Role of Nutrients in Critically Ill Patients

Wafaa Taha Salem, MD, Professor of Anesthesia, Intensive care Vice Dean of NCI, Cairo university

'critically-ill patient'

 The term 'critically-ill patient' refers to a group of patients with diverse diseases,



The population of critically-ill patients is not a homogeneous population

- surgical,
- trauma
- medical .

They have very different or even opposing metabolic responses.

Critically-ill patients is not a homogeneous population







Nutrients requirement critically-ill patient

Macronutrient

- **≻**Carbohydrate
- **≻**Lipids
- **≻**Proteins

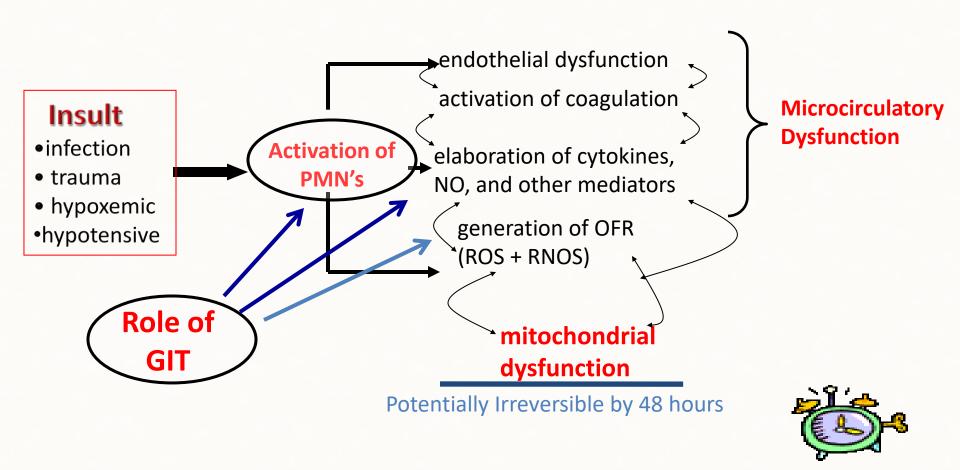
Micronutrient

- **≻**Vitamins
- **≻**Minerals
- ➤ Trace elements

Immunonutrients

- **≻**Glutamine
- ➤ Arginine
- ➤ Omega-3 fatty acids

Pathophysiology of Critical Illness 1



oxidative stress == cellular energetic failure== organ failure

Death

How can Nutrients help the critically ill?

- Provide nutritional substrates to meet protein and energy requirements
- Help protect vital organs and reduce break down of skeletal muscle
- ➤ To provide nutrients needed for repair and healing of wounds and injuries
- To maintain gut barrier function
- Modulate underlying pathphysiological processes in critically ill to improve outcome.

When to start and how to give?

Critically-ill patients who are not expected to receive a complete oral diet for at least 3 consecutive days should receive specialized nutritional support (C).

SEMICYUC: & SENPE Nutr Hosp 2011

Early administration, ranging from 24 to 72 h from admission to the ICU.

More (and Earlier) is Better!



If you feed them (better!)
They will leave (sooner!)

Early vs. Delayed EN: Effect on Infectious Complications



Review: Early Enteral Nutrition vs. delayed nutrient intake

Comparison: 01 Early EN vs. delayed nutrient intake

Outcome: 02 Infectious Complications

Study or sub-category	Early EN n/N	delayed n/N	RR (random) 95% CI	Weight %	RR (random) 95% Cl	Year
Moore	3/32	9/31	+	3.96	0.32 [0.10, 1.08]	1986
Singh	7/21	12/22		9.81	0.61 [0.30, 1.25]	1998
Kompan	9/27	16/25	-	12.53	0.52 [0.28, 0.96]	1999
Minard	6/12	7/15		8.48	1.07 [0.49, 2.34]	2000
Malhotra	54/100	67/100	-	33.98	0.81 [0.64, 1.01]	2004
Peck	12/14	11/13	+	27.04	1.01 [0.74, 1.39]	2004
Nguyen	3/14	6/14		4.21	0.50 [0.15, 1.61]	2008
Total (95% CI) Total events: 94 (Early EN),	220 128 (delayed)	220	•	100.00	0.76 [0.59, 0.98]	
Test for heterogeneity: Chi ² : Test for overall effect: Z = 2	= 9.22, df = 6 (P = 0.16), l ² = 34 .09 (P = 0.04)	.9%				
	······································		0.1 0.2 0.5 1 2	 5 10		
			Favours Early EN Favours dela			

Updated 2009 www.criticalcarenutrition.com

Early vs. Delayed EN: Effect on Mortality



Review: Early Enteral Nutrition vs. delayed nutrient intake

Comparison: 01 Early EN vs. delayed nutrient intake

Outcome: 01 Mortality

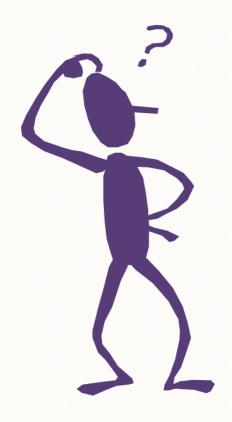
Study or sub-category	Early EN n/N	Delayed n/N	RR (random) 95% CI	Weight %	RR (random) 95% Cl	Year		
Moore	1/32	2/31	+	2.77	0.48 [0.05, 5.07]	1986		
Chiarelli	0/10	0/10			Not estimable	1990		
Eyer	2/19	2/19	+	- 4.44	1.00 [0.16, 6.38]	1993		
Chuntrasakul	1/21	3/17		3.24	0.27 [0.03, 2.37]	1996		
Singh	4/21	4/22		9.76	1.05 [0.30, 3.66]	1998		
Kompan 1999	0/14	1/14	+ +	1.57	0.33 [0.01, 7.55]	1999		
Minard	1/12	4/15	•	3.61	0.31 [0.04, 2.44]	2000		
Pupelis2000	1/11	5/18	-	3.77	0.33 [0.04, 2.45]	2000		
Pupelis	1/30	7/30		3.69	0.14 [0.02, 1.09]	2001		
Dvorak	0/7	0/10			Not estimable	2004		
Kompan 2004	0/27	1/25	• •	1.53	0.31 [0.01, 7.26]	2004		
Malhotra	12/100	16/100		31.58	0.75 [0.37, 1.50]	2004		
Peck	4/14	5/13		13.17	0.74 [0.25, 2.18]	2004		
Nguyen	6/14	6/14		20.86	1.00 [0.43, 2.35]	2008		
Total (95% CI) Total events: 33 (Early EN),		338	•	100.00	0.68 [0.46, 1.01]			
Test for heterogeneity: Chi ² Test for overall effect: Z = 1	= 6.36, df = 11 (P = 0.85), I² = 0 .90 (P = 0.06)	%						
			0.1 0.2 0.5 1 2	5 10				
Favours Early EN Favours delayed								

Updated 2009 www.criticalcarenutrition.com

Routes of Nutrients supply critically ill patient

Enteral route

- 1. Nasogastric tube
- 2. Gastrostomy tube
- 3. jejunostomy
- □ Routine or standard use of the naso-jejunal tube in critically-ill patients is not associated with increased efficacy in provision of enteral nutrition or a lower rate of infectious complications (A).
- ☐ Severe acute pancreatitis, Elevated gastric output. can be considered (C).



What if you can't provide adequate early enteral nutrition?

... to PN or not to PN, that is the question!

Complementary parenteral nutrition should be started when 60% of nutritional requirements are not met at the fourth day of admission, or for at least 2 consecutive days during the hospital stay (C).

Near-Target Caloric Intake in Critically Ill Medical-Surgical Patients Is Associated With Adverse Outcomes

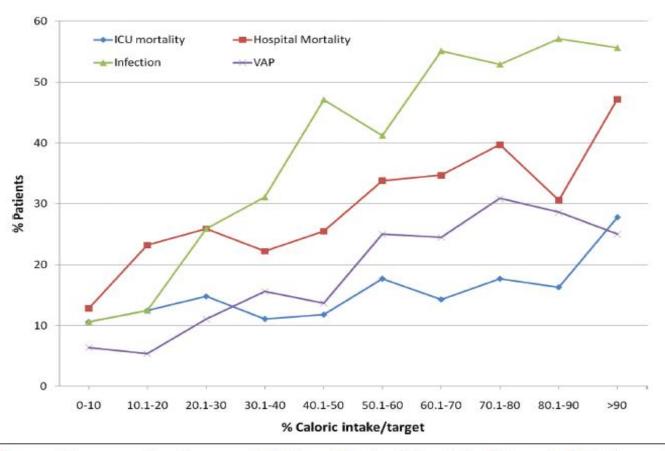
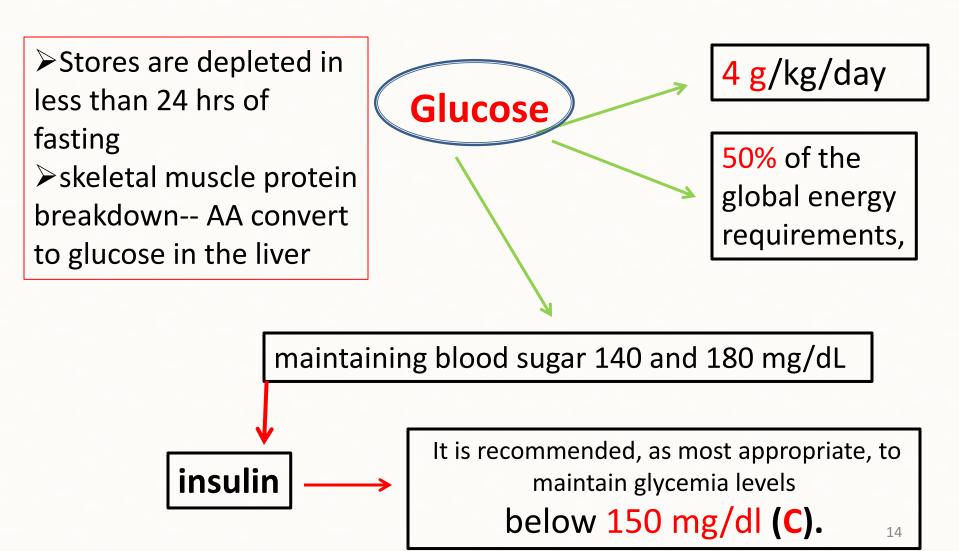


Figure 1. The association among intensive care unit (ICU) mortality, hospital mortality, ICU-acquired infections, and ventilator-associated pneumonia (VAP) rate and caloric intake/requirement.

The role of carbohydrates critically-ill, What type & amount should be supplied



The role of lipids in critically-ill, What type & amount should be supplied

- Providing energy
- concentrated
- isotonic
- non-glucose
- Prevent essential fatty acids deficiency
- Absorption of fat-soluble vitamins
- Maintain the structure of cell membranes
- Modulate intracellular signals
- Modulate immune cell function

lipids

- The recommended lipid supply is 0.7-1.5 g/kg/day
- \triangleright It is recommended to avoid single ω -6 supplies in critically-ill
- \triangleright a high ω -3 content from fish oil should be indicated for patients with acute lung injury (ALI) and (ARDS)
- ➤ The lipid emulsion with mixtures of (MCT), (FO), (OO) well tolerated and are used with preference over LCT
- Up to 40% of non-protein calories may be provided.

- ✓ Preferred concentrations of 30 or 20% vs 10%
- ✓ longer perfusions rather than in short periods



The role of protein in critically-ill, What type & amount should be supplied.

In critically-ill patients, no specific formulation of amino acids has been defined for generic use (C).

Branched chain amino acid's – support immune cell functions.

The there is not sufficient evidence to justify the use of formula with excess branched chain amino acids in critically ill specially in septic patients.

SEMICYUC: & SENPE Nutr Hosp 2011

In general, the supply must be adjusted to an amount of 1-1.8 g/kg/day (B).

1.5 g/kg/day decreases protein catabolism by 70%,

Estimate protein requirement

UUN : 5- 10 **level 1 stress** (1.2 –1.3 g protein/ kg BW)

UUN: 10-25 level 2 stress (2 g protein/ kg BW)

very high supplies are not recommended,

- ➤ 2 g/kg/day causes an increase in net protein degradation, drive urea genesis .
- ➤ No improvement in overall balance
- will not likely promote better nitrogen retention,

Kcal: N ratio

150: 1 (moderate stress) 80-100: 1 (severe stress)

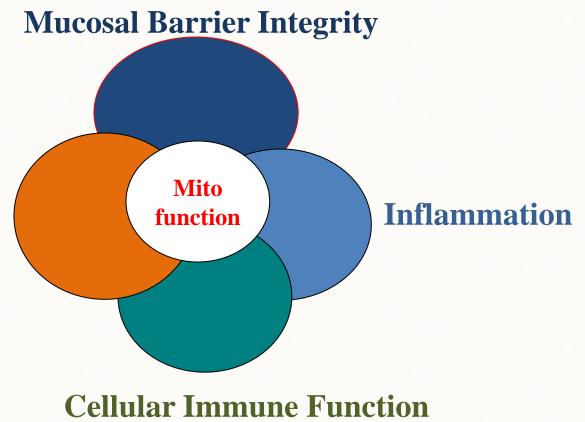
The Role of Immunonutrients in Critically ill

Specific nutrients found to have effects on immune system, metabolism, and GIT structure and function

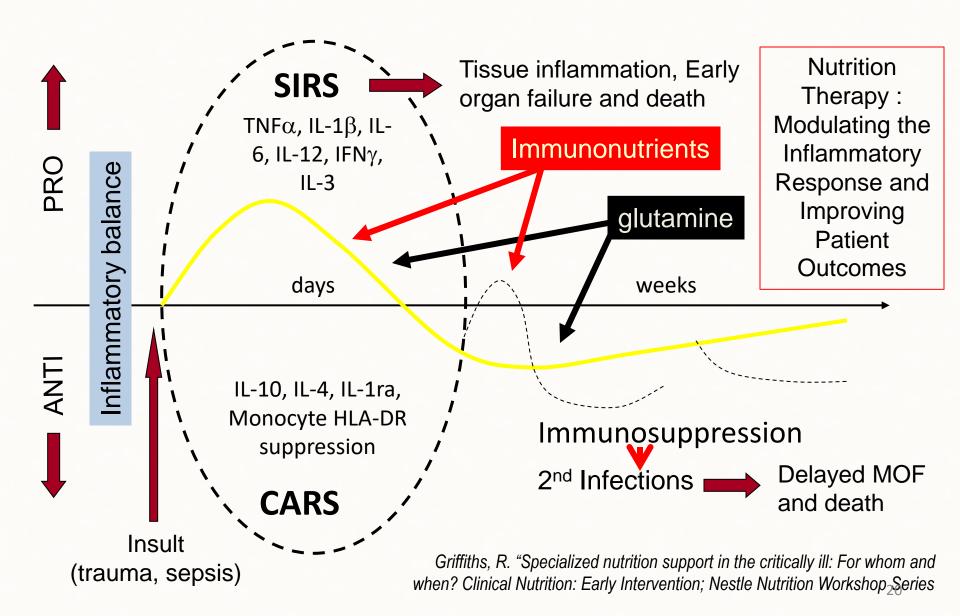


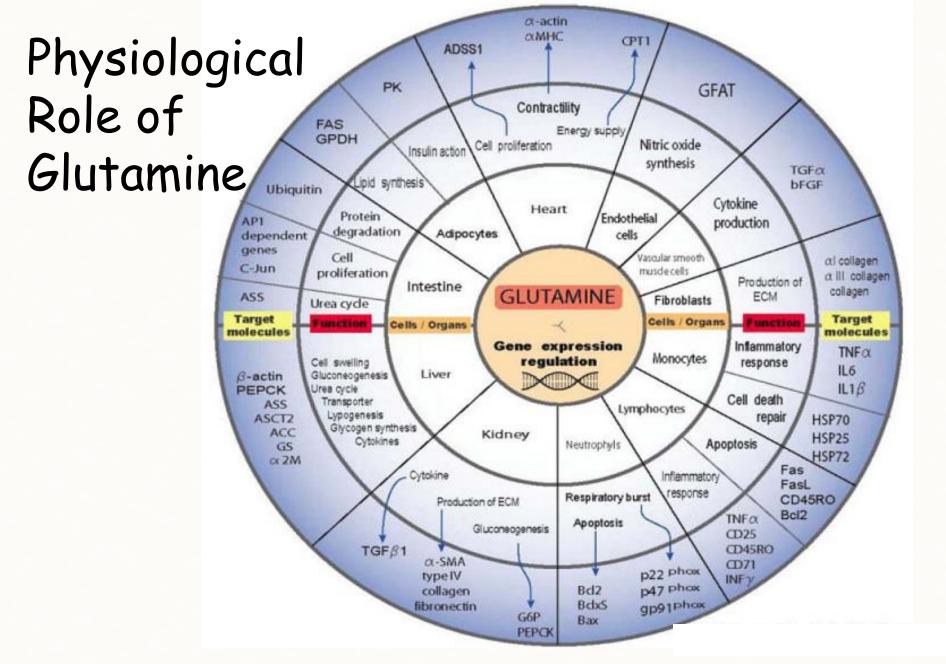
Oxidative stress

Glutamine
Arginine
Omega-3 fatty
acids



Inflammation and organ failure in the ICU





Curi R, et al, J Cell Physiol, 2005

Role of glutamine in Critically ill patients

Glutamine

Conditionally essential amino acid

Ischemia / reperfusion (Better cellular energetics)

Anti-inflammatory (Blunted cytokine burst)

Macrophages (Increased macrophage growth and ATP levels)

Lymphocytes (Increased bacterial killing - stimulated gluthatione)

Enterocytes (Better growth)

iNO synthase (Blunting of iNOs expression)

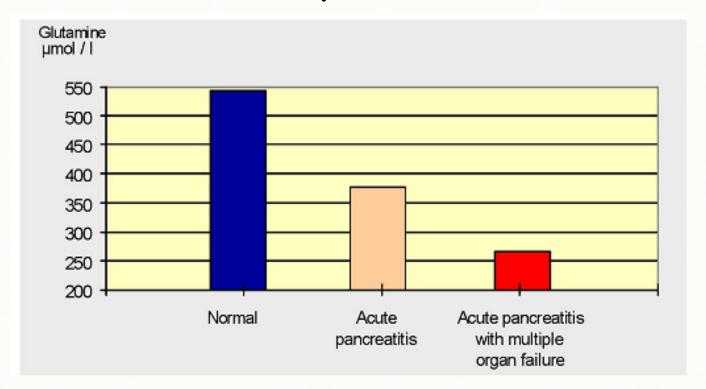
Cellular protection (Stimulated heat shock protein expression)

Oxydative stress (Improved antioxydant defenses)

Glucose metabolism (Reduced glucose resistance)

Bacterial metabolism (Possible bacterial growth in colon)

Glutamine depletion in Plasma and Severity of Disease



- The sicker the patient the higher the glutamine demand
- Low plasma glutamine at admission to ICU associated with the risk of poor outcome

¹ Roth et al. 1986, ² Oudemans-Van Straaten et al. 2001

Is glutamine administration of choice in Critically ill patients or sepsis?

In critically-ill patients intravenous administration of glutamine dipeptide (Ala-Gln) of 0.5 g/kg/day is recommended, complementing parenteral nutrition (A).

Although no studies have been performed in humans to evaluate the effect of glutamine on septic patients receiving PN When parenteral nutrition is indicated, it is recommended to use glutamine supplements (B).

When the patient is receiving EN It is recommend to give IV glutamine, as a supplement,

CLINICAL NUTRITION WEEK 2013 February 9-12, 2013

- •Do not exceed Glutamine recommended dose (up to 0.5 g/kg IBW) in critically ill patients
- •A combined enteral and intravenous administration of glutamine can be used as long as the total does not exceed 0.5 g/ kg IBW

i.v. glutamine-containing product should not be given in patients with

- renal insufficiency
- multi-organ failure incl. metabolic acidosis
- insufficient clinical nutrition

Role of Arginine in Critically ill patients

It is a non essential amino acid

It is reduced in trauma and sepsis.

- >an increase in acute phase reactants,
- ➤ It gives rise to an increase in nitrogenous compounds such as NO. with antibacterial activity
- It can increase substrates necessary for the synthesis of connective tissue (leads to wound healing).
- >action as bowel neurotransmitter
- rightarrows activity in insulin stimulation,
- > regulator of microcirculation
- improve immune function. And promoting cell growth and cell differentiation
- > modulation of cell signals

Are diets with mixtures of Pharmaconutrients indicated in critically ill (IMD)

There is a controversy about the outcomes and recommendations of the different meta analyses about (IMD)

(arginine, ω -3, nucleotides, antioxidants)

➤ In Sepsis; It may be associated with increased mortality.

Montejo JC, et al . Clin Nutr 2003

- There is sufficient evidence to use IMD in critically-ill patients, considering the benefits associated with their use and the lack of harmful effects

 Marik PE, et al Intensive Care Med 2008
- \triangleright A randomized, controlled, prospective study on PN vs EN enriched with Pharmaconutrients (mixture of arginine, ω -3 and antioxidants) in septic patients reported a greater intra-ICU mortality in the enteral group

- ❖ last meta analysis published, concluded that only in the group of patients with sepsis, septic shock, or acute respiratory distress syndrome (ARDS), the use of IMD was associated with a significant decrease of mortality, secondary infections, and stay at the ICU, but provided this formula contained fish oil.

 SEMICYUC: & SENPE Nutr Hosp 2011
- * IMD (arginine, ω-3, antioxidants) in septic patients in a critical condition. is associated with lower mortality compared with the use of a control diet SEMICYUC: & SENPE Nutr Hosp 2011

If unable to tolerate <700ml/d immune modulating formula should be stopped.

Antioxidants, vitamins and trace elements

The plasma concentration of micronutrients with antioxidant capacity decreases in critically-ill, particularly in septic patients

special attention should be paid to the supply of trace elements (particularly selenium, zinc and copper) and vitamins

The need for supplying micronutrients (vitamins and trace elements) is set (A),

But the amount cannot be established.

SEMICYUC: & SENPE Nutr Hosp 2011

- Meta-analysis of 15 randomized studies a combination of antioxidant vitamins and trace elements (selenium, zinc and copper)
 - Reduces mortality and the duration of mechanical ventilation
- Does not improve infectious complications or length of stay
 Canadian Clinical Practice Guidelines 2011

❖ The REDOX study,2013 on the potential beneficial effect of selenium for critically ill patients, Supplementation with antioxidants: selenium (i.v.) <u>plus</u> selenium, Vit. C+E, zinc, ß-carotene (enterally)

No significant difference found for 28-day mortality in any statistical analysis

High-dose selenium supplements alone may not be recommended routinely critically ill patients

Summary

The main Role of Nutrients is Improve Survival of Critically III Patients

- Energy: Carbohydrate and fat intake frees up protein (essential amino acids and nitrogen) so that it can be used for tissue building.
- Antioxidants, Vitamins and minerals: Control protein and energy metabolism through their coenzyme roles.
- Immunonutrients: Modulate underlying pathphysiological processes

