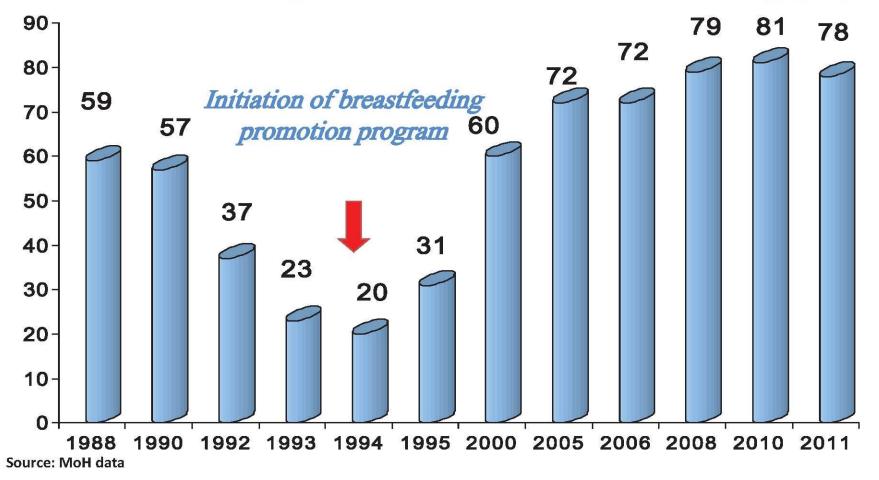
Comparison of cow-milk, breast milk and formula: nutritional, immunologic and developmental considerations

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#### Trends and prevalence of breastfeeding (%)



%	2005	2007	2010	2011
Exclusive breastfeeding for 6 months	48.4	52.2	62	59
Breastfeeding for one year and over	38	39	43	37

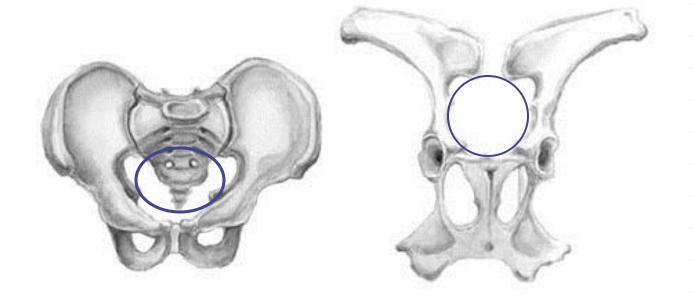
## Breast milk composition and human evolution

- Human and cows milk has evolved over millennia for the specific needs of each species
- In humans biped gate and brain size determined composition of human milk
  - Humans walked upright ~ 2 million years ago
  - Brain size increased to accommodate more intelligent behaviors and language in humans
  - Biped gate altered the structure of female pelvis from being round to wider width then length...

### **Human and Cow Pelvis**

Human female pelvis

Cow pelvis



Evolutionary solution was to deliver the fetus at decreased levels of maturity

## Human breast milk is adapted to needs of an immature newborn

Humans have adapted their
 breast milk to make up for the
 immaturity of the newborn by
 secreting bioactive components



**Duck-billed platypus** 



 Cow milk lacks these factors because there was no evolutionary pressure on the cow to develop them

# Comparison of nutritional and immune modulating factors in breast milk, cow milk and formula





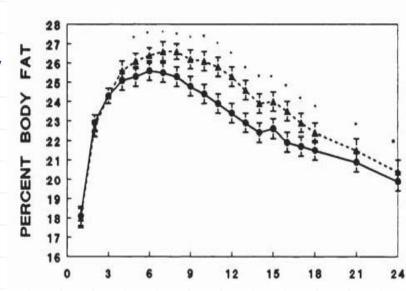


Growth differences between breastfed and formula fed infants → BREAST-FED

 Formula fed infants have significantly more body btw 3-18 mo

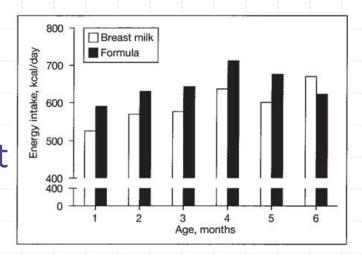
#### Possible causes

- 1. Growth factors in breast milk
- 2. Different endocrine responses to feeding
- 3. Control of food intake (self-regulation)
- 4. Nutrient composition of food-same in formula, different in breast milk



### Energy

 Equal energy density in breast milk and formula ~670 kcal/l



- Breast fed infant consume less energy
  - Not all CHO and proteins in breast milk are digestible
  - Breast fed infant take less volume (800 vs 1000 ml per day at 3 mo)
  - Breast fed infants can better regulate intake
- Energy requirement of infants might have overestimated the "true" requirements by 10-30%

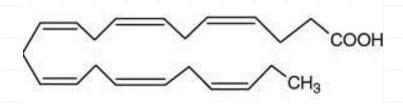
## Lipids

COOH

CH<sub>3</sub>

- Composition
  - Breast-milk
    - Colostrum—low
    - Early milk—3.5-4.0%
    - Mature milk—3.5-4.5%
  - Formula—3.5%

#### Arachidonic Acid [AA]

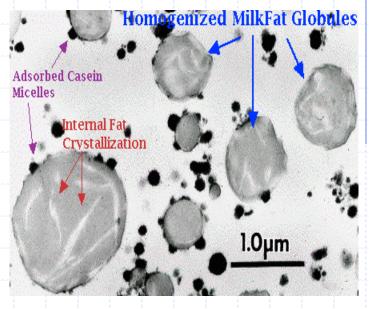


Docosahexaenoic acid [DHA]

- Differences btw breast milk and cow milk
  - Cow milk lacks long-chain polyunsaturated fatty acids (AA, DHA) because calf can synthesize them
  - AA and DHA are essential for humans and found in greatest proportion in early breast milk
  - AA and DHA improve vision and brain development problem solving at 10 mo of age
- AA and DHA are added to formula

## Lipid function-Milk Globule

- Non-polar core made of fatty acids on a triacylglycerol
- Fatty acids are arranged in specific order to allow easy digestion by lipases
- Bovine milk globule destroyed by milk pasteurization
- Results in prolonged digestion of formula (4 hrs) relative to breast milk (2 hrs)



#### Lactose in breast, bovine milk and formula

- Predominant carbohydrate in milk
  - 6.8 g/dL in human milk
  - 4.9 g/dL in bovine milk
- Structure: glucose +galactose
- Functions in humans
  - Major energy for brain--level of lactose correlates with brain size across species
  - Galactose is used to make galactolipids, especially cerebrosides needed for CNS development

#### Differences in protein composition

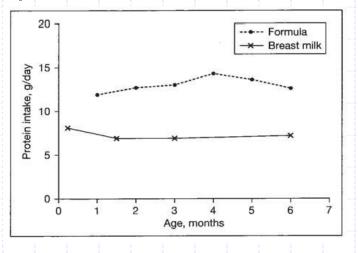
Quantity of protein in milk		
Milk type	g/L	
Colostrum	20-30	
Early Breast milk	9-11	
6 months Breast milk	8-10	
Formula	12-14	

Milk type	Whey(%)	Casein (%)
Cow	20	80
Early Breast	80	20
6 months	50	50
Formula	40	60

Differences in amino acids (**cow milk**) *More* sulfa-containing amino acids

Taurine—absent (added to formula)

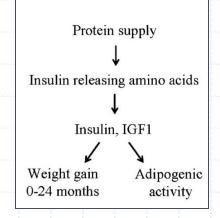
## Formula fed babies get more protein than breast fed

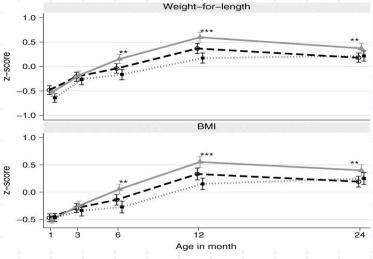


Lonnerdal B. Nestle Nutr Workshop Ser Pediatr Program. 2008;62:189-203

## Protein intake hypothesis

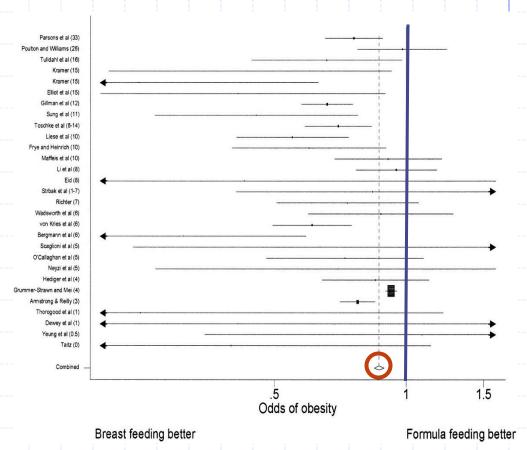
- Formula fed infants get 70% more protein than breast-fed
- Excess protein causes increased circulation of insulin releasing amino acids resulting in increased production of insulin, IGF-1 and programming for heavier infant and children later in life
- RCT of lower protein vs higher protein formula showed that at 2 yrs of age, infants on lower protein formula have lower BMI





## Odds of obesity of breast vs formula feeding, OR=0.87, 95%CI (0.85,0.89)

- Adjustment for confounding
  - Low SES
  - Maternal BMI
  - Smoking
- Reduced effect to 7%



## Breastfeeding and short term and long term offspring benefits?

Breast milk, the immune system and the gut

## Newborn's Immune System— The "Guard Dog"



Trained

Untrained

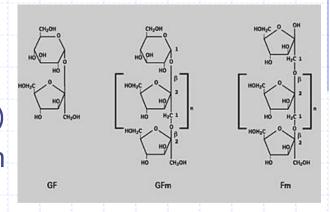
Guard dog	Immune system
Recognizing master	Recognizing self
Recognizing friend	Tolerance (allergens)
Biting thief	Killing pathogens





## Differences in gut ecology between breastfed and formula fed infants

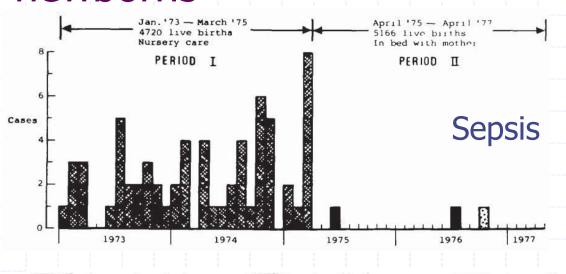
- During vaginal birth newborns GI tract is contaminated with maternal GI flora that has commensal bacteria (Bifidobacterium)
- These bacteria "train" the immune system to tolerate allergens (pollen, etc)

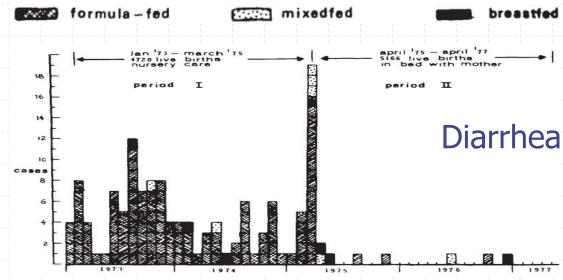


- Breast milk promotes the growth of bifidobacterium by providing prebiotic substrate for fermentation—fructo and galactooligosaccharides
- Intestine of newborn born vaginally and breastfed has more bifidobacterium and lactobacillus then formula fed infant

## Breastfeeding prevents death from sepsis and diarrhea in newborns

- Nursery in Phillippines
- Period 1—41%exclusivelybreast fed
- Period 2—85%exclusivelybreast fed





## Breastfeeding and asthma risk

#### B. Asthma/wheezing

	OR	969 Lower	6 CI Clooner
Positive family history			
Gruskay	0.85	0.21	2.62
Businco	0.26	0.03	1.28
Chandra	0.35	0.12	0.88
Hide	0.73	0.13	2.82
McCannochie	0.48	0.13	1 59
Fergusson	1.19	0.21	4.58
Marini	0.50	0.26	0.91
Subtotal	0.52	0.35	0.79

#### Negative family history or unstratified

Wilson	0.47	0.18	1.07
Oddy	0.80	0.65	0.98
Tang	0.55	0.37	0.82
Gordon	0.39	0.09	1.37
Wright	0.67	0.36	1.25
Gruskay	0.48	0.01	3.30
Hide	1 26	0.34	4 00
McConnochie	0.00	0.00	413
Fergusson	1 02	0.35	2.48
Subtotal	0.73	0.62	0.86

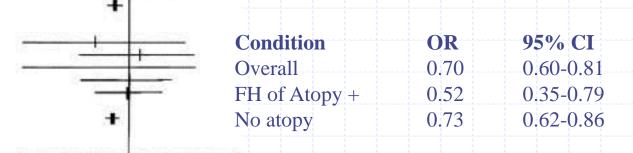
#### Children without a family history of atopy

Gruskay	0.48	0.01	3.30
Hide	1.26	0.34	4.00
McConnochie	0.00	0.00	413
Fergusson	1.02	0.35	2.48
Subtotal	0.99	0.48	2 03

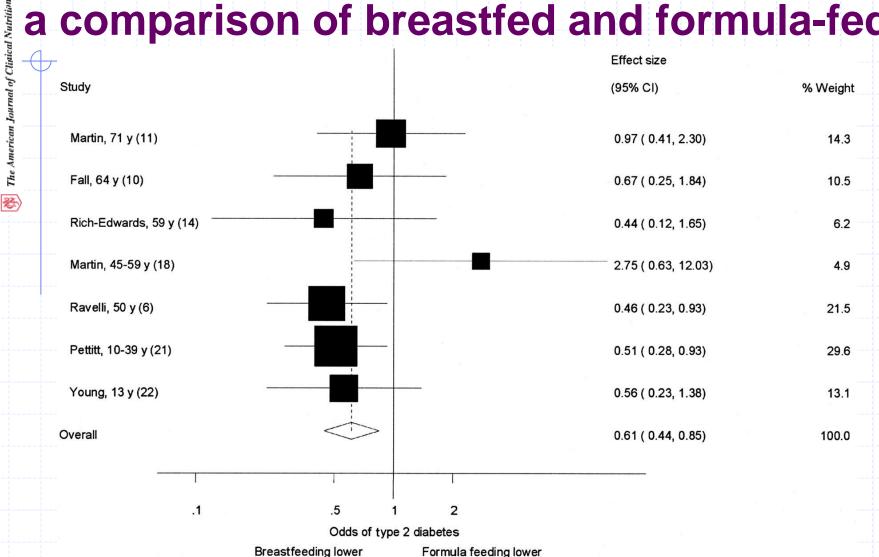
Total 0.70 0.60 0.6



 Analysis included 12 studies of >8000 children followed for 4.1 (1-8.4) years



## Odds ratios (95% Cls) of type 2 diabetes in a comparison of breastfed and formula-fed



### Breast-feeding and Inflammatory Bowel Disease

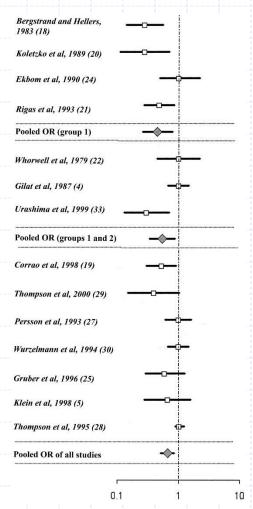
Crohn's Disease (CD), Ulcerative colitis (UC)

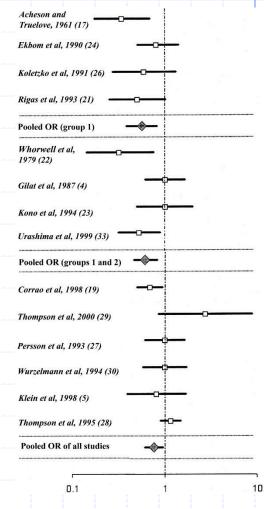
- Metaanalysis included17 studies
  - 6100 pts with CD
  - 7216 pts with UC
- Pooled estimate, random effects model
  - CD—0.67

(95% CI: 0.52, 0.86)

■ UC—0.77 (95% CI: 0.61, 0.96)

- Limitations
  - All but 2 studies were case-control studies
  - Breast-feeding definition





Klement E et al. Am J Clin Nutr 2004; 80:1342-1352 Crohn's Disease (CD)

**Ulcerative colitis (UC)** 

### Breast-feeding and cognitive development<sup>1</sup>

Metric	Effect of breastfeeding vs formula feeding	Comment
Head Circumference <sup>2</sup>	Less likely to have significant decrease in HC: OR 0.48, 95% CI 0.24, 0.99	Brazil, Dose dependent, Adjusted for SES
Brain white matter <sup>3</sup>	Dose dependent increase in CNS white matter development	
IQ <sup>4</sup>	Artificial feeding decreased IQ by 5.9 points at 6.5 yrs of age	PROBIT (Kramer in Belarus)
School achievement <sup>5</sup>	BF>9 mo associated with 0.5 to 0.8 more grades of schooling	Brazil, not associated with SES
Moving up social class <sup>6</sup>	41% (95% CI 10% to 82%) more likely to move up a social class	England, controlled for other SES variables

<sup>1.</sup> Tawia S. Breastfeeding Review 2013; 21(3): 15-20. 2. Ferreira H. Breastfeed Med. 2013;8:294-301. 3. Deoni SC. Neuroimage. 2013 Nov 15;82:77-86. 4. Kramer MS. Arch Gen Psychiatry 65: 578-584. 5. Victora S. Acta Paediatrica, 2005; 94: 1656-1660. 6. Martin RM. Arch Dis Child 92: 317-321

#### Conclusion

- Breastfeeding is "the gift that keeps on giving" throughout the entire lifecycle
- Breastfeeding is particularly important in the developing world because of:





- Prevention of infectious disease and related growth failure
- Prevention of obesity related adult disease whose prevalence is increasing
- Possible compensation for cognitive development problems conferred by poor nutrition
- Breastfeeding promotion is imperative regardless of any other interventions.