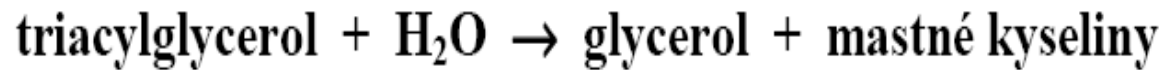


# Metabolismus lipidů

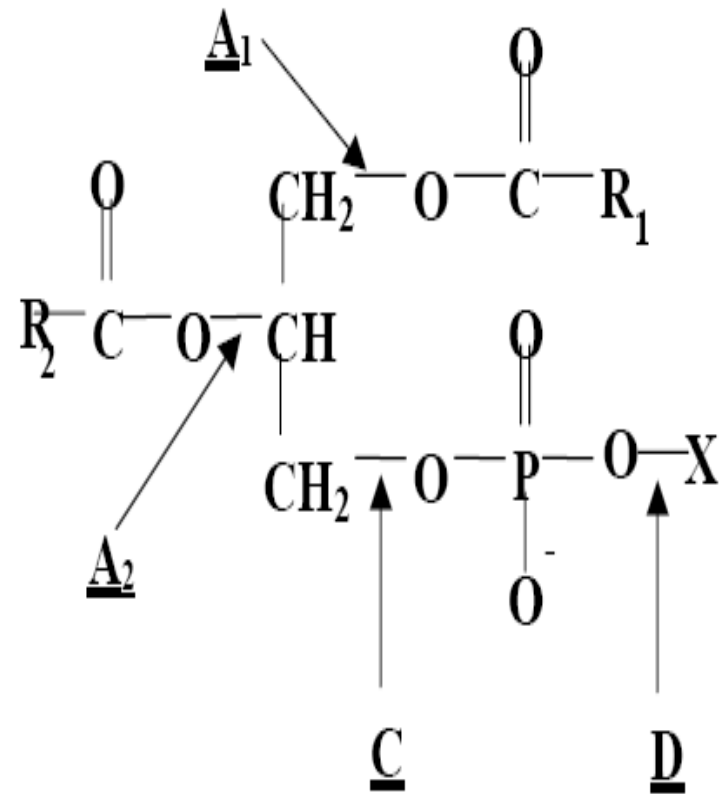
- Triacylglyceroly – 90 % potravních lipidů a zároveň hlavní zásobárnou energie – orgánový tuk
- Dvojnásobné množství energie  
CH<sub>2</sub> - CHOH
- Jako hydrofobní látky nejsou hydratovány
- Nejsou pohotovým zdrojem energie, spíše vhodné pro skladování.

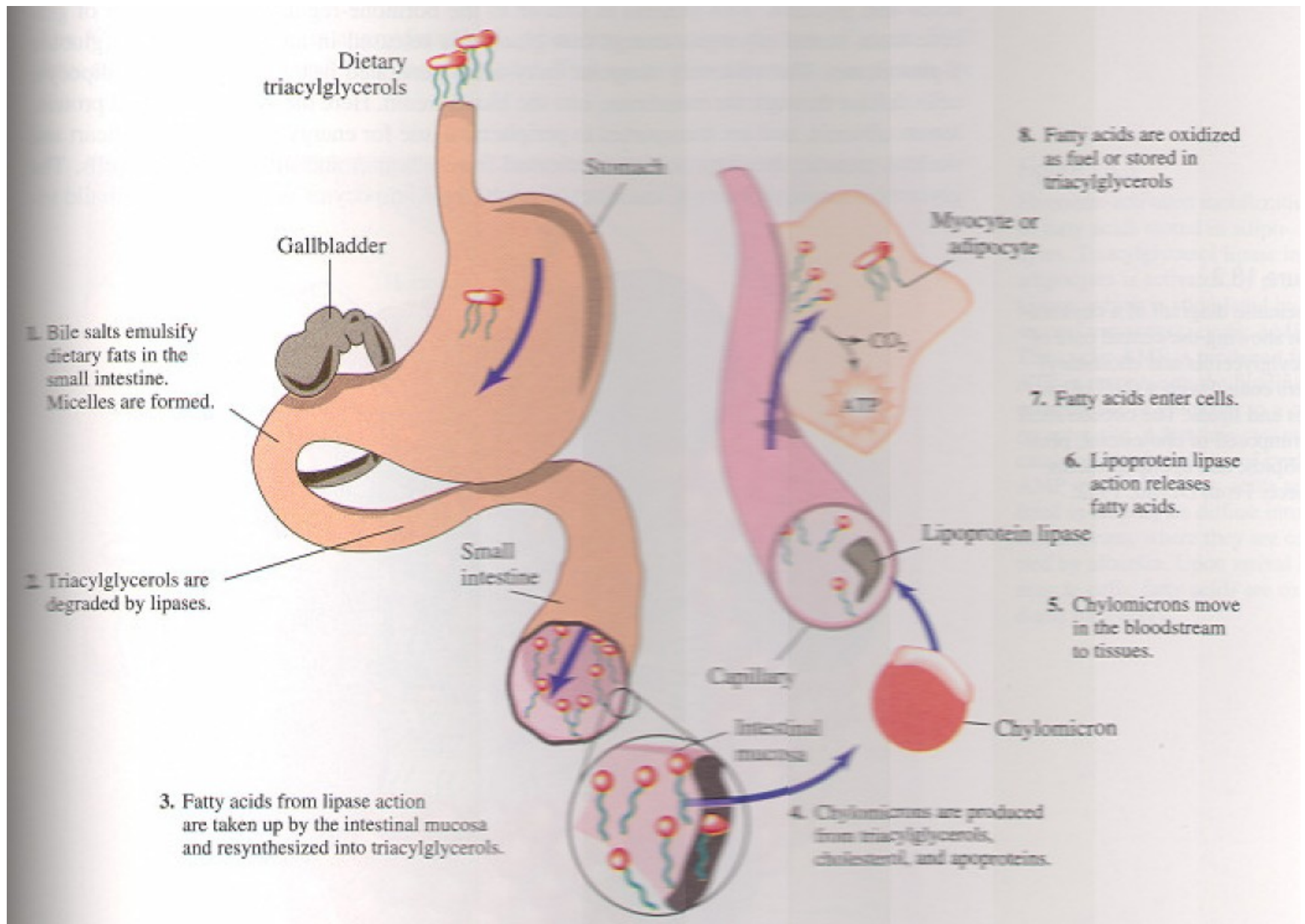
# METABOLISMUS LIPIDŮ

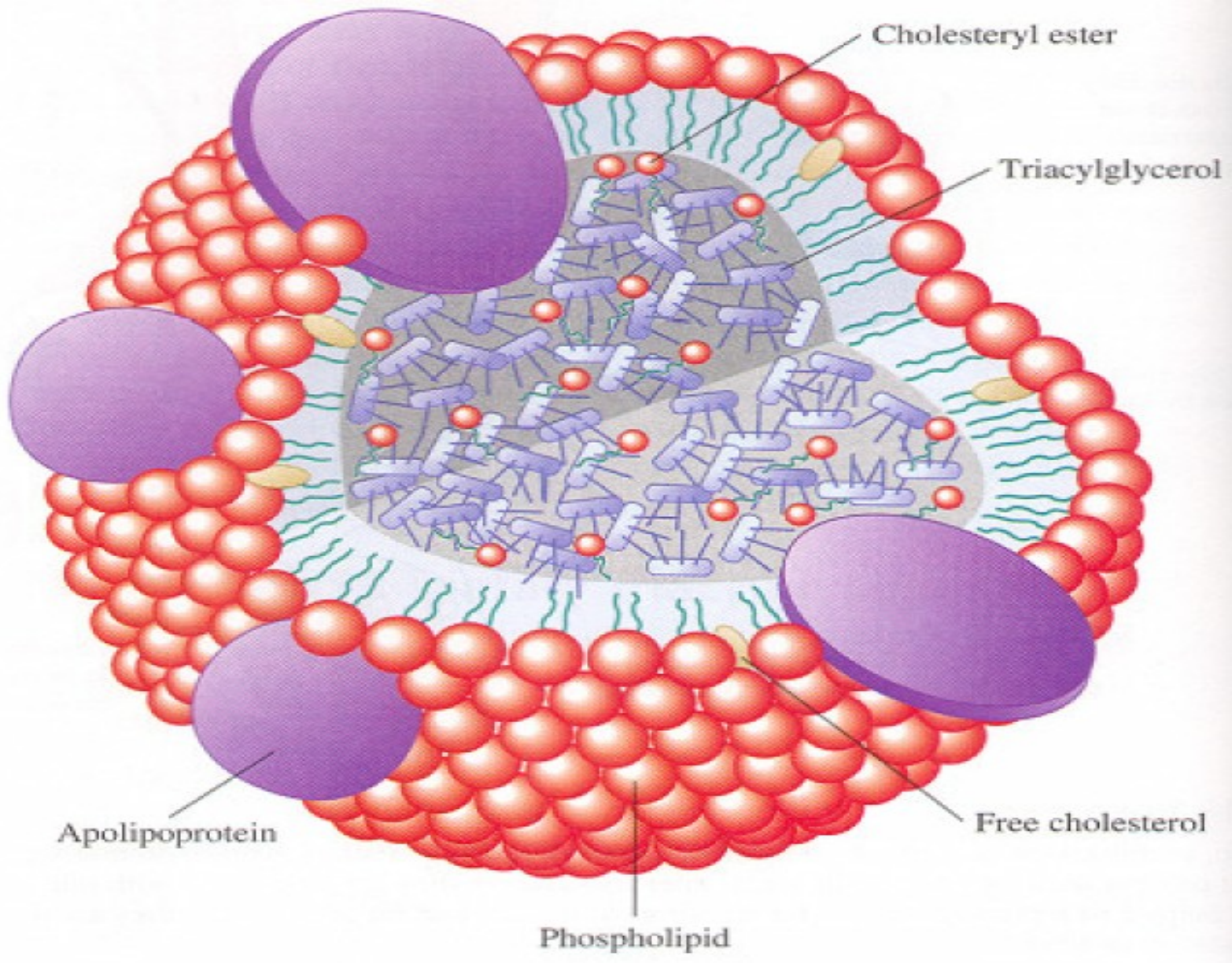
Lipasy - hydrolasy - karboxylesterasy



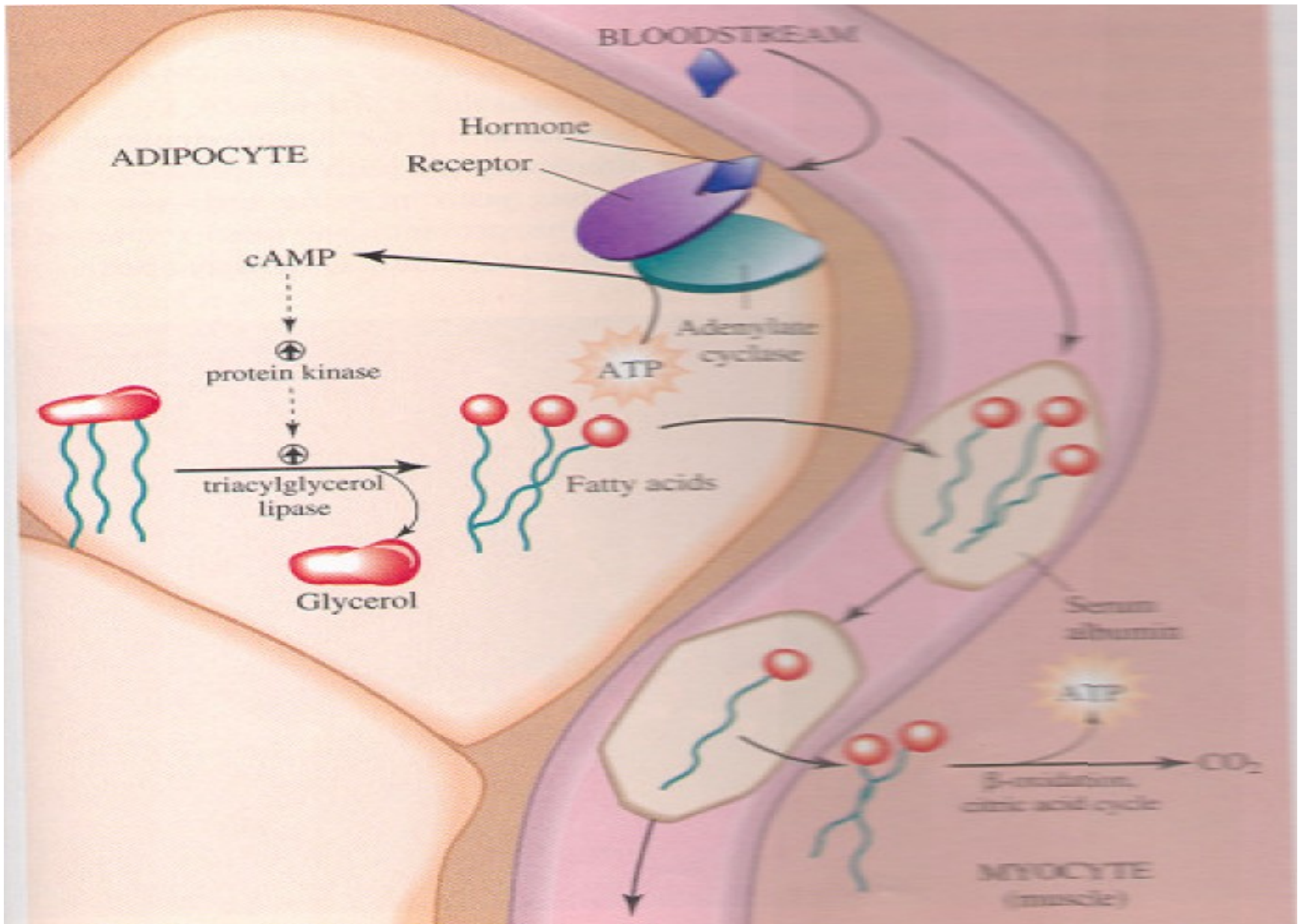
## Fosfolipasy









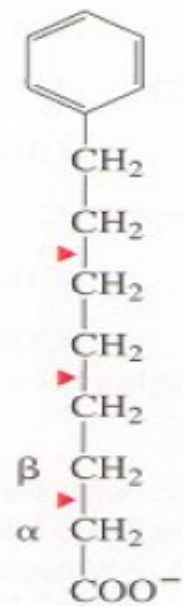


# Odbourávání mastných kyselin

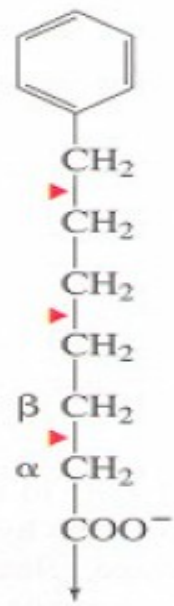
## $\beta$ oxidace

F.KNOOP 1909

F.LYNEN 1951



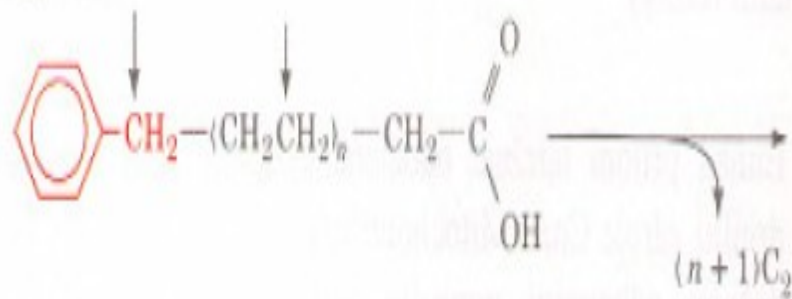
Phenylacetate  
(from even-numbered  
carbon chain)



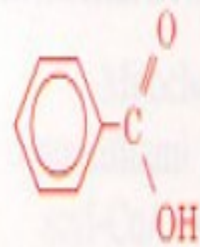
Benzoate  
(from odd-numbered  
carbon chain)



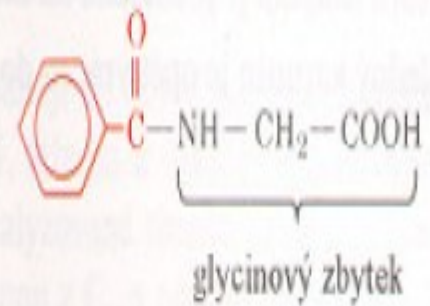
přidaná mastná  
kyselina



degradační produkty



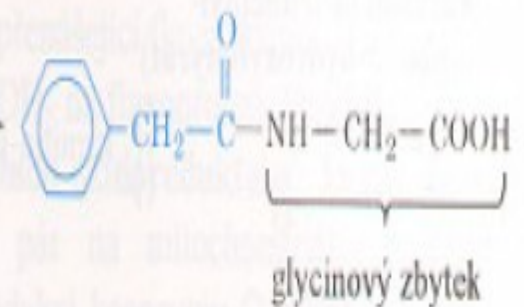
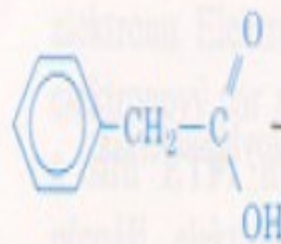
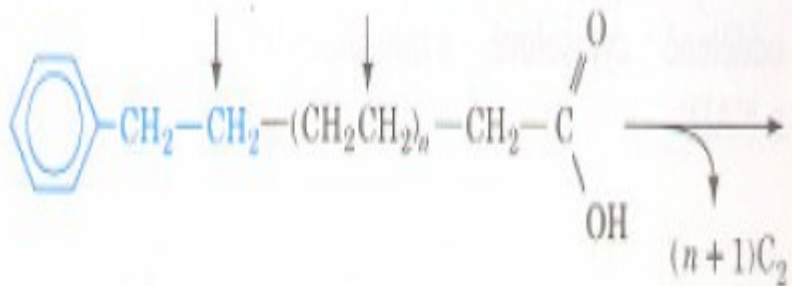
exkreční  
produkt



kyselina s lichým  
počtem C-atomů

benzoová kyselina

hippurová kyselina



kyselina se sudým  
počtem C-atomů

fenylactová kyselina

fenylaceturová kyselina

# Odbourávání mastných kyselin

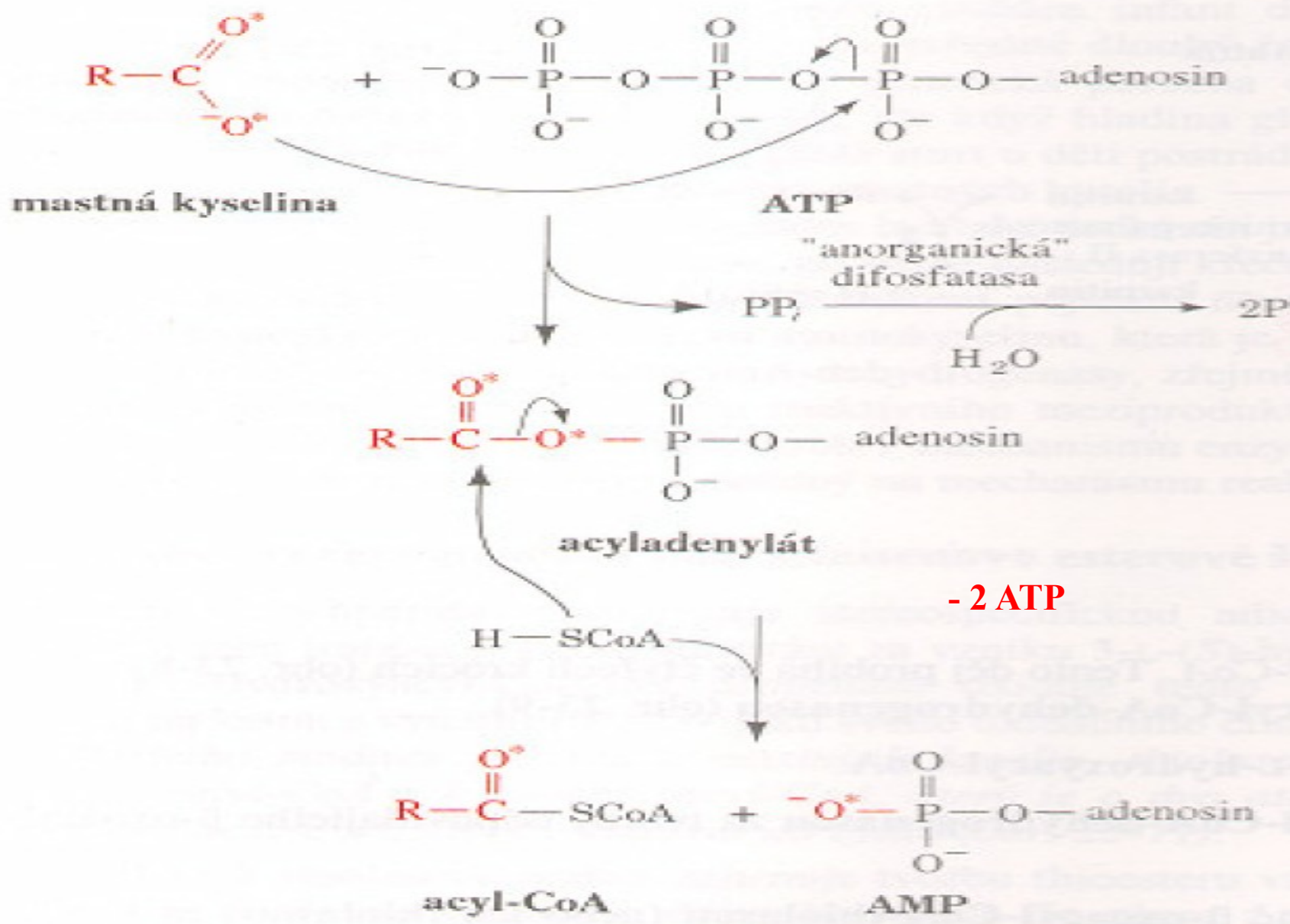
## $\beta$ oxidace

F.KNOOP 1909

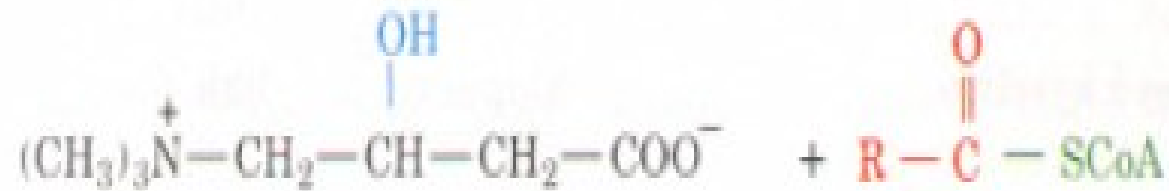
F.LYNEN 1951

### *A. Aktivace mastných kyselin*



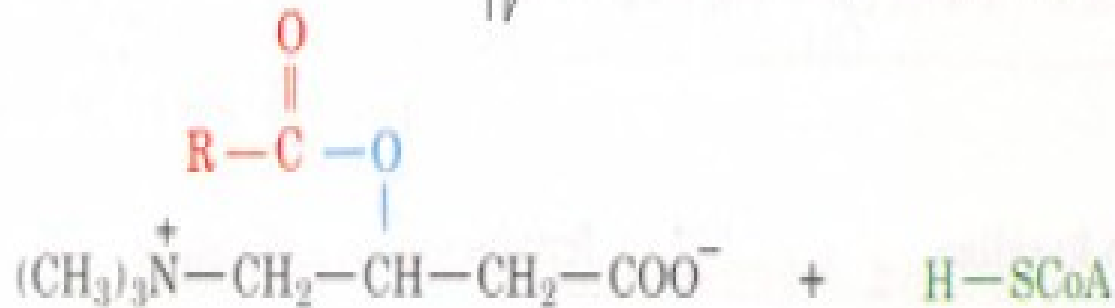


**B. Transport RCOSCoA - karnitinový člunek**



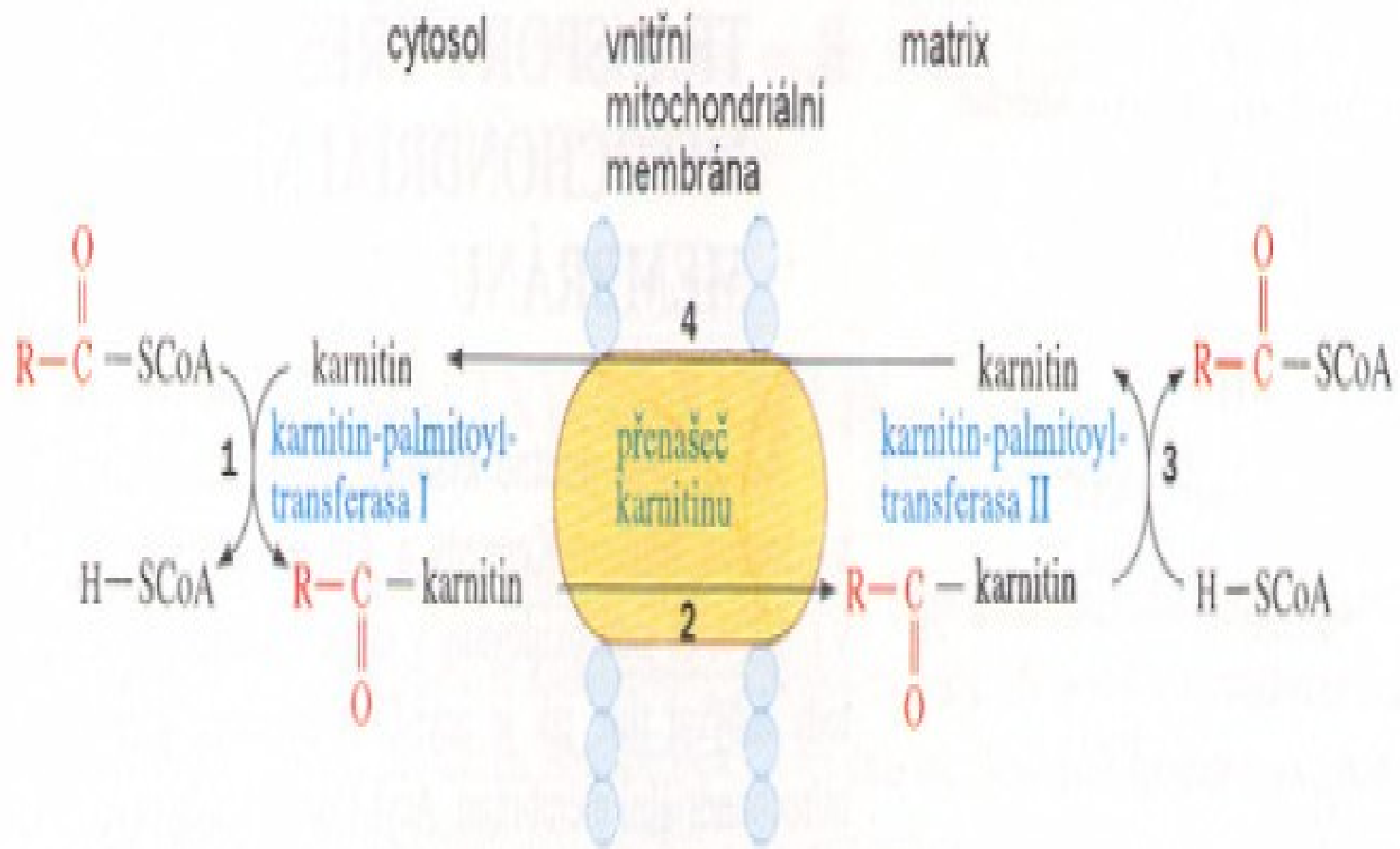
karnitin (4-trimethyl-  
amino-3-hydroxybutyrát)

⇌ karnitin-palmitoyltransferasa

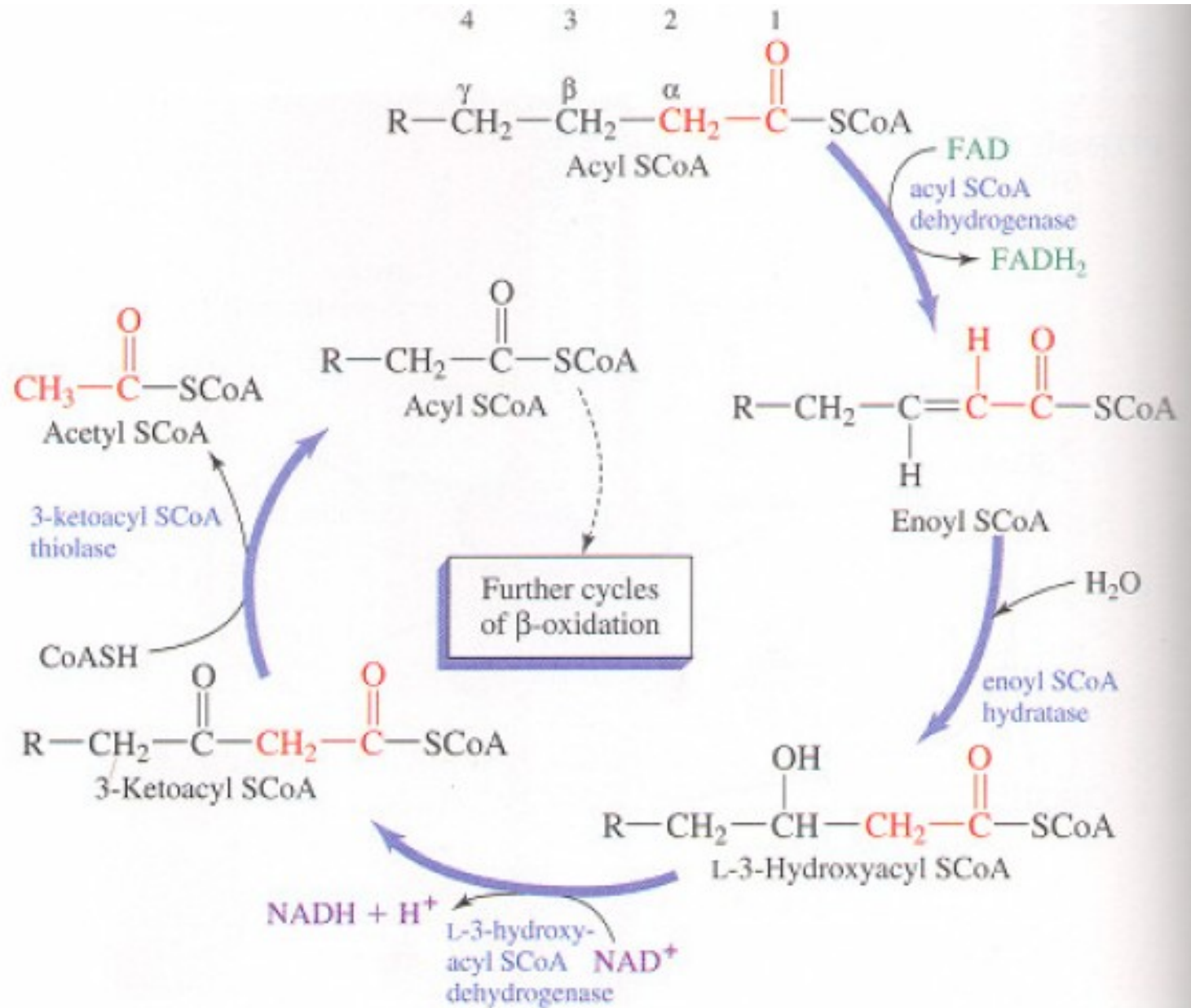


acylkarnitin

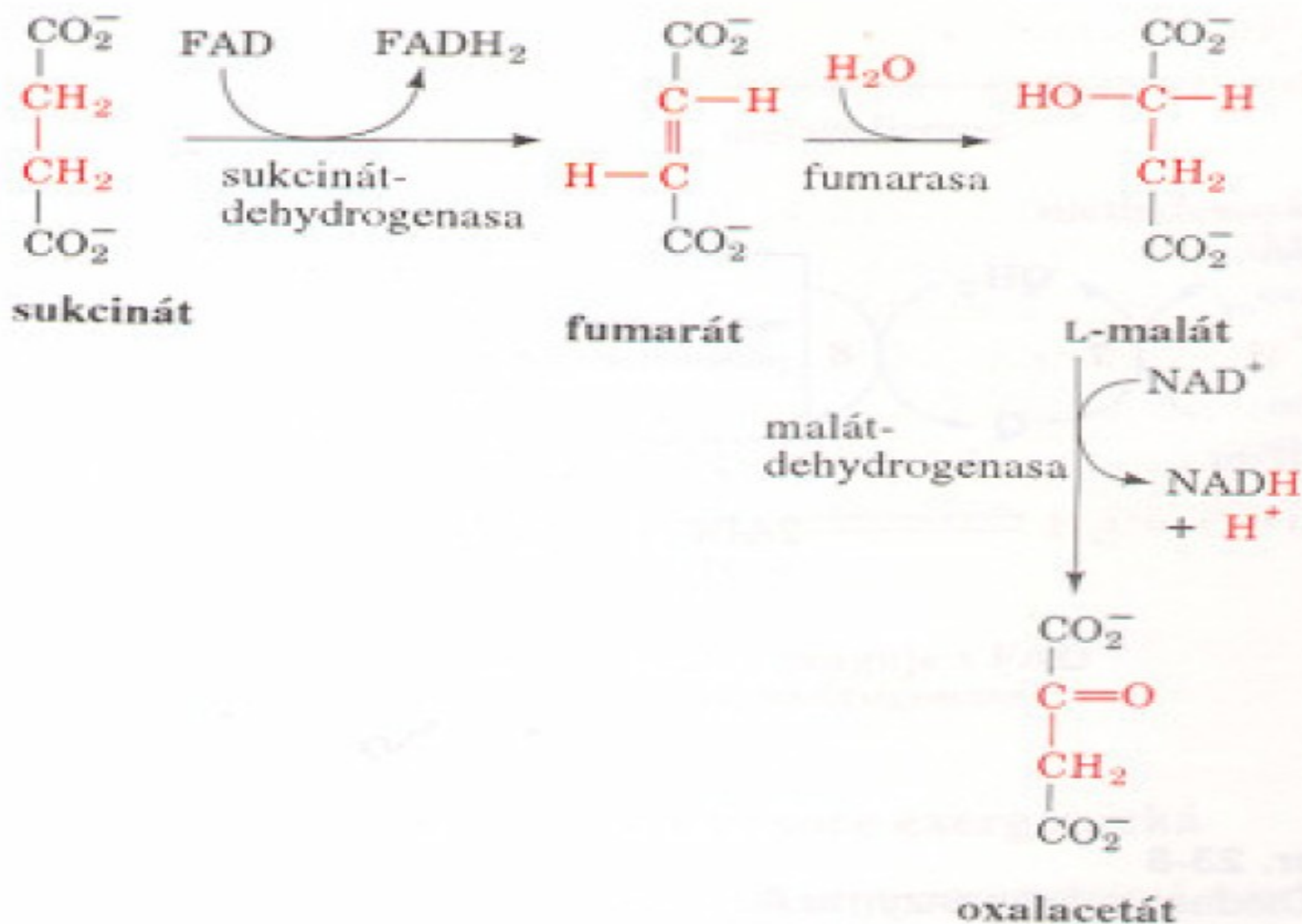
## B. Transport $RCOSCoA$ - karnitinový člunek

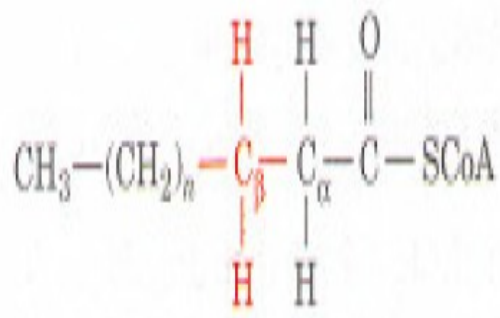


# C. $\beta$ oxidace

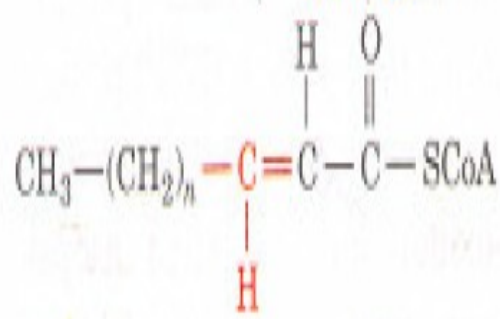
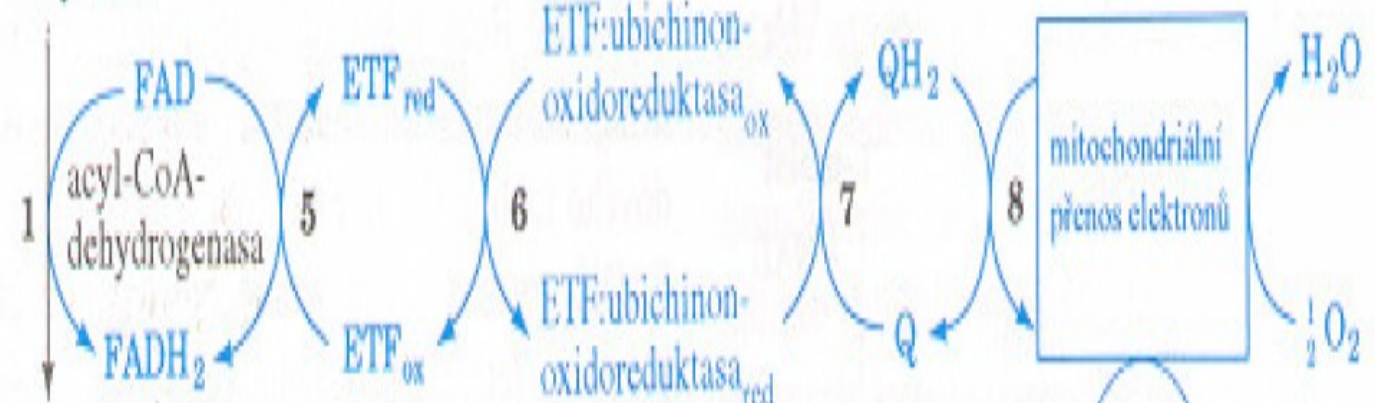








acyl-CoA

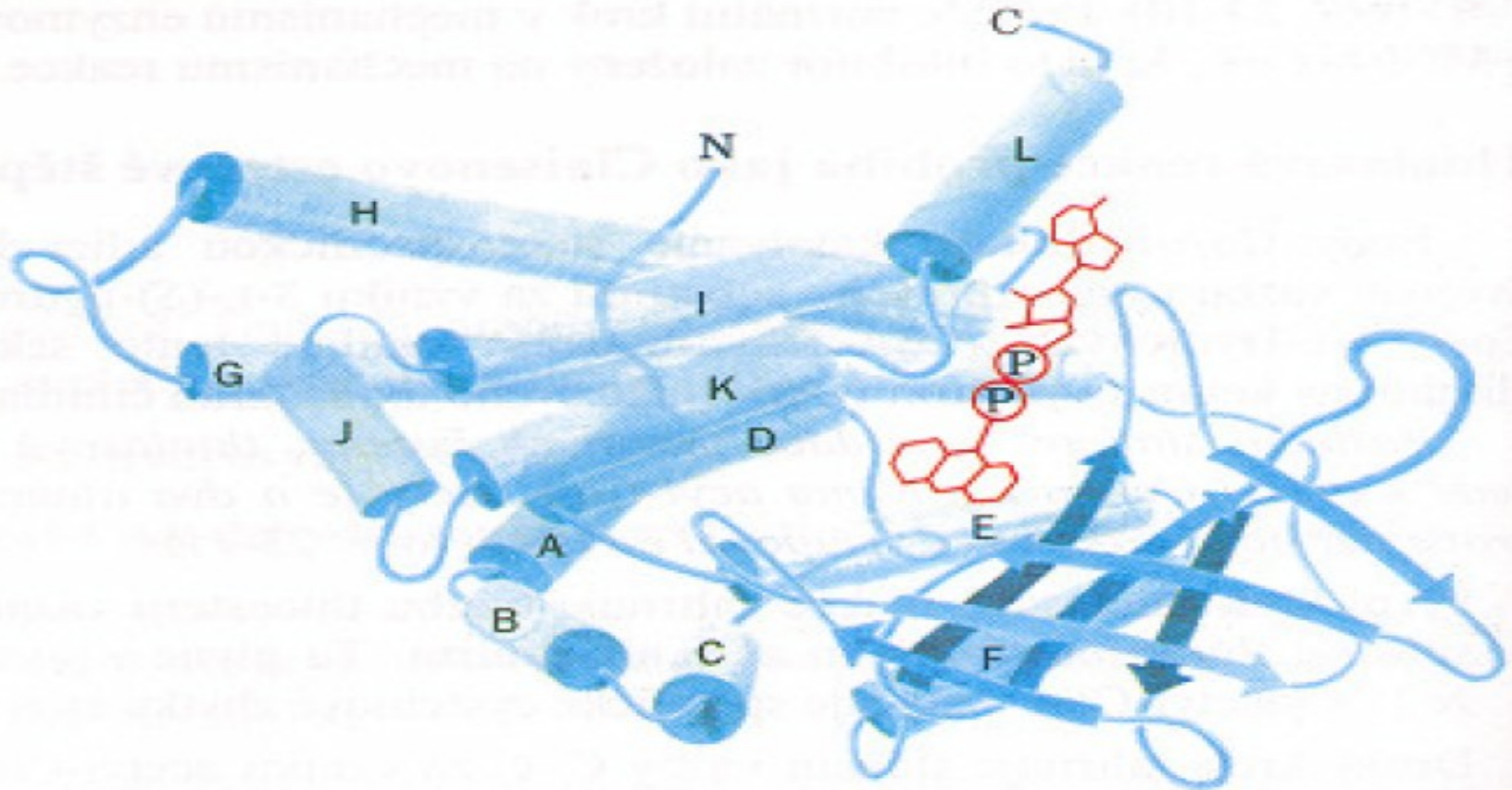


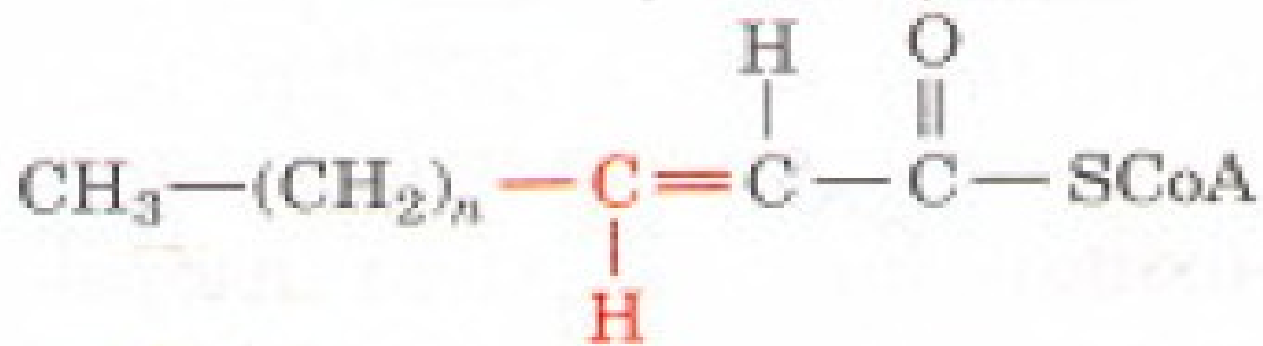
2ADP + 2P<sub>i</sub> → 2ATP

# Acyl-CoA DH

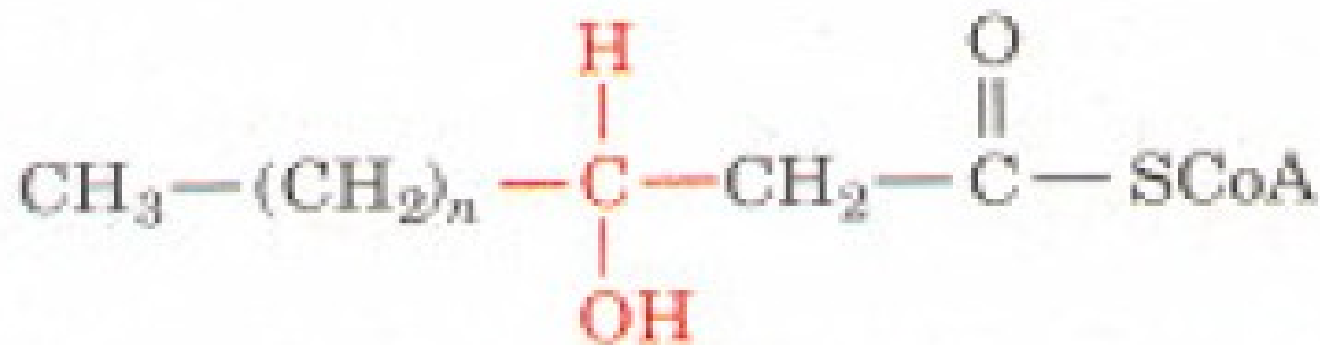
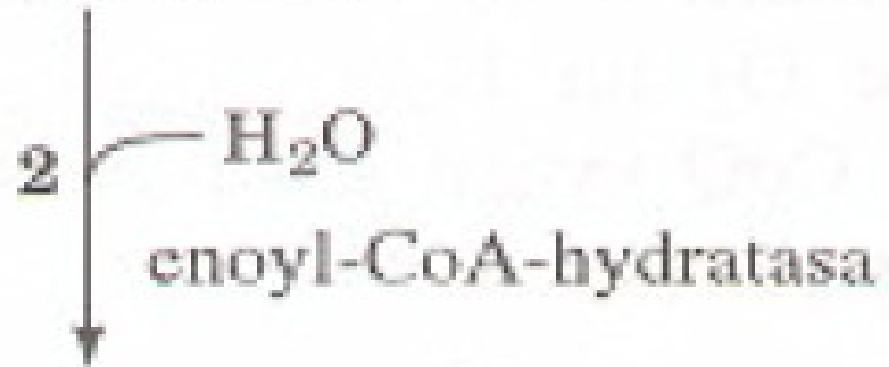
nedostatek

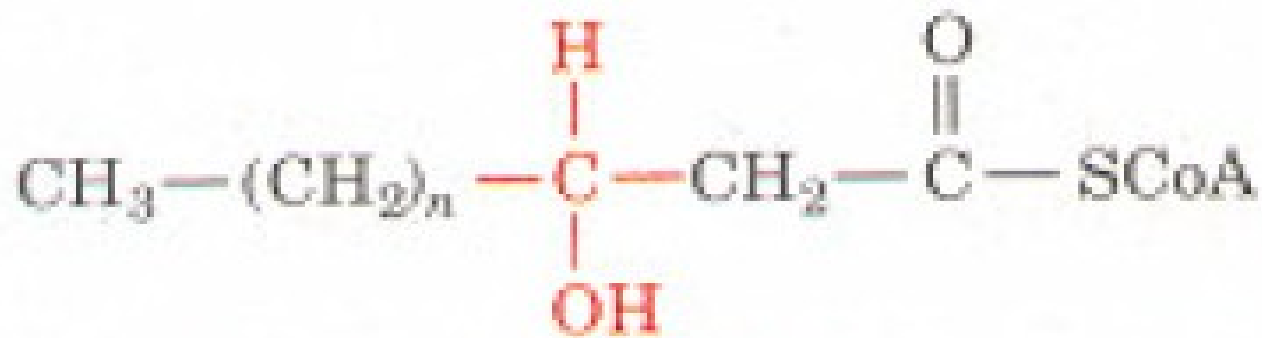
**syndrom náhlého dětského úmrtí (sudden infant death syndrome, SIDS)**



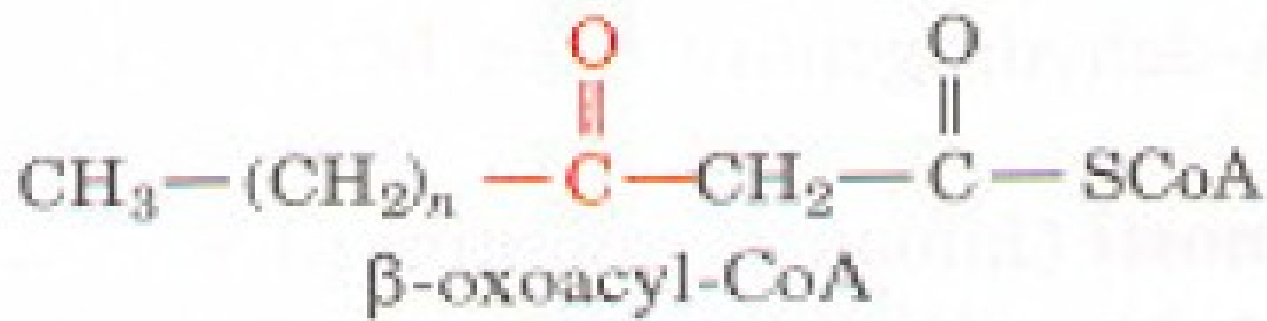
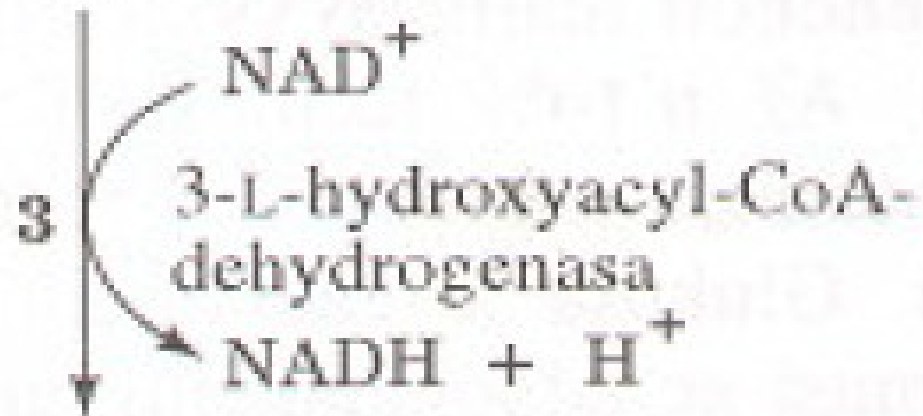


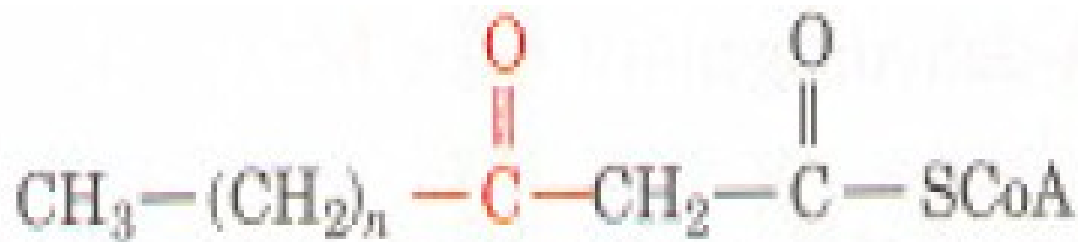
*trans*- $\Delta^2$ -enoyl-CoA





3-L-hydroxyacyl-CoA



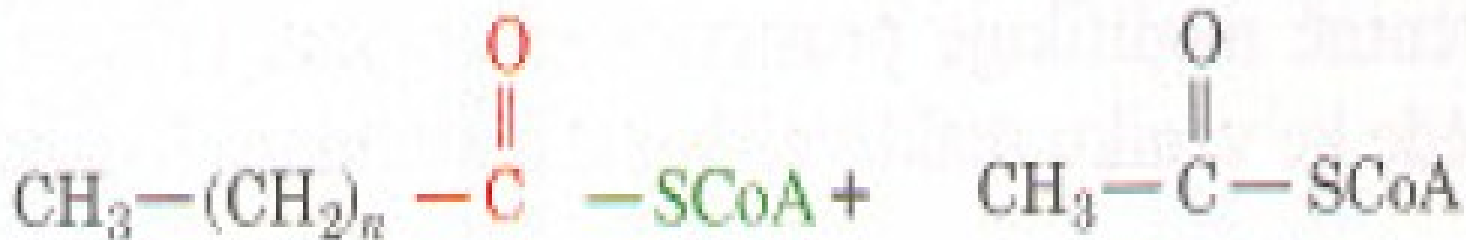


$\beta$ -oxoacyl-CoA



CoASH

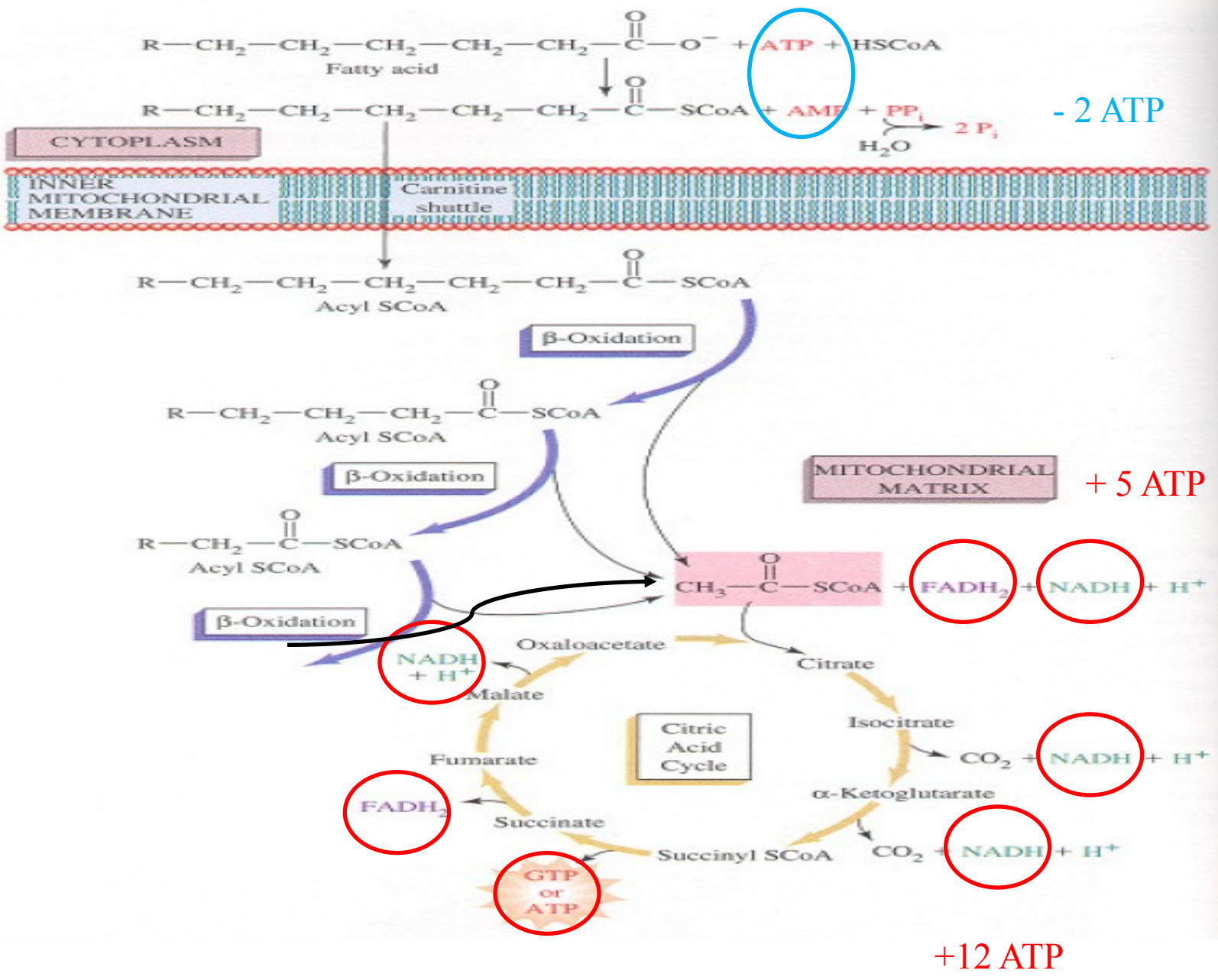
$\beta$ -oxoacyl-CoA-thiolasa



**acyl-CoA**  
(kratší o 2C)

**acetyl-CoA**



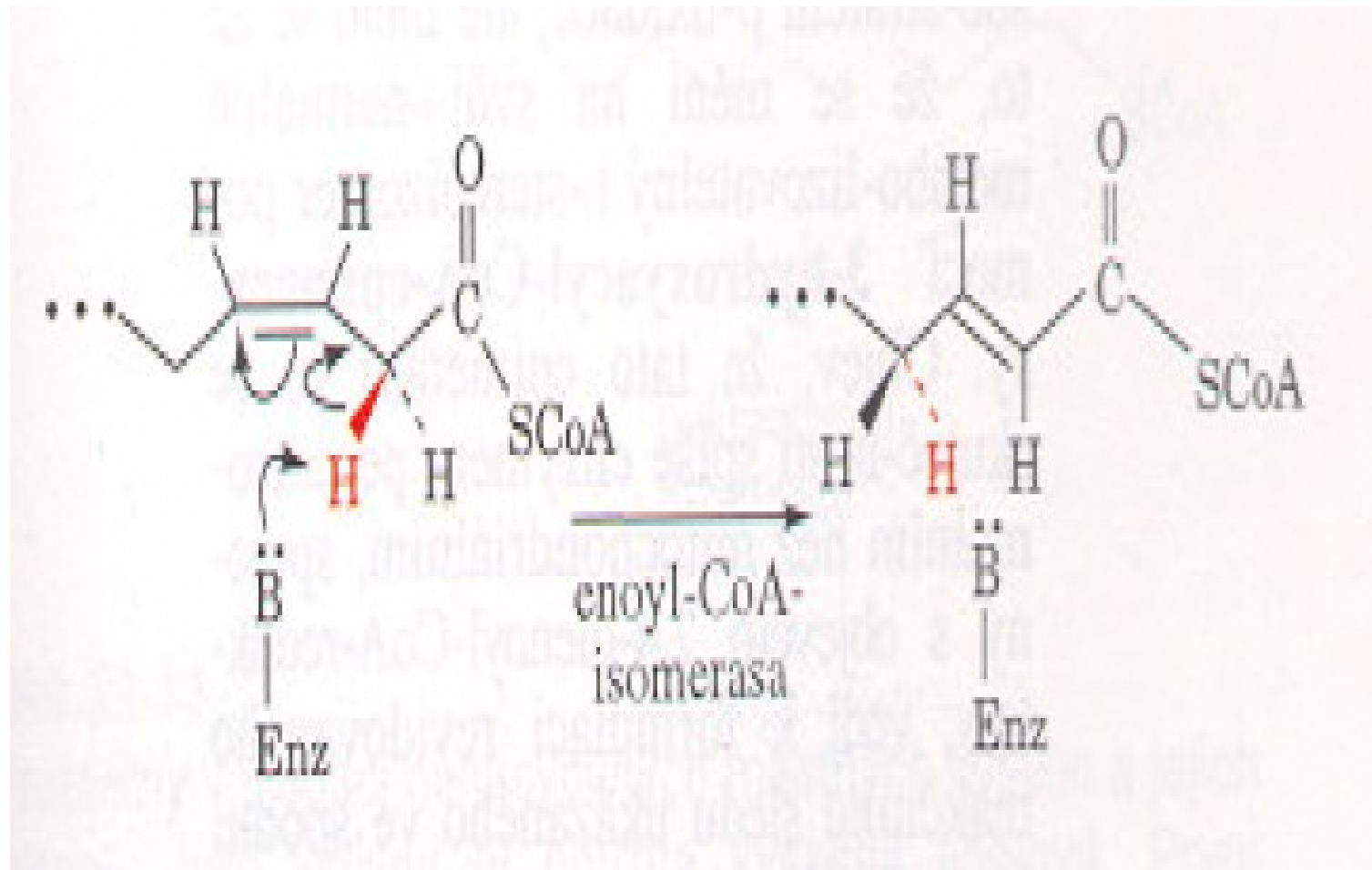


## Bilance $\beta$ oxidace :

1. cyklus - 1 FADH<sub>2</sub> (2 ATP) + 1 NADH (3 ATP) - 5 ATP  
acetylCoA (citrátový cyklus) - 12 ATP

na C<sub>16</sub> - 7 x  $\beta$  oxidace + 8 x citrátový cyklus - aktivace  
(7 x 5) + (8 x 12) - 2 ATP = 129 ATP

## Odbourávání nenasycených mastných kyselin





three cycles of  $\beta$ -oxidation



enoyl-CoA isomerase



complete  $\beta$ -oxidation cycle



acyl-CoA dehydrogenase



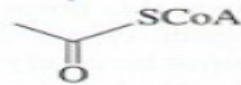
$\text{NADPH} + \text{H}^+ \rightarrow \text{NADP}^+$  (catalyzed by 2,4-dienoyl-CoA reductase)

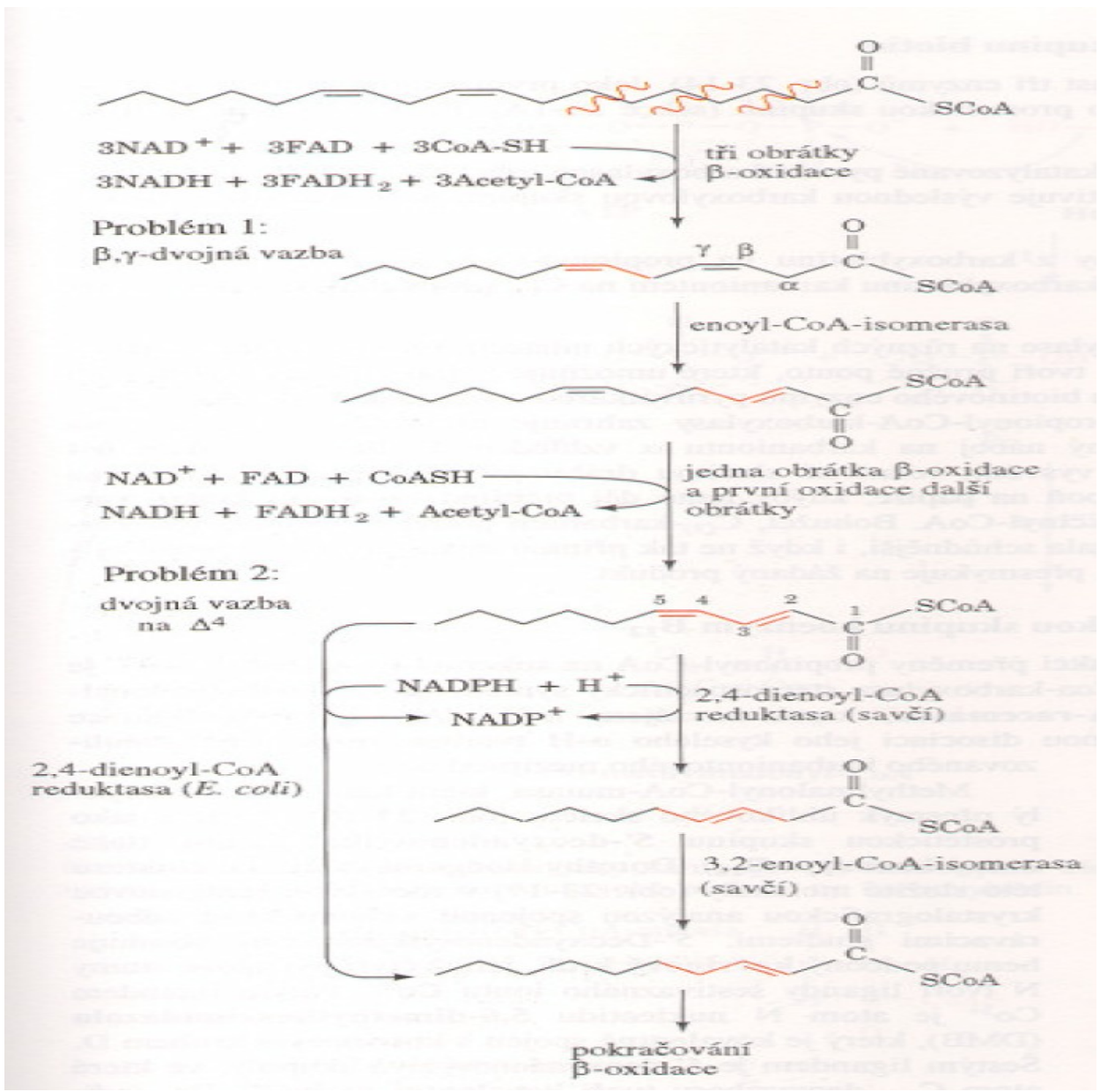


enoyl-CoA isomerase

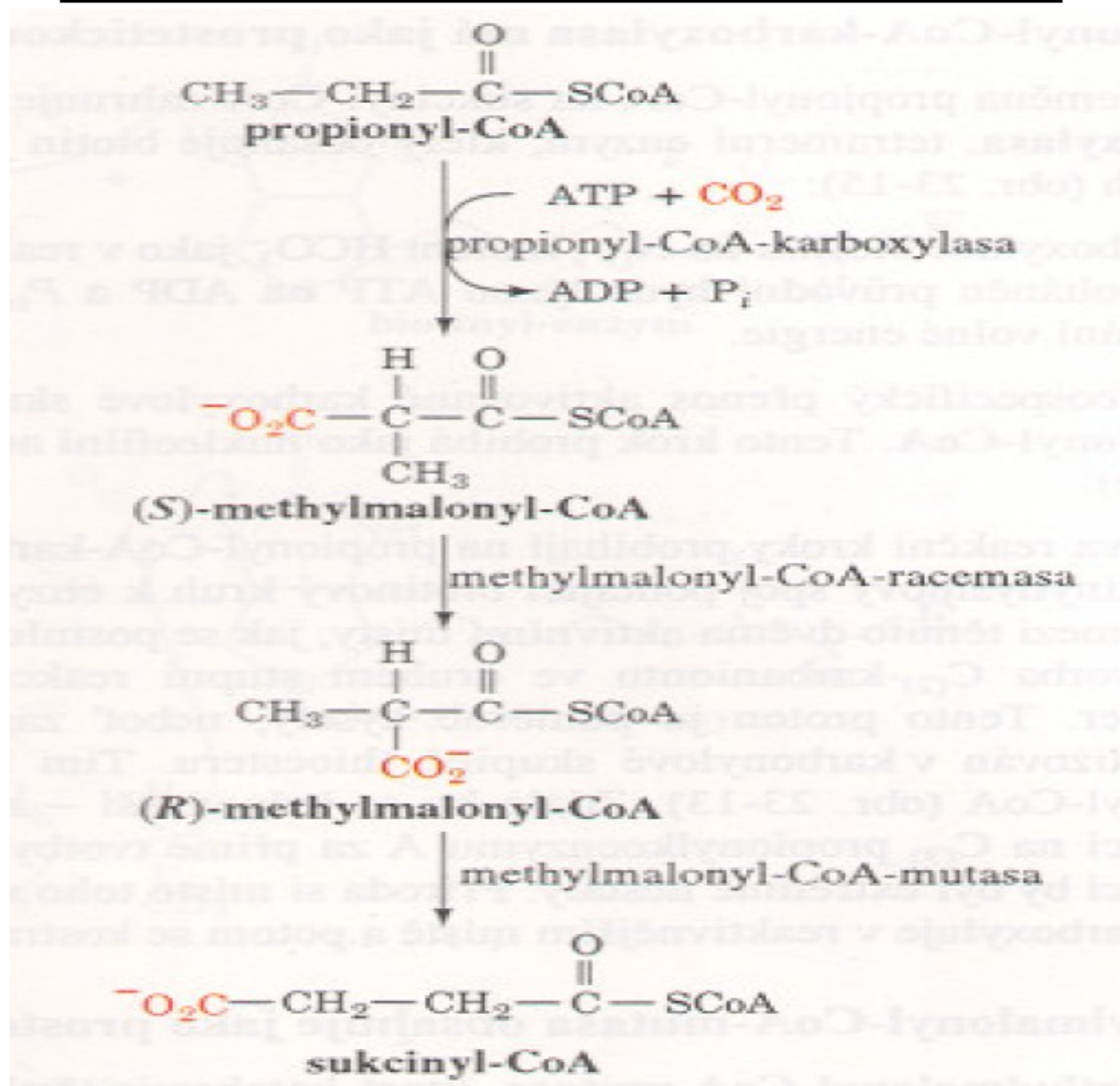


three cycles of  $\beta$ -oxidation



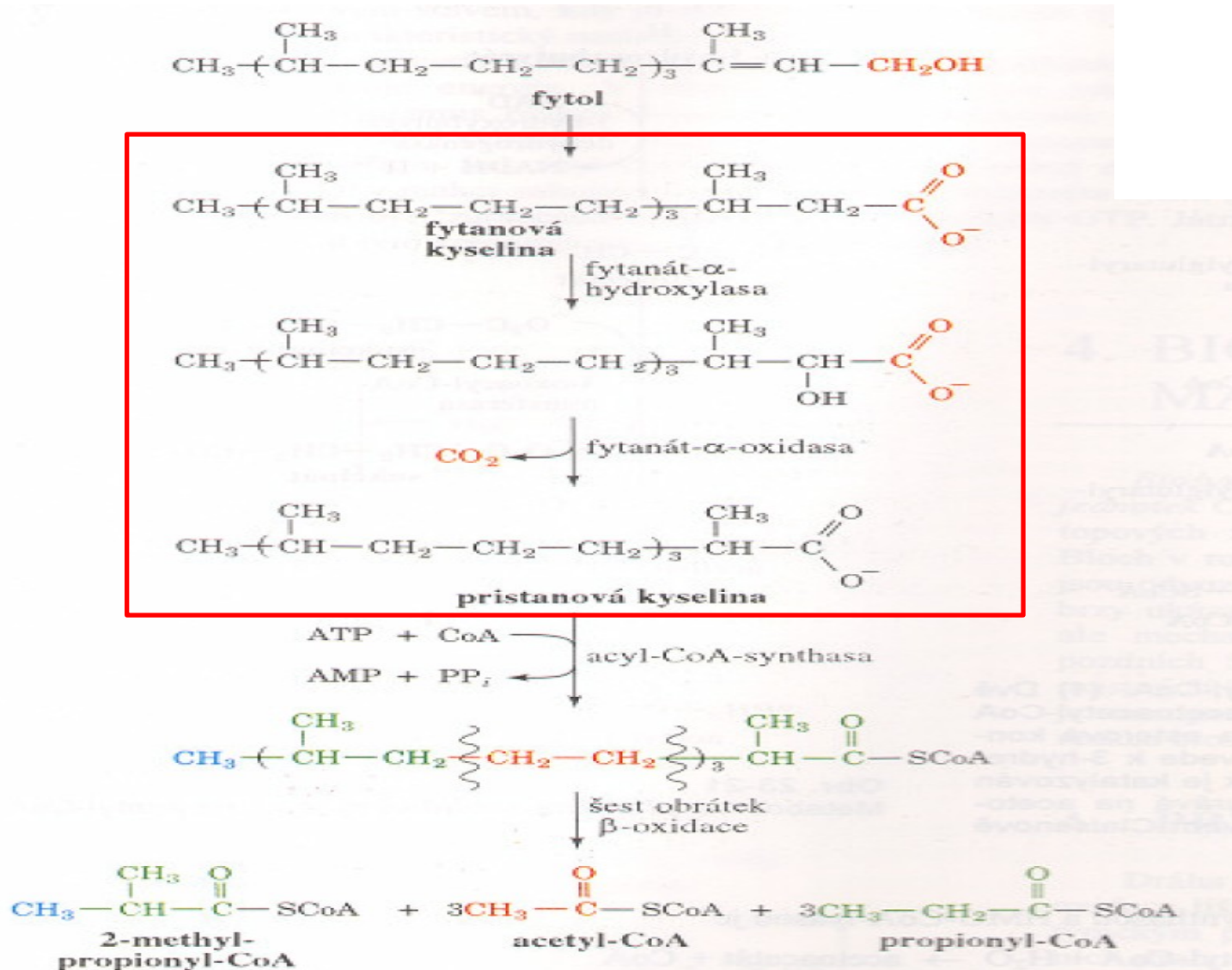


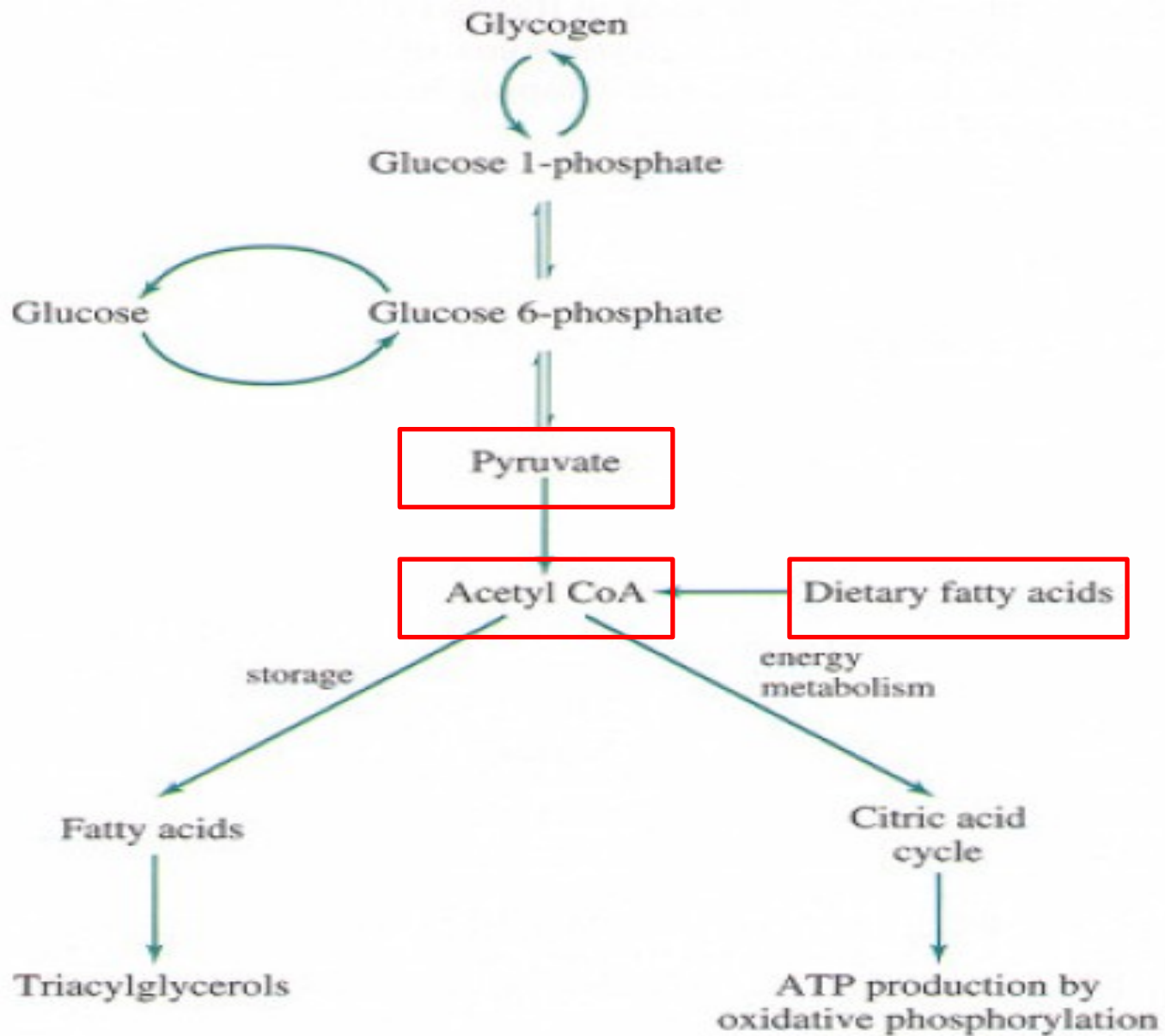
## Odbourávání mastných kyselin s lichým počtem C atomů



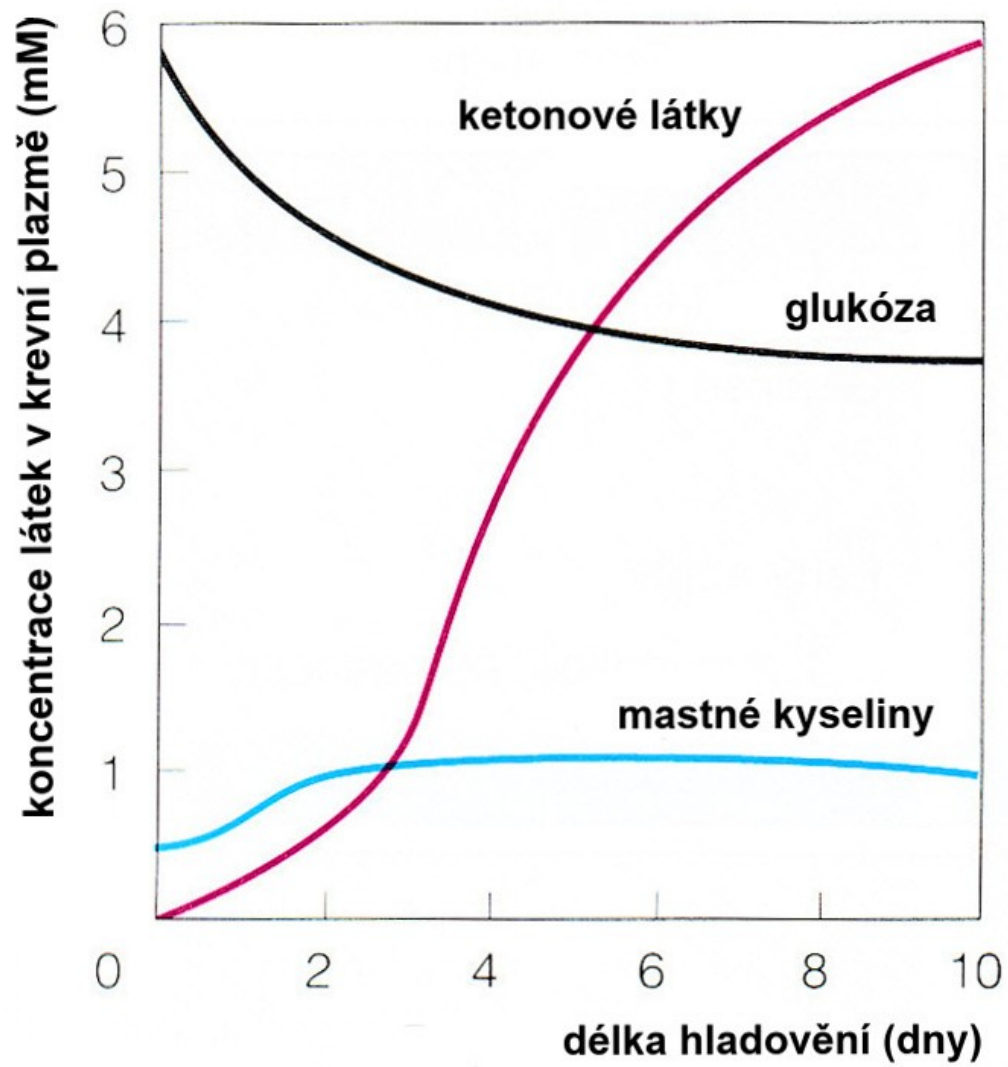


# $\alpha$ -oxidace

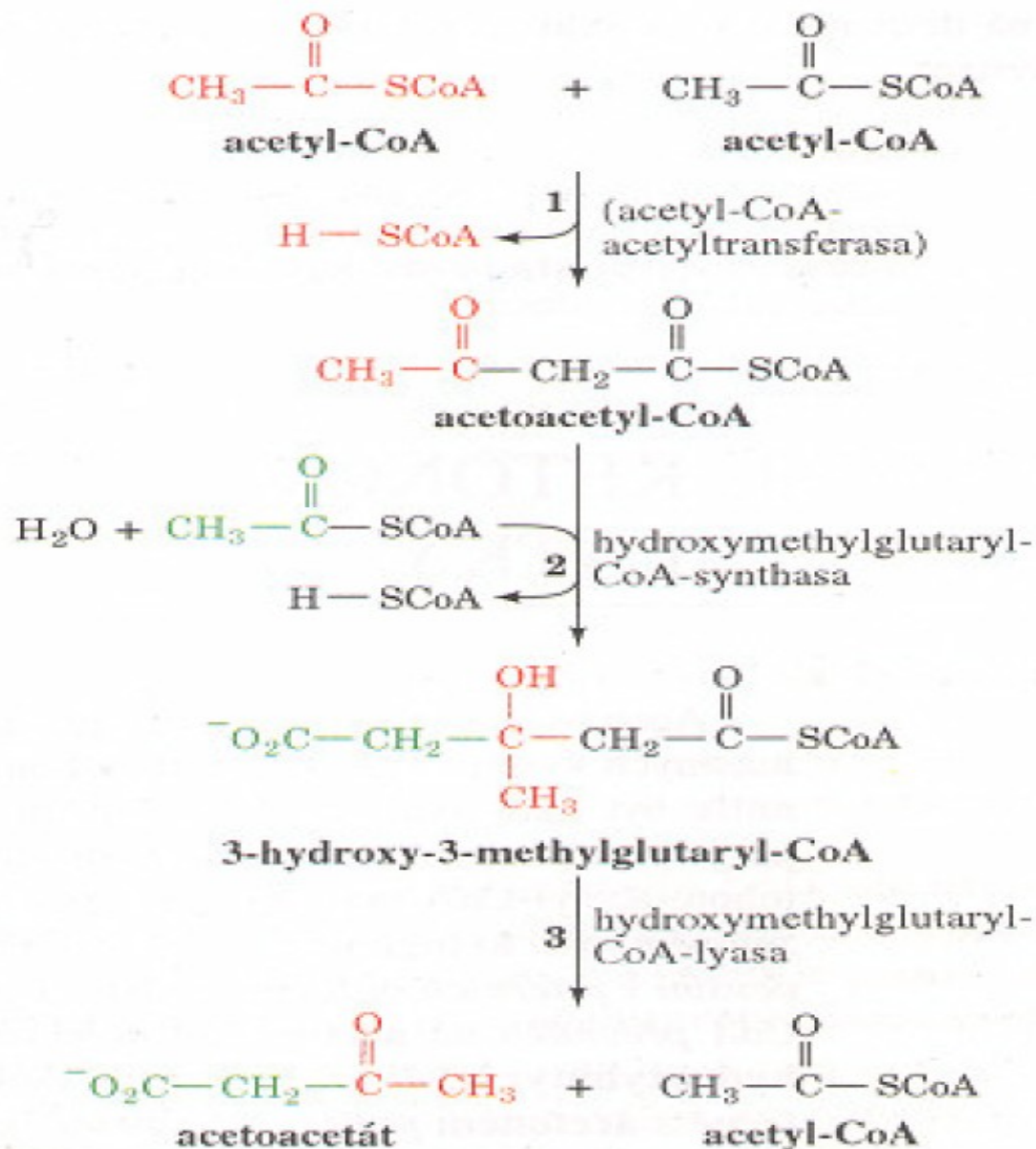




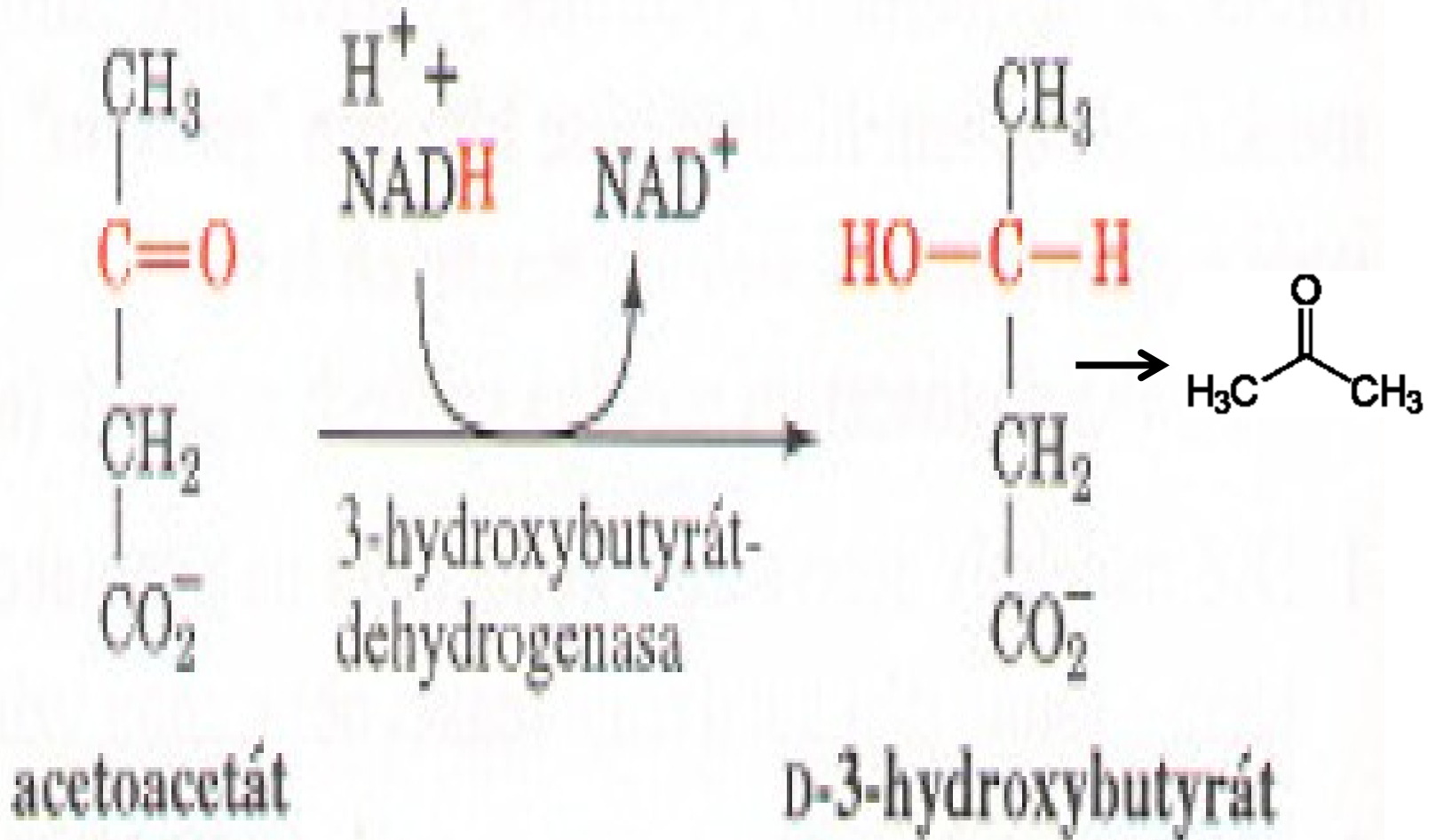




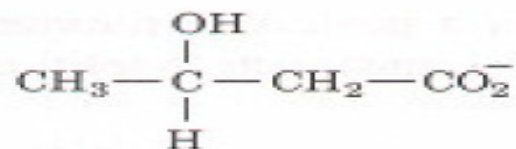
## Ketonové látky



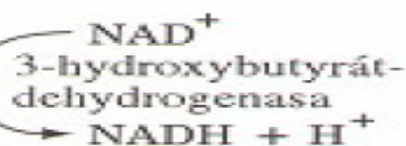
Ketonové látky



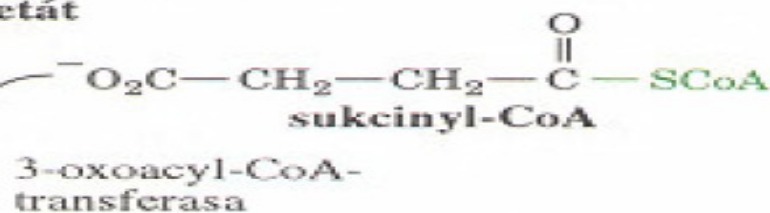
## Ketonové látky



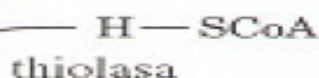
**D-3-hydroxybutyrát**



**acetoacetát**



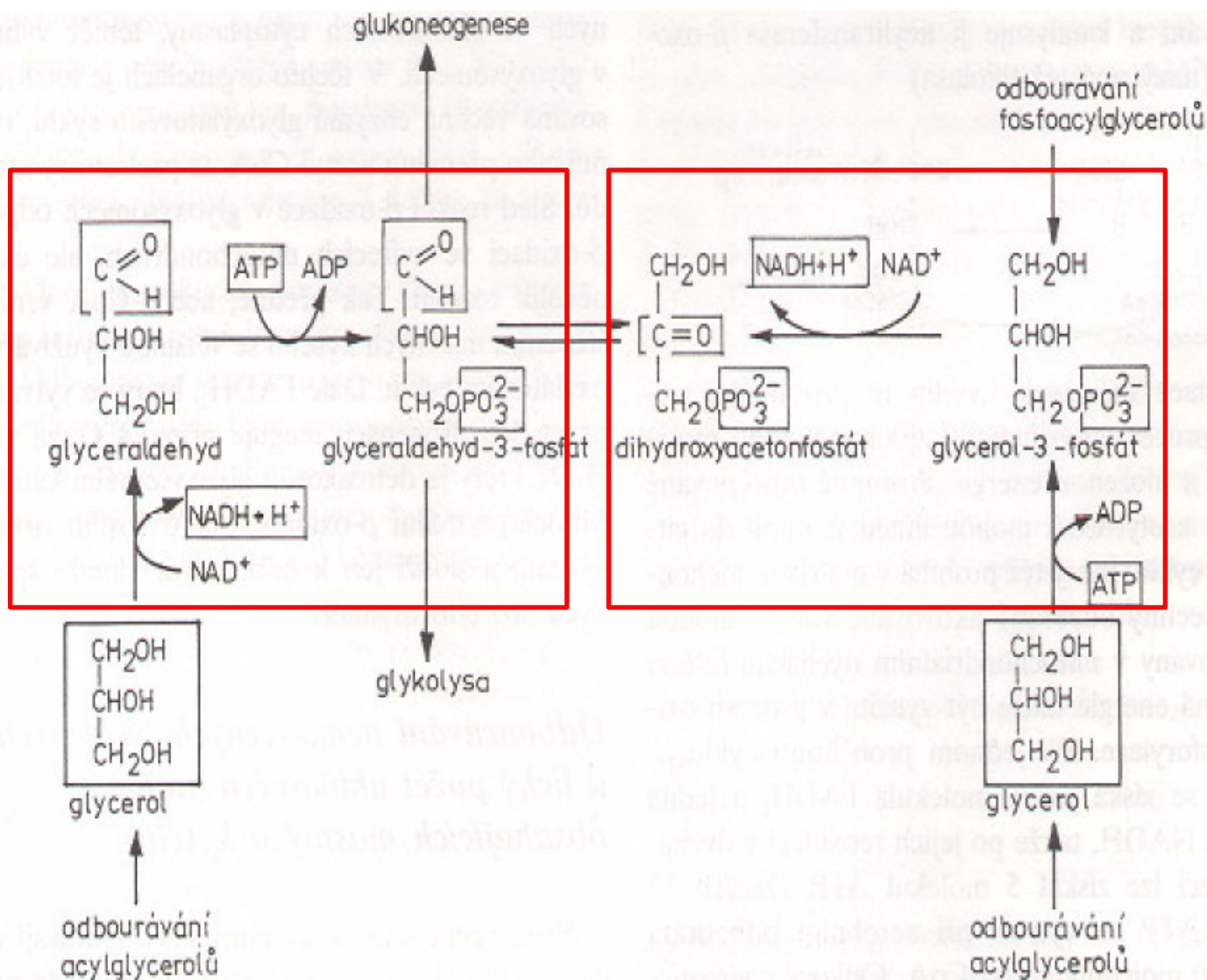
**acetoacetyl-CoA**



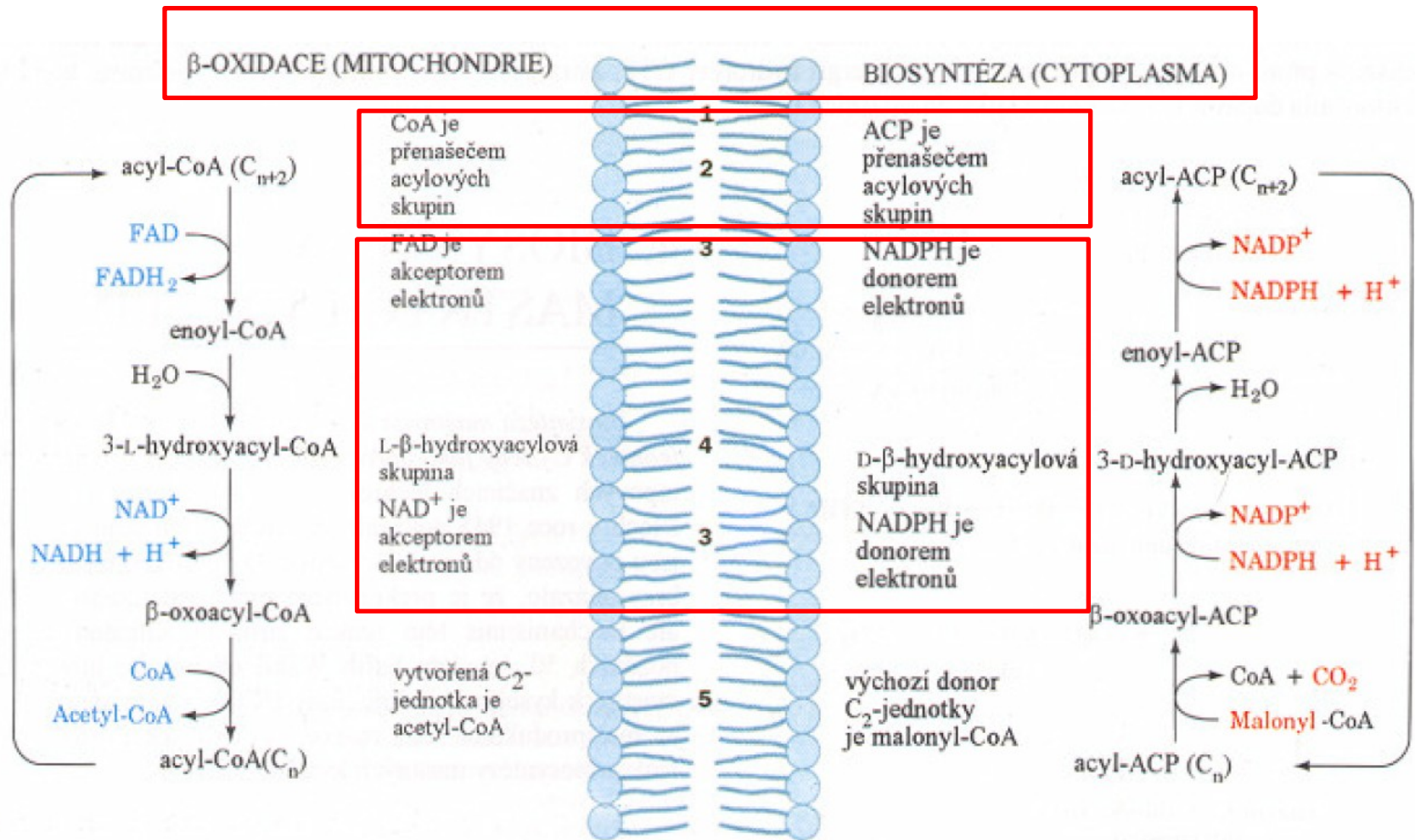
**acetyl-CoA**



# Metabolismus glycerolu



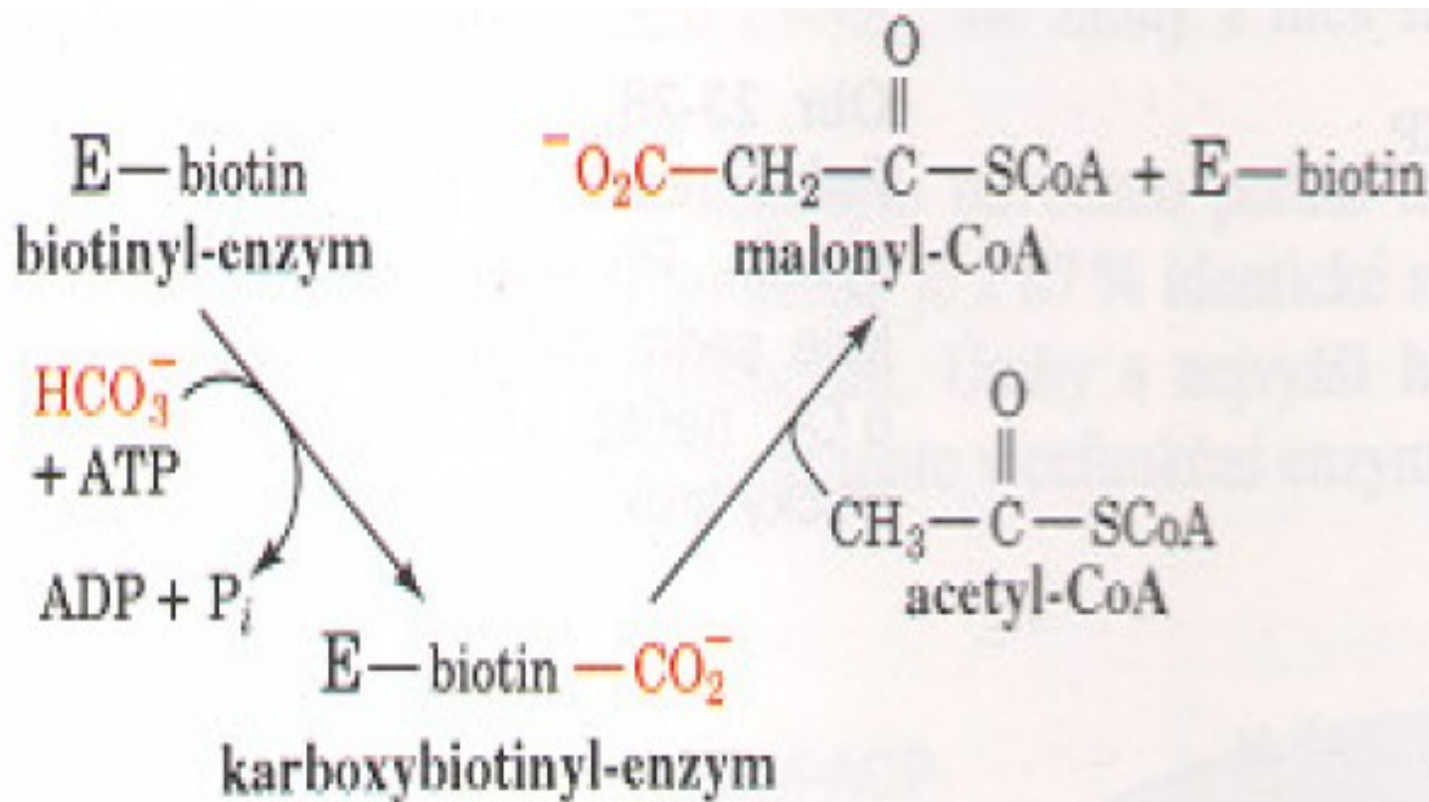
# Metabolismus versus biosyntéza MK





## Biosyntéza mastných kyselin

### *A. Syntéza malonylCoA*

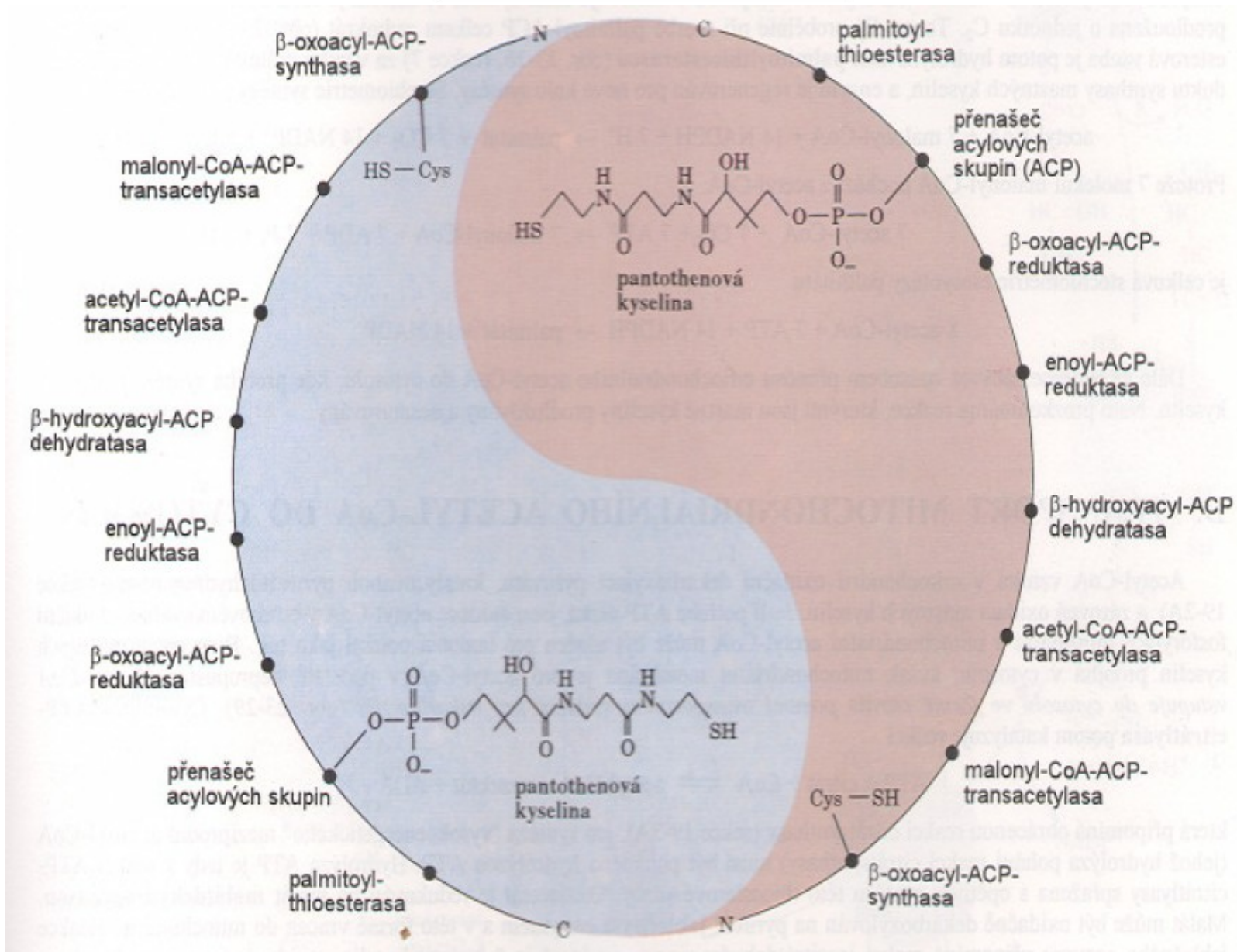




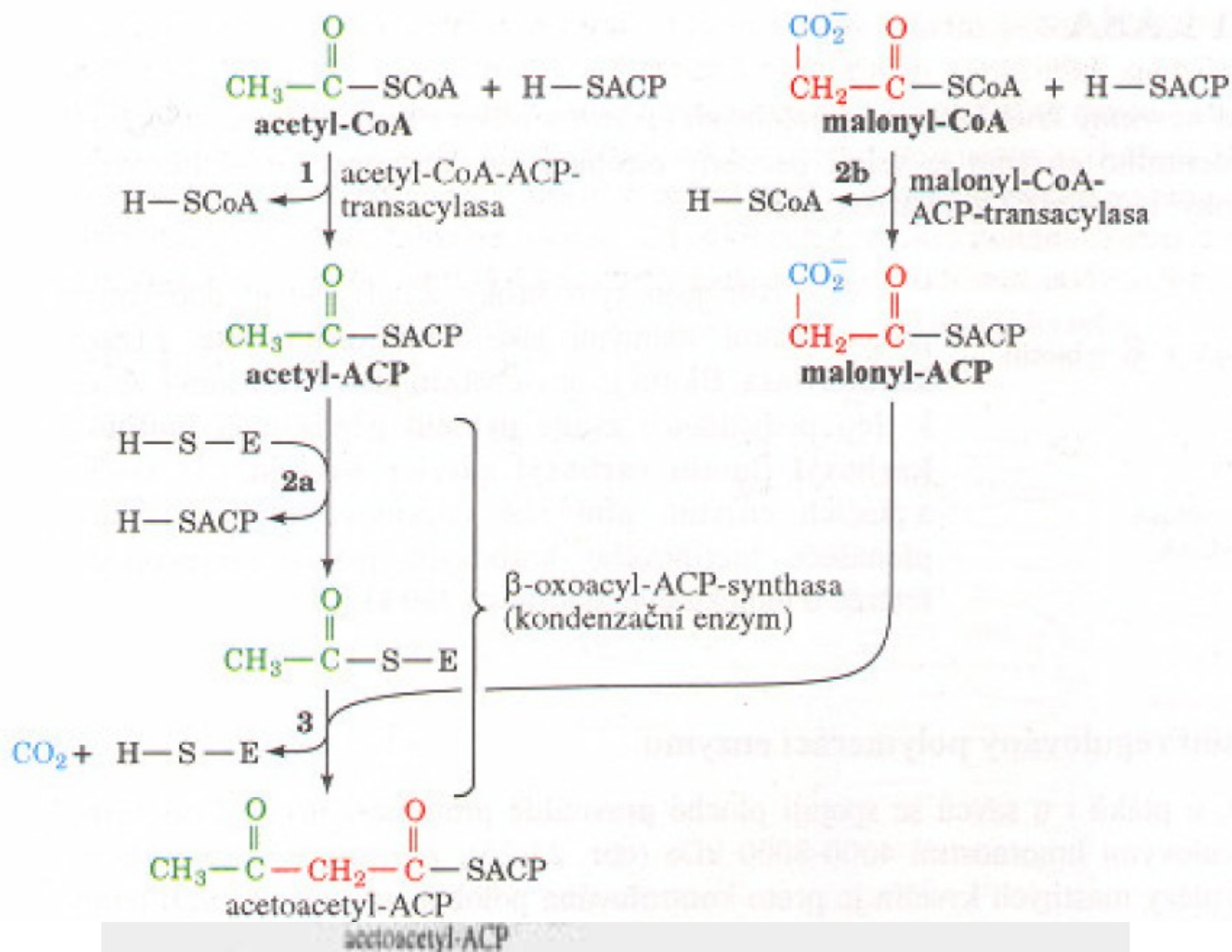
# Regulace Acetyl-CoA karboxylasy



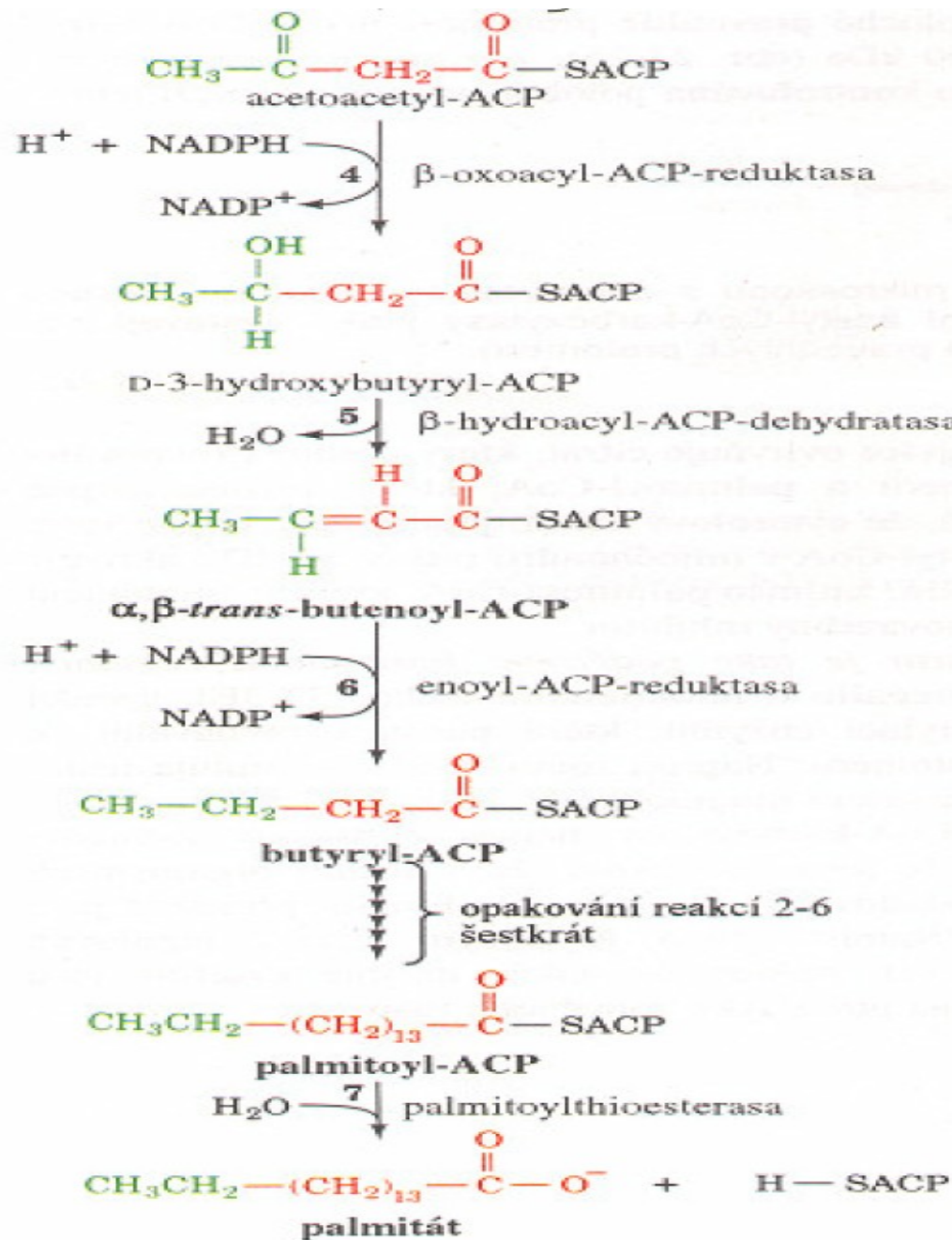
## Synthasa mastných kyselin



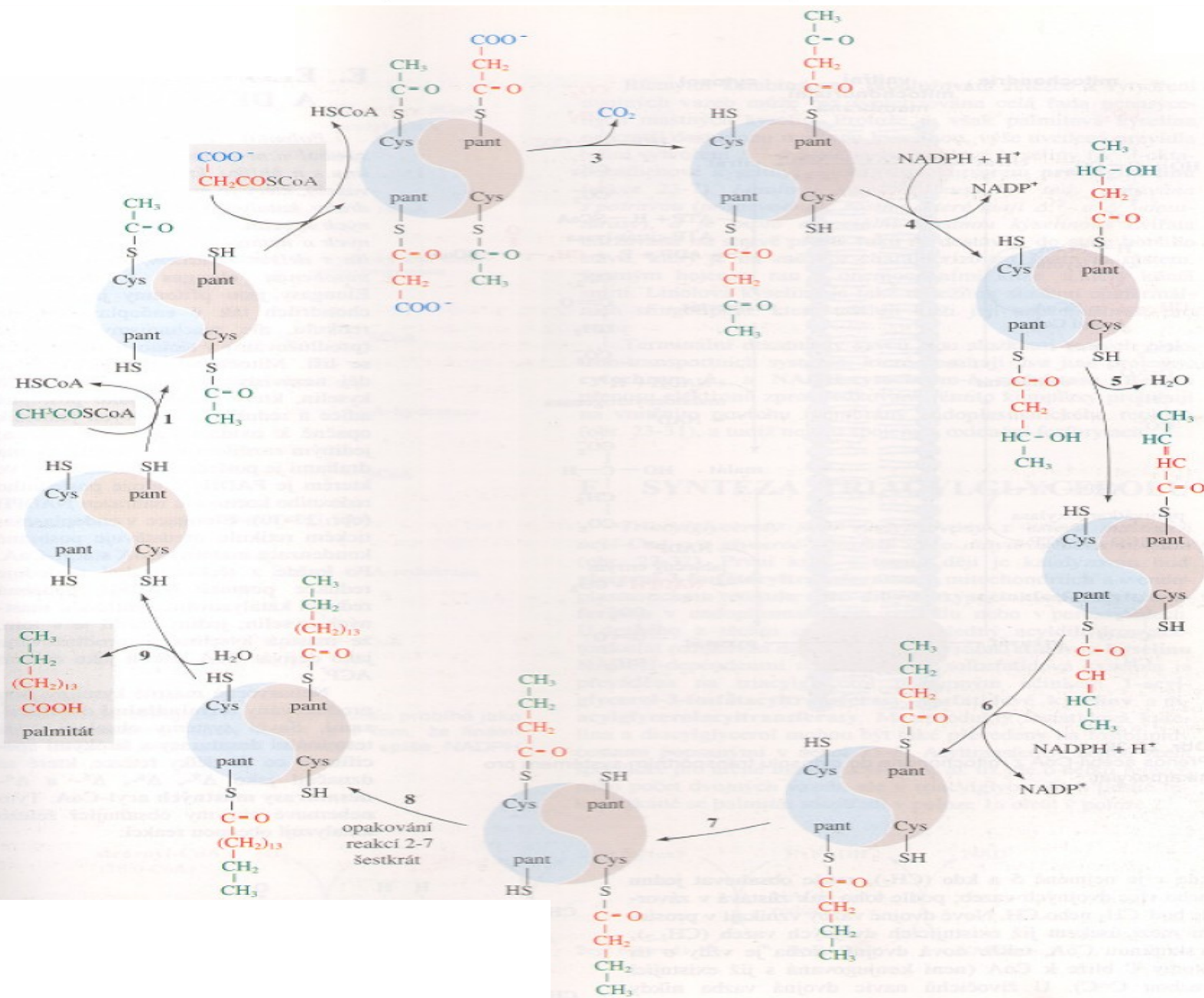
## B. Syntéza palmitové kyseliny







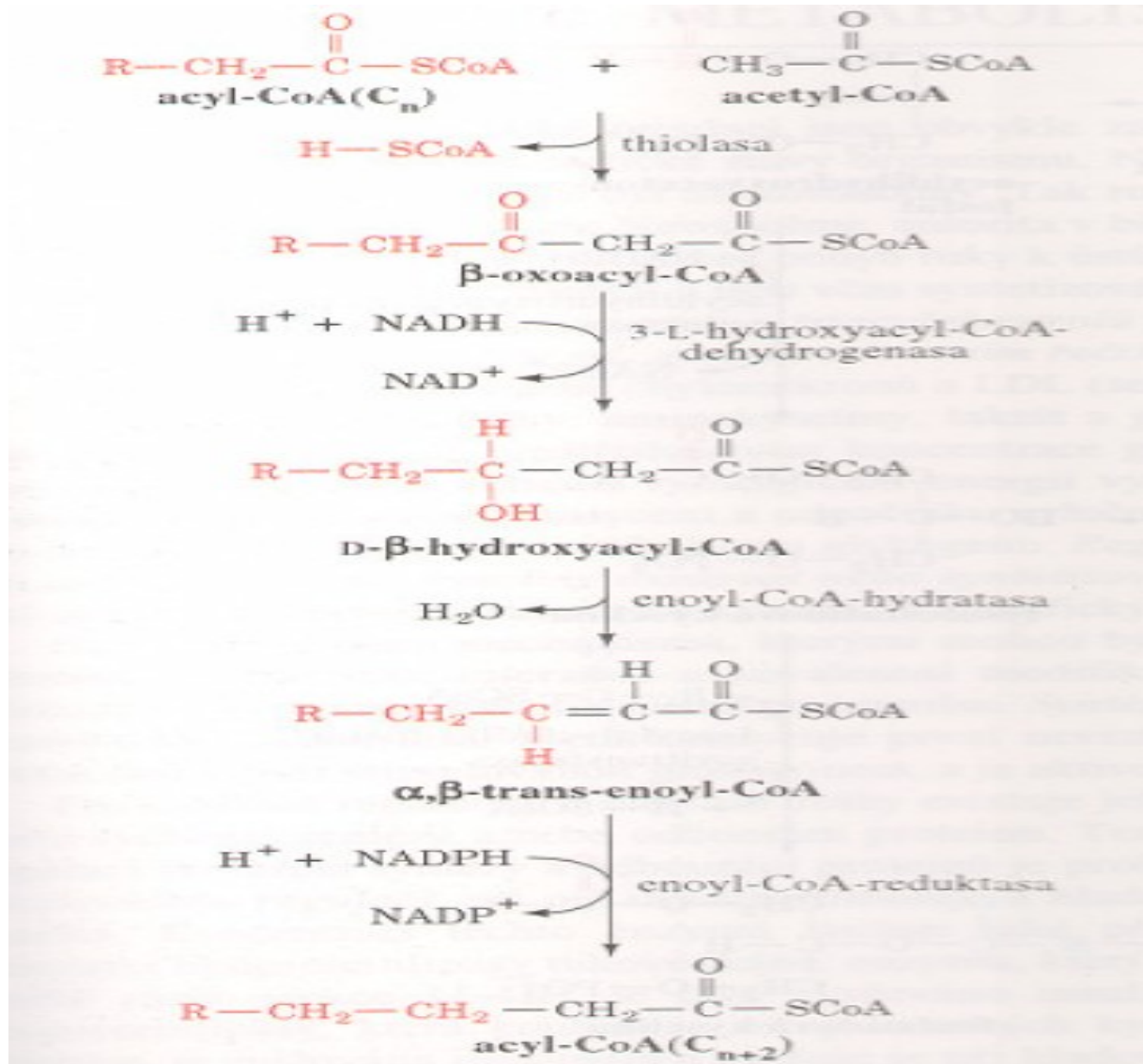
# Synthesa mastných kyselin



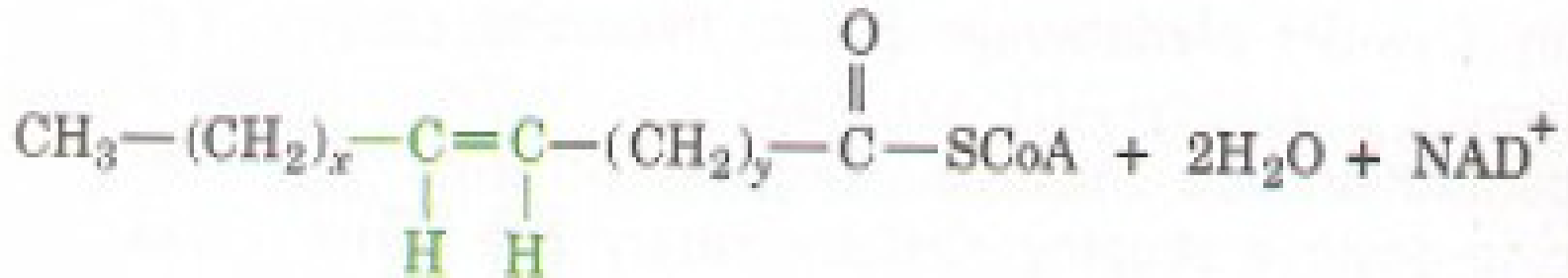
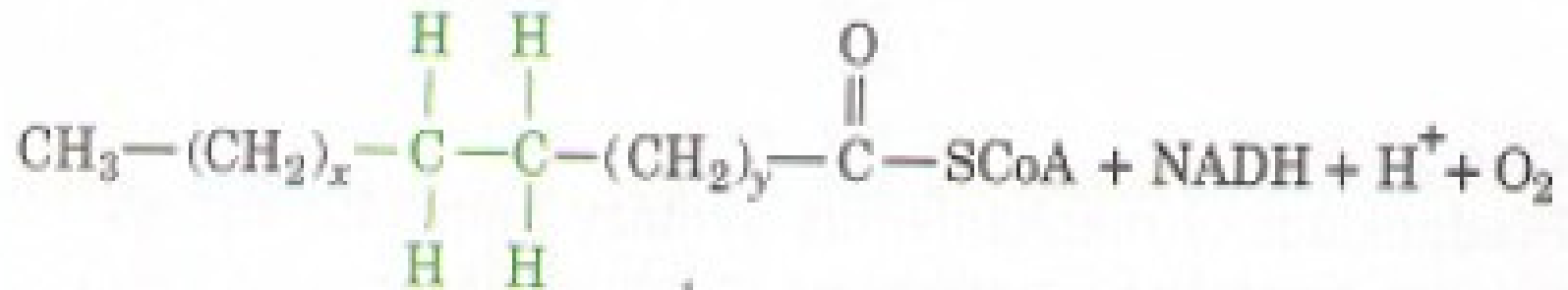
### *C. Další přeměny palmitové kyseliny*

- **prodlužování řetězce - elongace - elongasy**
- **dehydrogenace - desaturece - desaturasy**

# Elongace



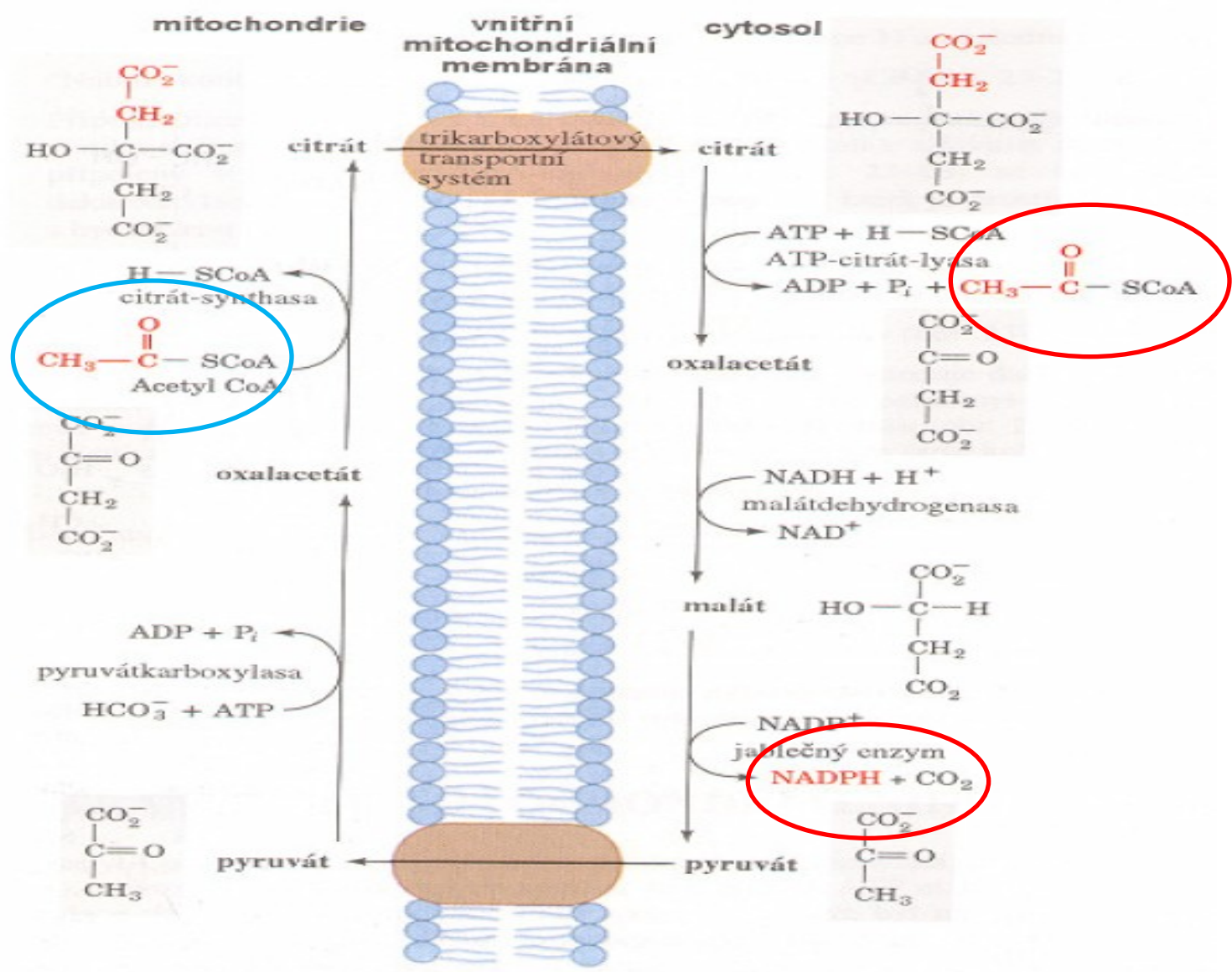
# Desaturase



**Bilance biosyntézy mastných kyselin :**

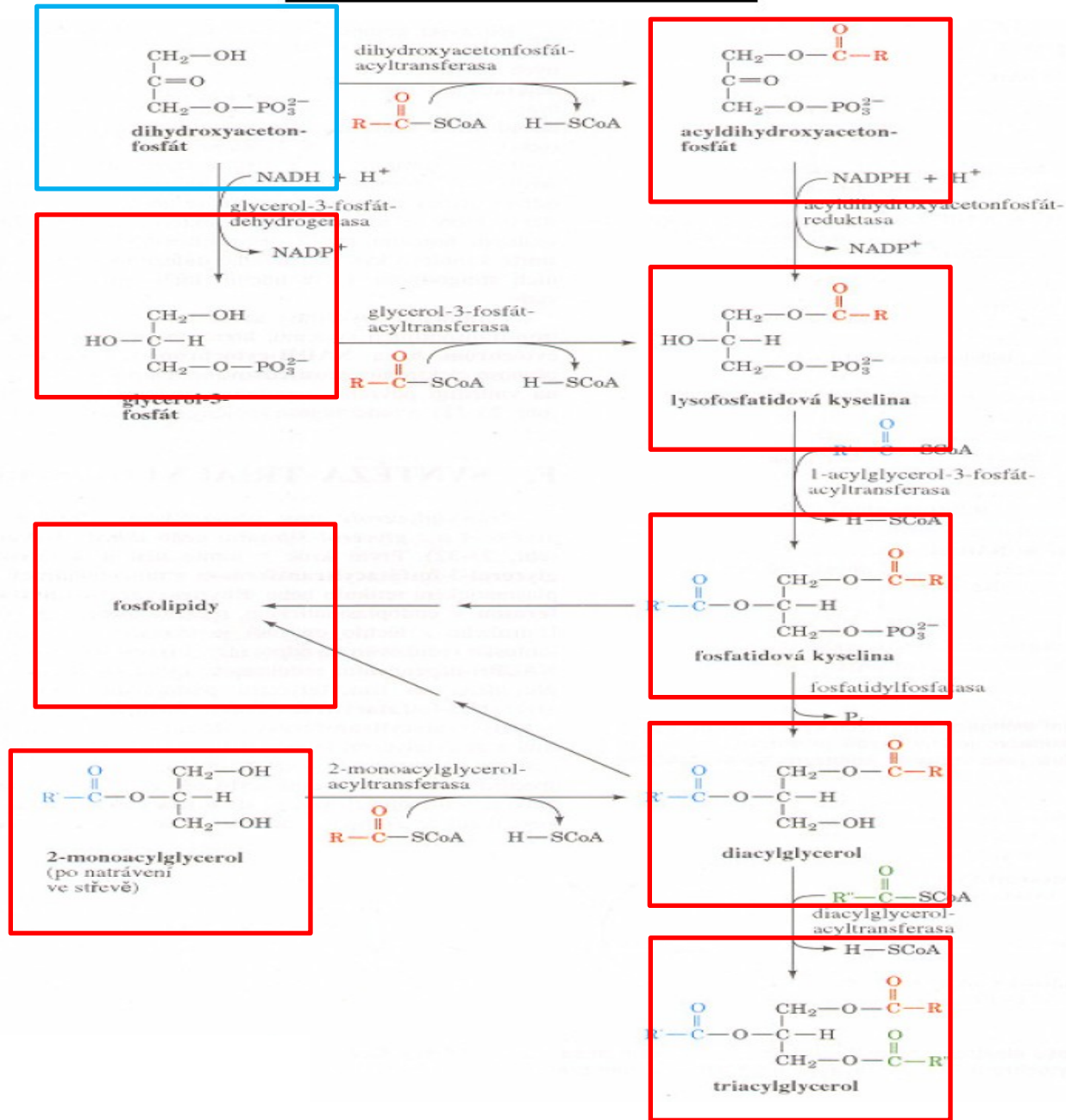
1. cyklus	syntéza malonylCoA	1 ATP
	2 NADPH na redukci	6 ATP
<hr/>		
na C <sub>16</sub>	7 x ( $\frac{16}{2} - 1$ )	49 ATP

# Přenos AcetCoA vně mitochondrie

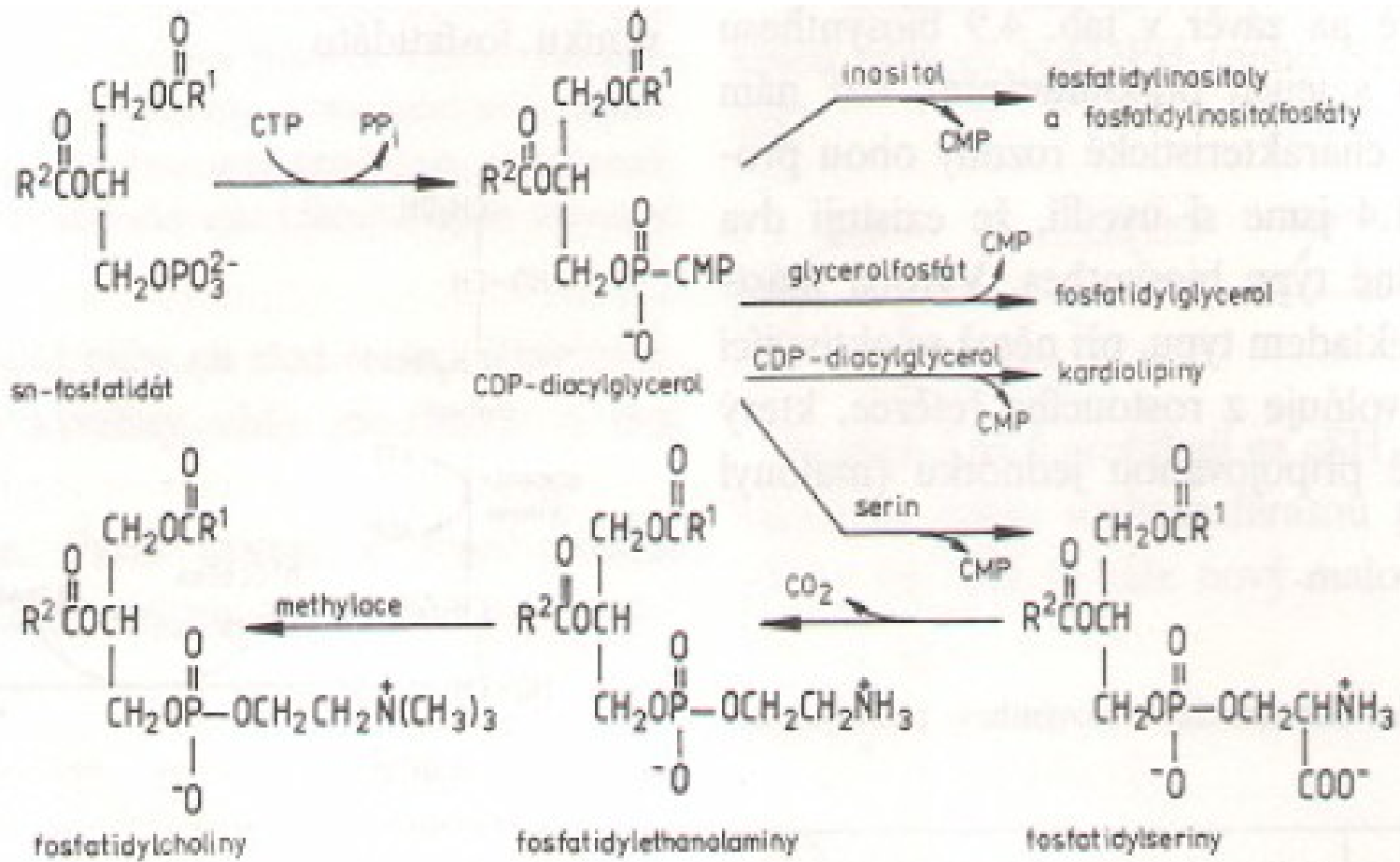




# Biosyntéza triacylglycerolů



# Biosyntéza fosfolipidů

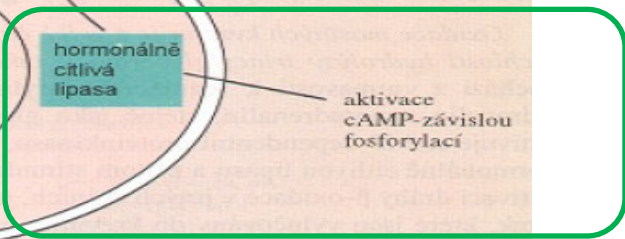
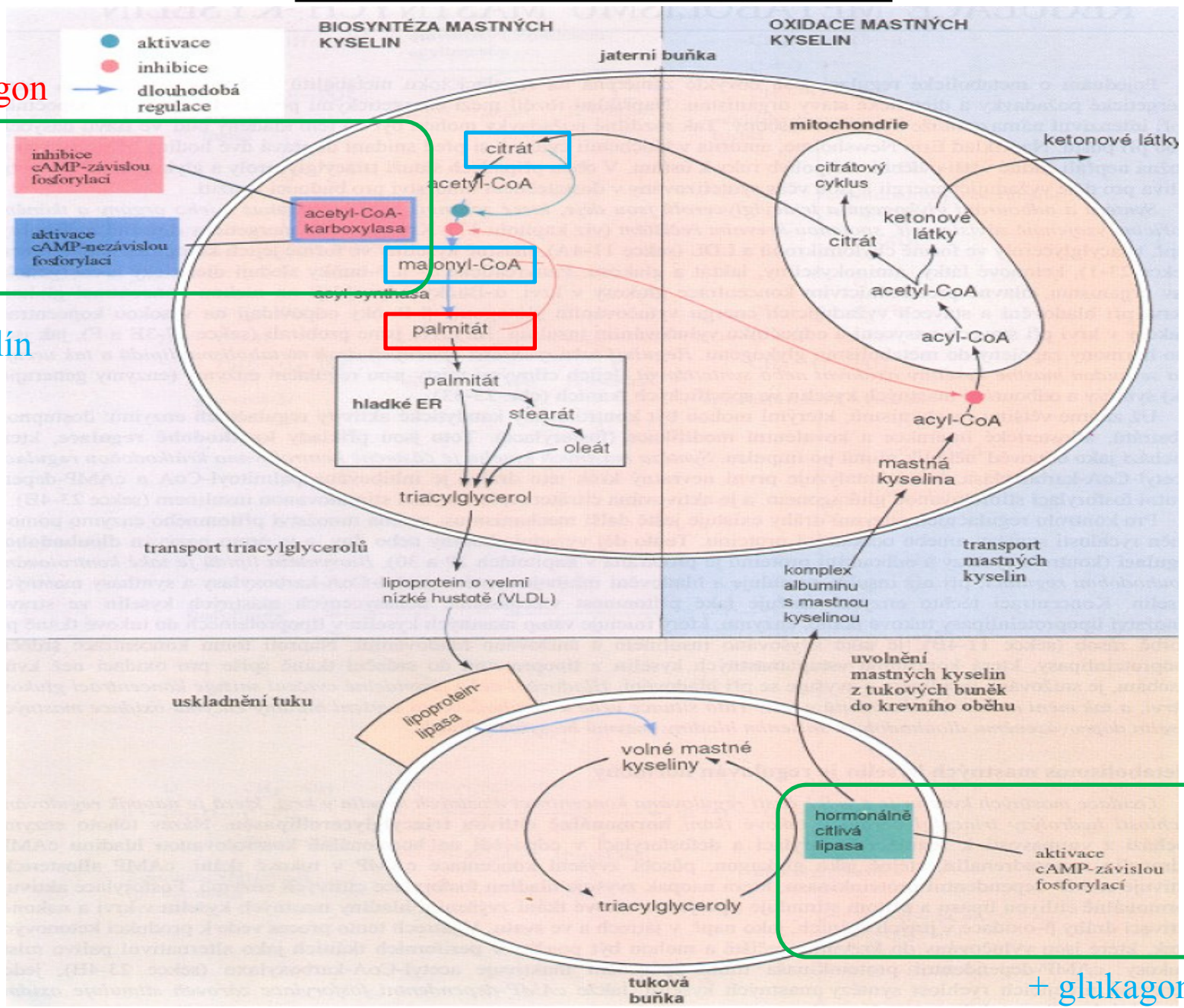
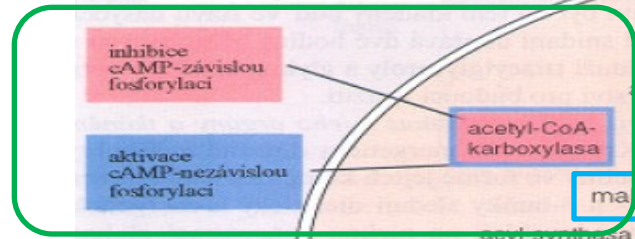


# Regulace metabolismu triacylglycerolů

- glukagon

+ inzulín

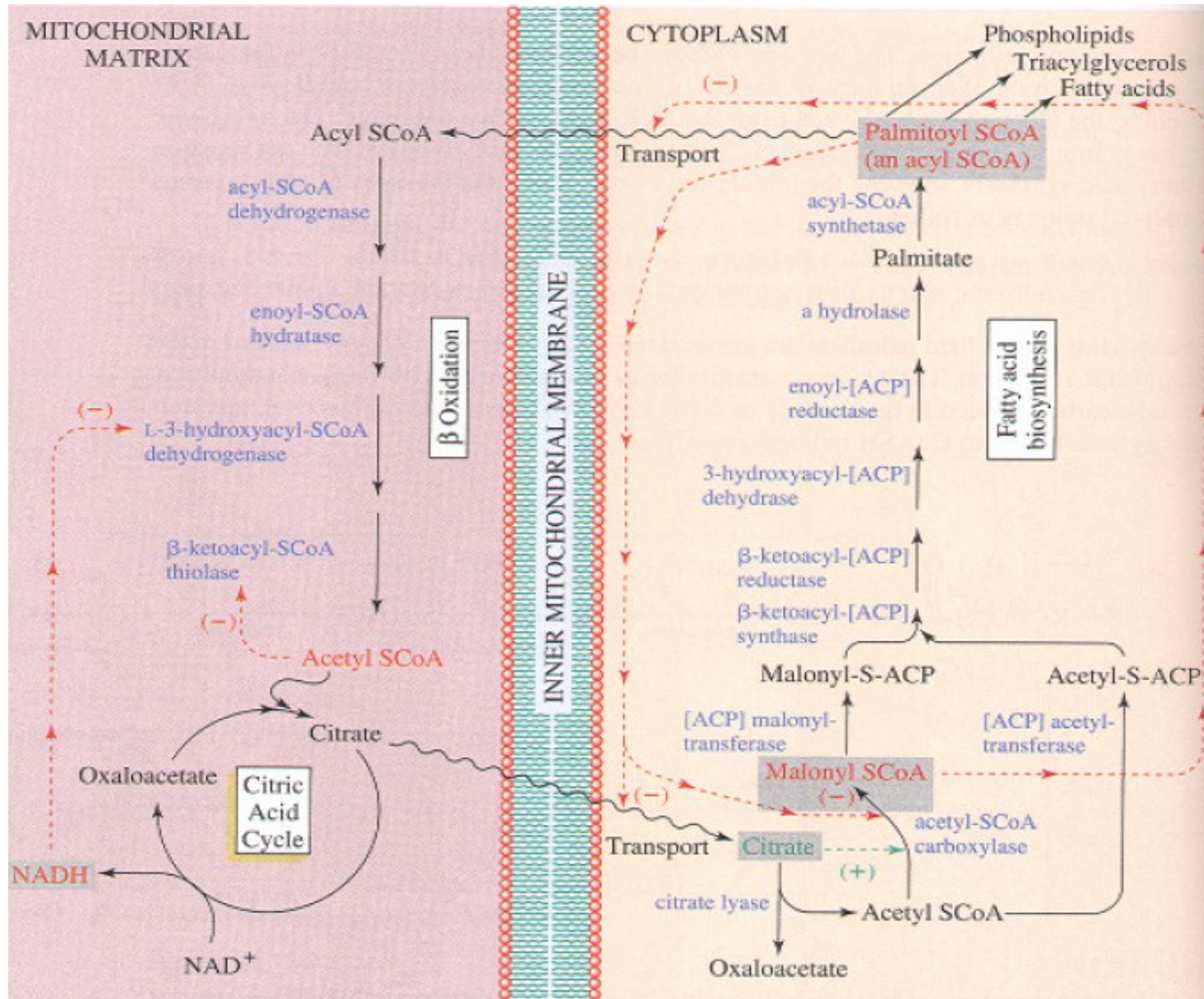
- aktivace
- inhibice
- dlouhodobá regulace



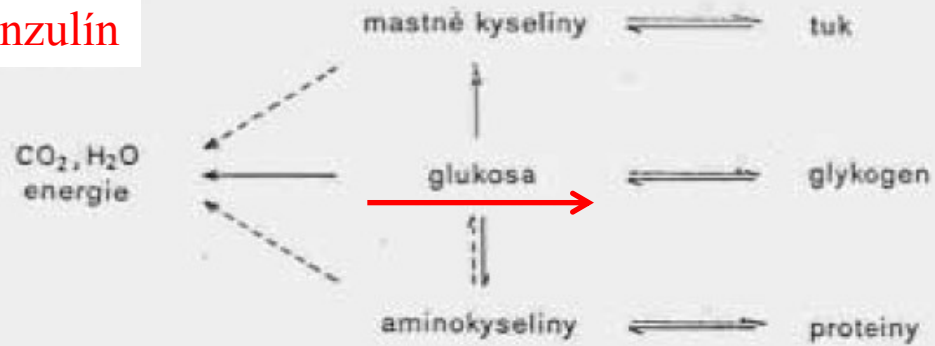
+ glukagon



## Regulate metabolismu triacylglycerolů



inzulín



glukagon

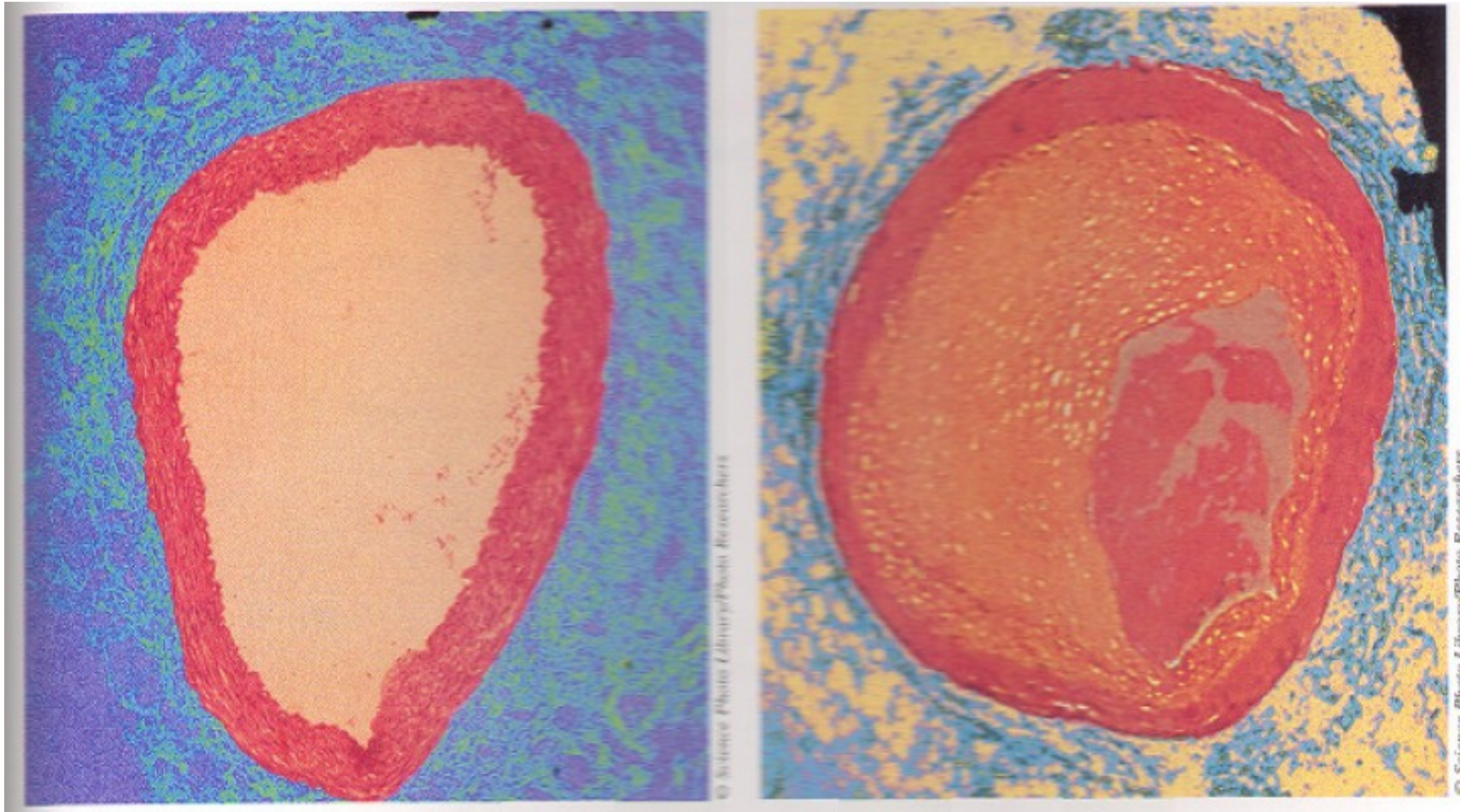


adrenalin

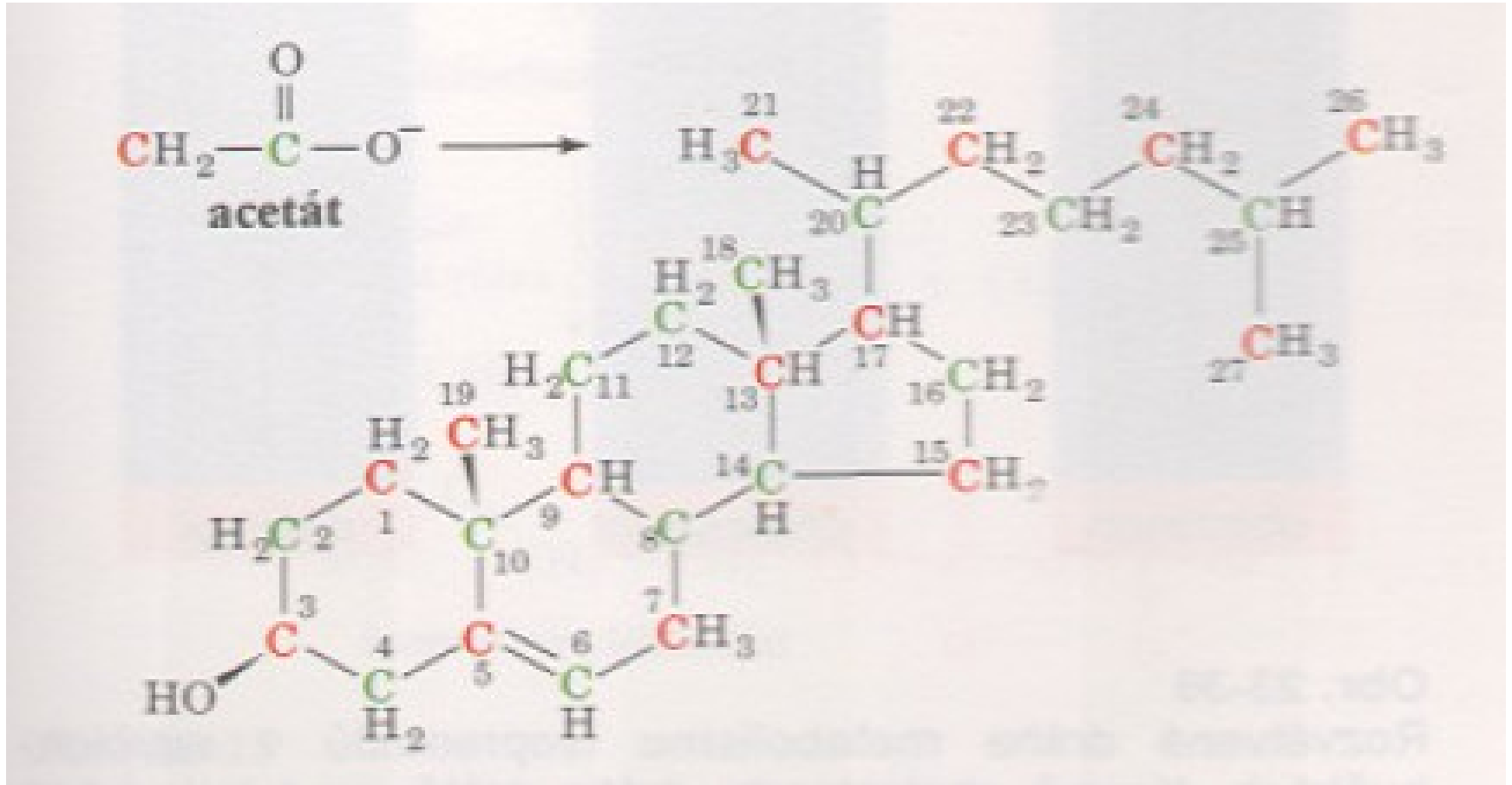




# Cholesterol



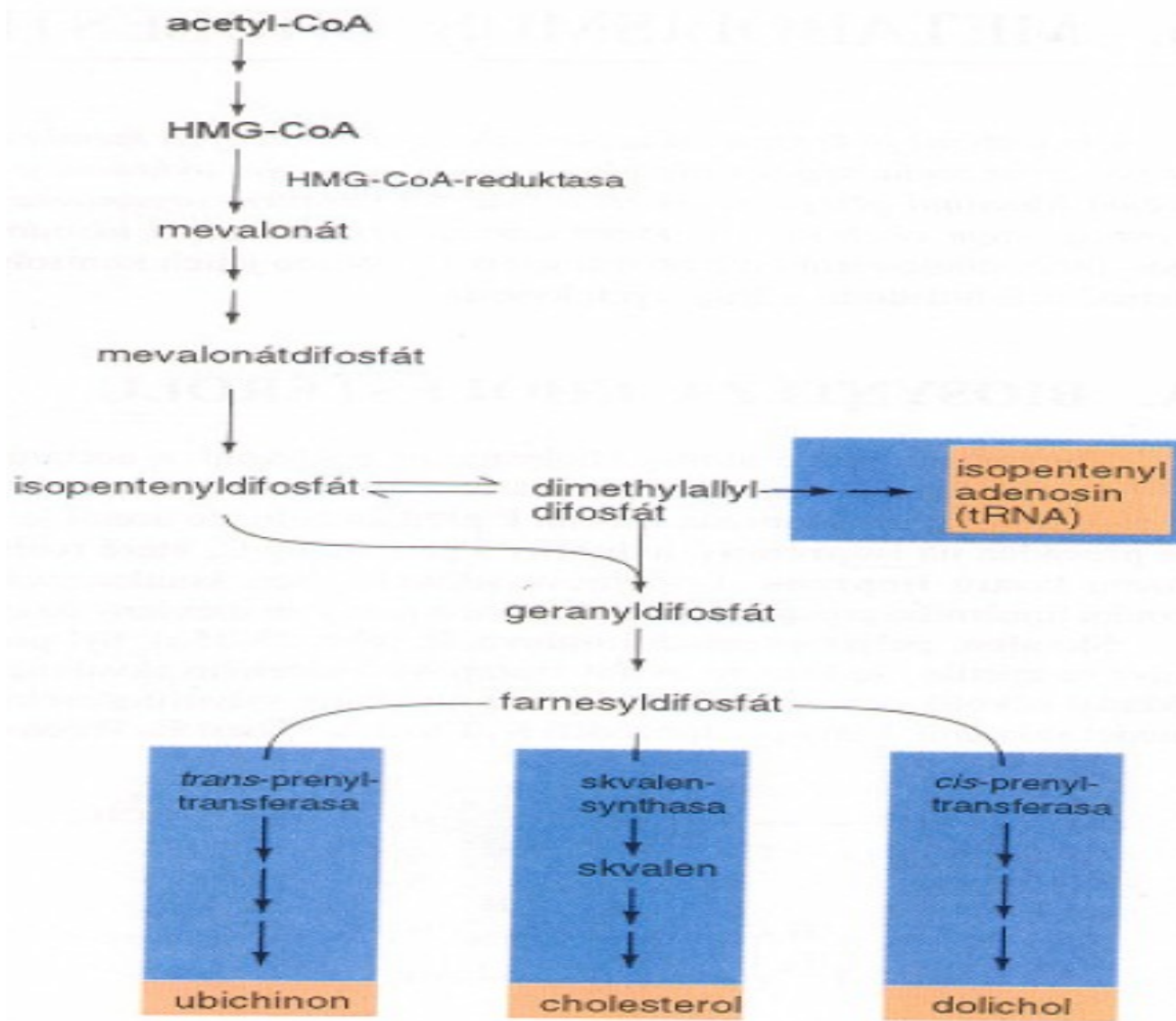
# Biosyntéza cholesterolu

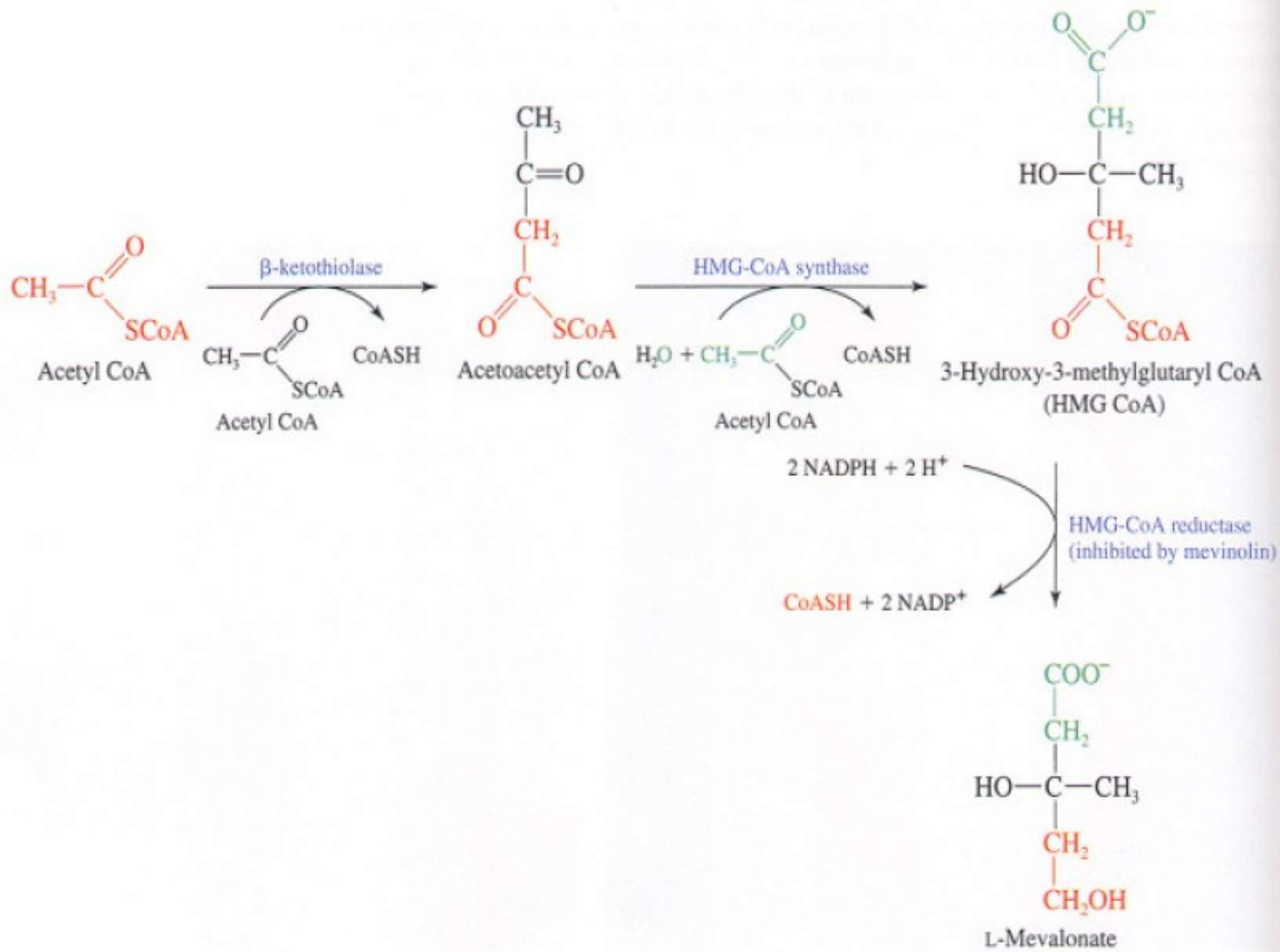


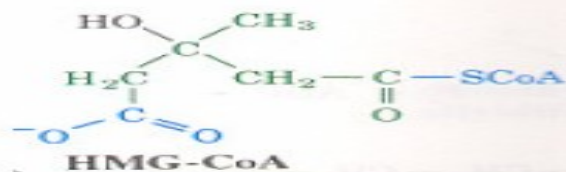
acetát → isoprenoidní intermediát → skvalen  
→ produkt cyklizace → cholesterol



# Biosyntéza cholesterolu



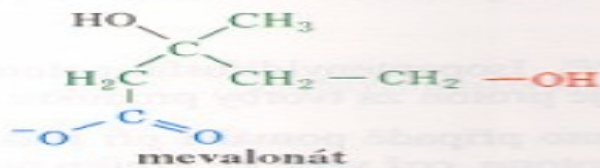




HMG-CoA-reduktasa

1

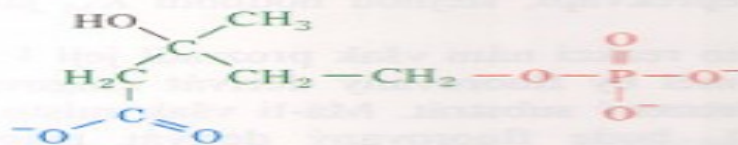
2NADPH → 2NADP<sup>+</sup> + CoA



mevalonát-5-fosfo-transferasa

2

ATP → ADP



fosfomevalonát-kinasa

3

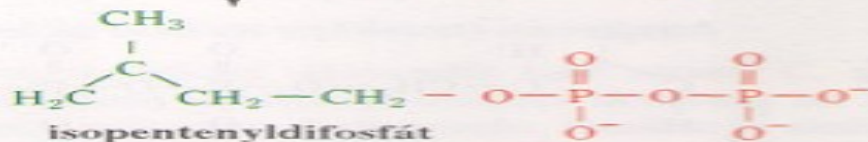
ATP → ADP

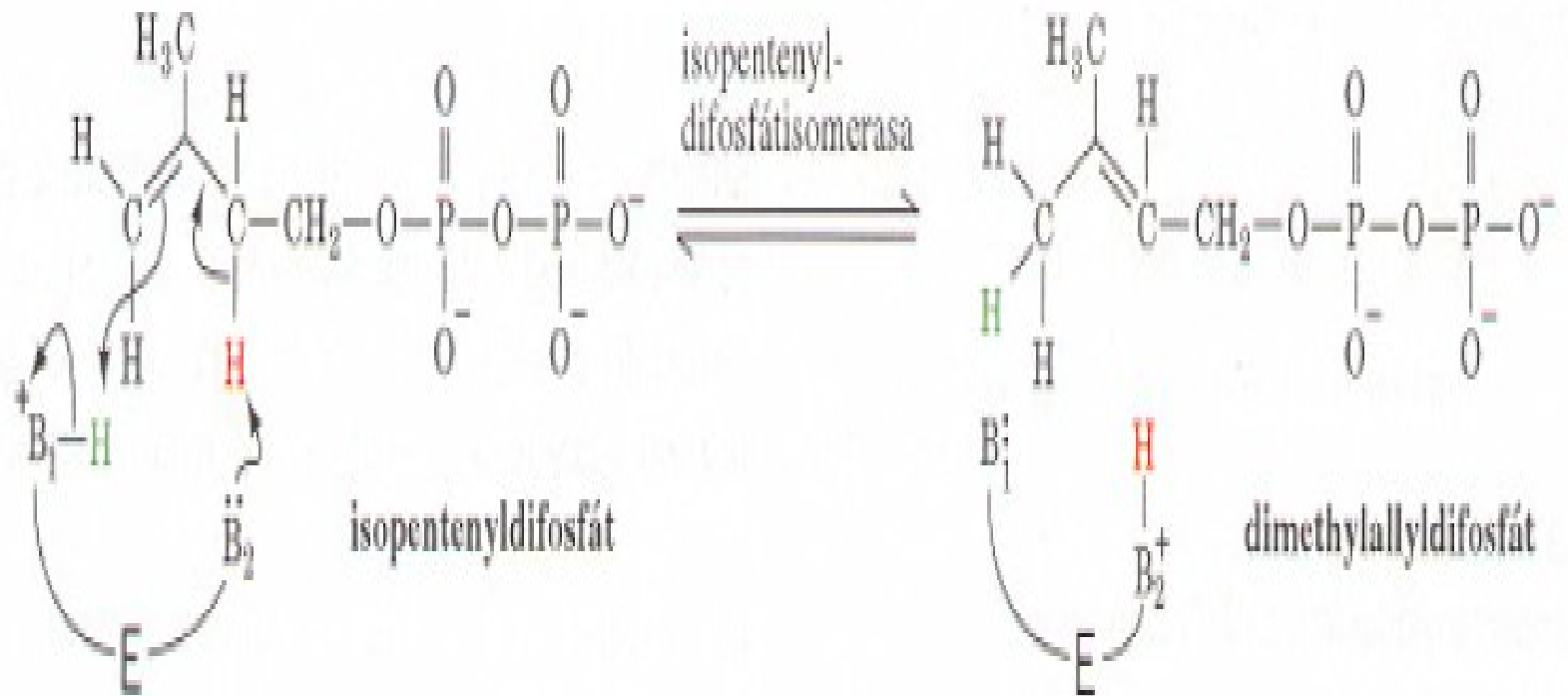


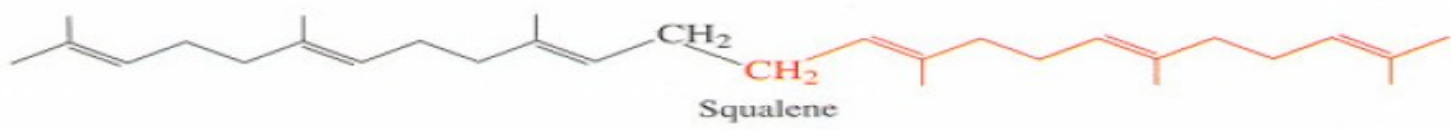
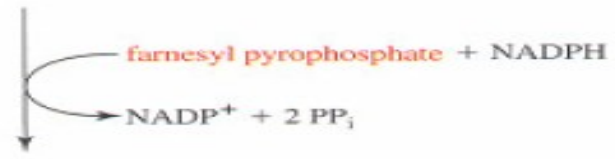
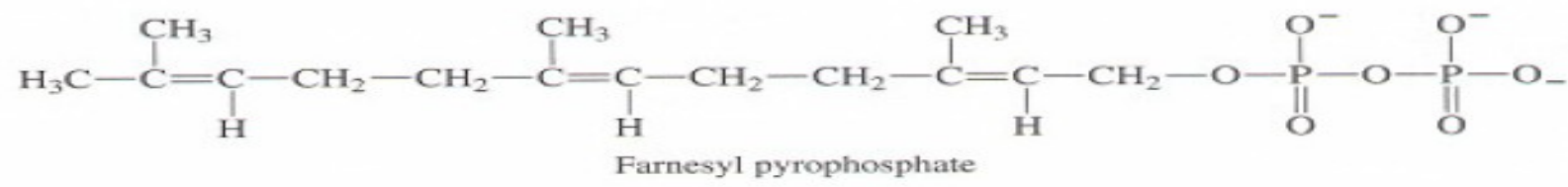
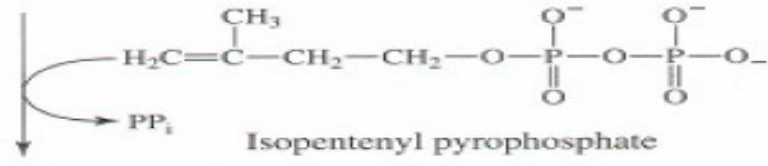
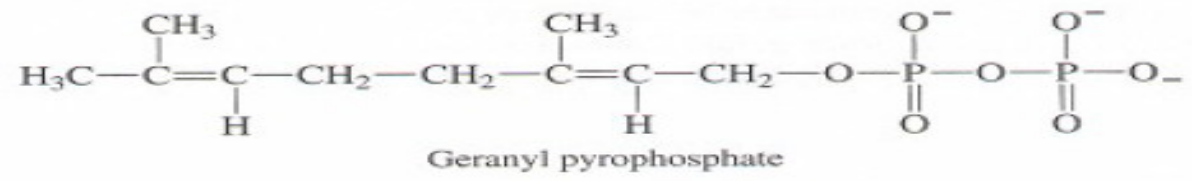
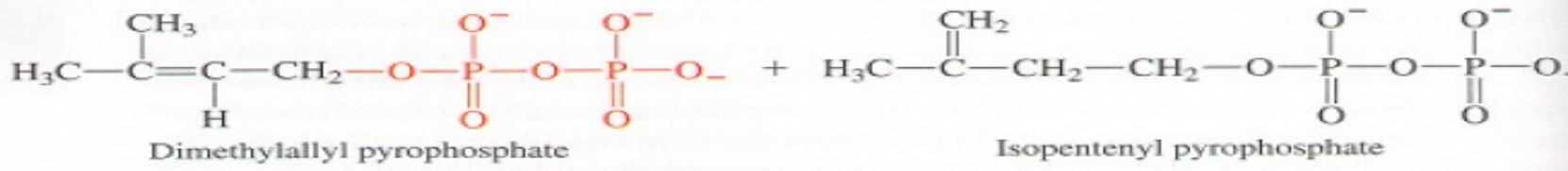
difosfomevalonát-dekarboxylasa

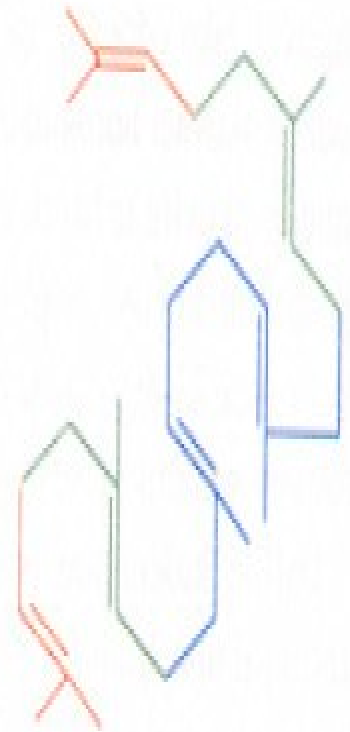
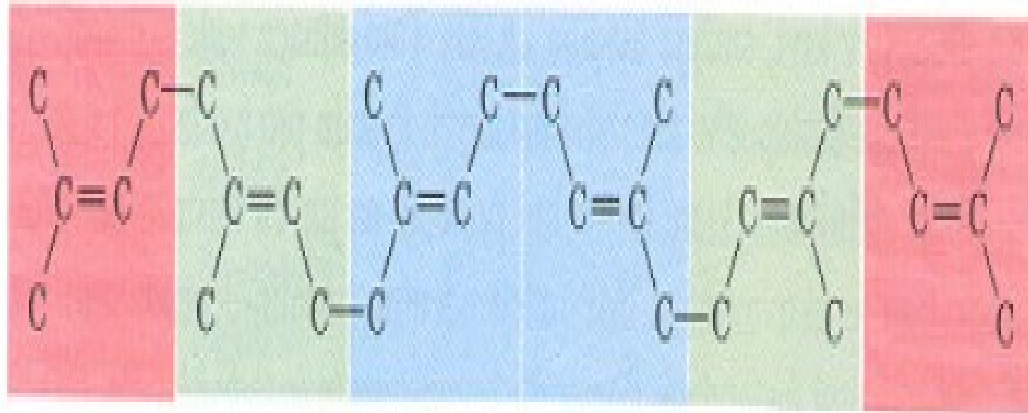
4

ATP → ADP + P<sub>i</sub> + CO<sub>2</sub>

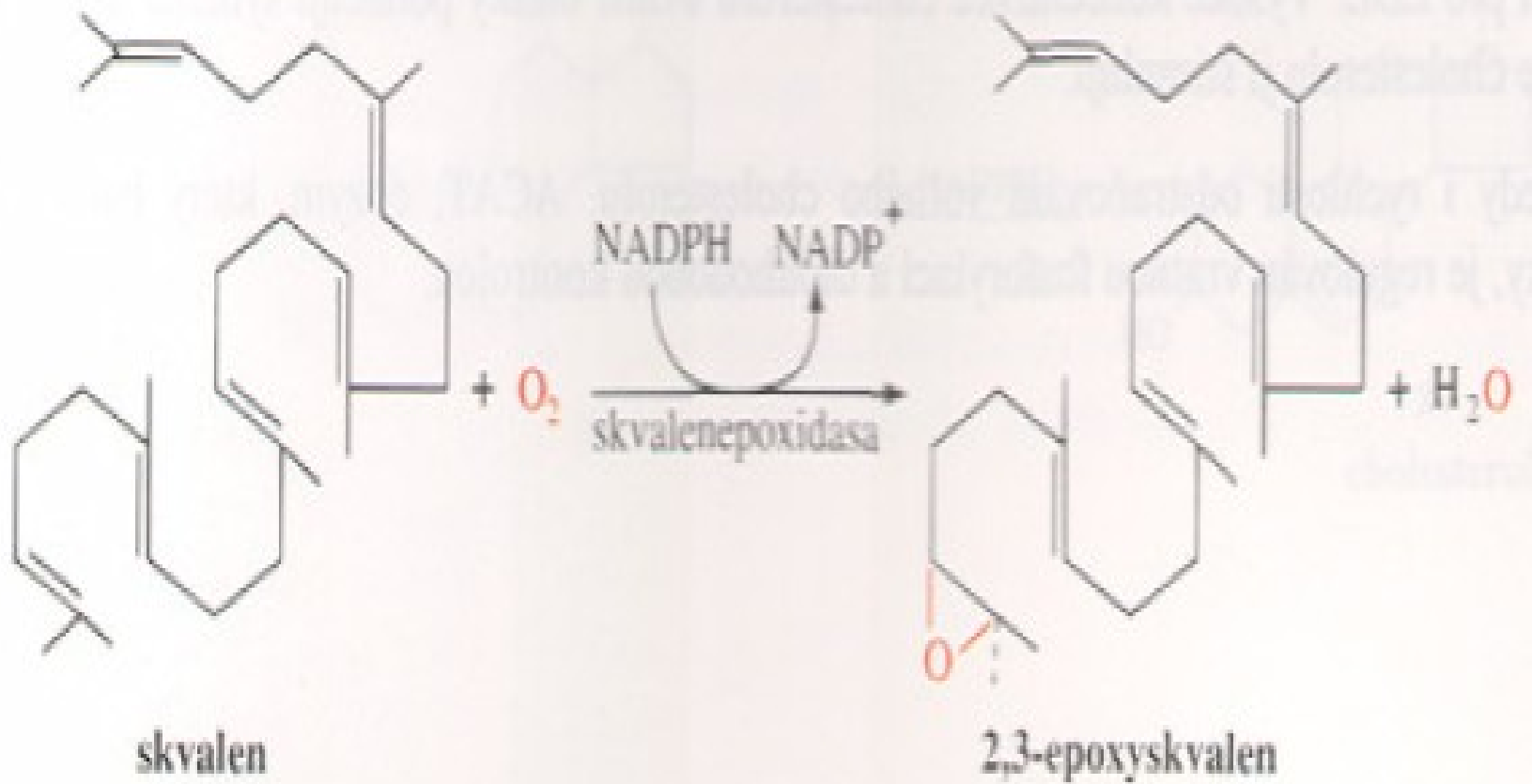


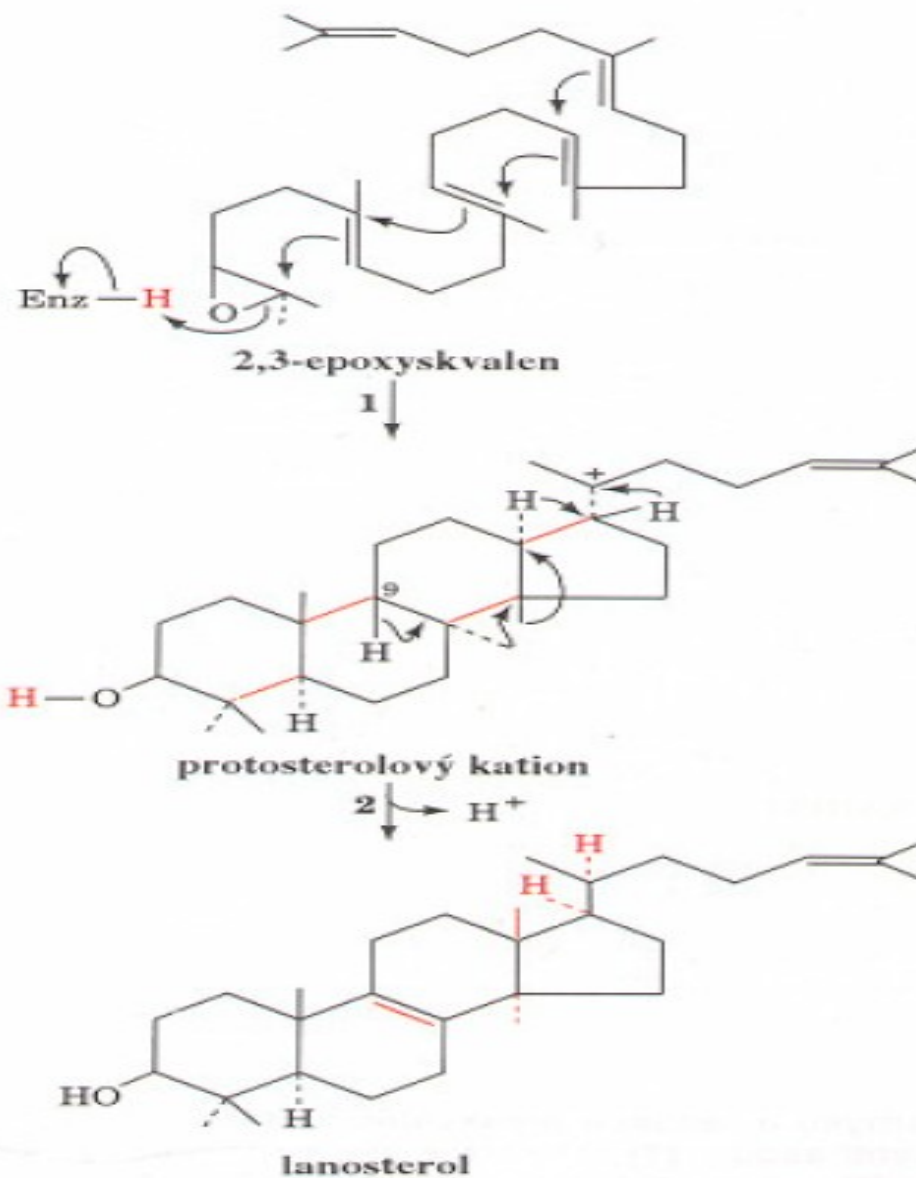


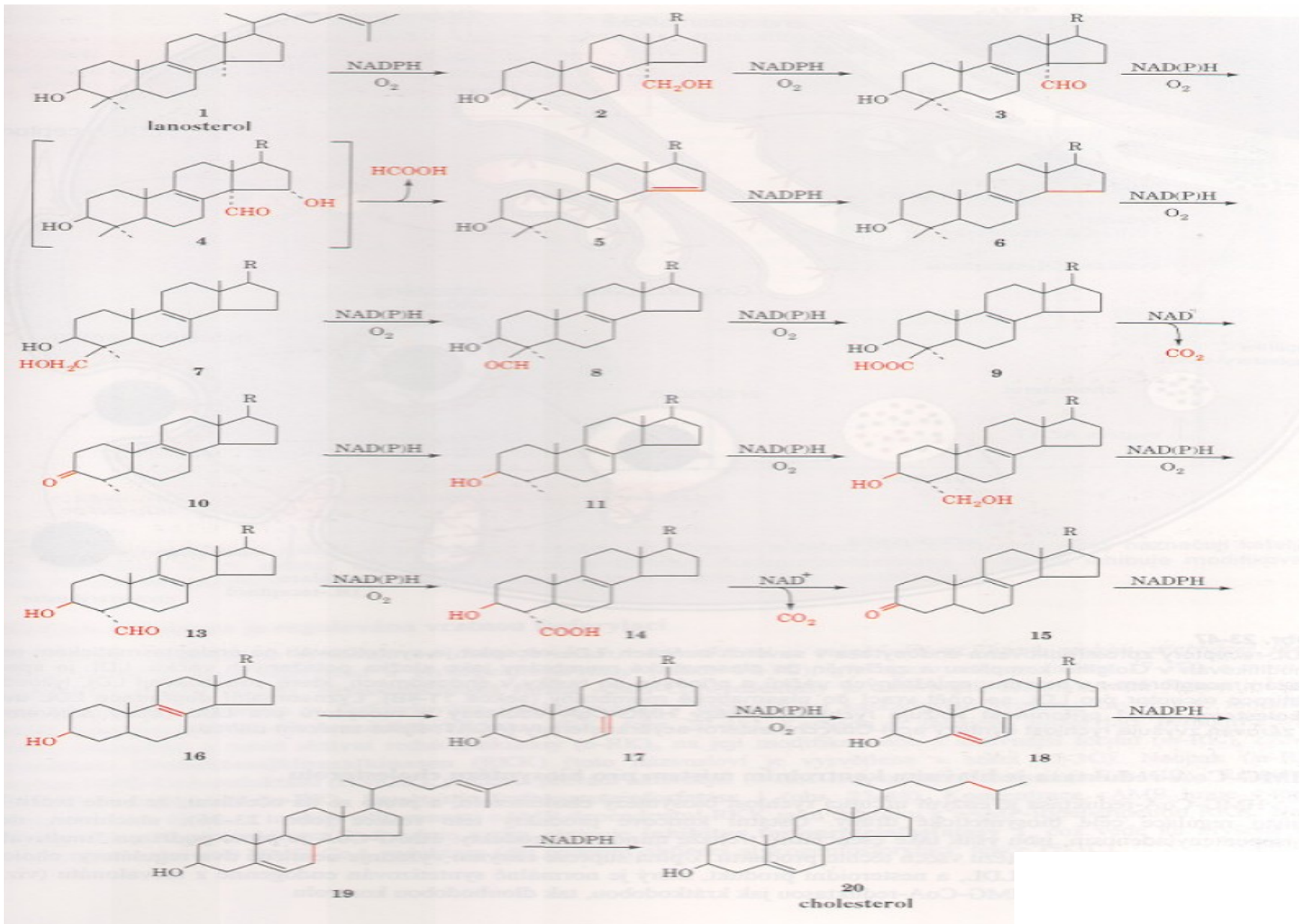












# Metabolismus bílkovin

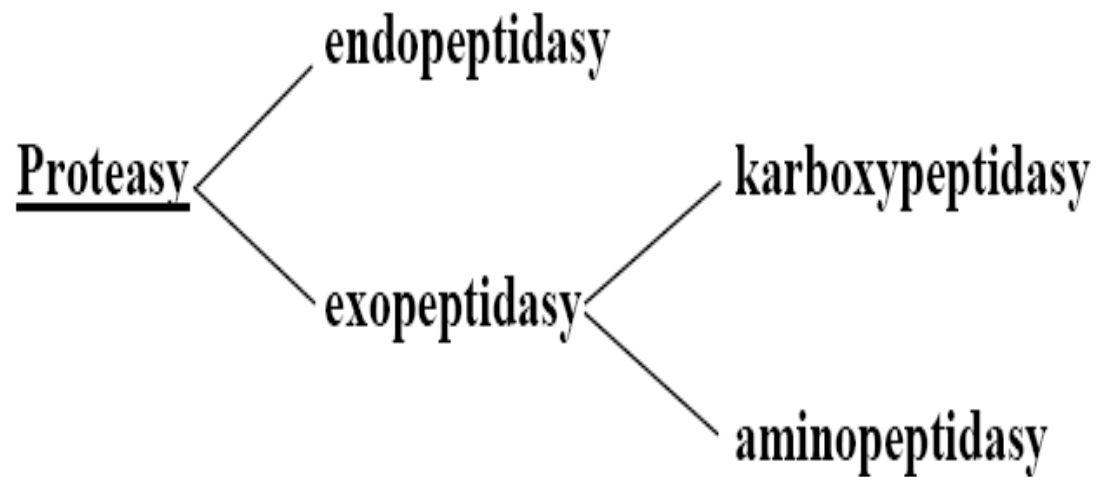
- Tuky, sacharidy – zásobárna energie, mohou se vzájemně zastupovat. Bílkoviny – tvorba tělních bílkovin, jsou zdrojem N pro heterotrofy
- V organismu neexistuje skladiště bílkovin
- U sacharidů (glykolýza, pentozový cyklus) a lipidů ( $\beta$  oxidace) jednotný metabolismus, AMK individuální metabolismus

# Metabolismus bílkovin

- AMK jsou prekurzory v různých metabolických drahách (puriny, pyrimidiny, protoporfiriny atd.)
- Biosyntéza bílkovin je geneticky řízena - proteosyntéza

# METABOLISMUS BÍLKOVIN

## Proteolýza





Proteasy - serinové

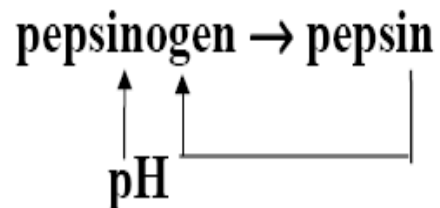
cysteinové

metaloproteasy

kyselé - aspartátové

## Aktivace zymogenů

*žaludek*



*slinivka břišní*

enterokinasa



trypsinogen → trypsin



chymotrypsinogen → chymotrypsin

proelastasa → elastasa

## 1. Žaludeční proteasy

- pepsin
- chymosin (renin, sýřidlo)

## 2. Pankreatické proteasy

- trypsin
- chymotrypsin
- elastasa
- karboxypeptidasa A,B

### 3. Proteasy střevní šťávy

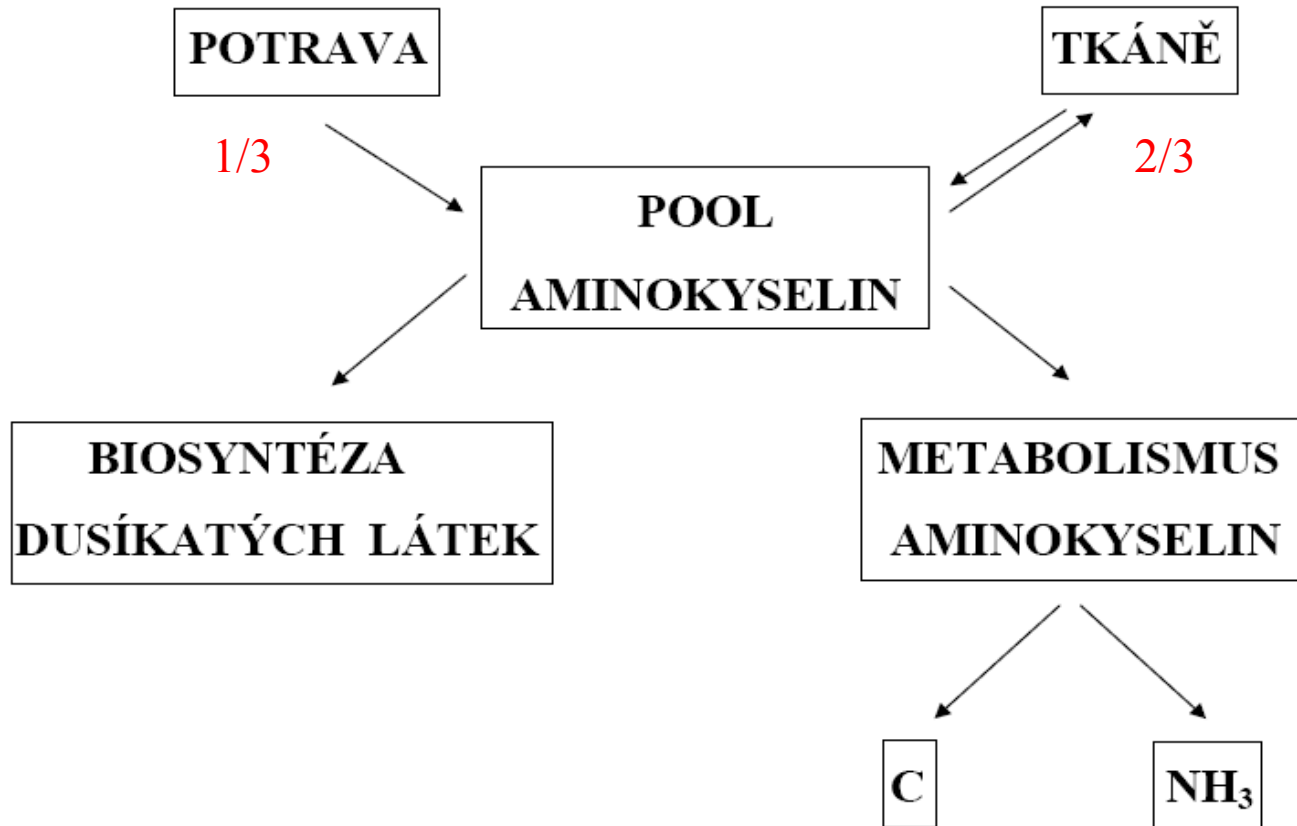
- aminopeptidasy
- dipeptidasy

### 4. Buněčné proteasy

- živočichové - kathepsiny B, D, L, H, M, S a T
- rostliny - papain
- bakterie - subtilisin, pronasa

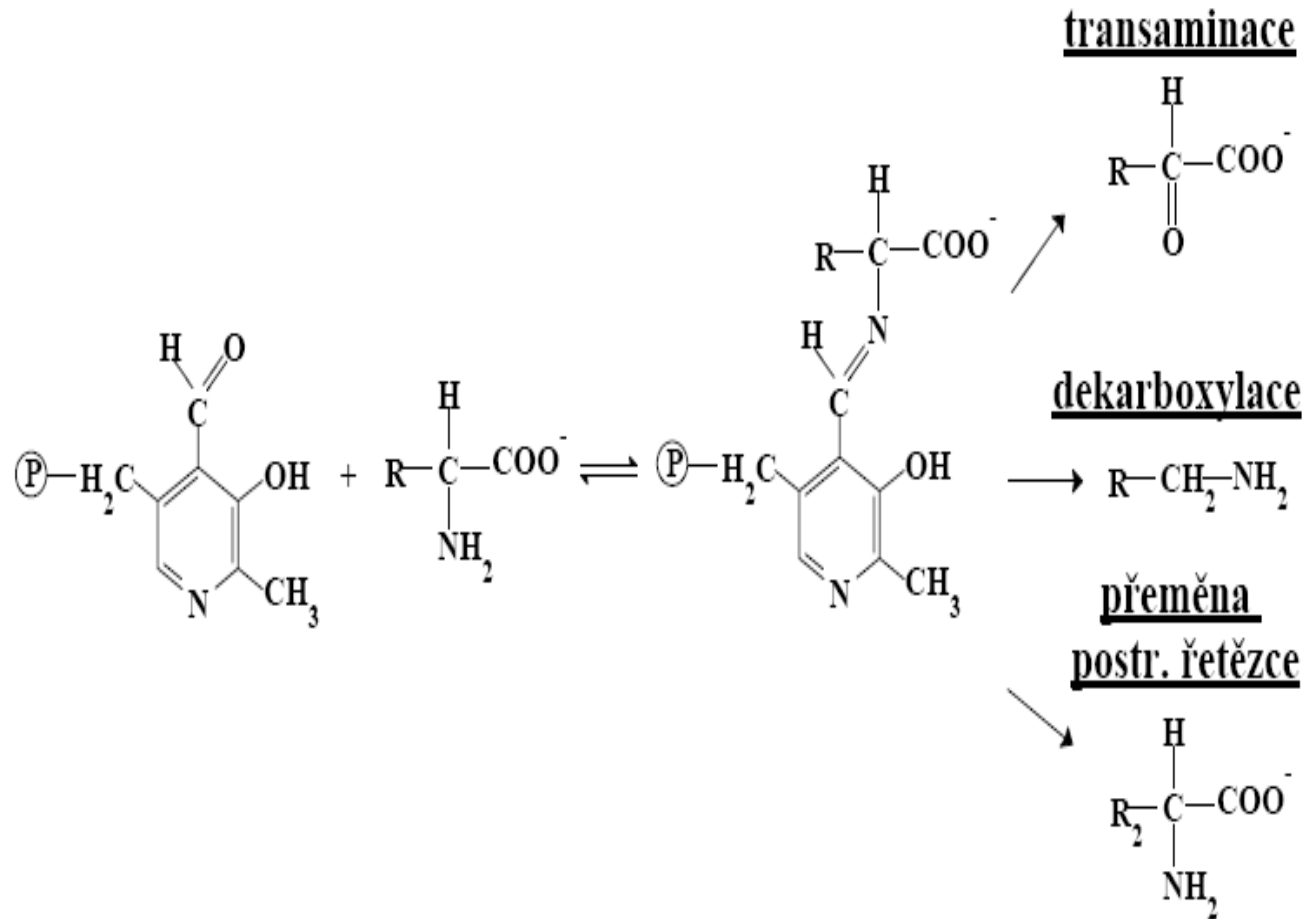
- Proteasy s jinou funkcí
- enterokinasa - aktivace zymogenů
- trombin - srážení krve

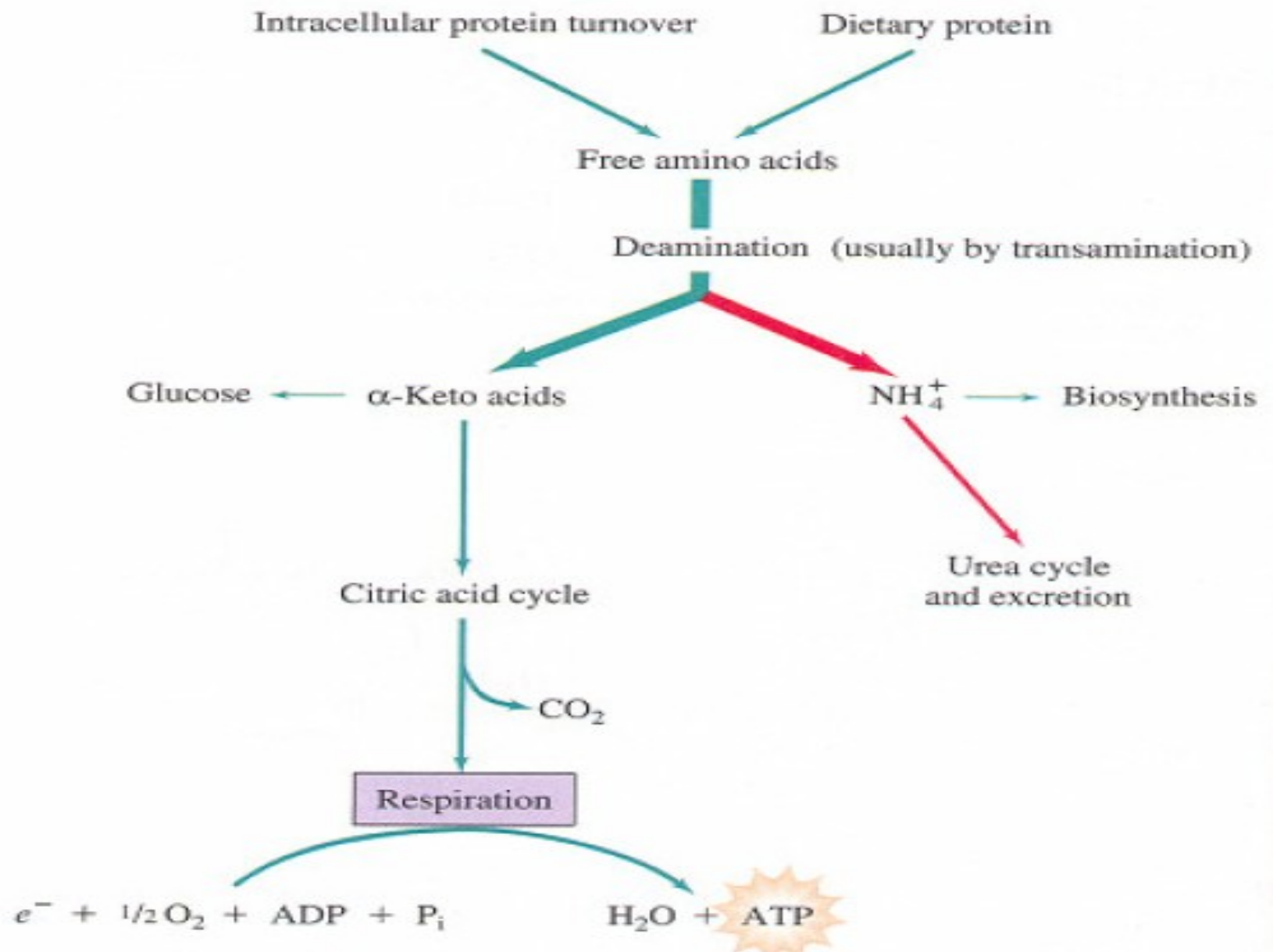
# Hotovost - pool aminokyselin



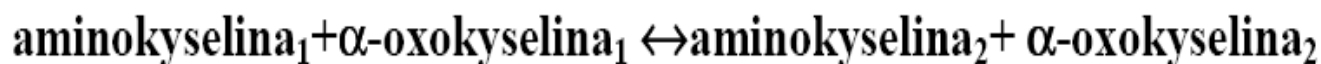


# METABOLISMUS AMINOKYSELIN



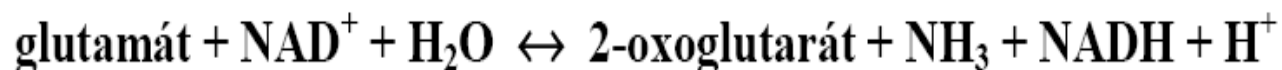


## Transaminace

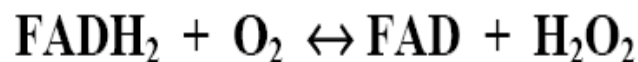
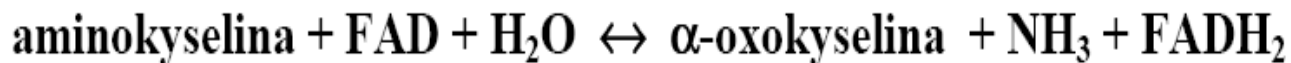


## Oxidační deaminace

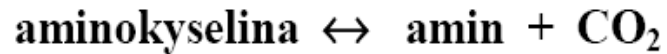
*savci*



*vejcorodí*

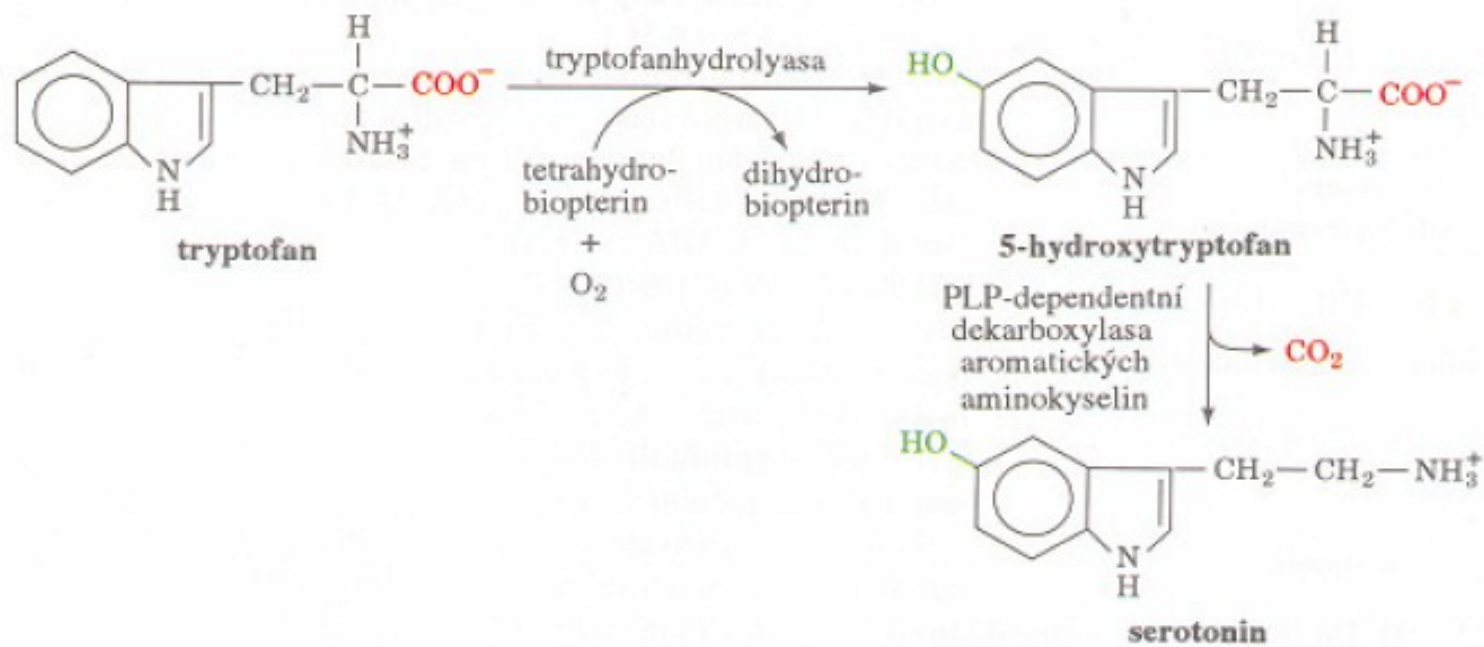
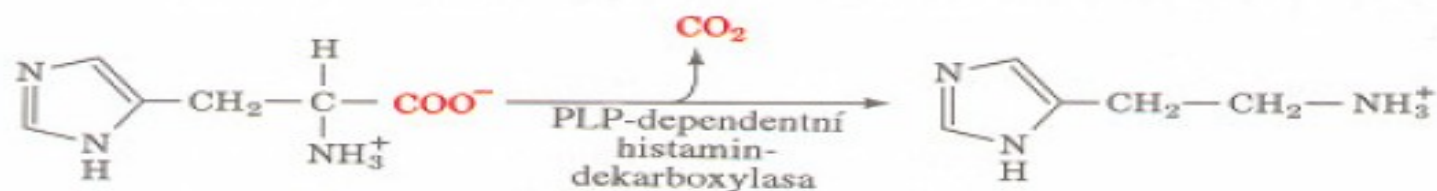
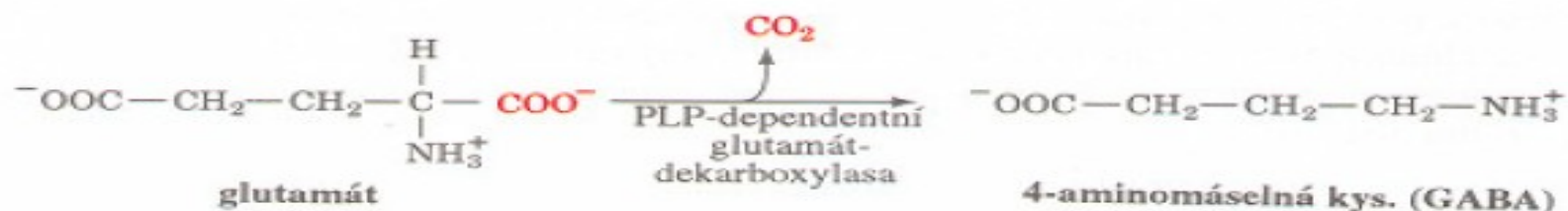


## Dekarboxylace



### Biogenní aminy

cystein	cystamin	CoA
k.asparagová	$\beta$ alanin	„
tyrosin	tyramin	tkáňový hormon
DOPA	dopamin	„
histidin	histamin	„
hydroxytryptofan	serotonin	„
k.glutamová	k. $\gamma$ -aminomáselná	neuromodulátor
serin	ethanolamin	fosfolipidy
methionin	spermin, spermidin	sperma



## Degradace uhíkových koster aminokyselin

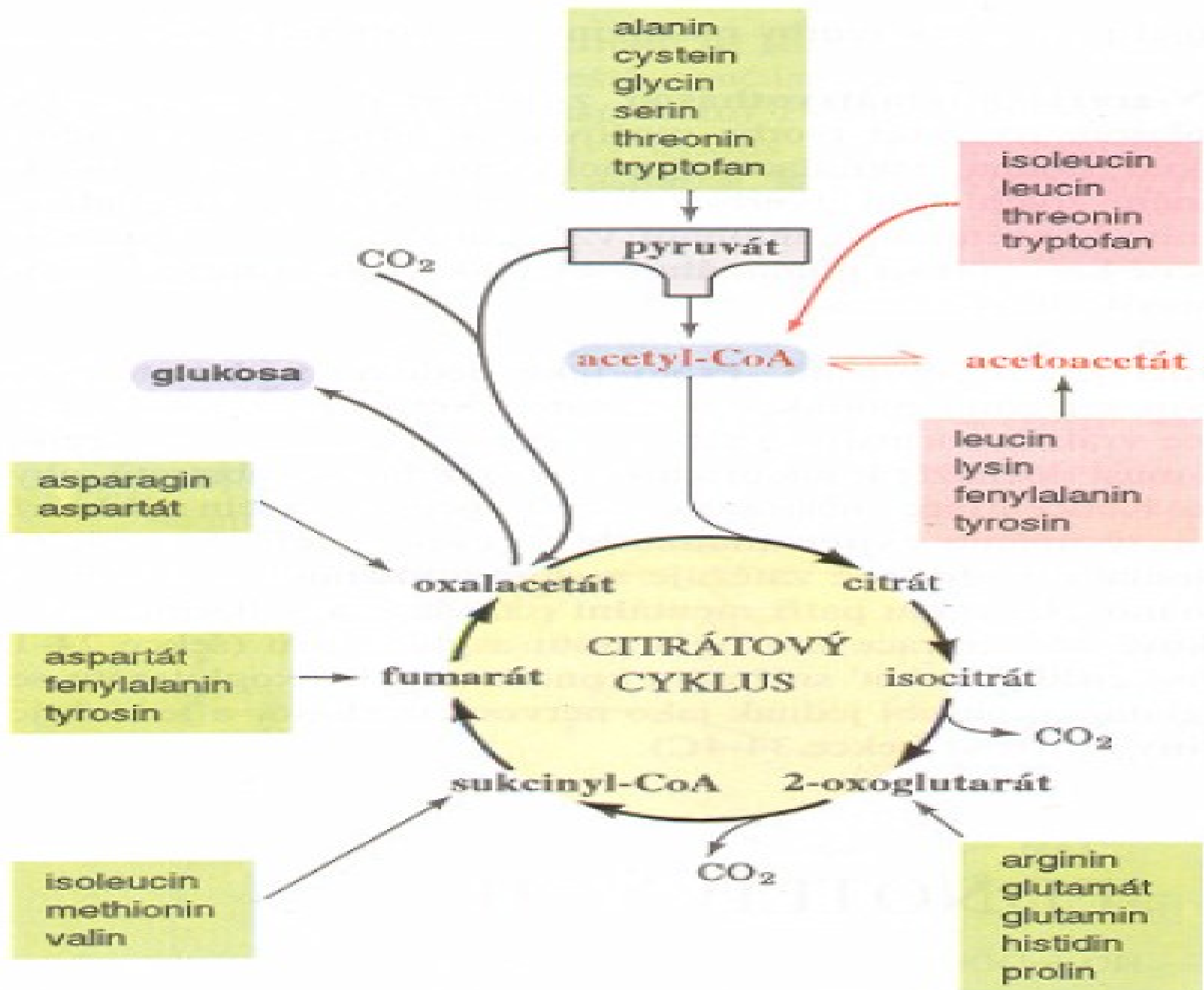
### 1. Glukogenní aminokyseliny - prekurzory sacharidů

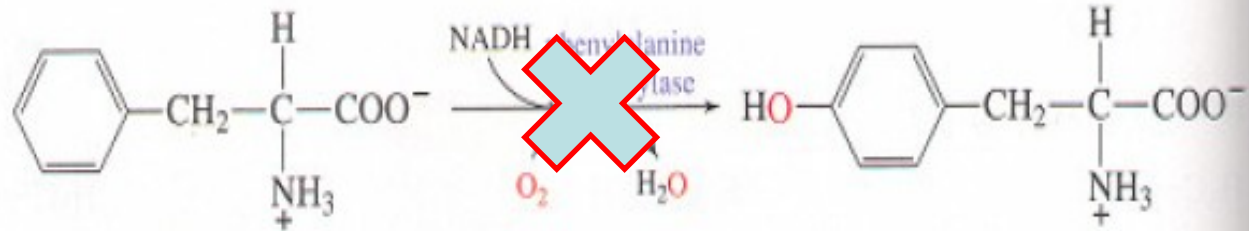
pyruvát	-	Ser, Ala, Cys, Gly, Thr, Met, Trp
2-oxoglutarát	-	Glu, Gln, Arg, Pro, His
oxalacetát	-	Asp, Asn
fumarát	-	Phe, Tyr
sukcinyl-CoA	-	Val, Ile, Met, Thr

### 2. Ketogenní aminokyseliny - prekurzory mastných kyselin

acetoacetát	-	Leu, Phe, Tyr, Lys, Trp
acetyl-CoA	-	Leu, Ile, Trp





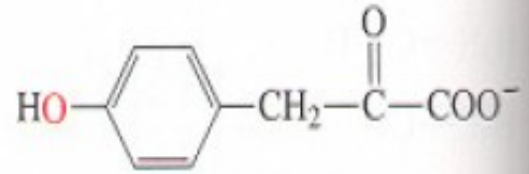


Phenylalanine

Fenylketonurie

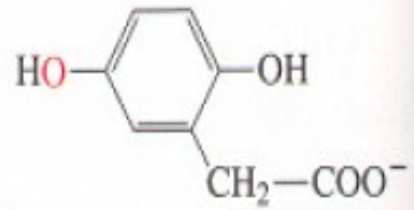
Tyrosine

aminotransferase



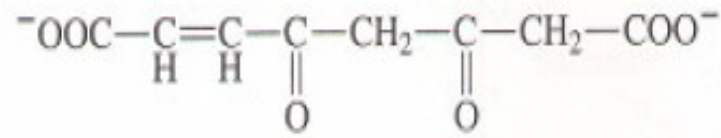
*p*-Hydroxyphenylpyruvate

↓



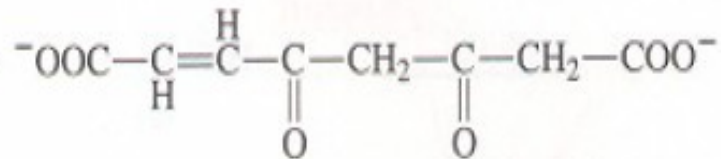
Homogentisate

homogentisate oxidase



4-Maleylacetoacetate

↑

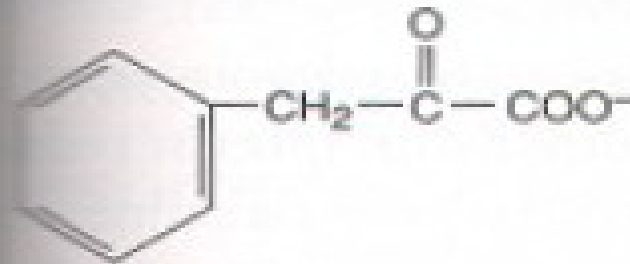


4-Fumarylacetoacetate

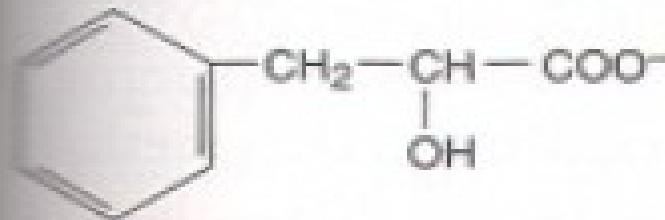
Fumarate + Acetoacetate

←

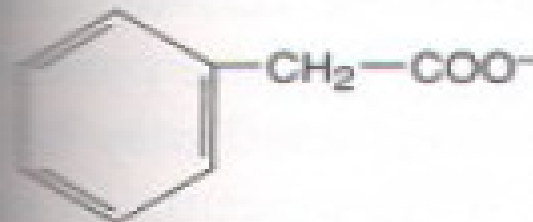
# Fenylketonurie



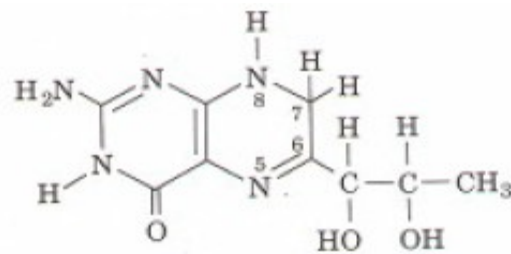
**Phenylpyruvate**



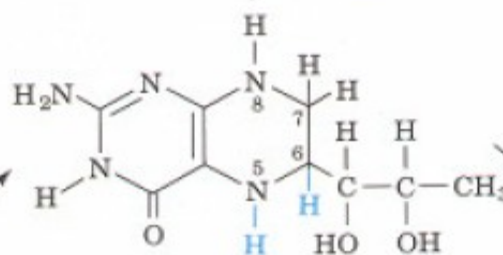
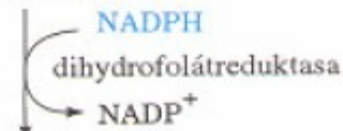
**Phenyllactate**



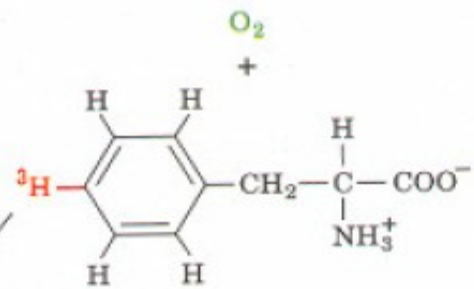
**Phenylacetate**



7,8-dihydrobiopterin

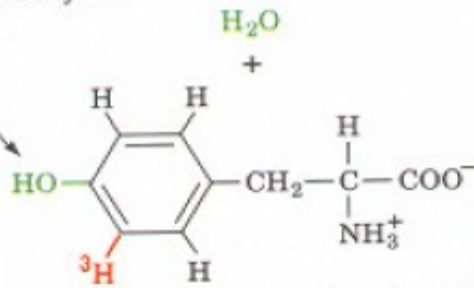


5,6,7,8,-tetrahydrobiopterin

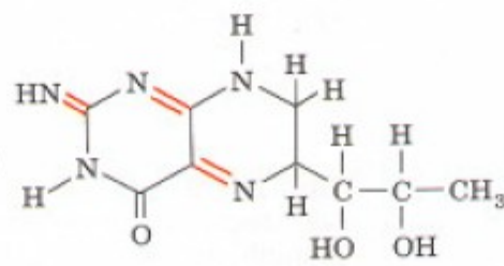


fenylalanin

fenylalanin-hydroxylasa



tyrosin

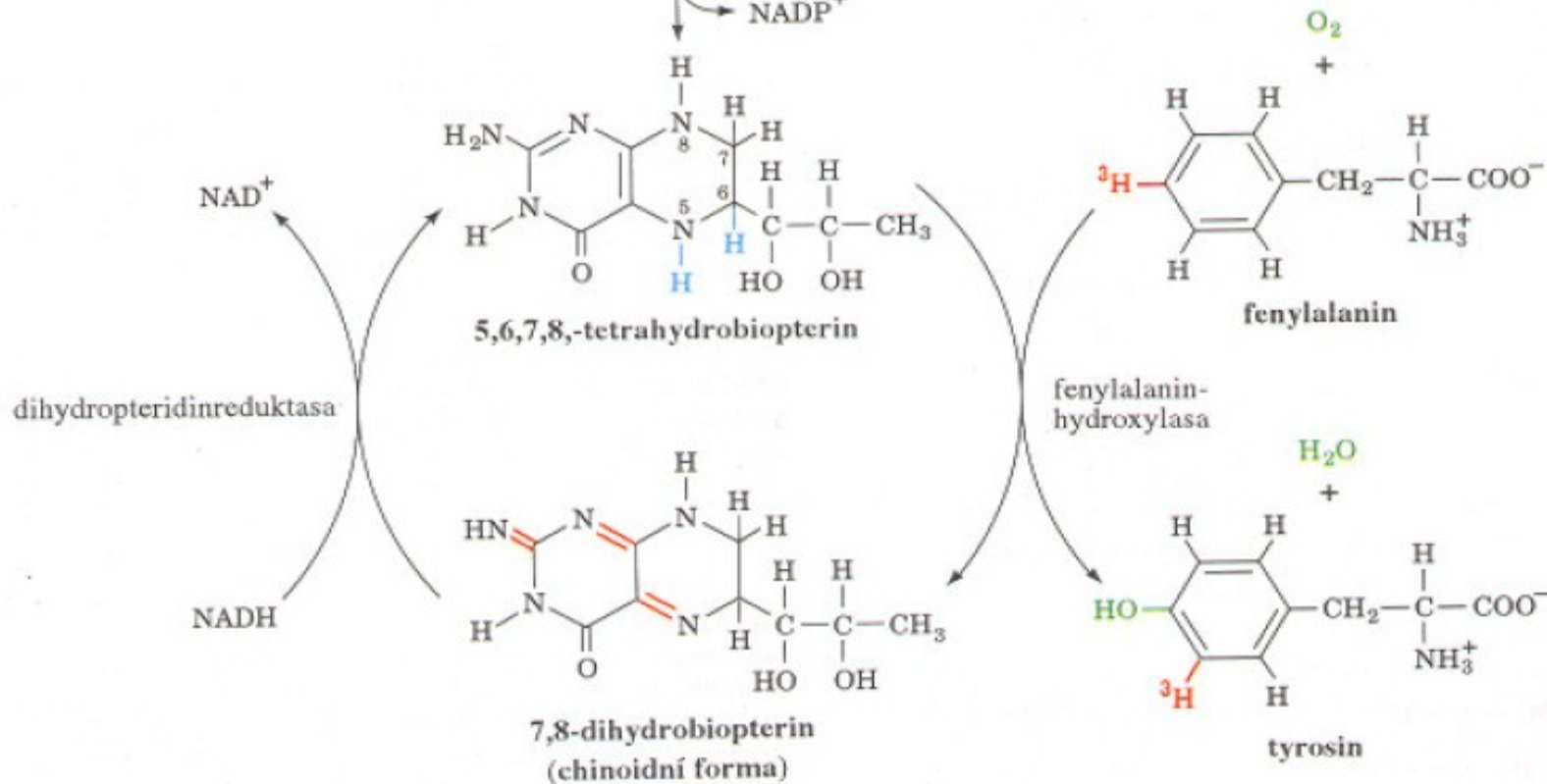


7,8-dihydrobiopterin (chinoidní forma)

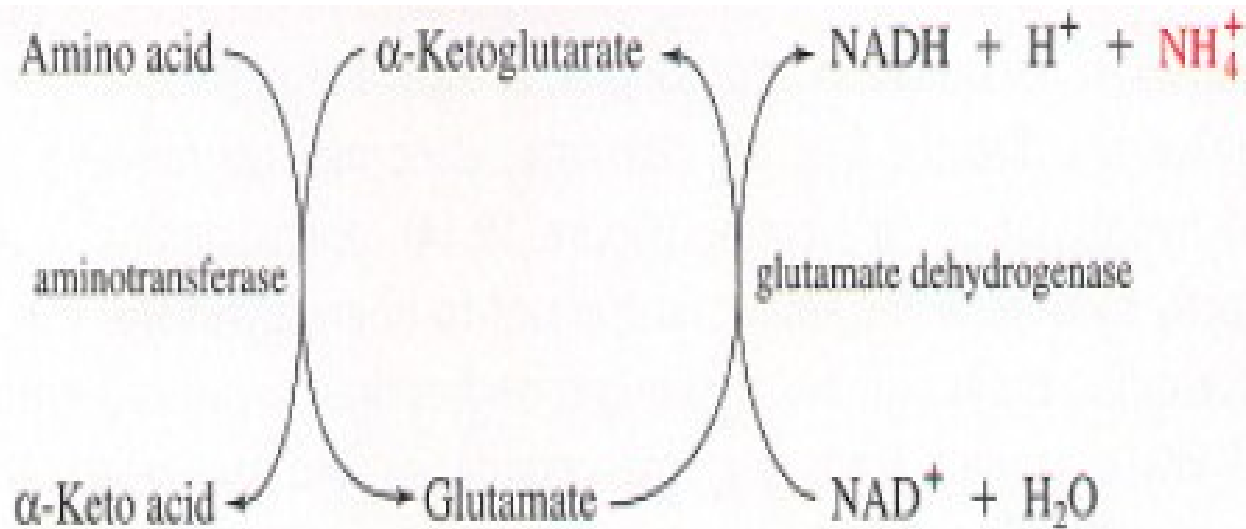
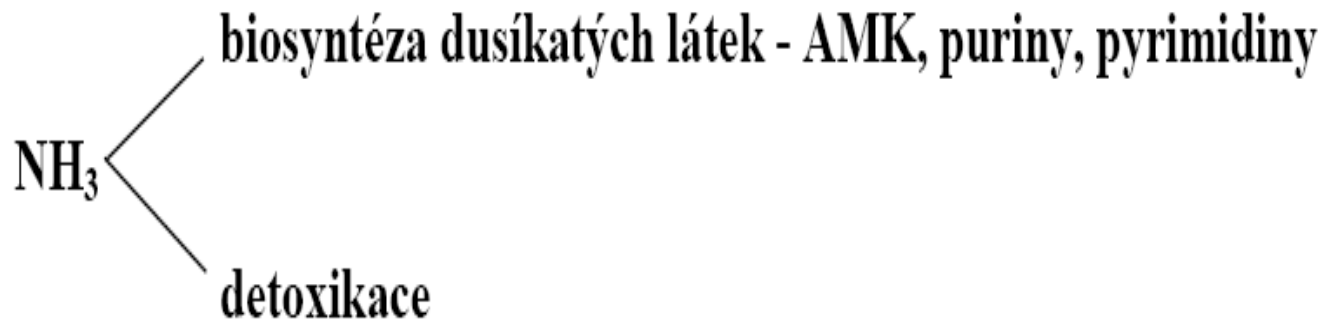
dihydropteridinreduktasa

NAD<sup>+</sup>

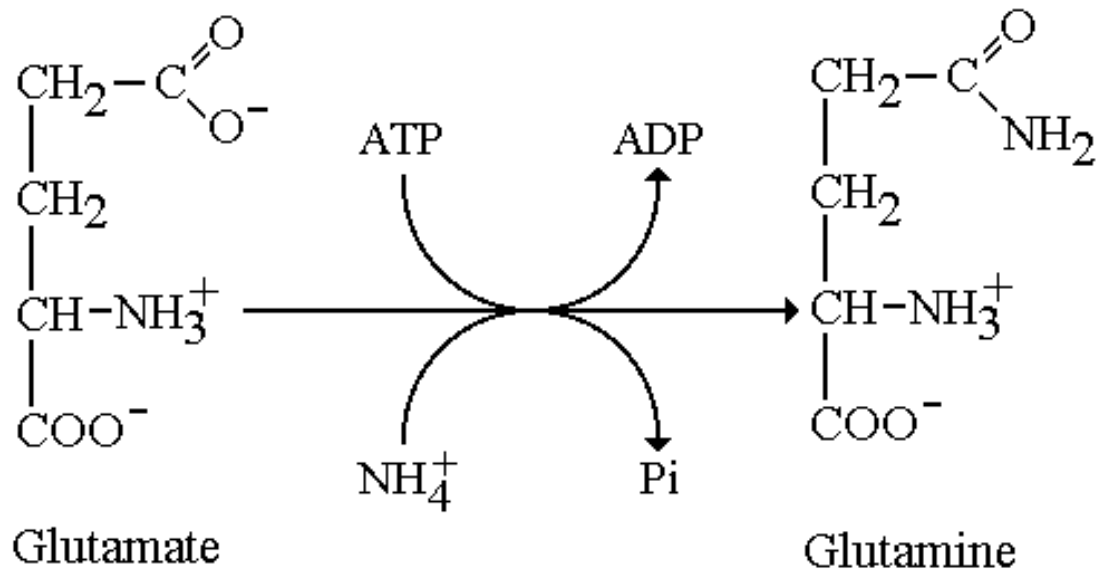
NADH



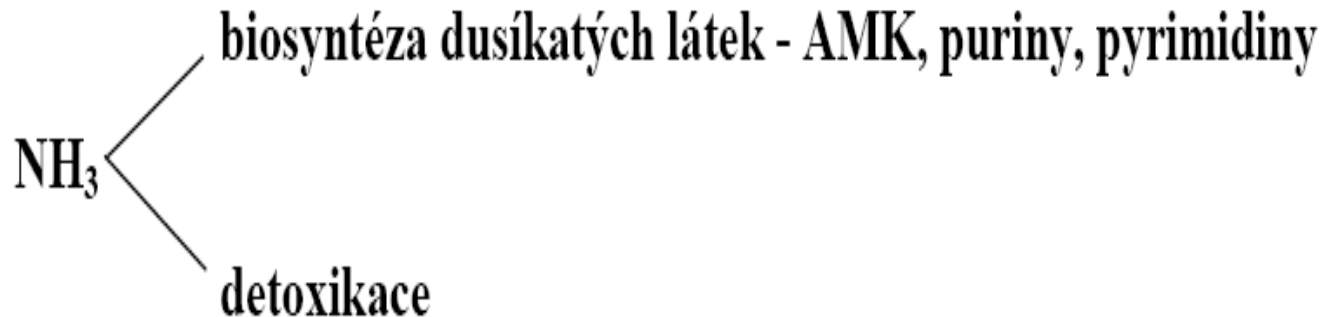
## Metabolismus amoniaku



# Glutamin syntethasa



## Metabolismus amoniaku



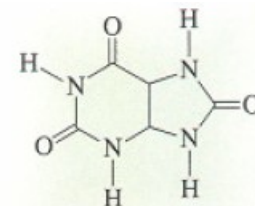
*Živočichové* - amonotelní -  $\text{NH}_3$  - vodní živočichové

- urikotelní - k.močová - vejcorodí

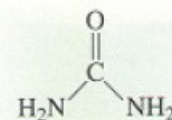
- ureotelní - močovina - placentálové

*Rostliny* - nevylučují  $\text{NH}_3$

$\text{NH}_4^+$   
(c) Ammonium ion



(b) Uric acid

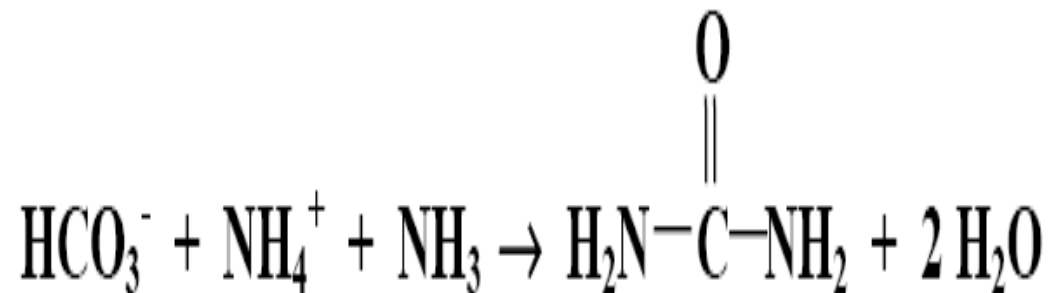


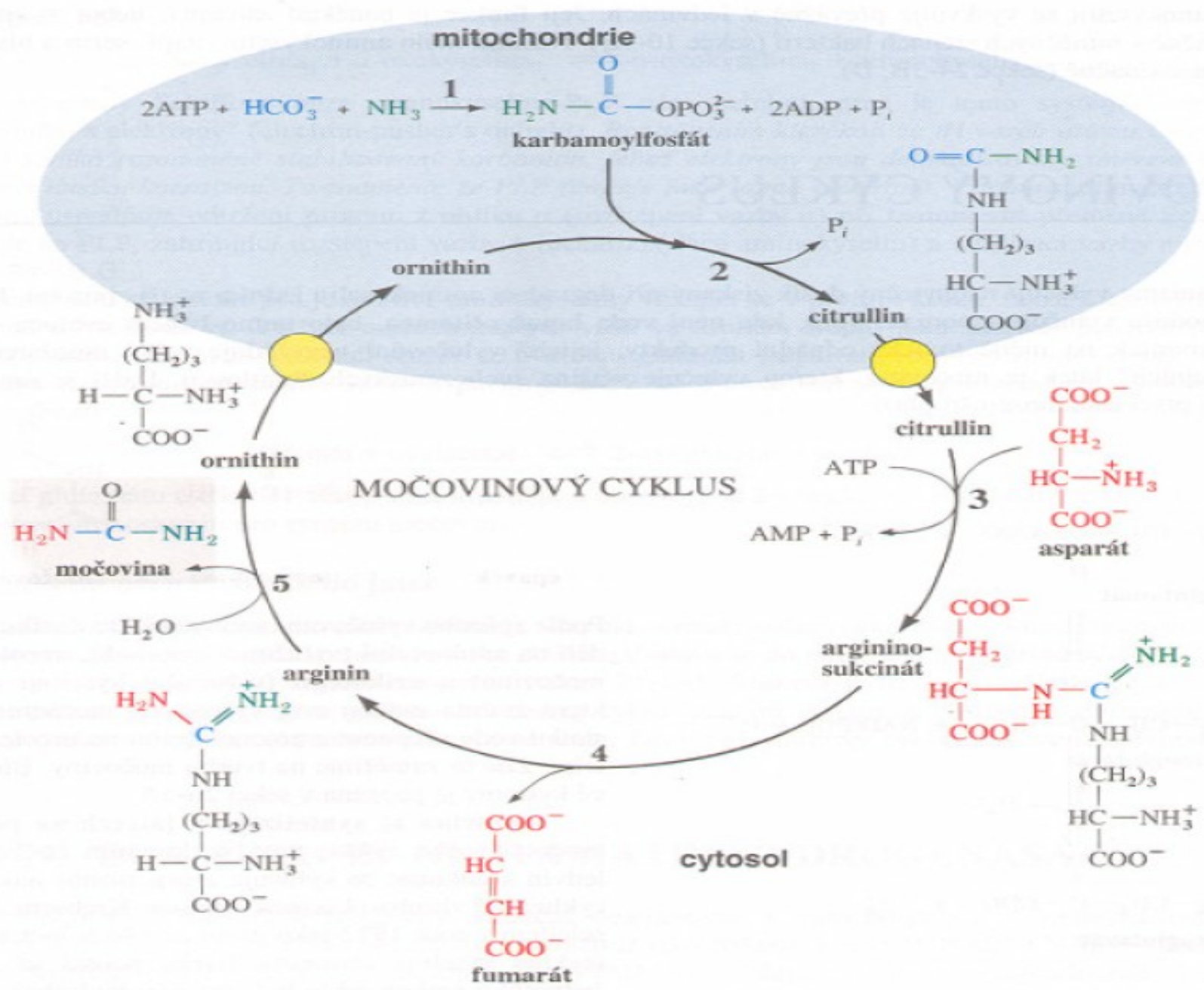
(a) Urea

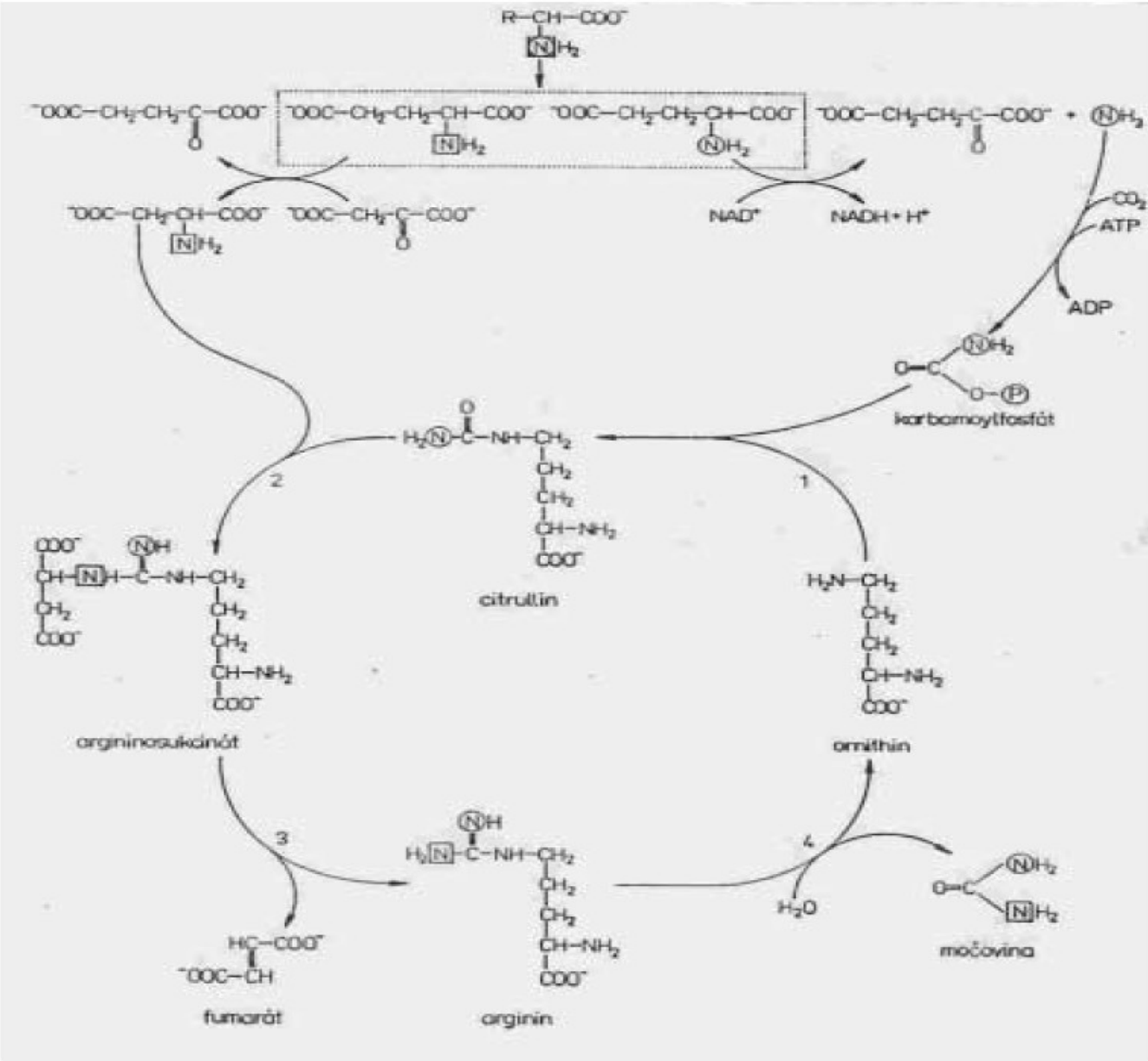


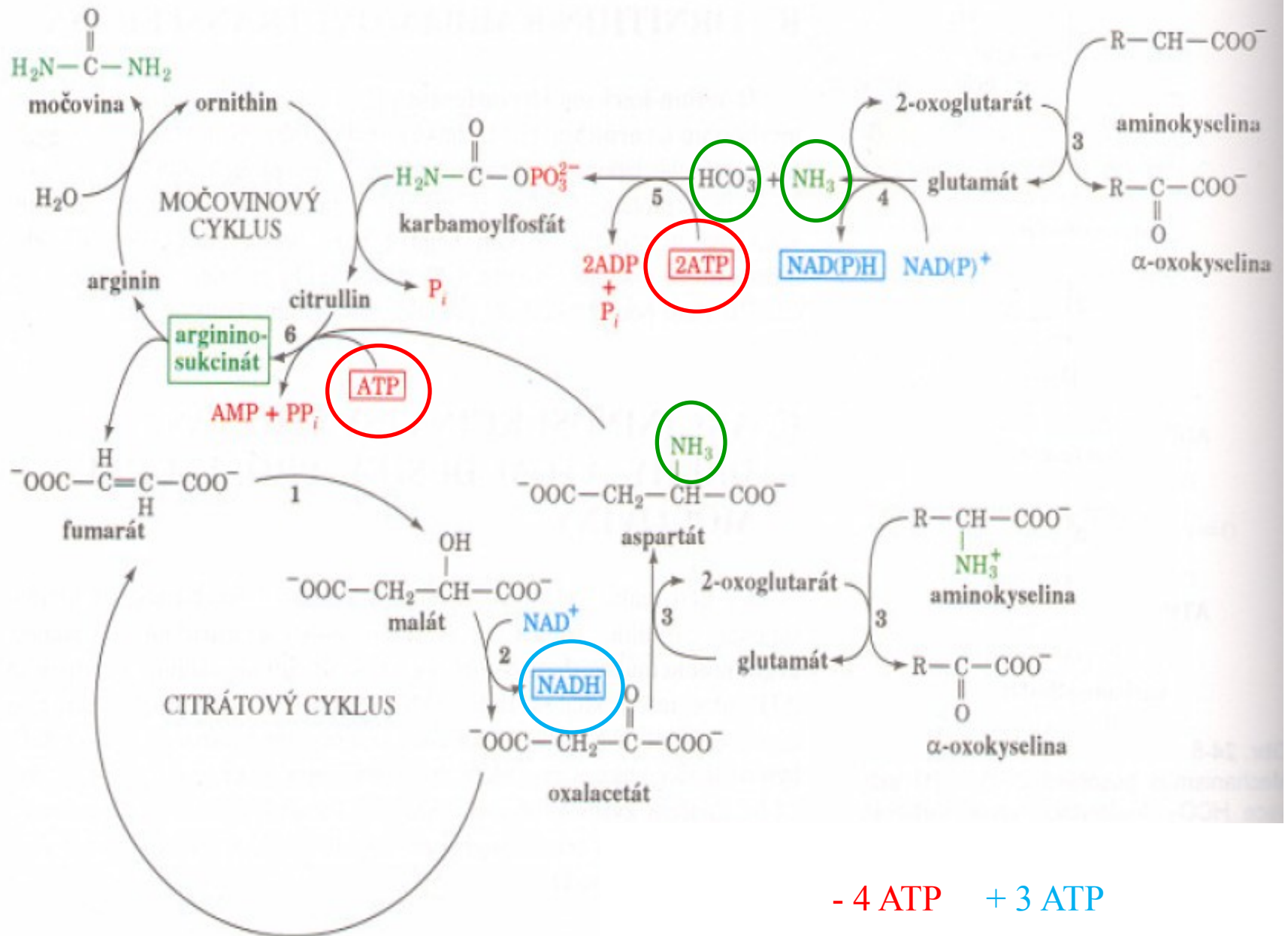
## Tvorba močoviny - ornitinový cyklus

H. KREBS, K. HENSELEIT - 1932

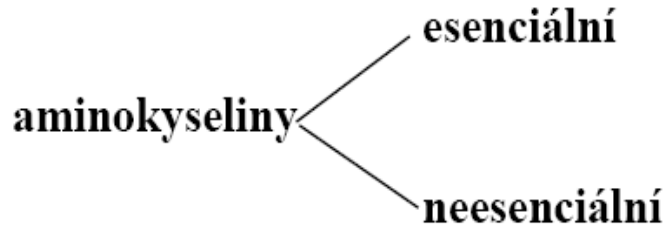






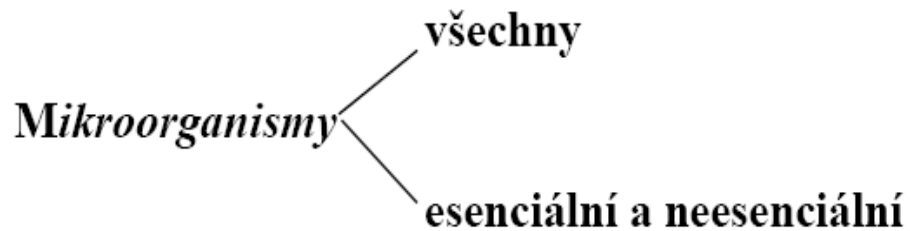


## Biosyntéza aminokyselin



*Rostliny* - všechny

$\text{NH}_2$ ,  $\text{NH}_4^+$ ,  $\text{NO}_3^-$ ,  $\text{NO}_2^-$ ,  $\text{N}_2$

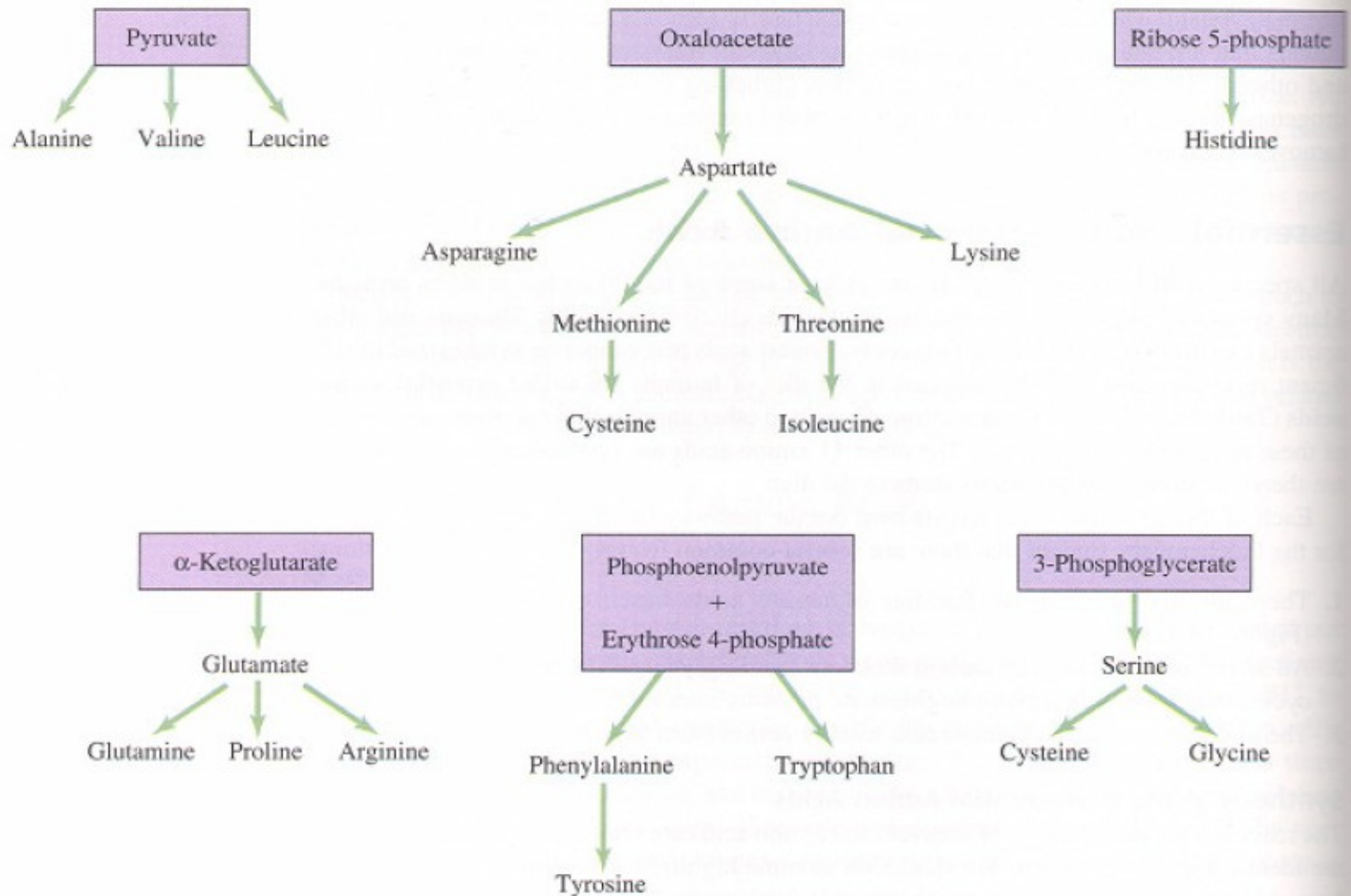


$\text{NH}_2$ ,  $\text{NH}_4^+$ ,  $\text{NO}_3^-$ ,  $\text{NO}_2^-$ ,  $\text{N}_2$

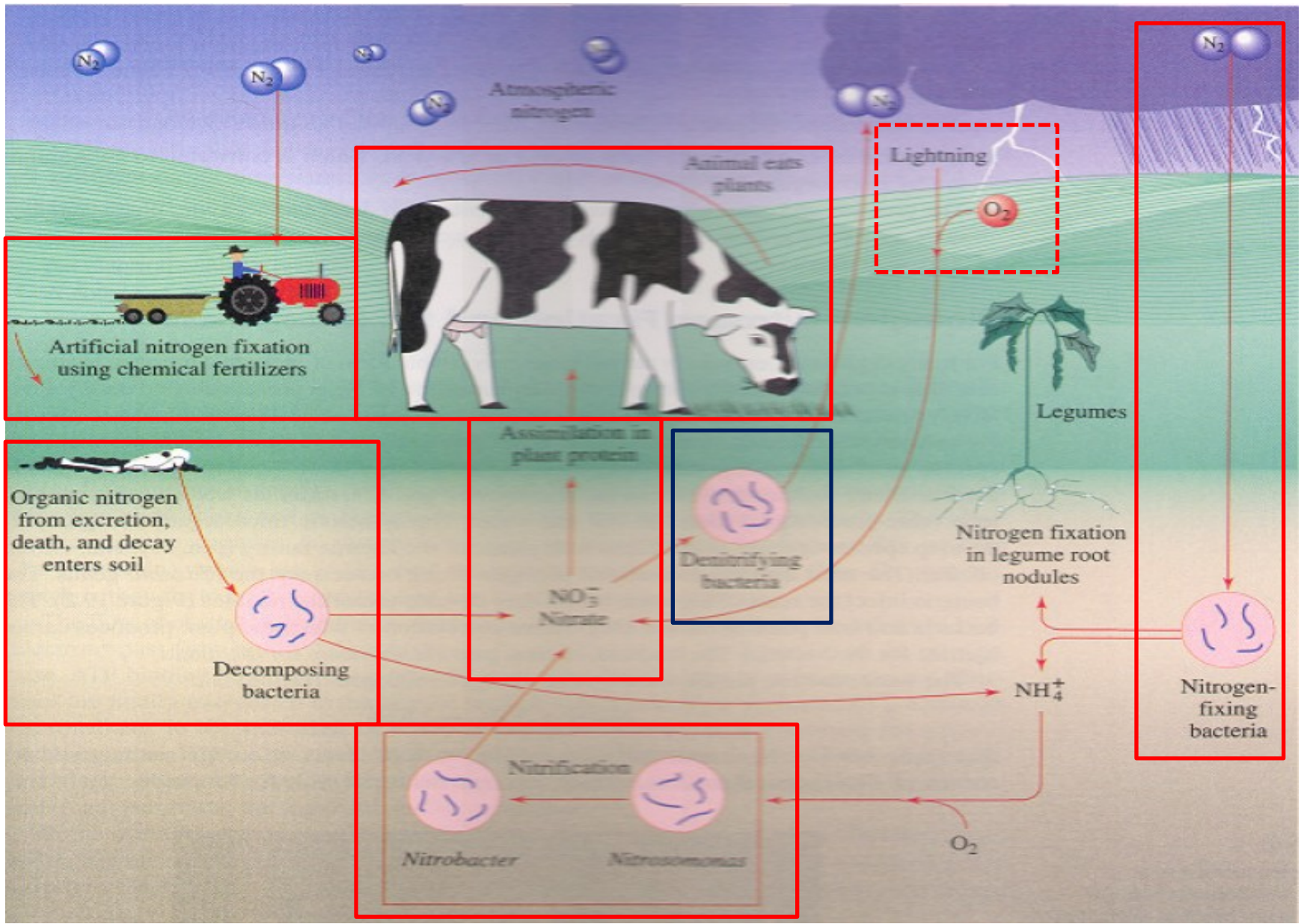
*Živočichové* - esenciální a neesenciální

$\text{NH}_2$

## Biosyntéza aminokyselin



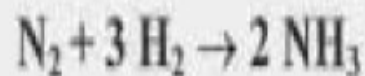




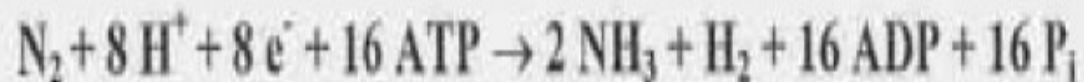


## Fixace N<sub>2</sub>

Chemická syntéza - Haber Bosch (500 °C, 300 atm, kat – Fe)

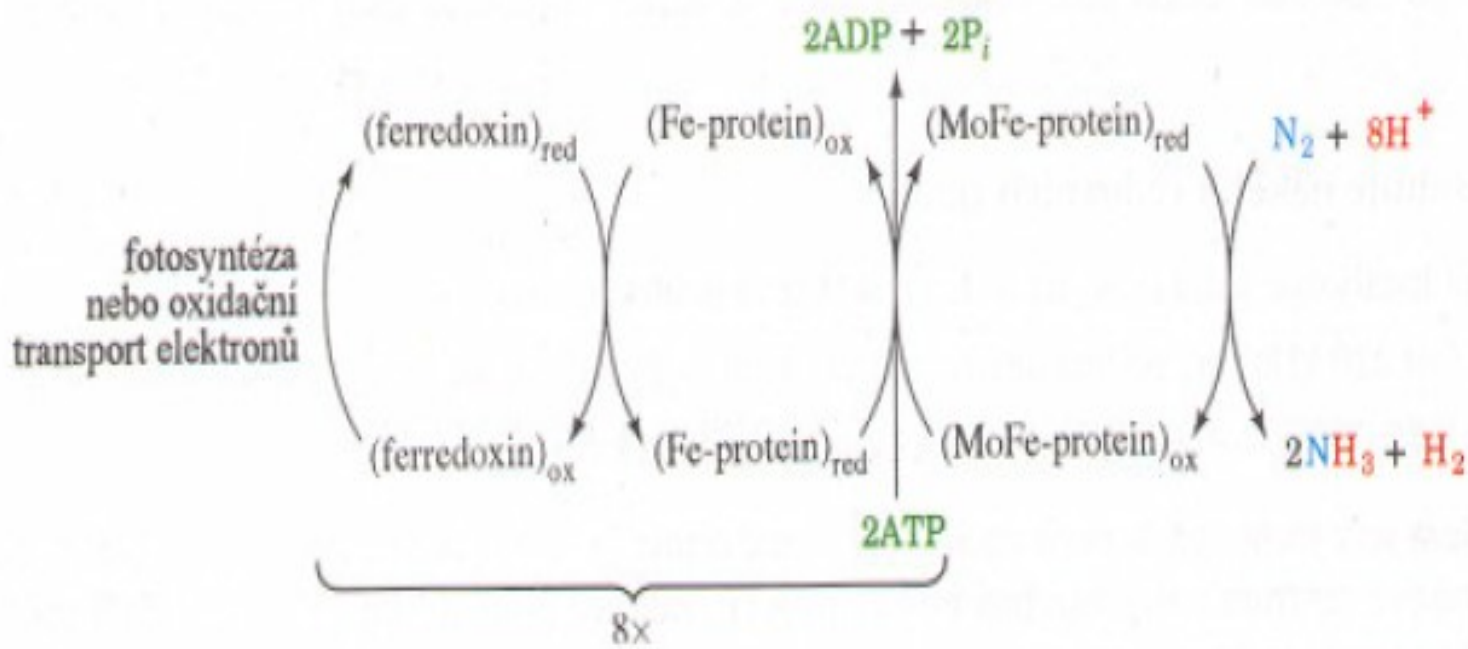
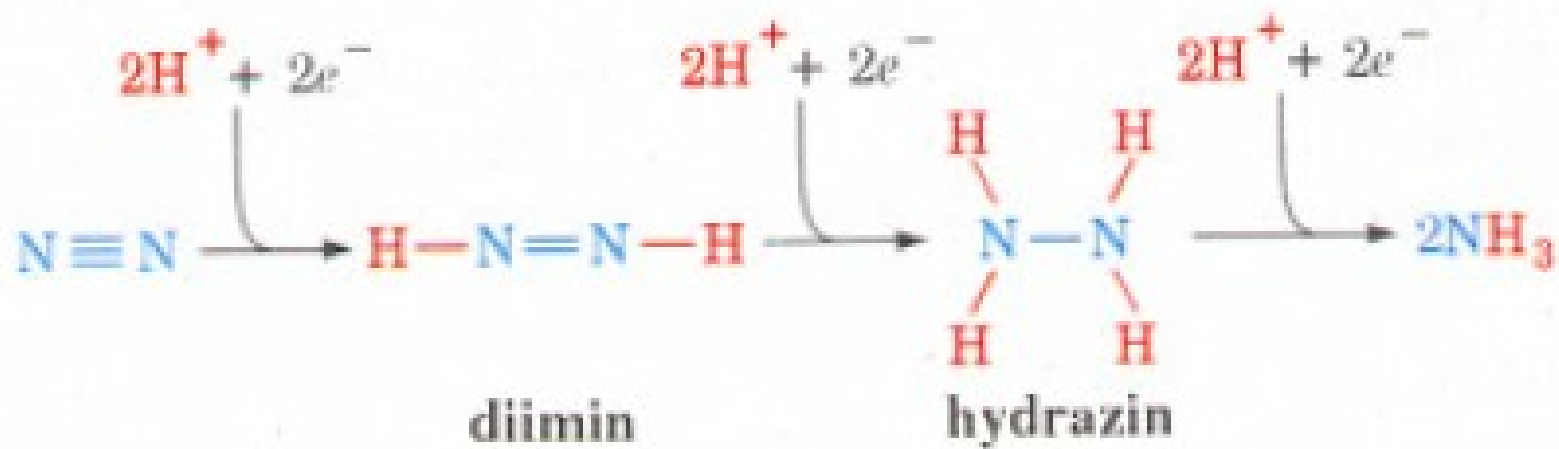


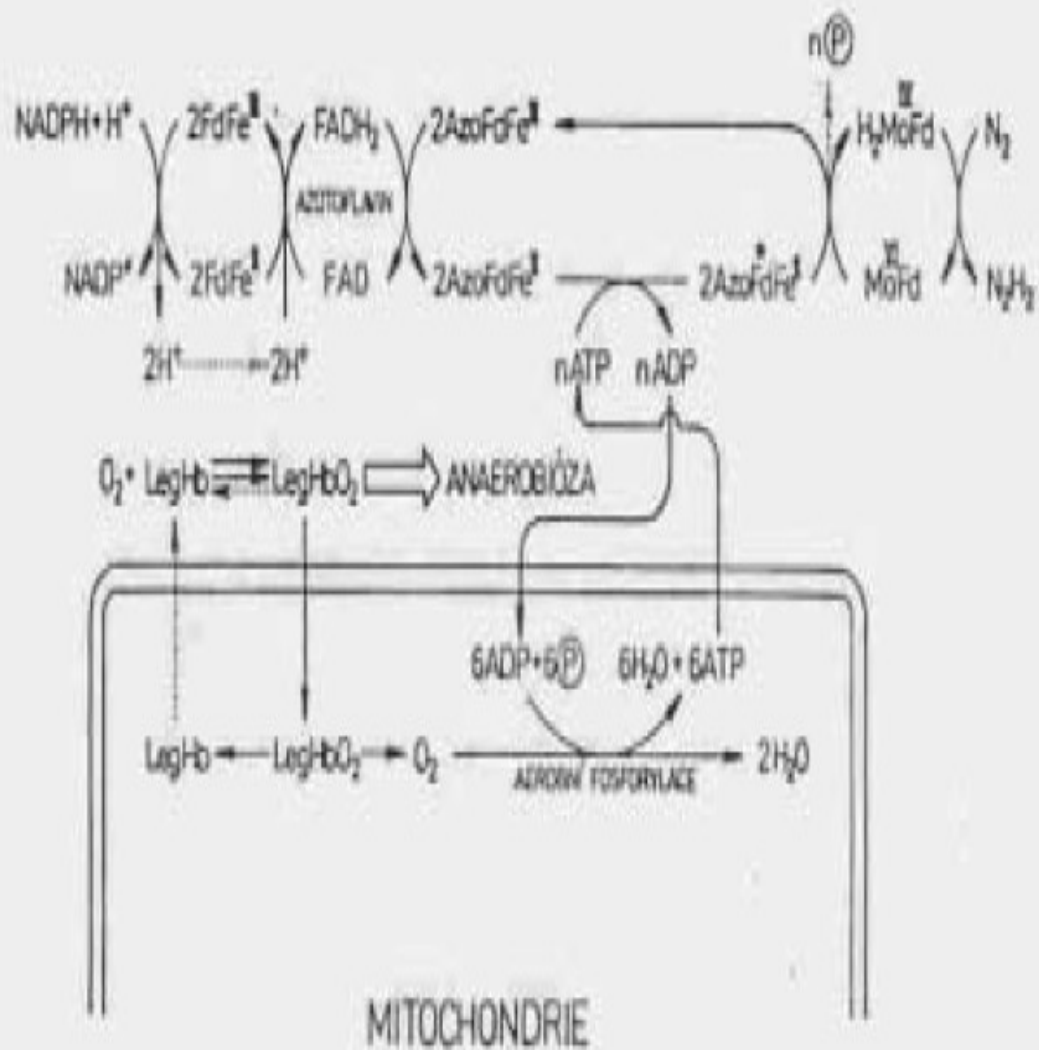
Biosyntéza – sinice, bakterie - *Rhizobium*, *Azotobacter*



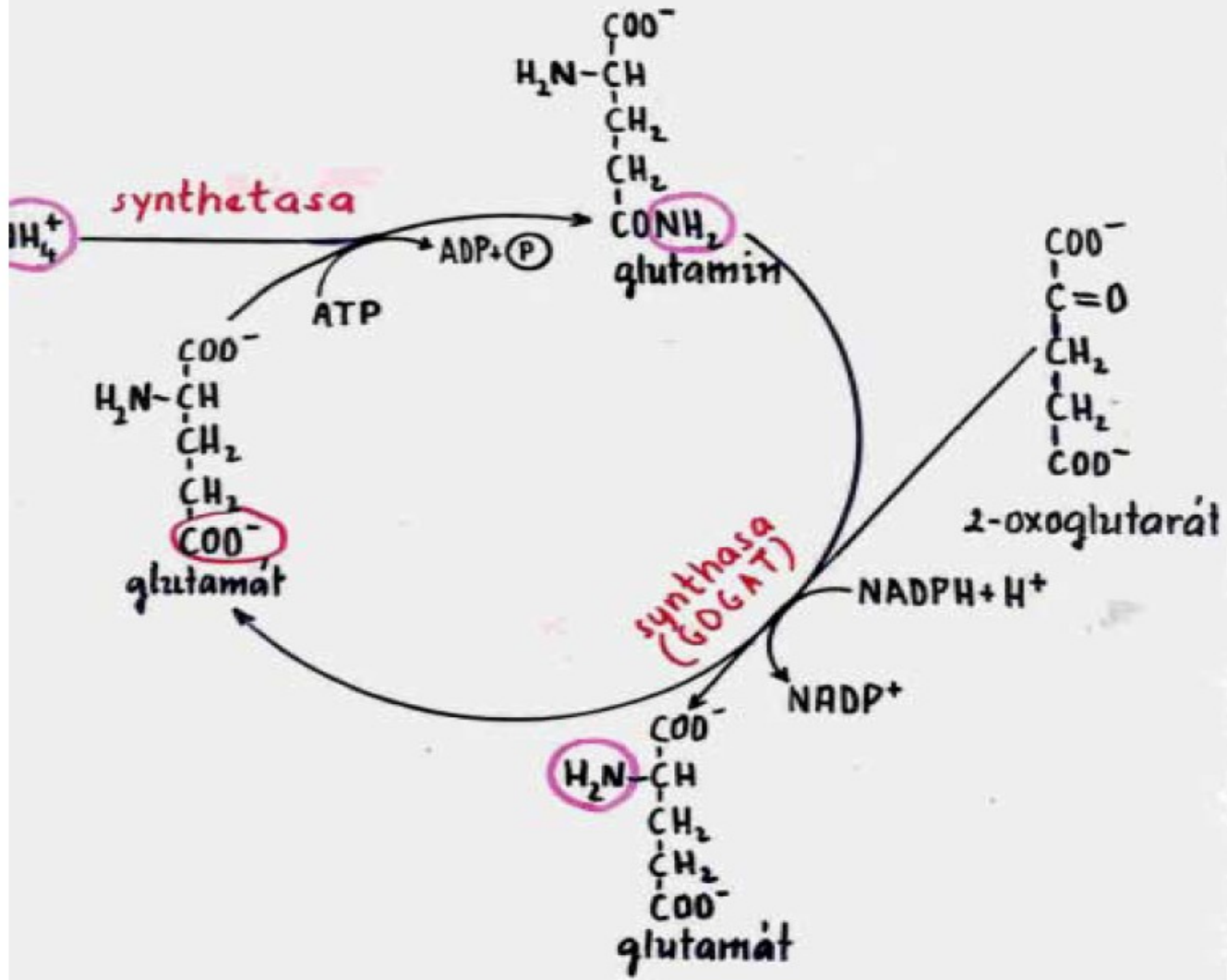
Nitrogenasa : 1. protein Fe  
2. protein MoFe







# INKORPORACE NH<sub>3</sub> U PROKARYOT



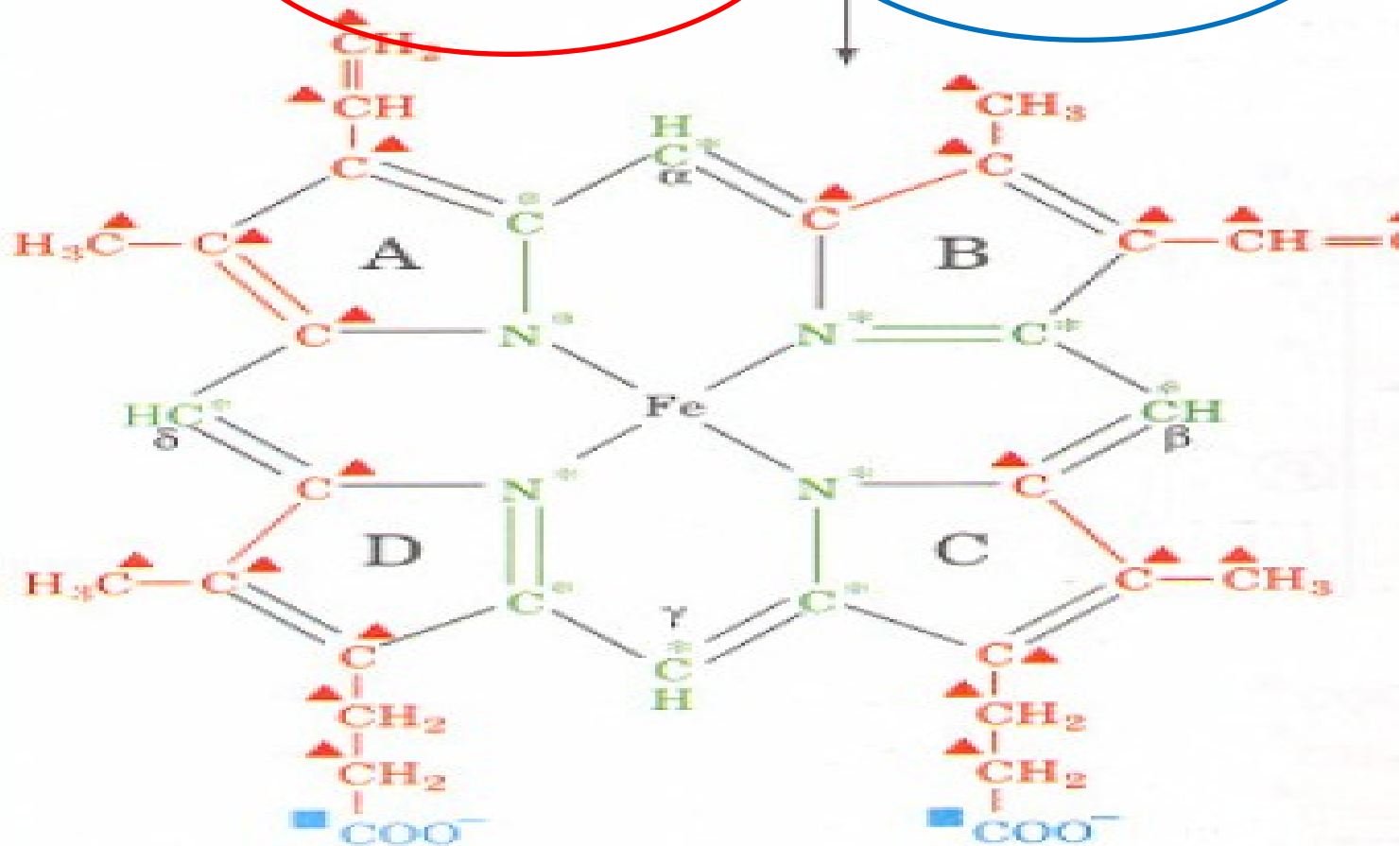
Eukaryota - GluDH

# AMK jako prekurzory

Bioactive Product	Biological Function	Amino Acid Precursor(s)
Alkaloids	Nitrogen bases in plants	Ornithine, Asp, Lys, Tyr, Trp, Phe, His
$\gamma$ -Aminobutyric acid (GABA)	Inhibitory neurotransmitter	Glu
Auxin	Plant growth hormone	Trp
Catecholamines	Neurotransmitters, hormones	Tyr, Phe
Glutathione	Redox tripeptide	Gly, Glu, Cys
Histamine	Allergic response, stomach acid secretion	His
Melanin	Skin pigments	Tyr, Phe
Melatonin	Regulates sleep cycles	Trp
Nitric oxide	Cell messenger	Arg
Phosphocreatine	Energy molecule in muscle	Gly, Arg, Met
Porphyryn	Heme and chlorophyll	Gly
Purine bases	RNA, DNA, cofactors	Asp, Gly, Gln
Pyrimidine bases	RNA, DNA, cofactors	Asp
Serotonin	Neurotransmitter (hormone)	Trp
Spermine, spermidine	DNA packaging	Met, ornithine
Thyroxine	Hormone	Tyr



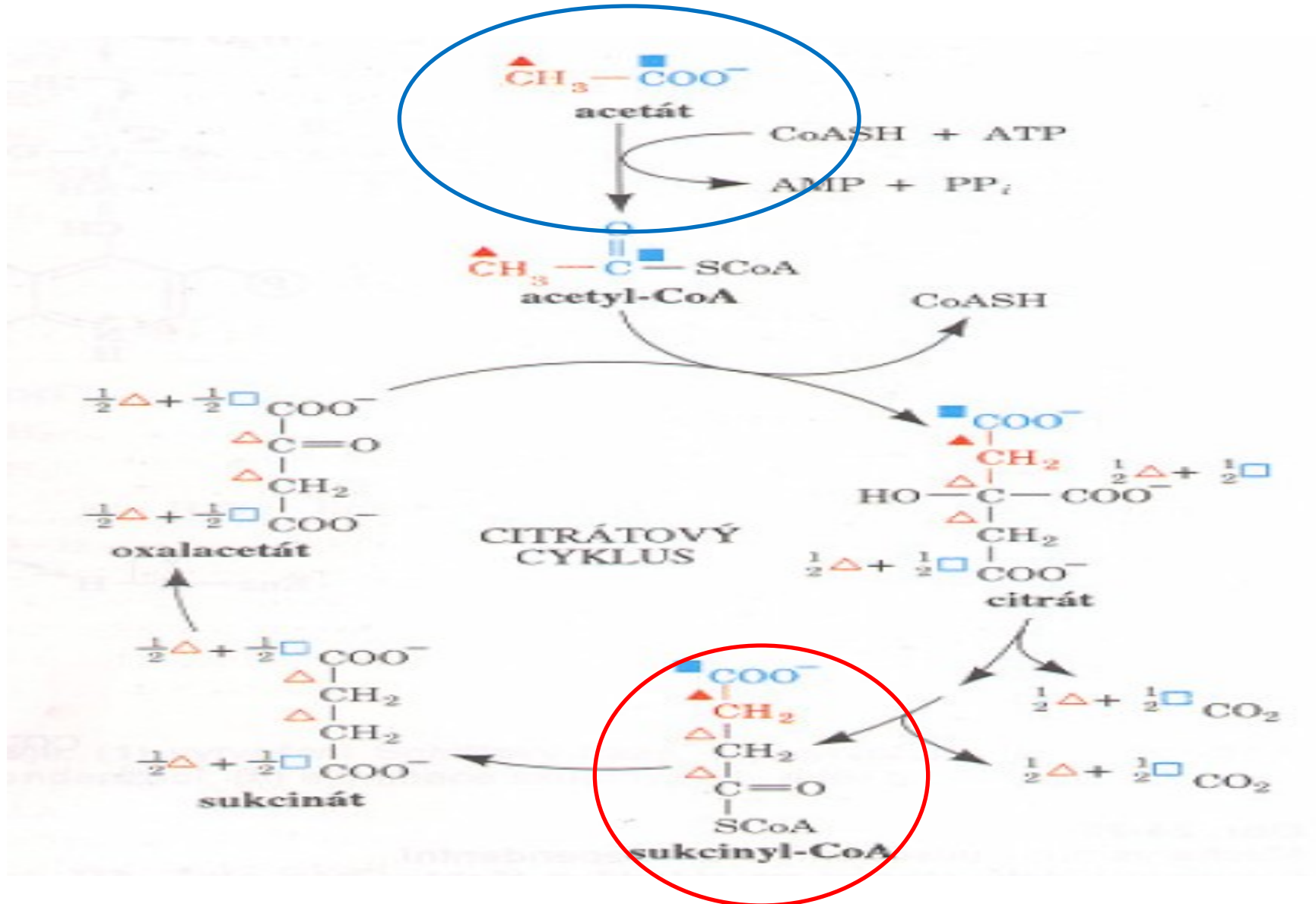
# Hem



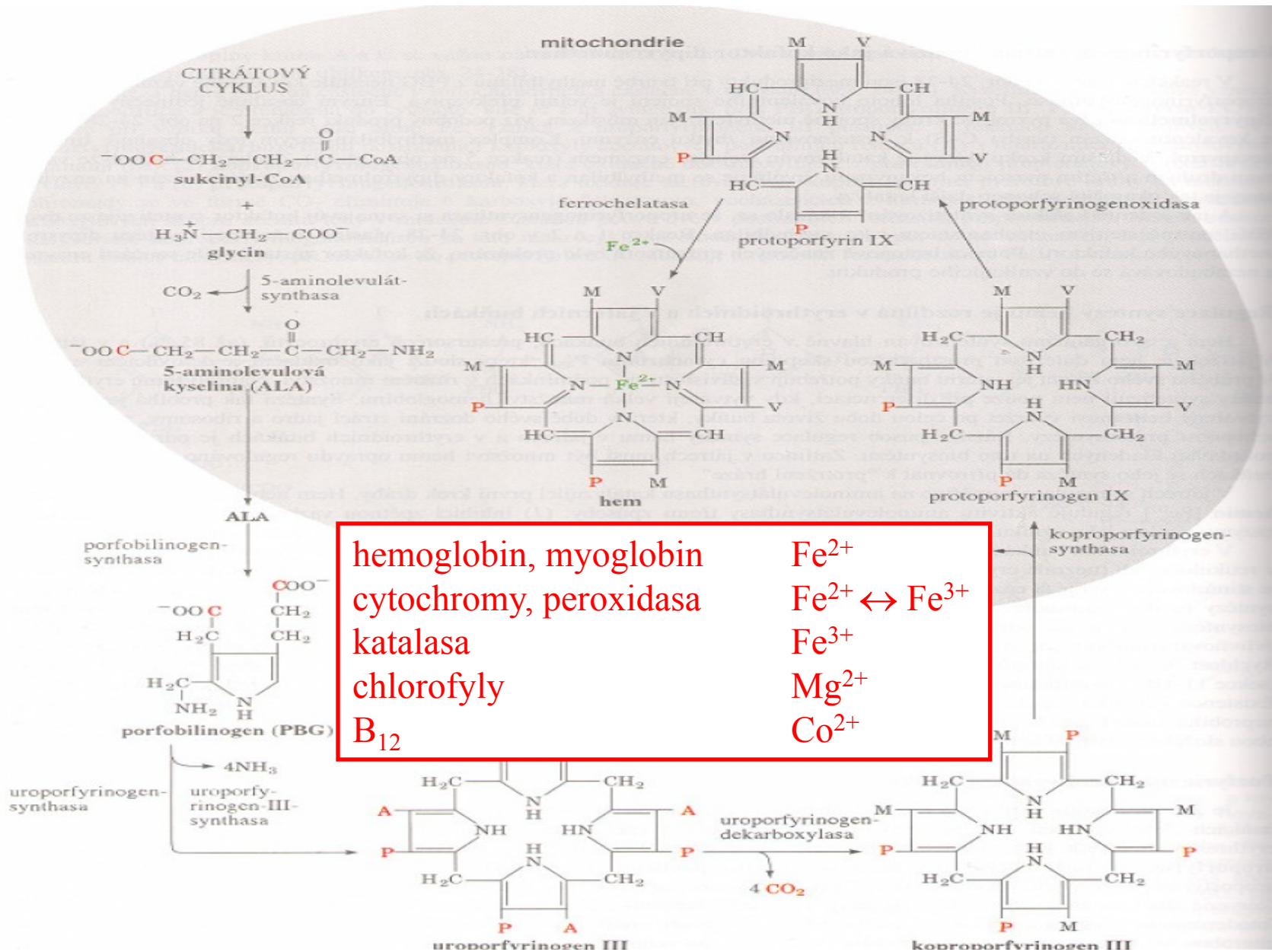
hem



# Hem



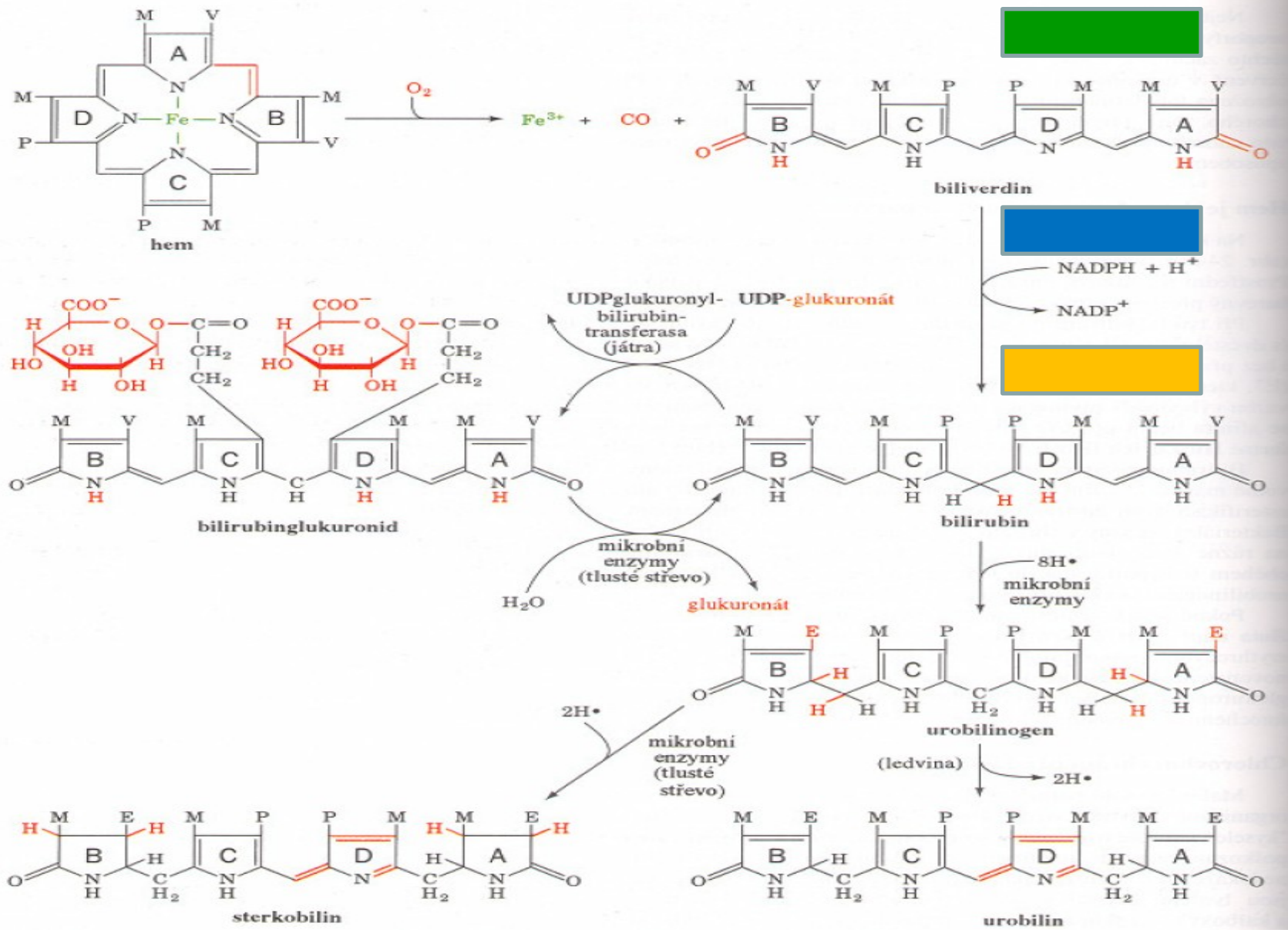
# Biosyntéza hemu



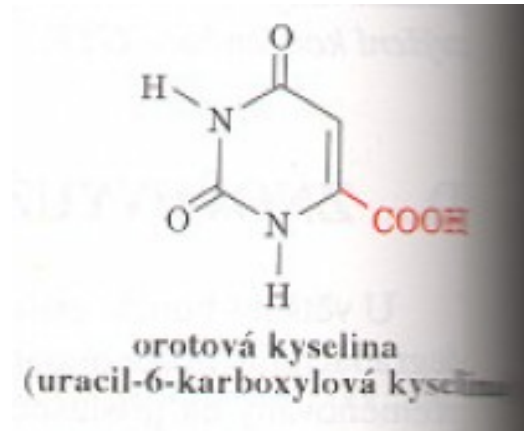
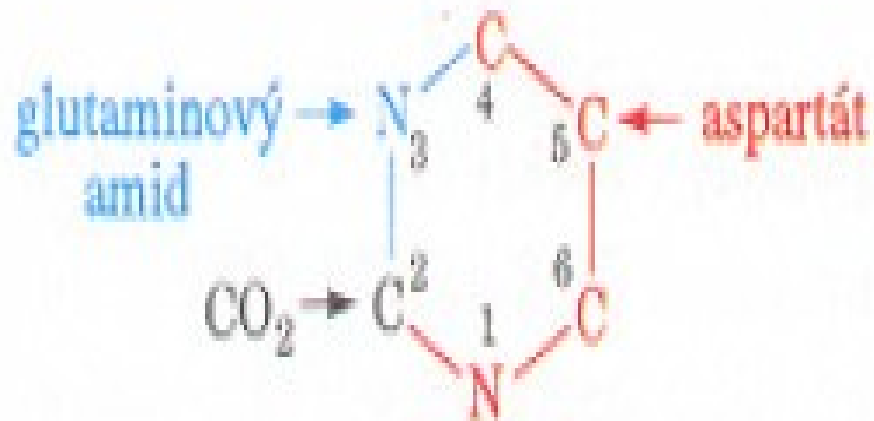
hemoglobin, myoglobin  
 cytochromy, peroxidasa  
 katalasa  
 chlorofyly  
 B<sub>12</sub>

Fe<sup>2+</sup>  
 Fe<sup>2+</sup> ↔ Fe<sup>3+</sup>  
 Fe<sup>3+</sup>  
 Mg<sup>2+</sup>  
 Co<sup>2+</sup>

# Degradace hemu

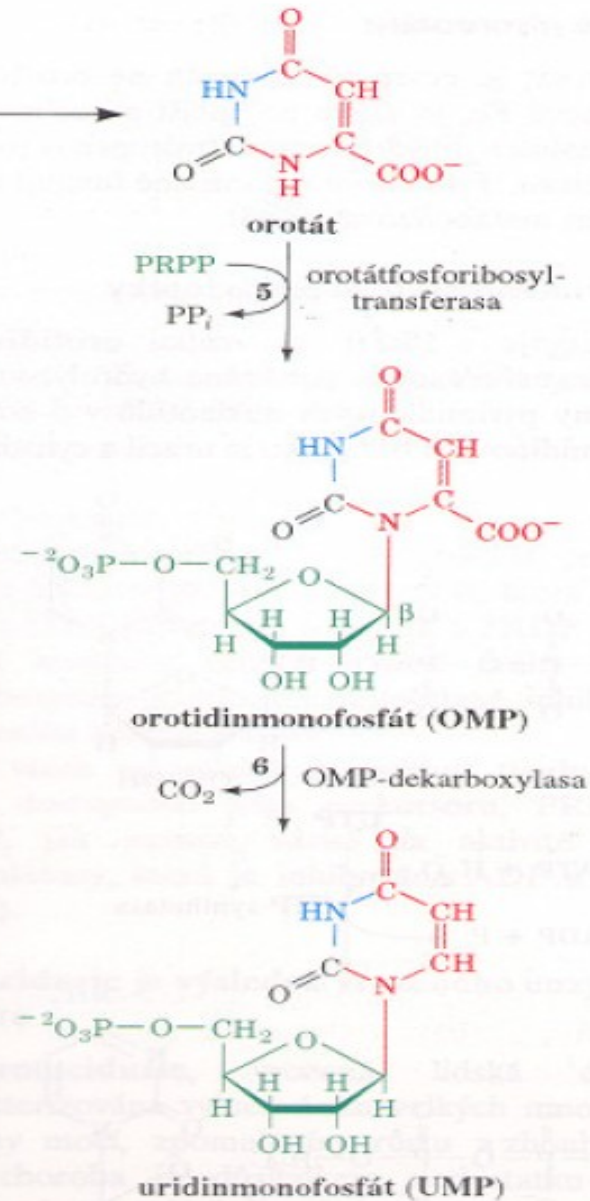
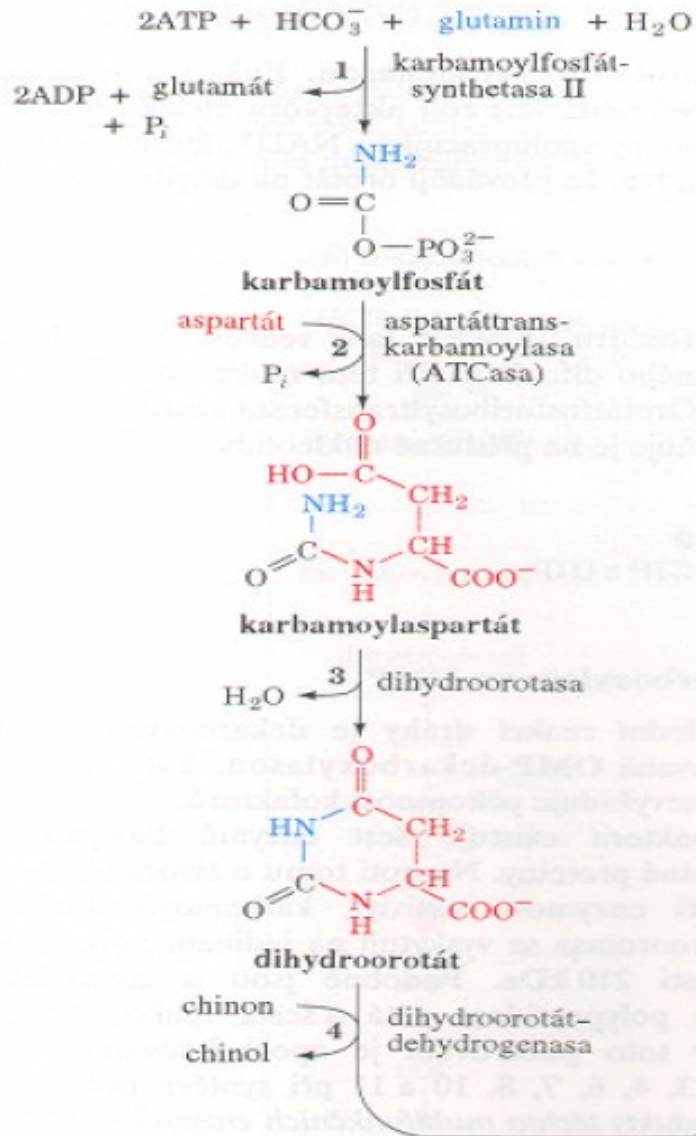


# Biosyntéza pyrimidinových bází

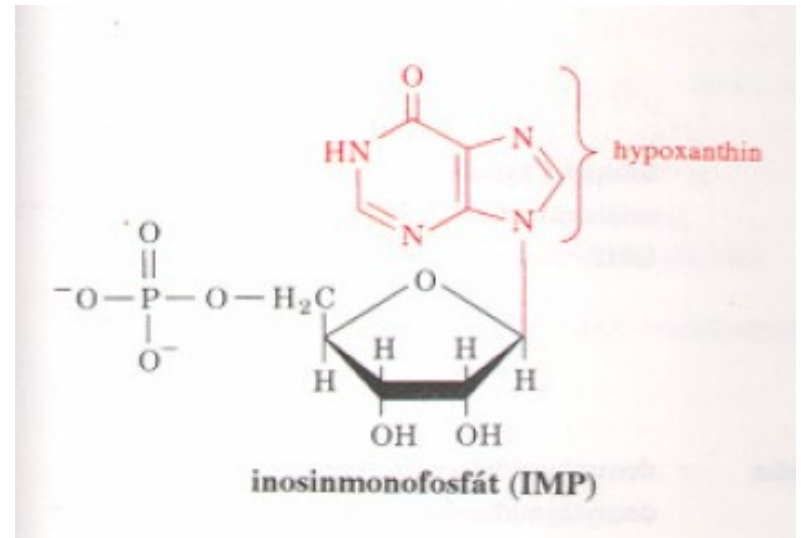
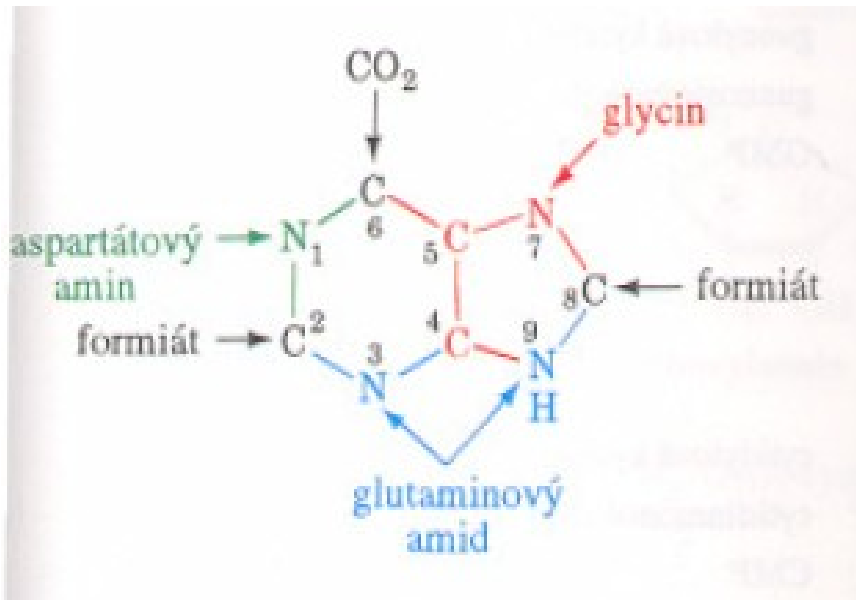




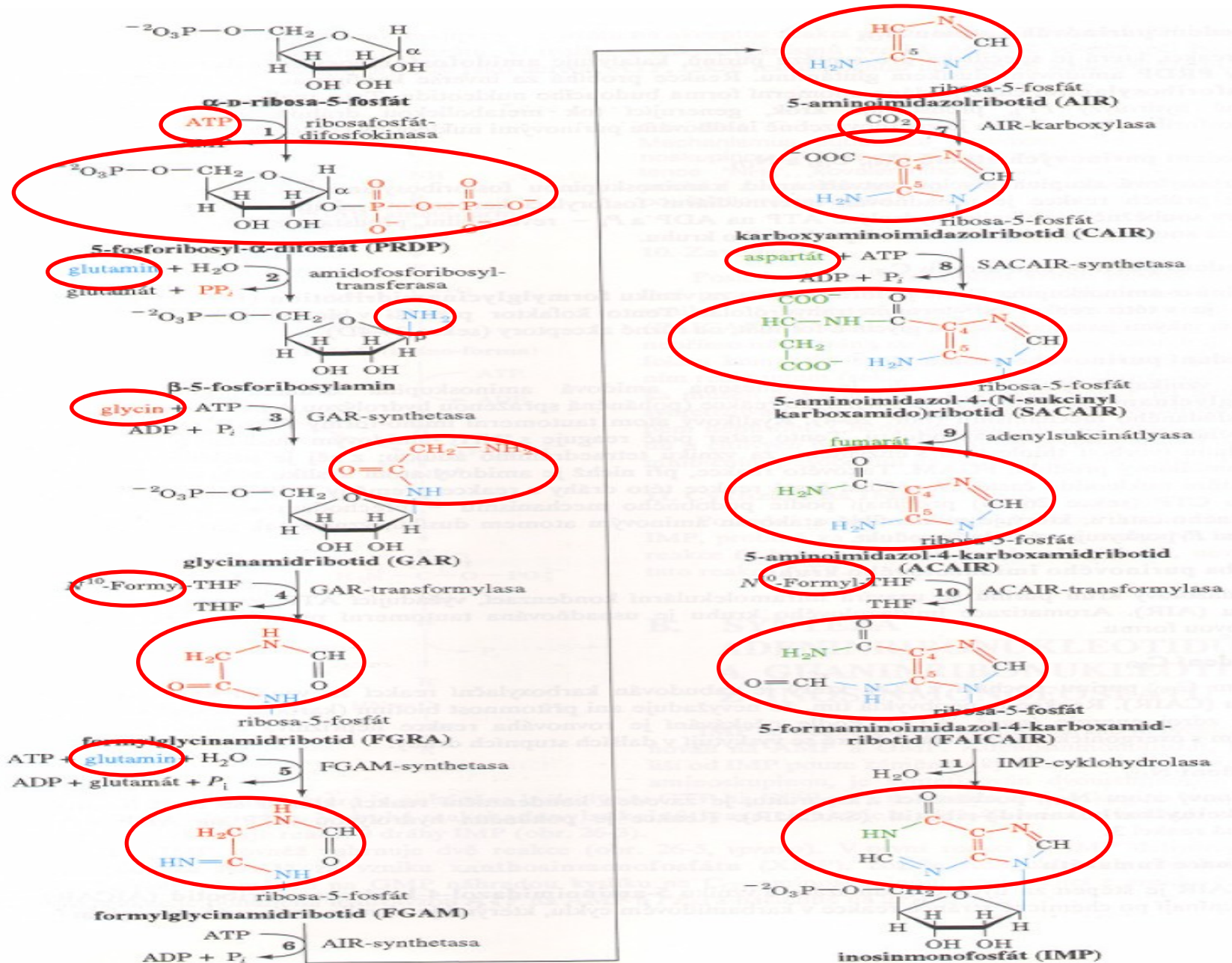
# Biosyntéza pyrimidinových bází



# Biosyntéza purinových bází

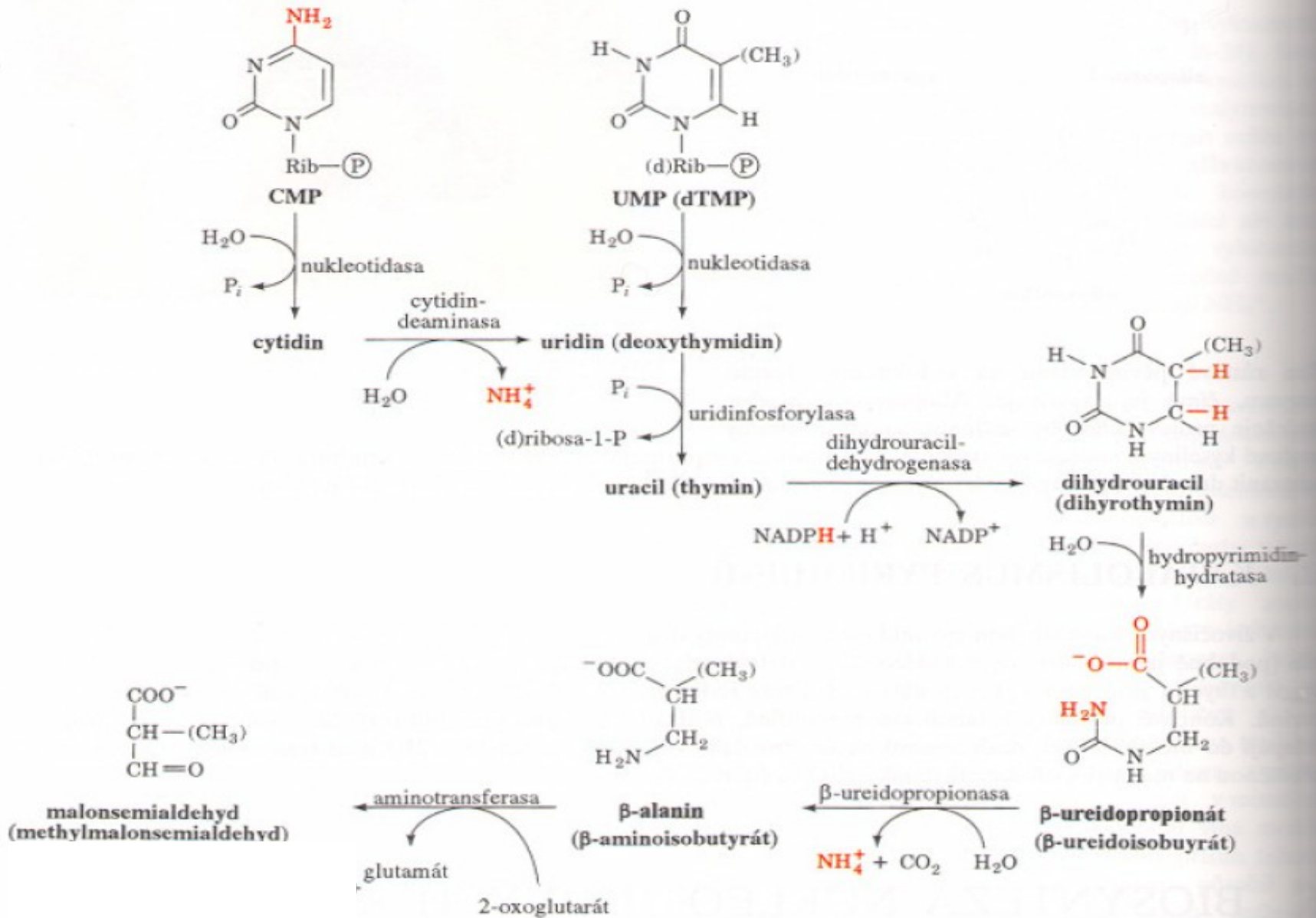


# Biosyntéza purinových bází

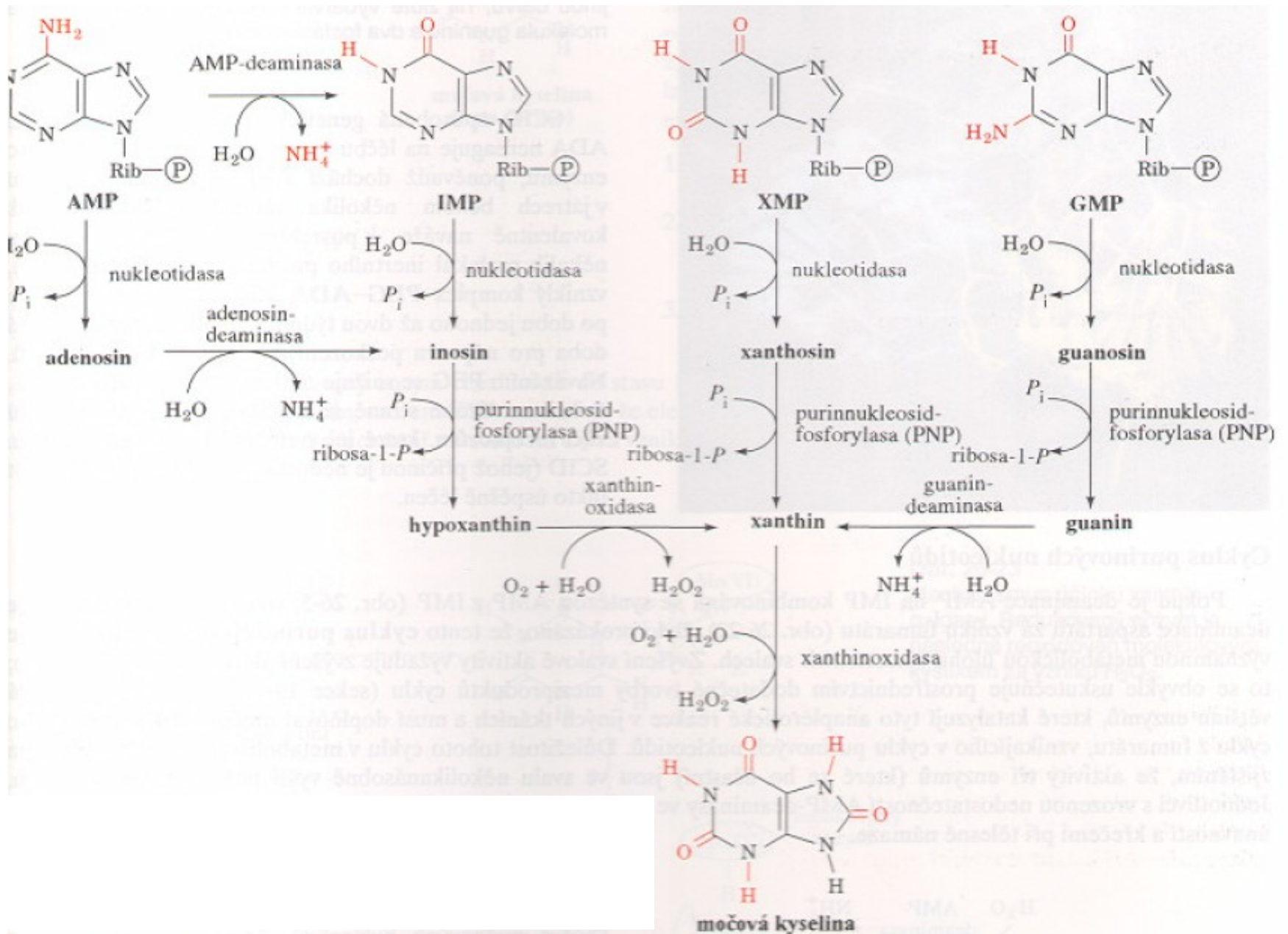




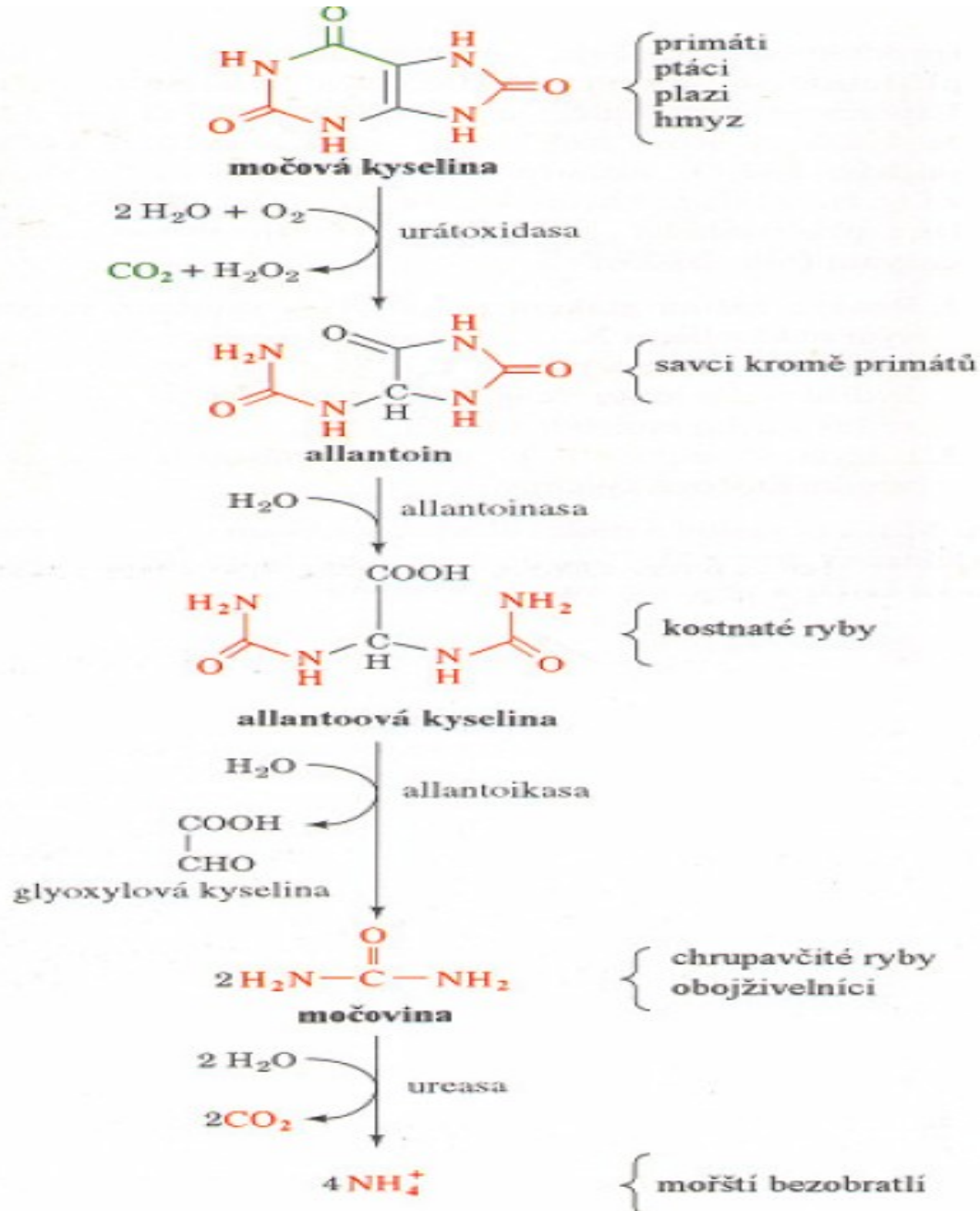
# Degradace pyrimidinových bází



# Degradace purinových bází



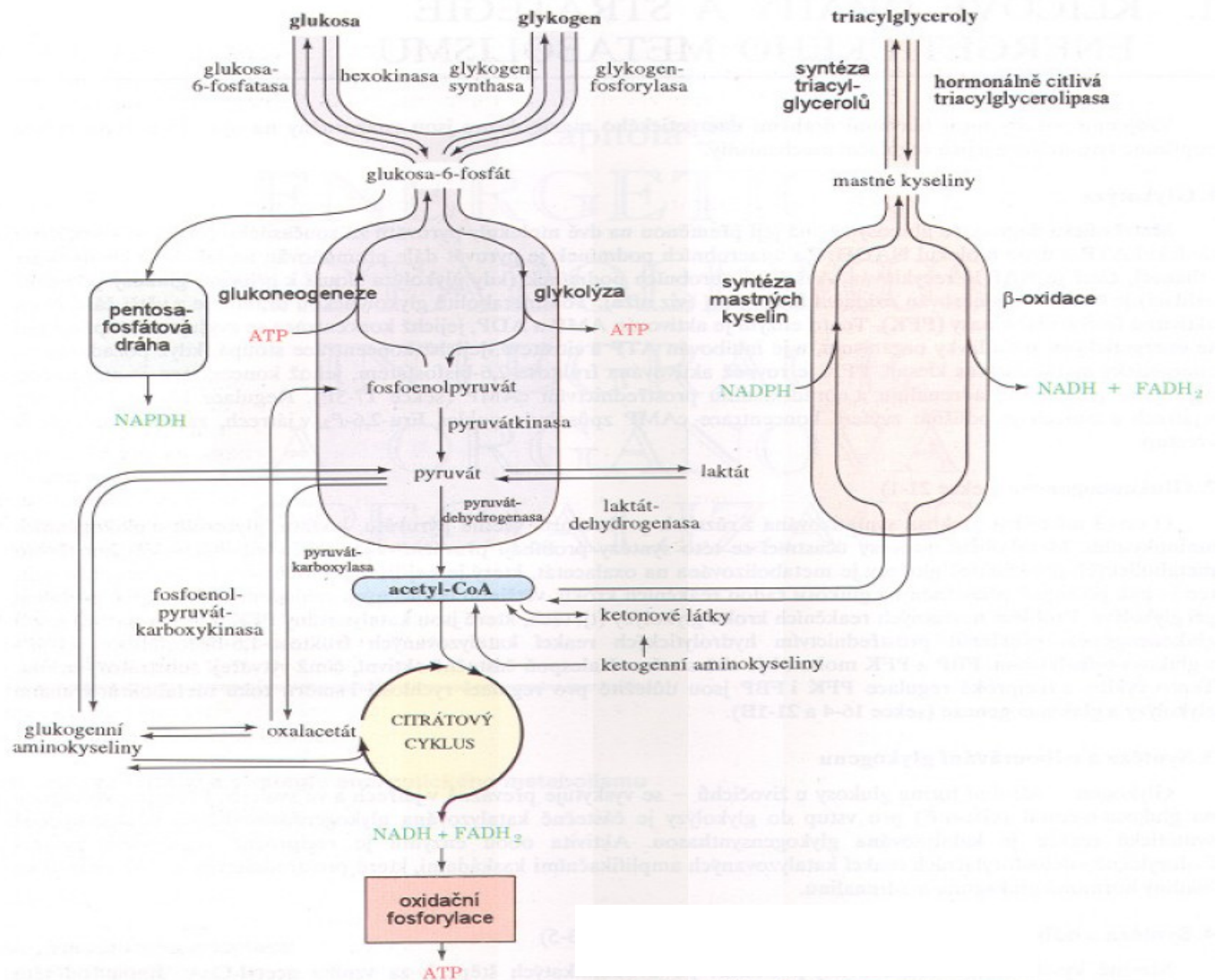
# Degradace purinových bází



# Metabolismus NK

- Žaludek – odštěpení proteinů pomocí HCl
- Nukleasa (fosfodiesterasa) – štěpení na oligo- a mononukletidy
- Mononukleotidasa – nukleosid +  $H_3PO_4$
- Nukleosidasa – cukr + basa

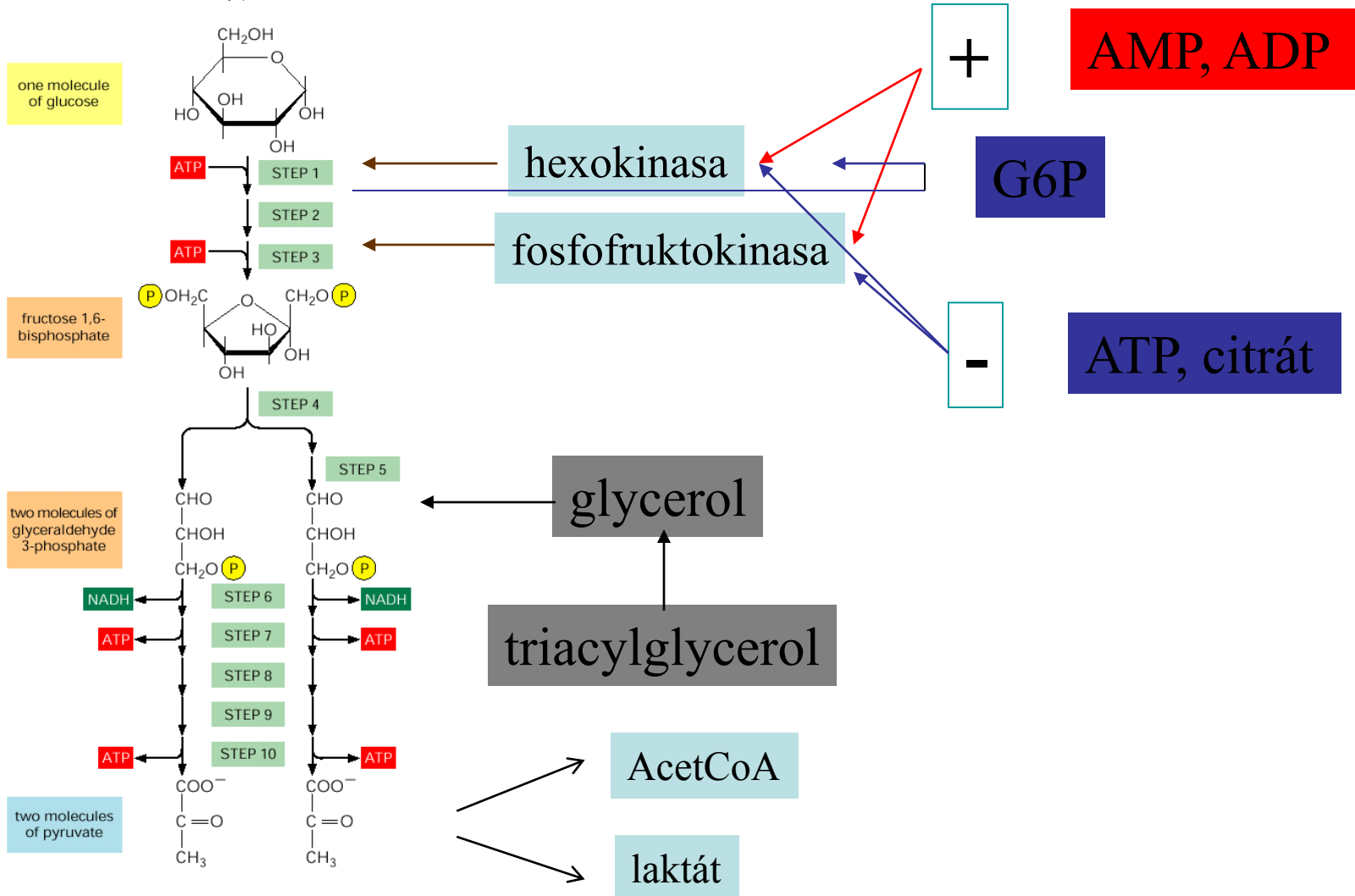
# Hlavní dráhy energetického metabolismu



glykogen

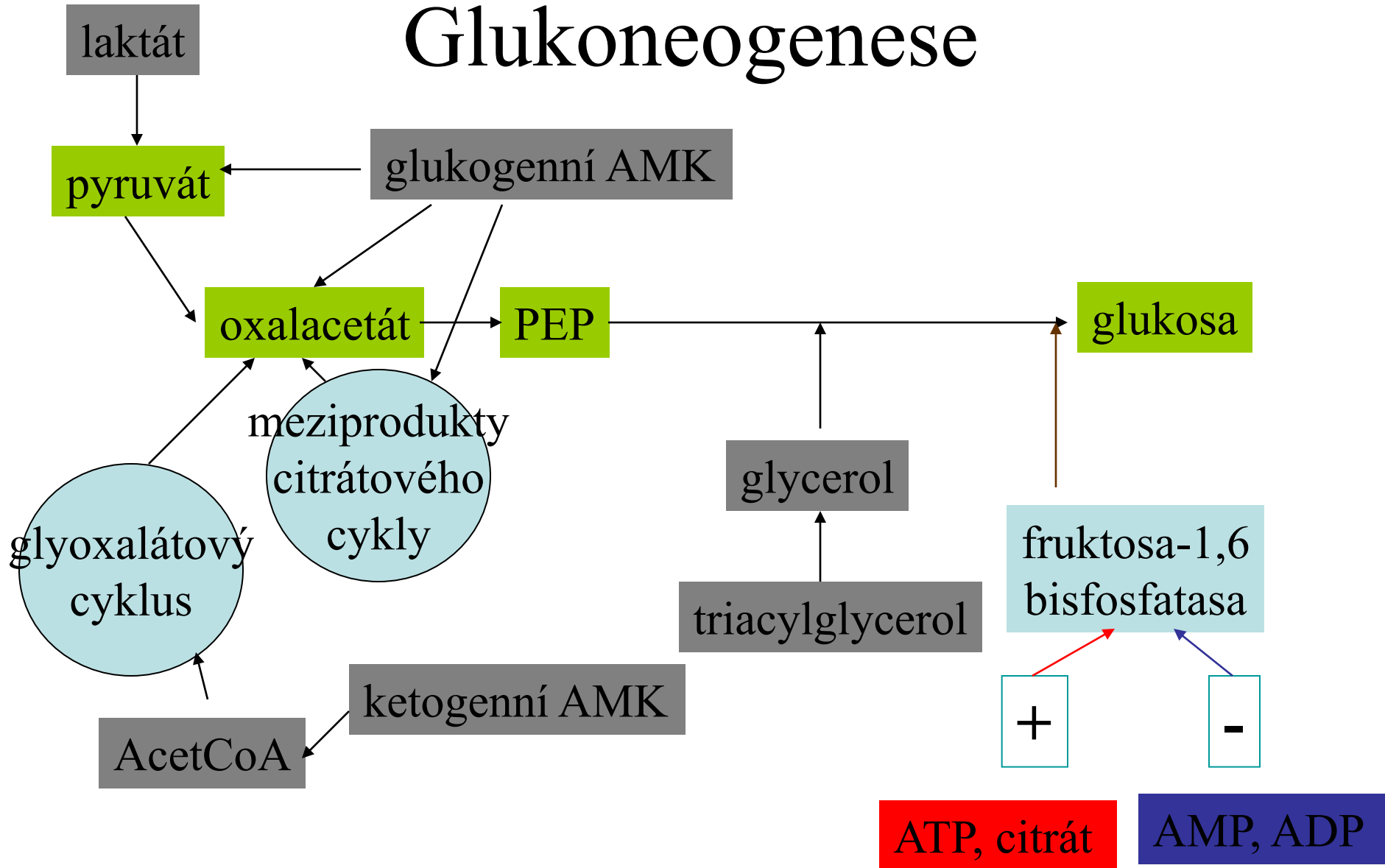
glukosa monosacharidy

# Glykolysa



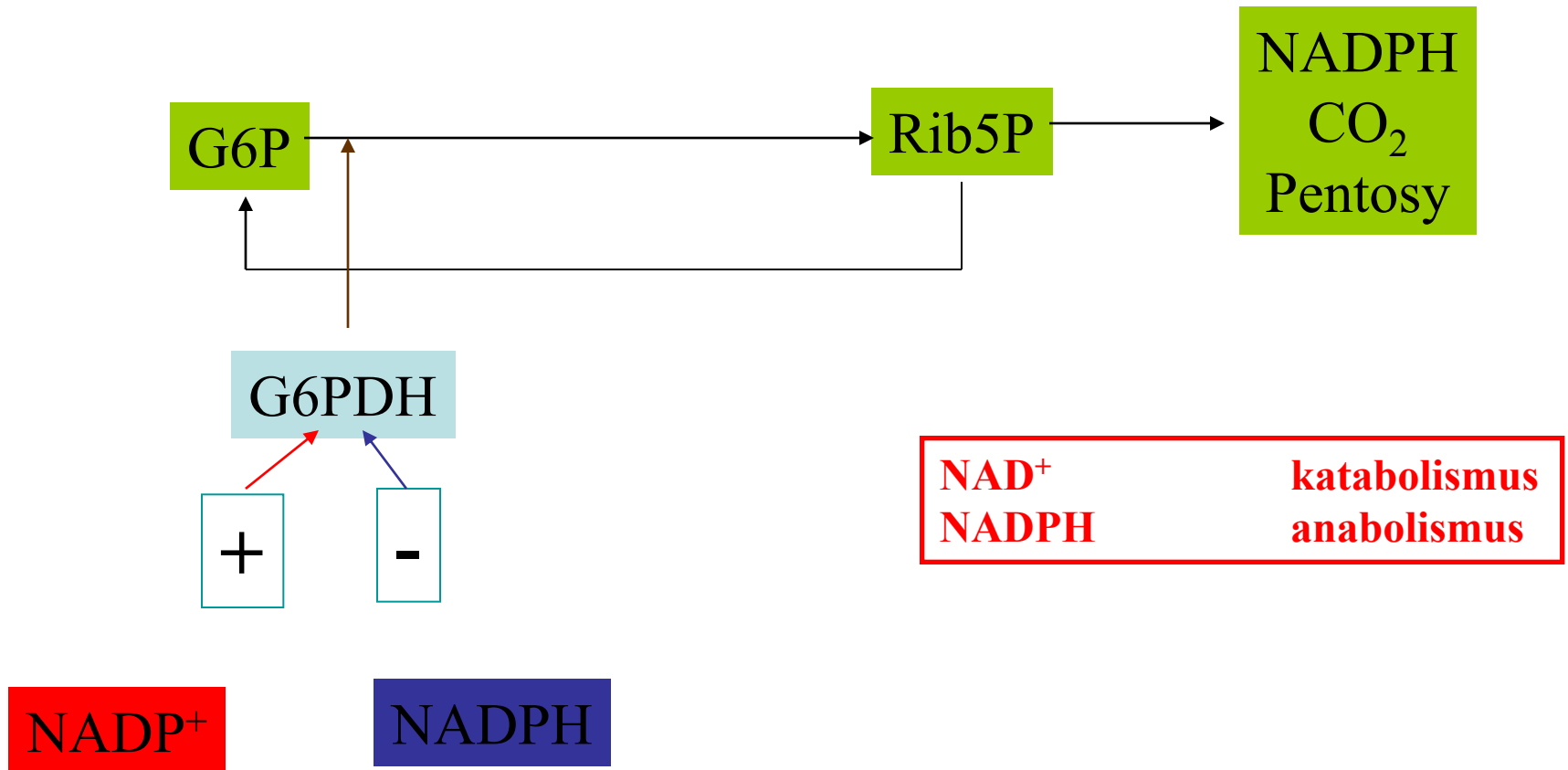


# Glukoneogenese

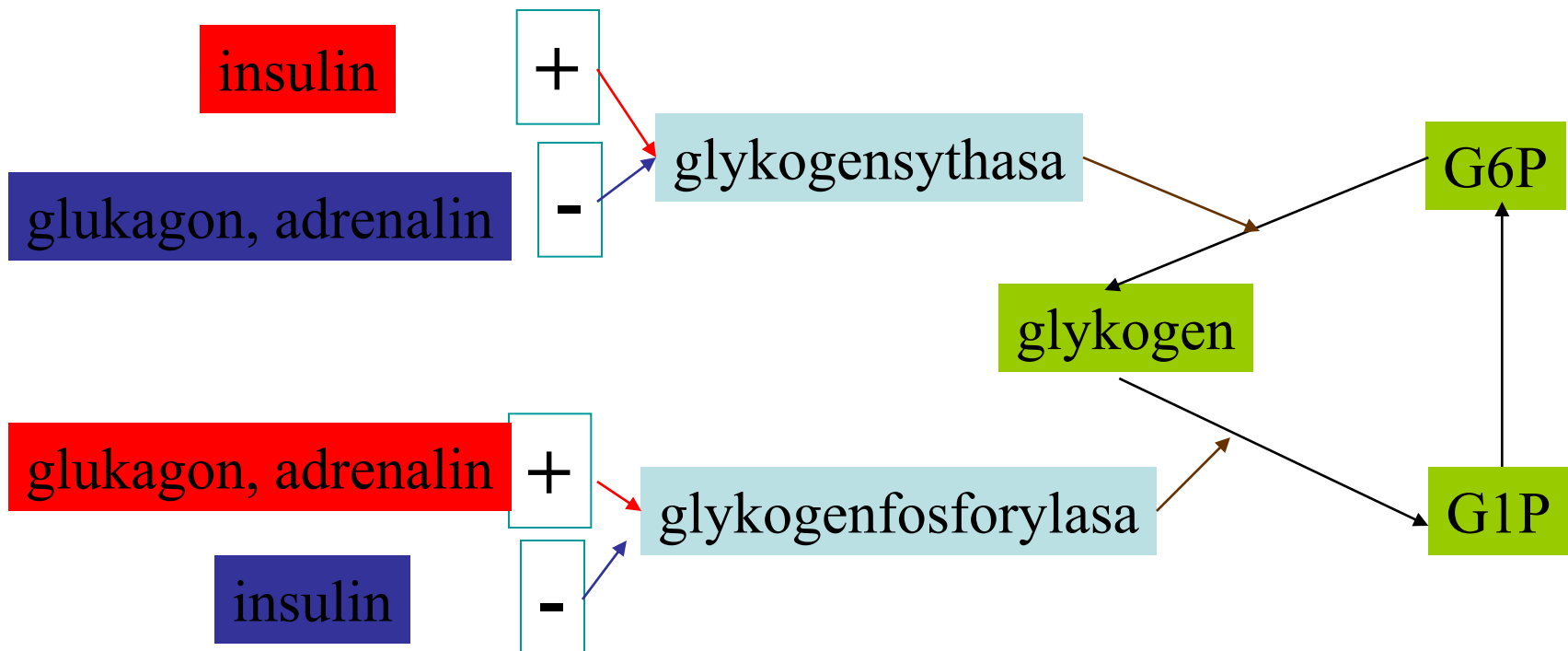




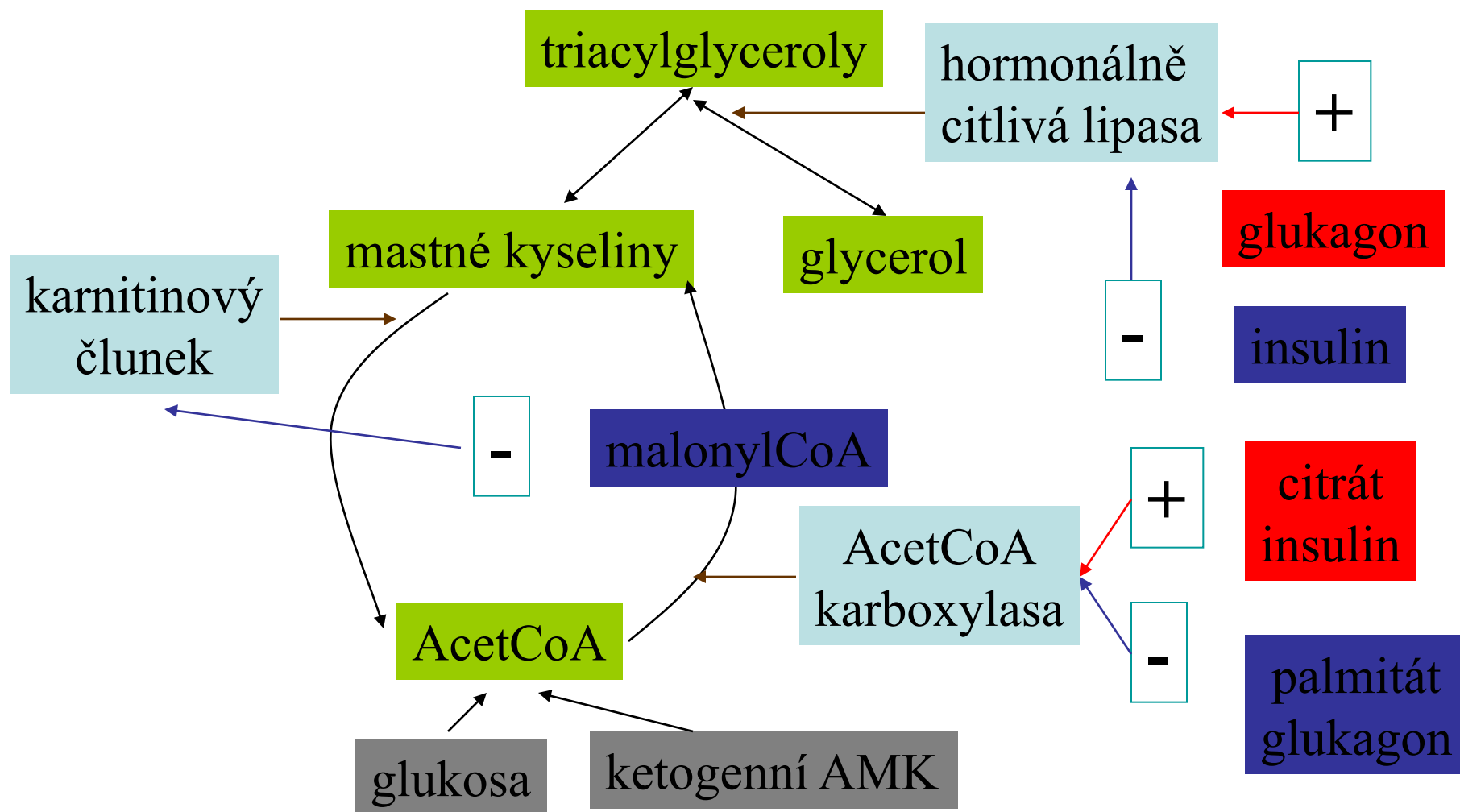
# Pentosový cyklus



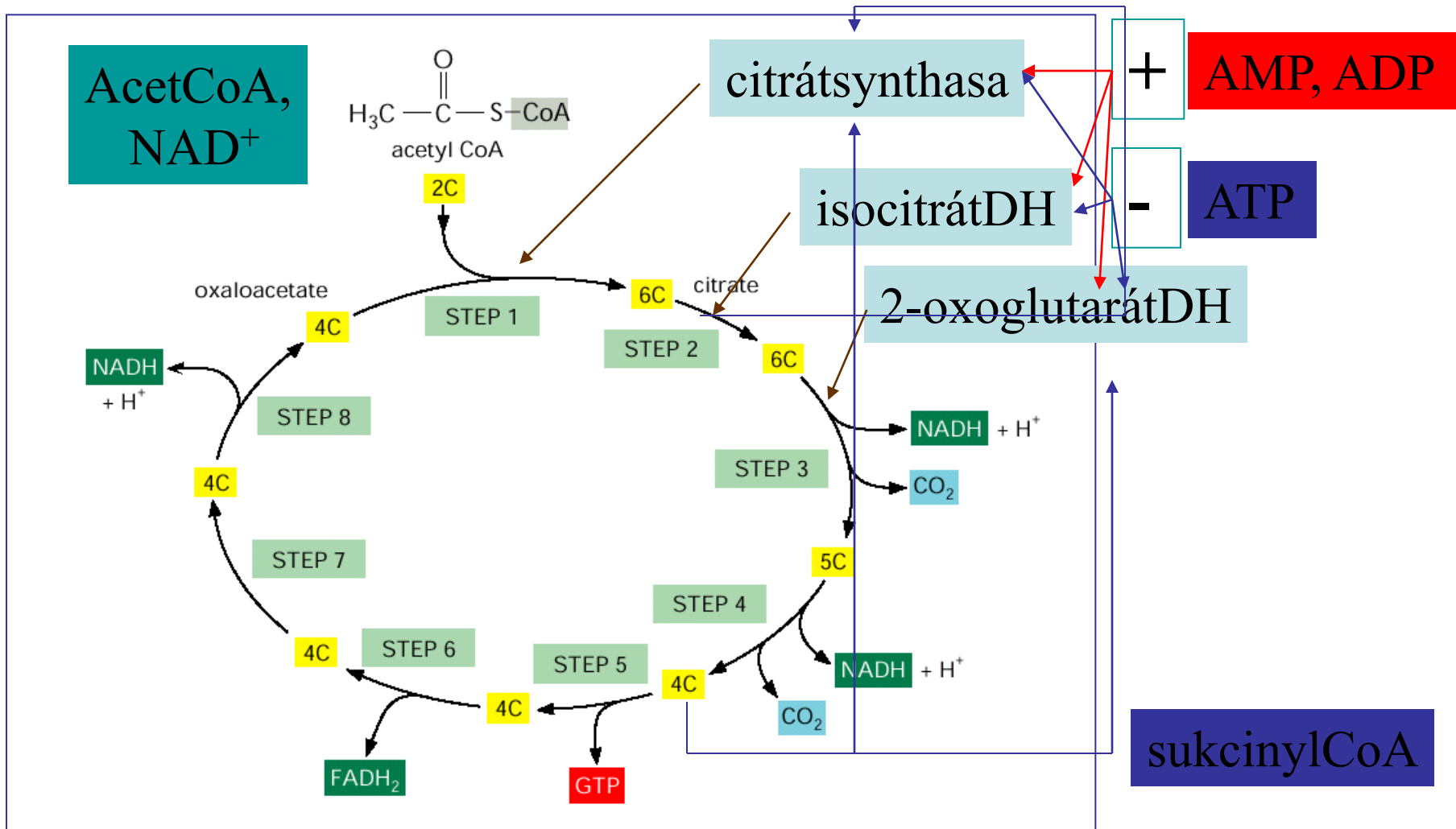
# Syntesa a odbourávání glykogenu



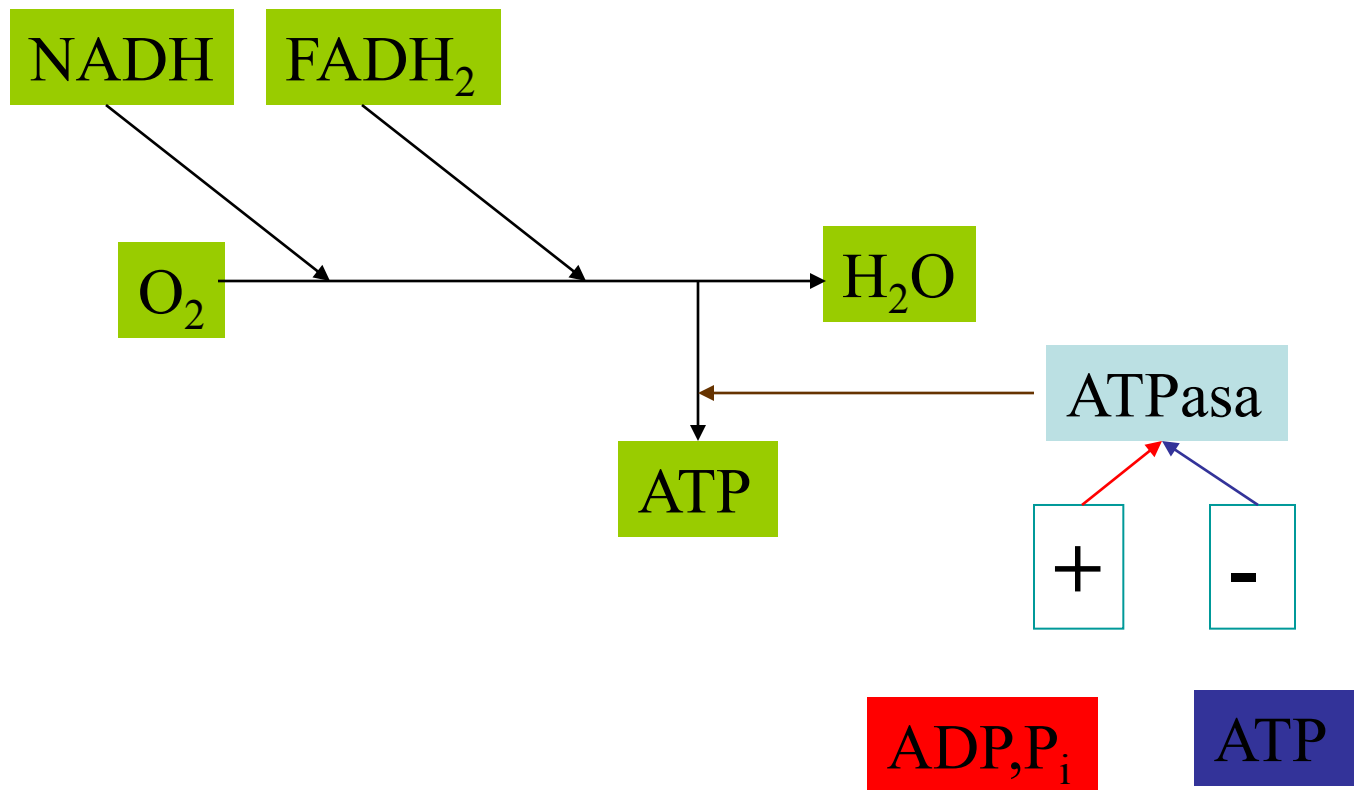
# Metabolismus triacylglycerolů



# Citrátový cyklus



# Oxidační fosforylace



# Orgánová specializace

