

**BIOGRAPHICAL SKETCH**

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NAME W. Allan Walker	POSITION TITLE Conrad Taff Professor of Nutrition and Pediatrics Chairman, Division of Nutrition, Harvard Medical School		
eRA COMMONS USER NAME WAWALKER			
EDUCATION/TRAINING <i>(Begin with baccalaureate or other initial professional education, such as nursing, and include postdoctoral training.)</i>			
INSTITUTION AND LOCATION	DEGREE <i>(if applicable)</i>	YEAR(s)	FIELD OF STUDY
DePauw University. Greencastle, Indiana	B.S. (magna cum Laude)	1959	Premedical Science English Literature
Washington University School of Medicine St. Louis, Missouri	M.D. (cum laude)	1963	Medicine

**A. Positions and Honors****Positions and Employment**

1963-64 Intern in Pediatrics, University of Minnesota Hospitals, Minneapolis, MN  
 1964-67 Resident and Chief Resident in Pediatrics, University of Minnesota Hospitals, Minneapolis, MN  
 1964-66 Wyeth Pediatric Fellowship  
 1967-69 Research Fellow in Immunologic Diseases (Drs. Robert Good/Richard Hong), Department of Pediatrics, University of Minnesota, Minneapolis, MN  
 1969-72 Research Fellow in Medicine, (Gastroenterology, Dr. Kurt Isselbacher), Massachusetts General Hospital, Harvard Medical School, Boston  
 1969- Chief, Mucosal Immunology Laboratory, Massachusetts General Hospital for Children  
 1970-71 Instructor in Medicine, Harvard Medical School  
 1972-75 Assistant Professor of Pediatrics, Harvard Medical School  
 1972-82 Chief of Pediatric Gastrointestinal and Nutrition Unit, Massachusetts General Hospital  
 1975; 1980 Associate Pediatrician and Pediatrician, Massachusetts General Hospital, Boston, MA  
 1975-82 Associate Professor of Pediatrics, Harvard Medical School  
 1982- Professor of Pediatrics, Harvard Medical School  
 1982-01 Chief, Combined Program in Pediatric Gastroenterology and Nutrition, Children's Hospital and Massachusetts General Hospital  
 1991- Conrad Taff Professor of Nutrition, Harvard Medical School  
 1993- Professor, Department of Nutrition, Harvard School of Public Health  
 1994- Principal Investigator, Harvard Clinical Nutrition Research Center (P30 DK40561)  
 1995- Chairman, Division of Nutrition, Harvard Medical School

**Other Experience and Professional Memberships**

1975-80 Academic Research Career Development Award, NIAMDD, NIH, Bethesda, MD  
 1980-87;1996-00 Member and Chairman, Gastrointestinal & Clinical Nutrition Study Section (GMA-2), NIH  
 1988-92 Member and Chairman for NIDDK Training Fellowship Committee  
 1989-2004 Editorial Boards: Gastroenterology, Nutrition Reviews, Pediatrics in Review, Nutritional Biochemistry, Pediatric Research; Editor- J of Pediatric Gastroenterology and Nutrition  
 1993-1996 Chairman Research Committee, American Gastroenterological Association  
 2001-2005 Member, Advisory Council, NIDDK, NIH

**Honors**

1984 Nutrition Award (Borden Award), American Academy of Pediatrics  
 1992 Fogarty International Fellowship (Sabbatical in Molecular Biology Department, Pasteur Institut with Professor Daniel E. Louvard)

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- 1995 Shwachman Award for outstanding achievement in Pediatric Gastroenterology, North American Society for Pediatric Gastroenterology and Nutrition
- 1997 MERIT AWARD- NICHD - R37 HD12437
- 1997 Hugh R. Butt Award for outstanding clinical research in nutrition, AGA
- 1998 Murray Davidson Award for excellence in gastroenterology, Am. Academy of Pediatrics

**B. Selected Peer-reviewed Publications or manuscripts in press 1993 to present (Selected from over 350 publications):**

1. Chu SW, **Walker WA**. Bacterial toxin interaction with the developing intestine: A possible explanation for toxigenic diarrhea of infancy. *Gastroenterology* 1993; 104:916-925.
2. Israel EJ, Simister N, Freiberg E, Caplan A, **Walker WA**. Immunoglobulin G binding sites on the human fetal intestine. *Immunology* 1993; 79:77-81.
3. Tanaka M, Lee K, Martinez-Augustin O, He Y, Sanderson IR, **Walker WA**. Exogenous nucleotides alter the proliferation, differentiation and apoptosis of human small intestinal epithelium. *J Nutr* 1996; 126:424-433.
4. Sanderson IR, Xu Z, Chu SW, Xie QY, Levine L, **Walker WA**. Developmental differences in the stimulatory GTP binding protein  $\alpha$  subunit (Gsa) for adenylate cyclase in the rat small intestine. *GUT* 1996; 38:853-858.
5. Sanderson IR, Ezzell RM, Keding M, Erlanger M, Xu Z, Pringault E, Leon-Robine S, Louvard D, **Walker WA**. Human fetal enterocytes *in vitro*: modulation of the phenotype by extracellular matrix. *PNAS* 1996; 93:7717-7722.
6. Nanthakumar N, Fusunyan R, Sanderson IR, **Walker WA**. Inflammation in the developing intestine: A possible pathophysiologic contribution to necrotizing enterocolitis. *PNAS* 2000; 97(11):6043-6048.
7. Fusunyan R, Nanthakumar N, Baldeon, ME, **Walker WA**. Evidence for an innate immune response in the immature intestine: Toll-like receptors on fetal enterocytes. *Pediatric Research* 2001; 49(4):589-593.
8. Savidge TC, Lowe DC, **Walker WA**. Developmental regulation of intestinal epithelial hydrolyase activity in human jejunal xenografts. *Ped Res* 2001; 50(2):196-202.
9. Claud E, **Walker WA**. Hypothesis: Inappropriate colonization of the premature intestine can cause neonatal necrotizing enterocolitis. *FASEB J* 2001; 15:1398-1403.
10. Dai D, Nanthakumar N, Savidge TC, Newburg DS, **Walker WA**. Region-specific ontogeny of  $\alpha$ 2.6 sialyltransferase during normal and cortisone-induced maturation in the mouse intestine. *Am J Physiology (Gastroenterology)* 2002; 282:G480-490.
11. Nanthakumar NN, Dai D, Newburg DS, **Walker WA**. (November 15, 2002) The role of indigenous microflora in the development of murine intestinal fucosyl- and sialyltransferases. *FASEB J*.(November 15, 2002) 10.1096/fj.02-003 1fje (summary: *FASEB J* 2003; 17:44-46.
12. Claud EC, Savidge T, **Walker WA**. Modulation of human intestinal epithelial cell interleukin-8 secretion by human milk factors. *Ped Res* 2003; 53(3):419-425.
13. Kohler H, McCormick BA, **Walker WA**. Bacterial-enterocyte crosstalk: Cellular mechanisms in health and disease. *J Pediatr Gastr Nutr* 2003; 36(2):175- 185.
14. Nanthakumar NN, Kloplic C, Fernandez I, **Walker WA**. Normal and glucocorticoid induced development of the human small intestinal xenograft. *Am J Physiol Regul Integr Comp Physiol* 2003; 285:R162-R170.
15. Lu L, Baldeon ME, Savidge T, Pothoulakis C, **Walker WA**. Development of microbial-human enterocyte interaction: Cholera toxin. *Ped Res* 2003; 54:212-218.
16. Claud EC, Lu L, Anton PM, Savidge T, **Walker WA**, Cherayil BJ. Developmentally-regulated I $\kappa$ B expression in intestinal epithelium and susceptibility to flagellin-induced inflammation. *PNAS* 2004; 101(19):7404-7408 .
17. Nanthakumar NN, Young C, Ko JS, Meng D, Chen J, Buie T, **Walker WA**. Glucocorticoid responsiveness in the developing human intestine: possible role in the prevention of necrotizing enterocolitis. *AJP: GI and Liver Physiology* 2005 288:G85-G92.
18. Nanthakumar NN, Dai D, Meng D, Chaudry N, Newburg DS, **Walker WA**. Regulation of intestinal ontogeny: effect of glucocorticoids and luminal microbes on galactosyltransferase and terehalase induction in mice. *Glycobiology* 2005; 15:221-232.
19. Lu L, Khan S, Lencer W, **Walker WA**. Endocytosis of cholera toxin by human enterocytes is

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developmentally regulated. Am J Physiology (Gastro) 2005; 298:G332-G341.

20. Sanderson IR, **Walker WA**. The role of TLRs/Nods in intestinal development and homeostasis. Am J Physiol (GI) 2006 (in press).

### **C. Research Support**

#### **Ongoing Research Support**

P30 DK40561-12 Walker (PI) 05/08/06-03/31/11

NIH

Clinical Research Nutrition Center

Specific aims of project: The major specific aims are: 1) to provide research in basic areas of biology relevant to problems in clinical nutrition; 2) to promote the study of clinical nutrition and application of knowledge derived there from within the HMS, MIT and HSHP community; 3) to promote interactions among scientists and clinical investigators to show relevance to advancing the science of clinical nutrition; and 4) to attract "basic" investigators into the domain of clinical nutrition to promote an environment and mechanism to develop new investigator-initiated research.

Role: Program Director and PI

5 RO1 DK70260-13 Walker (PI) 05/01/05-04/33/10

NIH-NICHD/NIDDK

Ontogeny of the Human Intestinal Mucosal Barrier and NEC

The major goals of this project are to study: 1) bacterial colonization and its pattern in NEC patients 2) the developmental regulation of antibacterial products of Paneth cells at various ages (fetal - childhood and in NEC patients) 3) the functional significance of the human fetal intestinal Fc receptor for human IgG and 4) selective aspects of intestinal inflammatory responses and immune function.

Role: PI

5 R37 HD12437-27 Walker (PI) 01/01/97-12/31/07

NIH

The Effect of Colostrum on Gut Maturation and Host Defense (MERIT)

The major goals of this project are to define the role of "growth factors" in amniotic fluid and in preterm/term on enterocyte glycosylation of MVM molecules and bacterial colonization. To establish an immortalized fetal human small intestinal cell line.

Role: PI

2 PO1 DK33506-22 Pothoulakis (PI) 09/30/05-08/31/10

NIH-NIDDK

Barrier Function of the GI Tract in Health and Disease

The major goals of this project are: 1) To develop a non-malignant human intestinal epithelial cell line and to examine its function as an antigen presenting cell. (a) by determining its capacity for MHC-class II, Ii, and B7 expression; (b) by defining its capabilities as an antigen presenting cell; (c) by comparing its antigen presentation capabilities to that of other antigens presenting cells. 2) To examine the basis for systemic non-responsiveness after oral administration of a soluble protein antigen (cytochrome C) in a T cell receptor transgenic mouse model. We will after oral administration of cytochrome C: (a) examine the sites and kinetics of induction of non-responsiveness in normal and TCR transgenic mice; (b) examine the contributions of clonal deletion, clonal anergy and active suppression to establishment of systemic tolerance in TCR transgenic mice; (c) determine which populations of lymphocytes are capable of transferring non-responsiveness and examine the phenotypic and functions characteristics of these cells.

Role: Program Director and PI

#### **Completed**

None