

## BIOGRAPHICAL SKETCH

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NAME Buffy Sue Ellsworth, Ph.D		POSITION TITLE Associate Professor of Physiology	
eRA COMMONS USER NAME (credential, e.g., agency login) bellsworth			
EDUCATION/TRAINING (Begin with baccalaureate or other initial professional education, such as nursing, include postdoctoral training and residency training if applicable.)			
INSTITUTION AND LOCATION	DEGREE (if applicable)	MM/YY	FIELD OF STUDY
South Dakota State University, Brookings, SD	B.S.	08/95	Microbiology
South Dakota State University, Brookings, SD	B.S.	08/96	General Chemistry
South Dakota State University, Brookings, SD	M.S.	07/97	Biology
Colorado State University, Fort Collins, CO	Ph.D.	10/02	Cell and Molecular Biology
University of Michigan, Ann Arbor, MI	Postdoctoral	09/07	Human Genetics

### A. Personal Statement

Our long-term goal is to identify the genes that, when mutated, cause pituitary hormone insufficiency and identify the mechanism of action of those genes. To expand the molecular diagnoses for pituitary hormone insufficiency, we have focused on a family of transcription factors referred to as forkhead factors. Forkhead factors are essential for diverse developmental processes and mutations in the genes encoding these factors are responsible for a number of human developmental disorders. Specifically, we investigate the contribution of forkhead factors to pituitary development. During my doctoral training, I showed that the forkhead factor, FOXL2, is part of a transcriptional complex that binds the gonadotropin-releasing hormone receptor gene, which is expressed in gonadotrope cells. As a postdoctoral fellow at the University of Michigan Medical School, I performed developmental mouse studies to demonstrate that FOXL2 protein is present in the prospective anterior lobe of the developing pituitary gland beginning at e11.5 and continuing through adulthood in gonadotrope and thyrotrope cells. I found that FOXL2 stimulates expression of the  $\alpha$ -subunit gene in cell culture studies and by over-expression of *Foxl2* in transgenic mice. In my current position as associate professor at Southern Illinois University, I organized a research team, which demonstrated that the forkhead factor, *Foxp3*, is required for normal gonadotrope function and that *Foxp3* mice exhibit hypothyroidism. We have also shown that loss of *Foxd1* causes reduced *Lhb* expression and increased pituitary cell proliferation. Finally, we investigated the expression patterns of FOXO1 during pituitary development and in the adult. We found that FOXO1 is present in the nucleus of pituitary cells starting at e14.5 and is present in an increased number of cells in *p27* null embryos, suggesting an intriguing negative feedback mechanism. Furthermore, our studies show that mice in which we have excised the *Foxo1* gene from the pituitary gland early in development using a *Foxg1*-cre mouse (*Foxo1* <sup>$\Delta$ pit</sup>) exhibit a drastic reduction in the number of GH-immunoreactive cells at e16.5 and e18.5, suggesting that FOXO1 is important for somatotrope specification/function. My expertise in both forkhead transcription factor function and pituitary development places me in a unique situation to address the roles of forkhead factors in pituitary development.

### B. Positions and Honors

#### Positions and Employment

1993-1995	Undergraduate Researcher, Microbiology, South Dakota State University
1995-1997	Graduate Research Assistant, Biology, South Dakota State University
1997-2002	Doctoral Candidate, Cell and Molecular Biology, Colorado State University
2002-2007	Postdoctoral Fellow, Human Genetics, University of Michigan Medical School
2007-2013	Assistant Professor, Physiology, Southern Illinois University School of Medicine, Carbondale
2013-present	Associate Professor, Physiology, Southern Illinois University School of Medicine, Carbondale

## Other Experience and Professional Memberships

- 1999-present Member, Endocrine Society
- 2002 **Embryonic Stem (ES) Cell Training** – Two week course, University of Michigan Transgenic Animal Model Core. General care and maintenance of ES cells for the purpose of gene targeting.
- 2003 **Developmental Biology Gordon Conference**
- 2004 **Mouse Embryo Microinjection Training** – One week course, University of Michigan Transgenic Animal Model Core. Egg collection, microinjection of DNA into 1-cell fertilized eggs, and transfer of 1-2 cell embryos into pseudopregnant female mice.
- 2004-5, 11-13 Member, Society for the Study of Reproduction
- 2005 **NIH/NICHD Postdoctoral Fellows' Career Workshop**
- 2009-2012 Women in Endocrinology, Communications Committee
- 2012-2014 Women in Endocrinology, Communications Committee, Chair
- 2008 Ad hoc reviewer for Developmental Biology
- 2009 Ad hoc reviewer for Development
- 2009 Ad hoc reviewer for Federation of American Societies for Experimental Biology Journal
- 2010 Ad hoc reviewer for Biology of Reproduction
- 2010 Ad hoc reviewer for Molecular Endocrinology
- 2011 Ad hoc reviewer for Human Genetics
- 2011-13 Biology of Reproduction, Editorial Board
- 2012-13 Frontiers in Genomic Endocrinology, Editorial Board
- 2011 Chair of Endocrine Society Symposium, "New Aspects of Signaling & Secretion in the Gonadotrope"
- 2012 Society for the Study of Reproduction, Program Committee, Module Session co-chair: Reproductive Endocrinology
- 2013 Chair of Endocrine Society Symposium, "Neuroendocrinology and Pituitary - Basic"

## Honors

- 1994 EPSCoR ADP Summer Research Stipend for Undergraduate Women and Minority Students
- 1996 Third Place in Raymond A. Moore Biostress Research Poster Contest
- 1996 Inducted into Gamma Sigma Delta -- The Honor Society of Agriculture
- 1997 Semi-Finalist in the SDSU Chapter of the Society of Sigma Xi 1997 Research Paper Award
- 1999 Colorado State University Graduate Program in Cell and Molecular Biology Annual Symposium Poster Contest Winner
- 2000 Travel Award from Women in Endocrinology
- 2000 Travel Award from Neuroendocrinology
- 1997-2000 USDA Biotechnology Training Grant
- 2001 Endocrine Society Travel Grant Award
- 2003 RSP Postdoctoral Fellowship, NIH T32 Training Grant
- 2004-2006 Ruth L. Kirschstein National Research Service Award (NRSA), NIH: National Institute of Child Health & Human Development
- 2008 McNair Scholar Program Outstanding Mentor
- 2008 SIUC Office of Research Development and Administration Faculty Seed Grant
- 2009 SIU Central Research Committee Starter Grant
- 2012 Invited talk: The Society for the Study of Reproduction Ann. Mtg. State College, PA. The Forkhead Transcription Factor, *Foxp3*, is Required for Normal Pituitary Function. Session: The Foxy Pituitary: Emerging Roles for Forkhead Transcription Factors in Gonadotrope Development and Function.
- 2012 SIU School of Medicine Excellence in Academic Medicine Award
- 2013 Invited talk: The Endocrine Society Ann. Mtg. San Francisco, CA. Forkhead Transcription Factors in Pituitary Development. Session: The Secret Lives of Gonadotropes.
- 2013 SIU School of Medicine Near Miss Award

**C. Publications**

\*Authors contributed equally to this publication.

1. Duval DL, **Ellsworth BS**, Clay CM 1999 Is gonadotrope expression of the gonadotropin-releasing hormone receptor gene mediated by autocrine/paracrine stimulation of an activin response element? *Endocrinology* 140:1949-1952.
2. **Ellsworth BS**, Burns AT, Escudero KW, Duval DL, Nelson SE, Clay CM 2003 The gonadotropin releasing hormone (GnRH) receptor activating sequence (GRAS) is a composite regulatory element that interacts with multiple classes of transcription factors including Smads, AP-1 and a forkhead DNA binding protein. *Mol Cell Endocrinol* 206:93-111.
3. \***Ellsworth BS**, \*White BR, Burns AT, Cherrington BD, Otis AM, Clay CM 2003 c-Jun N-terminal kinase (JNK) activation of AP-1 underlies homologous regulation of the gonadotropin-releasing hormone receptor gene in  $\alpha$ T3-1 cells. *Endocrinology* 144:839-849.
4. **Ellsworth BS**, Egashira N, Haller J, Butts DL, Cocquet J, Clay CM, Osamura RY, Camper SA. 2006 FOXL2 in the Pituitary: Genetic, Molecular and Developmental Analysis. *Mol Endocrinol* 20:2796-805.
5. Hertzano R, Dror A, Montcouquiol M, Ahmed Z, **Ellsworth B**, Camper S, Friedman TB, Kelley MW, Avraham K. 2007 *Lhx3*, a LIM domain transcription factor, is regulated by POU4F3 in the auditory, but not in the vestibular system. *Eur J Neurosci* 25:999-1005.
6. Tian G, Singh U, Yu Y, **Ellsworth BS**, Hemberger M, Geyer R, Stewart MD, Behringer RR, Fundele R. 2008 Expression and function of the LIM homeobox containing genes *Lhx3* and *Lhx4* in the mouse placenta. *Dev Dyn*, 237:1517-25.
7. \***Ellsworth BS**, \*Butts DL, Camper SA. 2008 Mechanisms Underlying Pituitary Hypoplasia and Dorsalization of Pre-gonadotropes in *Lhx3* Deficient Mice. *Dev Biol* 313:118-29. PMID: PMC2768753
8. Davis SW, Castinetti F, Carvalho LR, **Ellsworth BS**, Potok MA, Lyons RH, Brinkmeier ML, Raetzman LT, Carninci P, Mortensen AH, Hayashizaki Y, Arnhold IJ, Mendonca BB, Brue T, Camper SA. 2010 Molecular Mechanisms of pituitary organogenesis: In search of novel regulatory genes. *Mol Cell Endocrinol*, 323:4-19. PMID: PMC2909473
9. Carvalho LR, Brinkmeier ML, Castinetti F, **Ellsworth BS**, Camper SA. 2010 Corepressors TLE1 and TLE3 Interact with HESX1 and PROP1. *Mol Endocrinol*, 24:754-65. PMID: PMC2852357
10. Yang WH, Gutierrez N, Wang L, **Ellsworth BS**, Wang CM. 2010 Synergistic Activation of the Mc2r Promoter by FOXL2 and NR5A1 in Mice. *Biol Reprod*, 83:842.
11. Jung DO, Jasurda JS, Egashira N, **Ellsworth BS**. 2012 The Forkhead Transcription Factor, FOXP3, Is Required for Normal Pituitary Gonadotropin Expression in Mice. *Biol Reprod*, 86:144. PMID: PMC3364925
12. Gumbel JH, Patterson EM, Owusu SA, Kabat BE, Jung DO, Simmons J, Hopkins T, **Ellsworth BS**. 2012 The Forkhead Transcription Factor, *Foxd1*, is Necessary for Pituitary Luteinizing Hormone Expression in Mice. *PLoS ONE*, 7:e52156. PMID: PMC3526578
13. Majumdar S, Farris CL, Kabat BE, Jung DO, **Ellsworth BS**. 2012 Forkhead Box O1 is Increased in Embryonic Pituitary in the Absence of *p27<sup>Kip1</sup>*. *PLoS ONE*, 7:e52136. PMID: PMC3522653
14. Tran S, Zhou X, Lafleur C, Calderon MJ, **Ellsworth BS**, Kimmins S, Boehm U, Treier M, Boerboom D, Bernard DJ. 2013 Impaired Fertility and FSH Synthesis in Gonadotrope-Specific *Foxl2* Knockout Mice. *Mol Endocrinol*, 27:407. PMID: PMC3589670
15. **Ellsworth BS**. 2013 Obesity: A Somatotrope Perspective. *Endocrinology*, 154:1390.



Southern Illinois University School of Medicine      Ellsworth (PI)      03/1/2012 – 02/28/2013  
The Role of the Forkhead Transcription Factor, FOXG1, in Pituitary Development  
Excellence in Academic Medicine Award: The major goal of this project is to investigate the contribution of FOXG1 to pituitary gland development.  
Role: PI

Southern Illinois University School of Medicine      Ellsworth (PI)      11/01/2013-06/30/2014  
Mechanism by Which FOXO1 Regulates Somatotrope Differentiation and/or Function.  
Near-Miss Award: The major goal of this project is to obtain preliminary data for an NIH grant application.  
Role: PI