **Professor, Department of Neurological Surgery** 

#### **Exploration and Translation of Therapeutic Strategies to Repair the Injured Spinal Cord** and Brain

My laboratory focuses on several key aspects of CNS injury repair, including (1) the utility and clinical translation of exogenous and endogenously harnessed cell therapeutics (particularly when used in combinatory approaches), (2) understanding the role of, and developing therapies for, altered cyclic AMP (adenylyl cyclase, phosphodiesterases, and PKA) and MAPK signaling in neurons and glia after CNS injury, (3) the use of nanotherapeutics for multifunctional and sitedirected gene/drug targeting to the injured CNS, and (4) the application of methodologies for improved imaging of axonal regeneration and cell integration within the injured CNS such as 3D ultramicroscopy and diffusion tensor imaging.



JACQUELINE SAGEN, PH.D., M.B.A. **Professor, Department of Neurological Surgery** 

#### Cellular Implants for the Alleviation of Chronic Pain and CNS Injury

Our laboratory continues to explore novel and more effective strategies in the therapeutic management of chronic debilitating pain. Our recent research is focused on (1) identification of more effective analysesic agents and combinations for alleviating pain using SCI models and (2) development of emerging therapeutic interventions, including cell transplantation and gene therapy, which have the potential to provide long-term alleviation in people with intractable pain, overcoming the need for repeated pharmacologic administration.

THOMAS J. SICK, PH.D. Professor of Neurology and Physiology/Biophysics

#### Cellular and Neuronal Circuit Alterations after Traumatic Brian Injury That Contribute to **Cognitive Decline and Epilepsy**

My laboratory is conducting electrophysiological assessments of neuron and brain circuit alterations that occur after traumatic brain injury. Long-term clinical consequences of brain injury include declines in cognitive function and in many cases the development of epilepsy. We are trying to understand how circuits in the brain change over time after injury and how these changes might lead to alterations of brain function and behavior.



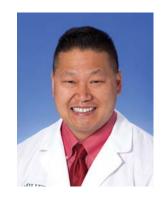


CHRISTINE K. THOMAS, PH.D. Professor, Department of Neurological Surgery Neuromuscular Weakness, Fatigue, Spasms, and Regeneration

Our laboratory is currently asking two main questions regarding SCI. First, in studies on people with SCI, we want to understand how well involuntary contractions of paralyzed muscles (spasms) are managed by exercise or by taking baclofen, a drug that is commonly used to control spasticity. Second, in our animal studies, we are exploring how to replace neurons that die because of SCI. Neuron death is common at the injury site and results in severe muscle weakness.

#### MICHAEL Y. WANG, M.D., F.A.C.S. Professor, Departments of Neurological Surgery and Rehabilitation Medicine **Spinal Cord Injury Outcomes**

My primary research has been in the investigation of SCI Outcomes. I work with Miami Project researchers Drs. Allan Levi and Barth Green in studying the clinical effects of Hypothermia. Currently, a multi-center randomized, prospective study on the effects of hypothermia in SCI is planned. In addition, I am studying the clinical application of SCI biomarkers to predict the effects of both injuries as well as therapeutic interventions with Drs. Dalton Dietrich and Ross Bullock.





NANCY L. BRACKETT, PH.D., H.C.L.D. Research Professor, Departments of Neurological Surgery and Urology Male Fertility following Spinal Cord Injury

Our research is focused on understanding and improving impairments to male fertility which occur following SCI. A major aim is to determine the cause of impaired semen quality in men with SCI. Our recent evidence indicates that the problem is related to the seminal plasma. Our current research is investigating inflammatory factors, including semen cytokine levels, as contributors to the problem. Our ultimate goal is to develop therapies to normalize semen quality in men with SCI, so that chances of biological fatherhood are increased.

#### JAMES D. GUEST, M.D., PH.D., F.A.C.S., F.R.C.S.(C) **Clinical Professor, Department of Neurological Surgery**

#### Augmented Recovery after SCI; Application of Therapeutic Combinations in Preclinical Studies, and Early Phase Clinical Trials

Recognizing that combined therapeutic approaches are needed to enhance recovery after traumatic SCI, we currently combine cell therapy with rehabilitation, and electrical stimulation. The lab group has members ranging from senior and junior medical faculty to postdoctoral students, medical, and undergraduate students. We use large animal models to increase the relevance of the preclinical testing to address key questions of efficacy and safety important to FDA. Therapeutics testing emulates human application as fully as possible; we use advanced histological, behavioral, electrophysiological, MRI, and ultrasound techniques. We design devices to deliver cells and therapeutics in a minimally injurious manner. Dr. Guest as also active in clinical trial design and execution through participation in the North American Clinical Trials Network and SCOPE (Spinal cord outcomes partnership endeavor).





GILLIAN A. HOTZ, PH.D. Research Professor, Department of Neurological Surgery **Director, KiDZ Neuroscience Center** Director, Concussion, WalkSafeTM & BikeSafeTMPrograms

As a behavioral neuroscientist my clinical interests have always been investigating the neurocognitive deficits of those individuals that have sustained a traumatic and acquired brain injury. I have co-authored two neurocognitive tests, The Brief Test of Head Injury for adults and the Pediatric Test of Brain Injury for children. My research has also focused on injury prevention, preventing brain and spinal cord injuries in children, and I have developed the WalkSafe program, which has been shown to decrease the number of elementary school age children that get hit by cars, and now the BikeSafe program to educate middle school age children on bicycle safety skills. As the Director of the Concussion Program we have a comprehensive countywide concussion care program including neurologic evaluation, neuroimaging, neuropharmacological management, and neuropsychological testing using ImPACT, a computerized neurocognitive screening measure.



ALBERTO MARTINEZ-ARIZALA, M.D. Clinical Professor, Departments of Neurology, Neurological Surgery, and Rehabilitation Medicine

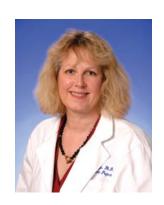
Chief, SCI Service Miami VA Medical Center

Pathophysiology and Treatment of Secondary Complications in Spinal Cord Injury My research interests focus on common complications that are seen following spinal cord injury: pain, spasticity, syringomyelia, and tethered cord syndrome. My interests include investigating the basis for the development of the different spasticity and pain profiles in the spinal cord injured population and to study potential novel treatments for those conditions.

EVA WIDERSTRÖM-NOGA, D.D.S., PH.D. Research Professor, Departments of Neurological Surgery, Rehabilitation Medicine, and **Health Scientist Veterans Affairs** 

#### **SCI-related Neuropathic Pain Phenotypes and Biomarkers**

My research program is focused on the identification of clinical correlates of underlying mechanisms of neuropathic pain associated with neurological trauma in order to facilitate the translation of basic research findings to treatments tailored to specific mechanisms. We are also examining the personal experiences of people living with chronic pain and SCI to increase our understanding about factors that help or hinder obtaining optimal pain relief. My research program is highly collaborative and includes extensive interdisciplinary protocols for a multimodal evaluation of self-reported pain symptoms and its psychosocial impact, quantitative assessment of neurological function, and biomarkers including non-invasive brain imaging and genetic polymorphism.





PATRICK M. WOOD, PH.D. Research Professor, Department of Neurological Surgery

Changes in the Molecular and Biological Properties of Human Schwann Cells Schwann cells have shown promise in animal studies in promoting recovery from SCI. We have developed protocols that allow the generation, from a small biopsy of human peripheral nerve, of large numbers of a person's own Schwann cells that can be transplanted back into their injured spinal cord. Efficient growth of human Schwann cells in culture requires the addition of recombinant neuregulin and the cAMP enhancer forskolin. To better understand the effects of these reagents on Schwann cells, we are performing basic research to determine the mechanisms by which neuregulin and cAMP enhancers promote interaction between axons and Schwann cells, including axon-induced proliferation and the formation of myelin sheaths.

NAGI AYAD, PH.D. Associate Professor, Department of Psychiatry and Behavioral Sciences Cell Cycle Transitions in the Developing and Diseased Nervous Systems

The research in my laboratory focuses on cell cycle transitions in the developing nervous system. My laboratory identified essential proteins required for controlling mitotic entry and exit and also demonstrated that cell cycle proteins are present and active in the developing nervous system and fully differentiated neurons. Specifically, the research group uncovered the mechanism through which the Anaphase Promoting Complex/Cyclosome (APC/C) and other cell cycle regulators such as Wee 1 control cell proliferation, cycle exit, and differentiation. Thus, my laboratory is searching for novel molecular pathways that control neural development and are targets in multiple human maladies.





HELEN M. BRAMLETT, PH.D. Associate Professor, Departments of Neurological Surgery and Psychology, Undergraduate Neuroscience Program Director, and Health Scientist Veterans Affairs

#### The Pathophysiology and Treatment of CNS Injury

The focus of my neurotrauma laboratory is to investigate both acute and long-term consequences of brain and spinal cord trauma. My current research interests are on the pathophysiology of traumatic brain and spinal cord injury with an emphasis on the pathogenesis of progressive white matter damage as well as the benefits of therapeutic hypothermia. My laboratory is also investigating mechanistic events leading to the development of posttraumatic epilepsy. Additionally, our current work is also focusing on complex traumatic brain injury models that mimic polytrauma as this type of injury has become more prevalent in combat areas.

#### JUSTIN C. SANCHEZ, PH.D.

Associate Professor, Department of Biomedical Engineering, Director, Neuroprosthetics Research Group; Program Manager, Defense Advanced Research Projects Agency **Neuroprosthetics and Neural Engineering** 

The goals of my research are to develop brain-machine interfaces (BMI) to restore communication and movement control to people with neurological impairments. The approach is to use technology to directly interact with the central and peripheral nervous system, interpret the internal coding of brain activity for intent, and send commands to bionic devices to trigger movements. The laboratory uses electrophysiological and neural computational tools to seamlessly interface these devices with the nervous system. I am interested in developing combined therapies (technology with rehabilitation and repair) to personalize therapeutic approaches for people living with disabilities.





PANTELIS TSOULFAS, M.D. Associate Professor, Departments of Neurological Surgery and Cell Biology & Anatomy **Neurotrophins: Specificity of Action** 

My laboratory is interested in two areas of neurobiology that are significant for developing new strategies for spinal cord injury repair. Over the past years, we have worked to modify neurotrophins that are better suited for use in SCI. We are also interested in understanding the processes involved in maintaining and differentiating neural stem cells.

#### KIM ANDERSON-ERISMAN, PH.D. Research Associate Professor, Department of Neurological Surgery Director of Education, The Miami Project to Cure Paralysis

#### Translational Investigations for Chronic Spinal Cord Injury

My research focuses on translational investigations and bridging the gap between basic science, clinical science, and the public community living with SCI. My current projects focus on 1) aging related changes in bladder health after SCI, 2) determining the minimum amount of exercise and locomotor training required for clinical trials targeting chronic SCI, and 3) identifying the facilitators and barriers to clinical trial participation from the SCI consumer perspective. In addition, I manage our Schwann cell clinical trial program and collaborate with Dr. Levi regarding our participation in an industry-sponsored stem cell trial.





IAN D. HENTALL, PH.D. Research Associate Professor, Department of Neurological Surgery **Brainstem Influences on Neurotrauma** 

Our research is centered on the general idea that serotonin-containing brainstem neurons influence natural repair processes following brain or spinal cord injury. We study in rats how these brainstem (raphé) neurons respond during injury, how raphé activity influences restorative molecular mechanisms in damaged regions, and how recovery from traumatic spinal cord or brain injury is improved by prolonged electrical stimulation of these nuclei or of their input areas. The procedure of deep brain stimulation has potential for treating early of chronic injury in man.

BRIAN R. NOGA, PH.D. Research Associate Professor, Department of Neurological Surgery **Brain and Spinal Mechanisms Controlling Walking** 

Our long-term goal is to develop and optimize treatments for spinal cord injury based on transmitter enhancement strategies that include deep brain stimulation, delivery of neurotransmitters or similarly acting drugs, or transplantation of cells secreting these substances. Of the many possible neurotransmitter candidates that could be used for this purpose, monoamines hold particular promise. We have concentrated our recent research effort on understanding the role monoamines play in the control of walking in normal and injured spinal cord.



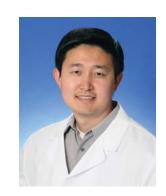


**COLEEN ATKINS, PH.D. Assistant Professor, Department of Neurological Surgery** 

Developing Novel Therapies for Traumatic Brain Injury and Spinal Cord Injury The research in my laboratory focuses on developing novel therapeutic interventions for traumatic brain injury (TBI) and spinal cord injury (SCI). The research goal of my laboratory is to enhance rehabilitation and recovery by manipulating synaptic plasticity at specific levels of the neuroaxis following TBI and SCI. We have found that specific synaptic plasticity signaling pathways are altered after TBI and we are currently using pharmacotherapies to target those pathways to improve behavioral recovery after TBI.

JAE K. LEE, PH.D. **Assistant Professor, Department of Neurological Surgery** Neutralizing Inhibitors of Axon Regeneration; Stimulating Plasticity

The long term research goal in my laboratory is to elucidate the mechanisms of cellular interactions in the injured adult spinal cord that creates an environment inhibitory to axonal growth. Currently, we know which cells can produce what types of inhibitory molecules, but we need a better understanding of how these cells interact and the relative significance of the different inhibitory molecules they produce in order to identify an effective therapeutic target to treat spinal cord injury and related neurological disorders.





KEVIN K. PARK, PH.D. **Assistant Professor, Department of Neurological Surgery Intrinsic Mechanisms of Axon Regeneration** 

My lab is interested in understanding the neuron's intrinsic mechanisms that account for failure of axon regeneration in the central nervous system (CNS). Previously, I and others have identified several key proteins that block axon regeneration, which are present in mature CNS neurons. In my current research, I will further extend my findings in order to better understand the mechanisms governing robust axon regeneration and also to explore the potential of developing therapeutic strategies for spinal cord injury and other neurodegenerative conditions.

#### ROBERTA BRAMBILLA, PH.D. Research Assistant Professor, Department of Neurological Surgery

Modulation of the Neuro-Immune Response in Neurologic Disease

The main focus of my research has been to understand the role of neuroinflammation in the pathophysiology of neurodegenerative disorders (e.g., spinal cord injury and multiple sclerosis), with an interest in the contribution of glial cells, specifically the astrocytes, which represent the most abundant cell population in the nervous system, and the oligodendrocytes, which are responsible for axon myelination. Currently, my laboratory is developing two specific lines of research in the area of neuroimmunology, which are focused on: (1) investigating the role of tumor necrosis factor in the processes of demyelination and remyelination, and (2) understanding how mitochondrial dysfunction in oligodendrocytes may be involved in the etiopathology of multiple sclerosis.





RACHEL E. COWAN, PH.D. Research Assistant Professor, Department of Neurological Surgery

Enhancement and Preservation of Maximal Transfer and Wheelchair Propulsion Ability Our first focus is defining what level of fitness and 'skill' are required to independently perform transfers to and from the bed, car, shower, and ground and if these are different for various levels of SCI. Our second focus is defining how changes in fitness and wheelchair configuration can meaningfully reduce the effort required to propel a manual wheelchair and how these changes may differ by level of SCI.

#### JUAN PABLO DE RIVERO VACCARI, PH.D. Research Assistant Professor, Department of Neurological Surgery

Underlying Mechanisms of the Innate Immune Response and Contributions to Various **CNS Diseases** 

My research focuses on understanding early inflammatory events in central nervous system (CNS) injury. Currently, my laboratory is studying the effects of pattern recognition receptor (PRR)-activation after spinal cord injury (SCI), traumatic brain injury (TBI), and stroke.





MOUSUMI GHOSH, PH.D. Research Assistant Professor, Department of Neurological Surgery

Altering Host Glial Responses following CNS Injury and Disease to Promote Repair My research interests are focused on altering the hostile environment of the injured or diseased CNS to one that is conducive to repair through altering inflammation. Specifically our work focuses on delineating the intrinsic and extrinsic signals present after injury that antagonize the conversion of activated microglia and macrophages to a reparative phenotype in experimental models of CNS injury and disease, such as Multiple Sclerosis. We are also interested in understanding how altering the immunophenotypical profile of macrophages and microglia can affect host glial responses, including glial scar formation, as well as influence the ability of transplanted cells, such as Schwann cells and stem cells, to mediate neurorepair.

#### HOWARD B. LEVENE, M.D., PH.D. **Assistant Professor, Department of Neurological Surgery Schwann Cell Transplantation after SCI**

One proposed therapy for spinal cord injury is to introduce cells to the injury site to help repair, restore, or support existing neurons. I work with my colleagues on a large animal model to study the effect and behavior of transplanted autologous Schwann cells. I have been involved in the refinement of this animal model. This approach allows for the scientific study of the behavior of implanted cells and generates the groundwork for clinical trials. Research utilizing this model is done in collaboration with clinicians and scientists at the Miami Project such as Drs. Guest, Solano, Pearse, Wood, Bunge, and many more.

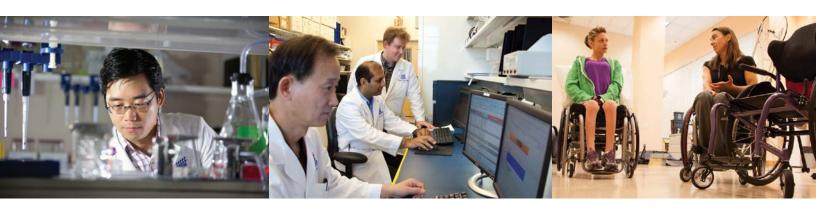




PAULA V. MONJE, PH.D. Research Assistant Professor, Department of Neurological Surgery

Molecular signals controlling Schwann cell proliferation and differentiation In vitro methods to enhance the Schwann cell's potency for CNS repair

My laboratory studies basic aspects of Schwann cell biology and their use in transplantation for CNS regeneration. Specifically, we are investigating the role of cAMP and growth factors in the reciprocal interactions between Schwann cells and neurons underlying the regulation of Schwann cell proliferation and myelination, as well as the initiation of Schwann cell dedifferentiation after injury. Our lab works intensively towards refining the use and developing new cell culture methods for the growth and assessment of function of both human and rodent Schwann cells. One important goal is to improve the quality of cultured adult Schwann cells for an intended use in clinical trials.







major role of The Miami Project is to provide education and training for the next generation of neuroscientists. Our Along-term educational goal is to increase the number of scientists and laboratories working on paralysis research and central nervous system disorders around the world. Students and young scientists beginning their careers gain skills from The Miami Project's state-of-the-art comprehensive research and academic environment.

In 2014, Drs. Anderson-Erisman and Dietrich embarked on year 2 of a 5-year NIH Summer Student Research Grant, which enables a handful of stellar undergraduate students the opportunity to work in the laboratory of a Miami Project faculty member for 10 weeks during the summer. Each week, the students attended 2 lectures and participated in 1 journal club, in addition to 37 hours of hands-on laboratory work (for a total of 40 hours each week). These 11 students wrote an abstract about their specific research project and presented a poster at the 4th Miami Project Summer Student Research Session on August 8, 2014.

C	Chadanta	and their R		Dunington
Summer	Students	and their K	esearch	Projects.

Name	Summer mentor	Summer project
Ali, Aleena	Dr. N. Brackett	Responders versus non-responders to penile vibratory stimulation in men with spinal cord injury
Carsello, Alex	Dr. B. Noga	Deep Brain Stimulation of the Mesencephalic Locomotor Region to Improve Locomotion in Animals with Spinal Contusion Injuries
Haase, Conner*	Dr. M. Bunge	Tumor Necrosis Factor Reduces Schwann Cell Migration into Astrocyte Regions in Culture
Khan, Alicia	Dr. J. Lee	SOCS3 deletion decreases the capacity of NG2 cells to differentiate into oligodendrocytes in the uninjured brain
Krajek, Alex*	Dr. R. Cowan	Assessment of a Novel Fitness Test in Males with Spinal Cord Injury
Purdy, Katherine	Dr. I. Hentall	Combinatorial strategies to treat spinal cord injury
Sam, Pingdewinde	Dr. R. Bullock	Quantitative Analysis Of FDA Approved Human Neural Stem Transplant Survival In Rats With Penetrating Ballistic Brain Injury (PBBI)
Srinivas, Shruthi*	Dr. P. Monje	Optimization of protocols for isolation and purification of rat Schwann cell cultures
Stafford, Carly	Dr. G. Hotz	Preventing Pediatric Pedestrian Trauma through Statewide Implementation of a Traffic Safety Program
Upadhyaya, Anjali	Dr. D. Pearse	In Vitro Assessment of Peripheral Nerve Schwann Cells and Fibroblasts on Neuronal Survival and Neurite Outgrowth
Zhou, Allen	Dr. V. Lemmon	Effect of multiple VP16 activation domains engineered to Krüppel-like factor 7 on neurite outgrowth

\*award winners of the poster session

## SCIENTIFIC PUBLICATIONS

# Scientifications Publications Acute Spinal Cord Injury Publications Scientifications Publications Publications

Published studies that have passed the test of peer review are the benchmark of scientific progress. Listed here are the 2014 research publications by Miami Project scientists and colleagues.

To find links to the abstracts and complete scientific publications listed here, visit the Research Publications section of our website at www.themiamiproject.org/researchpublications

Abeyruwan S, Vempati UD, Küçük-McGinty H, Visser U, Koleti A, Mir A, Sakurai K, Chung C, Bittker JA, Clemons PA, Brudz S, Siripala A, Morales AJ, Romacker M, Twomey D, Bureeva S, Lemmon V, Schürer SC (2014) Evolving BioAssay Ontology (BAO): modularization, integration and applications. *J Biomed Semantics*. 5(Suppl 1 Proceedings of the Bio-Ontologies Spec Interest G.

Adamczak S, de Rivero Vaccari JP, Dale G, Nonner D, Dahl G, Dietrich WD, Keane R (2014) Pyroptotic neuronal cell death mediated by the AIM2 inflammasome. *J Cerebral Blood Flow Metab* 34:621-629.

Ahmad FU, Levi AD (2014) Letters to the Editor: Hypothermia for spinal cord injury. *J Neurosurg Spine*. 21(5):843-5.

Ahmad FU, Wang MY (2014) Lateral mass of C1 fixation and ponticulus-posticus. *World Neurosurg.* 82(1-2):e145-6.

Ahmad FU, Wang MY (2014) The case for aggressive management in traumatic high-grade cervical dislocations. *World Neurosurg.* 82(6):1055-7.

Ahmad FU, Wang MY (2014) Use of anteroposterior view fluoroscopy for targeting percutaneous pedicle screws in cases of spinal deformity with axial rotation. *J Neurosurg Spine*. 21(5):826-32.

Berrocal YA, Almeida VW, Levi AD (2014) Cell transplantation. Response. *J Neurosurg.* 120(2):574.

Berrocal YA, Almeida VW, Puentes R, Knott EP, Hechtman JF, Garland M, Pearse DD (2014) Loss of central inhibition: implications for behavioral hypersensitivity after contusive spinal cord injury in rats. *Pain Res Treat*. 2014:178278.

Beurel E, Harrington LE, Buchser W, Lemmon V, Jope RS (2014) Astrocytes modulate the polarization of CD4+ T cells to Th1 cells. *PLoS One*. 9(1):e86257.

Blaya MO, Bramlett HM, Naidoo J, Pieper AA, Dietrich WD (2014) Neuroprotective efficacy of a proneurogenic compound after traumatic brain injury. *J Neurotrauma* 31:476-486.

Brambilla R, Morton PD, Ashbaugh JJ, Karmally S, Lambertsen KL, Bethea JR (2014) Astrocytes play a key role in EAE pathophysiology by orchestrating in the CNS the inflammatory response of resident and peripheral immune cells and by suppressing remyelination. *Glia*. 62(3):452-67.

Bramlett H, Dietrich WD (In press) Long-term consequences of traumatic brain injury: Current status of potential mechanisms of injury and neurologic outcomes. *J Neurotrauma*.

Bramlett HM, Dietrich WD, Marcillo A, Mawhinney LJ, Furones-Alonso O, Bregy A, Peng Y, Wu Y, Pan J, Wang J, Guo XE, Bauman WA, Cardozo C, Qin W (2014) Effects of low intensity vibration on bone and muscle in rats with spinal cord injury. Osteoporos Int. 25(9):2209-19.

Bullock MR, Schulman CI, Corry JJ, Pappas A (2014) Cooling strategies targeting trauma. Ther Hypothermia Temp Manag. 4(3):105-10.

Burks SS, Levi DJ, Hayes S, Levi AD (2014) Challenges in sciatic nerve repair: anatomical considerations. J*Neurosurg.* 121(1):210-8.

Cahill KS, Martinez JL, Vanni S, Wang MY, Levi AD (2014) Response. J Neurosurg Spine. 20(1):120-1.

Carballosa-Gonzalez MM, Vitores A, Hentall ID (2014) Hindbrain raphe stimulation boosts cyclic adenosine monophosphate and signaling proteins in the injured spinal cord. Brain Res. 1543:165-72.

Cardenas DD, Dalal K (2014) Spinal cord injury rehabilitation. Phys Med Rehabil Clin N Am. 25(3):xv-xvi.

Cardenas DD, Ditunno JF, Graziani V, McLain AB, Lammertse DP, Potter PJ, Alexander MS, Cohen R, Blight AR (2014) Two phase 3, multicenter, randomized, placebocontrolled clinical trials of fampridine-SR for treatment of spasticity in chronic spinal cord injury. Spinal Cord. 52(1):70-6.

Castle SM, Jenkins LC, Ibrahim E, Aballa TC, Lynne CM, Brackett NL (2014) Safety and efficacy of a new device for inducing ejaculation in men with spinal cord injuries. Spinal Cord. 52 Suppl 2:S27-9.

Daftarian MP, Vosoughi A, Lemmon V (2014) Gene-based vaccination and screening methods to develop monoclonal antibodies. Methods Mol Biol. 1121:337-46.

de Rivero Vaccari JC, Brand Iii F, Berti AF, Alonso OF, Bullock R, De Rivero Vaccari JP (In Press) Mincle signaling in the innate immune response after traumatic brain injury. J Neurotrauma.

de Rivero Vaccari JP, Brand FJ 3rd, Sedaghat C, Mash DC, Dietrich WD, Keane RW (2014) RIG-1 receptor expression in the pathology of Alzheimer's disease. J Neuroinflammation. 2;11:67.

de Rivero Vaccari JP, Dietrich WD, Keane RW (2014) Activation and regulation of cellular inflammasomes: gaps in our knowledge for central nervous system injury. J Cereb Blood Flow Metab. 34(3):369-75.

DeFazio MV, Rammo RA, Robles JR, Bramlett HM, Dietrich WD, Bullock MR (2014) The potential utility of blood-derived biochemical markers as indicators of early clinical trends following severe traumatic brain injury. World Neurosurg. 81:151-8.

Dellarole A, Morton P, Brambilla R, Walters W, Summers S, Bernardes D, Grilli M, Bethea JR (2014) Neuropathic pain-induced depressive-like behavior and hippocampal neurogenesis and plasticity are dependent on TNFR1 signaling. Brain Behav Immun. 41:65-81.

Dietrich, WD, Bullock PM, Lundbye JB (2014) Intraoperative temperature management. Ther Hypothermia *Temp Manag.* 4:67-71.

Dixon KJ, Theus MH, Nelersa CM, Mier J, Travieso LG, Yu TS, Kernie SG, Liebl DJ PhD (In press) Endogenous neural stem/progenitor cells stabilize the cortical microenvironment following traumatic brain injury. J Neurotrauma.

Dumas A, Amiable N, de Rivero Vaccari JP, Chae JJ, Keane RW, Lacroix S, Vallières L (2014) The inflammasome pyrin contributes to pertussis toxin-induced IL-1β synthesis, neutrophil intravascular crawling and autoimmune encephalomyelitis. PLoS Pathog. 29;10(5):e1004150.

Ellenbroek D, Kressler J, Cowan RE, Burns PA, Mendez AJ, Nash MS (In press) Effects of prandial challenge on triglyceridemia, glycemia, and pro-inflammatory activity in persons with chronic paraplegia. J Spinal Cord Med.

Field-Fote EC (2014) Saying what we mean, and meaning what we say. J Neurol Phys Ther. 38(4):205-6.

Field-Fote EC (2014) What happens in Vegas... J Neurol *Phys Ther.* 38(1):1.

Fillingim RB, Bruehl S, Dworkin RH, Dworkin SF, Loeser JD, Turk DC, Widerstrom-Noga E, Arnold L, Bennett R, Edwards RR, Freeman R, Gewandter J, Hertz S, Hochberg M, Krane E, Mantyh PW, Markman J, Neogi T, Ohrbach R, Paice JA, Porreca F, Rappaport BA, Smith SM, Smith TJ, Sullivan MD, Verne GN, Wasan AD, Wesselmann U (2014) The ACTTION-American Pain Society Pain Taxonomy (AAPT): an evidence-based and multidimensional approach to classifying chronic pain conditions. *J Pain.* 15(3):241-9.

Flight RM, Harrison BJ, Mohammad F, Bunge MB, Moon LD, Petruska JC, Rouchka EC (2014) categoryCompare, an analytical tool based on feature annotations. Front Genet.5:98.

Gajavelli S, Kentaro S, Diaz J, Yokobori s, Spurlock M, Diaz D, Jackson C, Wick A, Zhaol W, Leung LY, Shear D, Tortella F, Bullock R (In Press) Cerebral metabolism after penetrating brain injury. *Cerebral blood flow and metabolism*.

Gaunaurd IA, Gómez-Marín OW, Ramos CF, Sol CM, Cohen MI, Cahalin LP, Cardenas DD, Jackson RM (In press) Physical activity and quality of life improvements of patients with Idiopathic Pulmonary Fibrosis completing a pulmonary rehabilitation program. *Respir Care*. pii: respcare.03180.

Gilbert O, Croffoot JR, Taylor AJ, Nash MS, Groah SL, Schomer K (2014) Serum lipid concentrations among persons with spinal cord injury: A systematic review and meta-analysis of the literature. *Atherosclerosi.* 232:305-12.

Gomes-Osman J, Field-Fote EC (In press) Cortical vs. afferent stimulation as an adjunct to functional task practice training: A randomized, comparative pilot study in people with cervical spinal cord injury. *Clin Rehabil*.

Grossman RG, Fehlings MG, Frankowski RF, Burau KD, Chow DS, Tator C, Teng A, Toups EG, Harrop JS, Aarabi B, Shaffrey CI, Johnson MM, Harkema SJ, Boakye M, Guest JD, Wilson JR (2014) A prospective, multicenter, phase I matched-comparison group trial of safety, pharmacokinetics, and preliminary efficacy of riluzole in patients with traumatic spinal cord injury. *J Neurotrauma*. 31(3):239-55.

Guest J, Dietrich D (In press) Letter to the editor. Commentary regarding the recent publication by Tabarow et al, Functional regeneration of supraspinal connections in a patient with transected spinal cord following transplantation of bulbar olfactory ensheathing cells with peripheral nerve bridging. *J Neurotrauma*.

Hama A, Germano P, Varghese MS, Cravatt BF, Milne GT, Pearson JP, Sagen J (2014) Fatty acid amide hydrolase (FAAH) inhibitors exert pharmacological effects, but lack antinociceptive efficacy in rats with neuropathic spinal cord injury pain. *PLOS One* 9(5):e96396.

Hama A, Sagen J (2014) Effects of repeated dosing with mechanistically distinct antinociceptive ligands in a rat model of neuropathic spinal cord injury pain. *Pharmacol Res Persp* 2 (2): 1-9 e00034 DOI: 10.1002/prp2.34).

Hama A, Sagen J (2014) Selective antinociceptive effects of a combination of the N-methyl-D-aspartate receptor peptide antagonist [Ser1]histogranin and morphine in rat models of pain. *Pharmacol Res Persp* 2 (2): 1-11. E00032 DOI: 10.1002/prp2.32.

Hawryluk GWJ, Bullock Ross (In press) Design of acute neuroprotection studies. *Handbook of Clinical Neurology*.

Hittinger E, Raeburn J, Nash MS, Perry A, Signorile J, Kressler J, Jacobs K (2014) Ischemic preconditioning does not improve peak exercise capacity at sea level or simulated high altitude in trained male cyclists. *Applied Physiology, Nutrition, and Metabolism*, 10.1139/apnm-2014-0080

Hofstetter CP, Wang MY (2014) Burst fractures. *J Neurosurg Spine*. 20(2):148-9.

Hofstetter CP, Wang MY (2014) Diagnostic and therapeutic challenges of cervical myelopathy. *World Neurosurg.* 82(1-2):e173-5.

Hooshmand J, Hotz G, Neilson V, Chandler L (2014) BikeSafe: evaluating a bicycle safety program for middle school aged children. *Accid Anal Prev.* 66:182-6.

Ibrahim E, Castle SM, Aballa TC, Keane RW, de Rivero Vaccari JP, Lynne CM, Brackett NL (2014) Neutralization of ASC improves sperm motility in men with spinal cord injury. *Hum Reprod.* 29(11):2368-73.

Jackson DG, Wang J, Keane RW, Scemes E, Dahl G (2014) ATP and potassium ions: a deadly combination for astrocytes. *Sci Rep.* 4:4576.

Jackson RM, Gómez-Marín OW, Ramos CF, Sol CM, Cohen MI, Gaunaurd IA, Cahalin LP, Cardenas DD (2014) Exercise limitation in IPF patients: a randomized trial of pulmonary rehabilitation. *Lung.* 192(3):367-76.

Jayakumar AR, Tong XY, Ruiz-Cordero R, Bregy A, Bethea JR, Bramlett HM, Norenberg MD (2014) Activation of NF-κB mediates astrocyte swelling and brain edema in traumatic brain injury. *J Neurotrauma*. 31(14):1249-57.

Johnson JN, Cummock MD, Levi AD, Green BA, Wang MY (2014) Moderate hypothermia for intradural spinal tumor resection: a cohort comparison and feasibility study. *Ther Hypothermia Temp Manag.* 4(3):137-44.

Jordan LM, McVagh, J, Noga BR, Cabaj AM, Majczyński H, Sławińska U, Provencher J, Leblond, H, Rossignol S (2014) Cholinergic mechanisms in spinal locomotion – Potential target for rehabilitation approaches. In: Neuromodulatory Control of Spinal Function in Health and Disease. *Frontiers Neural Circuits*, 8:132. doi:10.3389/fncir.2014.00132.

Jose J, Smith MK, Kaplan L, Lesniak B, Levi AD (In press) Ultrasound-guided needle localization of the saphenous nerve for removal of neuroma in the infrapatellar branches: Technical Report. *Neurosurgery*.

Kanno H, Pearse DD, Ozawa H, Itoi E, Bunge MB (In press) Schwann cell transplantation for spinal cord injury repair: its significant therapeutic potential and prospectus. Reviews in the Neurosciences.

Kanno H, Pressman Y, Moody A, Berg R, Muir EM, Rogers JH, Ozawa H, Itoi E, Pearse DD, Bunge MB (2014) Combination of engineered Schwann cell grafts to secrete neurotrophin and chondroitinase promotes axonal regeneration and locomotion after spinal cord injury. J Neurosci. 34(5):1838-55.

Khan MF, Burks SS, Al-Khayat H, Levi AD (2014) The effect of steroids on the incidence of gastrointestinal hemorrhage after spinal cord injury: a case-controlled study. Spinal Cord. 52(1):58-60.

Kigerl KA, de Rivero Vaccari JP, Dietrich WD, Popovich PG, Keane RW (2014) Pattern recognition receptors and central nervous system repair. Exp Neurol. 258:5-16.

Knott EP, Assi M, Pearse DD (2014) Cyclic AMP signaling: a molecular determinant of peripheral nerve regeneration. Biomed Res Int. 2014:651625.

Kressler J, Cowan RE, Bigford GE, Nash MS (2014) Reducing cardiometabolic disease in spinal cord injury. Phys Med Rehabil Clin N Am. 25(3):573-604, viii.

Kressler J, Thomas CK, Field-Fote EC, Sanchez J, Widerström-Noga E, Cilien DC, Gant K, Ginnety K, Gonzalez H, Martinez A, Anderson KD, Nash MS (2014) Understanding therapeutic benefits of overground bionic ambulation: exploratory case series in persons with chronic, complete spinal cord injury. Arch Phys Med Rehabil. 95(10):1878-1887.e4.

Kressler J, Burns PA, Betancourt, L, Nash MS, (2014) Circuit training and protein supplementation in persons with chronic tetraplegia. Med Sci Sports Exer. 46(7):1277-84.

Kressler J, H Ghersin, MS Nash (2014) Use of quasisupervised home-based FES cycle ergometers by individuals with neurological impairments. Top Spinal Cord Inj Rehabi. 20(2):123-6.

Kressler J, Jacobs K, Burns P, Betancourt L, Nash MS (2014) Effects of circuit resistance training and timely protein supplementation on exercise-induced fat oxidation in tetraplegic adults. Top Spinal Cord Inj Rehabil. 20(2):113-22.

Kressler J, Cowan RE, Bigford G, Nash MS (2014) Reducing cardiometabolic disease in spinal cord injury. Phys Med Rehabil Clin North Am. 25(3):573-604.

Krishna V, Andrews H, Varma A, Mintzer J, Kindy MS, Guest J (2014) Spinal cord injury: how can we improve the classification and quantification of its severity and prognosis? J Neurotrauma. 31(3): 215-227.

Lee DH, Luo X, Yungher B, Bray E, Lee JK\*, Park, KK\* (2014) Mammalian target of rapamycin's distinct roles and effectiveness in promoting compensatory axonal sprouting in the injured CNS. Journal of Neuroscience. 34(46):15347-15355. \*co-corresponding authors

Lemmon VP, Abeyruwan S, Visser U, Bixby JL (2014) Facilitating transparency in spinal cord injury studies using data standards and ontologies. Neural Regen Res. 9(1):6-7.

Lemmon VP, Ferguson AR, Popovich PG, Xu X-M, Snow DM, Igarashi M, Beattie CE, Bixby JL, Abeyruwan SW et al (including Liebl DJ) (2014) Minimum Information About a Spinal Cord Injury Experiment (MIASCI) – a proposed reporting standard for spinal cord injury experiments. Journal of Neurotrauma. 31(15):1354-61.

Lerch JK, Martínez-Ondaro YR, Bixby JL, Lemmon VP (2014) cJun promotes CNS axon growth. Mol Cell Neurosci. 59:97-105.

Levene, HB, Ghobrial, GM, Jallo, JI, Lateral Extracavitary Approach, in Spine Surgery: Tricks of the Trade, A.R. Vaccaro and T.J. Albert, Editors. 2014, pending, Thieme Medical Publishers, Inc.: New York. p. TBD.

Levene, HB, Wang, MY, Green, BA, Emhoff, TA, Spinal Cord Trauma, in Irwin & Rippe's manual of intensive care medicine, R.S. Irwin, J.M. Rippe, and C.M. Lilly, Editors. 2014, Wolters Kluwer Health/Lippincott Williams & Wilkins: Philadelphia. p. 924-932.

Li L, Brockmeier AJ, Choi JS, Francis JT, Sanchez JC, Príncipe JC (2014) A tensor-product-kernel framework for multiscale neural activity decoding and control. Comput Intell Neurosci. 2014:870160.

Liu Y, Grumbles RM, Thomas CK (2014) Electrical stimulation of transplanted motoneurons improves motor unit formation. J Neurophysiol. 112(3):660-70.

Long J, Li B, Rodriguez-Blanco J, Pastori C, Volmar CH, Wahlestedt C, Capobianco A, Bai F, Pei XH, Ayad NG, Robbins DJ (2014) The BET bromodomain inhibitor I-BET151 acts downstream of smoothened protein to abrogate the growth of hedgehog protein-driven cancers. J Biol Chem. 289(51):35494-502.

Luo X, Yungher B, Park KK (2014) Application of tissue clearing and light sheet fluorescence microscopy to assess optic nerve regeneration in unsectioned tissues. Methods Mol Biol. 1162:209-17.

Lynne CM, Ibrahim E, Brackett NL (In press) Electroejaculation and penile vibratory stimulation. In: Case Studies in Male Infertility: Expert Analysis and Current Controversies. E Kim and R Brannigan (Eds) JP Medical Publishers, London.

Miranda RA, Casebeer WD, Hein AM, Judy JW, Krotkov EP, Laabs TL, Manzo JE, Pankratz KG, Pratt GA, Sanchez JC, Weber DJ, Wheeler TL, Ling GS (2014) DARPAfunded efforts in the development of novel brain-computer interface technologies. J Neurosci Methods. pii: S0165-0270(14)00270-2.

Morris-Blanco KC, Cohan CH, Neumann JT, Sick TJ, Perez-Pinzon MA (2014) Protein kinase C epsilon regulates mitochondrial pools of Nampt and NAD following resveratrol and ischemic preconditioning in the rat cortex. J *Cereb Blood Flow Metab.* 34(6):1024-32.

Nanayakkara, A, Santamaria, J, Benavides, F, Guada, L, Levene, HB, Solano, J, Guest, J (2014) The comprehensive mini pig model for spinal cord injury (SCI) research. in Canes Day. University of Miami.

Nash MS, Hicks A (2014) Letter: How "healthy" is circuit resistance training following paraplegia? Kinematic analysis associated with shoulder mechanical impingement risk. J Rehab Res Develop 51(2): vii-ix.

Nash MS (2014) Editorial Note on: Optimal scaling of weight and waist circumference to height for adiposity and cardiovascular disease risk in individuals with spinal cord injury. Spinal Cord October 21, 2014; doi:10.1038/ sc.2014.176

Nasirinezhad F, Gajavelli S, Priddy B, Jergova S, Zadina J, Sagen J (In press) Viral vectors encoding endomorphins and serine histogranin attenuate neuropathic pain symptoms after spinal cord injury in rats. Mol Pain.

Nielson JL, Guandique CF, Liu AW, Burke DA, Lash AT, Moseanko R, Hawbecker S, Strand SC, Zdunowski S, Irvine KA, Brock JH, Nout-Lomas YS, Gensel JC, Anderson KD, Segal MR, Rosenzweig ES, Magnuson DS, Whittemore SR, McTigue DM, Popovich PG, Rabchevsky AG, Scheff SW, Steward O, Courtine G, Edgerton VR, Tuszynski MH, Beattie MS, Bresnahan JC, Ferguson AR (2014) Development of a database for translational spinal cord injury research. J Neurotrauma. 31(21):1789-99.

Novrup HG, Bracchi-Ricard V, Ellman DG, Ricard J, Jain A, Runko E, Lyck L, Yli-Karjanmaa M, Szymkowski DE, Pearse DD, Lambertsen KL, Bethea JR (In press) Central but not systemic administration of XPro1595 is

therapeutic following moderate spinal cord injury in mice. J Neuroinflammation.

Pastori C, Daniel M, Penas C, Volmar CH, Johnstone AL, Brothers SP, Graham RM, Allen B, Sarkaria JN, Komotar RJ, Wahlestedt C, Ayad NG (2014) BET bromodomain proteins are required for glioblastoma cell proliferation. Epigenetics. 9(4):611-20.

Penas C, Ramachandran V, Simanski S, Lee C, Madoux F, Rahaim RJ, Chauhan R, Barnaby O, Schurer S, Hodder P, Steen J, Roush WR, Ayad NG(2014) Casein kinase 1δ-dependent Wee1 protein degradation. J Biol Chem. 289(27):18893-903.

Prasad A, Xue QS, Dieme R, Sankar V, Mayrand RC, Nishida T, Streit WJ, Sanchez JC (2014) Abiotic-biotic characterization of Pt/Ir microelectrode arrays in chronic implants. Front Neuroeng. 7:2.

Prins NW, Sanchez JC, Prasad A (2014) A confidence metric for using neurobiological feedback in actor-critic reinforcement learning based brain-machine interfaces. Front Neurosci. 8:111.

Sankar V, Patrick E, Dieme R, Sanchez JC, Prasad A, Nishida T (2014) Electrode impedance analysis of chronic tungsten microwire neural implants: understanding abiotic vs. biotic contributions. Front Neuroeng. 7:13.

Saunders NR, Noor, NM, Dziegielewska M, Wheaton BN, Liddelow SA, Steer DL, Ek CJ, Habgood MD, Wakefield M, Lindsay H, Truettner T, Miller RD, Smith AI, Dietrich WD (In press) Age-dependent gene and protein expression following transection of neonatal spinal cord of Monodelphis domestica (South American Grew Shorttailed opossum). PLoS One.

Scivoletto G, Glass C, Anderson KD, Galili T, Benjamin Y, Front L, Aidinoff E, Bluvshtein V, Itzkovich M, Aito S, Baroncini I, Benito-Penalva J, Castellano S, Osman A, Silva P, Catz A (In press) An international age- and gender-controlled model for the Spinal Cord Injury Ability Realization Measurement Index (SCI-ARMI). Neurorehabil Neural Repair.

Serak J. Wang MY (2014) Weighing the risks of perioperative aspirin. World Neurosurg. 82(6):928-9.

Sherman AL, Wang MY (2014) Hypothermia as a clinical neuroprotectant. Phys Med Rehabil Clin N Am. 25(3):519-29.

Shimoda K, Maeda T, Tado M, Yoshino A, Katayama Y, Bullock MR (In press) Outcome and surgical management for geriatric traumatic brain injury: Analysis of 888 cases

registered in the Japan Neurotrauma Data Bank. World Neurosurg.

Simmons OL, Kressler J, Nash MS (2014) Reference fitness values in the untrained spinal cord injury population. *Arch Phys Med Rehabil*.

Stathias V, Pastori C, Griffin TZ, Komotar R, Clarke J, Zhang M, Ayad NG (2014) Identifying glioblastoma gene networks based on hypergeometric test analysis. *PLoS One*. 9(12):e115842.

Theus MH, Liebl DJ (2014) Endogenous neurogenesis after traumatic brain injury. IN: Cellular Therapy for Stroke and CNS Injuries. Zhao LR, Zhang JH (eds). Springer Series in Translational Stroke Research. Chapter 10.

Theus MH, Ricard J, Glass SJ, Travieso LG, Liebl DJ (2014) EphrinB3 blocks EphB3 dependence receptor functions to prevent cell death following traumatic brain injury. *Cell Death Dis.* 5:e1207.

Thomas CK, Bakels R, Klein CS, Zijdewind I (2014) Human spinal cord injury: motor unit properties and behaviour. *Acta Physiol* (Oxf). 210(1):5-19. Thomas CK, Dididze M, Martinez A, Morris RW (2014) Identification and classification of involuntary leg muscle contractions in electromyographic records from individuals with spinal cord injury. *J Electromyogr Kinesiol*. 24(5):747-54.

Thomas CK, Grumbles RM (2014) Age at spinal cord injury determines muscle strength. *Front Integr Neurosci*.8:2.

Titus DJ, Furones C, Atkins CM, Dietrich WD (In press) Emergence of cognitive deficits after mild traumatic brain injury due to hyperthermia. *Exp. Neurol.* 

Titus DJ, Oliva AA, Wilson NM, Atkins CM (In press) Phosphodiesterase inhibitors as therapeutics for traumatic brain injury. *Curr. Pharm.* 

Valle EJ, Van Haren RM, Allen CJ, Jouria JM, Bullock MR, Schulman CI, Namias N, Livingstone AS, Proctor KG (2014) Does traumatic brain injury increase the risk for venous thromboembolism in polytrauma patients? *J Trauma Acute Care Surg.* 77(2):243-50.

Wang MY (2014) The enemy of good is perfect. *World Neurosurg*. pii: S1878-8750(14)01064-X.

Wang MY (2014) Development of a systematic algorithm to triage patients with low back pain to a physician. *Neurosurgery.* 75(6):N11-2.

Wang MY (2014) Miniopen pedicle subtraction osteotomy: surgical technique and initial results. *Neurosurg Clin N Am*. 25(2):347-51.

Wang MY (2014) Physician protect thyself. *World Neurosurg*. pii: S1878-8750(14)01063-8.

Wang MY, Mummaneni PV, Fu KM, Anand N, Okonkwo DO, Kanter AS, La Marca F, Fessler R, Uribe J, Shaffrey CI, Lafage V, Haque RM, Deviren V, Mundis GM Jr; Minimally Invasive Surgery Section of the International Spine Study Group (2014) Less invasive surgery for treating adult spinal deformities: ceiling effects for deformity correction with 3 different techniques. *Neurosurg Focus*. 36(5):E12.

Wang MY, Vasudevan R, Mindea SA (2014) Minimally invasive lateral interbody fusion for the treatment of rostral adjacent-segment lumbar degenerative stenosis without supplemental pedicle screw fixation. *J Neurosurg Spine*. 21(6):861-6.

Widerström-Noga E, Biering-Sørensen F, Bryce TN, Cardenas DD, Finnerup NB, Jensen MP, Richards JS, Siddall PJ (2014) The International Spinal Cord Injury Pain Basic Data Set (version 2.0). *Spinal Cord*. 52(4):282-6.

Widerström-Noga EG, Cruz-Almeida Y, Felix ER, Pattany PM (In press) Somatosensory phenotype is associated with thalamic metabolites and pain intensity after spinal cord injury. *Pain*.

Widerström-Noga EG (In press) The assessment and treatment of pain syndromes in neurorehabilitation. *Oxford Textbook of Neurorehabilitation* (Volker Dietz/Nick Ward eds.) Oxford University Press (UK), Oxford.

Winslow J, Martinez A, Thomas C (In press) Automatic identification and classification of muscle spasms in long-term EMG recordings. *IEEE J Biomed Health Inform.* 2014 Apr 29.

Wu X, Liu J, Tanadini LG, Lammertse DP, Blight AR, Kramer JL, Scivoletto G, Jones L, Kirshblum S, Abel R, Fawcett J, Field-Fote E, Guest J, Levinson B, Maier D, Tansey K, Weidner N, Tetzlaff WG, Hothorn T, Curt A, Steeves JD (In press) Challenges for defining minimal clinically important difference (MCID) after spinal cord injury. *Spinal Cord*.

Zhu Y, Soderblom C, Krishnan V, Ashbaugh J, Bethea JR, Lee JK (In press) Hematogenous macrophage depletion reduces the fibrotic scar and increases axonal growth after spinal cord injury. *Neurobiology of Disease*.

