UNIVERSITY OF SOUTH FLORIDA

Division of Allergy and Immunology
Department of Internal Medicine
Joy McCann Culverhouse Airway Disease Research Center
and The James A. Haley V.A. Medical Center
Tampa, Florida

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2012 - 2013
Annual Report

http://health.usf.edu/medicine/internalmedicine/allergy/index.htm
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I. GREETINGS!

The late Samuel C. Bukantz, MD, founded the University of South Florida College of Medicine, Department of Internal Medicine, Division of Allergy and Immunology in 1972. Richard F. Lockey, M.D. succeeded Dr. Bukantz in 1983 and is the current Director of the Division. Mrs. Joy McCann Culverhouse endowed the Division in 1997 and The Joy McCann Culverhouse Airway Disease Research Center was dedicated in February 1998. In 1998, Mabel and Ellsworth Simmons endowed the Division with a grant for education and research.

The goals of the Division are: first, to provide care to patients with allergic and immunologic diseases at the University of South Florida College of Medicine, Tampa General Hospital, James A. Haley V.A. Medical Center, All Children’s Hospital, and H. Lee Moffitt Cancer Center; second, to train students, residents, and fellows in the subspecialty of allergy and immunology; and third, to conduct basic and clinical research in allergy, asthma, and immunology.

Individuals interested in collaborating with members of the Division may contact Richard F. Lockey, M.D. or any faculty member at (813) 972-7631 (email: rlockey@health.usf.edu).

Mark Ballow, M.D., Professor, Department of Pediatrics and Internal Medicine, Division of Allergy & Immunology, may be contacted at (727)-553-3533 (email: mballow@health.usf.edu)

Richard F. Lockey, MD
Distinguished University Health Professor
Professor of Medicine, Pediatrics & Public Health
Joy McCann Culverhouse Chair of Allergy and Immunology
Director, Division of Allergy and Immunology
University of South Florida Morsani College of Medicine
Department of Internal Medicine
and
Chief, Section of Allergy/Immunology
James A. Haley Veterans’ Hospital
Tampa, Florida
II. FACULTY AND STAFF

Core Faculty

* Richard F. Lockey, M.D., University Distinguished Health Professor; Professor of Medicine, Pediatrics, and Public Health; Division Director; Joy McCann Culverhouse Chair of Allergy and Immunology
  Thomas B. Casale, M.D., Professor of Medicine
* Roger W. Fox, M.D., Professor of Medicine, Pediatrics and Public Health
* Mark C. Glaum, M.D., Ph.D., Associate Professor of Medicine and Pediatrics
* Dennis K. Ledford, M.D., Professor of Medicine and Pediatrics; Mabel & Ellsworth Simmons Professor
  Narasaiah Kolliputi, Ph.D., Associate Professor of Medicine & Pediatrics
  Michael Teng, Ph.D., Associate Professor of Medicine & Pediatrics
  Jia-Wang Wang, Ph.D., Assistant Professor of Medicine & Pediatrics

Joint Faculty

  Mark Ballow, M.D., Professor of Pediatrics and Internal Medicine; Director, Pediatric Allergy & Immunology
  Blanca Camoretti-Mercado, Ph.D., Assistant Professor of Personalized Medicine and Medicine
* Sandra G. Gompf, M.D., Associate Professor of Medicine
  Jennifer Leiding, M.D., Assistant Professor of Pediatrics and Medicine
  Gary W. Litman, Ph.D., Professor of Pediatrics and Medicine; University Distinguished Health Professor; Andrew and Ann Hines Chair in Pediatrics
  Mandel R. Sher, M.D., Clinical Professor of Pediatrics and Medicine
  Panida Sriaroon, M.D., Assistant Professor of Pediatrics and Medicine
  Nathan Tang, M.D., Associate Professor of Pediatrics and Medicine

Clinical Faculty

  Enrique Fernandez-Caldas, Ph.D., Clinical Professor of Medicine
* Monroe J. King, D.O., Adjunct Clinical Associate Professor of Medicine
* Brett E. Stanaland, M.D., Clinical Associate Professor of Medicine
* G. Edward Stewart II, M.D., Clinical Associate Professor of Medicine
* Hugh H. Windom, M.D., Clinical Associate Professor of Medicine
  Rosa Codina, Ph.D., Clinical Assistant Professor of Medicine
  Mary L. Jelks, M.D., Clinical Assistant Professor of Medicine
* Ronald T. Purcell, M.D., Clinical Assistant Professor of Medicine
  Glenn Whelan, Pharm D, Clinical Assistant Professor of Medicine

* Have joint appointments in the Section of Allergy and Immunology, Department of Internal Medicine, James A. Haley Veterans’ Hospital, Tampa, Florida
Richard F. Lockey, M.D., M.S.

Dr. Richard F. Lockey received his B.S. degree from Haverford College, Haverford, Pennsylvania; M.D. from Temple University, Philadelphia, Pennsylvania (Alpha Omega Alpha); M.S. from the University of Michigan in Ann Arbor, Michigan where he trained in Internal Medicine and Allergy/Immunology (A/I) and was a Major and Chief of A/I at Carswell Air Force Base, Fort Worth, Texas, from 1970-1972. He received a medal from the Florida Academy of Sciences, Tallahassee, Florida, in 2000, for his dedication and work to improve the health and well-being of the community and citizens of Florida. He was also the recipient of the Southern Medical Society, Dr. Robert D. and Alma W. Moreton Orginal Research Award in 2012. The American Academy of Allergy Asthma and Immunology presented him with a Special Recognition Award in 1993, Distinguished Service Award in 1999, and Distinguished Clinician Award in 2008. He has the honor of authoring, co-authoring or editing over 600 publications and 35 books or monographs with colleagues and has lectured on numerous occasions nationally and internationally. He is the co-editor of two books and an encyclopedia of allergy/immunology, with Dennis K. Ledford, MD, published in 2014: Asthma, Comorbidities, Co-Existing Conditions, and Differential Diagnoses, Oxford University Press; Allergens and Allergen Immunotherapy: Subcutaneous, Sublingual and Oral, 5th edition, CRC Press/Taylor & Francis Group; and Encyclopedia of Infection and Immunity, Springer, Inc. Professional honors include President of the American Academy of Allergy Asthma and Immunology (1992), past Director of the American Board of Allergy and Immunology (1993-1998) and President of the World Allergy Organization (2010-2012). He has served as co-editor or participant of two WHO reports and served on many journal editorial boards.

Over 90 physician specialists and 50 international post-graduate PhDs or MDs in basic and clinical research and medicine, many of whom have assumed leadership positions in medicine throughout the world, have been trained in the Division. The Division's staff consists of 5 clinicians, 3 basic scientists, and approximately 60 other healthcare professionals including physicians, support, and laboratory personnel.

Areas of expertise and research: insect allergy; allergen immunotherapy; asthma; inflammatory lung diseases; pulmonary fibrosis; co-morbid conditions of asthma; and respiratory syncytial virus vaccine development.
Thomas B. Casale, M.D.

Before joining USF in October 2013, as Professor of Medicine and Chief of Clinical and Translational Research, Dr. Thomas Casale was Professor of Medicine and Medical Microbiology and Immunology and Chief of Allergy/Immunology at Creighton University, Omaha, Nebraska. He did an allergy/immunology fellowship at the National Institutes of Health, Bethesda, MD, where he was chief medical staff fellow. From 1984 to 1996 he was at the University of Iowa where he attained the rank of Professor of Medicine and Director of Allergy/Immunology.

Dr. Casale is a member of the American Thoracic Society and served on their Board of Directors; American Society for Clinical Investigation; and a Fellow of the American College of Physicians and both the American College and American Academy of Allergy Asthma and Immunology. He is a Past President of the American Academy of Allergy Asthma and Immunology and the current Executive Vice President. He is a past member of the Board of Directors of the World Allergy Organization. He also served on the American Board of Allergy and Immunology and was Chair from 2005-2006.

Dr. Casale’s clinical and basic research interests are directed toward the determination and treatment of the pathophysiologic mechanisms involved in asthma and allergic diseases. He has published over 300 scientific papers, reviews and chapters on these topics.
Roger W. Fox, M.D.

After receiving his medical degree from St. Louis University School of Medicine, Dr. Roger W. Fox completed his 3 years of internal medicine and 2 years in allergy and immunology at the University of South Florida, Morsani College of Medicine, Tampa, FL. He joined the Division’s faculty in July, 1980. He is a Fellow of the American College of Physicians and the American Academy of Allergy Asthma and Immunology.

He has been elected to “The Best Doctors in America” for the past decade. He serves as the Director of Education of the allergy/immunology fellowship training program and has helped train over 75 physicians in this specialty. He has published extensively and presented at local, national and international medical meetings and has served on various boards, including the Hillsborough County Medical Association, the Florida Allergy Asthma and Immunology Society of which he has been president, and numerous committees in the American Academy of Allergy Asthma and Immunology.

Dr. Fox enjoys being a clinician and mentor and in that capacity, sees patients at the University of South Florida Morsani Medical Clinics, the Veterans’ Administration Hospital Allergy Clinic, as well as the other clinics affiliated with the University of South Florida. Dr. Fox is an attending physician at the James A. Haley Tampa VA Hospital and he has staff privileges at Tampa Genera Hospital, Moffitt Cancer Hospital, and Florida Hospital.

His research interests include vocal cord dysfunction, urticaria and angioedema, comorbid conditions of asthma, allergic drug reactions and atopic eczema.
Mark C. Glaum, M.D., Ph.D.

Dr. Mark Glaum was born and raised in the Philadelphia area of southeastern Pennsylvania. He earned his B.A. in psychology from Fordham University, New York City. Following graduation, he returned to Philadelphia, PA, and enrolled in the Graduate School at Hahnemann University where he was awarded a Master of Science in Physiology. Immediately following graduate school, he enrolled in Hahnemann Medical School, Philadelphia, PA, and was accepted into their combined MD/PhD program, where he was awarded an MD degree as well as a PhD in immunology. He stayed at Hahnemann Hospital to complete his internship and residency in internal medicine and then completed a fellowship in allergy and immunology at the University of Pennsylvania, Philadelphia, PA, where he received the Stanley E. Bradley Award for Bench Research in Internal Medicine. Dr. Glaum is board certified in both internal medicine and allergy and immunology and is a fellow of the American Academy of Allergy Asthma and Immunology.

Dr. Glaum is an associate professor of medicine and pediatrics at the Morsani College of Medicine, University of South Florida. As a faculty member in the Division of Allergy and Immunology at the College of Medicine, Dr. Glaum is interested in research and education with teaching responsibilities for medical students, residents and fellows who are training to become allergy and immunology specialists.

In addition to teaching, Dr. Glaum’s research interests include mast cell biology, the study of chronic sinusitis with nasal polyps and pollen identification using novel molecular techniques.
Dennis K. Ledford, M.D.

Dr. Dennis Ledford received his medical degree from the University of Tennessee Center for Health Sciences, Nashville, Tennessee. He completed his internal medicine residency there and served as chief medical resident for Dr. Gene Stollerman, M.D., Chairman of Internal Medicine at this same university. A fellowship in rheumatology and immunology followed at New York University and Bellevue Hospital in New York as well as a fellowship in allergy and immunology at the University of South Florida. He joined the faculty at USF Morsani College of Medicine and achieved the rank of professor of medicine in 2000.

Local and regional activities include past service as President of the USF Medical Faculty for the Morsani College of Medicine and President of the Florida Allergy Asthma and Immunology Society. He is current Head of the Allergy/Immunology Section, Florida Hospital, Tampa, FL. National contributions include current service as an associate editor of the *Journal of Allergy and Clinical Immunology* and chair of the Steering Committee for the Allergy Asthma and Immunology Education and Research Trust Fund. He also served as President of the American Academy of Allergy Asthma and Immunology, Co-Chair of the American Council of Graduate Medical Education (ACGME), Allergy/Immunology Residency Review Committee and Director of the American Board of Allergy and Immunology.

Clinical responsibilities and student and resident teaching are combined with research interests in severe, glucocorticoid-dependent asthma, allergen characterization, the association of gastroesophageal reflux and upper airway disease, and eosinophilic esophagitis.
**Narasaiah Kolliputi, Ph.D.**

Dr. Narasaiah Kolliputi is an associate professor (tenured) and Division Director of Research Education for the Division of Allergy and Immunology, Department of Internal Medicine, at the USF Morsani College of Medicine, Tampa, FL. He received his postdoctoral training at Massachusetts General Hospital, Harvard Medical School, Boston, MA. Prior to that time, he received his BS in biology and chemistry in 1997 followed by an MS in biochemistry in 1999 at Sri Venkateswara University, Tirupati, India. He then went on to do his PhD in biochemistry at Osmania University, Hyderabad, India, which he completed in 2004.

Dr. Kolliputi has published 25 papers, including a paper in *Circulation Research and Immunology*. He currently serves as a grant reviewer for the National Institutes of Health, VA Merit Grants, USA Department of Defense and the American Heart Association. He is an associate editor for *Frontiers in Pharmacology, Frontiers in Physiology, and Frontiers in Genetics*, and is a guest associate editor for *Frontiers in Mitochondrial Physiology*. He is also an editorial board member for *Translational Medicine, Virology & Mycology* and *Journal of Biocatalysis & Biotransformation*. Dr. Kolliputi’s research is funded by an NIH RO1 and an American Heart Association Scientist Developmental grant.

He is working on translational strategies to attenuate oxidative stress mediated acute lung injury (ALI), pulmonary fibrosis and pulmonary arterial hypertension.
Dr. Michael Teng received his Ph.D. in immunology from the University of Chicago in 1993. He trained as a postdoctoral fellow studying viral pathogenesis at The Scripps Research Institute in La Jolla, CA. Subsequently, he became a research fellow at the National Institute of Allergy and Infectious Diseases, investigating the molecular biology of respiratory syncytial virus (RSV) and RSV vaccine development. In 2002, he accepted a faculty appointment in the Department of Biochemistry and Molecular Biology at the Pennsylvania State University, University Park, PA, where his laboratory continued studies on RSV pathogenesis.

Dr. Teng joined the faculty of the Division of Allergy and Immunology at USF Morsani College of Medicine in 2010 and is director of the basic research program in the Division. He holds joint appointments in the Departments of Pediatrics, Molecular Medicine, and Pharmaceutical Sciences (College of Pharmacy). Dr. Teng currently serves as a grant reviewer for the National Institutes of Health and the American Heart Association. His past and present research funding includes grants from the National Institutes of Health, the American Heart Association, and contracts with pharmaceutical companies.

Dr. Teng’s research focuses on the host-virus interactions important for RSV pathogenesis. In particular, he is interested in the mechanisms by which RSV inhibits innate immune responses to enhance viral replication. Understanding the interplay between RSV proteins and innate immunity may lead to the development of more immunogenic RSV vaccine candidates. Additionally, Dr. Teng studies the interactions between cellular signal transduction machinery and viral proteins, with a view to discovering potential targets for antiviral therapy.
Jia-Wang Wang, Ph.D.

Dr. Jia-Wang Wang received an M.S. degree from Sichuan University, Chengdu, China and a Ph.D. degree from Wuhan University, Wuhan, China. He conducted postdoctoral research at the Chinese Academy of Medical Sciences, Beijing, China and at the Perelman School of Medicine, University of Pennsylvania, Philadelphia, PA. He then worked as a research associate at the H. Lee Moffitt Cancer Center & Research Institute, Tampa, FL, before he joined the Division of Allergy and Immunology. Dr. Wang is a member of the American Academy of Allergy, Asthma & Immunology (AAAAI) and an editorial board member of the *Austin Journal of Clinical Immunology*.

He has a strong background in genetics, immunology and molecular biology, discovered the lipopolysaccharide-responsive beige-like anchor (LRBA) gene, mutation of which causes immunodeficiency and autoimmunity, and contributed fifteen GenBank sequence entries. He has published more than 24 papers, some of which are in high profile journals including *Science*. He has more than fourteen conference abstracts including featured posters and oral presentations. He has three approved patents, one copyright computer program and three patents pending. Most of these intellectual properties are on microRNAs (miRNA), which have great promise as biomarkers and therapeutics for human diseases.

He also has extensive experience working with mouse models to study human diseases. He has successfully generated SHIP gene null and conditional knockout mouse models, two LRBA knockout mouse models, and four miRNA transgenic overexpression mouse models.
DIVISION OF ALLERGY AND IMMUNOLOGY

FELLOWS-IN-TRAINING (2012 to 2015)

Adam Updegraff, DO, 1st year fellow, is scheduled to graduate in June, 2015. Dr. Updegraff received his medical degree from Michigan State University College of Osteopathic Medicine in East Lansing, Michigan. He completed a combined residency in internal medicine and pediatrics from William Beaumont Hospital, Royal Oak, Michigan. Dr. Updegraff’s research interests include asthma, atopic dermatitis, and food allergy.

Michael Balduzzi, MD, 1st year fellow, is scheduled to graduate in June 2015. Dr. Balduzzi received his medical degree from New York Medical College, Valhalla, NY. He completed his residency in internal medicine at the University of South Florida Morsani College of Medicine, Tampa, FL where he also spent time in his final year of residency as one of the program’s chief resident. Dr. Balduzzi has research interest in local mucosal factors playing a role in chronic allergic and nonallergic rhinitis and improving asthma outcomes by controlling upper airway comorbid conditions.

Gregory Cowan, MD, 2nd year fellow, is scheduled to graduate in June, 2014. Dr. Cowan received his medical degree from the University of South Florida Morsani College of Medicine, Tampa, Florida. He completed his residency in pediatrics at Phoenix Children’s Hospital/Maricopa Medical Center, Phoenix, Arizona. His research interest in allergy & immunology involves primary immune deficiencies and asthma in children. His long term goal is to improve patient and parent understanding of how to manage asthma and improve long-term outcomes.

Nauman Salim, MD, 2nd year and chief fellow, is scheduled to graduate in June, 2014. Dr. Salim received his medical degree from Allama Iqbal Medical College, Lahore, Pakistan. He completed his residency in internal medicine and pediatrics from Indiana University School of Medicine, Indianapolis, Indiana. His research interest includes blood and bone marrow transplantation with emphasis on graft versus host disease. His long term goal is to practice allergy and immunology and provide immunologic expertise for patients undergoing stem cell transplantation.
RESEARCH STAFF MEMBERS

Lakshmi Galam, Ph.D., Research Instructor
Jutaro Fukumoto, M.D., Ph.D., Postdoctoral Fellow
Kunyu Li, Biological Scientist
Kim Teng, Senior Biological Scientist
Michelle Reiser, M.S., Research Technician
Eileen Rifkin, Research Technician

STUDENTS AND VISITING RESEARCH SCHOLARS

Ruan Cox, Graduate Student
Sara Garcia, Research Student
Matthew Ho, Research Student
Amanda Hodgkins, Research Assistant/Student
Bao Huynh, Research Assistant/Student
Michelle Kaminsky, Research Assistant/Student
Tucker Maute, Research Student
Anu Stephen, Research Student
Tran Tran, Graduate Student
Jillian Whelan, Graduate Student
Olivia Smith, Volunteer Research Student
Bangmei Wang, Volunteer Research Student
Rebekah Cook, Volunteer Research Student

ADMINISTRATIVE PERSONNEL

* Peggy Hales, Program Assistant
* Rebecca Carter, Administrative Secretary
* Geeta Gehi, Administrative Secretary
* Also James A. Haley Veterans’ Hospital, Tampa, FL
CLINICAL RESEARCH UNIT

Carlotta Riner, CRC, Clinical Research Administrator
Shawna Ogilbee, CRC, Unit Manager
Jeaneen Ahmad, BA, ALA Coordinator
Sarah Croker, RN, CRC, BA, Clinical Research Coordinator
Rebecca McCrery, AS, Regulatory Specialist, Coordinator

ALL CHILDREN’S HOSPITAL

Amy Baldwin, Administrative Assistant to Dr. Mark Ballow
Marjorie Peak, Pediatric Allergy and Immunology Fellowship Coordinator
III.  JOY MCCANN CULVERHOUSE
AIRWAY DISEASE RESEARCH CENTER

A.  Basic Research Projects

1.  Akt as a therapeutic target for respiratory syncytial virus (PI: Teng)

Two aspects of Akt activity in RSV infection are being explored. First, RSV phosphoprotein has been identified as a target for Akt phosphorylation and the role of this phosphorylation on viral replication is under investigation. Second, the process of understanding the viral factors responsible for Akt activation during RSV infection is being researched. These studies are being done in collaboration with Dr. Biao He (University of Georgia, Athens, Georgia).

2.  Structural determinants of NS2 for pathogenic functions (PI: Teng)

We have previously published that NS2 blocks interferon induction by binding to RIG-I. In addition NS2 appears to have additional functions associated with viral pathogenesis, including NFκB induction and STAT2 degradation. Trying to separate these activities by mutagenesis to understand how NS2 accomplishes each function is under investigation. The focus is on differentially altering the functions to develop an attenuated RSV vaccine candidate that maintains its immunogenicity.

3.  Enhancing immunogenicity of RSV vaccines by altering NS1 function (PI: Teng)

This is part of a program project in collaboration with Dr. Mark Peeples (Nationwide Children’s Hospital, Columbus, OH). The hypothesis that decreasing the ability of NS1 to inhibit interferon responses can enhance the immunogenicity of RSV vaccine candidates is being researched.

4.  Mechanism of RSV temperature sensitivity due to cis-acting sequences (PI: Teng)

Previous studies show that a single nucleotide change in the M2 transcription start sequence is sufficient to confer temperature sensitivity to recombinant RSV. The mechanism is being investigated by which this mutation affects RSV replication and transcription at non-permissive temperatures.

5.  RSV M protein trafficking and virus assembly (PI: Teng)

This is a long-term collaboration with Drs. David Jans and Reena Ghildyal (Monash University, Melbourne, AUS) to determine the role of M protein trafficking in RSV morphogenesis and the importance of nuclear translocation in M function.
6. **Effect of NALP-3 inflammasome on epithelial permeability (PI: Kolliputi)**

Our previous reports demonstrate the inflammasome, a proinflammatory cytokine processing complex, plays an important role in the production of early inflammatory cytokines associated with edema. Ceramide is a critical mediator of pulmonary edema, however the ability of ceramide to activate the inflammasome has not been elucidated. Utilizing macrophages in vitro, we discovered that ceramide induced inflammasome activation results in significant cytokine secretion. Genetic silencing of inflammasome components abolished the ability of ceramide to induce inflammasome activation, and a rescue of the barrier integrity of alveolar epithelial cell (AEC) in co-culture was observed. These novel results reveal that ceramide induced cytokine secretion and AEC permeability occurs through an inflammasome dependant mechanism.

7. **Effect of microRNA 16 on epithelial sodium channel in human alveolar epithelial cells (PI: Kolliputi)**

Removal of edema from the air spaces of the lung is a critical function of the epithelial sodium channel (ENaC) and also involves the serotonin (5HT) transport system. Recent studies suggest that microRNA-16 (miR-16) targets the serotonin transporter (SERT). However, the role of miR-16 on its targets SERT and ENaC have not been studied. The expression patterns of miR-16, SERT, ENaC and serotonin are being investigated in mice exposed to room air and hyperoxia. The effects of miR-16 overexpression are being observed in vitro. MiR-16 and ENaC down regulation in mice exposed to hyperoxia correlates with an increase in SERT expression and pulmonary edema. Overexpression of miR-16 in alveolar epithelial cells suppresses SERT and increases ENaCβ levels. These data suggest that miR-16 upregulates ENaC, a major sodium channel involved in resolution of pulmonary edema in acute lung injury (ALI).

8. **Role of enhancer of zeste homolog 2 on pulmonary artery smooth muscle cell proliferation (PI: Kolliputi)**

Pulmonary arterial hypertension (PAH) is characterized by excessive proliferation of the pulmonary arterial smooth muscle cells (PASMCs). EZH2 regulates cancer cell proliferation; however, the role of EZH2 in the proliferation of PASMCs is not clear. Therefore, the expression of EZH2 is being investigated in normal and hypertensive mouse PASMCs. The effects of EZH2 overexpression on the proliferation of human PASMCs also are being tested. EZH2 protein expression in mouse PASMCs correlates with decreased right ventricular function. The overexpression of EZH2 in human PASMCs enhances proliferation and migration and decreases the rate of apoptosis. EZH2 transfected cells demonstrated an increase in proliferation and a significant decrease in apoptosis. These findings show that EZH2 plays a role in the migration and proliferation of PASMCs. More importantly, EZH2 may serve as a potential target for new therapies for PAH.
9. **Effect of mir-206 on pulmonary artery smooth muscle cell proliferation and differentiation (PI: Kolliputi)**

Pulmonary arterial hypertension (PAH) is a progressive devastating disease characterized by excessive proliferation of the pulmonary arterial smooth muscle cells (PASMCs). MicroRNA-206 (miR-206) is known to regulate proliferation however, the role of miR-206 in PAH has not been studied. Therefore, the expression patterns of miR-206 are being investigated in normal and hypertensive mouse PASMCs. The effects of miR-206 on cell proliferation, apoptosis and smooth muscle cell marker expression in human PASMCs also are being measured. MiR-206 expression in mouse PASMCs correlates with an increase in right ventricular systolic pressure. Reduction of miR-206 levels in hPASMCs causes increased proliferation and reduced apoptosis. These results suggest that miR-206 is a potential regulator of proliferation, apoptosis and differentiation of PASMCs, which could yield a novel treatment strategy in PAH.

10. **MicroRNAs as biomarkers and therapeutics for asthma (PI: Wang)**

MicroRNAs (miRs) are ~22 nucleotides long non-coding RNAs that inhibit mRNA translation by the base pairing rule at the accuracy of one base. It is believed that most human genes and the entire spectrum of biological pathways are tightly and delicately controlled by the miRNome. Deregulation of miRs may contribute to various diseases. The mechanism underlying miR regulations of immunity is under investigation. Developing miR biomarkers and therapeutics for inflammatory diseases, such as asthma, using cell culture and mouse models are goals of this research.


Lipopolysaccharide (LPS)-responsive beige-like anchor (LRBA) is a novel gene essential for the normal function of the immune system. It is the eighth common variable immunodeficiency (CVID) gene, mutation of which causes CVID and autoimmunity. LRBA is involved in some critical cellular processes such as cell proliferation, apoptosis and autophagy. It may interact with multiple important signal transduction pathways. The molecular mechanism by which LRBA regulates the immune system is being explored.

**B. Clinical Research Projects**

1. **Effects of pine cone extract on IgE levels in patients with allergic rhinitis (PI: Ledford)**

Pine cones and their aqueous extracts (PCE) were thought to have medicinal properties as far back as 2000 years ago in Japanese populations. Anecdotal reports suggest that the use of PCE improves allergic rhinitis symptoms; it significantly reduces serum IgE levels in mouse models. The purpose of this study is to determine if oral PCE extract administered in a double-blind fashion significantly reduces IgE levels in patients with evidence of perennial allergic rhinitis.
2. Pollen and mold counts and immunochemical quantification of outdoor allergens (PI: Glaum)

Particles other than pollen which transport aeroallergens have been described. The Division, which houses the Pollen and Mold Counting Station for Tampa, has two collectors adapted to collect both pollen and pollen aeroallergens. The collectors are located on the roof of the James A. Haley V.A. Medical Center Research Building. Pollen counts are performed twice weekly, disseminated to local media once weekly and to the Internet twice weekly. Dr. Mary Jelks reads and interprets the slides. The purpose of this study is to determine whether or not there are molecular methods by which aeroallergens can be quantitated and compare them to present day standards assessing pollen counts.

3. Prevalence of food allergy in adult patients with eosinophilic esophagitis (PI: Ledford)

Food allergies are known to play a significant role in children with eosinophilic esophagitis. Little is known about the prevalence of food allergies in adult patients with this disease. The purpose of this study is to determine the prevalence of food allergies in a cohort of adult patients with eosinophilic esophagitis, results of which will be compared to findings in adult patients with gastroesophageal reflux disease.

4. Effect of oxymetazoline hydrochloride in combination with nasal glucocorticoid on the apnea hypopnea index (AHI) (PI: Lockey)

“Nocturnal Oxyhemoglobin Saturation, Snoring, and Sleep Quality in Subjects with Persistent Nasal Congestion. A Double Blinded, Placebo Control, Cross Over Prospective Trial” is designed to evaluate the effectiveness on the apnea/hypopnea index (AHI) of adding oxymetazoline to intranasal mometasone, and on other sleep parameters. The study will be carried out in subjects with persistent nasal congestion secondary to allergic or non-allergic rhinitis despite treatment with the highest recommended doses of intranasal mometasone.

5. A study of function of respiration and cognition in the elderly (The FORCE study) (PI: King)

The purpose of this study is first, to determine which diagnostic tool, impulse oscillometry (IOS), a measurement of respiratory impedance using sound impulses or spirometry, is more feasible for the elderly population to measure pulmonary function; second, to determine if cognitive performance is related to the ability to perform spirometry and IOS; and third, to determine the relationship between biomarkers, cognitive performance, and asthma diagnosis.

6. Evaluation of calcium and vitamin D intake in children on inhaled or intranasal corticosteroids compared to normal children (PI: Lockey)

The specific aims of this project are to evaluate the dietary calcium intake of asthmatic children (4-17 years) who are receiving long-term treatment with inhaled or intranasal corticosteroids versus healthy controls using a validated food frequency questionnaire.
7. **Identification of plasma miRNAs as potential biomarkers in asthma exacerbation (PI: Lockey)**

It is hypothesized that there is a statistically significant difference in miRNA profiling and expression in subjects with asthma exacerbation compared to a baseline level or following effective treatment of an exacerbation of asthma. Therefore, plasma miRNA profiling may provide highly specific and sensitive biomarkers for asthma exacerbation detection and treatment follow-up.

8. **Differences in mold counts from January 1995 to December 2011 (PI: Lockey)**

Allergic diseases are due to complex interactions between genetic and environmental factors. Airborne mold and pollens are known to trigger allergic respiratory disease in sensitive individuals. Yet little is known about possible changes as related to climate change in pollen and mold counts over the last 16 years. Daily pollen and fungal spores sample data between January 1995 to December 2010 are available for Sarasota, FL. These data will be compared to weather data for Sarasota available from the National Climatic Data Center. The objectives of this study are to determine if changes in pollen/mold counts can be correlated with climate changes.

C. **Clinical Research Unit (CRU)**

The University of South Florida, Asthma, Allergy and Immunology CRU was established in 1977 to improve the treatment of patients who suffer from asthma, allergic and immunologic diseases.

The CRU provides quality research in a variety of clinical areas which include the following: allergic conjunctivitis; allergen immunotherapy; allergen skin testing; allergic rhinitis; asthma; atopic eczema; bronchitis, acute and chronic; contact dermatitis; chronic obstructive pulmonary disease; exercise induced asthma; headache (migraine and tension); hereditary angioedema; immunodeficiency diseases; insect allergy; intravenous immunoglobulin; nasal polyps; sinusitis, acute and chronic; temporomandibular joint disease; urticaria and vasomotor rhinitis.

Studies funded by pharmaceutical companies are conducted at the Division's CRU. The CRU is also a member of the American Lung Association’s Asthma Clinical Research Center network, one of 19 centers throughout the United States.
IV. BASIC AND CLINICAL RESEARCH SUPPORT

ENDOWMENTS

Joy McCann Culverhouse Endowment and Chair in Allergy and Immunology
Mabel and Ellsworth Simmons Professorship for Asthma Research

EXTRAMURAL FUNDING

Government Funding
National Institutes of Health (NIAID)
National Heart, Lung and Blood Institute

Non-Profit Funding
American Lung Association
American Heart Association
American Academy of Allergy Asthma and Immunology, ARTrust

Pharmaceutical Funding (past or present)
Ablynx, NV
Almirall Pharmaceuticals
CSL Behring
Cytos
Dyax Corporation
Forest Laboratories
Genentech Inc
GlaxoSmithKline
Jerini, US
Merck and Co., Inc.
MedImmune
Novartis Pharmaceuticals
Pfizer
Pharming Inc.
Roche
Sanofi-Aventis Pharmaceuticals
Schering-Plough Corporation
Shire
Teva Pharmaceuticals
Viropharma
V. PUBLICATIONS FROM THE DIVISION

BOOKS PUBLISHED OR IN PRESS


BOOK CHAPTERS PUBLISHED OR IN PRESS


**SCIENTIFIC ARTICLES PUBLISHED OR IN PRESS**

Lockey RF (Contributor): Writing Committee for the American Lung Association Asthma Clinical Research Centers: Lansoprazole for children with poorly controlled asthma. *JAMA* 2012; 307(4): 373-381.


**REVIEW ARTICLES/EDITORIALS PUBLISHED OR IN PRESS**


Stokes J, Casale TB: The relationship between IgE and allergic disease. Up To Date, in press.

INTERNET PUBLICATIONS


VI. FACULTY AND STAFF AWARDS

Richard F. Lockey, MD
“Member of Honor” award, presented at the Adjuvant & Allergen Vaccines 2012 International Conference by the Cuban Society of Immunology, Havana, Cuba, May 10, 2012

Richard F. Lockey, MD
Southern Medical Society, Dr. Robert D and Alma W. Moreton Original Research Award, 2012.

Dennis K. Ledford, MD
Mabel & Ellsworth Simmons Professor of Allergy and Immunology, 2012

Dennis K. Ledford, MD
Listed as “Most Compassionate Doctor” – Vitals, 2012

Kolliputi, Narasaiah, PhD
WAO Outstanding Abstract Award
XXIII World Allergy Organization International Scientific Congress (WISC), Hyderabad, India, December 6–9, 2012

Kolliputi, Narasaiah, PhD
World Allergy Organization (WAO) Junior Member Travel Award
Hyderabad, India, December, 2012

Dennis K. Ledford, MD
Elected to the Board of Directors, 2013
Joint Council of Allergy Asthma and Immunology

Dennis K. Ledford, MD
President of Medical Faculty, 2012-2013
University of South Florida Morsani College of Medicine

Dennis K. Ledford, MD
“25 Years of Service” Award
Tampa General Hospital, 2013

Mark C. Glaum, MD
American Academy of Allergy Asthma and Immunology (AAAAI)
ARTrust Mini Grant, February 25, 2013.
VII. VISITING PROFESSOR EDUCATIONAL PROGRAM

Adam Wanner, MD, Joseph Weintraub Professor of Medicine, University of Miami Miller School of Medicine, Division of Pulmonary & Critical Care Medicine, Department of Medicine, Miami, Florida. “The Airway Circulation in Asthma”, February 16, 2012

Blanca Camoretti-Mercado, PhD, Department of Medicine, Section of Pulmonary & Critical Care, University of Chicago, Chicago, IL. “Airway Smooth Muscle and Inflammation: Beyond Contraction”, August 2, 2012

Prof. Cezmi A. Akdis, MD, Director, Swiss Institute of Allergy and Asthma Research, Davos, Switzerland. “Tissue Factors that Play a Role in Chronicity and Immune Regulation in Asthma”, September 24, 2012

Hendrik Nolte, MD, PhD, Senior Director, Respiratory and Immunology, Merck Research Laboratories, Kenilworth, New Jersey. “What is the Future of Sublingual Immunotherapy In North America?”, July 25, 2012

Lewis J. Smith, MD, Professor, Pulmonary and Critical Care Medicine, Northwestern University Feinberg School of Medicine, Chicago, Illinois. “The American Lung Association Asthma Clinical Research Network – Accomplishments and Future Studies”, November 15, 2012

Rosa Codina, PhD, Senior Research Scientist, Greer Laboratories, Inc. Lenoir, NC; Clinical Assistant Professor of Medicine, University of South Florida Morsani College of Medicine, Division of Allergy and Clinical Immunology, Department of Internal Medicine, Tampa, FL. “An Introduction to Fungal Allergenic Extracts” and “A Review of a Few Questions to Prepare for the Board Exam”, November 29, 2012

Robert Q. Lanier, MD, Executive Medical Director, American College of Allergy, Asthma & Immunology; Clinical Professor of Pediatrics, University of North Texas Health Science Center, Fort Worth, Texas. “Steroid Footprint: the Cumulative Effects of Steroids in Atopic Patients”, October 10, 2013.

Bruce D. Levy, MD, Associate Professor of Medicine, Harvard Medical School; Associate Physician, Brigham and Women’s Hospital, Boston, Massachusetts. “Pre-resolving mediators in acute lung injury and inflammation”, October 11, 2013.

Dr. Jose Laerte Junior Boechat Morandi, Adjunct Professor, Section of Clinical Immunology, Department of Internal Medicine, Universidade Federal Fluminense, Rio de Janeiro, Brazil. “Eosinophilic Esophagitis”, November 14, 2013.