

 Excellence is just the beginning.

Rush System for Health  
**Effects of Infant-Driven Feeding in the Neonatal Intensive Care Unit (NICU) for Premature Infants with Additional Morbidities**

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April 26<sup>th</sup>, 2024 | Alexa Rogers, Anna Marilla, Lisa LaGorio  
Co-contributors

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**Disclosure**

There are no relevant financial or non financial relationships to disclose.



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**1 Statement of the Problem**

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Swallowing is a complex process that can be disrupted by preterm birth

Physiologic stability, sensorimotor pathway functioning, and respiration

Premature birth is highly associated with comorbid conditions

High risk of neurologic, cardiac, respiratory, and/or gastrointestinal disorders

Comorbidities exacerbate dysphagia and burden

Infant-driven feeding has been emphasized in the literature to reduce these implications

Prolonged hospitalization/length of stay

Full oral feeding is a common final barrier preterm infants face in order to be discharged from the NICU

Delany and Anvedson (2008)  
Lau (2016)  
Mwantham and Jacobshe (2020)

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### Limitations of Current Literature

Dodrill et al. (2008)

Volume-driven feeding only

Did not specify which morbidity had the greatest impact

Jackson et al. (2015)

Only included late preterm infants (i.e., 32-36 weeks)

Only 9% of the infants had a comorbidity

Park et al. (2015)

Only included extremely preterm infants (i.e., <28 weeks)

Volume-driven feeding only

Morbidities were analyzed in isolation, despite a majority of infants having more than one

Rogers (2016)

Volume-driven feeding only

Manilla (2018)

Interactions between GA at birth, comorbidities, and feeding method on time to full oral feeding was not analyzed

Osman et al. (2021), Lane et al. (2021)

Exclusion of premature infants with various morbidities (e.g., IVH grade III or IV, cardiac anomalies, Glanomalies)

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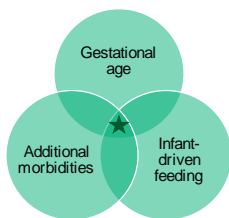
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### Interactions between gestational age, additional morbidities, and infant-driven feeding practices



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# 2 Normal Infant Swallowing

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**Phases of Swallowing**  
 Infants complete this process at a rate of one suck per second

1 Oral Phase	2 Pharyngeal Phase	3 Esophageal Phase
<ul style="list-style-type: none"> <li>• Expression</li> <li>• Suction</li> <li>• Containment</li> <li>• Propulsion to the pharynx</li> </ul>	<ul style="list-style-type: none"> <li>• Swallowing reflex</li> <li>• Pharyngeal constriction</li> <li>• Transportation to and through the upper esophageal sphincter (UES)</li> </ul>	<ul style="list-style-type: none"> <li>• Transportation through the esophagus and into the stomach</li> </ul>

**All phases are reflexive and mediated by a central pattern generator**

Delany and Arvidson (2008)  
 Daniels and Goss (2010)  
 Lau (2016)  
 Vanneman and Jäschke (2022)

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# 3 Swallowing Development in Utero

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### Oral-motor Movements

<b>13-15 weeks</b> <ul style="list-style-type: none"> <li>Suckling</li> <li>Swallowing</li> </ul>	➔	<b>Adaptive reflexes*</b> <ul style="list-style-type: none"> <li>Rooting</li> <li>Suckling</li> </ul>
<b>21 weeks</b> <ul style="list-style-type: none"> <li>Tongue thrust</li> </ul>		<b>Protective reflexes*</b> <ul style="list-style-type: none"> <li>Tongue protrusion</li> <li>Tongue lateralization</li> <li>Phasic bite</li> <li>Gag reflex</li> <li>Coughing</li> </ul>
<b>28 weeks</b> <ul style="list-style-type: none"> <li>Tongue cupping</li> </ul>		<p>*emerge during the third trimester of pregnancy</p>

Delany and Arvedson (2008)  
Dorfl and Giza (2015)  
Gottlieb et al. (2006)

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### Gastrointestinal and Respiratory Systems

Emerge around 4 weeks gestation as a result of cellular division

<b>Foregut</b> <ul style="list-style-type: none"> <li>Esophagus</li> <li>Somach</li> <li>Duodenum</li> </ul>	<b>Midgut</b> <ul style="list-style-type: none"> <li>Jejunum &amp; Ileum</li> <li>Mecum &amp; Ascending Colon</li> <li>Proximal Transverse Colon</li> </ul>	<b>Hindgut</b> <ul style="list-style-type: none"> <li>Distal Transverse Colon</li> <li>Descending &amp; Sigmoid Colon</li> <li>Rectum</li> </ul>
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Schematic from Achilli and Gweil (2008)

Image from Mullerberry and Smith (2015)

Lenfesty and Neu (2018)  
Wadsworth (2017)

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# 4 Effects of Prematurity on Swallowing

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### Classifications of Prematurity According to the World Health Organization (WHO)

Extremely Preterm	Very Preterm	Moderate to Late Preterm
<28;0 weeks' gestation	28;0-31;6 weeks' gestation	32;0-36;6 weeks' gestation

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### Dysphagia in Healthy Premature Infants

Prevalence: ~10.5% to ~26%

1 Signs of Oral Dysphagia	2 Signs of Pharyngeal Dysphagia	3 Signs of Esophageal Dysphagia
<ul style="list-style-type: none"> <li>Absent or reduced oral reflexes</li> <li>Impaired suction/expiration</li> <li>Irregular lingual movements</li> <li>Nasal regurgitation</li> <li>Delays in bolus formation</li> <li>Posterior loss</li> </ul>	<ul style="list-style-type: none"> <li>Delayed pharyngeal response</li> <li>Decreased pharyngeal pressure</li> <li>Incoordination between the pharyngeal reflex and UES distention</li> </ul>	<ul style="list-style-type: none"> <li>Impaired pressure gradients</li> <li>Esophageal immobility</li> <li>Abnormal peristalsis</li> </ul>

DeWitt (2011) | Jadhava (2016) | Hwang et al. (2021) | Rein et al. (2011) | Jadhava and Jadhava (2020) | Effects of Infant-Driven Feeding in the Neonatal Intensive Care Unit (NICU) for Premature Infants with Additional Morbidities | Rush System for Health | 3/15/2024 | 14

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## 5 Effects of Additional Comorbidities on Swallowing

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**Morbidity Overview**

Morbidity risk significantly increases as gestational age decreases

Allen et al. (2011)  
 Quillen (2011)  
 Engle (2011)

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**Morbidity Overview**

**Neurologic**

- Dysfunctional sensorimotor integration
- Dysfunctional coordination

**Cardiac**

- Uncoordinated SSB patterns
- Desaturation fatigue
- Oral stasis
- Anterior spillage
- Intestinal deficits, d/t lack of blood flow

**Respiratory**

- Oxygen desaturations
- Prolonged deglutition apnea
- Uncoordinated SSB patterns
- Atypical sucking
- Weaker suck

**Gastrointestinal**

- Feeding aversion
- Frequent regurgitation
- Discomfort following feedings
- Apnea, desaturations, bradycardia

Carter (2007) | Remani and Ambalavanan (2013)  
 Eichenwald (2018) | Scher and Pawlow (2008)

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**6 Feeding Factors that Impact Swallowing in Premature Infants**

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### Volume-Driven Feeding

#### Premise

Superior feeders – consume larger volumes  
Poor feeders – consume smaller volumes

#### Characteristics

Strategies to "empty the bottle"

- Increasing flow rate
- Prodding
- Pulling the infant's head back
- Unswaddling

#### Drawbacks

Unnecessary stress  
Risk of developing oral aversion  
Risk of long-term feeding problems

#### Signs of stress during feeding:

- Change in alertness
- Change in cardiorespiratory behaviors
- Change in postural control
- Fingers splayed
- Extension
- Loss of bolus
- Coughing/choking

#### Strict feeding schedule

Continue feeding despite mild signs of stress, incoordination, and disengagement

Lane et al. (2021)  
Shaker (2012)  
Shaker (2013)



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### Infant-Driven Feeding

#### Premise

Dynamic process between caregiver and infant during feeding  
Highly individualized process based on the infant's communication  
Quality > quantity

#### Characteristics

Evaluating readiness to feed  
Altering flow rate  
Side-lying  
Swaddling  
Pacing techniques  
Calm or re-alerting

#### Benefits

Supports neurodevelopmental maturation  
Safer feeding  
Achieve full oral feeding sooner  
Fewer adverse events  
Improved SSB patterns at discharge  
Shorter hospitalization

#### Signs of stability during feeding:

- Regular respirations
- Hands to midline
- Good postural control
- Organized, calm
- Alert

Lane et al. (2021)  
Lobbie (2018)  
Newland et al. (2013)  
Shaker (2012)  
Shaker (2013)  
Whitton (2016)



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### Research Question

- 1 What are the interactions between gestational age, comorbidities, and feeding practice (i.e., infant-driven versus volume-driven approaches) in the NICU on time to full oral feeding?



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# 7 Methods of the Current Study

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**Methods**

- 1 Retrospective chart review of 120 premature infants admitted to the Rush University Medical Center NICU born between Jan 1, 2018, and Jan 1, 2020, who received infant-driven feeding interventions
- 2 Eliminated infants from the data set based on predetermined inclusionary and exclusionary criteria
- 3 Analyzed the data according to each hypothesis (n = 88)
- 4 Combined the new data set with previous data from Manilla's (2018) study
- 5 Compared the results to Rogers' (2016) study to investigate feeding outcomes depending on feeding intervention (i.e., volume-driven versus infant-driven) (n = 268)

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**Data Collected from Chart Review**

Sex	Volume of 1 <sup>st</sup> oral feed	PMA at discharge
Race	Date of 1 <sup>st</sup> oral feed	Weight at discharge
Date of birth	PMA at exclusive oral feeding	Length of hospitalization (days)
Gestational age at birth	Weight at exclusive oral feeding	Participation in a Videofluoroscopic Swallowing Study
Postmenstrual age (PMA) at time of 1 <sup>st</sup> exposure to oral feeding	Volume at exclusive oral feeding	Diagnoses/morbidities
Weight at 1 <sup>st</sup> oral feed		

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### Morbidity Assessment Index of Newborns (MAIN)

Item No.	Item name	Distinction value <sup>a</sup>	$\chi^2$ statistic <sup>a</sup>	Sub-score <sup>b</sup>																
1	Poor suckling within 24 h	0.14	16.3	81	<table border="1"> <thead> <tr> <th>Score</th> <th>Conclusion</th> <th>Corresponding range of morbidity</th> </tr> </thead> <tbody> <tr> <td>&lt;500</td> <td>Score &lt; 6% (of score)</td> <td>Very mild to mild</td> </tr> <tr> <td>500-600</td> <td>Score &lt; 20% (of score)</td> <td>Mild</td> </tr> <tr> <td>600-2000</td> <td>Score &lt; 10% (of score)</td> <td>Mild to moderate</td> </tr> <tr> <td>&gt;2000</td> <td>Score &lt; 6% (of score)</td> <td>Moderate to severe</td> </tr> </tbody> </table>	Score	Conclusion	Corresponding range of morbidity	<500	Score < 6% (of score)	Very mild to mild	500-600	Score < 20% (of score)	Mild	600-2000	Score < 10% (of score)	Mild to moderate	>2000	Score < 6% (of score)	Moderate to severe
Score	Conclusion	Corresponding range of morbidity																		
<500	Score < 6% (of score)	Very mild to mild																		
500-600	Score < 20% (of score)	Mild																		
600-2000	Score < 10% (of score)	Mild to moderate																		
>2000	Score < 6% (of score)	Moderate to severe																		
1	Apnea with gasp/straw/murmur <sup>†</sup>	0.39	26.6																	
1	Apnea without 17% $\text{SpO}_2$ $\text{cmO}_2$	0.33	31.6																	
2	Poor suckling at 24 h to 7 days	1.46	5.3	98																
3	Severe bilirubin > 20 mg/dL (or need for phototherapy)	0.61	21.3	103																
4	Assist ventilation within 24 h <sup>†</sup>	0.19	63.1																	
4	Apnea and need for oxygen <sup>†</sup>	1.49	6.7	115																
5	Assist ventilation beyond 24 h	0.76	26.2	117																
6	Poor suckling beyond 7 days	1.15	12.9	119																
7	Heart rate $> 150$ beats/min	0.80	2.9	120																
8	Apical score $< 7$ at 1 min	0.51	16.1	125																
9	Resuscitation at birth with intubation	0.99	8.0	127																
10	Hypotonia at 1-20 h	1.12	8.8	129																
11	Mechanical ventilation within 24 h	1.27	2.8	130																
12	Respiratory rate $< 30$ /min or $> 60$ /min at 3-24 h	0.67	16.8	131																
13	Mechanical ventilation at 24 h to 7 days	1.47	4.5	135																
14	Respiratory rate $< 30$ /min or $> 60$ /min at day 7-32 <sup>†</sup>	0.19	17.2	136																
14	Low systemic blood pressure <sup>†</sup>	1.00	7.9	136																
15	Low mean Hg for 32-42 wk:																			
15	$< 40$ mm Hg for 32-42 wk	0.66	11.8	136																
16	Pharyngeal feeding	1.08	0.6	137																
17	Taxonom or single seizure	0.02	2.2	137																
18	Apnea corrected by resuscitation	1.19	3.4	140																
18	Respiratory rate $> 60$ /min at 1-24 h	0.50	6.5	140																
20	Low urine output ( $< 2$ mL/kg per hour <sup>†</sup> )	1.47	1.5	141																
21	Altered color: Central cyanosis or dusky	0.81	17.8	145																
22	Thrombocytopenia or bleeding skin, lungs, or gastrointestinal tract	0.94	2.3	147																
23	Cord blood pH $< 7.1$	0.54	9.6	151																

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### Statistical Analysis by Hypothesis

#### Hypothesis 1

Effect of age on time to full oral feeding

- Age as a continuous variable: Pearson correlation
- Age as a categorical variable: point-biserial correlation and ANOVA

#### Hypothesis 2

Effect of additional morbidities on time to full oral feeding

- One-way independent ANOVA with the binary presence or absence of morbidity as the independent variable

#### Hypothesis 3

Effect of specific morbidities on time to full oral feeding

- ANOVA was used to compare results among the four morbidity groups
- Pearson correlations were used to compare results among MAIN severity score

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### Statistical Analysis by Hypothesis

#### Hypothesis 4

Effect of morbidity on time to full oral feeding in extremely premature infants

- One-way independent ANOVA with the binary presence or absence of morbidity as the independent variable

#### Hypothesis 5

Effect of feeding protocol on time to full oral feeding

- Independent t-tests to compare the mean time to full oral feeding between the two feeding method groups

#### Hypothesis 6

Interaction between gestational age, morbidity, and feeding method

- ANCOVA

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# 8 Results

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**Hypothesis 1 (n = 88)**  
**Effect of Gestational Age at Birth on Time to Full Oral Feeding**

**Pearson correlation coefficient**  
**Strong negative correlation** between gestational age at birth and time from birth to full oral feeding [ $r = -0.856$ ,  $p < 0.0001$ ].  
**Moderately negative correlation** between gestational age at birth and time from first to full oral feeding [ $r = -0.567$ ,  $p < 0.0001$ ].

**Univariate ANOVA**  
**Statistically significant** difference between gestation age group at birth and time from:

1. Birth to first oral feeding [F(2, 85) = 200.469,  $p < 0.0001$ ]
2. First to full oral feeding [F(2, 85) = 20.123,  $p < 0.0001$ ]
3. Birth to full oral feeding [F(2, 85) = 121.30,  $p < 0.0001$ ]

**Bonferroni's multiple comparisons post-hoc analysis**  
 Mean value of time from birth to first oral feeding was **significantly different** ( $p < 0.0001$ ) between the following age groups: <28.0 weeks' gestation and <28.0 weeks' gestation and 28.0-31.6 weeks' gestation and 28.0-31.6 weeks' gestation and 32.0-36.6 weeks' gestation and 32.0-36.6 weeks' gestation.

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**Hypothesis 2 (n = 88)**  
**Effect of Morbidities on Time to Full Oral Feeding**

**One-way independent ANOVA**  
**Statistically significant** between group difference among premature infants diagnosed with at least one additional morbidity (n = 76) compared to those without (n = 12) on time from:

1. Birth to full oral feeding [F(1, 74) = 13.877,  $p < 0.0001$ ]
2. First to full oral feeding [F(1, 86) = 19.634,  $p < 0.0001$ ]

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### Hypothesis 3 (n = 88)

#### Effect of Specific Diagnoses on Time to Full Oral Feeding

##### ANOVA

Statistically significant between group difference among specific diagnoses on time from:

1. Birth to full oral feeding [F(8, 79) = 16.693, p < 0.0001]
2. First to full oral feeding [F(8, 79) = 8.120, p < 0.0001]

Diagnosis	n	M	SD	Diagnosis	n	M	SD
Time from Birth to Full Oral Feeding (days)				Time from First to Full Oral Feeding (days)			
Respiratory	45	26.67	22.798	Respiratory	45	15.0	11.695
Cardiac	2	20.0	5.657	Cardiac	2	19.0	5.657
GI	1	32.0	—	GI	1	1.0	—
Neurologic and Respiratory	3	36.67	20.744	Neurologic and Respiratory	3	21.67	9.815
Respiratory and Cardiac	3	53.33	12.662	Respiratory and Cardiac	3	22.67	24.947
Neurologic, Respiratory, and Cardiac	2	119.50	38.891	Neurologic, Respiratory, and Cardiac	2	54.50	3.536
Neurologic, Respiratory, and GI	1	93.0	—	Neurologic, Respiratory, and GI	1	44.0	—
Neurologic, Respiratory, Cardiac, and GI	2	95.50	7.778	Neurologic, Respiratory, Cardiac, and GI	2	28.50	6.364

Note. Each diagnostic category represents the number of infants with a single morbidity (e.g., respiratory, cardiac, etc.) and infants with multiple morbidities (e.g., neurologic + respiratory + cardiac).



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### Hypothesis 3 (n = 88)

#### Effect of Specific Diagnoses on Time to Full Oral Feeding

##### Pearson correlation

Moderately positive correlation between MAIN severity and time from:

1. Birth to full oral feeding [r = 0.546, p < 0.0001]
2. First to full oral feeding [r = 0.518, p < 0.0001]



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### Hypothesis 4 (n = 88)

#### Interaction Between Gestational Age and Morbidity in the Youngest Premature Infants (i.e., <28.0 weeks')

##### One-way independent ANOVA

Planned; however, analysis could not be performed because all infants born <28.0 weeks' gestation had at least one additional morbidity.

##### ANOVA

No statistically significant difference between the morbidities of the youngest premature infants on time from:

1. Birth to full oral feeding [F(4, 6) = 1.553, p = 0.300]
2. First to full oral feeding [F(4, 6) = 1.743, p = 0.258]



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### Hypothesis 5 (n = 268)

#### Effect of Infant-Driven Feeding Protocol on Time to Full Oral Feeding

##### Independent t-test

No statistically significant difference between feeding methods on time from:

- 1. Birth to full oral feeding (t(267) = -0.454, p = 0.650)
- 2. First to full oral feeding (t(267) = -1.382, p = 0.168)

##### ANOVA

No statistically significant interactions between WHO (2023) GA group and feeding method on time from:

- 1. Birth to full oral feeding [F(2, 262) = 1.187, p = 0.307]
- 2. First to full oral feeding [F(2, 262) = 0.560, p = 0.572]

##### Main effect of age for time from:

- 1. Birth to full oral feeding [F(2, 2) = 269.555, p = 0.004]
- 2. First to full oral feeding [F(2, 2) = 64.430, p = 0.015]

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### Hypothesis 5 (n = 268)

#### Effect of Infant-Driven Feeding Protocol on Time to First Full Oral Feeding

Infants were reclassified into four groups based on gestational age at birth



##### Univariate ANOVA

No statistically significant difference between the reclassified gestational age groups on time from:

- 1. Birth to full oral feeding [F(3, 260) = 1.701, p = 0.167]
- 2. First to full oral feeding [F(3, 260) = 1.256, p = 0.290]

##### Main effect of age for time from:

- 1. Birth to full oral feeding [F(4, 4.139) = 109.815, p < 0.0001]
- 2. First to full oral feeding [F(4, 4.585) = 16.026, p = 0.006]

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### Hypothesis 6 (n = 268)

#### Interaction Between Gestational Age at Birth, Comorbidities, and Feeding Method

##### ANOVA

No statistically significant interactions between WHO (2023) GA group, morbidity, and feeding method on time from:

- 1. Birth to full oral feeding [F(1, 258) = 1.033, p = 0.310]
- 2. First to full oral feeding [F(1, 258) = 0.548, p = 0.460]

Main effect of morbidity on time from first to full oral feeding [F(1, 1) = 366.933, p = 0.033]

##### ANOVA

No statistically significant interactions between reclassified GA group, morbidity, and feeding method on time from:

- 1. Birth to full oral feeding [F(1, 256) = 1.133, p = 0.288]
- 2. First to full oral feeding [F(1, 256) = 0.553, p = 0.458]

Infants 23:0-24:6 weeks' gestation with at least one additional morbidity in the volume-driven feeding group:

- 12.29 more days from birth to full oral feeding
- 18.43 more days from first to full oral feeding

RUSH

Effects of Infant-Driven Feeding in the Neonatal Intensive Care Unit (NICU) for Premature Infants with Additional Morbidity | Rush System for Health | 3/15/2024 36

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### Hypothesis 6 (n = 268)

#### Interaction Between Gestational Age at Birth, Comorbidity Severity, and Feeding Method

##### ANOVA

No statistically significant interactions between WHO (2023) GA group, morbidity severity, and feeding method on time from:

- 1. Birth to full oral feeding [F(4,249) = 0.531, p = 0.713]
- 2. First to full oral feeding [F(4,249) = 0.899, p = 0.465]

**Main effect of age and MAIN severity for time from birth to full oral feeding**

- 1. Age [F(2, 1,850) = 0.691, p = 0.005]
- 2. MAIN severity [F(3, 12,185) = 274.181, p = 0.007]

##### ANOVA

No statistically significant interactions between reclassified GA group, morbidity severity, and feeding method on time from:

- 1. Birth to full oral feeding [F(4, 245) = 0.531, p = 0.726]
- 2. First to full oral feeding [F(4, 245) = 0.386, p = 0.819]

**Main effect of age for time from birth to full oral feeding**[F(3,5,018) = 65.014, p < 0.0001]

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# 8 Discussion

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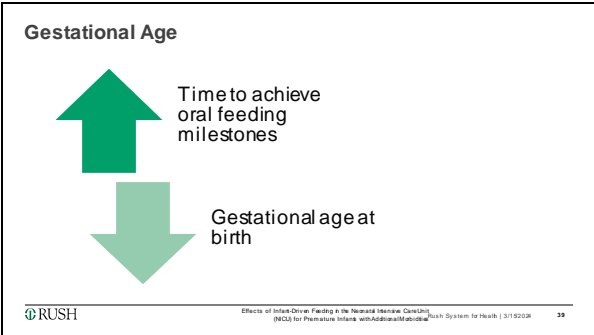
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### Comorbid Medical Diagnoses

Presence of at least one morbidity

Time to achieve oral feeding milestones

RUSH Effects of Infant-Driven Feeding in the Neonatal Intensive Care Unit (NICU) for Premature Infants with Additional Morbidities Rush System for Health | 3/1/2023 40

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### Specific Morbidity Diagnoses

Morbidity severity

Time to achieve oral feeding milestones

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### Specific Morbidity Diagnoses

Neurologic

Respiratory

Cardiac

Time to achieve oral feeding milestones

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### Extreme Prematurity and Comorbidities

All infants <28;0 weeks' gestation had at least one concomitant morbidity  
Highlights how prevalent comorbidities are among extremely preterm infants

No difference on time to full oral feeding between morbidities  
Highlights the inevitability of delayed attainment of oral feeding milestones in this GA group regardless of morbidity diagnoses

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### Feeding Protocol

Gestational Age Categorized According to the WHO (2023)  
No statistically significant interaction may have been due to the wide range of ages lumped into the <28-week group and the extreme values → regression to the mean

WHO (2023) GA categories were created at a time when infants born <28 weeks did not survive

#### Survival Rates



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### Feeding Protocol

Proposing a new premature infant GA classification schema:



Youngest infants reached full oral feeding almost 20 days sooner when infant-driven feeding was implemented

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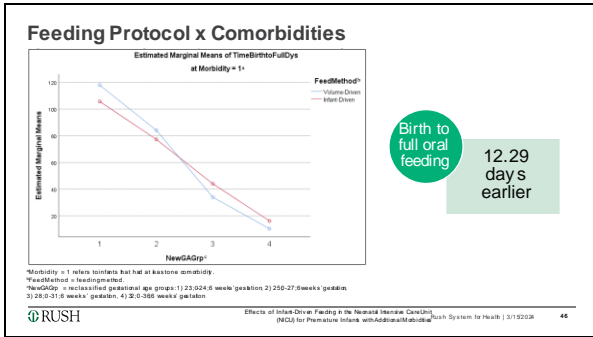
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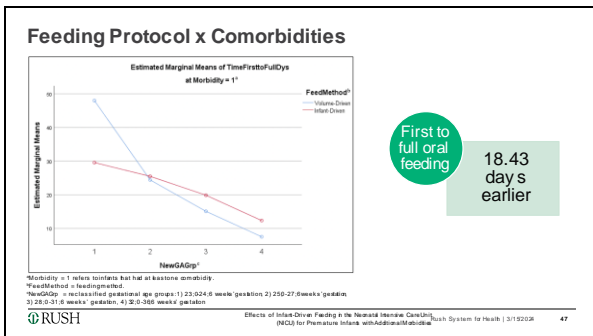
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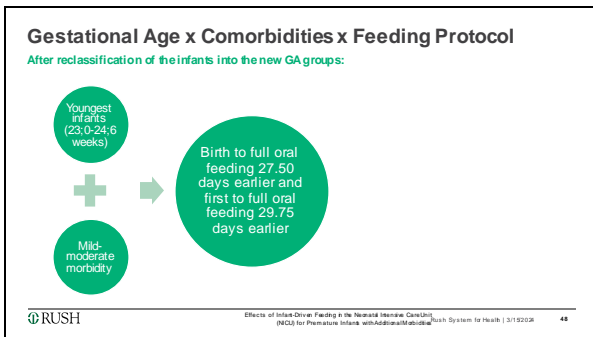
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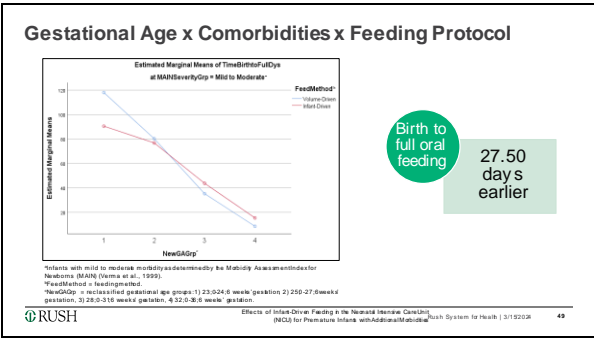
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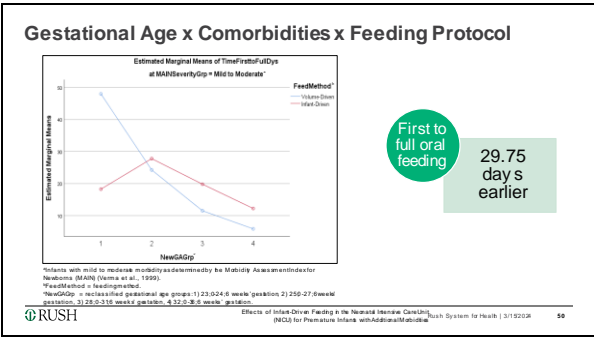
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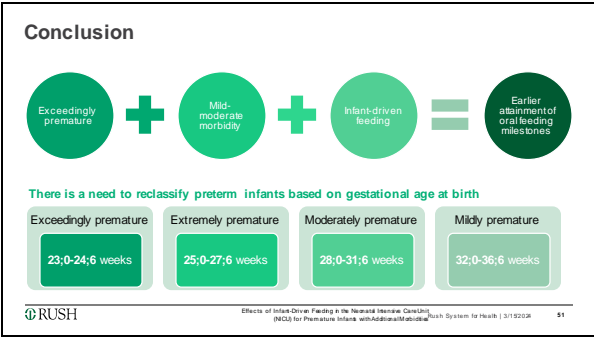
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Thank you.

