

# Electron Transport Chain (Respiratory Chain)

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1 Lecture

Respiratory Block

# Objectives

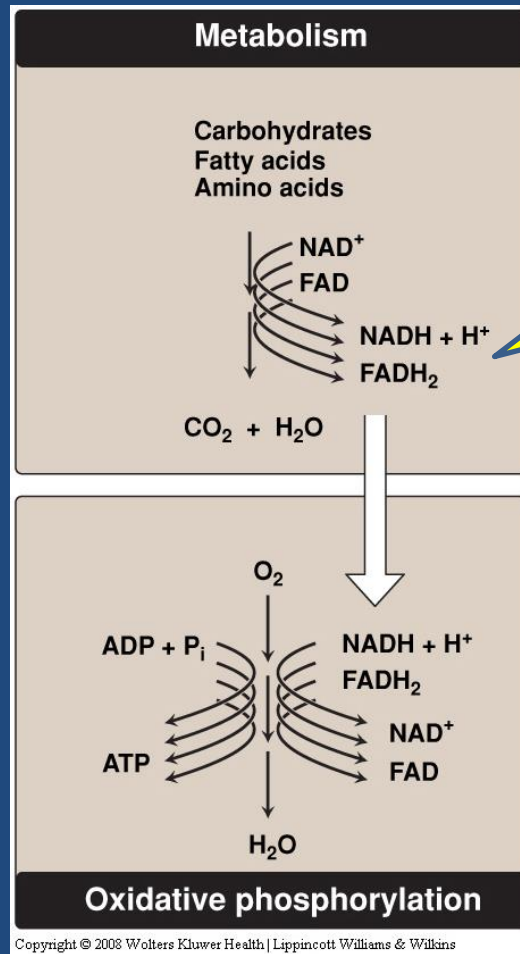
- Understand how energy-rich molecules including glucose are metabolized by a series of oxidation-reduction reactions ultimately yielding  $\text{CO}_2$  and water
- Explain the process of electron transport chain that releases free energy, which is used for ATP synthesis and heat production
- Recognize the reactions of electron transport chain taking place in mitochondria that are coupled to oxidative phosphorylation

# Electron Transport Chain (ETC)

- A system of electron transport that uses respiratory  $O_2$  to finally produce ATP (energy)
- Located in the inner mitochondrial membrane
- Final common pathway of metabolism
- Electrons from food metabolism are transported to  $O_2$
- Uses maximum amount of body's oxygen

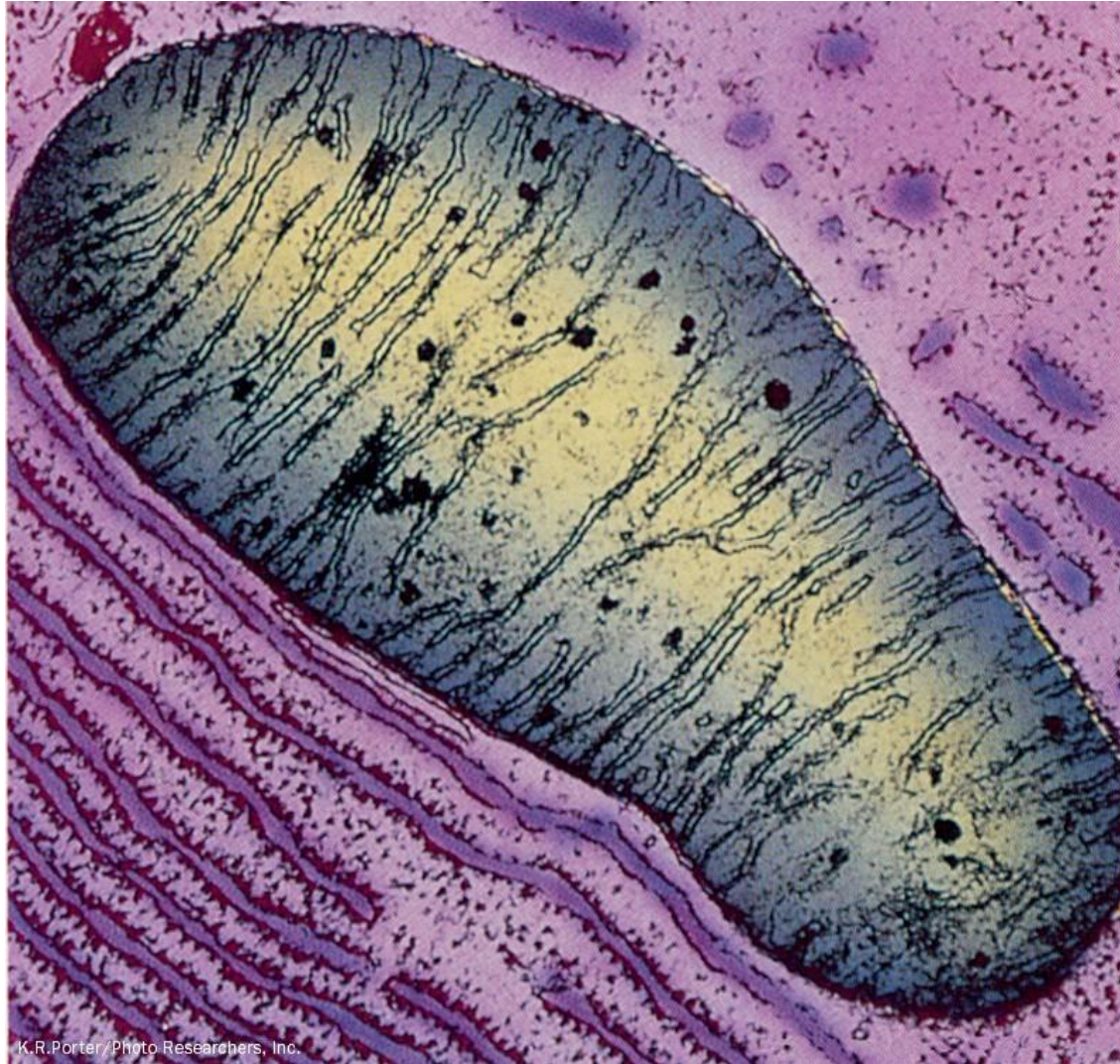
# Metabolic breakdown of energy-yielding molecules

Electrons ( $e^-$ ) lose their free energy

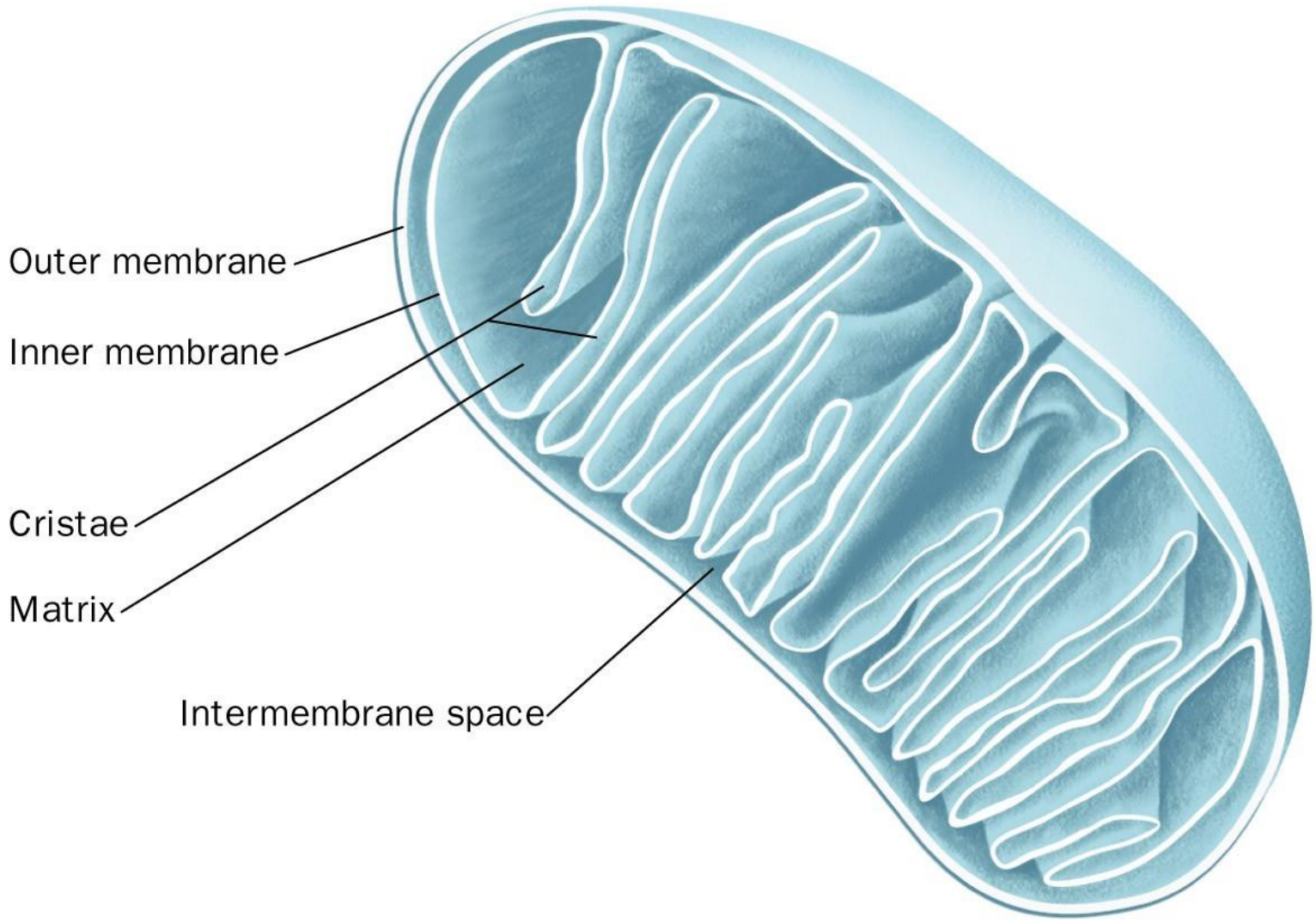


Energy-rich reduced coenzymes

Excess energy generates heat



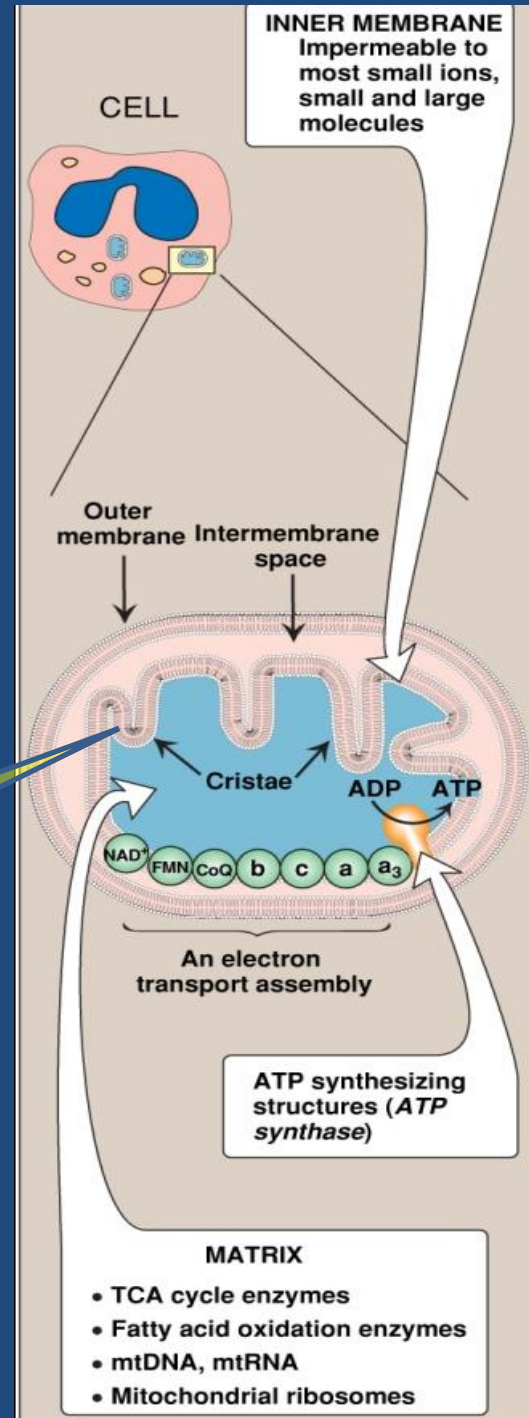
An electron micrograph of an animal mitochondrion



Cutaway diagram of a mitochondrion

# Mitochondrion

Cristae increase the surface area



# Components of ETC

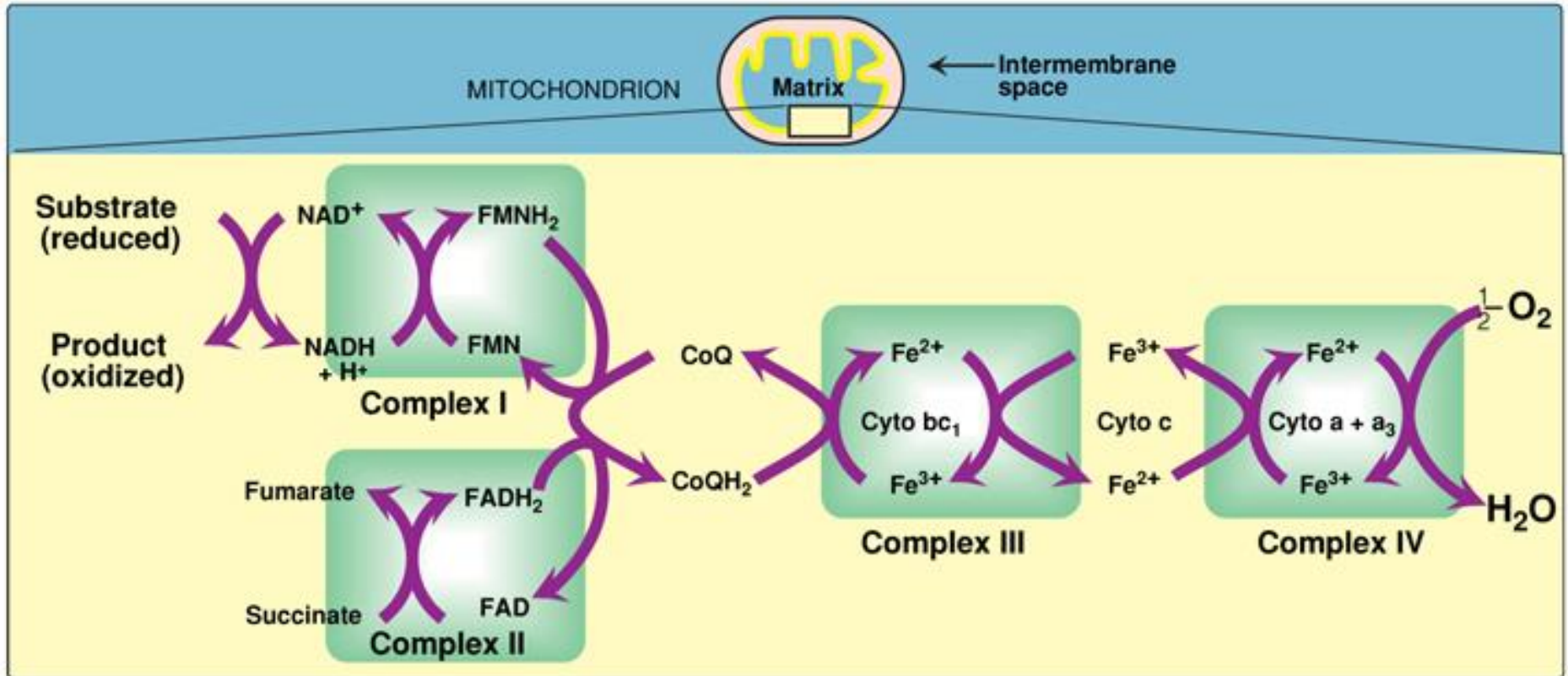
- All members/components are located in the inner mitochondrial membrane (IMM)
- IMM contains 5 complexes:
  - Complex I, II, III, IV (part of ETC)
  - Complex V (ATP synthase: catalyzes ATP synthesis)
  - Mobile electron carriers
    - CoQ
    - Cytochrome c



# Organization of ETC

- Each complex accepts or donates electrons to mobile carriers
- Carriers accept electrons from donors and then donate to the next carrier in chain
- Electrons finally combine with oxygen and protons to form water
- Oxygen is required as a final acceptor (respiratory chain)

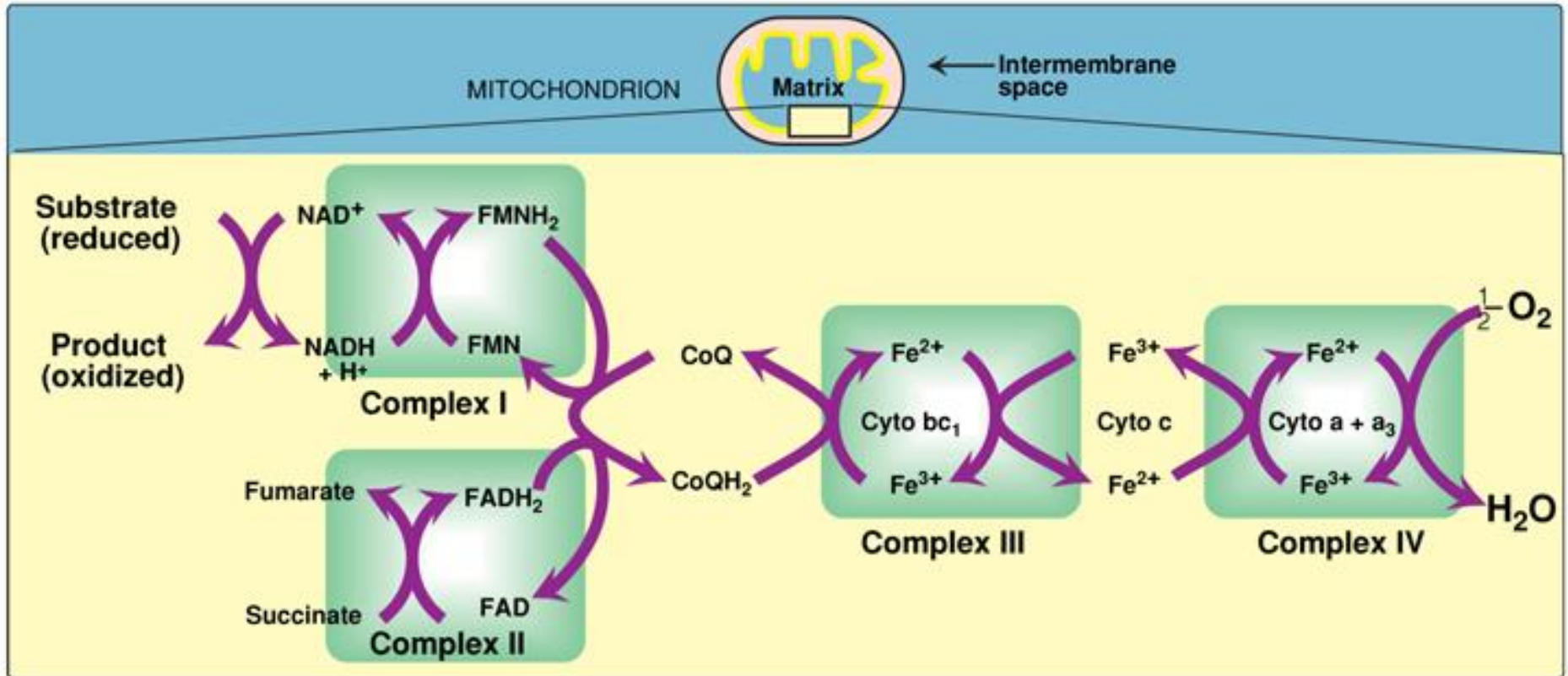
# Electron Transport Chain



# Complex I – NADH Dehydrogenase

- This complex collects the pair of electrons from NADH and passes them to CoQ

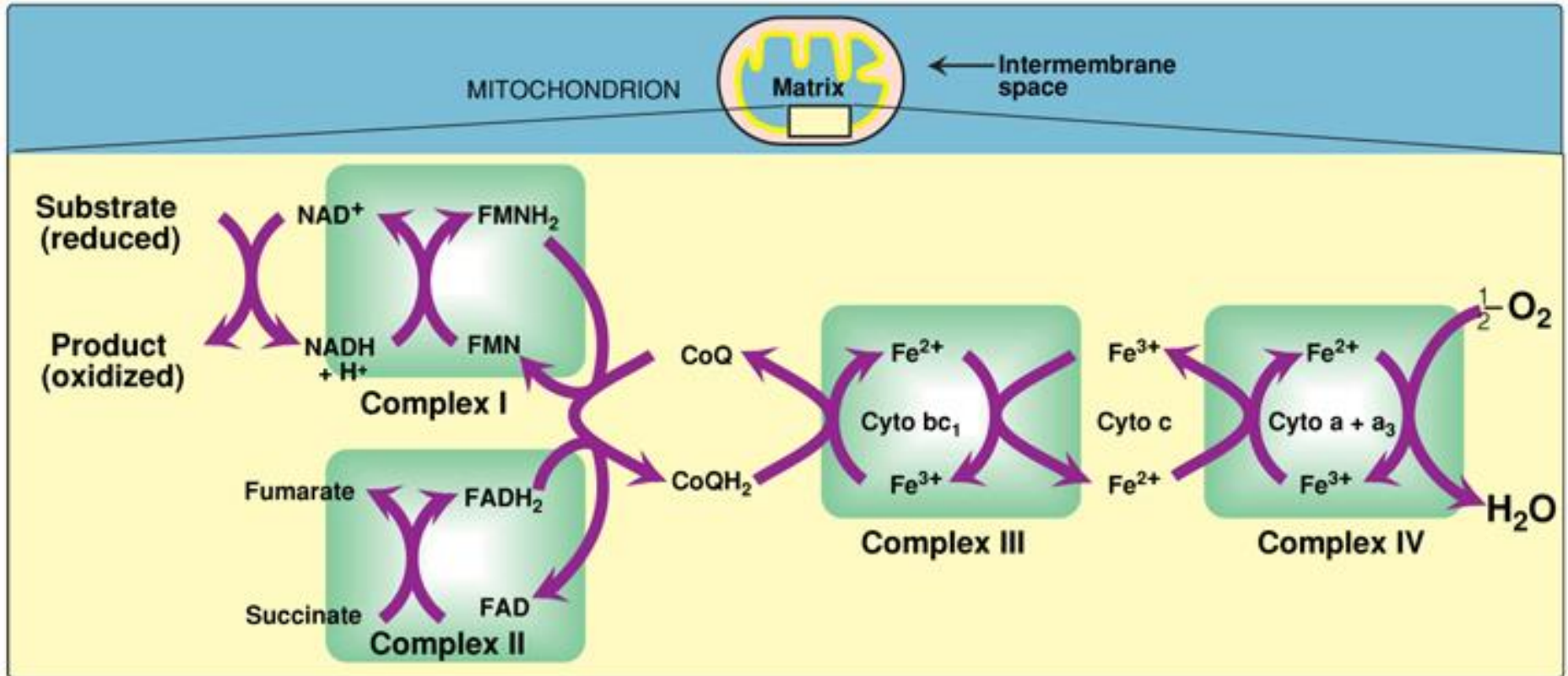
# Electron Transport Chain



# Complex II – Succinate dehydrogenase

- It is also a part of the Krebs cycle
- Transfers electrons to CoQ

# Electron Transport Chain



# Coenzyme Q (CoQ)

- Also called ubiquinone (ubiquitous in biological systems)
- A non-protein member of the ETC
- Lipid soluble and mobile

# Cytochromes

- Each cytochrome is a protein that contains
  - Heme group (porphyrin ring + iron in  $\text{Fe}^{3+}$  state)
- When cytochromes accept electron
  - $\text{Fe}^{3+}$  (ferric) is converted to  $\text{Fe}^{2+}$  (ferrous)
  - $\text{Fe}^{2+}$  is reoxidized to  $\text{Fe}^{3+}$  when it donates electrons to the next carrier



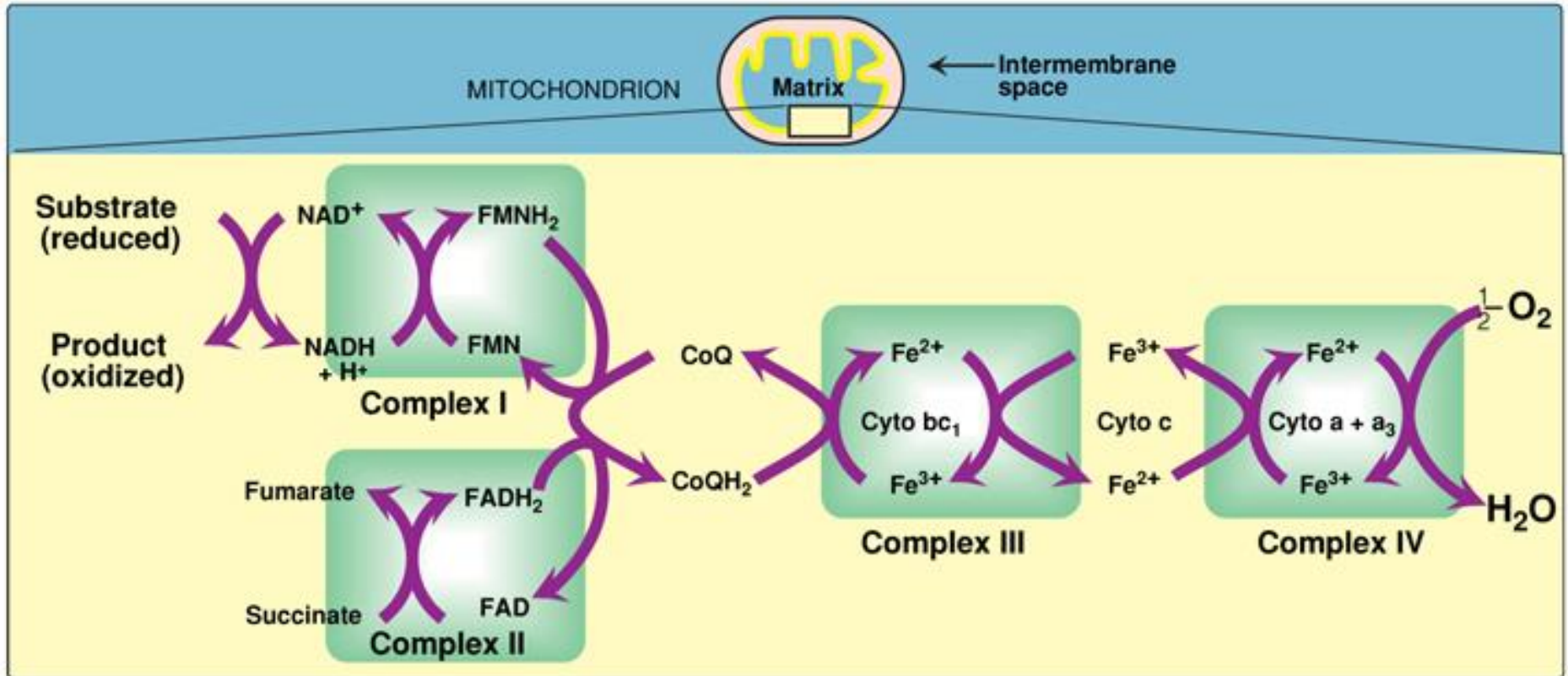
# Complex III and IV

- Complex III: Cytochrome bc<sub>1</sub>
- Complex IV: Cytochrome a + a<sub>3</sub>

Electrons flow from:

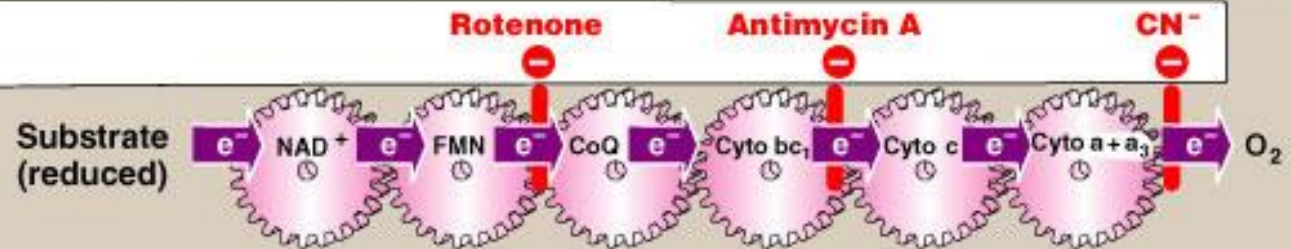
- CoQ → Complex III → Cyt. c → Complex IV

# Electron Transport Chain



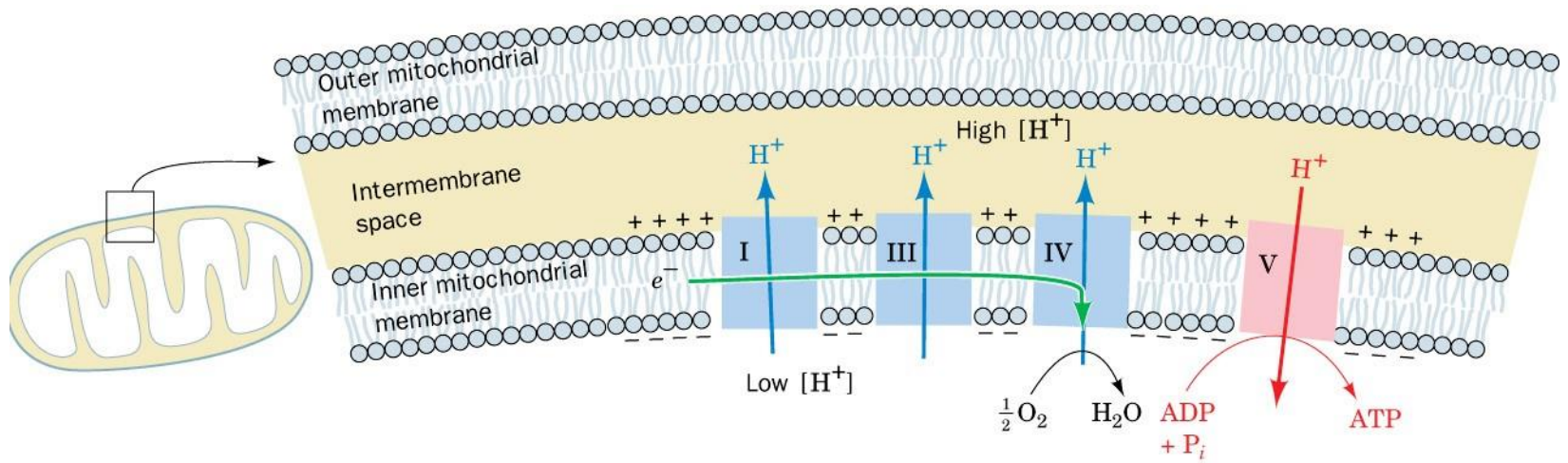
# Site-specific inhibitors of ETC

Blocking electron transfer by any one of these inhibitors stops electron flow from substrate to oxygen because the reactions of the electron transport chain are tightly coupled like meshed gears.



# ETC is coupled to proton transport for ATP synthesis

- The energy of electron transfer is used to drive the protons out of the matrix
- It is done by complexes I, III and IV (proton pumps)
- This creates a proton gradient across the IMM to synthesize ATP

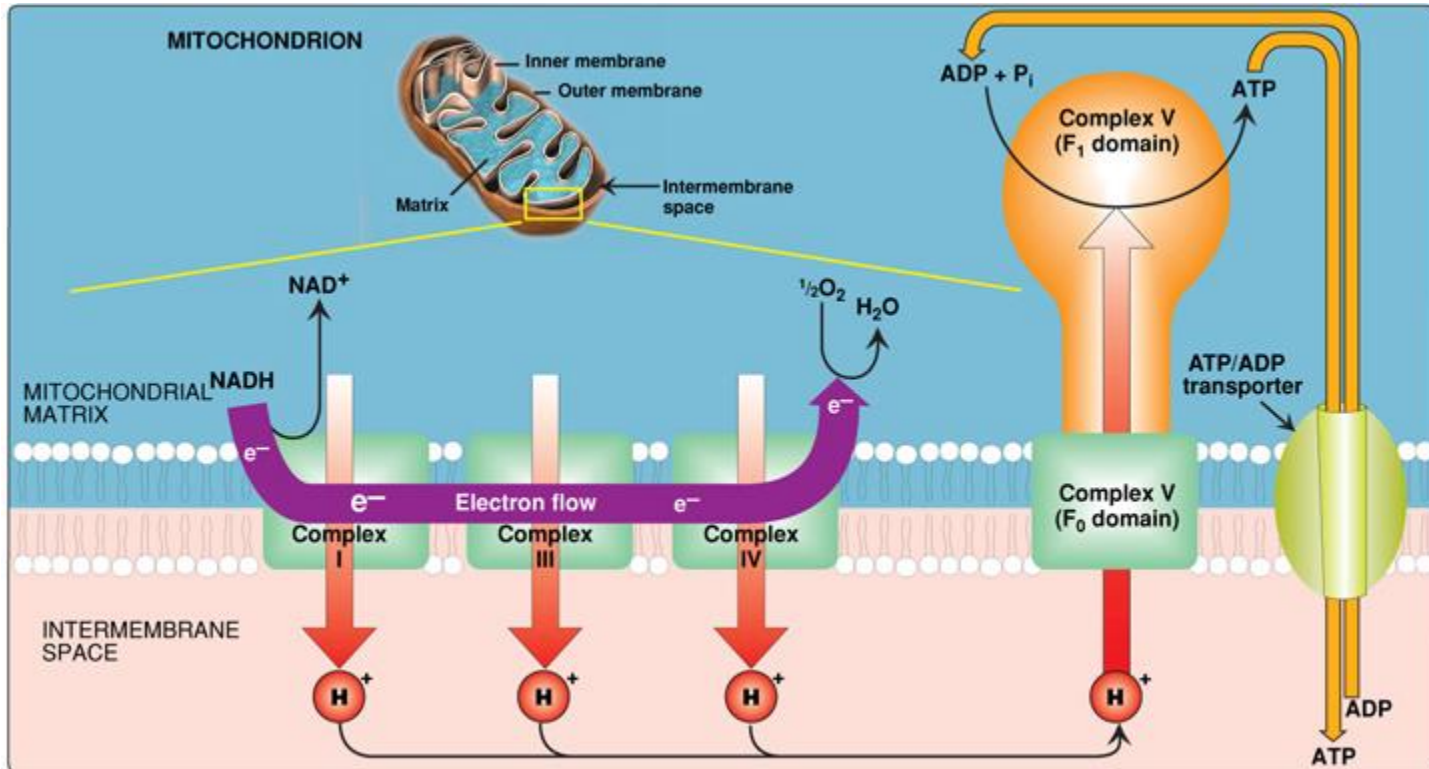


Coupling of electron transport (*green arrow*) and ATP synthesis

# ATP synthase

- ATP synthase (Complex V) synthesizes ATP
- Consists of two domains:
  - $F_0$  – membrane spanning domain
  - $F_1$  – extramembranous domain

# Transport of protons



# Energetics of ATP synthesis

- The energy required for phosphorylation of ADP to ATP = 7.3kcal/mol
- Energy produced from the transport of a pair of electrons from NADH to O<sub>2</sub> = 52.58 kcal
- No. of ATP molecules produced is 3 (NADH to O<sub>2</sub>)
- Excess energy is used for other reactions or released as heat



# P:O ratio

- ATP made per O atom reduced
  - For NADH
    - P:O = 3:1
  - For FADH<sub>2</sub>
    - P:O = 2:1

# Inhibitors of ATP synthesis

- Oligomycin:
  - Binds to  $F_0$  domain of ATP synthase and closes the  $H^+$  channel
- Uncoupling proteins (UCPs):
  - Create proton leaks (allow protons to reenter the matrix without ATP synthesis)
  - Energy is released as heat (nonshivering thermogenesis)

# Take home message

- ETC is a common pathway of transferring energy-rich electrons from metabolism finally yielding CO<sub>2</sub> and water
- The energy of the electrons transferred is used for ATP synthesis and heat production

# References

- Lippincott's Illustrated Reviews, Biochemistry, 6th Edition, Denise R. Ferrier, Lippincott Williams & Wilkins, USA, pp. 73-79.

Uncoupling proteins create a "proton leak," allowing protons to reenter the mitochondrial matrix without capturing any energy as ATP.

