

بسم الله الرحمن الرحيم



Biochemistry Team 43'





Color index: Doctors slides Doctors' notes Extra information

Musculoskeletal block

Objectives:

- ◆ Recognize the importance of ATP as energy source in skeletal muscle
- Compare three systems of energy transfer in the body
- ◆ Differentiate between energy metabolism in red and white muscle fibers
- ✤ Understand how skeletal muscles derive ATP from aerobic and anaerobic metabolism
- $\boldsymbol{\diamondsuit}$ Discuss the importance of Cori and glucose-alanine cycles in energy metabolism

Overview:

- ✤ Three systems of energy transfer
- ✤ ATP as energy source
- ✤ Aerobic metabolism: red muscle fibers
- ✤ Anaerobic metabolism: white muscle fibers
- ✤ Cori cycle Glucose-alanine cycle
- ✤ Muscle fatigue and endurance in athletes

Systems of energy transfer



Notice that the three systems are overlapping

ATP as an energy source

The nucleotide coenzyme adenosine triphosphate (ATP) is the most important form of chemical energy stored in cells

Breakdown of ATP into ADP+PO4 releases energy

This energy is used for all body functions (biosynthesis, membrane transport, muscle contraction, etc.)

The main pathway for ATP synthesis is oxidative phosphorylation catalyzed by the respiratory chain ATP synthase catalyzes the synthesis of ATP

ADP+Pi ---> ATP

ATP SYNTHASE

ENERGY METABOLISM IN MUSCLE

- Muscle contraction requires high level of ATP consumption
- Without constant resynthesis, the amount of ATP is used up in less than 1 sec. of contraction



Overview of Energy Metabolism in Skeletal Muscle



Aerobic metabolism in red muscle fibers

- Red muscle fibers are suitable for prolonged muscles activity
- Their metabolism is mainly Aerobic and Depends on adequate supply of Oxygen .
- They obtain ATP mainly from fatty acids.
- Fatty acids are broken down by b-oxidation,Krebs cycle, and the respiratory chain.
- The Red color of the muscle is due to myoglobin. "It has a lot of mitochondria as well"
- Myoglobin has higher oxygen affinity Than hemoglobin
- It releases oxygen when its level drops
- 1) Fatty acid are broken down by B-oxidation to produce acetyl CoA
- 2) Acetyl CoA enters the tricarboxylic acid cycle and produce NADH
- 3) NADH enters the respiratory chain
- 4) In the respiratory chain the final acceptor of the electron is oxygen which gets converted to water and makes ATP
- Another way to produce ATP molecule : (produce ATP in less amounts)
- Adenylate kinase (1) takes 2ADP joins them to make 1ATP and 1AMP



the process of degrading fatty acids by cutting the chains into units made of 2 carbons, making acetyl CoA

Anaerobic Metabolism in White Muscle Fibers

- White muscle fibers are suitable for fast, strong contractions
- They mainly obtain ATP from anaerobic glycolysis.
- During intense muscle activity (weightlifting, etc.) O2 supply from blood quickly drops, then the muscle will use anaerobic glycolysis.
- They have supplies of glycogen that is catabolized and undergoes glycolysis.
- Glycogen → glucose-1-PO4 → glucose-6-PO4 → glycolysis → ATP
- NADH+H+ is **re-oxidized** to maintain glucose degradation and ATP formation.
- Anaerobic glycolysis produces lactate
- Lactate is resynthesized into glucose in the liver by Cori cycle



The Cori Cycle : the resynthesize of Lactate into glucose in the liver

In anaerobic glycolysis

Glucose is transported back to muscles to be reused





skeletal muscles can 't produce new glucose from lactate because:

Gluconeogenesis requires much more ATP than is supplied by glycolysis in muscle تصنيع الجلركرز يتطلب طاقة عالية ما تقدر تصنعها عمليات الجلايكرلس بالمحنلات

O2 deficiencies do not arise in the liver even during intense exercise

Therefore, liver always has sufficient ATP for gluconeogenesis

ولان الكبد ما يحصل فيه نقص بالاكسجين حتى خلال التمارين المكثفة فهو مصدر عالى للطاقة وهو المكان المناسب لتصنيع الجلوكوز من اللاكتيت

The glucose-alanine cycle



Liver Blood Muscle

- Glucose

Glycogen

- α-Amino acid

transamination

× α-Ketoacid

Pvruvate

Alanine

Exercise and AMPK

In exercise, the metabolic enzymes are regulated through phosphorylation by AMP-activated protein kinase (AMPK)





AMPK activation shuts down ATP-requiring processes and stimulates ATP-producing processes

Muscle fatigue and endurance in athletes

• **Muscle fatigue:** Inability of muscles to maintain a particular strength of contraction over time.



- Athletes are able to change the proportions of red and white muscle fibers by targeted training.
- The expression of muscle proteins (and enzymes) can also change during the course of training.
- This provides them with:
- High endurance during muscle activity.
- Efficient energy producing and consumption.
- Delayed fatigue.

<u>436:</u>

Athletes should train to have more red muscle fibers also they can convert white muscle fibers to red The red muscle fibers are better for them because it is suitable for prolonged effort + it is an aerobic which doesn't cause lactic acid formation which leads to muscle fatigue.

Take home messages

- ATP is an important source of chemical energy needed by the cells to perform body functions
- Muscular activity requires constant supply of ATP for energy either from aerobic or anaerobic metabolism
- Cori and glucose-alanine cycles play an important role in regenerating glucose for energy
- Athletes are able to change proportions of their red and white muscle fibers with appropriate training

Summary

Three systems of energy transfer

Type of system:	Immediate system (ATP-PCr)	<u>Short term</u> <u>system</u> (glycolysis)	<u>Long term</u> <u>system</u> (aerobic)
Type of metabolis m:	Anaerobic	Anaerobic	Aerobic
Type of energy source:	Phosphocreat ine (PCr)	Glucose	Fatty acid
type of exercise:	High intensity exercise	High intensity exercise	Continuous exercise
Duration	3-15 sec	15 sec. to 2 min.	hours

<u>ATP</u>

- The main pathway for ATP synthesis is oxidative phosphorylation catalyzed by the respiratory chain
- The nucleotide coenzyme adenosine triphosphate (ATP) is the most important form of chemical energy stored in cells
- Without constant re synthesis, the amount of ATP is used up in less than 1 sec. of contraction.

Energy metabolism in muscle



there are 4 ways for muscle fibers to get energy

1- aerobic by red fibers 2-anaerobic by white fibers 3-creatine phosphate 4- by the enzyme adenylate cyclase; (will take two ADP and produce 1 ATP & 1AMP)

The Cori Cycle In anaerobic glycolysis, the glucose is converted to lactate, lactate in muscle is released into blood, transported to the liver, liver converts lactate into glucose via gluconeogenesis, the newly formed glucose is transported to muscles to be used for energy again.

Exercise and AMPK exercise, the metabolic enzymes are regulated thru phosphorylation by AMP-activated protein kinase (AMPK)

<u>Muscle fatigue</u> Inability of muscles to maintain a particular strength of contraction over time. Causes: muscle damage, accumulation of lactic acid.

MCQs

1. Which one is the shortest system of energy transfer	4' B
	3. B
A) ATP-PCr B) glycolysis C)aerobic	2. B
	A .1
2. Fatty acid are broken down by	: sıəwanA
A) alpha oxidation B) beta oxidation C) gamma oxidation	

9[.] B

- 3. Red color in red muscle fiber due to the rich of
- A) hemoglobin B) myoglobin C) Oxygen
- 4. What is the normal fate of lactate in muscle
- A) gluconeogensis in muscle B) released into bloodstream C) accumulate in muscle
- 5. Pyruvate and is converted to alanine
- A) NH3 B) NH2 C) NH

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