#### **Preterm Dietary Supplements**



#### **Dr Umesh Vaidya**

#### IAP Neocon, Mumbai 2015



**Preterm VLBW Nutrition : Ideal practice** 

- Minimal enteral feeds (10 ml / kg / day)
- Human breast milk
- Feed advancement @ 20 ml / kg / day
- Human milk fortification 100 ml / kg /day
- Parenteral nutrition (ELBW < 1000 g )</th>Aminoacids 1 1.5 g / kg / dayDay 1Lipids1 2 g / kg / dayDay 1-3

Ziegler EE – J Mat – Fed – Neonatal Med, Mar 2009

## Case, Baby of MI 28 wks, BW 1280 gms

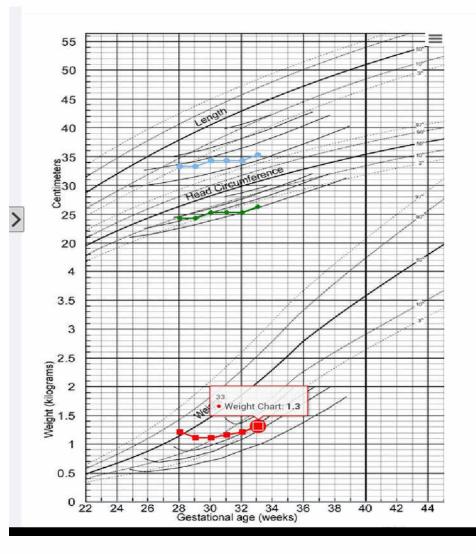
#### DAYS NUTRITION

- DAY 1 TPN (1.5 g amino acids; 1.5 g lipids)
- DAY 2 Minimal Enteral Feeding
- DAY 2-8 Grading up of feeds HMF started
- DAY 9-30 Fortified milk- full feeds
- DAY 33-34 Sodium supplementation
- DAY 30-35 Transition to oral feeds- discharge

Weight gain: Day 9 - 30 - 10 gms / day



Initial birth weight: 1280 gms Birth weight regained on day 34 Birth centile : 30 th (Fenton) Discharge centile : 3 rd

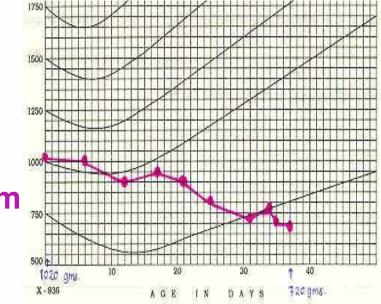


## **Extrauterine growth retardation – EUGR**



#### Born at 28 wks, BW 1020 gm

#### **RDS / Pulmonary morbidity**



#### Wt at 1 month 720 gm

## **EUGR - A serious problem in preterms**

- Preterms 23 34 weeks, 24371 Preterms
- Growth < 10<sup>th</sup> percentile at 40 weeks

WT	28 %
LNTH	34 %
HC	16 %



#### Possible neurologic and sensory handicaps

Clark RH, Thomas P, Peabody J, Pediatrics 2003

**CONSEQUENCES OF INADEQUATE EARLY NUTRITION** 

#### **Post-natal Growth Restriction - a global concern**

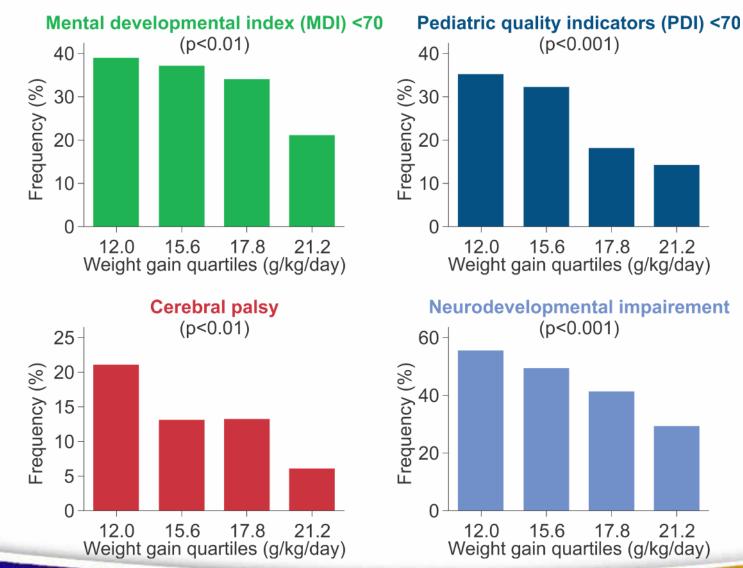
#### **Vulnerable periods**

# Nutritional insults – impaired somatic growth Impaired neuro-cognitive development

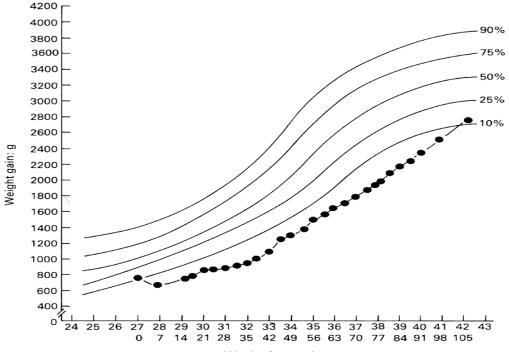
Ehrenkranz et al Pediatrics 2006



# In-hospital Growth Velocity and Neurodevelopmental Outcomes



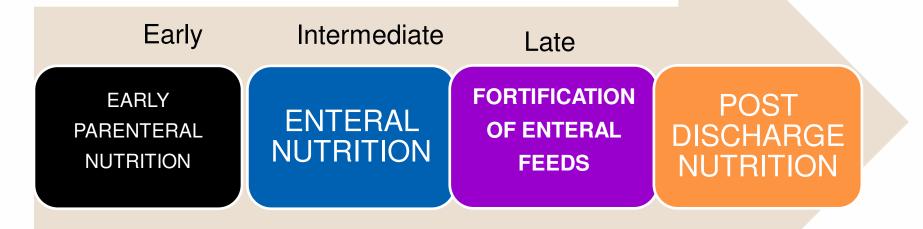
# **DEFINING GROWTH TARGETS**



Week of gestation

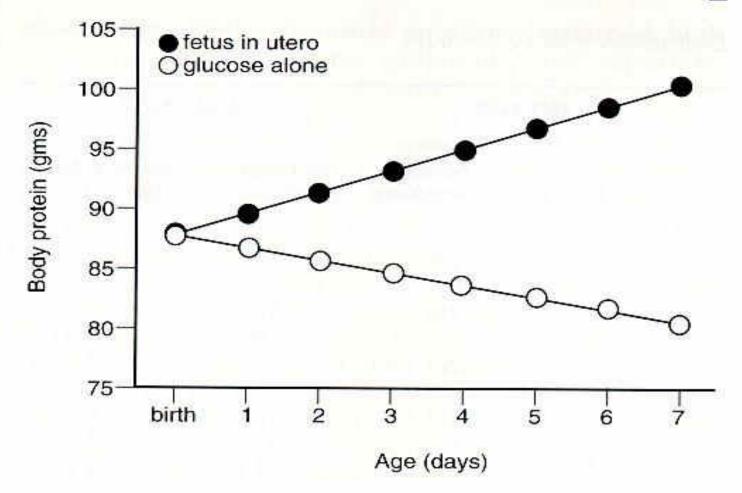
Weight gain 10-15 gm/kg/day Length gain 0.75-1 cm/wk HC gain 0.75 cm/wk

# **NUTRITIONAL CARE PLAN**



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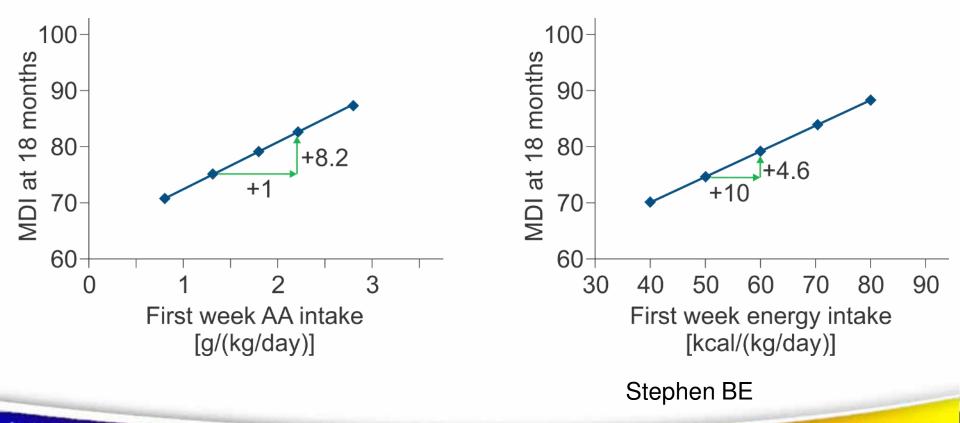
# GLUCOSE ALONE AS NUTRITION (26 weeks, 1000g)



#### Denne SC, J Clin Invest 1996

# First Week Protein and Energy Intake and Neurodevelopmental Outcomes

- Retrospective study of 124 ELBW infants at 18 months CA
- AA intake  $1^{st}$  week:  $1.8 \pm 0.4$  g/kg/day
- Energy intake 1<sup>st</sup> week: 60±8 kcal/kg/day



#### EARLY AGGRESSIVE PN

#### **Current recommendation**

Day 1 Min AA 1.5 g/kg (3 g / kg preferred) Started within 1 hr of birth Increase to 4 g / kg within few days LIPIDS 1 – 2 g / kg Day 1 Several studies (AA 3.5 g / kg , Lipids 3 g / kg) No increase in BUN, lipids, acidosis

**Uhing MR Clin Perinatol 2009** 

# MINIMAL ENTERAL FEEDING

- EBM 8 -12 ml / kg , 3-6 hrly, starting 1-3 hrs after birth
- Preferably with EBM / Donor Human milk

# **GRADING UP FEEDS**

Increments of at least 20 ml / kg / day (Full feeds Day 7 ) Caution in babies < 750 g and SGA infants as data limited

(Cochrane Systematic Review 2011)



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### Human Milk Banking A National mission

#### **Network of Human Milk Banking**



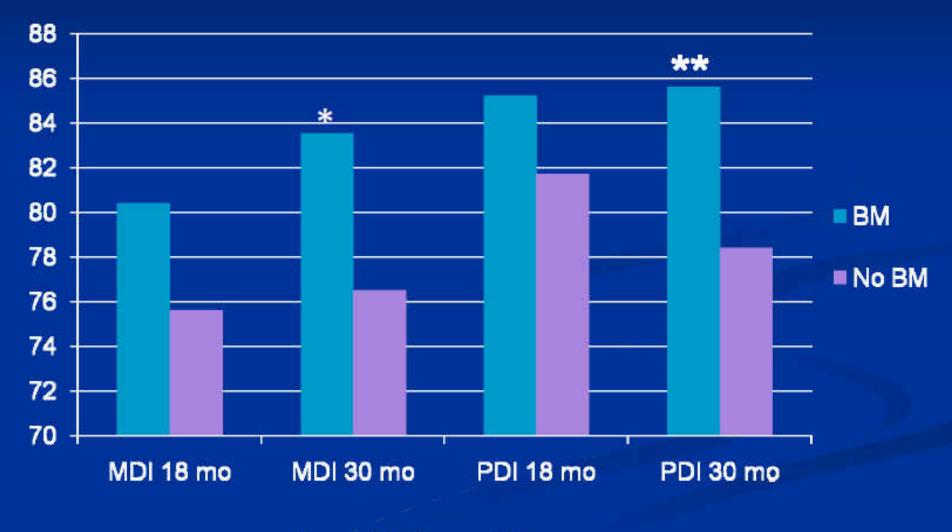


# Effect of human milk feeding on morbidity & hospital stay

	Human Milk > 50 ml/kg/d	Human milk and formula	Preterm formula
No.infants	62	63	42
Human milk intake	96 + 23	20 <u>+</u> 15	0
NEC n(%)	1(2%)	16(25%)	6(13%)
Late onset sepsis (LOS) n (%) **	19(31%)	29(45%)	22(48%)
LOS & NEC n(%)	19(31%)	35(56%)	25(54%)
Hosp stay days ***	73 <u>+</u> 19	87 <u>+</u> 43	88 <u>+</u> 47

Schanler , et al . Pediatrics 1999; 103: 1150 – 57 B. Wt . 1000 g GA < 30 wks, \*p < 0.01, \*\* p < 0.07, \*\*\*p < 0.05

# Mean MDI and PDI scores for ELBW infants at 18 & 30 months age



\* p = .03, \*\* p = .008

#### Enteral nutrient supply for preterm Infants : Commentary ESPGHAN Committee on Nutrition

Fluid: 110-200 ml/kg/d
Energy intake 110-135 kcal/kg/d
Protein intake 3.5 –4.5 g/kg/d (much higher)
Fat intake 4.4 – 6.6 g/kg/d
CHO intake 11.5 - 16 g/kg/d

J Pediatric Gastroenterology & Nutrition 50 : 1-9 2010

### **PRETERM : ADEQUACY OF BREAST MILK ?**

NUTRIENT	PRETERM MILK (PER 100ML)	EBM 200 ml / kg	Requirement	DEFICIT
Protein (g)	1.1 – 1.5	2.2 - 3.0	3.5 – 4	1.3 – 1
Calcium (mg)	20	40	120 – 140	80 – 100
Phosphorus (mg)	15	30	60 – 90	30 – 60
Zinc (mcg)	295	590	800 – 1200	500 – 700
Vitamin A (IU)	250	500	1400 – 2500	1000 – 2000
Vitamin D(IU)	2.2	4.4	800 – 1000	800 - 1000

## **Types of Fortification**

Monocomponent Fortification

Carbohydrates Proteins Fats Calcium PO4 Iron Vitamins Multi-component Fortification

Combination of 2 or more nutrients

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# **MONO- COMPONENT FORTIFICATION**

Carbohydrates

no evidence available

Fats (MCT oil)

Little evidence (1 RCT) showed no effect

#### **Proteins**

4 studies 90 babies, Better short term growth Long term effects ?

# MULTI-COMPONENT FORTIFICATION : The Evidence

13 studies, > 600 patients

Short term outcome



Increased wt gain 3.6 g /kg / d

Increased LNTH increment 0.12 cm / week

Increased HC 0.12 cm / week

Long term outcome

No difference in WT./LNTH/HC at 12 & 18 mths No effects on neurodevelopmental outcome No effects on bone mineral content

Cochrane database 2009

#### **Problems with Standard Fortification**

58 % VLBW infants receiving fortified EBM have extra – uterine growth retardation at discharge

Henrikson C et al 2009

#### **Standard Fortification – Protein deficits**

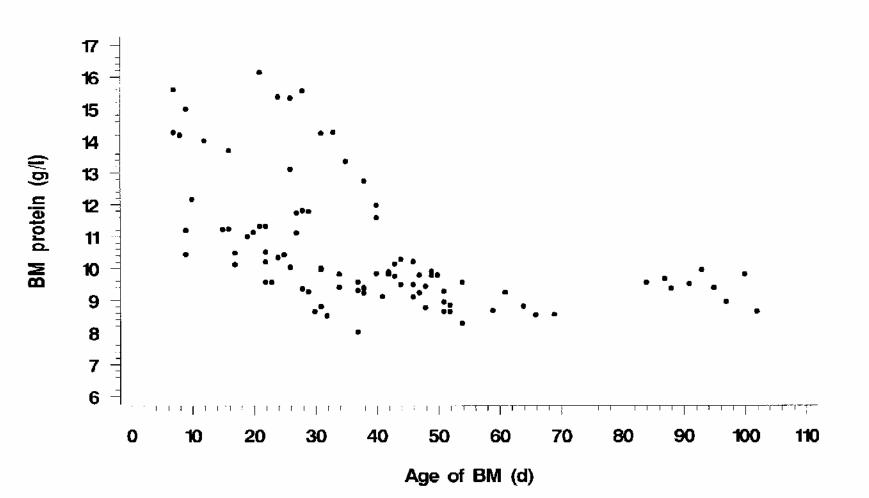


# Proteins is the issue !!

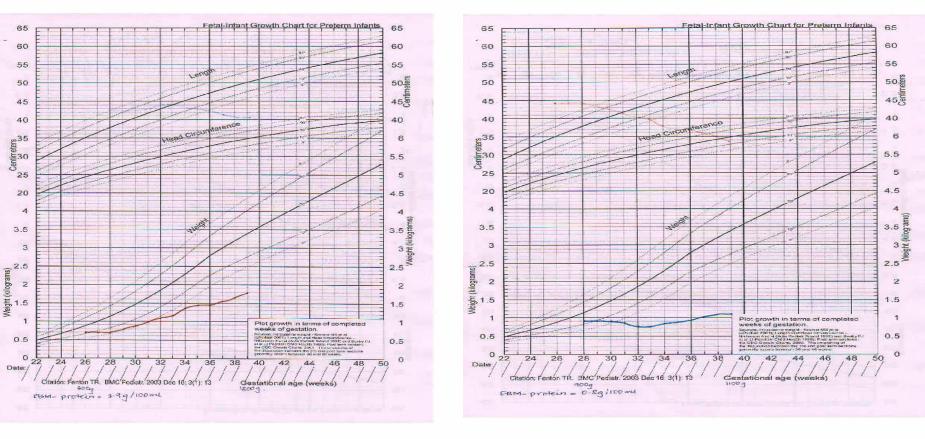
#### LOW PROTEIN INTAKE IS THE PRIMARY LIMITING FACTOR FOR GROWTH FAILURE

- Assumed higher protein content of human milk
- Low protein content of fortifiers
- •Transition from high protein PN solutions to lower protein enteral feeds

#### Variable Protein Content of EBM



### **Variable Protein content of EBM**



#### Baby G EBM protein 1.9 g / 100 ml

Baby S EBM protein 0.8 g / 100 ml





- Human milk analysis is essential to the health and growth of preterm babies.
- Miris Human Milk Analyser helps clinicians manage preterm nutrition quickly and simply



# Protein content of Fortifiers (per gm)

Lactodex HMF (Raptakos)	0.1
Hijam (Endocura)	0.25
Enfamil HMF (MJ)	0.27
Similac HMF (Abbott)	0.25
Aptamil BMF (Milupa)	0.2
FM 85 (Nestle)	0.2

# **Novel Methods of Fortification**

### Focus on more protein

#### **Standard Fortification**

Pre-determined amount of fortifier added to EBM

#### **Targeted Fortification**

#### **Adjustable Fortification**

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Analyse EBM for proteins and a add desired amount of fortifier to reach 3.5 g / kg /day

Assess protein intake by evaluation of infant's metabolic response by checking BUN (Increase dose of fortifier if BUN < 9 mg %, maintain between 9 - 14)

Recommendation & Guidelines for perinatal practice Arslanoglou S, Moro GE, Ziegler E, J. Perinatal Med 2010

# **Adjustable Fortification : Turkey study**

58 preterm VLBWs (<32 weeks,<1500 g)

Adjustable Fortification based on Blood urea levels

BUN< 9 mg %</th>Increase Protein by 0.55 g14-20 mg %Decease Protein by 0.55 g> 20 mg %Stop Protein supplement

Study group 4g/kg/day versus Control group 2.78 g/kg/day

**Result : Significant improvements in WT,LNTH,HC** 

Alan S et al, Early Human Dev 2013

Is there an alternative Protein supplement ? Skimmed Milk Powder (SMP) Each 100 g contains

Energy	360 kcal
Proteins	34.5 g
Fats	1 g
CHO	52 g
Calcium	1250 mg
PO4	970 mg
Sodium	500mg

# SMP FORTIFICATION (2 gm / 100 ml @ 170 ml / kg / day)

Energy kcal	126	(110 – 135)
Proteins (g)	3	(3.5 – 4)
CHO (g)	13.7	(11.6 – 13.2)
Fats (g)	7.2	(4.8 – 8.8)
Calcium (mg)	102	(120 – 140)
Phosphorus (mg)	58.5	(60 – 90)
Sodium (mg)	25.5	(69 – 115)



# WHAT WE DO ???

# NUTRITIONIST-GUIDED ENTERAL FEEDING

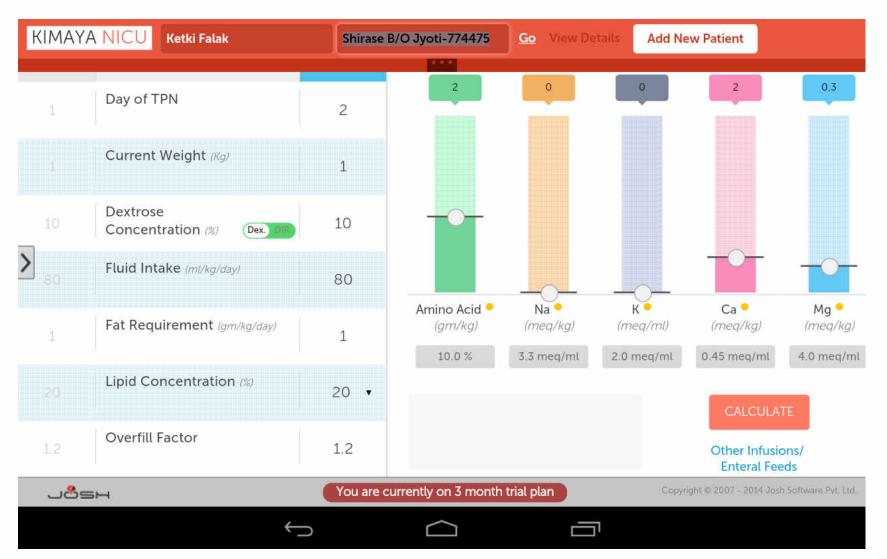
- Early TPN for all < 1500 g
- Minimal Enteral Nutrition with Colostrum
- Use of Expressed Breast milk /Donor milk
- Rapid grading up in stable prems (20-30 ml/kg/day)
- Fortification at 100 ml/kg/day with HMF/SMP
- Optimal calories, proteins, Vitamin D, Calcium PO4, Iron

# **Bedside Nutrition Management Tool**

#### **Kimaya NICU Nutrition Software**

- Bedside nutrition plan
- Individualized calculations for Enteral & Parenteral Nutrition
- Linkage between EN & PN
- Helps in optimizing calorie & protein intake
- Growth chart interface

# **TPN Calculation**



### **Enteral Calculation**

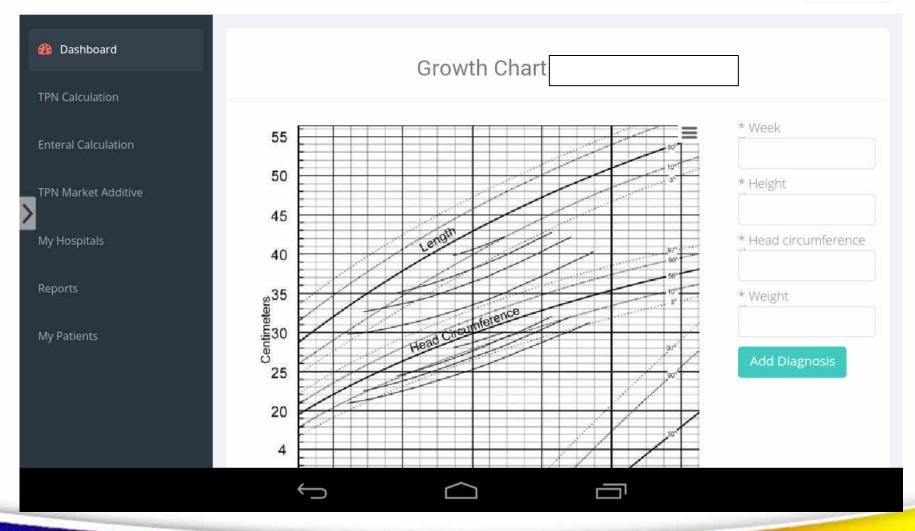
Intake Day of life     Additives Intake     HMF HMF     Additives Intake (gm/day)     Additives Intake (gm/day)     Additives Intake (gm/day)     SMP SMP     CALCULATE     Day of life     Day of life     Total Proteins     Total calories     Intake (gm/day)     SMP     SMP     CALCULATE     Day of life     Day of life     Intake     Intake		Day Of Life Intake <i>(ml)</i> Type of	9 Current ( <i>Kg</i> ) 17 Intake Fr ( <i>per hour</i> )	equency 2 h V	400 sanger 200 0 4	6 7	8 9	
Additives MCT intake (gm/day)   MCT Oil MCT intake (gm/day)   Additives Intake (gm/day)   SMP SMP   SMP SMP intake gm	1	Intake	Additives			Proteins 📕 Tot	al calories	
SMP       SMP intake gm         CALCULATE       SMP intake gm    Total Fluid 204 ml/day 170 ml/kg/day Total Calories 165.21 kcal 137.68 kcal/kg Total Proteins 3.66 gm 3.05 gm/kg		🔲 MCT Oil	MCT Oil 🔹	MCT intake gm			Report	
		SMP			Total Calories	165.21 kcal	137.68 kcal/kg	
Corones.			CALCULATE		Total Proteins Calories	3.66 gm	3.05 gm/kg	

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# **Growth chart**

#### ∃ KIMAYA NICU

Ketki Falak 🔻



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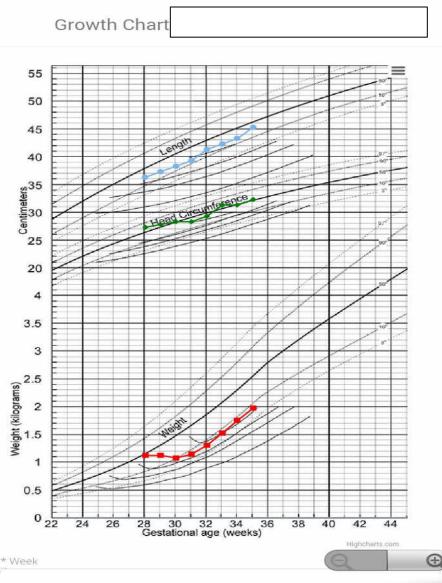
# **Daily Nutrition Plan**

Date		Fluid (ml/kg/day)	Proteins (gm/kg/day)	Calories (kcal/kg/day)	Total Fluid (ml/kg/day)	Total Proteins (gm/kg/day)	Total Calories (kcal/kg/day)
2015-10-10	Enteral TPN	175	2.86	128.50	175	2.86	128.5
2015-10-06	Enteral TPN	177	2.89	129.85	177	2.89	129.85
2015-10-02	Enteral TPN	177	2.89	129.85	177	2.89	129.85
2015-09-30	Enteral TPN	175	2.90	129.02	175	2.9	129.02
2015-09-29	Enteral TPN	175	2.94	129.57	175	2.94	129.57
2015-09-23	Enteral TPN	163	2.90	122.52	163	2.9	122.52
2015-09-13	Enteral TPN	175	3.17	132.25	175	3.17	132.25
2015-09-04	Enteral TPN	173	3.10	130.18	173	3.1	130.18
2015-09-02	Enteral TPN	179	3.37	137.05	179	3.37	137.05
2015-08-23	Enteral TPN	140	3.46	117.06	140	3.46	117.06
2015-08-18	Enteral	174	3.49	135.32	174	3.49	135.32

### **Tailor-made Nutrition**

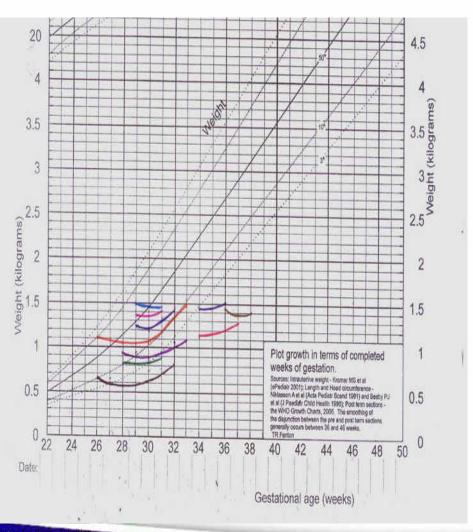


 Growth and metabolic monitoring (Proteins, Hb, Ferritin, Ca PO4,alk PO4)

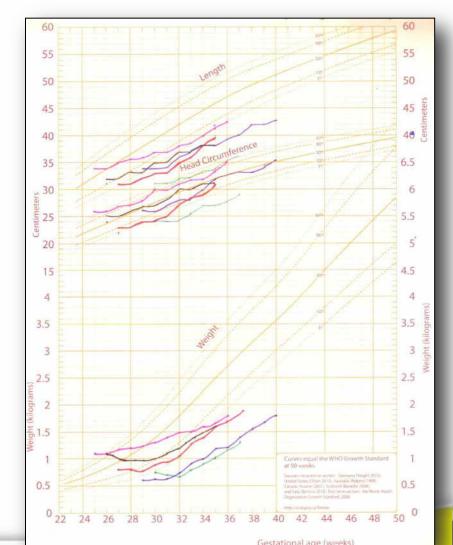


# **Growth of KEM babies**

#### 2010



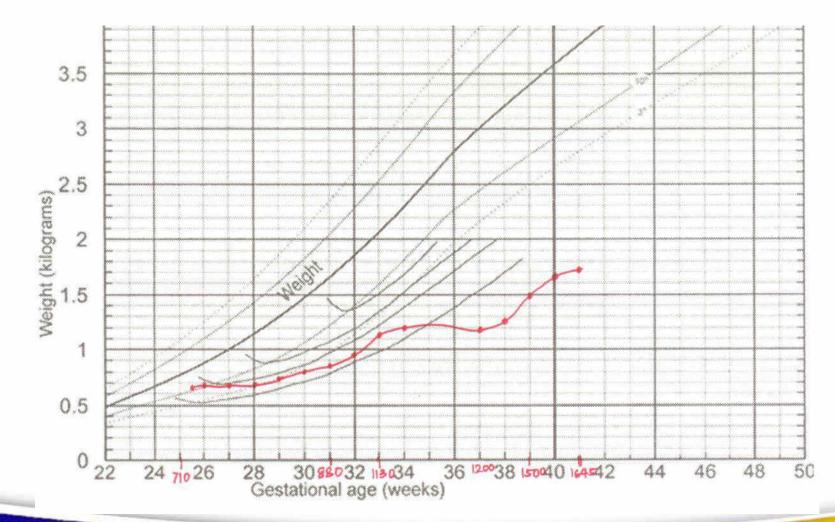




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#### **POST-DISCHARGE NUTRITION**

#### Baby K, GA 25 weeks Triplet, Weight 710 gms



### What could be the reasons for poor growth?

Inadequate feeding due to poor suck and swallow Inadequate breast milk

- Cold stress
- Exaggerated anaemia of prematurity

Inadequate calorie / protein intake due to discontinuation of fortification

Sepsis/sickness in the baby

Electrolyte imbalance (Late onset Hyponatraemia)

# 3. Human milk fortification after Discharge

#### **Powdered Human Milk Fortifier (HMF)**

**Proper training of mother important** 

Estimate approx milk intake and calculate dose of HMF

Express milk to mix HMF and then can breastfeed

2 gm in 50 ml EBM (1 gm HMF in 25 ml EBM)

Fortification with MCT oil – not recommended due to lack of scientific evidence (Cochrane Review 2009)

Special post – discharge formulae (72–80 kcal/dl)

4. Post-Discharge Nutritional Supplementation

### Stable, full feeds

Multivitamin with Zinc (? Continue till 1 year) Calcium : Phosphorus (continue till term / 3 kg) Vitamin D 800 IU per day (ESPGHAN 2010) (continue till 1 year)

#### 4 -6 weeks

Start Iron supplementation Continue till one year



# Nutritional supplementation for preterms

• Vitamin A - 1500 IU per day

(Dose of multivitamin drops to be titrated accordingly)

- **Zinc** 2.5 mg 3 mg / d
- Vitamin D: 800-1000 IU /day (ESPGHAN 2010)
- Calcium 150 mg /kg
- Phosphorus 75 mg /kg

### **MULTIVITAMIN DROPS (per 1 ml)**

	Brand A	Brand B	Brand C
A	2500 IU	1000 IU	2750 IU
$D_3$	200 IU	400 IU	250 IU
E	2.5 mg	1.5 mg	5 mg
С	40 mg	40 mg	40 mg
B <sub>1</sub>	1 mg	2 mg	1 mg
B <sub>6</sub>		1 mg	1 mg
K			
DOSE:	0.5 ml	1 ml	0.5 ml

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# **Iron supplementation**

- < 1000g
- 1000-1500g
- 1500-1800g
- >1800g

- 4 mg / kg / d
- 3-4 mg / kg / d
- 2-3 mg / kg / d
- 2 mg / kg / d

- Start at 4-6 weeks, Continue till 12-15 months
- For babies <1500 gm early supplementation (2 weeks) may be considered

# **IRON DROPS (per ml)**

J
g

mg Fe / drop

1

0.3

# 6. When to start complementary foods?

• Complementary food should be started about at corrected age of 4-6 months for preterm infants

Deborah L, Unger S. Seminars in fetal and neonatal medicine, (2013)

- Generally, most babies would be around 5 kg, would have some neck support and tongue-thrust reflex would have diminished
- Complementary foods are chosen as per recommendation for term infants

# SUMMARY

- Postnatal growth retardation common in preterm LBWs
- Optimal growth targets unclear, but prevention of EUGR is mandatory
- Current trends favour aggressive nutrition strategies to improve neurologic outcome
- Early PN, use of human milk, fortification of enteral feeds to achieve adequate calories and proteins



# SUMMARY (Contd)

Human Milk Banking now a national mission

Donor Human milk has both short and long-term benefits

Protein deficits are contributing to growth failure

Recent guidelines suggest higher protein intakes for for babies < 1500 g

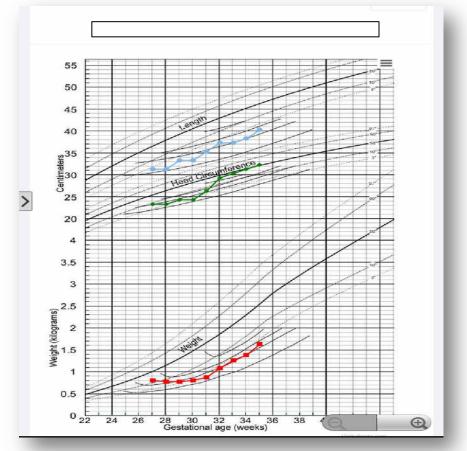
Novel methods for protein fortification are being studied

# SUMMARY (Contd)

Essential to drive and achieve optimal growth amongst all odds

Bedside management tool facilitates better nutrition by providing the interface between nutrition delivery and growth outcome

### LET US PROVIDE THE BEST NUTRITION !!!



### **OPTIMAL GROWTH IS**

**TOMORROW'S OUTCOME!**