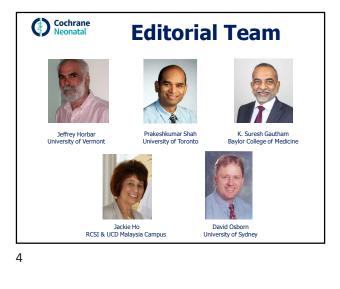




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Cochrane Neonatal

Feeding strategies and methods: Evidence from Cochrane systematic reviews

Disclosure

Roger F. Soll, M.D. is the Vice President of the Vermont Oxford Network and the Coordinating Editor of Cochrane Neonatal

No other relevant financial issues to disclose

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Goal of Nutrition for Preterm Infants

Most neonatologists have accepted the recommendation of the American Academy of Pediatrics that growth of the postnatal preterm infant, both their anthropometric indices and body composition, should be the same as the normal fetus of the same gestational age growing in its mother's uterus.

In reality, this proves to be a challenge.....

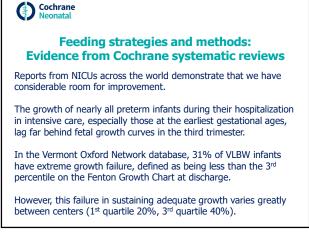
American Academy of Pediatrics Committee on Nutrition: Nutritional needs of low-birth-weight infants. Pediatrics. 1985;76:976–986.



Feeding strategies and methods: Evidence from Cochrane systematic reviews

To develop an understanding of the strengths and weaknesses of evidence provided by systematic reviews and metaanalyses to inform our practice of neonatalperinatal medicine.

Today's focus will be on optimizing feeding strategies and methods.



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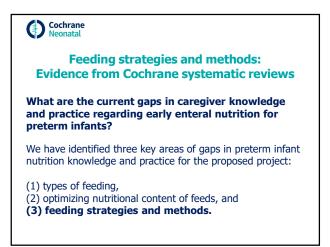
Feeding strategies and methods: Evidence from Cochrane systematic reviews

Variation in practice

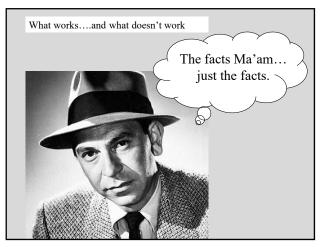
A great deal of variation is seen in feeding practices worldwide. (Klingenberg and colleagues. Arch Dis Child 2012).

For example, in infants 25 to 27 weeks GA, 100% of units in Scandinavia routinely initiate feeds in the first 24 hours of life, whereas only 15% of units in Australia report routine use.

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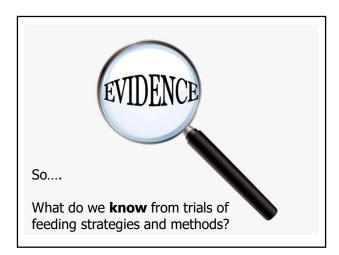
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What are the downstream effects of undernutrition and growth failure?

Numerous studies have shown that failure to meet the preterm infant's nutritional goals at critical stages of development produces serious problems for the preterm infant, including:

- short stature,
- growth failure,
- abnormalities of brain growth and development.





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Feeding strategies and methods: Evidence from Cochrane systematic reviews

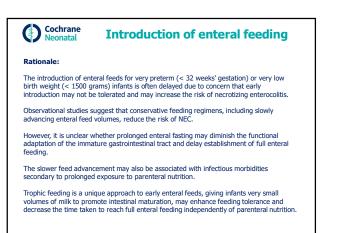
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- · Promoting enteral feeding
- Methods of enteral feeding
- Enteral feeding strategy

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Feeding strategies and methods: Evidence from Cochrane systematic reviews

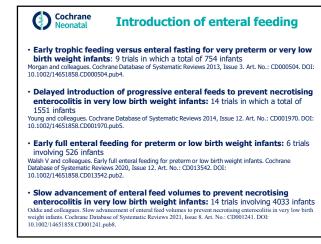
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Cochrane Neonatal Introduction of enteral feeding Effect on necrotizing enterocolitis				
Intervention	Studies	Infants	Results	
Early trophic feeding	9 studies	748 infants	RR 1.07 (95% CI 0.67 to 1.70)	
Delayed introduction of progressive enteral feeds	13 studies	1507 infants	RR 0.81 (95% CI 0.58 to 1.14)	
Early full enteral feeding	6 studies	522 infants	RR 0.98 (95% CI 0.38 to 2.54)	
Slow advancement of enteral feed volumes	14 studies	4026 infants	RR 1.06 (95% CI 0.83 to 1.37)	
No evidence of effect on necrotizing enterocolitis				

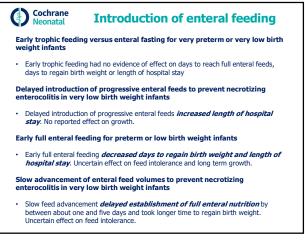


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Cochrane Neonatal Introduction of enteral feeding What exactly is the intervention?				
Intervention	Definition			
Early trophic feeding	Early trophic feeding (milk volumes up to 24 ml/kg/day introduced before 96 hours postnatal age and continued until at least one week after birth) versus a comparable period of enteral fasting			
Delayed introduction of progressive enteral feeds	Delayed (four or more days after birth) versus earlier introduction of progressive enteral feeds			
Early full enteral feeding	Early full feeding (60 mL/kg to 80 mL/kg on day one after birth) with minimal enteral feeding (typically 20 mL/kg on day one) supplemented with intravenous fluids.			
	Feed volumes were advanced daily as tolerated by 20 mL/kg to 30 mL/kg body weight to a target steady-state volume of 150 mL/kg to 180 mL/kg/day.			
Slow advancement of enteral feed volumes	Trials typically defined slow advancement as daily increments of 15 to 24 mL/kg, and faster advancement as daily increments of 30 to 40 mL/kg).			

Effect on mortality					
Intervention	Studies	Infants	Results		
Early trophic feeding	8 studies	558 infants	RR 0.66 (95% CI 0.41 to 1.07)		
Delayed introduction of progressive enteral feeds	12 studies	1399 infants	RR 0.97 (95% CI 0.70 to 1.36)		
Early full enteral feeding	6 studies	522 infants	RR 0.78 (95% CI 0.36 to 1.70)		
Slow advancement of enteral feed volumes	13 studies	3860 infants	RR 1.13 (95% CI 0.91 to 1.39)		
No evidence of effect on mortality					

Cochrane Neonatal Introduction of enteral feeding Effect on invasive infection					
Intervention	Studies	Infants	Results		
Early trophic feeding	3 studies	237 infants	RR 1.06 (95% CI 0.72 to 1.56)		
Delayed introduction of progressive enteral feeds	7 studies	872 infants	RR 1.44 (95% CI 1.15 to 1.80)*		
Early full enteral feeding	4 studies	359 infants	RR 0.72 (95% CI 0.36 to 1.46)		
Slow advancement of enteral feed volumes	11 studies	3583 infants	RR 1.14 (95% CI 0.99 to 1.31)*		
*Increased risk of infection with delayed introduction and slow advancement of feeds?					



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Effect on length of hospital stay (days)				
Intervention	Studies	Infants	Results	
Early trophic feeding	4 studies	341 infants	MD -3.9 days (95% CI -11.5 to 3.8 days)*	
Delayed introduction of progressive enteral feeds	4 studies	368 infants	MD 4.6 days (95% CI 1.5 to 7.6 days)*	
Early full enteral feeding	5 studies	436 infants	(MD -3.1 days (95% CI -4.1 to -2.0 days)*	
Slow advancement of enteral feed volumes			Variable (2 studies no difference, 2 studies report longer duration of hospital stay among infants in the slow advancement group)	
*Decreased duration of hospitalization?				

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Cochrane Introduction of enteral feeding

Rationale

Avoidance of bottles during the establishment of breast feeds in preterm infants

Preterm infants often start milk feeds by gavage tube. As they mature, sucking feeds are gradually introduced. Women with preterm infants may not always be in hospital to breastfeed their baby and need an alternative approach to feeding. Most commonly, milk (expressed breast milk or formula) is given by bottle. Whether using bottles during establishment of breastfeeds is detrimental to breastfeeding success is a topic of ongoing debate.

Non-nutritive sucking for increasing physiologic stability and nutrition in preterm infants

Sucking on a pacifier (non-nutritive sucking) during gavage feeding may encourage the development of sucking behavior and improve digestion of the feeding % f(x)

Oral stimulation for promoting oral feeding in preterm infants

A range of oral stimulation interventions may help infants to develop sucking and oromotor co-ordination, promoting earlier oral feeding and earlier hospital discharge.

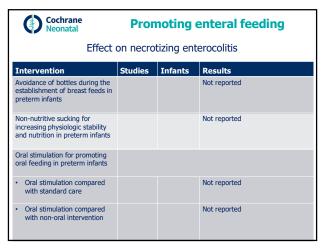
Cochrane Neonatal Introduction of enteral feeding					
What exactly is the intervention?					
Intervention	Definition				
Avoidance of bottles during the establishment of breast feeds in preterm infants	Avoidance of bottle feeds during establishment of breastfeeding included studies used a cup feeding strategy (n=5), a tube feeding strategy (n=1) and a novel teat when supplements to breastfeeds were needed (n=1).				
Non-nutritive sucking for increasing physiologic stability and nutrition in preterm infants	Sucking on a pacifier (non-nutritive sucking) during gavage feeding				
Oral stimulation for promoting oral feeding in preterm infants	A range of oral stimulation interventions may help infants to develop sucking and oromotor co-ordination, promoting earlier oral feeding and earlier hospital discharge				
Oral stimulation compared with standard care	Comparison: standard care				
Oral stimulation compared with non-oral intervention	Comparison: non-oral intervention (e.g. body stroking protocols or gavage adjustment protocols)				

	Cochrane A Neonatal	Avoidance of bottles during the establishment of breastfeeds in preterm infants.			
d	Dutcome	Studies	Infants	Relative risk (RR)	
	Full breastfeeding				
•	At discharge home	6 studies	1074 infants	RR 1.47 (95% CI 1.19 to 1.80)	
•	Three months post discharge	4 studies	986 infants	RR 1.56 (95% CI 1.37 to 1.78)	
•	Six months post discharge	3 studies	887 infants	RR 1.64 (95% CI 1.14 to 2.36)	
	Any breastfeeding				
•	At discharge home	6 studies	1138 infants	RR 1.11 (95% CI 1.06 to 1.16)	
•	Three months post discharge	5 studies	1063 infants	RR 1.31 (95% CI 1.01 to 1.71)	
•	Six months post discharge	3 studies	886 infants	RR 1.25 (95% CI 1.10 to 1.41)	
-					
	There were no harms including length of hospital stay (MD 2.25 days, 95% CI -3.36 to 7.86 days; 4 studies, 1004 infants				

Cochrane Neonatal Promoting enteral feeding Effect on length of hospital stay (days)				
Intervention	Studies	Infants	Results	
Avoidance of bottles during the establishment of breast feeds in preterm infants	4 studies	1004 infants	Mean length of hospital stay was 2.3 days higher (95% CI -3.4 lower to 7.9 days higher)	
Non-nutritive sucking for increasing physiologic stability and nutrition in preterm infants	12 studies	825 infants	Mean length of hospital stay in the intervention group was 6.8 days lower (95% CI 7.6 lower to 5.9 days lower)*	
Oral stimulation for promoting oral feeding in preterm infants				
Oral stimulation compared with standard care	5 studies	249 infants	Mean length of hospital stay in the intervention group was 4.3 days lower (95% CI 6.0 lower to 2.7 days lower)*	
Oral stimulation compared with non-oral intervention	10 studies	591 infants	Mean length of hospital stay in the intervention group was 6.1 days lower (95% CI 8.6 lower to 3.7 days lower)*	

Cochrane Neonatal	Promoting enteral feeding		
Effect on	at discharge		
Intervention	Studies	Infants	Results
Avoidance of bottles during the establishment of breast feeds in preterm infants	6 studies	1074 infants	RR 1.47 (95% CI 1.19 to 1.80)*
Non-nutritive sucking for increasing physiologic stability and nutrition in preterm infants	1 studies	303 infants	RR 1.08 (95% CI 0.88 to 1.33)
Oral stimulation for promoting oral feeding in preterm infants			
Oral stimulation compared with standard care	1 studies	59 infants	RR 1.83 (95% CI 0.96 to 3.48)
Oral stimulation compared with non-oral intervention	3 studies	301 infants	RR 1.03 (95% CI 0.86 to 1.23)

Cochrane Neonatal	Promoting enteral feeding		
	Effect on	weight g	ain
Intervention	Studies	Infants	Results
Avoidance of bottles during the establishment of breast feeds in preterm infants			Variably reported No proven effect
Non-nutritive sucking for increasing physiologic stability and nutrition in preterm infants	5 studies	221 infants	Mean weight gain (g/day) in the intervention group was 4.1 higher (95% CI 2.9 to 5.2 higher)
Oral stimulation for promoting oral feeding in preterm infants			
Oral stimulation compared with standard care			Not reported
Oral stimulation compared with non-oral intervention			Not reported

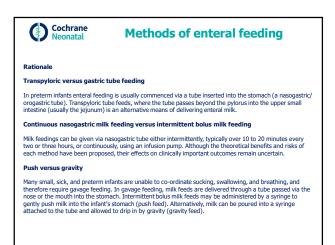


Cochrane Neonatal

Feeding strategies and methods: Evidence from Cochrane systematic reviews

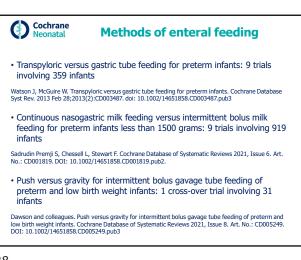
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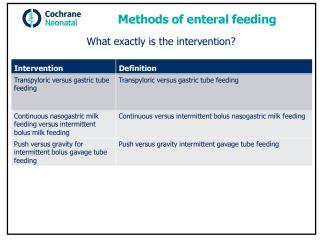


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Cochrane Neonatal Methods of enteral feeding Effect on necrotizing enterocolitis					
Intervention Studies Infants Results					
Transpyloric versus gastric tube feeding	7 studies	298 infants	RR 0.63 (95% CI 0.26 to 1.53)		
Continuous nasogastric milk feeding versus intermittent bolus milk feeding	4 studies	372 infants	RR 1.19 (95% CI 0.67 to 2.11)		
Push versus gravity for intermittent bolus gavage tube feeding			Not reported		
Effect on gastrointestinal disturbance					
Intervention	Studies	Infants	Results		
Transpyloric versus gastric tube feeding	7 studies	297 infants	RR 1.48 (95% CI 1.05 to 2.09)* RD 0.09 (95% CI 0.02 to 0.17)		



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Cochrane Neonatal	Methods of enteral feeding			
	Effect on mortality			
Intervention	Studies	Infants	Results	
Transpyloric versus gastric tube feeding	6 studies	245 infants	RR 2.46 (95% CI 1.36 to 4.46)* RD 0.16 (95% CI 0.07 to 0.26)	
Continuous nasogastric milk feeding versus intermittent bolus milk feeding			Not reported	
Push versus gravity for intermittent bolus gavage tube feeding			Not reported	
*Note: The increased mortality associated with transpyloric feeding may be due to selective allocation of the less mature and sicker infants to transpyloric feeding in the trial that contribute most weight to the meta- analysis				

Cochrane Neonatal	Methods of enteral feeding						
Effect on growth							
Intervention	Studies	Infants	Results				
Transpyloric versus gastric tube feeding	4 studies	93 infants	Mean weight gain (g/week) in the intervention group was 5.5 lower (95% CI 26.9 lower to 15.9 higher)				
Continuous nasogastric milk feeding versus intermittent bolus milk feeding	5 studies	433 infants	Standardized MD 0.09 higher (95% CI 0.27 lower to 0.46 higher)				
Push versus gravity for intermittent bolus gavage tube feeding			Not reported				
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Enteral feeding strategy
 High versus standard volume enteral feeds to promote growth in preterm or low birth weight infants: 3 trials involving 347 infants
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Cochrane Neonatal Enteral feeding strategy What exactly is the intervention?				
Intervention	Definition			
High versus standard volume enteral feeds to promote growth in preterm or low birth weight infants	In infants who were fed fortified human milk or preterm formula, high and standard volume feeds were defined as > 180 mL/kg/day and ≤ 180 mL/kg/day, respectively. In infants who were fed unfortified human milk or term formula, high and standard volume feeds were defined as > 200 mL/kg/day and ≤ 200 mL/kg/day, respectively			
Responsive versus scheduled feeding for preterm infants	Policy of feeding preterm infants on a responsive basis versus feeding prescribed volumes at scheduled intervals			
Short versus long feeding interval for bolus feedings in very preterm infants	Short (e.g. one or two hours) versus long (e.g. three or four hours) feeding intervals			





Cochrane Neonatal	Enteral feeding strategy Effect on growth				
Intervention	Studies	Infants	Results		
High versus standard volume enteral feeds (g/kg/day)					
 with fortified human milk or preterm formula 	2 studies	271 infants	MD 2.6 g/kg/day higher (95% CI 1.4 g/kg/day higher to 3.8 g/kg/day higher)*		
 with unfortified human milk or term formula 	1 study	61 infants	MD 6.2 g/kg/day higher (95% CI 2.7 g/kg/day higher to 9.3 g/kg/day higher)*		
Responsive versus scheduled feeding (g/kg/day)	4 studies	305 infants	Mean weight change during study period in the intervention group was 1.4 g/kg/day lower (95% CI 0.3 to 2.4 g/kg/day lower)*		
Short versus long feeding interval for bolus feedings			Not reported		

Effect on length of hospital stay (days)						
Intervention	Studies	Infants	Results			
High versus standard volume enteral feeds						
with fortified human milk or preterm formula	2 studies	271 infants	The mean duration of hospital stay in the intervention group was 1.0 day longer (95% CI 3.5 days shorter to 5.5 days longer)			
with unfortified human milk or term formula			Not reported			
Responsive versus scheduled feeding	2 studies	145 infants	The mean duration of hospital stay in the intervention group was 1.0 days shorter (95% CI 9.4 days shorter to 7.3 days longer)			
Short versus long feeding interval for bolus feedings	2 studies	207 infants	The mean duration of hospital stay in the intervention group was 3.4 days shorter (95% CI 9.2 days shorter to 2.5 days longer)			



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Cochrane Neonatal **Enteral feeding strategy** Effect on necrotizing enterocolitis Studies Infants Results Intervention High versus standard volume enteral feeds with fortified human milk or 2 studies 283 infants RR 0.74 (95% CI 0.12 to 4.51) preterm formula with unfortified human milk or term formula Not reported Responsive versus scheduled Not reported feeding Short versus long feeding 4 studies 417 infants RR 1.07 (95% CI 0.54 to 2.11) interval for bolus feedings

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Questions regarding optimizing feeding strategies and methods in preterm infants....

Where does the evidence take us?

What are best "practices" regarding optimizing feeding strategies and methods in preterm infants?

What future research is urgently needed?



