# **Human Milk:** The Ultimate Superfood & **Personalized Medicine**

#### Meghan Azad, PhD

 ${\it Canada\ Research\ Chair,\ Developmental\ Origins\ of\ Chronic\ Disease}$ Co-Director, Manitoba Interdisciplinary Lactation Centre (MILC) Scientific Director, International Milk Composition Consortium (IMiC) Deputy Director, CHILD Cohort Study











#### Disclosures

#### Research grants:

- Canadian Institutes of Health Research
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   Network of Centers of Excellence
- Canadian Lung Association
- Research Manitoba
- Children's Hospital Foundation of Manitoba
- Prolacta Biosciences
- Bill and Melinda Gates Foundation
- Canada Foundation for Innovation
- W. Garfield Weston Foundation
- Health Data Research UK
- Canadian COVID Immunity Task Force

#### Unrestricted research funding:

- Canada Research Chairs Program
- Healthy Child Manitoba Office, Government of Manitoba
- Canadian Institute for Advanced Research (CIFAR)

#### Advisory Committee Membership:

■ Malaika Vx

#### Conference presentation honoraria:

Prolacta Biosciences

Last 3 years (Apr 2021)

#### personalized medicine

Medhan Alad

"Human breastmilk is not only a perfectly adapted nutritional supply for the infant, but probably the most specific **personalized medicine** that he or she is likely to receive, given at a time when gene expression is being **fine tuned for life**."

Victora et al.

The Lancet

Breastfeeding Series (2016)

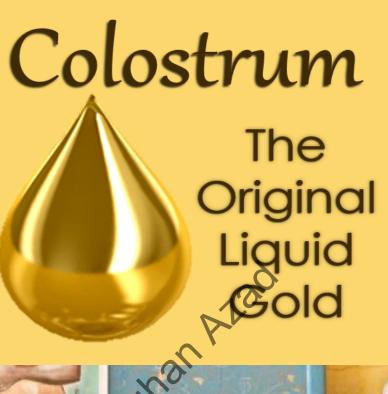


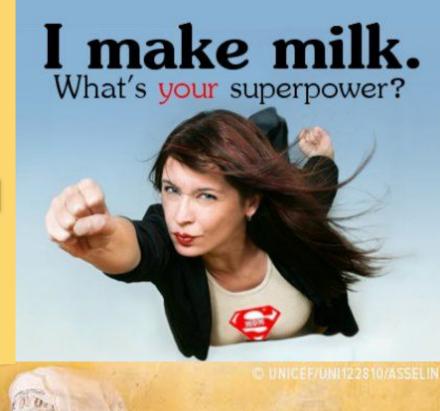
CESAR VICTORA, WINNER OF THE JOHN DIRKS CANADA GAIRDNER GLOBAL HEALTH AWARD

Breastmilk is Superfood for Babies

naturemoms.com









# Breastmilk is Superfood for Babies

"Milk is really a genius fluid that was outrageously understudied. If we can identify components of human breast milk that are important, then we can understand the wisdom of milk-and take advantage of them."

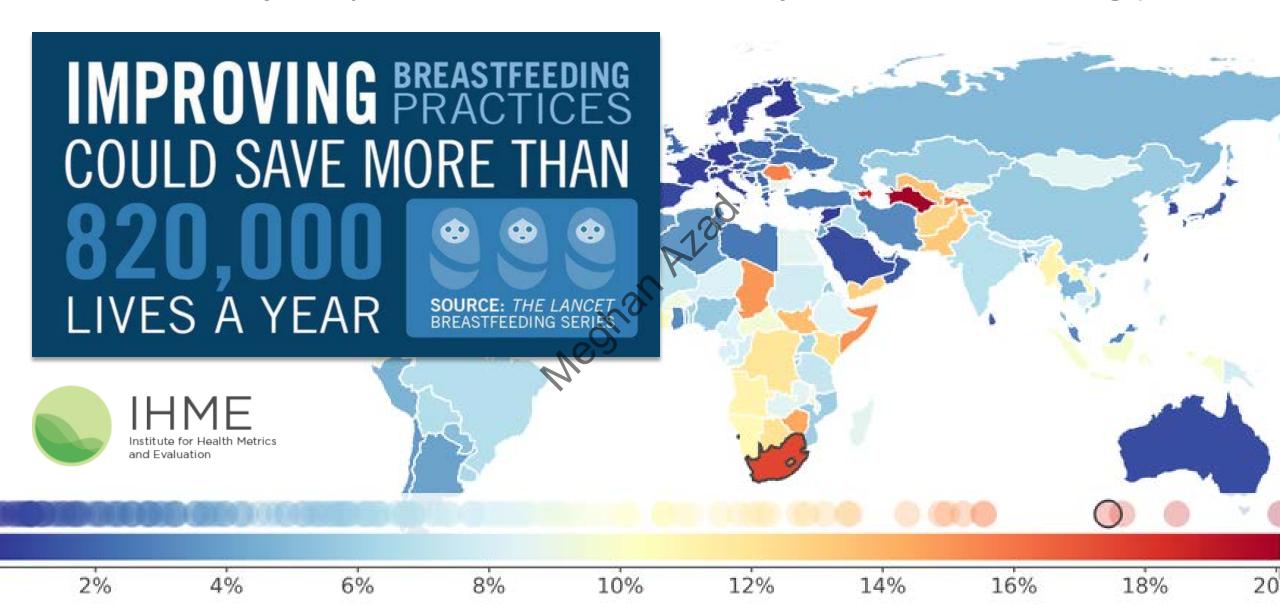
David Mills, UC Davis



#### Nature's first functional food

Breast milk feeds helpful microbes, fights harmful ones, provides immunity, and jump-starts a newborn's life ing explored once again. Sor have focused on making be microbiome fostered by milk have documented how bre more than feed a newborn bacteria. Mother's milk also

#### Child mortality (<5 years) attributable to suboptimal breastfeeding (2013)



#### **Eight Great Reasons to Breastfeed Your Baby**

**#1 It's Convenient -** Breast milk is always quick, clean, the right temperature and available.

**#2 It's Perfect Food For Babies -** Breast milk gives your baby exactly what he or she needs to grow and develop.

**#3 Breastfed Babies Are Healthier -** Breast milk contains antibodies that help a baby have fewer infections and allergies. It is easier to digest so your baby will have less diarrhea and constipation.

**#4 Moms Are Healthier, Too -** Medical research has linked breastfeeding to lowered risk of breast and ovarian cancer.

#### #5 Mom Gets Back Into Shape Faster -

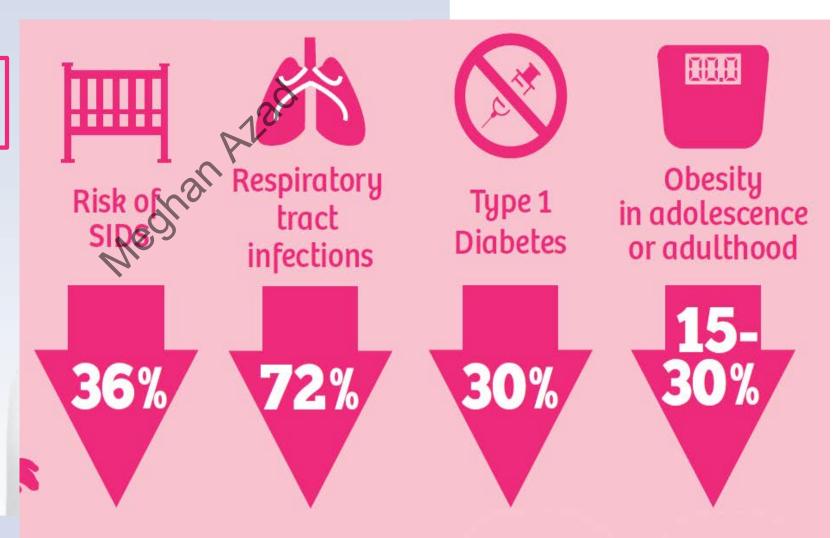
Breastfeeding burns extra calories and triggers the uterus to contract and return to its normal shape more quickly.

**#6 It's Better For The Planet -** Breastfeeding is never wasteful. Formula feeding results in discarded cans and bottles.

**#7 Breastfeeding Builds Love -** Skin-to-skin contact brings you and your baby close. It helps your baby feel secure.

**#8 Breastfeeding Saves Money -** Breastfeeding helps the family budget, by saving over \$1000 a year compared to formula.





# **Eight Great Reasons to Breastfeed Your Baby**



nature How early life shapes the infant gut and risk of disease SECURITY BLANKET REACHING FOR

Maternal BMI Maternal pregnancy wtgain Maternal diabetes medication Maternal diabetes-Maternal preeclampsia Maternal probiotic Birth mode Preterm Sex **Breast milk** Solid food **Probiotic** Vitamin D Geographical location Household siblings Household furry pets Child lives on farm Daycare exposure **Antibiotics** Coeliac disease Any acute disease Any chronic disease-

Azad et al. 2013 - CHILD

Stewart et al. 2018 - TEDDY

### **Developmental Origins of Health and Disease (DOHaD)**

and the Microbiome



"Human breastmilk is not only a perfectly adapted nutritional supply for the infant, but probably the most specific **personalized medicine** that he or she is likely to receive, given at a time when gene expression [and the microbiome] is being **fine tuned for life**."

Victora et al. The Lancet Breastfeeding Series (2016)



# Human Milk is

#### **Personalized Nutrition**

Matches the changing nutritional needs of the (human!) infant

## **Personalized Medicine**

Delivers (optimized!) immunity while infant immune system develops

## **Dynamic**

Varies diurnally, within feedings, across lactation, and among mothers

#### DID YOU EVER WONDER WHAT'S IN ... ?

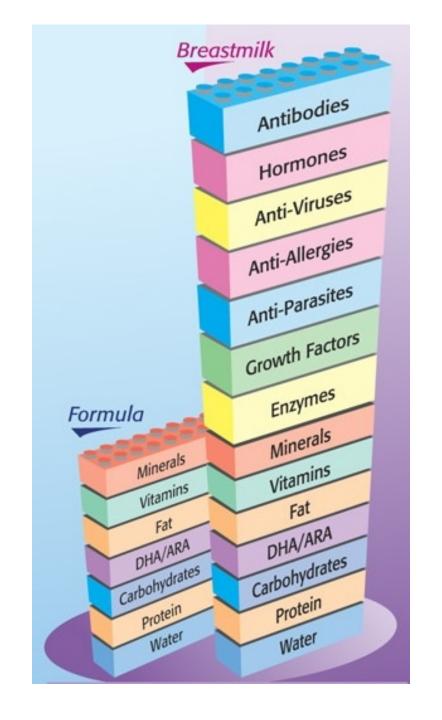


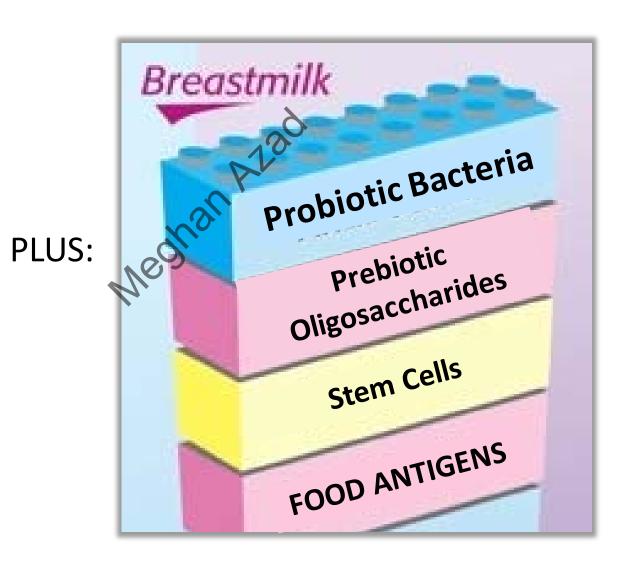
Name something found in human milk that is not found in infant formula

#### DID YOU EVER WONDER WHAT'S IN ... ?



Developed as a student project for the Breadfeeding Course for Health Care Providers, Douglas College, New Westminster, BC, Canada - © 2007 by Cecily Heslett, Sherri Hedberg and Haley Rumble.

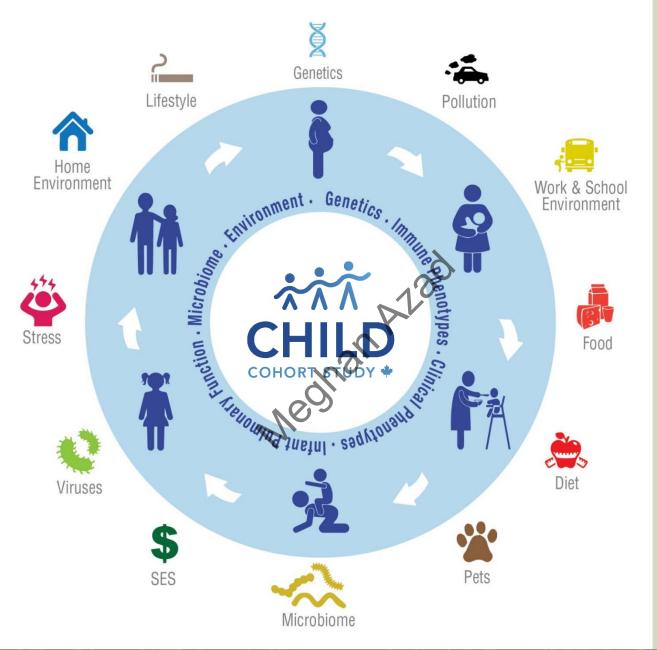




# The Canadian Healthy Infant Longitudinal Development (CHILD) Study

How do genes and the environment influence child health and development?





\$56M Invested 500,000 Samples:

Blood, Urine, <u>Stool</u>, Nasal Swabs, Dust,

**Breast Milk** 

200,000 Questionnaires

**3600** Families

**40+** Researchers

**20+** Disciplines

**8(+)** Years Follow-Up

93% Retention





# Human Milk \*\*\* CHILD

#### **Completed or Underway**

- Endocrine Hormones (C. Field, U. Alberta) Chan et al. Int J Obesity 2017
- Oligosaccharides (L. Bode, UC San Diego) Azad et al. Am J Clin Nutr 2018
- Microbiota (E. Khafipour, U. Manitoba) Moossavi et al. Cell Host & Microbe 229
- Fatty Acids (C. Field, U. Alberta) Miliku et al. Am J Clin Nutr 2019
- Cytokines (J. Marshall, Dalhousie)
- Peptides (K. Hettinga, Wageningen University)

#### **Coming Soon**

- Bioactive Proteins (L. Bode, UC San Diego)
- Antigen-specific Immunoglobulins (T. Eiwegger, SickKids & ADI Inc.)
- Proteomics (Precision Biomarker Laboratory)
- Untargeted Metabolomics (M. Jain, UC San Diego)
- Targeted Metabolomics (Biocrates)
- Micro & Macro nutrients (L. Allen, USDA)

#### Wish List (Your name here! ☺)

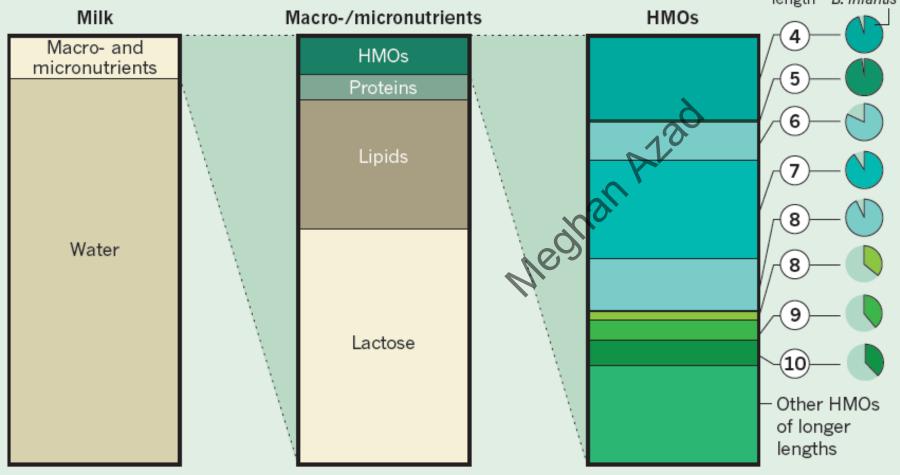
Toxins, Food Antigens, Artificial Sweeteners, Choline, Cortisol...



#### WHAT'S IN HUMAN MILK

Human milk oligosaccharides (HMOs) are food for friendly bacteria like *Bifidobacterium*infantis. Shorter chain HMOs in particular are almost entirely consumed by this microbe.

Chain eaten by length *B. infantis* 



## **HMOs**

Non-digestible carbohydrates
Food for microbes (not for baby!)

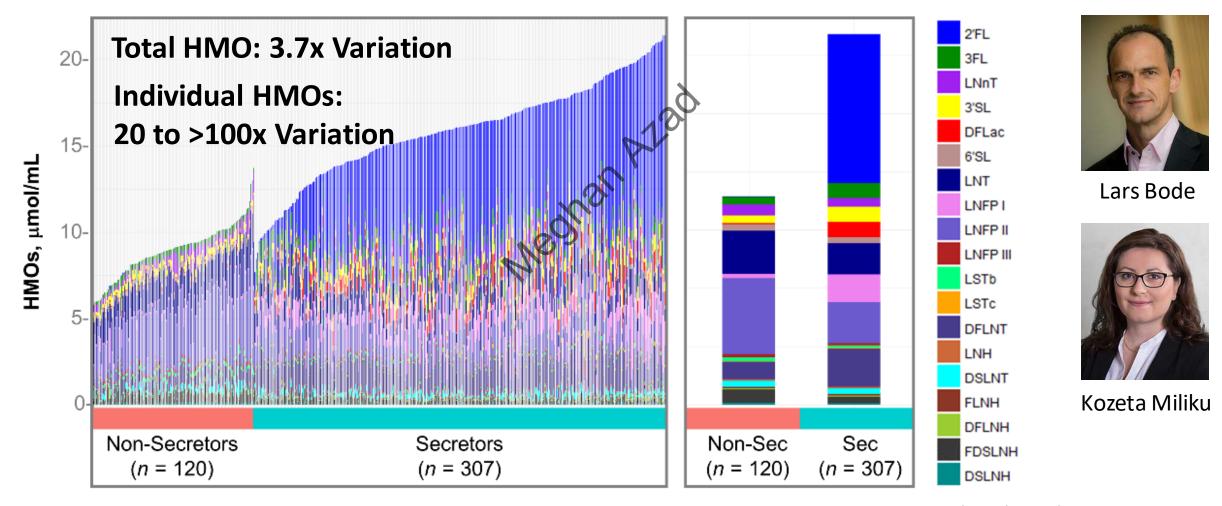
& species-specific Cows ~40 vs. Humans > 100

Highly variable (How & Why??)

Health Impact?

#### HMOs in the CHILD Cohort

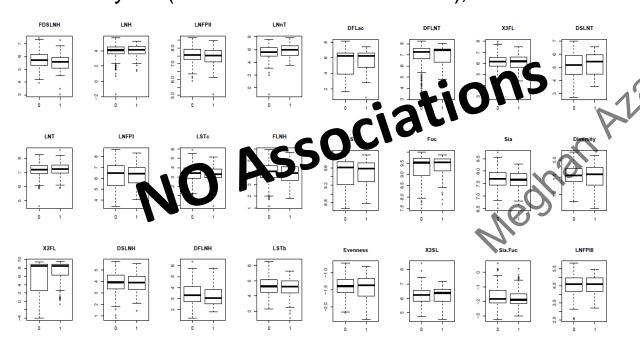




#### **HMOs & Infant Food Sensitization**



#### N=430 dyads (59 sensitized vs. 362 controls); 19 HMOs

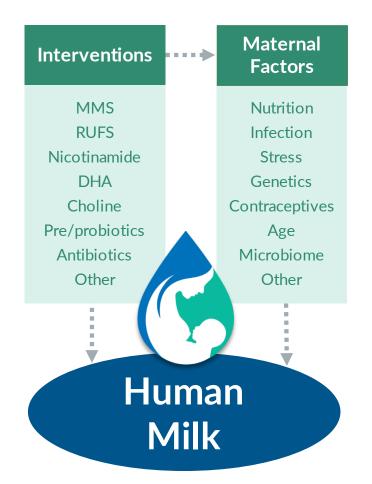


While no <u>single</u> HMO predicts food sensitization, the **overall HMO profile** does.

# What is the MECHANISM?

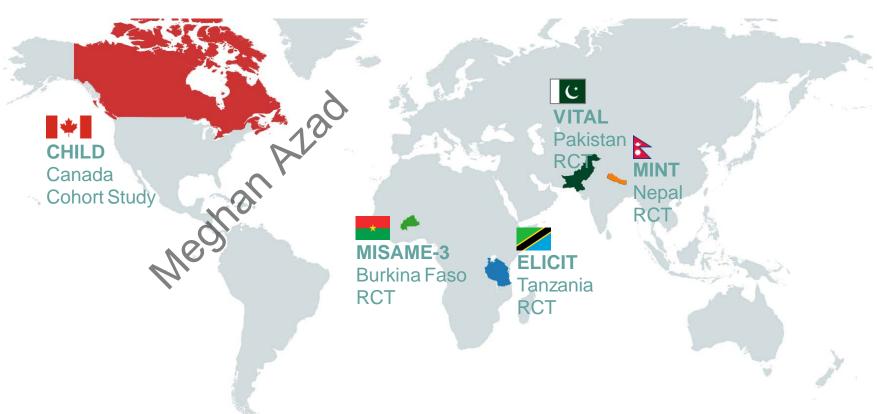
What about OTHER milk components?

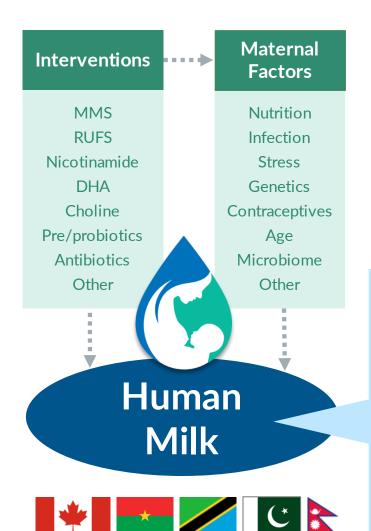
**OTHER conditions?** 











BILL&MELINDA GATES foundation



# **Laboratory Partners**

#### **Macronutrients**

Fat. Protein. Carbohydrates

#### **Micronutrients**

Vitamins, Minerals

**Bioactives** 

Microbiota

Bacteria, Fungi

mmunoglobulins

Oligosaccharides

Lactoferrin

Lysozyme

**Cytokines** 

**Hormones** 

miRNAs

**Peptides** 

**Immune Cells** 



Data Science **Partners** 



#### **Infant / Child** Health

**Growth & Body** Composition

> **Nutritional Status**

**Gut Health** 

Neuro **Development** 

Immune Development

N = 1000 dyads









#### Nima Aghaeepour **Stanford Medicine**

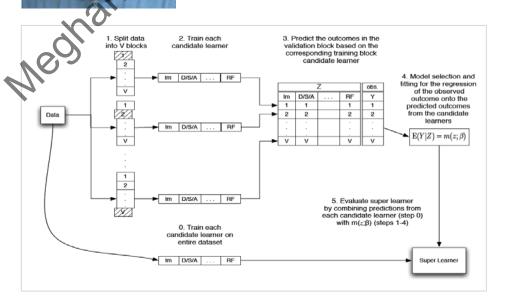
- Bioinformatics
- Multiomics
- Algorithm benchmarking
- Explainable artificial intelligence

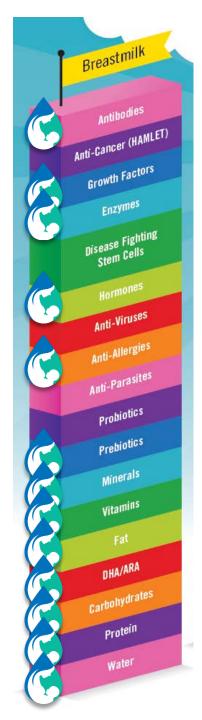




#### Alan Hubbard **Berkeley Public Health**

- Causal inference
- Machine learning
- Epidemiology
- Precision medicine







**Meghan Azad** PhD MSc Human Milk & Epidemiology



Nathan Nickel MPH PhD
Public Health & Biostatistics





Infant Feeding & Lactation Database

- All mothers in Manitoba
- Report infant feeding at routine vaccination visits
- Data faxed to MILC using Teleform



Human Milk Biorepository

- General population at selected clinics & hospitals
- Targeted clinical studies
- Provide milk + additional research data
- Macronutrient profile + long-term storage



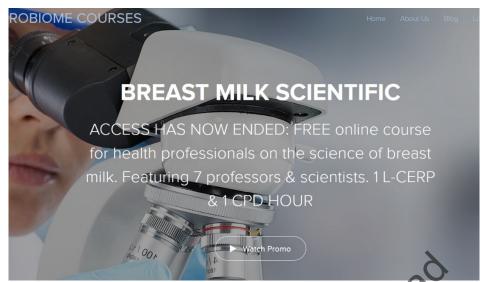
Manitoba Population Research Data Repository

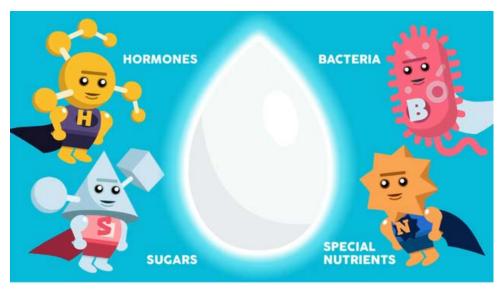
- All mothers-infant dyads in Manitoba
- Administrative health + sociodemographic data
- Linked to all MILC data and samples



## Knowledge **Translation**









The National Academies of SCIENCES · ENGINEERING · MEDICINE

Advising the Nation

**COMMITTEE ON SCANNING FOR NEW EVIDENCE ON THE NUTRIENT CONTENT OF HUMAN MILK** 

KATHLEEN M. RASMUSSEN (Chair), Cornell University, Ithaca, New York MEGHAN AZAD, University of Manitoba, Canada LARS BODE, University of California, San Diego MICHELLE MCGUIRE, University of Idaho, Moscow LAURIE NOMMSEN-RIVERS, University of Cincinnati, Ohio IAN J. SALDANHA, Brown University School of Public Health, Rhode Island

#### THE CONVERSATION





Community support programs for breastfeeding have changed under COVID-19 restrictions. (Shutterstock

Breastfeeding is recommended during pandemic, but coronavirus has changed support systems

May 7, 2020 3.03pm EDT



Merilee Brockway, Sarah Turner, University of Manitoba

# Breastmilk is Superfood for Babies

"Milk is really a genius fluid that was outrageously understudied. If we can identify components of human breast milk that are important, then we can understand the wisdom of milk—and take advantage of them."

David Mills, UC Davis

Breastfeeding Medicine, Vol. 14, No. 3 | Basic Science

# A Pilot Study of Human Milk to Reduce Intestinal Inflammation After Bone Marrow Transplant

Pooja Khandelwal ☑, Heidi Andersen, Lindsey Romick-Rosendale, Cynthia B. Taggart, Miki Watanabe, Adam Lane,
Christopher E. Dandoy, Kelly E. Lake, Bridget A. Litts, Ardythe L. Morrow, Martin L. Lee, David B. Haslam, and Stella M. Davies

Published Online: 11 Apr 2019 | https://doi.org/10.1089/bfm.2018.0199



Human milk oligosaccharide supplementation in irritable bowel syndrome patients: A parallel, randomized, double-blind, placebo-controlled study

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Cristina Iribarren<sup>1,2</sup>  | Hans Törnblom<sup>2</sup>  | Imran Aziz<sup>2,3</sup>  | Maria K. Magnusson<sup>1</sup>  | Johanna Sundin<sup>2</sup>  | Louise K. Vigsnæs<sup>4</sup>  | Ingvild Dybdrodt Amundsen<sup>4</sup>  | Bruce McConnell<sup>4</sup>  | Dorthe Seitzberg<sup>4</sup>  | Lena Öhman<sup>1,2</sup>  | Magnus Simrén<sup>2,5</sup>
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#### Characterization of SARS-CoV-2 RNA, Antibodies, and Neutralizing Capacity in Milk Produced by Women with COVID-19

D Ryan M. Pace, D Janet E. Williams, Kirsi M. Järvinen, Mandy B. Belfort, Christina D. W. Pace, Kimberly A. Lackey, Alexandra C. Gogel, Phuong Nguyen-Contant, Preshetha Kanagaiah, Theresa Fitzgerald, Rita Ferri, Bridget Young, Casey Rosen-Carole, Nichole Diaz, Courtney L. Meehan, Beatrice Caffé, Mark Y. Sangster, David Topham, Mark A. McGuire, Antti Seppo, Michelle K. McGuire

#### RESEARCH LETT

JAMA May 18, 2021 Volume 325, Number 19

#### SARS-COV-2-Specific Antibodies in Breast Milk After COVID-19 Vaccination of Breastfeeding Women

Apr 24, 2020, 03:26pm EDT | 271,423 views

#### Breast Milk Studied As Potential Coronavirus Treatment



Marla Milling Contributor (1) Health

I am a Forbes.com Contributor specializing in geriatric health and women's health articles.



# **Breastfeeding** and COVID-19

Breastfeed to protect your infants and children from getting sick and for their healthy growth and development.

Breastfeeding is particularly effective against infectious diseases because it strengthens the immune system by transferring antibodies from you.







# DO YOU DO RESEARCH IN HUMAN MILK AND LACTATION? JOIN ISRHML TODAY!

#### WHAT IS ISRHML?

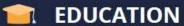
A vibrant and growing non-profit organization dedicated to the promoting excellence in research and the dissemination of research findings in the field of human milk and lactation from cell biology to public policy.

#### WHO CAN BE A MEMBER?

Members are affiliated with universities, medical centers, governmental agencies, NGOs and industry. ISRHML has four membership categories.

- Full Active Lesearchers
- Associate Alied professionals
- Traine Researchers in training
- Emeritus –Retired researchers

# WHAT ARE THE MEMBER BENEFITS?





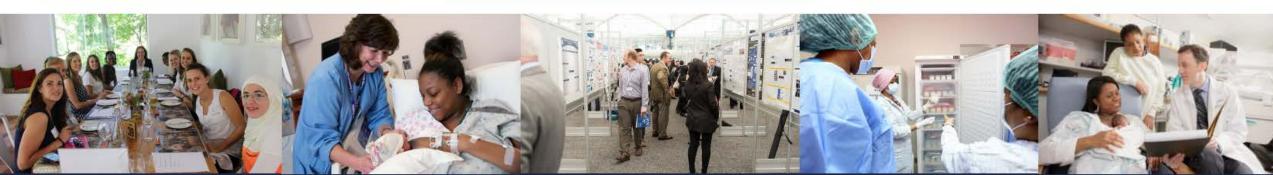


**CAREER GROWTH** 

TRAINEE EXPANSION PROGRAM (TEP)

SAVINGS

\$10k Travel Awards











Natalie Rodriguez Tricia Choquette Rilwan Azeez Stephanie Goguen Nicole Fiorentino Sarah Turner Kelsey Fehr Allison Daniel

Larisa Lotoski Bridget McGann Merilee Brockway (Kozeta Miliku)



Manitoba: Elinor Simons, Allan Becker, Scarlet Salas & Team

National: PJ Subbarao, Stuart Turvey & Team

Collaborators: Lars Bode (UC San Diego, MoMI CORE)



Nathan Nickel (Manitoba Centre for Health Policy)
Drs. Raimondi & Kearns (Winnipeg Breastfeeding Centre)
MILC Club members



























# Questions?

#### Meghan Azad, PhD

Canada Research Chair in Developmental Origins of Chronic Disease
Deputy Director, CHILD Cohort Study
Director, International Milk Composition (IMiC) Consortium
co-Director, Manitoba Interdisciplinary Lactation Centre (MILC)
co-Lead, international Perinatal Outcomes in the Pandemic (iPOP) Study
Children's Hospital Research Institute of Manitoba
Department of Pediatrics & Child Health, University of Manitoba, Canada

www.azadlab.ca www.childstudy.ca www.milcresearch.com www.ipopstudy.com

meghan.azad@umanitoba.ca @MeghanAzad











International Perinatal Outcomes in the Pandemic Study