

**Helping Parents Navigate Food Allergies** 



# Helping Parents Navigate Food Reactivity and Reach Their Feeding Goals

Differentiate between IgE and non-IgE mediated **food allergies** & intolerances





Understand how dietary foods transfer to the breast and maternal health plays a role

Comprehend the various **elimination diet and reintroduction** strategies





## Identifying Food Reactivity in Children

Skin Rash

**Chronic Congestion** 

Sleep Disturbances

Reflux/Vomiting

**Tummy Pain** 

Slow Weight Gain

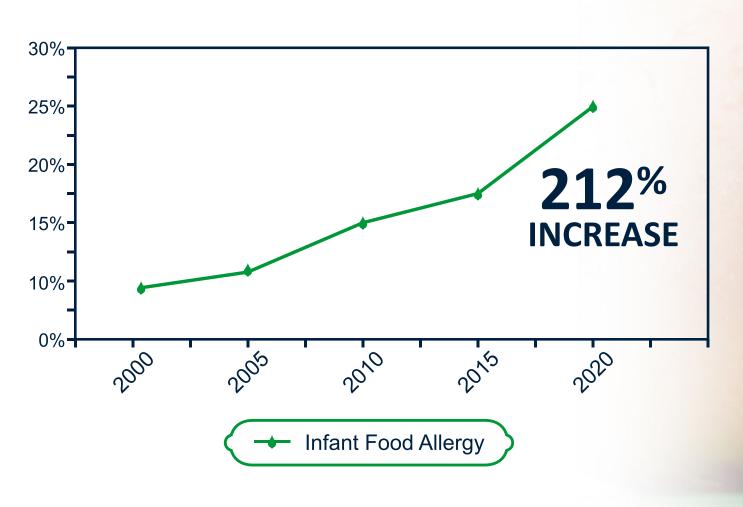
**Feeding Aversions** 

**Abnormal Stools** 

Lethargy

Colic

# Global Food Reactivity 35 Million Infants/Year





#### Food Intolerance

Inability to properly digest a food, often due to an enzyme deficiency

#### **Proteases**

Break down protein into amino acids and peptides

#### Lipases

Break down fat into three fatty acids and a glycerol

#### **Amylases**

Break down carbs into simple sugars

Can be diagnosed via blood, breath, &/or biopsy testing

#### **Lactose Intolerance**

Inability to properly break down lactose due to lactase deficiency

- Congenital: incredibly rare (only reported in a few infants worldwide) & caught immediately
- Neonatal: developmental amylase deficiency until at least 34 weeks gestation
- Secondary: caused by invasive GI surgery or severe malnutrition seen in developing countries

## Allergy Categories



**IgE** 

Immunoglobulin E





Mixed IgE & Non-IgE

**EoE**Eosinophilic Esophagitis

AEDS
Atopic Eczema/Dermatitis
Syndrome



Non-IgE

**FPIES** 

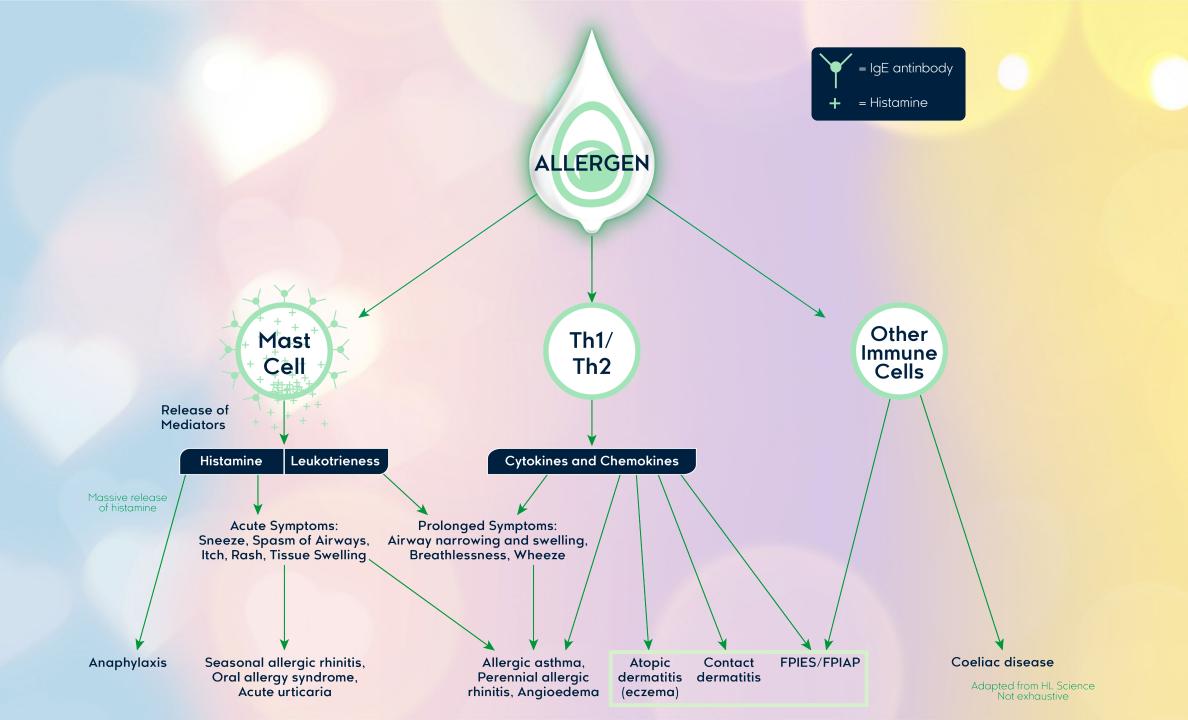
Food Protein Induced Enterocolitis Syndrome

**FPIAP** 

Food Protein Induced Allergic Proctocolitis

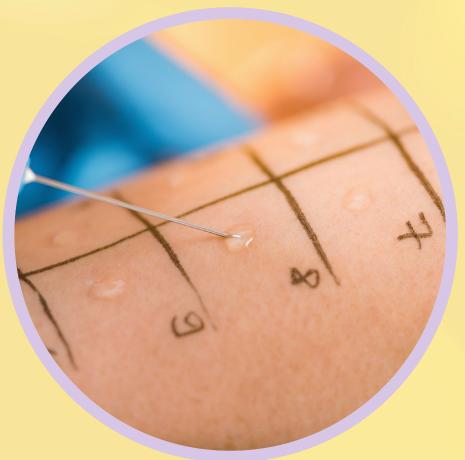
FPE

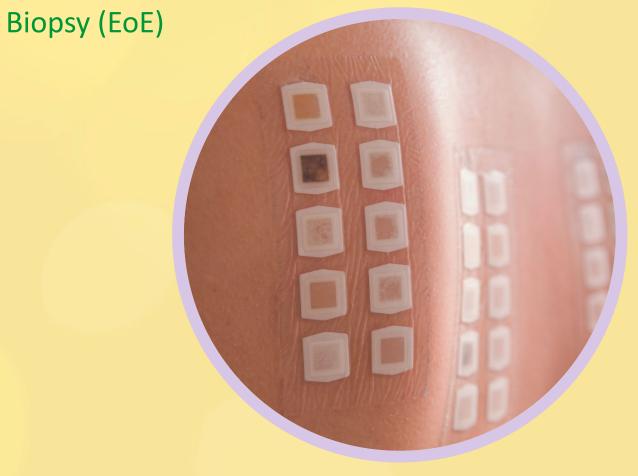
Food Protein Induced Enteropathy



### **Diagnostic Tests**

Skin prick or patch (IgE)
Blood test (IgE, EoE)
Oral food challenge (FPIES, FPIAP, FPE)





# IgE Mediated Allergy

Skin: "hives" (red blotches or welts that itch), swelling

Eyes: tearing, redness, itch

Nose: clear discharge, itch, congestion

Mouth: itch, lip swelling, tongue swelling

Throat: tightness, trouble speaking, trouble inhaling

Lungs: shortness of breath, rapid breathing, wheeze

Stomach: vomiting, nausea, abdominal pain, diarrhea

**Heart and circulation:** weak pulse, loss of consciousness

**Brain:** anxiety, agitation, loss of consciousness



A Phase 1, Randomized Study Evaluating the Safety Tolerability, Pharmacokinetics (PK) and Pharmacodynamics (PD) of Single Ascending Doses of Epinephrine Prodrug 109 Sublingual Film (AQST-109) in Healthy Male Volunteers

John Oppenheimer, MD FAAAAI 

Steve Wargacki Rajesh Kainthan, PhD Shawn Berg Cathie Leister

Mark Lepore, MD FAAAAI



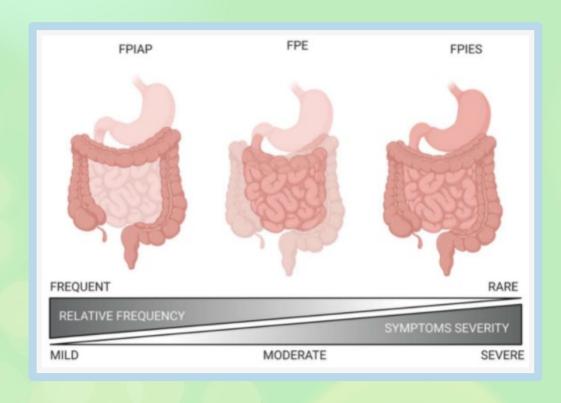
#### Sublingual

Rapid onset of action with direct entry into systemic circulation

**Epiphast by Aquestive** 

#### Non-IgE Mediated Allergy (GI)

EoE: vomiting, abdominal pain, difficulty swallowing, gagging, reflux



Secondary symptoms:
Weight gain issues, "colic",
feeding aversions, sleep
disturbance, lethargy,
dehydration, shock

FPIAP & FPE: bloody stools, mucous stool, diarrhea FPIES: diarrhea, vomiting, reflux, bloody stool

#### **Upper GI Reactivity**

Deciphering between anatomical or lactation issues versus allergy

Causes physical discomfort

Large volumes (one large or many small)

Accompanied by other symptoms

Delayed by an hour or more

Contains significant amounts of mucus

Mimicking factors: oral restrictions, oversupply, congenital defects, poor latch, teething, illness

#### Lower GI Reactivity

Deciphering between anatomical or lactation issues versus allergy

Causes physical discomfort

Diarrhea (8+ BM/Day)

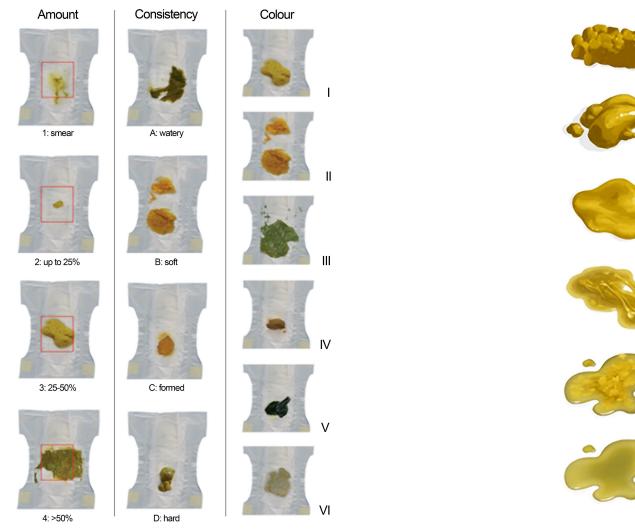
Constipation (Painful/Hard)

6-48 hours post-ingestion

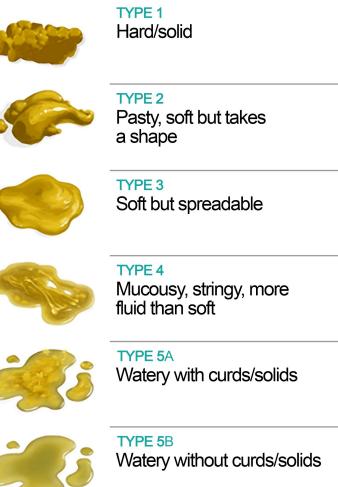
Contains significant amounts of mucus &/or blood

Mimicking factors: oral restrictions, oversupply, congenital defects, poor latch, teething, illness

#### Infant Stool Charts



**Amsterdam Stool Form** 



**Diapered Infant Stool Scale** 

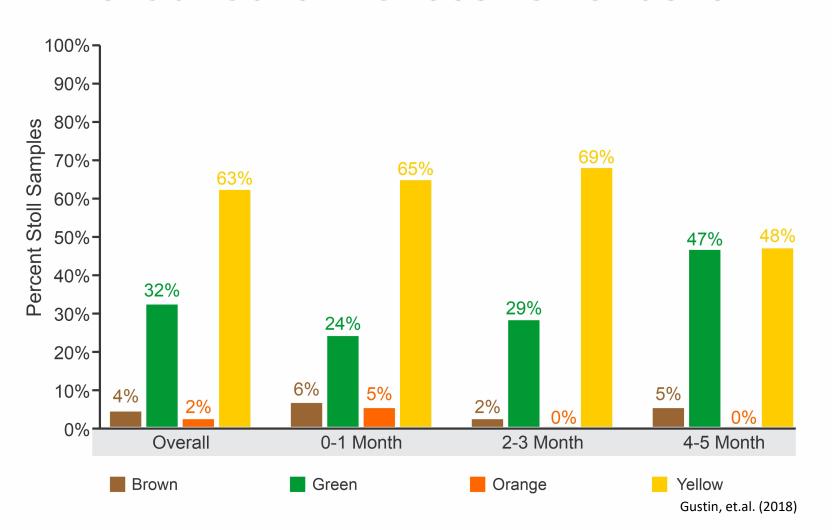
Gustin, et.al. (2018)

## "Normal" Infant Stool?

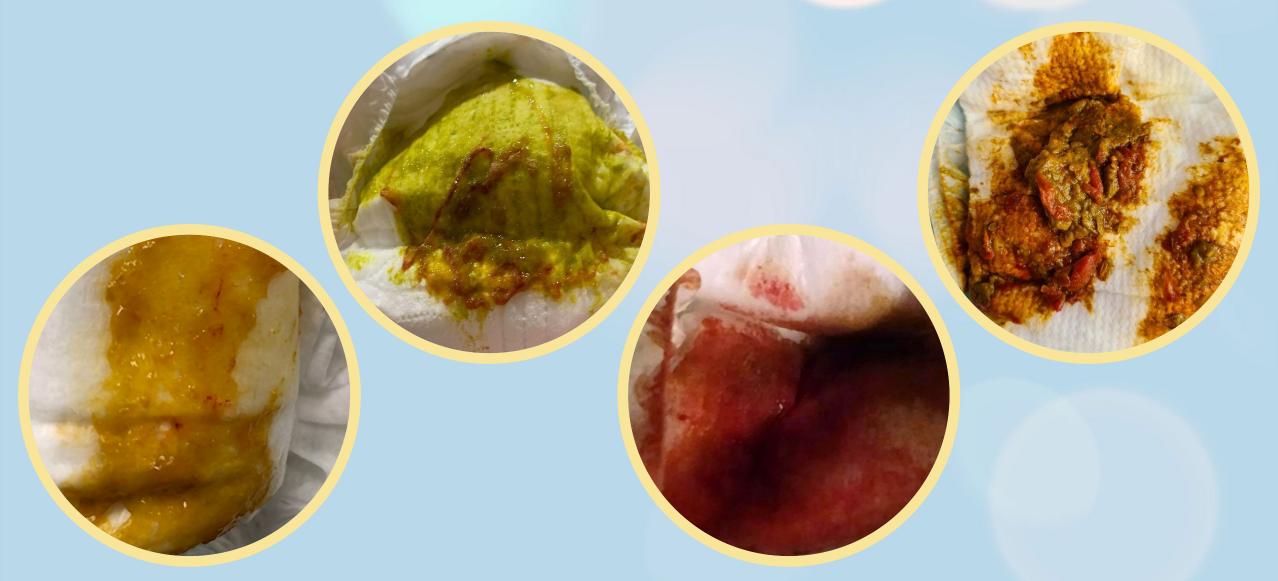


#### **Infant Stool Color**

#### STOOL COLOR ACROSS AGE GROUPS

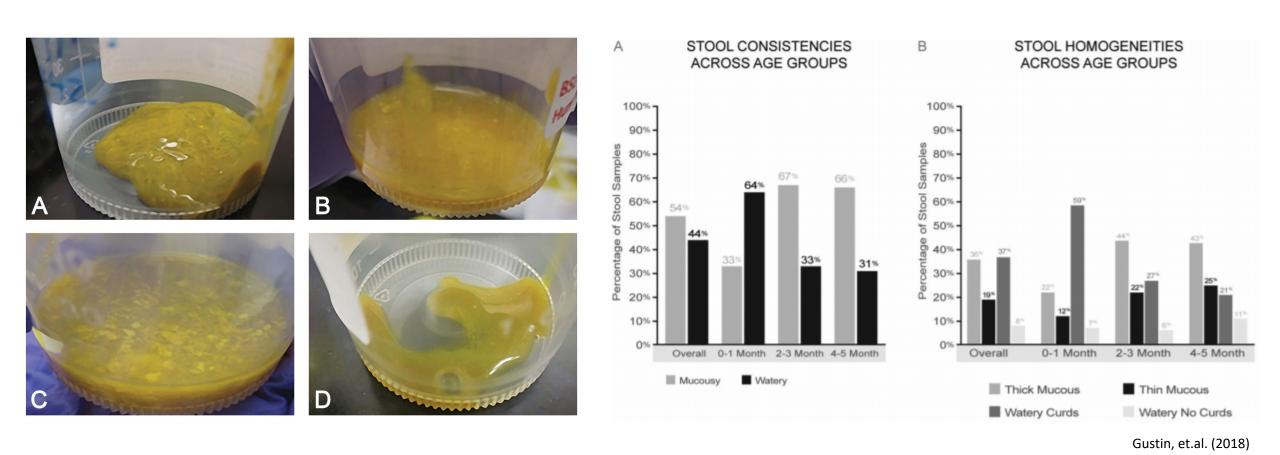


#### Infant Stool Color - Red



Mimicking factors: cracked nipples, anal fissure, food particles, "seedy" stool

## **Infant Stool Consistency**



Mimicking factors: oral restrictions, oversupply, congenital defects, poor latch, teething, illness

# **Infant Stool Consistency**



## **Infant Stool Reaction Examples**



#### **Fecal Testing Options**

- Fecal occult blood test (FOBT) will tell you if there is blood that cannot be seen with the naked eye.
- Calprotectin looks for white blood cells in the stool which is also indicative of inflammation.
- Fecal fat stain informs doctors if there is a malabsorption issue based on the amount of fat left in stool.
- Other various analysis can be completed for parasites, infections, bacteria, and more.





Mimicking factors: environmental reactivity, teething, heat, irritation

#### Symptom Takeaways

All journeys are valid & will naturally vary





Most children will not present with all the symptoms discussed

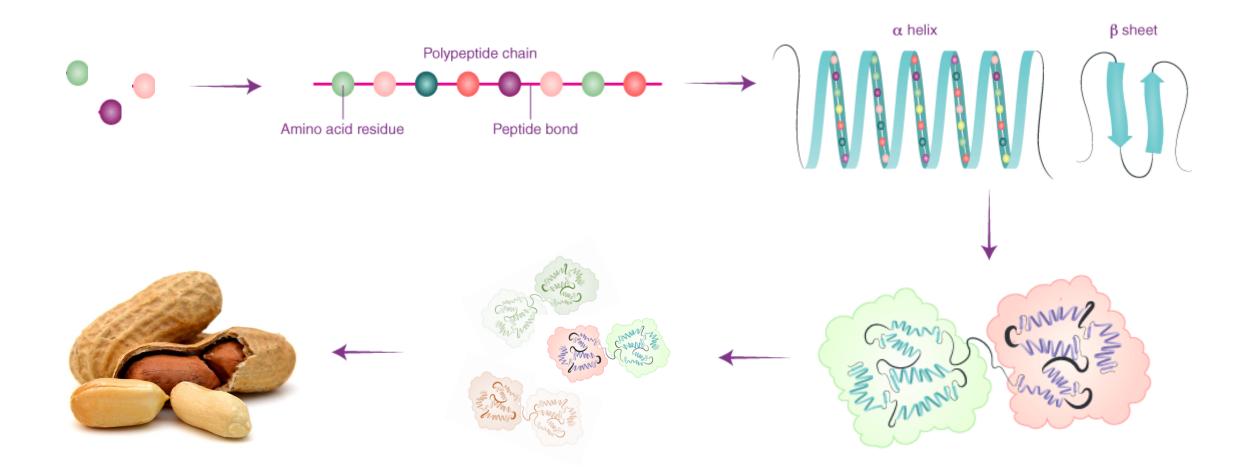
Any food can be a trigger, and each can elicit a different response



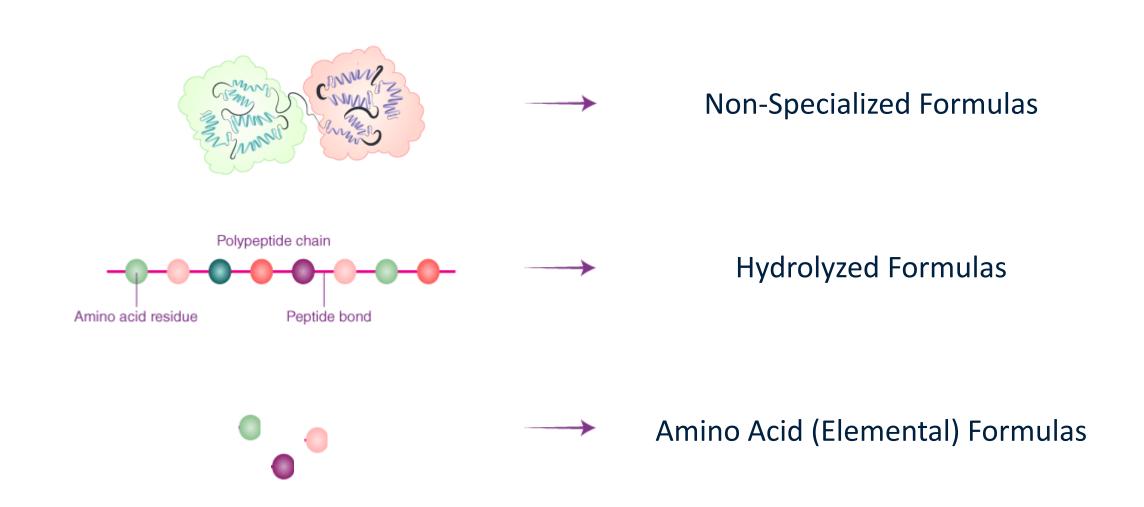


There is no one thing that rules out food reactivity IE weight gain

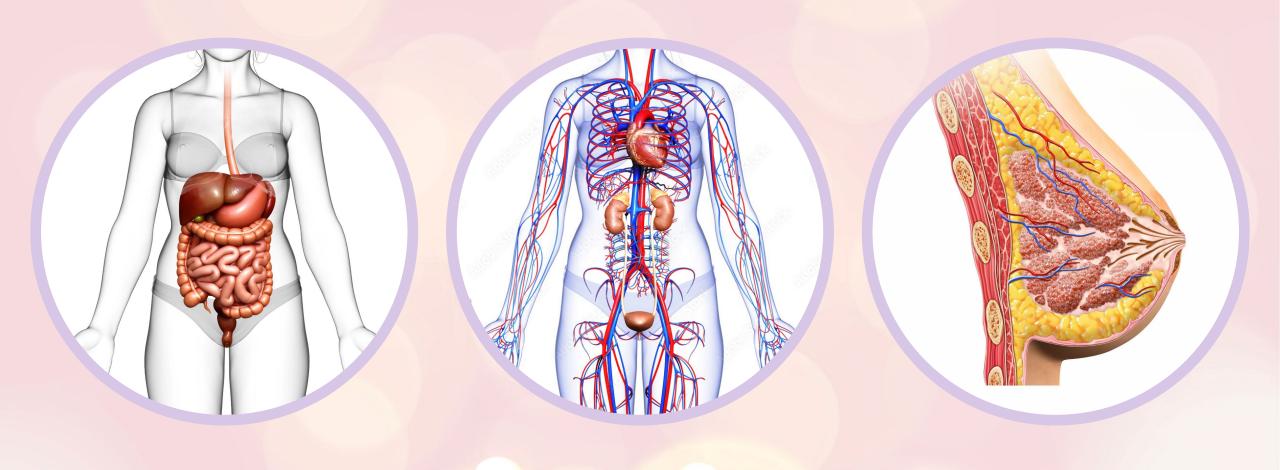
## **Triggering Components**



### **Triggering Components in Formula**

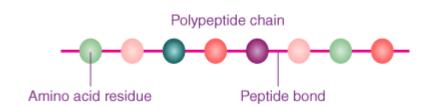


## Pathway from Lactating Parent to Child

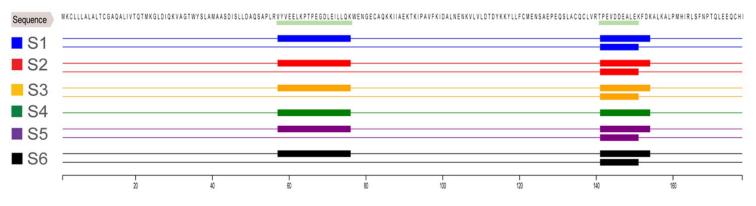


#### What is Found in the Breast?





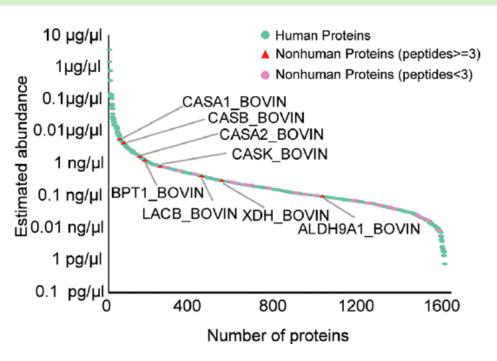
#### sp|P02754|LACB\_BOVIN Beta-lactoglobulin OS=Bos taurus GN=LGB PE=1 SV=3

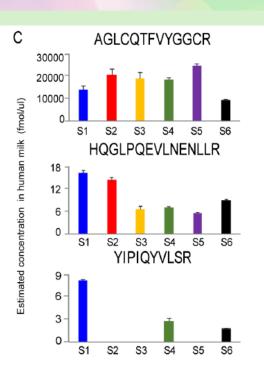


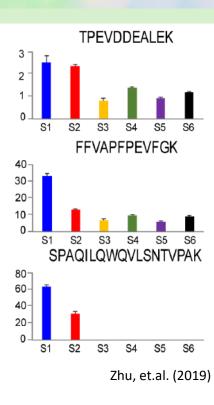




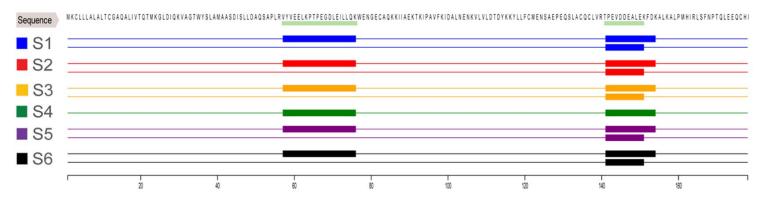
#### **Peptide Concentrations**







sp|P02754|LACB\_BOVIN Beta-lactoglobulin OS=Bos taurus GN=LGB PE=1 SV=3



## **Transferability Timing**

#### Molecular Nutrition Food Research

Review 🗈 Open Access 💿 🕦 😑 💲

Breastfeeding: Maternally Transferred Allergens in Breast Milk: Protective or Sensitizing?

Frauke Schocker 🔀 Uta Jappe

First published: 26 May 2022 | https://doi.org/10.1002/mnfr.202200066

Table 2. Synopsis of studies demonstrating allergens in human milk measured by sensitive and specific detection techniques.

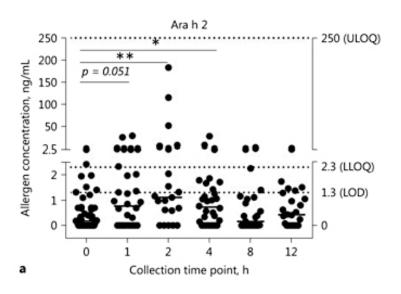
Allergen source/allergen	Concentration	Time course	Frequency	Included subjects (n)	References
MILK/ß-Lactoglobulin	110 pg–6.4 ng mL <sup>-1</sup>	4–6 h	53%	19	Kilshaw and Cant <sup>[52]</sup>
	0.01–7.84 μg mL <sup>-1</sup> (1 h)/0.01–2.34 μg mL <sup>-1</sup> (2 h)	1–2 h	75%	53	Sorva et al. <sup>[53</sup> ]
	5–800 μg mL <sup>-1</sup>	No data	40%	25	Axelsson et al.[54]
	0.9–150 μg mL <sup>-1</sup>	4–24 h	95%	20	Host et al.[55]
	0.1–16.5 μg mL <sup>-1</sup>	No data	62.5%	24	Fukushima et al.[56]
	0.41-3.80 ng mL <sup>-1</sup>	3, 6, 24 h; 3 days, 7 days	100%	15	Matangkasombut et al.[57]
WHEAT/Gliadin	5–95 ng mL <sup>-1</sup>	2–6 h	68%	53	Troncone et al.[58]
	5-1,200 ng mL <sup>-1</sup>	No data	100%	49	Chirdo et al.[59]
EGG/Ova	0.2–4.0 ng mL <sup>-1</sup>	2–6 h	74%	19	Cant et al.[60]
	0.1–1,200 ng mL <sup>-1</sup>	No data	35%	68	Vance et al.[61]
	0.12-1,258 ng mL <sup>-1</sup>	No data	35%	68	Palmer et al. <sup>[62]</sup>
	0.2–4.0 ng mL <sup>-1</sup>	2–6 h	74%	19	Kilshaw and Cant[52]
	0.7 ng mL <sup>-1</sup>	1–3, 4–8, and 9–15 h	8.3%	2	Fukushima et al. <sup>[56]</sup>
	0.25-14.03 ng mL <sup>-1</sup> (ingestion of 1 cooked egg)	2–8 h	68%	41	Palmer et al. <sup>[63]</sup>
	150 pg mL <sup>-1</sup> (median after 3 months) 173 pg mL <sup>-1</sup> (median after 6 months)	No data	49% 45%	88	Verhasselt et al. <sup>[64]</sup>
Ovomucoid	0.32-37 ng mL <sup>-1</sup>	No data	32%	37	Hirose et al.[65]
PEANUT/peanut allergens (not specified)	120–430 ng mL <sup>-1</sup>	1–3 h	48%	23	Vadas et al. <sup>[66]</sup>
Ara h 6	10-3,370 pg mL <sup>-1</sup>	10 min - 26 h	100%	2	Bernard et al.[67]
Ara h 2 and DRP*-Ara h 2	46–2,602 ng mL <sup>-1</sup>	1–12 h	26%	32	Schocker et al.[68]
Ara h 2, Ara h 6	<b>Ara h 2</b> : 2.3–184 ng mL <sup>-1</sup> <b>Ara h 6</b> : 1.1–9.7 ng mL <sup>-1</sup>	<b>Ara h 2</b> : 1–12 h <b>Ara h 6</b> : 1–12 h	Ara h 2: 35% Ara h 6: 22.5%	40	Schocker et al. <sup>[69]</sup>

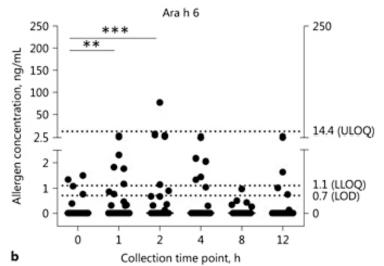
## **Transferability Timing**

# Detection of the Peanut Allergens Ara h 2 and Ara h 6 in Human Breast Milk: Development of 2 Sensitive and Specific Sandwich ELISA Assays

Frauke Schocker<sup>a</sup> Alexandra Scharf<sup>a</sup> Skadi Kull<sup>a</sup> Uta Jappe<sup>a, b</sup>

<sup>a</sup>Division of Clinical and Molecular Allergology, Research Center Borstel, Priority Research Area Asthma and Allergy, Airway Research Center North (ARCN), German Center for Lung Research (DZL), Borstel, and <sup>b</sup>Interdisciplinary Allergy Outpatient Clinic, Department of Internal Medicine, University of Lübeck, Lübeck, Germany



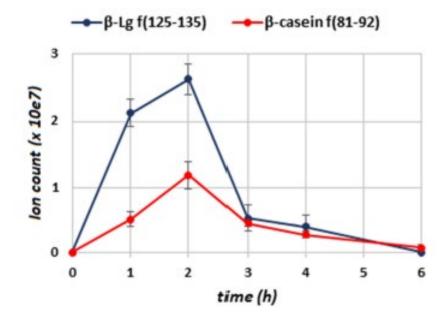




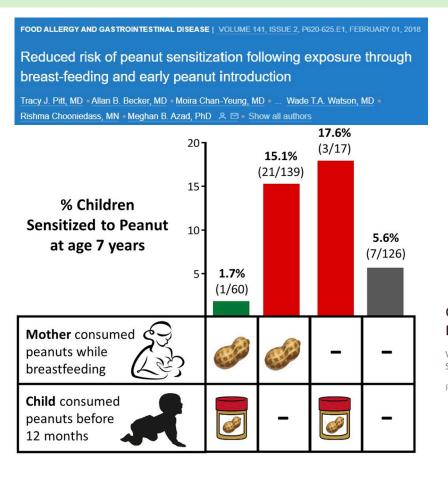
ORIGINAL RESEARCH published: 12 March 2019 doi: 10.3389/fnut.2019.00025

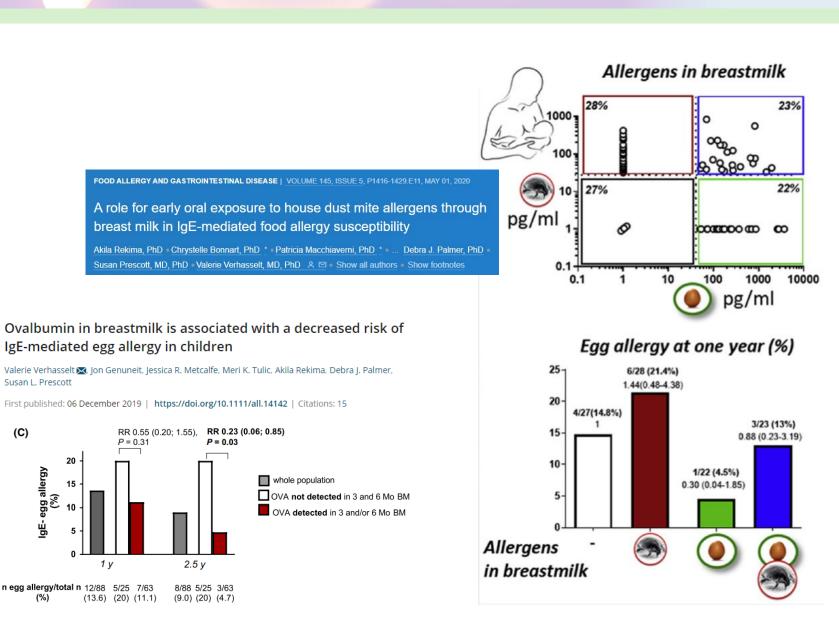
#### **Excretion of Dietary Cow's Milk Derived Peptides Into Breast Milk**

Gianluca Picariello 1\*, Maristella De Cicco 1, Rita Nocerino 2.3, Lorella Paparo 2.3, Gianfranco Mamone 1, Francesco Addeo 4 and Roberto Berni Canani 2.3,5,6\*



#### Tolerance Induction through Transfer





#### More to the Story

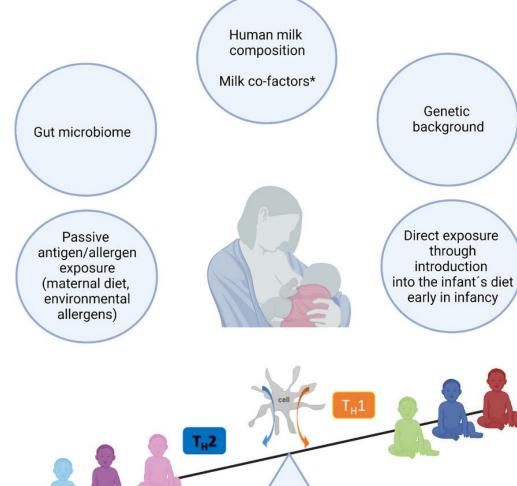
#### Molecular Nutrition Food Research

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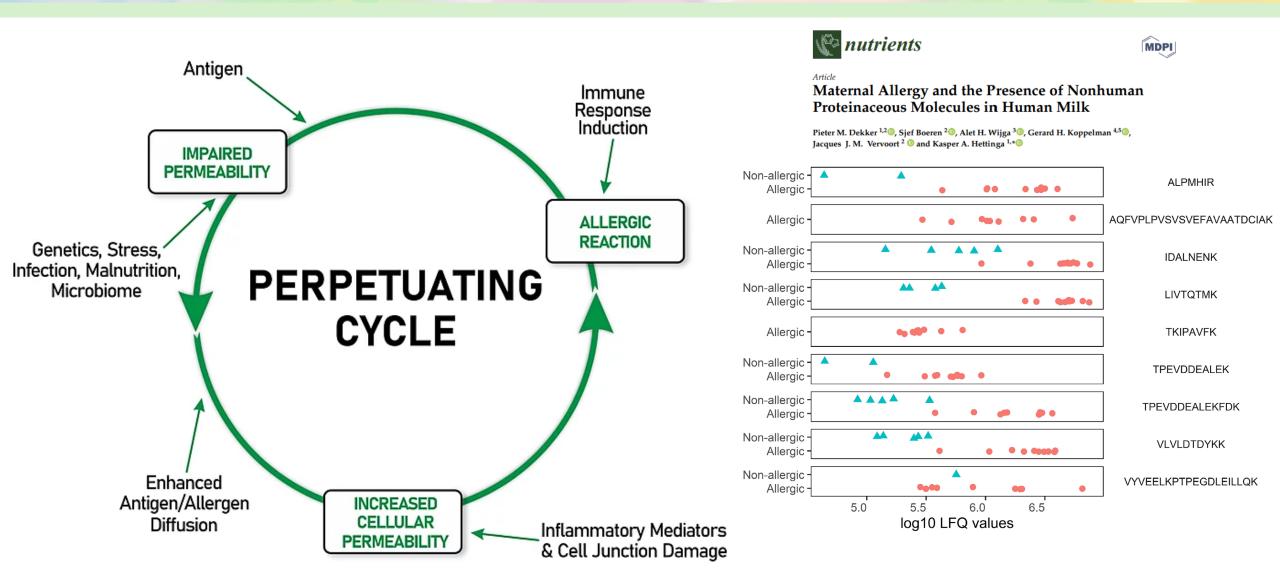
First published: 26 May 2022 | https://doi.org/10.1002/mnfr.202200066



sensitization

tolerance development

#### **Factors Involved**



### Transfer Factor Takeaways

It is <u>NOT</u> the parents fault this is occurring





Cortisol levels play an important role so consider stress reduction practices

Parent needs to consider limiting foods that elicit them a response





Minimum parent intake of 100 ounces of water each day

# Diet to Breast Transfer Timing



Ingestion

Parent consumes food

Peak

Protein concentration peaks in breast milk

Clear

Single servings typically clear in 8 hours Recommendation

Standard recommendation is to wait 24 hours for full clearance due to differences in metabolism



# Infant Reactivity & Healing

0

~0-2 hours

~6-48 hours

Post-48 hours



Ingestion



IgE Mediated & Non-IgE Acute



Non-IgE Chronic



Slow Improvement Day over Day

### Elimination Diet

Eliminating food groups/items from lactating parents diet

Allows for removal of that item from breast milk

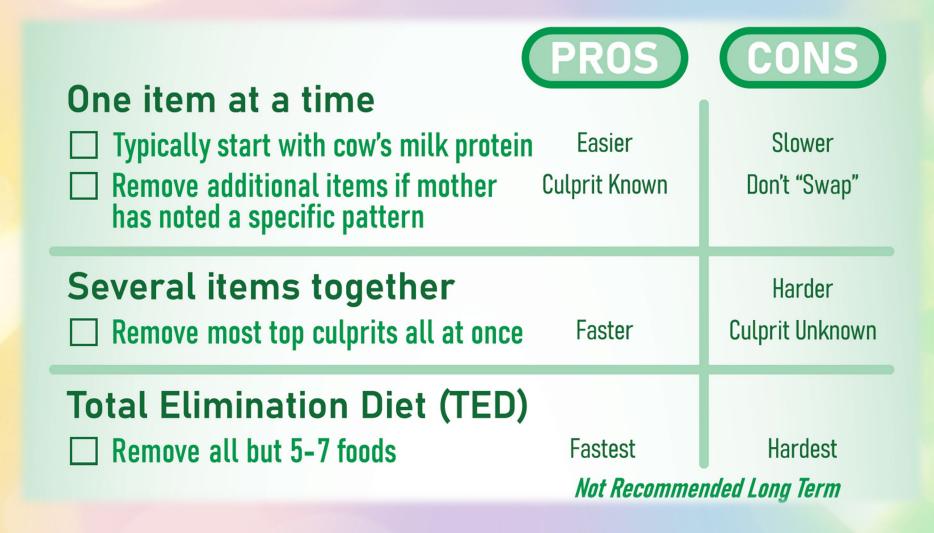
### Strategies:

- One item at a time
- Several items together
- Total Elimination Diet (TED)

Goal: Reach "baseline"



# Elimination Strategies



# Which Elimination Strategy?

### Symptomology

- FPIES, FPIAP, EoE, IgE, other
- Mild versus severe

### **Dietary Ability**

Nutritional needs and availability

#### Mental Health

Dietary restriction vs. symptoms



Mental health considerations

Postpartum depression & anxiety

Obsessive compulsive disorder

Food allergy PTSD

History of disordered eating

Family & professional support



## Light at the End of the Tunnel

Approximately 80% of food allergies occur before first birthday Many non-IgE cases will resolve themselves within 9-12 months

#### Study of 117 non-IgE milk allergic infants

- 56% tolerant by 1
- 87% tolerant by 3
- 95% tolerant by 5

IgE allergies are much less likely to develop tolerance

 Do not reintroduce IgE mediated triggers (get testing)

Wood, et.al. (2003)

### Reintroduction

Bringing foods back into the diet after a period of removal (for non-IgE)

#### Confirmation Reintroduction:

Occurs after baseline is achieved to pinpoint triggers

#### **Trigger Reintroduction:**

Occurs after proper elimination period

Different from starting solids



### Reintroduction Ladders



## Reintroduction Strategies

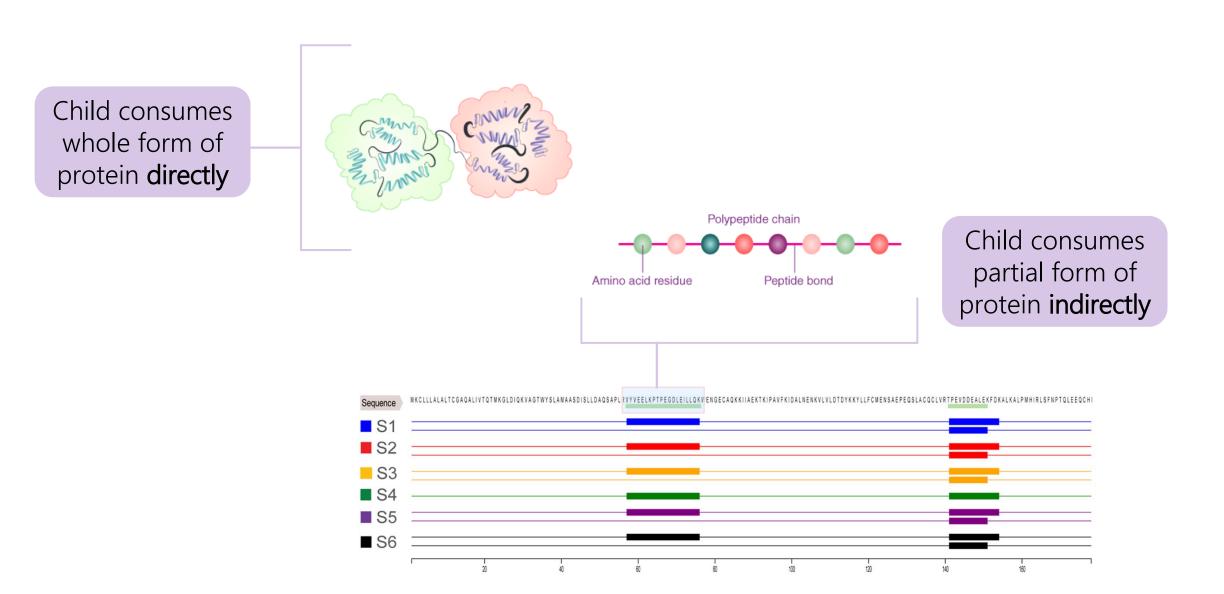
#### **Direct Introduction**

- Trigger item given directly to child
- Pro: Exact time and concentration known
- Con: Child won't eat solids, possibly stronger reaction

#### Indirect Introduction

- Trigger item trialed through breast milk
- Pro: Breastfeeding child doesn't need to do anything new
- Con: Unknown if item made it to breast milk

### Direct versus Indirect Proteins

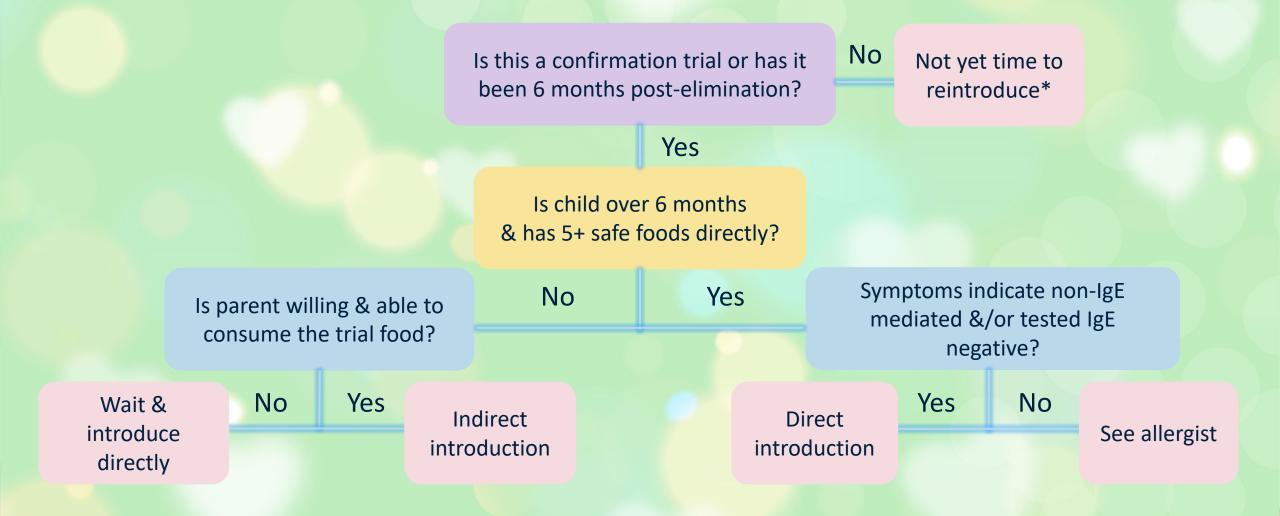


Indirect pass ≠ Direct pass



Direct pass = Indirect pass

### Reintroduction Decision Tree



## Elimination & Introduction Strategies: Takeaways



Child presents with one or more food allergy symptoms



Rule out other possible contributing factors for those symptoms



Determine best elimination diet strategy & timeline



Provide resources for tracking, nutrition, & community



At baseline, complete confirmation reintro if necessary



Re-evaluate nutrients post-confirmation & establish solids plan





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**Helping Parents Navigate Food Allergies** 



### References

- Andıran, F., S. Dayı, and E. Mete. "Cows milk consumption in constipation and anal fissure in infants and young children." Journal of paediatrics and child health 39.5 (2003): 329-331.
- Allen, Katrina J., et al. "Allergen reference doses for precautionary labeling (VITAL 2.0): clinical implications." Journal of allergy and clinical immunology 133.1 (2014): 156-164.
- Bartnikas, Lisa M., et al. "The evolution of food protein–induced enterocolitis syndrome: From a diagnosis that did not exist to a condition in need of answers." *Annals of Allergy, Asthma & Immunology* 126.5 (2021): 489-497.
- Cianferoni, Antonella. "Non-IgE mediated food allergy." Current Pediatric Reviews 16.2 (2020): 95-105.
- Chirdo, F. G., et al. "Presence of high levels of non-degraded gliadin in breast milk from healthy mothers." Scandinavian journal of gastroenterology 33.11 (1998): 1186-1192.
- Chalmers, Joanne R., et al. "Daily emollient during infancy for prevention of eczema: the BEEP randomised controlled trial." *The Lancet* 395.10228 (2020): 962-972.
- Fischler, Björn, and Thierry Lamireau. "Cholestasis in the newborn and infant." Clinics and research in hepatology and gastroenterology 38.3 (2014): 263-267.
- Gustin, Jennifer, et al. "Characterizing exclusively breastfed infant stool via a novel infant stool scale." Journal of Parenteral and Enteral Nutrition 42 (2018): S5-S11.
- Luccioli, Stefano, et al. "Maternally reported food allergies and other food-related health problems in infants: characteristics and associated factors." *Pediatrics* 122.Supplement\_2 (2008): S105-S112.
- Maciag, Michelle C., et al. "A Slice of Food Protein–Induced Enterocolitis Syndrome (FPIES): Insights from 441 Children with FPIES as Provided by Caregivers in the International FPIES Association." *The Journal of Allergy and Clinical Immunology: In Practice* 8.5 (2020): 1702-1709.
- Ollivier-Bousquet, Michele. "Transferrin and prolactin transcytosis in the lactating mammary epithelial cell." Journal of mammary gland biology and neoplasia 3.3 (1998): 303-313.
- Picariello, Gianluca, et al. "Excretion of dietary cow's milk derived peptides into breast milk." Frontiers in Nutrition 6 (2019): 25.
- Walker-Smith, J. A. "Diagnostic criteria for gastrointestinal food allergy in childhood." Clinical & Experimental Allergy 25 (1995): 20-22.
- Zhu, Jing, et al. "Discovery and quantification of nonhuman proteins in human milk." Journal of proteome research 18.1 (2018): 225-238.