

## Hormonal Regulation of Metabolism

	<b>Insulin</b>	<b>Glucagon</b>	<b>Epinephrine</b>	<b>Cortisol</b>	<b>Growth Hormone</b>
<b>Liver</b>					
Glycogen breakdown	↓	↑	↑		
Glycogen synthesis	↑	↓	↓	↑	
Gluconeogenesis	↓	↑	↑	↑	
Glycolysis	↑	↓	↓		
Glucose release	↓	↑	↑		
Glucose uptake	↑	↓	↓		
Glucagon receptor					↑
<b>Skeletal Muscle</b>					
Glycogen breakdown	↓		↑		
Glycogen synthesis	↑		↓		
Glycolysis			↑†	↓	
Glucose uptake	↑		†	↓	
Protein catabolism	↓			↑	
Amino acid uptake	↑			↓	
Amino acid release	↓			↑	
<b>Adipose Tissue</b>					
Lipolysis	↓		↑	↑	↑
Glucose uptake	↑			↓	
<b>Pancreas</b>					
Insulin release	↓	↑	↓	↓	
Glucagon release	↓	↓	↑		
<b>Systemic Effects</b>					
Insulin action	↑*			↓	↓

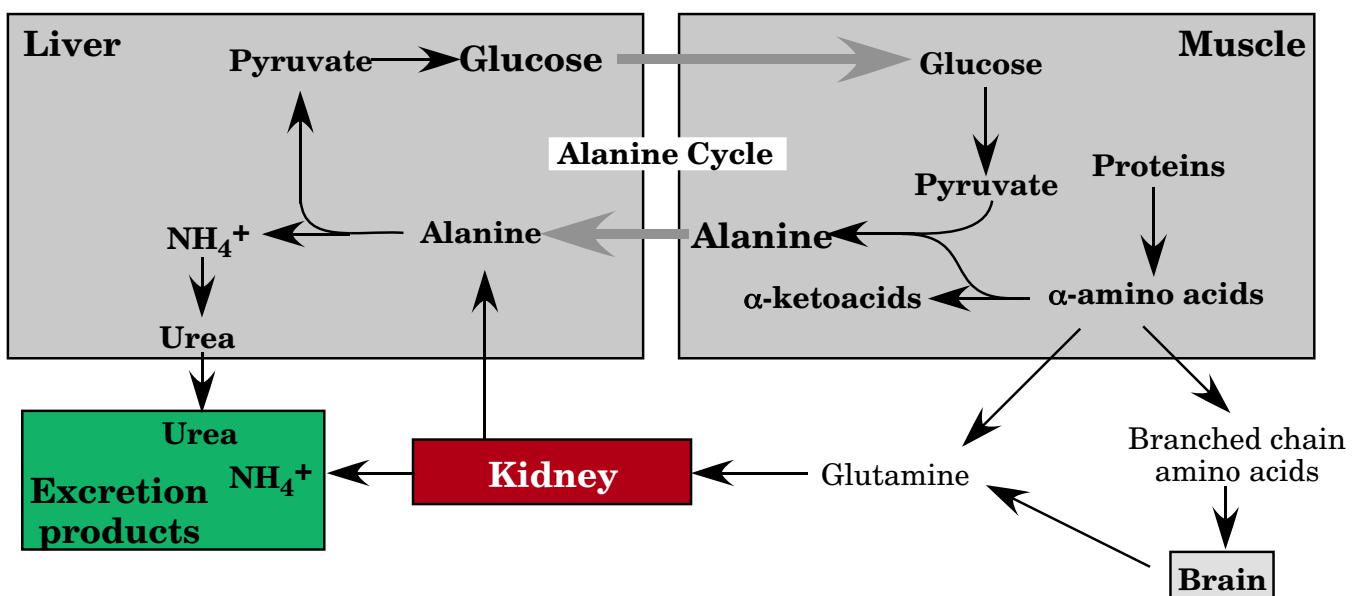
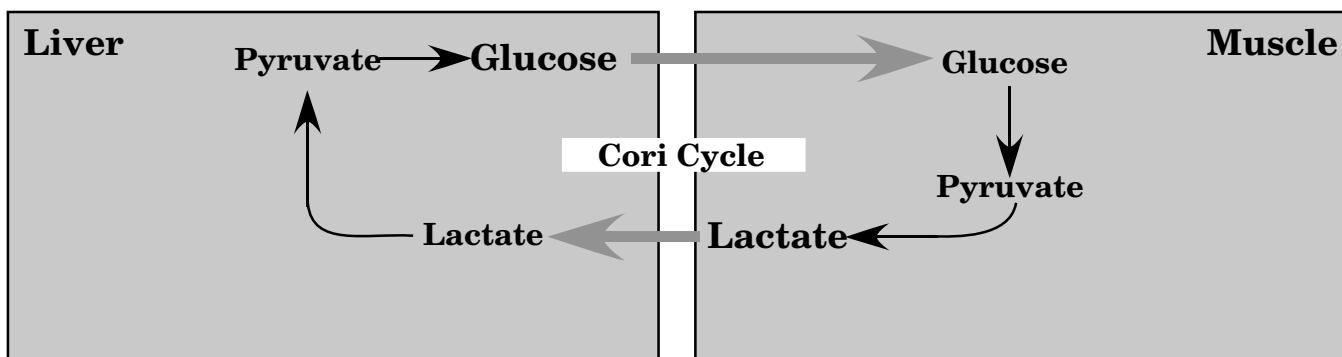
\*Note: Insulin obviously has “insulin actions”, hence the ↑; however, prolonged high levels of insulin decrease the insulin response in target tissues.

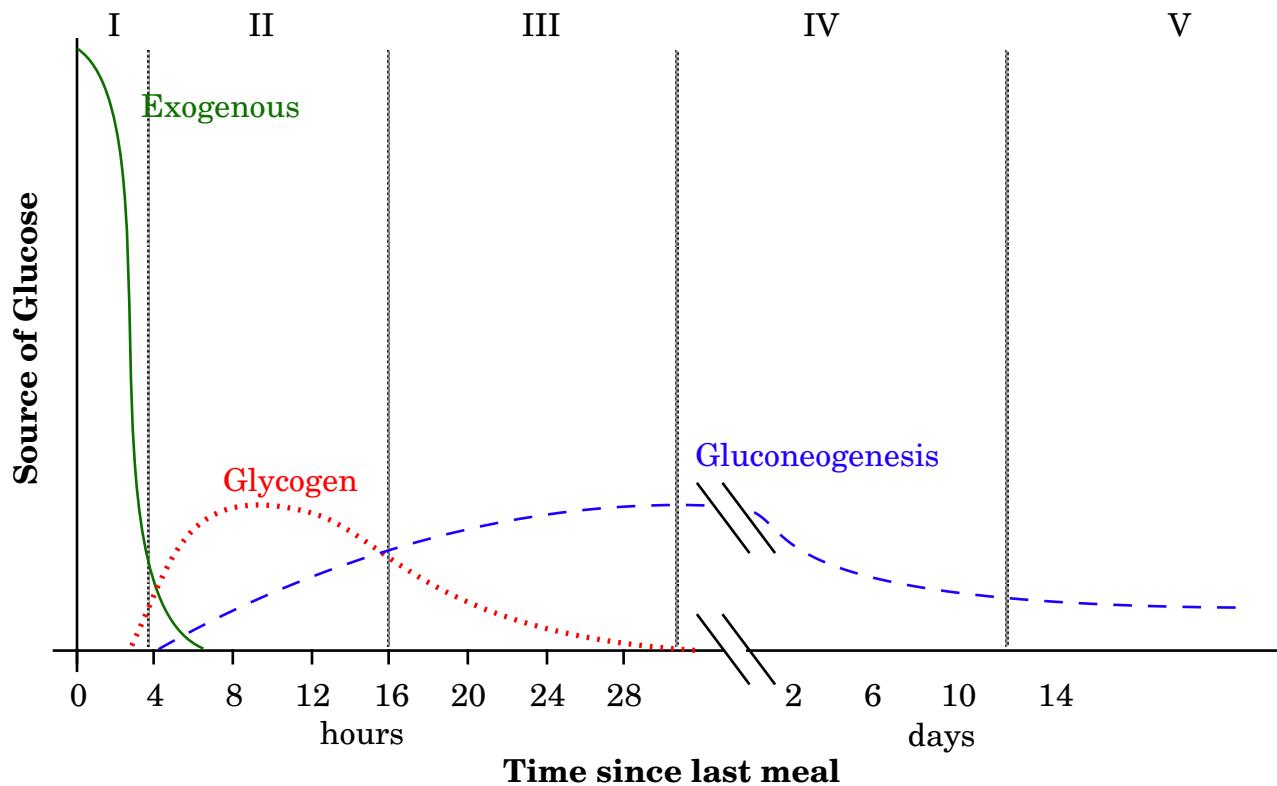
†Note: epinephrine-induced effects on muscle glucose metabolism are relatively small in the absence of exercise. Glucose uptake in muscle is stimulated by exercise, but is probably not directly affected by epinephrine.

### Fuel reserves of “typical” 70 kg individual

Organ	Available energy (kcal)		
	Glucose/glycogen	Triacylglycerols	Degradable Protein
Brain	8	0	0
Blood	60	45	0
Liver	400	450	400
Muscle	1200	450	24,000
Adipose tissue	80	135,000	40

(modified from Stryer (1995) *Biochemistry*, 4th Ed.)





	I	II	III	IV	V
<b>Origin of glucose</b>	Diet	Glycogen, Hepatic Gluconeogenesis	Hepatic Gluconeogenesis, Glycogen	Hepatic Gluconeogenesis, Renal Gluconeogenesis	Hepatic Gluconeogenesis, Renal Gluconeogenesis
<b>Major Brain fuel source</b>	Glucose	Glucose	Glucose, branched chain AA	Glucose, AA, Ketone bodies	Ketone bodies, Glucose
<b>Tissues using glucose</b>	All	All except liver	All except liver	Brain, Blood cells, muscle (for exercise)	Brain, Blood cells
<b>Hormonal response</b>	Insulin ↑	Insulin ↓ Glucagon ↑	Insulin ↓ Glucagon ↑ Cortisol ↑	Insulin ↓ Glucagon ↑ Thyroid hormone ↓	Insulin ↓ Glucagon ↑ Thyroid hormone ↓
<b>Change in Muscle protein</b>	↑	—	↓↓↓	↓↓	↓
<b>Change in Adipose stores</b>	↑	↓	↓↓	↓↓↓	↓↓

## **Diabetes Mellitus: Type I**

Insulin-dependent diabetes mellitus (IDDM)  
“Juvenile-onset” diabetes

## **Diabetic Ketoacidosis (DKA)**

Frequent urination (Polyuria) – Osmotic diuresis

Extreme thirst (Polydipsia)

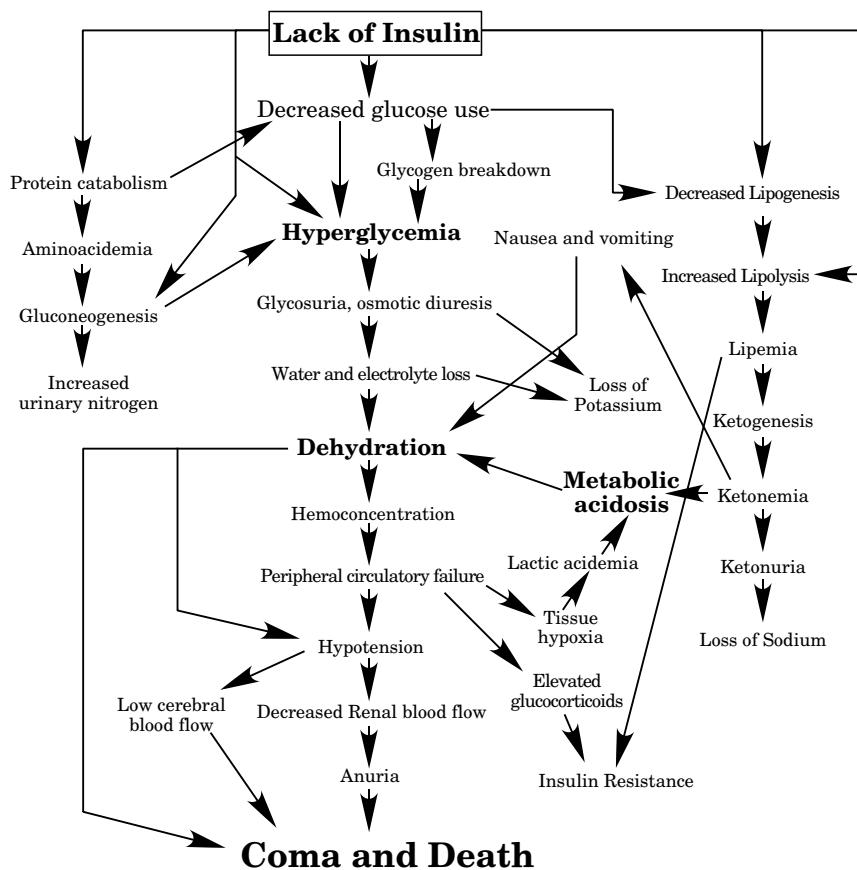
Extreme hunger (Polyphagia)

Peripheral neuropathy

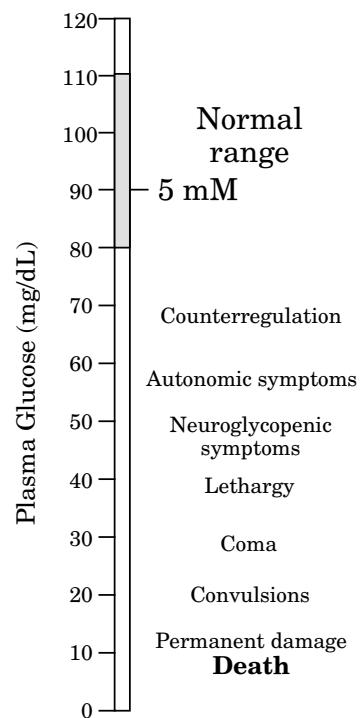
Weakness

Kussmaul respiration

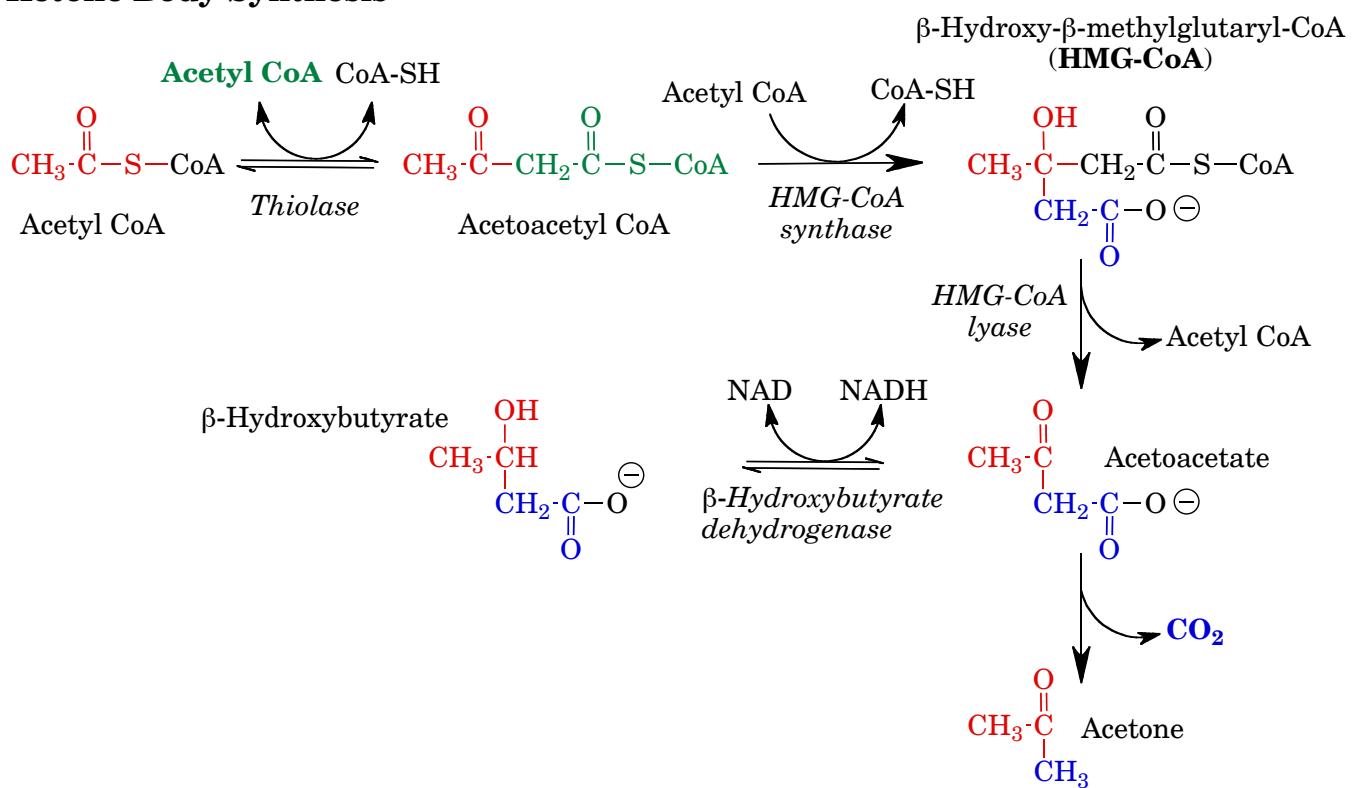
Vision alterations



## **Too much insulin**



## Ketone Body Synthesis



## Diabetes Mellitus: Type II

Non-insulin-dependent diabetes mellitus (NIDDM)  
“Adult-onset” diabetes

**Hyperglycemic HyperOsmotic Non-Ketotic Coma (HONK)**

## Oral Antihyperglycemic Drugs

