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- The Acute Inflammatory Response
  - Cellular activation
  - Inflammatory mediators (TNF, IL1, etc)
  - Paracrine Vs endocrine effects

- The Endothelium
  - Selectins, Integrins, and ICAMs
  - Nitric Oxide
  - Tissue Factor

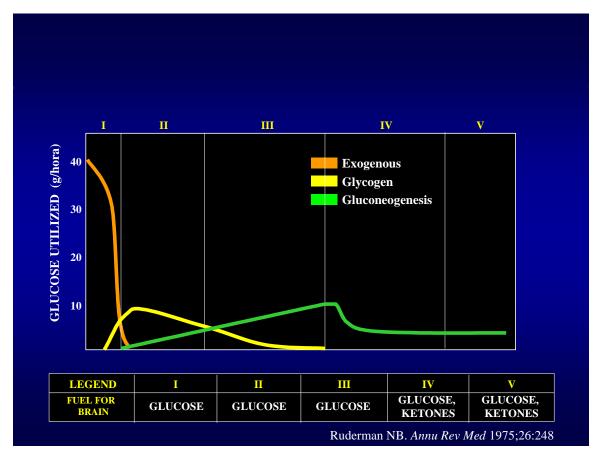
- Afferent Nerve Stimulation
  - Sympathetic Nervous System
  - Adrenal Gland Medulla

- The Endocrine System
  - Pituitary Gland (GH, ACTH, ADP)
  - Adrenal Gland (Cortisol, Aldosterone)
  - Pancreatic (Glucagon,  $\downarrow$  Insulin)
  - Others (Renin, Angiotensin,  $\downarrow$  Sex hormones,  $\downarrow$  T4)

#### Consequences of the Response

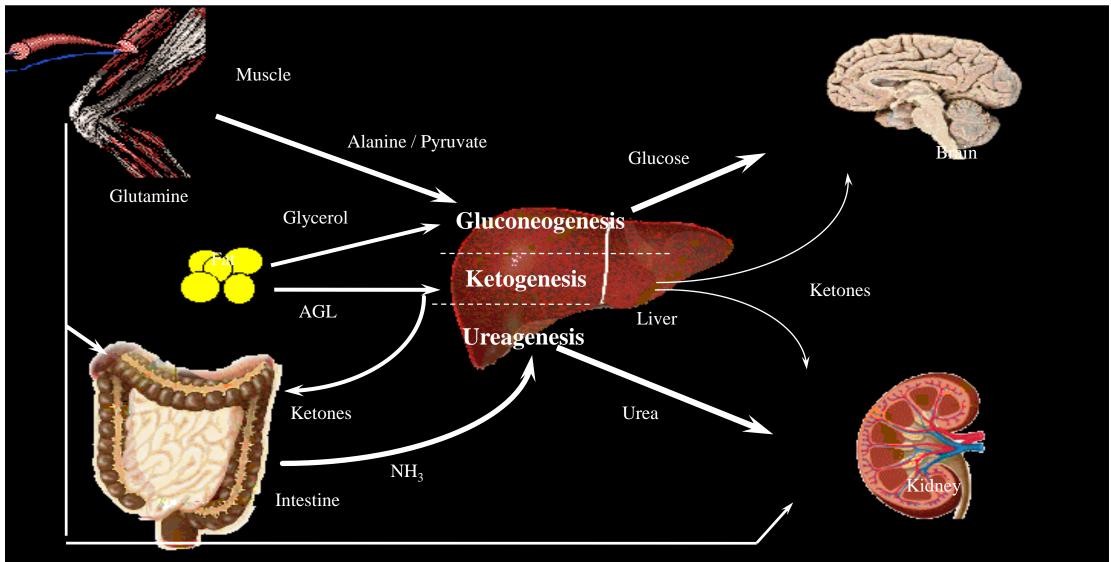
- Limiting injury
- Initiation of repair processes
- Mobilization of substrates
- Prevention of infection
- Distant organ damage

#### Metabolic Response to Fasting

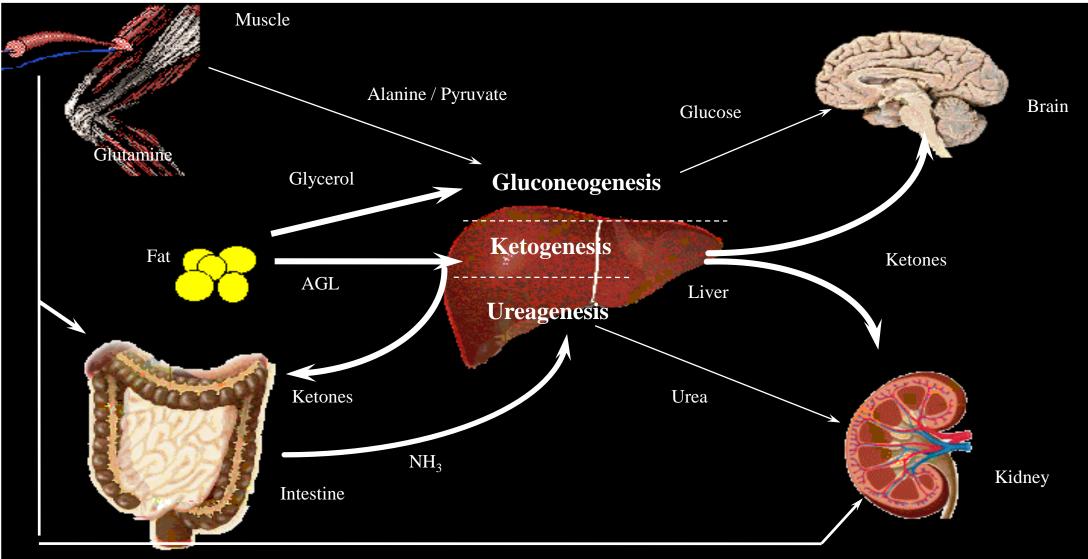


 The carbohydrate deposits of the body last about 18 to 20 hours and new glucose is produced through gluconeogenesis of amino acids from the lean body mass

#### Starvation – Early Stage



#### Starvation – Late Stage



#### Metabolic Response to Starvation

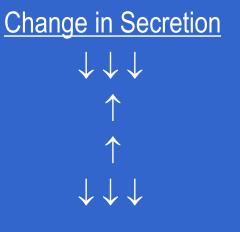
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Hormone
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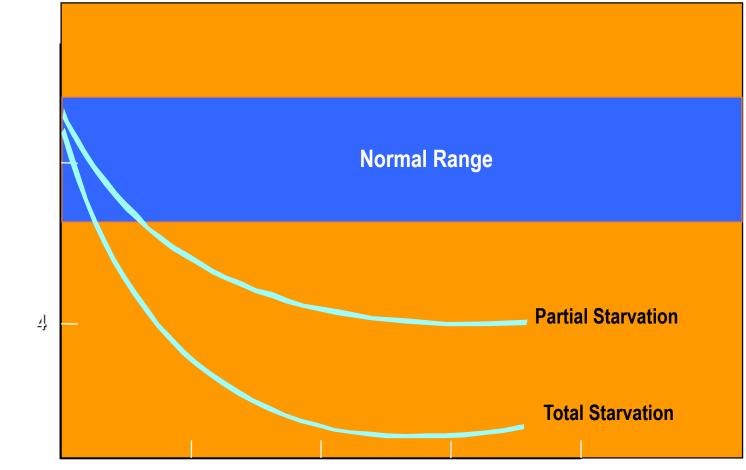
Norepinephrine Norepinephrine Epinephrine Thyroid Hormone T4

#### <u>Source</u>

Sympathetic Nervous System Adrenal Gland Adrenal Gland Thyroid Gland (changes to T3 peripherally)

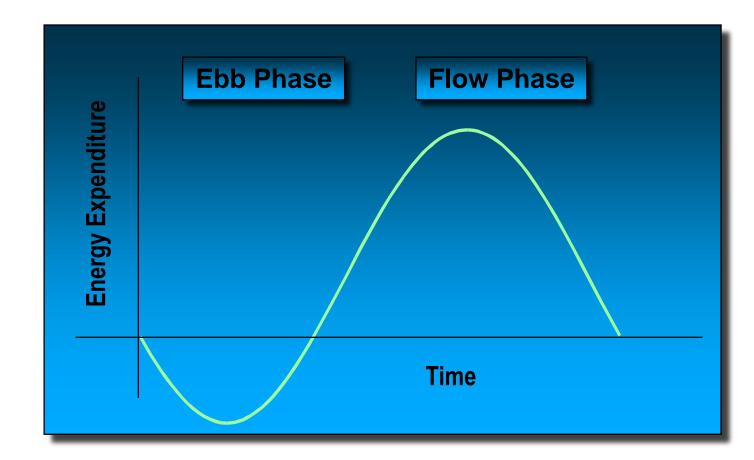


#### Energy Expenditure in Starvation



**Days** Long CL et al. *JPEN* 1979;3:452-456

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Cutherbertson DP, et al. Adv Clin Chem 1969;12:1-55

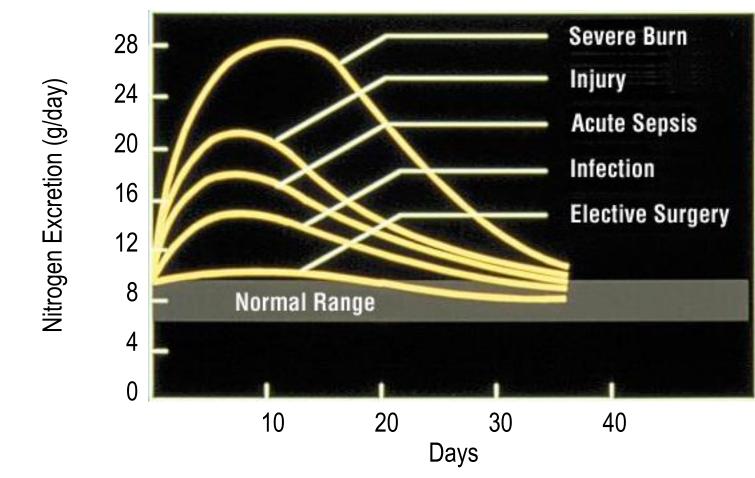
### Metabolic Response to Injury: Ebb Phase

- Characterized by hypovolemic shock
- Priority is to maintain life/homeostasis
- $\downarrow$  Cardiac output
- $\downarrow$  Oxygen consumption
- $\downarrow$  Blood pressure
- $\downarrow$  Tissue perfusion
- $\downarrow$  Body temperature
  - $\downarrow$  Metabolic rate

### Metabolic Response to Injury: Flow Phase

- $\uparrow$  Catecholamines
- $\uparrow$  Glucocorticoids
- $\uparrow$  Glucagon
- Release of cytokines, lipid mediators
- Acute phase protein production



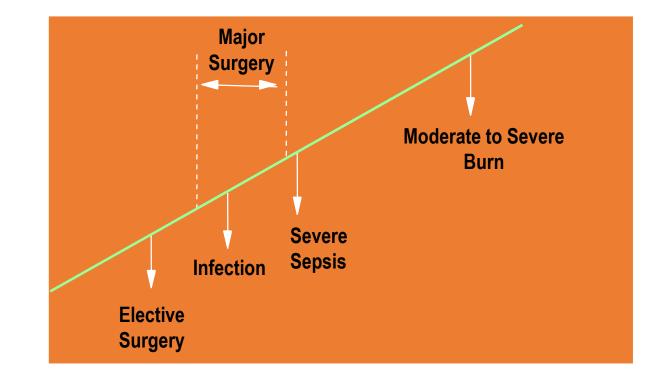


Long CL, et al. *JPEN* 1979;3:452-456

# Severity of Injury: Effects on Nitrogen Losses and Metabolic Rate

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Nitrogen Loss in Urine



**Basal Metabolic Rate** 

Adapted from Long CL, et al. JPEN 1979;3:452-456

#### Comparing Starvation and Injury

StarvationTrauma or DiseaseMetabolic rate↓↑ ↑Body fuelsconservedwastedBody proteinconservedwastedUrinary nitrogen↓↑ ↑Weight lossslowrapid

The body adapts to starvation, but not in the presence of critical injury or disease.

Popp MB, et al. In: Fischer JF, ed. Surgical Nutrition. 1983.

### Modifying the Response

- Medication (before or after injury)
- Nutritional status
- Severity of injury
- Temperature
- Anesthetic technique

#### Summary

- Injury (Trauma or Surgery) leads to a metabolic response
- Metabolic response to injury is an adaptive response
- Metabolic response could overwhelm the body and lead to increased morbidity and mortality
- We can modify the metabolic response before and sometimes after injury

Questions