# David G. Meckes, Jr.

Department of Biomedical Sciences

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Research Interests: Virus-host interactions

Extracellular vesicles (exosomes and microvesicles)

Cancer cell biology, signal transduction, vesicular trafficking

Proteomics and protein-protein interactions Mechanisms of protein transport and targeting

**Education:** 

2003-2008 The Pennsylvania State University

**College of Medicine** 

Hershey, PA

Doctor of Philosophy

Microbiology and Immunology

1999-2003 University of the Sciences in Philadelphia

Philadelphia, PA

Bachelor of Science (cum laude)

Major: Microbiology

Positions:

2013-present **Assistant Professor** 

Florida State University College of Medicine

Department of Biomedical Sciences

2009-2013 Postdoctoral Fellow

> University of North Carolina at Chapel Hill Lineberger Comprehensive Cancer Center

### Selected Academic Awards and Honors:

2015	Outstanding Junior Faculty Investigator, FSU College of Medicine
2012	Priscilla Schaffer Award for top oral presentation by a postdoctoral fellow at the
	International Herpesvirus Workshop
2011	Joseph S. Pagano Award for top paper by a postdoctoral fellow
2011-2013	American Cancer Society Postdoctoral Fellowship (PF-11-158-01-MPC)
2011	NCI F32 Postdoctoral Fellowship– Awarded (declined)
2009-2011	UNC Cancer Center T32 Postdoctoral Training Fellowship from the NCI
2009	Enders' Award for Scholarly Research Achievement
2008	The 33rd International Herpesvirus Workshop Travel Award (for top oral
	presentation)
2006-2008	NCI T32 Predoctoral Training Fellowship: Viruses and Cancer
2003	Dr. Louis Gershenfeld Memorial Prize for Excellence in Microbiology
2001-2003	NETS Science Scholarship
1999-2003	Dean's List (7 semesters)
1999-2003	L. N. P. Rudolph Academic Scholarship
1999	Alpha Lambda Delta National Academic Honor Society

## Research Experience:

May 2009-April 2013 University of North Carolina at Chapel Hill

**Lineberger Comprehensive Cancer Center** 

Chapel Hill, NC Postdoctoral Scholar

Laboratory of Nancy Raab-Traub, Ph.D.

Study of the content and function of exosomes released from Epstein-Barr virus infected cancer cells

2008- May 2009 The Pennsylvania State University College of Medicine

**Hershey, PA**Postdoctoral Fellow

Laboratory of John W. Wills, Ph.D.

Mechanisms of UL16 Packaging into Herpes Simplex Virus

2004-2008 The Pennsylvania State University College of Medicine

Hershey, PA

Doctoral Thesis Research

Laboratory of John W. Wills, Ph.D.

The Dynamic Herpesvirus Tegument: Receptor Binding Induced Release of UL16 from the Capsid of Herpes

Simplex Virus

2001-2003 The University of the Sciences in Philadelphia

Philadelphia, PA

Undergraduate Research

Laboratory of James R. Johnson, Ph.D.

Mutagenesis and Penicillin Selection for Methionine Auxatrophs of Staphylococcus Aureus to Study its Methionine

**Biosynthesis** 

## Teaching Experience:

2015-present Co-Course Director

The Immune Response to Infection and Cancer

Delivered lectures and facilitated small group discussions of primary research articles

Florida State University College of Medicine

2014-present **Assistant Course Director** 

Medical Microbiology

Delivered lectures and facilitated small group clinical case studies for medical students

Florida State University College of Medicine

2013-2014 Content Expert and Small Group Facilitator

Medical Microbiology

Delivered lectures and facilitated small group clinical case studies for medical students

Florida State University College of Medicine

2014-present Content Expert

Chromatin Structure, Epigenetics & Human Health

Delivered lectures on cancer epigenetics and facilitated group discussions

Florida State University College of Medicine

2013-present **Directed Independent Studies – Cancer Biology** 

Trained undergraduate students in laboratory science and directed their research studies

Florida State University College of Medicine

2009-2013 Graduate and Undergraduate Research Advisor

Trained undergraduate students in laboratory science and managed their projects and experiments

Supervised and trained graduate students

University of North Carolina

2005-2009 Graduate Research Assistant

Trained and supervised junior graduate students in laboratory science

The Pennsylvania State University College of Medicine

2005-2007 Graduate Teaching Assistant

Medical Microbiology Laboratory for Medical Students The Pennsylvania State University College of Medicine

2001-2003 Teaching Assistant

Microbiology Laboratory

University of the Sciences in Philadelphia

# Publications (17) h-index (13)

http://scholar.google.com/citations?user=n9llb6AAAAAJ&hl=en&oi=ao https://www.researchgate.net/profile/David Meckes

Hurwitz, S.N., M.M. Conlon, M.A. Rider, **D.G. Meckes, Jr.** 2016. Nanoparticle analysis sheds budding insights into genetic drives of extracellular vesicle biogenesis. *J. Extracell. Vesicles.* **5**:31295

Rider, M.A., S.N. Hurwitz, **D.G. Meckes, Jr.** 2016. ExtraPEG: A Polyethylene Glycol-Based Method for Enrichment of Extracellular Vesicles. *Sci. Rep.* **6**:2397.

Meckes, D.G., Jr. 2015. Exosomal Communication Goes Viral. J Virol. 10:5200-3. Epub 2015 March 4 (Listed on JV most read articles for March and April)

**Meckes, D.G., Jr.** 2014. Affinity purification combined with mass spectrometry to identify herpes simplex virus protein-protein interactions. *Methods Mol. Biol.* 1144:209-22.

Meckes, D.G., Jr., H. P. Gunawardena, R.M. Dekroon, P.R. Heaton, R. Hood Edwards, S. Ozgur, J.D. Griffith, B. Damania, N. Raab-Traub. 2013. Modulation of B-cell exosome proteins by gamma herpesvirus infection. *Proc Natl Acad Sci USA*. 31:2925-33. Cited 61 times Special Commentary: Pegtel, D.M., Oncogenic Herpesviruses sending mixed signals. *Proc Natl Acad Sci U S A*. 31:12503-4

**Meckes, D. G., Jr.,** N.F. Menaker, and N. Raab-Traub. 2013. Epstein-Barr Virus LMP1 Modulates Lipid Raft Microdomains and the Vimentin Cytoskeleton for Signal Transduction and Transformation. *J Virol*. 3:1201-11. Epub 2012 Nov 14

Meckes, D.G., Jr. and N. Raab-Traub. 2011. Mining Epstein-Barr Virus LMP1 Signaling Networks. J Carcinogene Mutagene.

**Meckes, D.G., Jr.** and N. Raab-Traub. 2011. Microvesicles and Viral Infection. *J Virol.* **24**:12844-54. Epub 2011 Oct 5. Cited 148 times (Listed on JV most read articles for October through April)

P.C. Yeh, J. Han, P. Chadha, **D.G. Meckes, Jr.,** M.D. Ward, O.J. Semmes, J.W. Wills. 2011. Direct and Specific Binding of the UL16 Tegument Protein of Herpes Simplex Virus to the Cytoplasmic Tail of Glycoprotein E. *J Virol.* **18**:9425-36. Epub 2011 Jul 6.

Han, J., P. Chadha, **D.G.**, **Meckes**, **Jr.**, N.L. Baird, J.W. Wills. 2011. Interaction and Interdependent Packaging of Tegument Protein UL11 and Glycoprotein E of Herpes Simplex Virus. *JVirol*. **18**:9437-46. Epub 2011 Jul 6.

\*Kung, C.P., \***D.G. Meckes, Jr.,** and N. Raab-Traub. 2011. EBV Latent Membrane Protein 1 (LMP1) Activates STAT3 and ERK through effects on EGFR and PKCδ. *JVirol.* **9**:4399-408. Epub 2011 Feb 9. \*These authors contributed equally to the work. (Journal of Virology Spotlight Article)

**Meckes, D.G., Jr.,** A.R. Marquitz, K.H. Shair, R.H. Edwards, C.P. Kung, and N. Raab-Traub. 2010. A Human Tumor Virus Utilizes Exosomes for Intercellular Communication. *Proc Natl Acad Sci U S A.* 47:20370-5. Epub 2010 Nov 8. Cited 219 times

Cited in: *Nature* Research Highlights. "Communicators key for cancer virus." Vol. 468, Page:349. 2010 Nov 18.

Science Daily, "Cellular Communicators for Cancer Virus Identified." (Nov 9, 2010); also featured in 10 other science/medical news outlets.

Priority Paper Evaluation: Middeldorp J.M., D.M. Pegtel. 2011. A human tumor virus extends its reach. Future Virol. 4:413-415.

**Meckes, D.G., Jr.**, J.A. Marsh and J.W. Wills. 2010. Complex Mechanisms for the Packaging of the UL16 Tegument Protein into Herpes Simplex Virus. *Virology.* **2**:208-213. Epub 2010 Jan 3.

\*Harper, A.L., \*D.G. Meckes, Jr., \*J.A. Marsh, N.L. Baird, P.C. Yeh, C.B. Wilson, and J.W. Wills. 2009. Interaction Domains of the UL16 and UL21 Tegument Proteins of Herpes Simplex Virus. *J Virol.* **6**:2963-71 Epub 2009 Dec 30. \*These authors contributed equally to the work.

**Meckes, D.G., Jr.** and J.W. Wills. 2008. Structural Rearrangement within an Enveloped Virus upon Binding to the Host Cell. *J Virol.* **21**:10429-35. Epub 2008 Aug 20.

(Journal of Virology Spotlight Article) (Selected for Faculty of 1000 Biology, must read)

Yeh, P.C., **D.G. Meckes**, **Jr.**, and J.W. Wills. 2008. Analysis of the Interaction between the UL11 and UL16 Tegument Proteins of Herpes Simplex Virus. *J Virol*. **21**:10693-700. Epub 2008 Aug 20.

**Meckes, D.G., Jr.** and J.W. Wills. 2007. Dynamic Interactions of the UL16 Tegument Protein with the Capsid of Herpes Simplex Virus. *J Virol.* **23**:13028-36. Epub 2007 Sep 12.

#### Service:

## **Ad Hoc Reviewer**

Journal of Virology

Journal of General Virology

Proceedings of the National Academy of Sciences (PNAS)

PLOS One Methods

BMC Cancer

**BioMed Research International** 

NIH Study Section ZDA1 JXR-G (13) – Special Emphasis Panel, "Extracellular Vesicles in HIV/AIDS and Substance Abuse" (R01, R21)

Danish Council for Independent Research Medical Research Council, United Kingdom

Ghent University

Alzheimer's Association

## **University Committees**

2015-present Fulbright Faculty Committee

## **Departmental and College Committees**

2016 Faculty Council Outstanding Junior Investigator Award Committee
2016 Faculty Council Outstanding Junior Educator Award Committee
2016 Curriculum Committee Review of Pre-clerkship Course

2015-present Curriculum Committee Year 1 and 2
2015-present By-Laws and Policy Committee
2015-present Faculty Recruitment Committee
2015 Graduate Program Committee
2015 Promotion and Tenure Committee

2013-present Curriculum Redesign Committees - Medicine 2, Host Defense, and Hematology teams 2014-present Speaker and organizer for BMS Public Information Session "Current Health Issues"

2014 Faculty Development and Mentoring Committee

#### **Volunteering**

2015-present "Science Night" Gilchrist Elementary School – organized science experiments and educational activities for students
2014-present First Friday's at Railroad Square "Ask a Scientist" – A monthly event organized by FSU Professors to promote science in

the community

Fundraising for the Muscular Dystrophy Association

2013 Panel Speaker for FSU Postdoc Symposium session "Challenges Facing Junior Scientists in the 21st century."

## Society Memberships:

International Society for Extracellular Vesicles

American Society for Exosomes and Microvesicles

American Society for Microbiology

International Association for Research on Epstein-Barr virus and Associated Diseases

# Extramural Funding:

## Current

1RO1CA204621- National Institutes of Health 4/1/2016 - 3/31/2021

National Cancer Institute Total Award: \$1,704,214 Title: Modulation of Host Cell Exosome Content and Function by EBV LMP1

Role: PI

The overall goal of these studies is to determine the mechanisms that LMP1 drives exosome content reorganization and alters the functions of exosomes. We hypothesize that LMP1 exosomal trafficking modulates the components and biological properties of exosomes by altering

endocytic routes and membrane microdomains. To test this, we aim to: 1.) investigate the mechanism through which LMP1 alters exosome components; 2.) determine the functions of LMP1-modified exosomes in intracellular communication and cellular transformation.

1R15CA188941- National Institutes of Health 5/1/2015 – 4/30/2018

National Cancer Institute Total Award: \$414,462

Title: Exosome-Dependent Trafficking of Epstein-Barr Virus LMP1

Role: PI

The overall goal of this project is to uncover the molecular basis controlling LMP1 exosome trafficking and release from the cell. We hypothesize that LMP1 traffics from the Golgi to the site of exosome formation (mutltivesicular bodies, MVBs) in Rab31/VAMP4 containing vesicles in complex with CD63. To test this hypothesis, we have proposed two focused specific aims that are appropriately tailored for the funding period and total award amount for the R15 funding mechanism. Using dominant negative constructs and siRNAs directed against Rab31 and VAMP4 we will monitor the localization of LMP1 to MVBs and exosomes using western blot and fluorescence-based assays. In this study, we will also further investigate the CD63-LMP1 interactions important for exosomal trafficking.

6AZ11 – Alzheimer's Research Program 1/1/2016 – 1/31/2018 Florida Department of Health Total Award: \$81,499

Title: Blood Exosomes and Neurodegenerative Disease

Role: Pl

Our overall hypothesis is that pathological factors that contribute to the course of AD and can be used for early detection of the disease are resident in brain derived exosomes of the circulation. Therefore, our objects for this research proposal are twofold - 1) to develop techniques for identifying the tissue origins of circulating exosomes; and 2) to characterize specifically the neuronal exosomes present in human blood samples the blood from mouse models of AD.

## Completed

 $\begin{array}{lll} 4BB05 - Bankhead\text{-}Coley \ Program & 12/1/2013 - 11/30/2015 \\ Florida \ Department \ of \ Health & Total \ Award: \ \$396,328 \\ Title: \ Proteomic \ Analysis \ of \ Cancer \ Exosomes \ for \ Diagnostic \ and \ Therapeutic \ Targets \end{array}$ 

Role: PI

The major goal of this project is to utilize exosome purification strategies that we have developed together with advanced quantitative proteomics techniques to define the protein composition of exosomes secreted from a diverse set of human cancer cell lines (the National Cancer Institute, NCI-60). The completion of this project will reveal a common set of proteins found in cancer exosomes that are likely important for their formation and function. Exosomal proteins expressed only in specific cancer types (e.g., breast, prostate and colon) may represent potential diagnostic biomarkers that will be further explored with patient samples. Overall, this project aims to understand the composition and function of exosomes secreted from cancer cells with the goal of discovering novel therapeutic and diagnostic targets.

2011-2013: American Cancer Society Fellowship

Title: "Molecular Properties of Exosomes Secreted from Cancer Cells Expressing LMP1"

Total Award: \$150,000

# Intramural Funding:

1/9/15-1/8/16: FSU Equipment and Infrastructure Enhancement Grant

Title: "Enhancing Biomedical Research by Confocal Microscopy with Quantitative Capability"

Total: \$85,000 (Co-PI)

6/01/14-5/30/15: FSU Equipment and Infrastructure Enhancement Grant

Title: "Enhancing Cell Biology Research Through Automated Cell Counting and Fluorescent-Based Assays"

Total: \$29,618 (PI)

### Abstracts:

Hurwitz, S.N., Nkosi, D., Conlon, M.M., **Meckes, Jr., D.G.** 2016 Tetraspanin protein CD63 mediates exosomal packaging of Epstein Barr virus LMP1. Keystone Symposia Exosomes/Microvesicles: Novel Mechanisms of Cell-Cell Communication. Keystone, CO. Poster Presentation

Hurwitz, S.N., Conlon, M.M., Rider, M.A., Brownstein, N.C., **Meckes, Jr., D.G.** 2016. Nanoparticle tracking analysis of cancer cell vesicles sheds budding insights into exosome and microvesicle biogenesis. Keystone Symposia Exosomes/Microvesicles: Novel Mechanisms of Cell-Cell Communication. Keystone, CO. Poster Presentation

Hurwitz, S.N., Nkosi, D., Conlon, M.M., **Meckes, Jr., D.G.** 2016 Tetraspanin protein CD63 mediates exosomal packaging of Epstein Barr virus LMP1. 41st International Herpesvirus Workshop. Madison, WI. Poster Presentation

Nkosi, D., Howell, L.A., Conlon, M.M., Tremblay, D. C., **Meckes, Jr., D.G.** 2015. Transmembrane Domains Mediate Intra- and Extracellular Trafficking of Epstein-Barr Virus LMP1. 40<sup>th</sup> Annual International Herpesvirus Workshop. Boise, ID. Oral Presentation

Rider, M.A. and **Meckes, D.G., Jr.** The Interactome of the Epstein-Barr Virus Oncoprotein Evaluated by the Proximity-Based BioID Approach. 40<sup>th</sup> Annual International Herpesvirus Workshop. Boise, ID. Poster Presentation

**Meckes, D.G., Jr.,** H.P. Gunawardena, R.M., Dekroon, P.R. Heaton, R.H. Edwards, S. Ozgur, J.D. Griffith, B. Damania, and N. Raab-Truab. 2013. Modulation of B-Cell Exosome Proteins by Gammaherpesvirus Infection. 38<sup>rd</sup> Annual International Herpesvirus Workshop. Grand Rapids, MI. Oral Presentation

**Meckes, D.G., Jr.,** N. Menaker, R.H. Edwards, and N. Raab-Traub. 2012. Reorganization of Lipid Raft Microdomains by EBV Latent Membrane Protein 1 (LMP1) Contributes to its Signaling and Transformation Capabilities. 37<sup>rd</sup> Annual International Herpesvirus Workshop. Calgary, Canada. Oral Presentation

Meckes, D.G., Jr., N. Menaker, R.H. Edwards, and N. Raab-Traub. 2012. Reorganization of Lipid Raft Microdomains by EBV Latent Membrane Protein 1 (LMP1) Contributes to its Signaling and Transformation Capabilities. International Congress on Oncogenic Herpesviruses and Associated Diseases. Philadelphia, PA. Poster Presentation

Meckes, D.G., Jr., A.R. Marquitz, K.H. Shair, R.H. Edwards, C.P. Kung, and N. Raab-Traub. 2010. Epstein-Barr Virus Utilizes Exosomes for Intercellular Communication. International Association for Research on Epstein-Barr virus and Associated Diseases. Birmingham, England. Oral Presentation

**Meckes, D.G., Jr.** and J.W. Wills. 2008. Rearrangement of the Tegument upon Herpesviruses Binding to their Host Cells. 33<sup>rd</sup> Annual International Herpesvirus Workshop. Estoril, Portugal. Oral Presentation

**Meckes, D.G., Jr.** and J.W. Wills. 2008. Rearrangement of the Tegument upon Herpesviruses Binding to their Host Cells. 8<sup>th</sup> Annual Herpesvirus Symposium. Philadelphia, PA. Oral Presentation

**Meckes, D.G., Jr.** and J.W. Wills. 2007. Dynamic Interactions of the UL16 Tegument Protein with the Capsid of Herpes Simplex Virus. 32<sup>nd</sup> Annual International Herpesvirus Workshop. Asheville, NC. Poster Presentation

**Meckes, D.G., Jr.** and J.W. Wills. 2007. Dynamic Interactions of the UL16 Tegument Protein with the Capsid of Herpes Simplex Virus. 7<sup>th</sup> Annual Herpesvirus Symposium. Philadelphia, PA. Oral Presentation

**Meckes, D.G., Jr.,** P.C. Yeh, R.J. Courtney, and J.W. Wills. 2005. Localization and Virion Incorporation of the Herpes Simplex Type 1 Tegument Protein, UL16. International Union of Microbiological Societies. XIII International Congress of Virology. San Francisco, CA. Poster Presentation

#### **Invited Seminars:**

Modulation of the Components and Functions of Exosomes by Tumor Viruses. Florida State University 4th Annual Life Sciences Symposium. Feb. 13, 2014

Oncogenic herpesviruses modify the cargo and functions of exosomes released from host cells. Florida State University Institute of Molecular Biophysics Structural Biology/Biochemistry Seminar. Nov. 12, 2013

The Far-Reaching Effects of Epstein-Barr Virus Oncoprotein LMP1. The Pennsylvania State University, Department of Biochemistry and Molecular Biology. February 23, 2012.

The Far-Reaching Effects of Epstein-Barr Virus Oncoprotein LMP1. The University of Iowa College of Medicine, Department of Microbiology. January 30, 2012.

# Trainees (name, years, current position):

#### **Undergraduate**

Antonia Veltcheva, 2015-present Marius Kostelic, 2015-present Alexandra Dolan, 2015-present Timothy Bobroskie, 2014-present Natalie Marenghi, 2013-2014 (medical student, FSU College of Medicine)

#### Medical

Stephanie Hurwitz, 2014 Maria Raye Anne Ng, 2016

#### Graduate

Stephanie Hurwitz (MD/PhD), 2015-present Meghan Conlon, 2014-2016 Lauren Howell, 2013- 2016 Sara York, 2016-present

#### Postdoctoral

Mark Rider, 2013-present
Dingani Nkosi, 2013-present
Mujeeburahiman Cheerathodi, 2016-present

#### Research Scientist

Xia Lui, 2014-present

Deanna Tremblay, 2013-2014 (Postdoctoral recruiter, St. Jude Children's Research Hospital)

# **Graduate Student Committees (name, years, program)**

Alyssa Rolfe, 2016-present (Biomedical Sciences) Brittany Brewers, 2015-present (Biology) Stephanie Hurwitz, 2015-present (Biomedical Sciences) Meghan Conlon, 2015-2016 (Biomedical Sciences) Emily Lee, 2014-present (Biology) Siming Ma, 2014-present (Biology) Lauren Howell, 2014-2016 (Biomedical Sciences)

## **Undergraduate Honor Thesis Committees**

Allaura Sherman 2015 (Biology) Marius Kostelic 2016 (Chemistry)