

Biochemistry
Team 434

CREATINE METABOLISM

Musculoskeletal Block

Color index

Red= Important

Purple= Addition

Orange= Explanation

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Objectives:

- 1.To study the importance of creatine in muscle as a storage form of energy
- 2.To understand the biosynthesis of creatine
- 3.To study the process of creatine degradation and formation of creatinine as an end product
- 4.To understand the clinical importance of creatinine as a sensitive indicator of kidney function
- 5.To study different types of creatine kinase (CK) and their clinical importance

creatine synthesis

◆ the synthesis of creatine requires 3 amino acids : shown in green

- Glycine
- Arginine
- Methionine (as S-adenosylmethionine → is the activation form of methionine)

*this process happens in the **liver** and **kidney** [NOTE:the liver is the creatine synthesis site]

*From liver, transported to other tissues

***98%** are present in skeletal and heart muscles

*In **Muscle**, gets converted to the high energy source creatine phosphate (NOTE:

in presence of ATP and Creatine Kinase)

Kidneys

Arginine + Glycine

Ornithine

Amidino-
transferase

Guanidinoacetate

★ SAM

★ SAH

Methyltransferase

Creatine

Muscle

ATP

ATP

ADP

ADP + H⁺

Creatine Kinase

Creatine phosphate

*s.adenosylmethionine *s.adenosylhomocystine

Creatine Phosphate

High energy phosphate compound

Acts as **storage form of energy** in muscle

Provides small but ready form of energy
(during first minutes of intense muscular contraction)

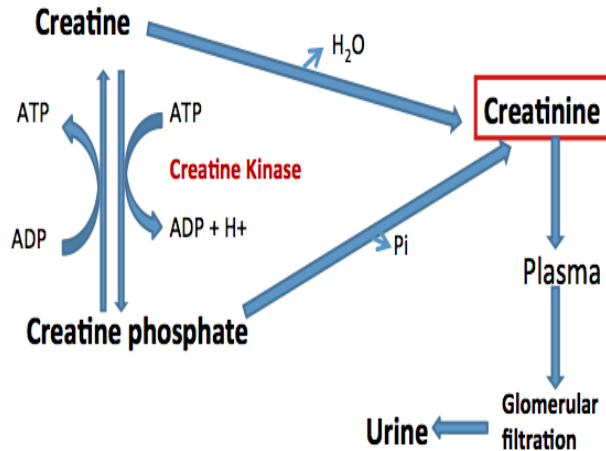
Amount of Creatine Phosphate in body *~proportional to~* Muscle mass

Urinary Creatinine

(1) Typical male excretes about **15 mmol** of Creatinine per day

(2) **Decrease** in muscle mass >> **Decrease** in level of Creatinine in urine
(due to muscular dystrophy or paralysis)

(3) Amount of Creatinine in urine is used as an indicator for the proper collection of 24 hours urine sample



Creatine Degradation

(1) **Creatine + Creatine Phosphate** >spontaneously form> **Creatinine** (end product)
(NOTE:spontaneously means in absence of any enzyme)

(2) Creatinine **excreted in urine**

(3) Serum Creatinine: -Sensitive **indicator of kidney disease** (*Kidney Function Test*)
-**Increases** with impairment of kidney function

*(NOTE: creatinine is called **anhydrous creatine** because it does not have a water)*

Creatine Kinase (CK)

CK levels are changed in disorders of *cardiac* and *skeletal muscles*

Responsible for **generation of energy**
(in *contractile muscular tissue*)

Has **3 Isoenzymes**:
CK-MM (mainly in *Skeletal Muscles*)
CK-MB (mainly in *Heart Muscles*)
CK-BB (mainly in *Brain*)

Required for conversion of
Creatine *>into>* *Creatine Phosphate*

Serum Total **CK increased** in:
Crush Injuries (damage of skeletal muscles)
Myocardial Infarction (damage of heart muscle)

Summary

CK has 3 isoenzymes:
CK-BB in Brain
In skeletal muscle CK-MM
CK-MB
In heart muscle.
Serum total CK is increased in Crush injuries, Myocardial infarction

is responsible for the generation of energy in contractile muscular tissues

Three amino acids are required:

Glycine
Arginine
Methionine (as S-adenosylmethionine)
Site of biosynthesis:
Step 1: Kidneys
Step 2: Liver

biosynthesis

Distribution

Storage form of energy

Degradation

98% are present in skeletal and heart muscles

Creatine Phosphate Acts as a storage form of energy in the muscle

Creatinine is end product of both **creatine** and **creatine phosphate**. Creatinine is excreted in the **urine**

Creatine Phosphate

Creatine kinase

creatine

Creatinine level in serum indicate kidney function it will increase in case of **impairment kidney function**.

The amount of creatine phosphate in the body is proportional to the muscle mass

◆ Quiz your self

<https://www.examttime.com/en-US/p/1809224>

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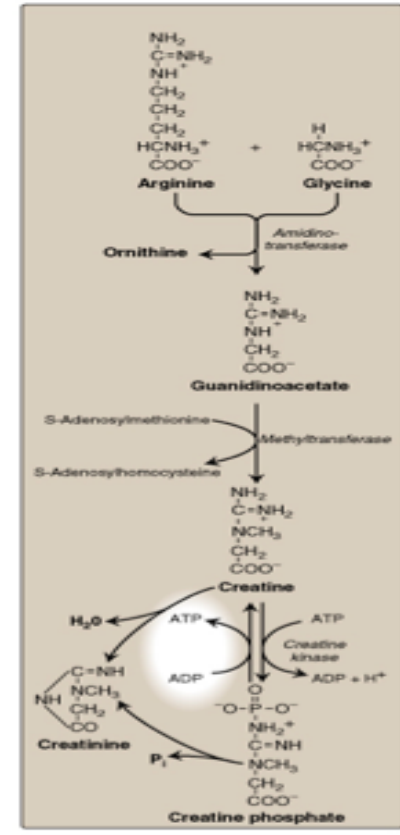


Figure 21.16
Synthesis of creatine.