#### **Cellular Respiration**





February 27<sup>th</sup>, 2012

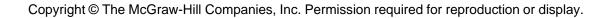
# Outline

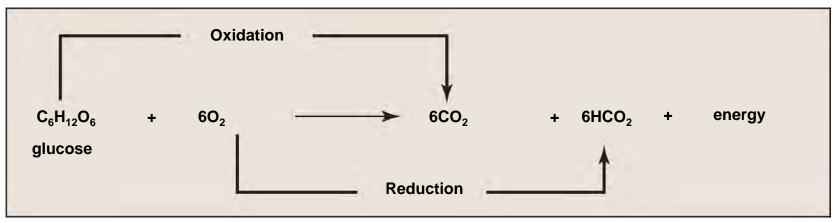
- Cellular Respiration
  - NAD+ and FAD
  - Phases of Cellular Respiration
- Glycolysis
- Fermentation
- Preparatory Reaction
- Citric Acid Cycle
- Electron Transport System
- Metabolic Pool
  - Catabolism
  - Anabolism

## **Cellular Respiration**

- A cellular process that breaks down carbohydrates and other metabolites with the concomitant buildup of ATP
- Consumes oxygen and produces carbon dioxide (CO<sub>2</sub>)
  - Cellular respiration is an aerobic process.
- Usually involves breakdown of glucose to CO<sub>2</sub> and water
  - Energy extracted from glucose molecule:
    - Released step-wise
    - Allows ATP to be produced efficiently
  - Oxidation-reduction enzymes include NAD<sup>+</sup> and FAD as coenzymes

#### **Glucose Breakdown: Summary Reaction**





 Electrons are removed from substrates and received by oxygen, which combines with H+ to become water.

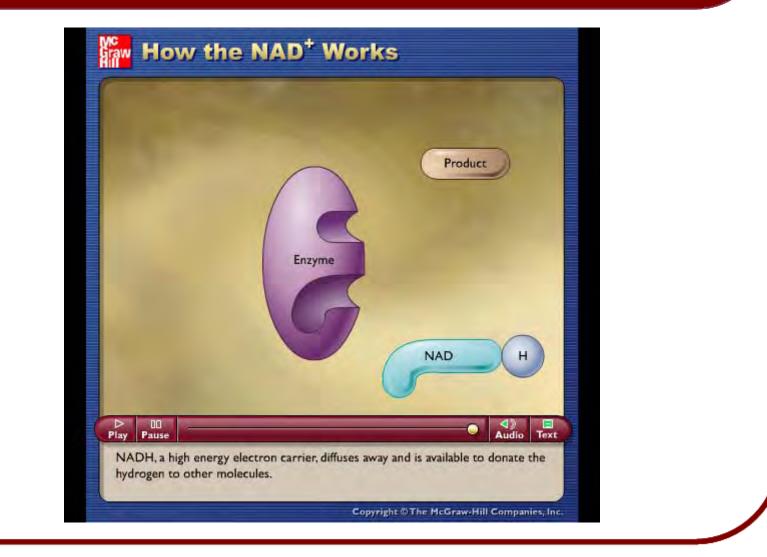
Glucose is oxidized and O<sub>2</sub> is reduced

#### NAD<sup>+</sup> and FAD

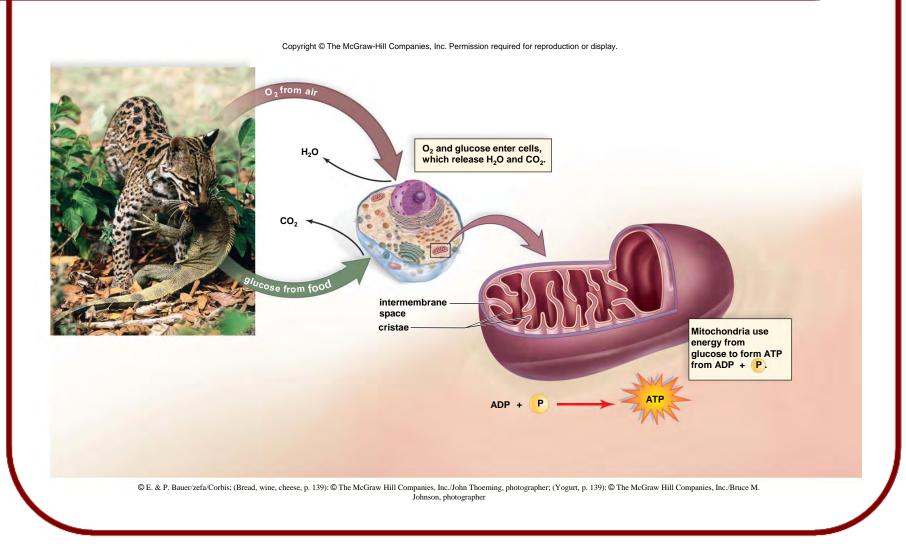
#### NAD<sup>+</sup> (nicotinamide adenine dinucleotide)

- Called a coenzyme of oxidation-reduction. It can:
  - Oxidize a metabolite by accepting electrons
  - Reduce a metabolite by giving up electrons
- Each NAD<sup>+</sup> molecule used over and over again
- FAD (flavin adenine dinucleotide)
  - Also a coenzyme of oxidation-reduction
  - Sometimes used instead of NAD<sup>+</sup>
  - Accepts two electrons and two hydrogen ions (H<sup>+</sup>) to become FADH<sub>2</sub>

#### Animation



#### **Cellular Respiration**



# Phases of Cellular Respiration

- Cellular respiration includes four phases:
  - Glycolysis is the breakdown of glucose into two molecules of pyruvate
    - Occurs in cytoplasm
    - ATP is formed
    - Does not utilize oxygen
  - Transition (preparatory) reaction
    - Both pyruvates are oxidized and enter mitochondria
    - Electron energy is stored in NADH
    - Two carbons are released as CO<sub>2</sub> (one from each pyruvate)

#### Phases of Cellular Respiration

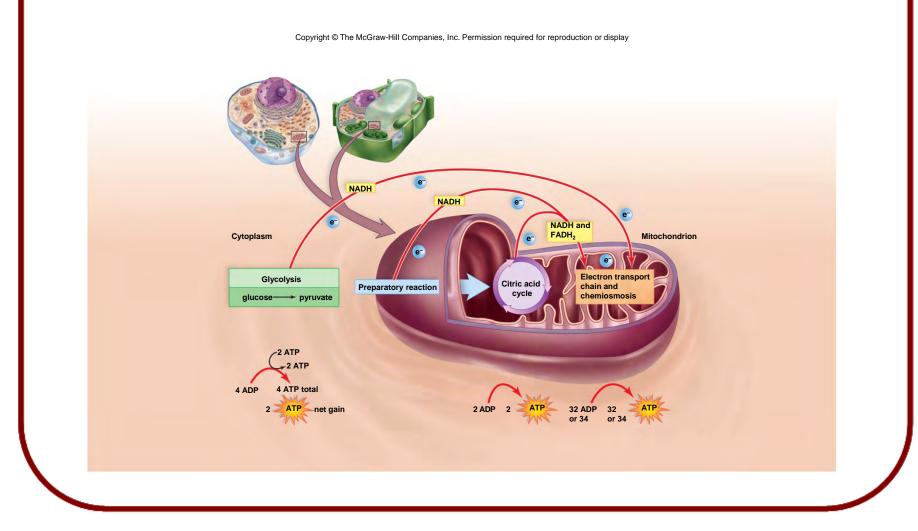
#### Citric acid cycle

- Occurs in the matrix of the mitochondrion and produces NADH and FADH<sub>2</sub>
- In series of reaction releases 4 carbons as CO<sub>2</sub>
- Turns twice (once for each pyruvate)
- Produces two immediate ATP molecules per glucose molecule

#### Electron transport chain

- Extracts energy from NADH & FADH<sub>2</sub>
- Passes electrons from higher to lower energy states
- Produces 32 or 34 molecules of ATP

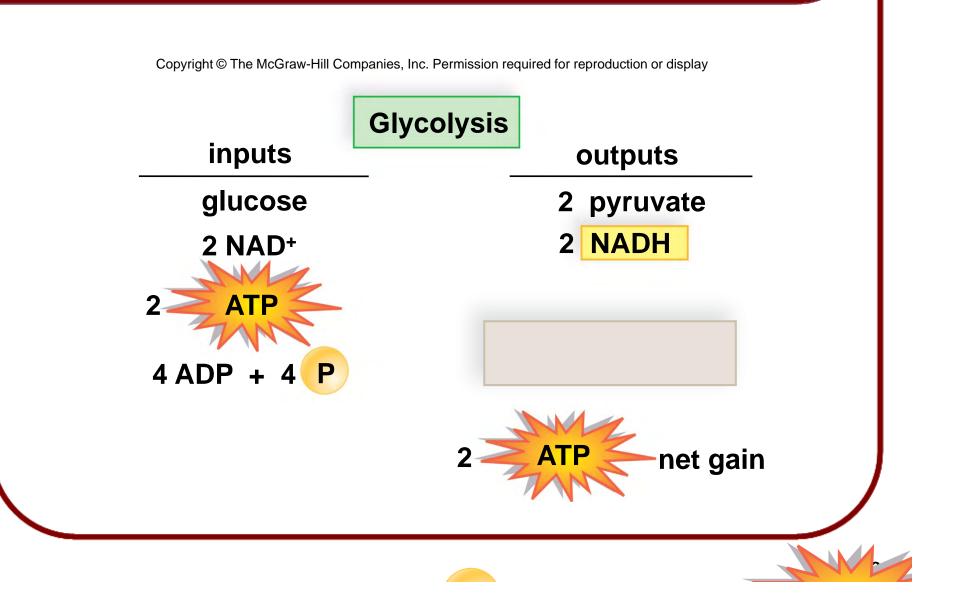
#### Glucose Breakdown: Overview of 4 Phases



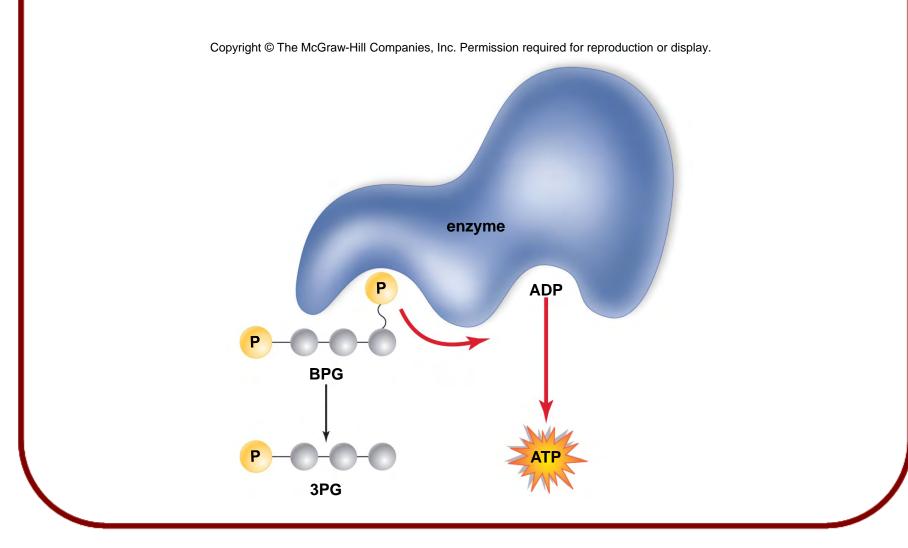
# Glucose Breakdown: Glycolysis

- Occurs in cytoplasm outside mitochondria
- Energy Investment Steps:
  - Two ATP are used to activate glucose
  - Glucose splits into two G3P molecules
- Energy Harvesting Steps:
  - Oxidation of G3P occurs by removal of electrons and hydrogen ions
  - Two electrons and one hydrogen ion are accepted by NAD<sup>+</sup> resulting two NADH
  - Four ATP produced by substrate-level phosphorylation
  - Net gain of two ATP
  - Both G3Ps converted to pyruvates

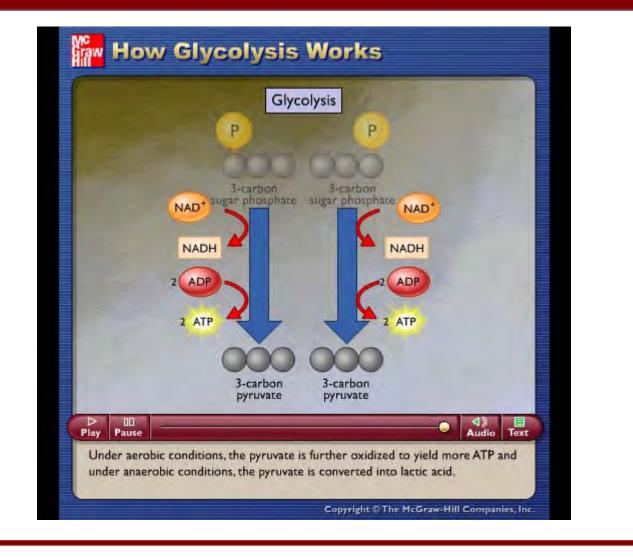
# **Glycolysis: Inputs and Outputs**



# Glycolysis



#### Animation



#### Pyruvate

- Pyruvate is a pivotal metabolite in cellular respiration
- If O<sub>2</sub> is not available to the cell, fermentation, an anaerobic process, occurs in the cytoplasm.
  - During fermentation, glucose is incompletely metabolized to lactate, or to CO<sub>2</sub> and alcohol (depending on the organism).
- If O<sub>2</sub> is available to the cell, pyruvate enters mitochondria by aerobic process.

#### Fermentation

- An anaerobic process that reduces pyruvate to either lactate or alcohol and CO<sub>2</sub>
- NADH passes its electrons to pyruvate
- Alcoholic fermentation, carried out by yeasts, produces carbon dioxide and ethyl alcohol
  - Used in the production of alcoholic spirits and breads.
- Lactic acid fermentation, carried out by certain bacteria and fungi, produces lactic acid (lactate)
  - Used commercially in the production of cheese, yogurt, and sauerkraut.
- Other bacteria produce chemicals anaerobically, including isopropanol, butyric acid, proprionic acid, and acetic acid.

#### Fermentation

- Advantages
  - Provides a quick burst of ATP energy for muscular activity.
- Disadvantages
  - Lactate is toxic to cells.
  - Lactate changes pH and causes muscles to fatigue.
  - Oxygen debt and cramping
- Efficiency of Fermentation
  - Two ATP produced per glucose of molecule during fermentation is equivalent to 14.6 kcal.

# **Products of Fermentation**

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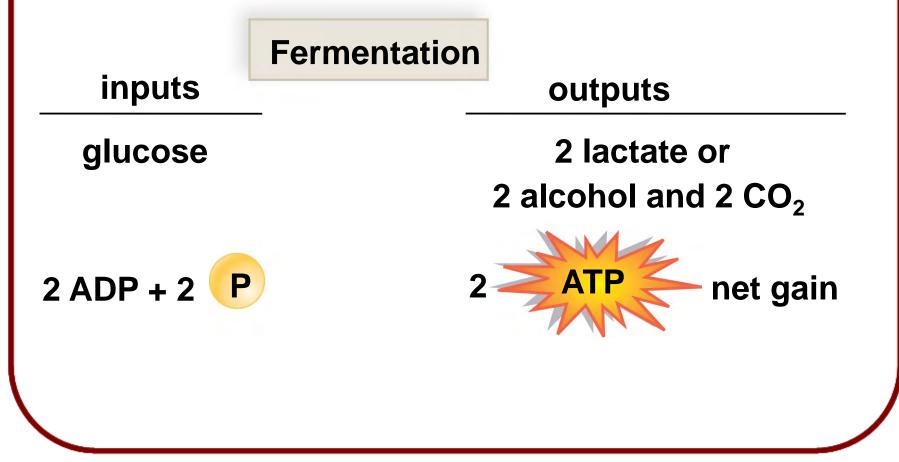
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# **Efficiency of Fermentation**

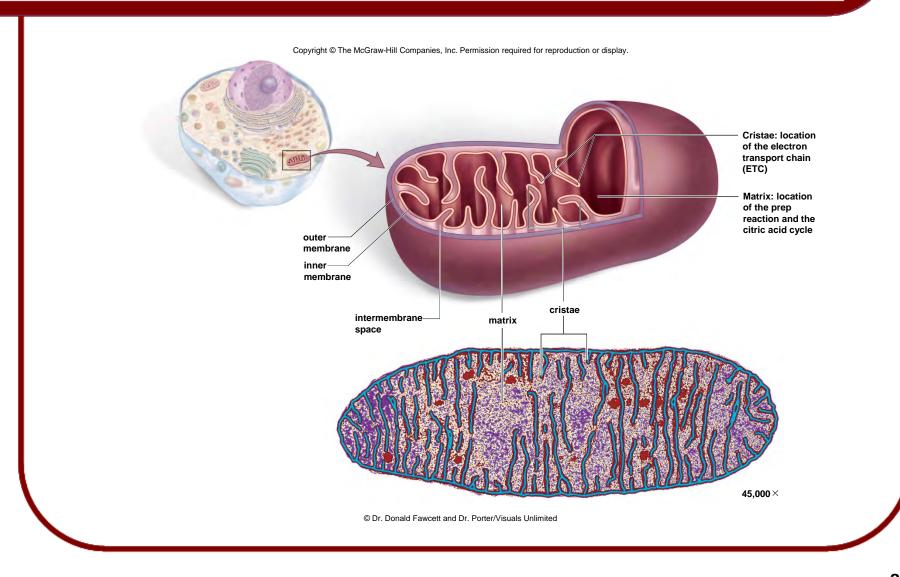
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#### The Preparatory (Prep) Reaction

- Connects glycolysis to the citric acid cycle
- End product of glycolysis, pyruvate, enters the mitochondrial matrix
- Pyruvate converted to 2-carbon acetyl group
  - Attached to Coenzyme A to form acetyl-CoA
  - Electron picked up (as hydrogen atom) by NAD<sup>+</sup>
  - CO<sub>2</sub> released, and transported out of mitochondria into the cytoplasm

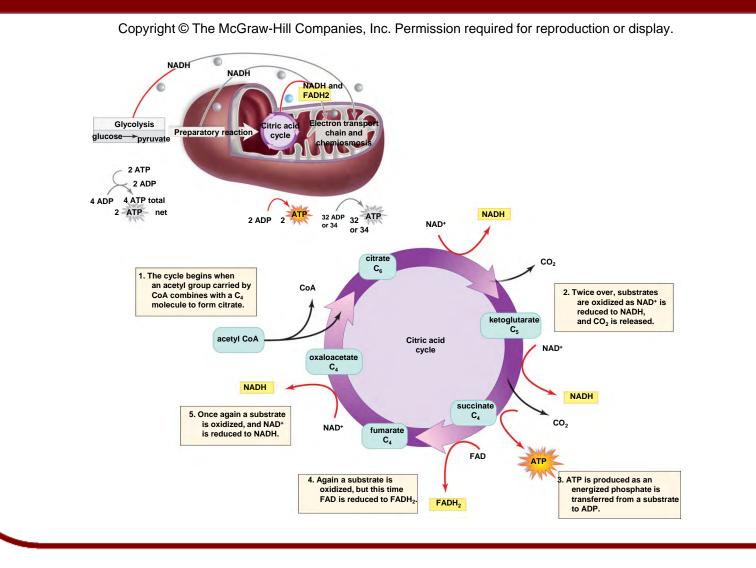
#### Mitochondrion: Structure & Function



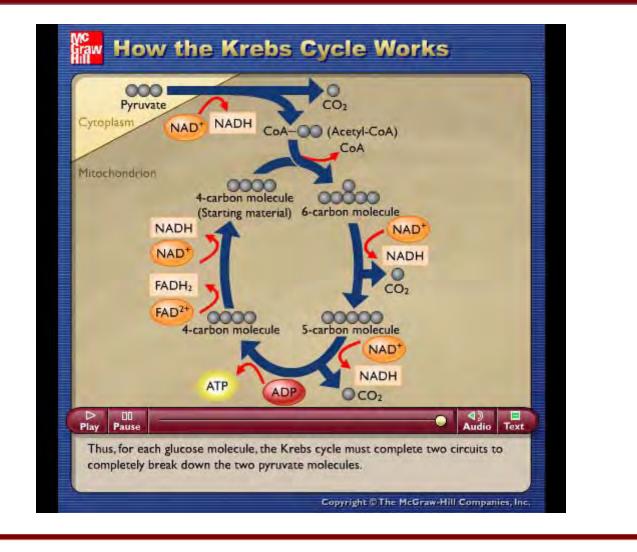
#### Glucose Breakdown: The Citric Acid Cycle

- A.K.A. Krebs cycle
- Occurs in matrix of mitochondria
- Begins by the addition of a two-carbon acetyl group to a four-carbon molecule (oxaloacetate), forming a six-carbon molecule (citric acid)
- NADH, FADH<sub>2</sub> capture energy rich electrons
- ATP formed by substrate-level phosphorylation
- Turns twice for one glucose molecule.
- Produces 4 CO2, 2 ATP, 6 NADH and 2 FADH2 (per glucose molecule)

#### The Citric Acid Cycle

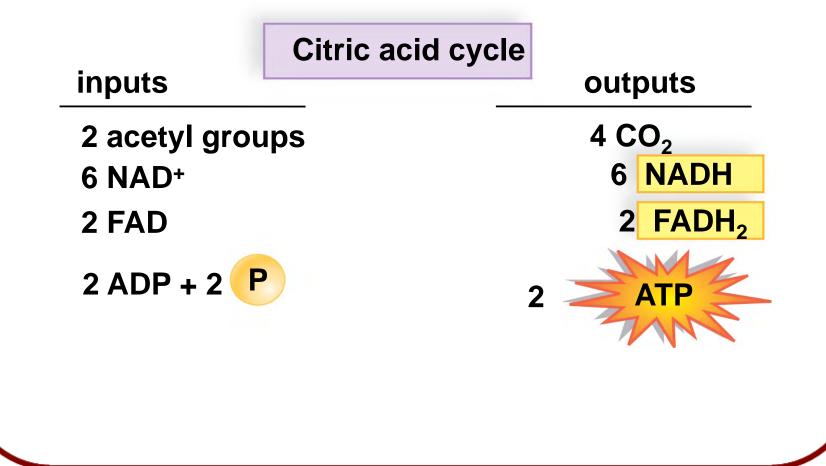


#### Animation



#### Citric Acid Cycle: Balance Sheet

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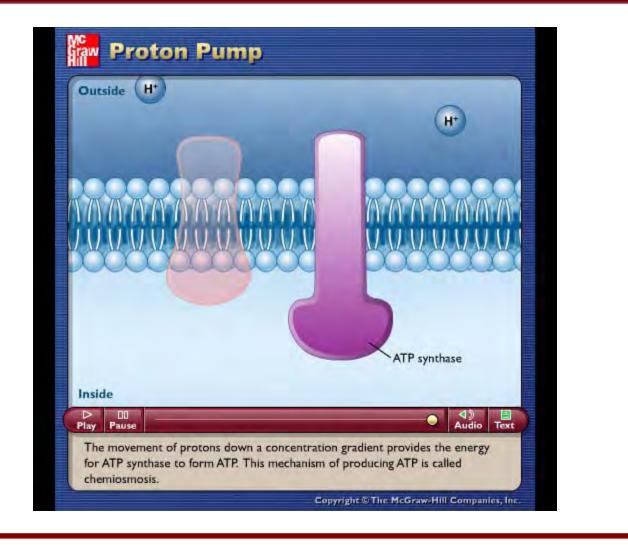


# **Electron Transport Chain**

#### • Location:

- Eukaryotes: cristae of the mitochondria
- Aerobic Prokaryotes: plasma membrane
- Series of carrier molecules:
  - Pass energy rich electrons successively from one to another
  - Complex arrays of protein and cytochromes
    - Cytochromes are respiratory molecules
    - Complex carbon rings with metal atoms in center
- Receives electrons from NADH & FADH<sub>2</sub>
- Produce ATP by oxidative phosphorylation
- Oxygen serves as a final electron acceptor
  - Oxygen ion combines with hydrogen ions to form water

#### Animation



# **Electron Transport Chain**

- The fate of the hydrogens:
- Hydrogens from NADH deliver enough energy to make 3 ATPs
  - Those from FADH<sub>2</sub> have only enough for 2 ATPs
  - "Spent" hydrogens combine with oxygen
- Recycling of coenzymes increases efficiency
  - Once NADH delivers hydrogens, it returns (as NAD<sup>+</sup>) to pick up more hydrogens
  - However, hydrogens must be combined with oxygen to make water
  - If O<sub>2</sub> not present, NADH cannot release H
  - No longer recycled back to NAD<sup>+</sup>

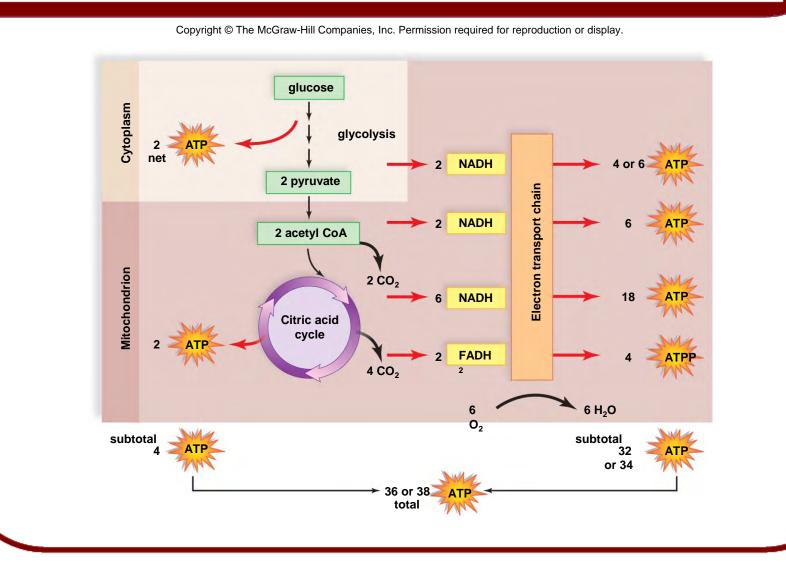
#### Animation



#### Glucose Catabolism: Overall Energy Yield

- Net yield per glucose:
  - From glycolysis 2 ATP
  - From citric acid cycle 2 ATP
  - From electron transport chain 32 ATP
- Energy content:
  - Reactant (glucose) 686 kcal
  - Energy yield (36 ATP) 263 kcal
  - Efficiency 39%; balance is waste heat

#### Overall Energy Yielded per Glucose Molecule



#### Metabolic Pool: Catabolism

- Glucose is broken down in cellular respiration.
- Fat breaks down into glycerol and three fatty acids.
- Amino acids break down into carbon chains and amino groups
  - Deaminated (NH<sub>2</sub> removed) in liver
    - Results in poisonous ammonia (NH<sub>3</sub>)
    - Quickly converted to urea
  - Different R-groups from AAs processed differently
  - Fragments enter respiratory pathways at many different points

#### Metabolic Pool: Anabolism

- All metabolic reactions part of metabolic pool
- Intermediates from respiratory pathways can be used for anabolism
- Anabolism (build-up side of metabolism):
  - Carbs:
    - Start with acetyl-CoA
    - Basically reverses glycolysis (but different pathway)
  - Fats
    - G3P converted to glycerol
    - Acetyls connected in pairs to form fatty acids
    - Note dietary carbohydrate RARELY converted to fat in humans!

#### Metabolic Pool: Anabolism

#### Anabolism (cont.):

- Proteins:
  - Made up of combinations of 20 different amino acids
  - Some amino acids (11) can be synthesized from respiratory intermediates
    - Organic acids in citric acid cycle can make amino acids
    - Add NH<sub>2</sub> transamination
  - However, other amino acids (9) cannot be synthesized by humans
    - Essential amino acids
    - Must be present in diet or die