

Infant malnutrition and growth failure in the community: Exploring new nutritional management strategies

Presenter: Liz Bacon, MS, RD, LD, CSP - Medical Science Liaison, Nutricia

Live event date: September 16, 2021 - Recording on NutriciaLearningCenter.com within ~2 weeks of live event



Learning Objectives:

- Identify infant malnutrition and growth failure
- Discuss common interventions for infant malnutrition and growth failure
- List the risks associated with concentrating and fortifying formula
- Identify a novel way of managing term infants with malnutrition and growth failure

Notes:			

Nutricia North America supports the use of breast milk wherever possible.

A.S.P.E.N. Definition of Malnutrition

"an imbalance between nutrient requirement and intake, resulting in cumulative deficits of energy, protein or micronutrients that may negatively affect growth, development and other relevant outcomes."

Mehta, et al. J Parenter Enteral Nutr 2013;37(4):460-81

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Prevalence of Growth Failure 10% of infants in primary care settings 4% of hospitalized infants² 1. Daymont C, et al. Acad Pediatr. 2020;20:405-12. 2. Carvalho-Salemi, et al. J Acad Nutr Diet. 2018;118:40-51.e7.

Inadequate Intake	Increased Energy Needs	Inadequate Absorption
Reflux	Chronic lung disease	GI conditions
Supply concerns	Congenital heart disease	Cystic Fibrosis

Improper mixing Renal failure Milk protein allergy

Feeding difficulties Hyperthyroidism Inborn errors of metabolism

Neglect/abuse Chronic infection

Ftiologies of Growth Failure

Acute illness

Inadequate energy intake is the **most common** cause of growth failure for children with and without chronic disease

Carvalho-Salemi J, et al. J Acad Nutr Diet. 2018;118(1):40-51.

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Risk Factors for Growth Failure

Medical

- Low birth weight
- Reflux
- Congenital anomalies
- Chronic illness
- Acute illness

Psychosocial

- Poor feeding techniques
- Post-partumDepression (PPD)
- Poverty
- Abuse

Homan. American Family Physician. 2016;94(4)295-300.

Primary indicators	When Single Data	Points Available		
	Weight-for-height z score	BMI-for-age z score	Length-/height-for-age z	Mid-upper arm circumference
Mild Malnutrition	-1 to -1.9 z score	-1 to -1.9 z score	No data	≥ -1 to -1.9 z score
Moderate Malnutrition	-2 to -2.9 z score	-2 to -2.9 z score	No data	≥ -2 to -2.9 z score
Severe Malnutrition	-3 or greater z score	-3 or greater z score	-3 z score	≥ -3 z score
			0.2 30010	2 -0 2 30016
Primary Indicators	When <u>Two or More</u> Weight gain velocity (<2 years of age)		Deceleration in weight- for-length/height z score	Inadequate nutrient intake
Primary Indicators	When Two or More Weight gain velocity	Points Available:	Deceleration in weight-	Inadequate nutrient
	When Two or More Weight gain velocity (<2 years of age) <75%1 of the norm2	Points Available: Weight loss (2-20 years of age)	Deceleration in weight- for-length/height z score	Inadequate nutrient intake 51-75% estimated

Implications of Growth Failure

"The first 1000 days"

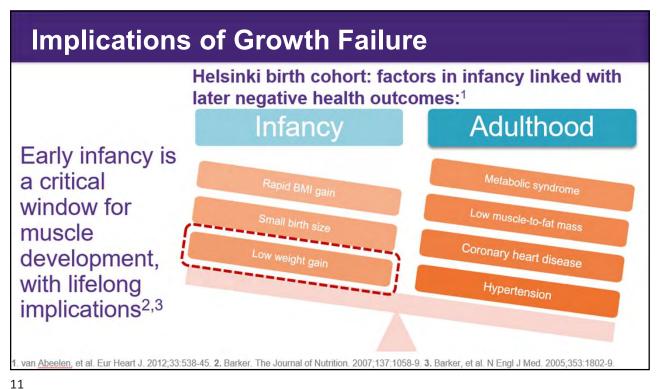
Critical period of brain growth and development

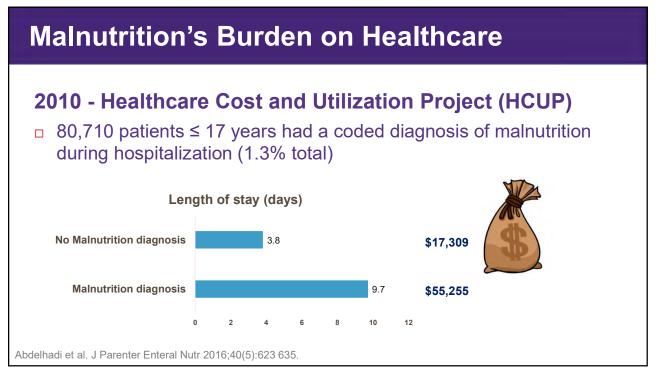
Inadequate nutrition support:

- Cognitive developmental delays
- Decreased growth potential
- Decreased immune function



Georgieff MK et al, Acta Paediatr. 2018:107:1310-21





Case Study – Baby Sophie

History

- · Healthy, term, infant
- · 3 months old
- · Hx of adequate growth
- Breastfed

Diagnosis

- RSV
- Acutely ill
- · Decreased appetite
- · Difficulty feeding due to fatigue

Anthropometrics

Weight/length z-score -1.1 (mild malnutrition)

Mom's Request

· Mom wants to continue breastfeeding

Hx = history; RSV = respiratory syncytial virus



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Troubleshooting Growth Failure

Troubling trends on growth curve

OR

Meets criteria for malnutrition diagnosis

- Does child have an acute or chronic illness?
- What is infant's feeding regimen?
- Has volume, frequency, or type of feeds changed?
- If mother is breastfeeding or pumping, how is her supply and/or feeding quality?
- Signs of intolerance?
- How are caregivers preparing formula?
- Is infant on a non-standard formula recipe?
- Is socio-economic status a concern?

Current Practices for Infants with Growth Failure

- Feed more volume, more often.
- Concentrate formula.
- Use a higher-calorie formula at standard concentration.
- Start with a "base" and then supplement or fortify.



Simental S. J Pediatr Gastroenterol Nutr. 2020;71:S453.

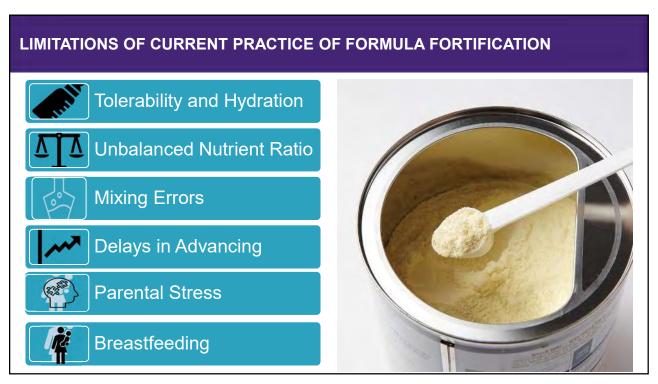
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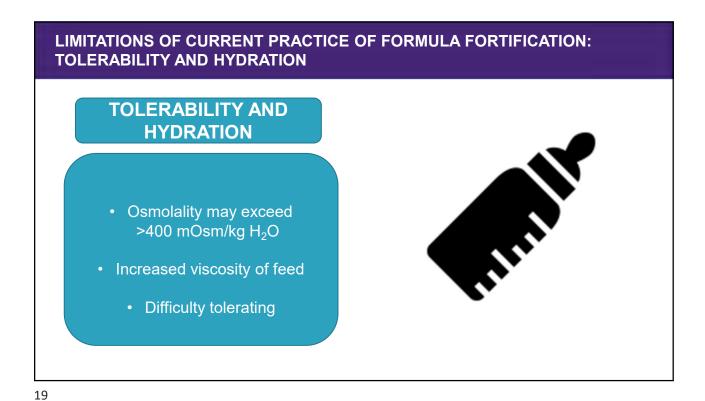
RDs often use formula and/or modulars to feed hospitalized infants with FTT¹ Common enteral nutrition sources (n=25: each chose top 3) Modular(s) sometimes used to help meet 100% of protein +/or energy goals (n=21: each chose all that apply) 84% % RESPONDENTS 56% % RESPONDENTS 36% Standard eHF AAF Breast Preemie Preemie Use 1+ Use Use Use CHO+ Use PRO+ Use milk formula HP modulars **PRO** FAT All 3 AAF = amino acid-based formula; CHO = carbohydrate; eHF = extensively hydrolyzed formula; Prem-HP = high-protein premature formula; PRO = protein; RD = registered dietitian. Nutricia supports the use of breast milk wherever possible. 1. Simental. Poster at NASPGHAN Annual Meeting. J Pediatr Gastroenterol Nutr. 2020;71:S453(684).

POLL – respond in the right-hand panel

Which practice do you use most often for feeding infants with growth failure?

- A. Increase volume and frequency of feeds
- B. Concentrate powdered formula
- C. Use a higher-calorie formula at standard concentration
- D. Start with a "base" and then supplement or fortify



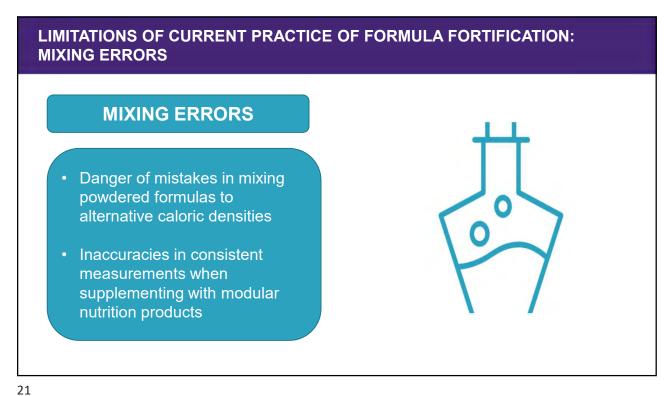


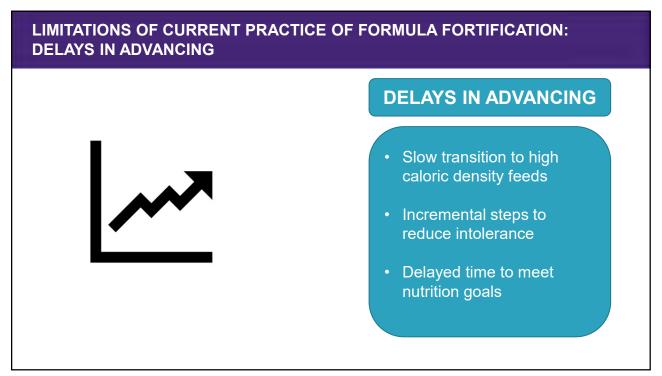
LIMITATIONS OF CURRENT PRACTICE OF FORMULA FORTIFICATION:
UNBALANCED NUTRIENT RATIO

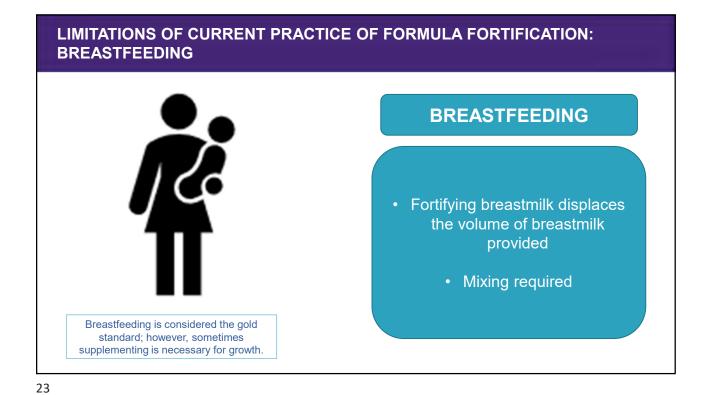
UNBALANCED NUTRIENT RATIO

Increasing calories can skew ratio of other components

Macronutrients not tailored for target protein needs of 9-12% of calories







Parental Stress

Parental Stress

Additional burden on caregivers or staff

Understanding complex mixing instructions

Additional time needed to prepare feeds

POLL – respond in the right-hand panel

Which limitation of fortifying formula most concerns you?

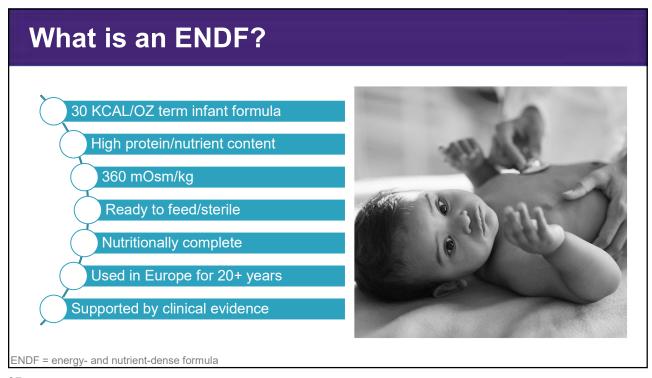
- A. Poor tolerance
- B. Inadequate protein
- C. Inadequate hydration
- D. Mixing errors
- E. Delays advancing
- F. Parental stress
- G. Displaces breastmilk intake

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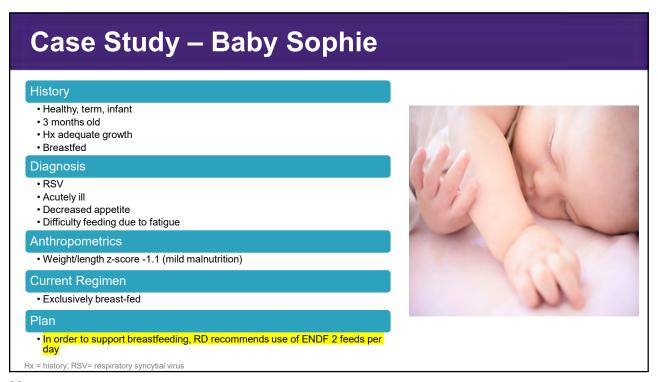
A novel way of managing infants with high-calorie needs

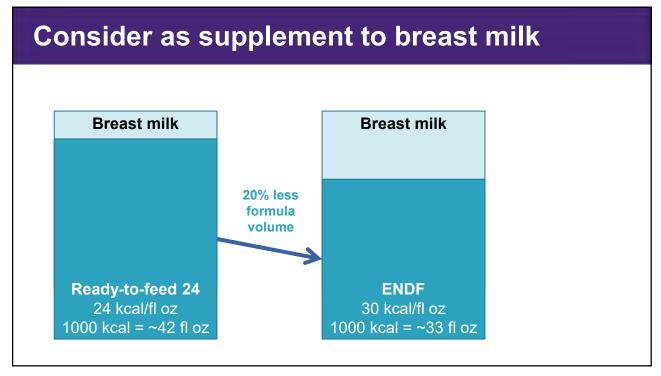
Energy- and nutrient-dense infant formulas (ENDF)



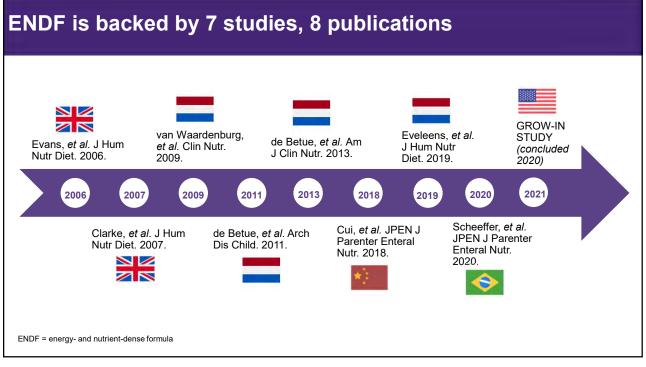












Earliest studies established ENDF as safe, well-tolerated, and supportive of catch-up growth in infants with growth failure

Randomized comparison of a nutrient-dense formula with an energy-supplemented formula for infants with faltering growth

2007 - Clarke et al.



Should high-energy infant formula be given at full strength from its first day of usage?

2006 – Evans et al



ENDF = energy- and nutrient-dense formula

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Clarke et al. aimed to compare energy- and nutrient-dense formula (ENDF) to the current practice of energy supplemented formulas (ESF)

Design

- Randomized, open-label, controlled trial
- 6-week intervention
- At least 80% of kcal from study formula
- Study formulas

Intervention (n = 26)	Control (n = 23)
ENDF • 1 kcal/mL • 2.6g protein/ 100kcal (10.4% PE)	ESF • 1 kcal/mL • 1.4g protein/ 100kcal (5.5% PE)
Birmingham Children'	s Hospital, 1997 - 98

Study Population

- Enterally fed infants <12 months old with diagnosis of FTT due to organic or non-organic causes
- Diagnoses in study population:
 - Congenital heart disease: 47% (n=23)
 - Gl/surgical patients: 31% (n=15)
 - Cystic Fibrosis: 10% (n = 5)
 - Other: 12% (n=6)
- No significant differences in anthropometry, sex, or biochemistry

Clarke et al. noted a trend toward better growth for males in the ENDF group



Outcomes & Results: Anthropometry

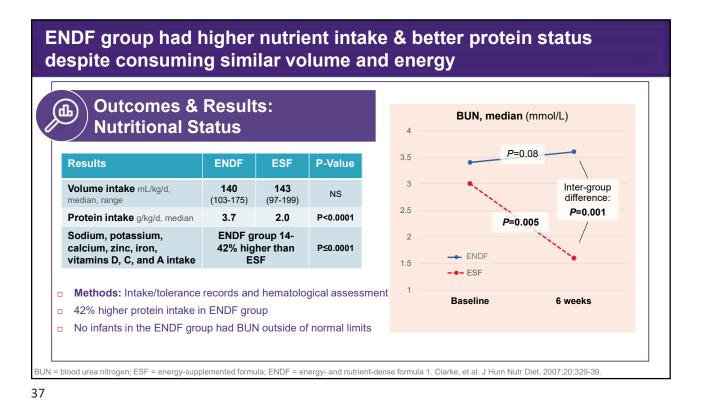
- Methods: Length, weight, MUAC measurements done by a single trained observer
- Outcomes: Significant improvement in WFA & MUAC z-score overall for both groups
- □ LFA z-score trended negatively for both groups (>males)
- Only significant for males in ESF group



ESF = energy-supplemented formula; ENDF = energy- and nutrient-dense formula WFA = weight for age; MUAC = mid-upper arm circumference; LFA = length for age; . 1. Clarke

et al. J Hum Nutr Diet. 2007:20:329-39

ENDF su	ıpports grow	th				
	Within ENDF group (n=26) (n=14 male) (n=12 female)*	Within ESF group (n=23) (n=12 male) (n=11 female)*	Between groups†	ESF = energy- supplemented formula; ENDF = energy- and nutrient-dense		
	WFA z-scores			formula WFA weight for age		
Q 07	0.29 P = 0.007	0.49 P = 0.006	P = 0.26	*Within-group differences: Wilcoxon signed rank test. †Between- group differences:		
MALES	0.21 P = 0.02	0.40 P = 0.24	P = 0.98			
Q FEMALES	0.32 P = 0.16	0.86 P = 0.01	P = 0.12	differences: Mann–Whitnetest. Female lengt		
	Length-for-age z-scores					
Q 07	Q -0.18 -0.28 P = 0.01					
MALES	-0.16 P = 0.42	-0.80 P = 0.002	P = 0.02	J Hum Nutr Diet. 2007;20:329- 39.		



ENDF was as safe and well-tolerated as ESF



- No difference in number of daily emesis or stools
- No safety concerns noted for either formula

Results	ENDF	ESF	P-Value
Stools/d median, range	2.5 (0.7-4.5)	2.5 (1.0-4.8)	NS
Emesis/d median, range	0.33 (0-1.8)	0.30 (0-5)	NS
Volume intake mL/kg/d, median, range	140 (103-175)	143 (97-199)	NS

ESF = energy-supplemented formula; ENDF = energy- and nutrient-dense formula 1. Clarke, et al. J Hum Nutr Diet. 2007;20:329-39.

Evans et al. compared tolerance of ENDF introduced at full strength versus by graded introduction

Randomized comparison of a nutrient-dense formula with an energy-supplemented formula for infants with faltering growth

2007 - Clarke et al.



ENDF = energy- and nutrient-dense formula

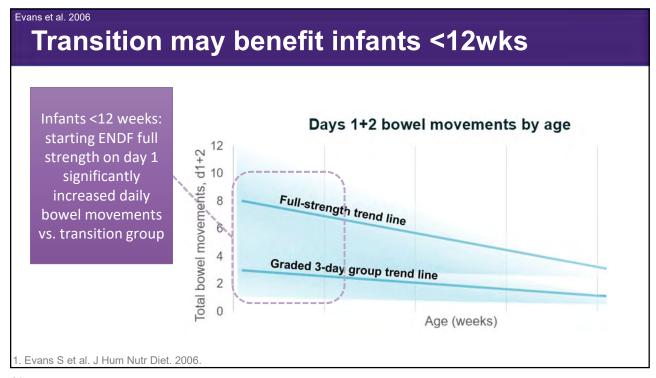
Should high-energy infant formula be given at full strength from its first day of usage?

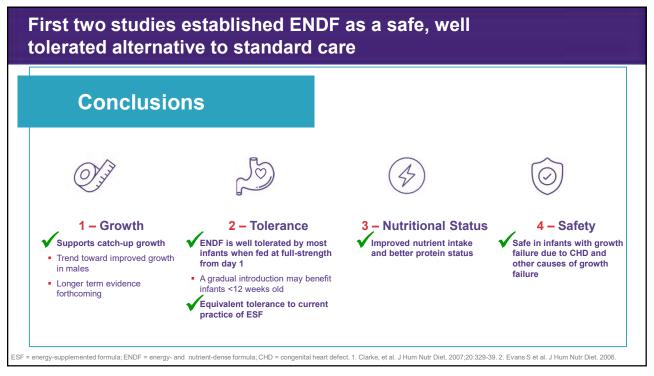
2006 – Evans et al

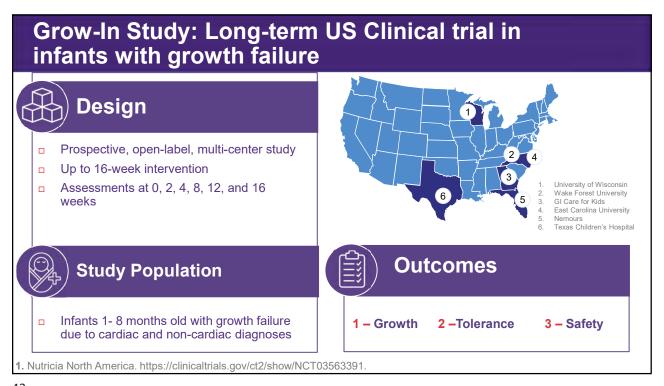


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Some infants may benefit from transitioning onto ENDF Full Strength (FS) Start **Graded Start** Design (n = 18)(n = 12)Full strength at day 1 ENDF diluted with water Day 1: 80% (24 kcal/ fl. oz) Randomized, controlled trial (non-blinded) Day 2: 90% (27 kcal/ fl. oz) 2-week intervention with ENDF Day 3:100% (30 kcal/ fl. oz) Birmingham Children's Hospital **Outcomes & Results** Study Population 1 - Tolerance: Emesis & Stool 2 - Anthropometry: Infants with diagnosis of failure to thrive Weight, Length, HC, MUAC 73% (n=22) of the infants had CHD ↑ stool frequency in first 2 Non-significant trend days for FS group to higher weight for FS 93% (n=28) standard formula (20 kcal/fl oz) group (p=0.02), inversely previously correlated with age No differences in emesis ENDF = energy- and nutrient-dense formula; CHD = congenital heart defect; FS = full strength. 1. Evans S et al. J Hum Nutr Diet. 2006





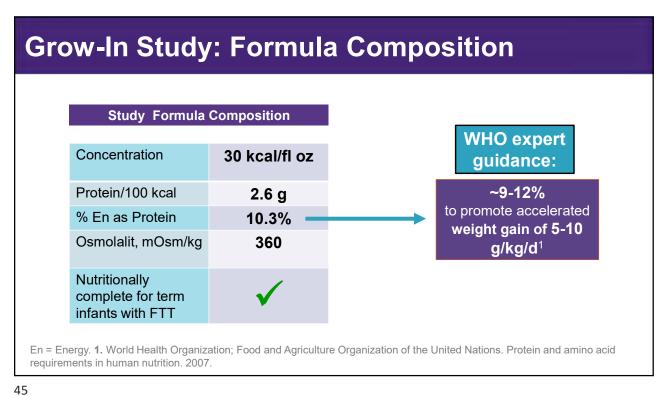


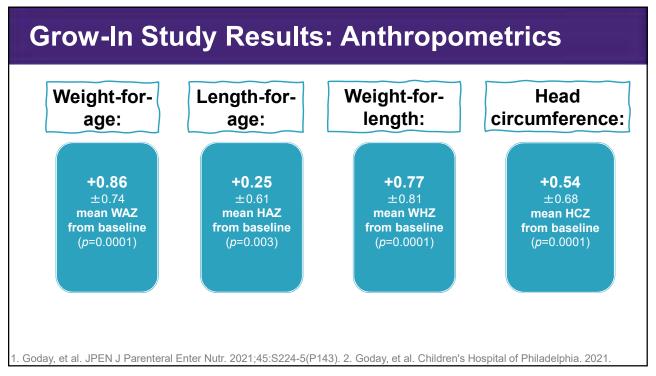
Grow-In Study: Population Characteristics

Variable	Result
Gender: n (%)	
male	16 (61.5%)
female	10 (38.5%)
Gestational age*	37.4 ± 3.2
Age at Visit 1**	22.2 ± 10.5
WAZ at birth (mean)	-0.19
WAZ at baseline (mean)	-2.92

N=26 Per Protocol subjects. WAZ = weight-for-age z-score *Median weeks **Mean Weeks.

1. Goday, et al. JPEN J Parenteral Enter Nutr. 2021;45:S224-5(P143). 2. Goday, et al. Children's Hospital of Philadelphia. 2021.





Grow-In Study: Rate of catch-up growth

Percentage of infants who achieved weight gain velocity >WHO median:

CA.

83%

At ≥1 time point

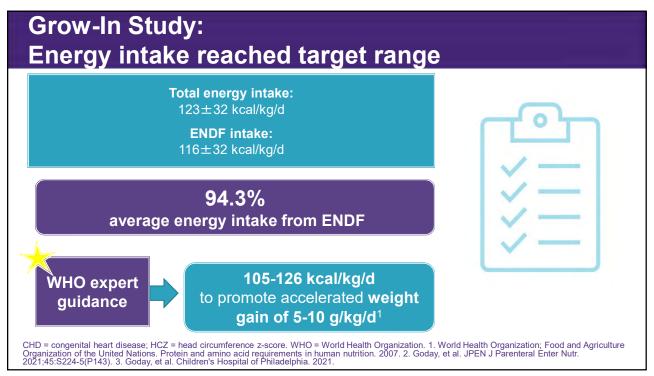
67%

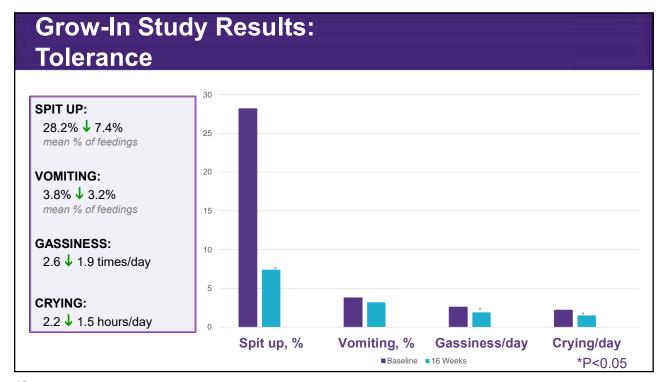
For overall study period

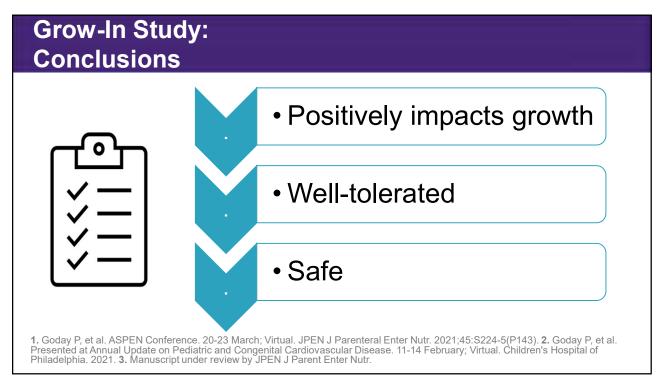
13%

Met criteria for "early success"* No difference in time to meeting weight gain velocity target by presence of CHD

*Early success = WHZ ≥0 or weight velocity > +2 z-score for age at two consecutive visits: transitioned to lower energy-density formula. Two infants in the PP group withdrew early and are not included. 1. Goday, et al. JPEN J Parenteral Enter Nutr. 2021;45:S224-5(P143). 2. Goday, et al. Children's Hospital of Philadelphia. 2021.







ENDF is well tolerated and supports nutrient intake, growth and anabolism								
2006 Evans	2007 Clarke	2009 2011 2013 van de Betue de Betue Waardenburg		2018 Cui	2019 Eveleens	2020 Scheeffer	2021 Goday	
Clinical trial	Clinical trial	Clinical trial 3 publications from 1 trial		Clinical trial	Retrospective review	Clinical trial	Clinical trial	
1	1	✓		1	1	1	1	
Not studied	1	✓		1	Not studied	Not studied	Not studied	
Not studied	Not measured	✓		1	Not studied	Not studied	Not studied	
Not designed to study growth	1	Too brief to study growth		Too brief to study growth	1	1	1	
Not studied	Not studied	✓		1	Not studied	Not studied	Not studied	
	1				1	1	1	1
	2006 Evans Clinical trial Not studied Not studied Not designed to study growth	2006 Evans Clinical trial Clinical trial Not studied Not studied Not measured Not designed to study growth	Clinical trial Not studied Not study growth Not study growth Not study growth Clinical trial Not designed to study growth	Clinical trial Not studied Not studied Not designed to study growth	2006 Evans Clarke Vaan Waardenburg Clinical trial Clinical trial Clinical trial Not studied Not measured Not designed to study growth	2006 Evans Clarke Van Waardenburg Clinical trial Clinical trial Clinical trial 3 publications from 1 trial Not studied Not measured Not designed to study growth Too brief to study growth	2006 Evans Clarke Vaar Waardenburg Clinical trial Too brief to study growth Clinical trial Retrospective review Not studied Not studied Not studied Too brief to study growth Too brief to study growth	2006 Evans Clarke Van Waardenburg Clinical trial Clinical trial 3 publications from 1 trial Not studied Not measured Not study growth Too brief to study growth 2018 2019 Eveleens Scheeffer Clinical trial Clinical trial Retrospective Clinical trial review Not studied Not studied Not studied Not study growth Too brief to study growth

POLL – respond in the right-hand panel

Which finding related to energy- and nutrient-dense formula is most relevant to your practice?

- A. WELL TOLERATED: 30 kcal/fl oz ENDF is tolerated as well as 20 kcal/fl oz formula and energy-supplemented formula
- B. SUPPORTS ANABOLISM: ENDF supports positive nitrogen balance and anabolism
- C. FOR CHD INFANTS: 30 kcal/fl oz formula has been studied in infants with congenital heart defects
- **D. HIGHER NUTRIENT INTAKE**: 30 kcal/fl oz ENDF supports higher protein and nutrient intake than standard formula

USE OF ENERGY NUTRIENT DENSE FORMULAS

WHEN ENERGY NUTRIENT DENSE FORMULAS ARE BENEFICIAL

Increased energy and protein requirements

Catch-up growth for undernourished or at risk of growth failure

Demonstrating poor growth on current regimen

Achieve protein-calorie goals earlier during period of fluid restriction

Higher Protein: Energy ratio to promote positive nitrogen balance during illness

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ENDF PATIENT POPULATION Term infants with: Congenital Chronic Lung Heart Disease Growth Failure Disease At-Risk of Growth Failure Neurological Cystic Fibrosis Syndromes Increased Energy Requirements Non-Disease-□ Fluid Restriction Respiratory Related Failure Syncytial Virus to Thrive

INTRODUCING ENERGY NUTRIENT DENSE **FORMULAS (ENDF)**

GUIDELINES TO SUCCESSFULLY INTRODUCE ENERGY NUTRIENT DENSE FORMULAS (ENDF)

Infants <12 weeks of age:

Administer full strength formula and alternate with current feed or dilute to 24 kcal/oz and grade to full strength over 3 days to avoid frequent stools

Infants >12 weeks to 18 months:

Full strength and full transition from day 1

Alternate breast milk/breast feeding and ENDF feeding

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Case Study – Baby Sophie

History

- · Healthy, term, infant
- · Adequate growth
- Breastfed

Diagnosis

- RSV
- · Acutely ill
- · Decreased appetite
- · Difficulty feeding due to fatigue

Anthropometrics **Section**

• Weight/length z-score -1.1 (mild malnutrition)

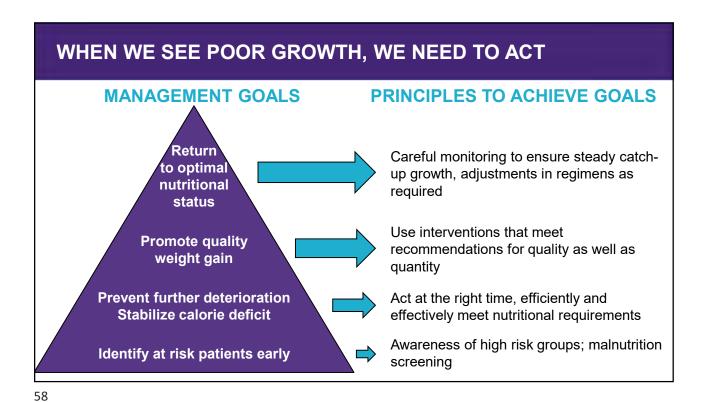
Mom's Request

· Mom wants to continue breastfeeding

• In order to support breastfeeding, RD recommends use of ENDF 2 feeds per day

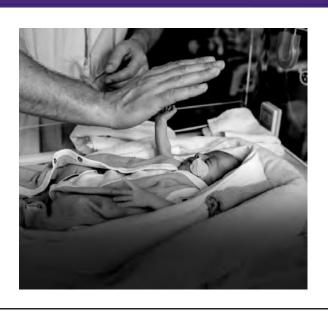
- 3 weeks later weight/length z-score is 0.3
 ENDF supported growth and well tolerated
- Infant able to resume exclusive breastfeeding





SUMMARY

- Malnutrition has serious implications for infants. It's critical to prevent, identify and resolve malnutrition in a timely and safe manner.
- ENDF provide optimal energy, protein, and micronutrients to support lean tissue gain for catch-up growth and support increased protein needs during critical illness.
- ENDF have equivalent tolerability to 20 kcal/oz infant formulas.
- Clinical research has shown ENDF promote catch-up growth in disease and non-disease related growth failure.



ENDF = energy- and nutrient-dense formula

References

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 - 25. World Health Organization; Food and Agriculture Organization of the United Nations. Protein and amino acid requirements in human nutrition. 2007.

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1. Access survey and provide feedback to receive Certificate of Attendance:

Aim your smartphone camera at this > QR code



OR

access the survey at: bit.ly/infant_growth OR

the survey will pop-up when you exit the live event

- 2. Find the event code at end of survey
- 3. Visit www.NutriciaLearningCenter.com Enter event code into your NLC Dashboard Certificate of Attendance added to your NLC profile!

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