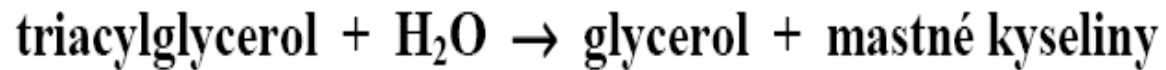


Metabolismus lipidů

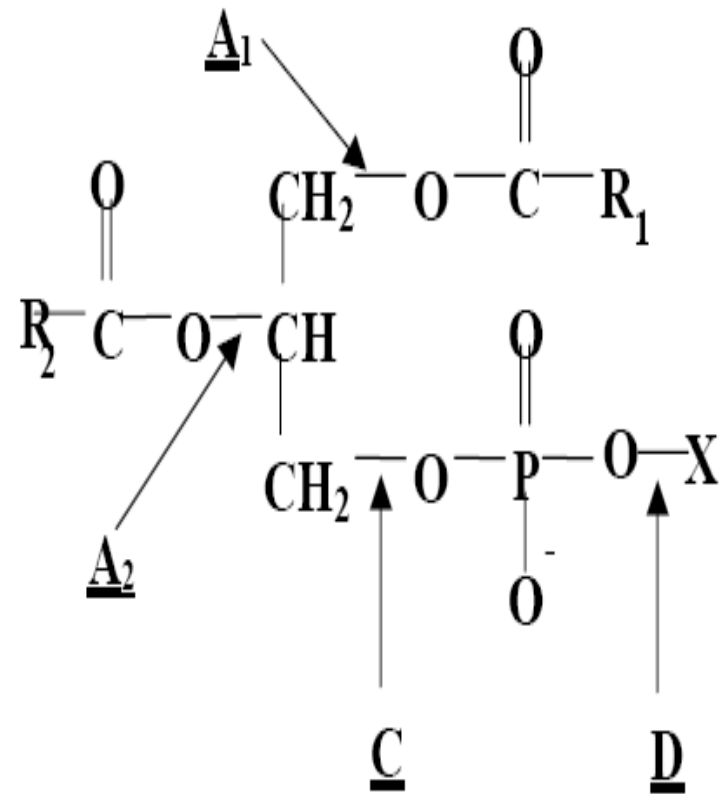
- Triacylglyceroly – 90 % potravních lipidů a zároveň hlavní zásobárnou energie – orgánový tuk
- Dvojnásobné množství energie
CH₂ - CHOH
- Jako hydrofobní látky nejsou hydratovány
- Nejsou pohotovým zdrojem energie, spíše vhodné pro skladování.

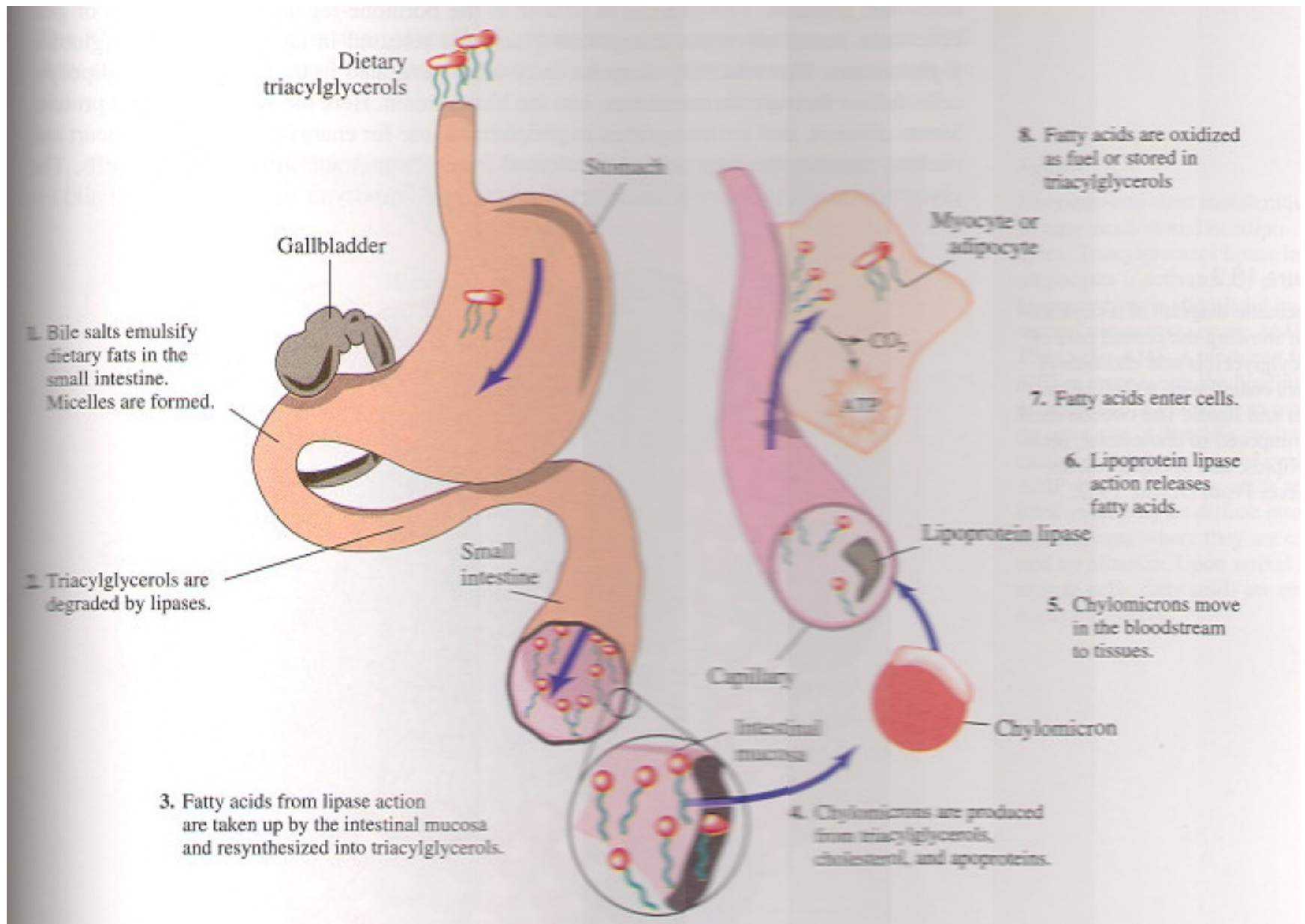
METABOLISMUS LIPIDŮ

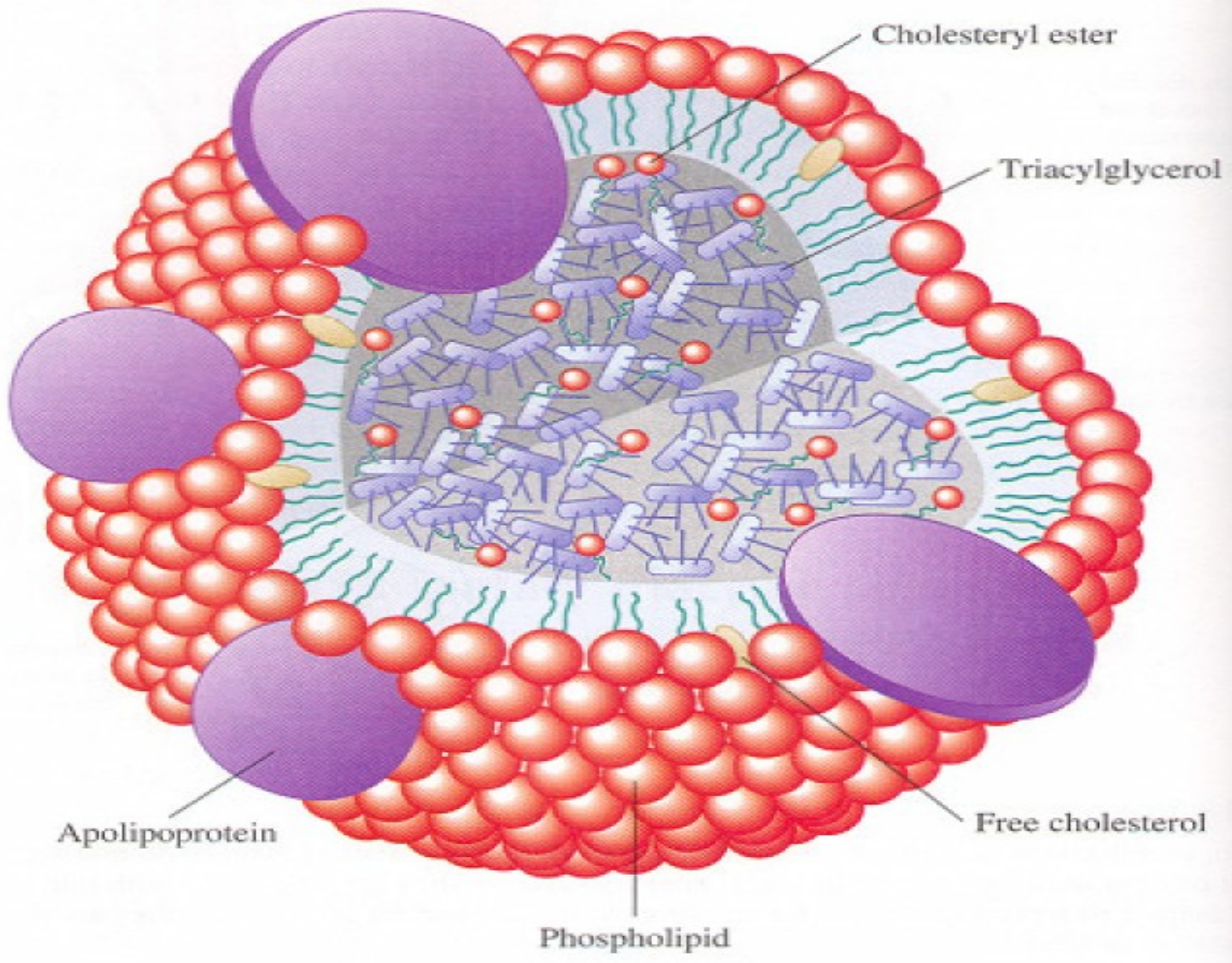
Lipasy - hydrolasy - karboxylesterasy

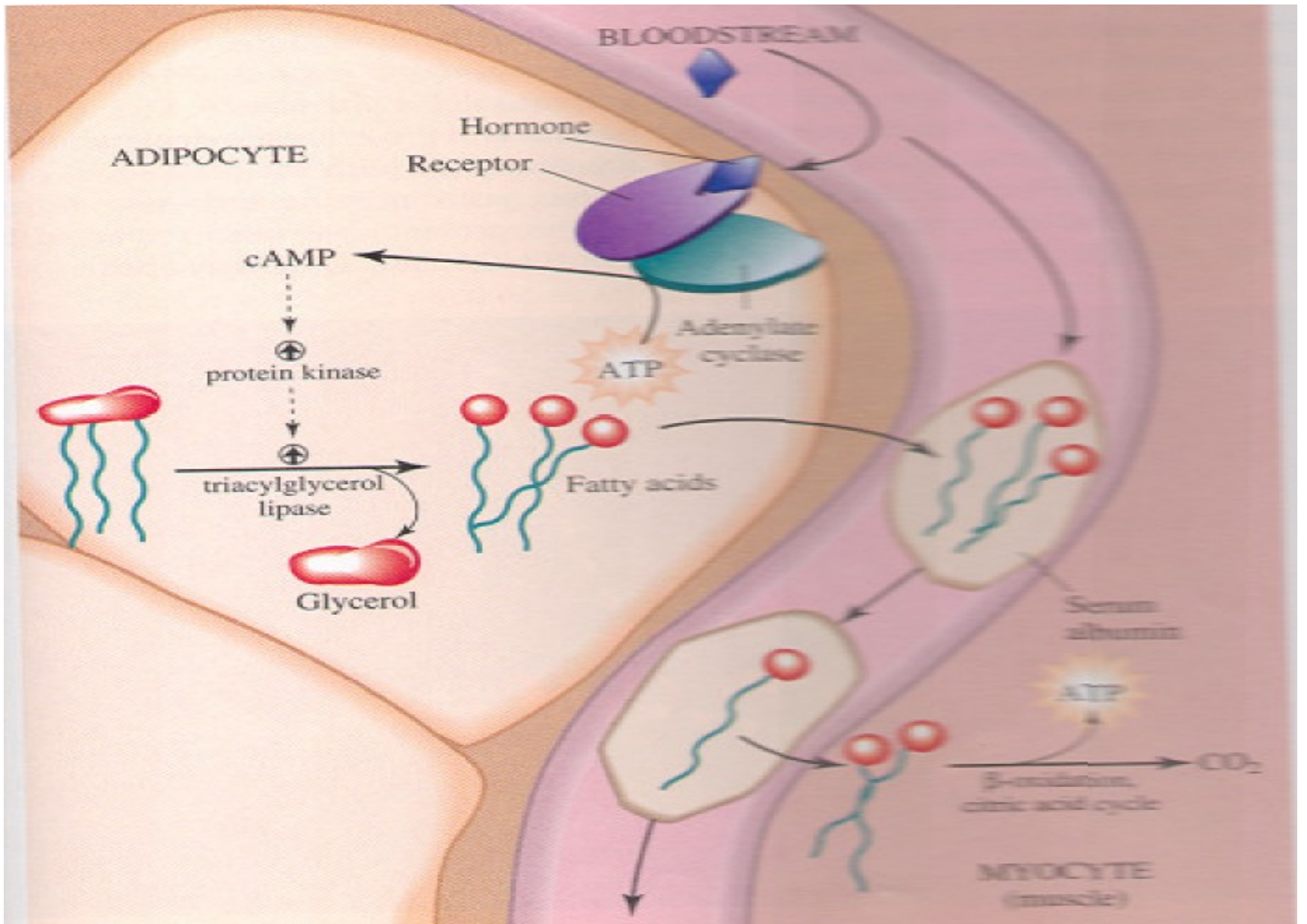


Fosfolipasy







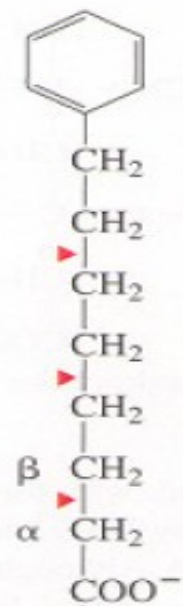


Odbourávání mastných kyselin

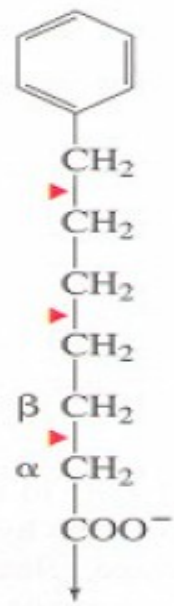
β oxidace

F.KNOOP 1909

F.LYNEN 1951

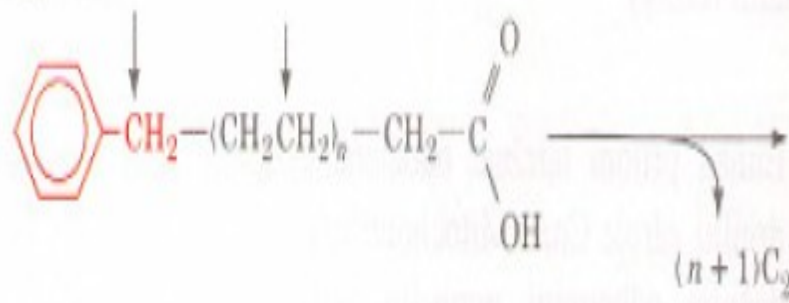


Phenylacetate
(from even-numbered
carbon chain)

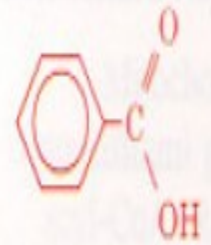


Benzoate
(from odd-numbered
carbon chain)

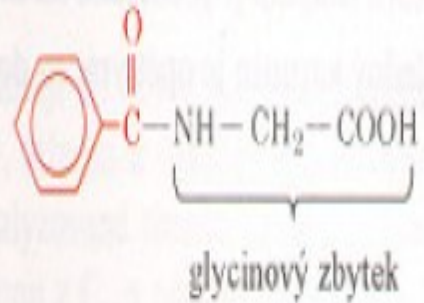
přidaná mastná kyselina



degradační produkty



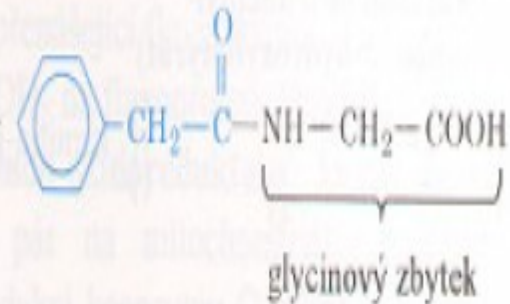
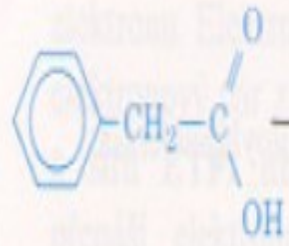
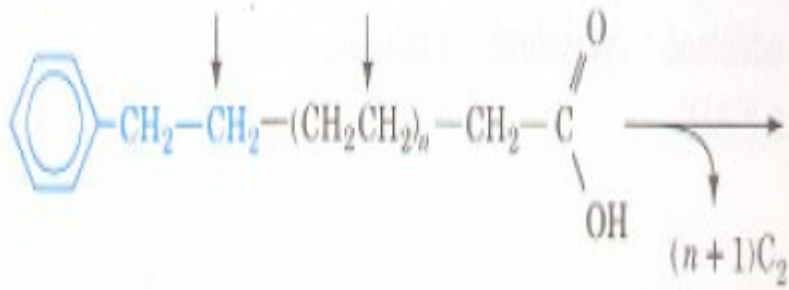
exkreční produkt



kyselina s lichým počtem C-atomů

benzoová kyselina

hippurová kyselina



kyselina se sudým počtem C-atomů

fenylactová kyselina

fenylaceturová kyselina

Odbourávání mastných kyselin

β oxidace

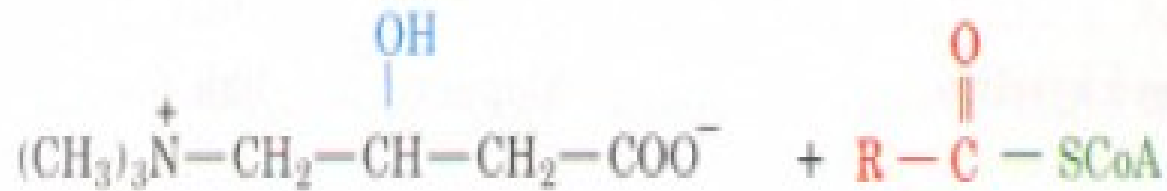
F.KNOOP 1909

F.LYNEN 1951

A. Aktivace mastných kyselin

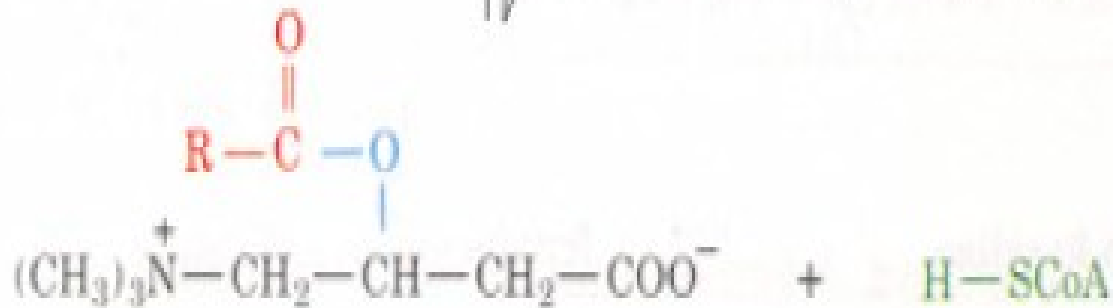


B. Transport RCOSCoA - karnitinový člunek



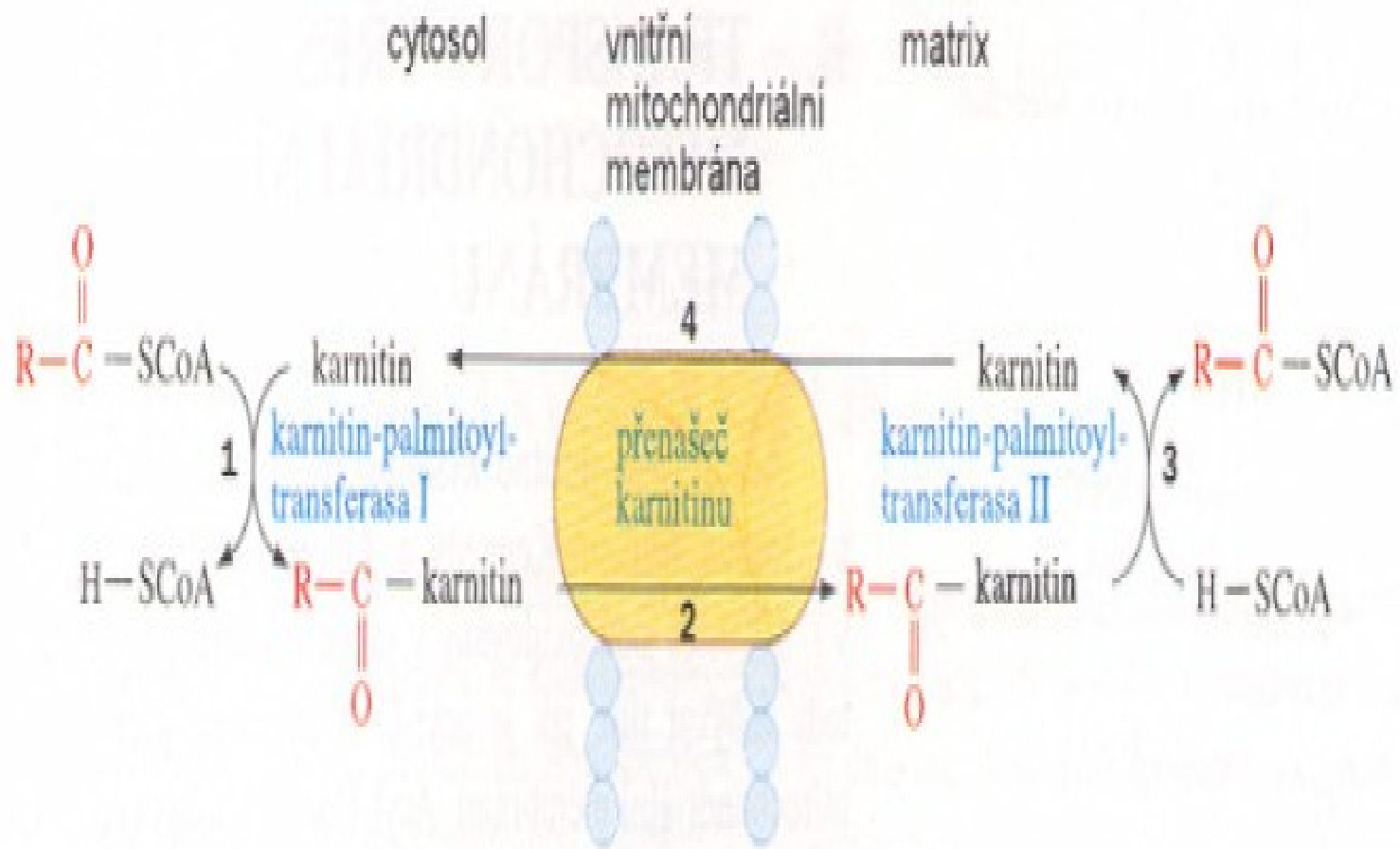
karnitin (4-trimethyl-
amino-3-hydroxybutyrát)

⇌ karnitin-palmitoyltransferasa

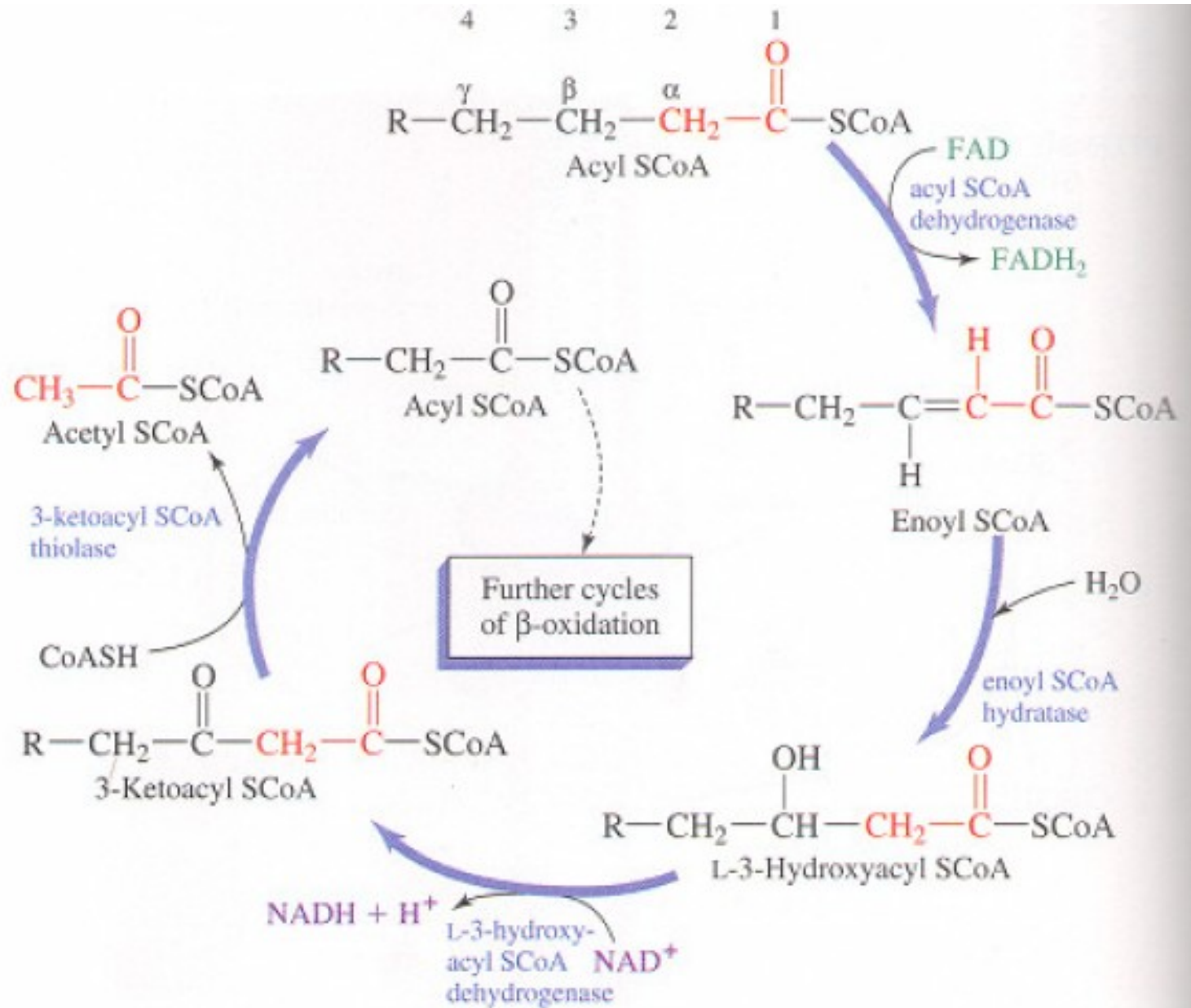


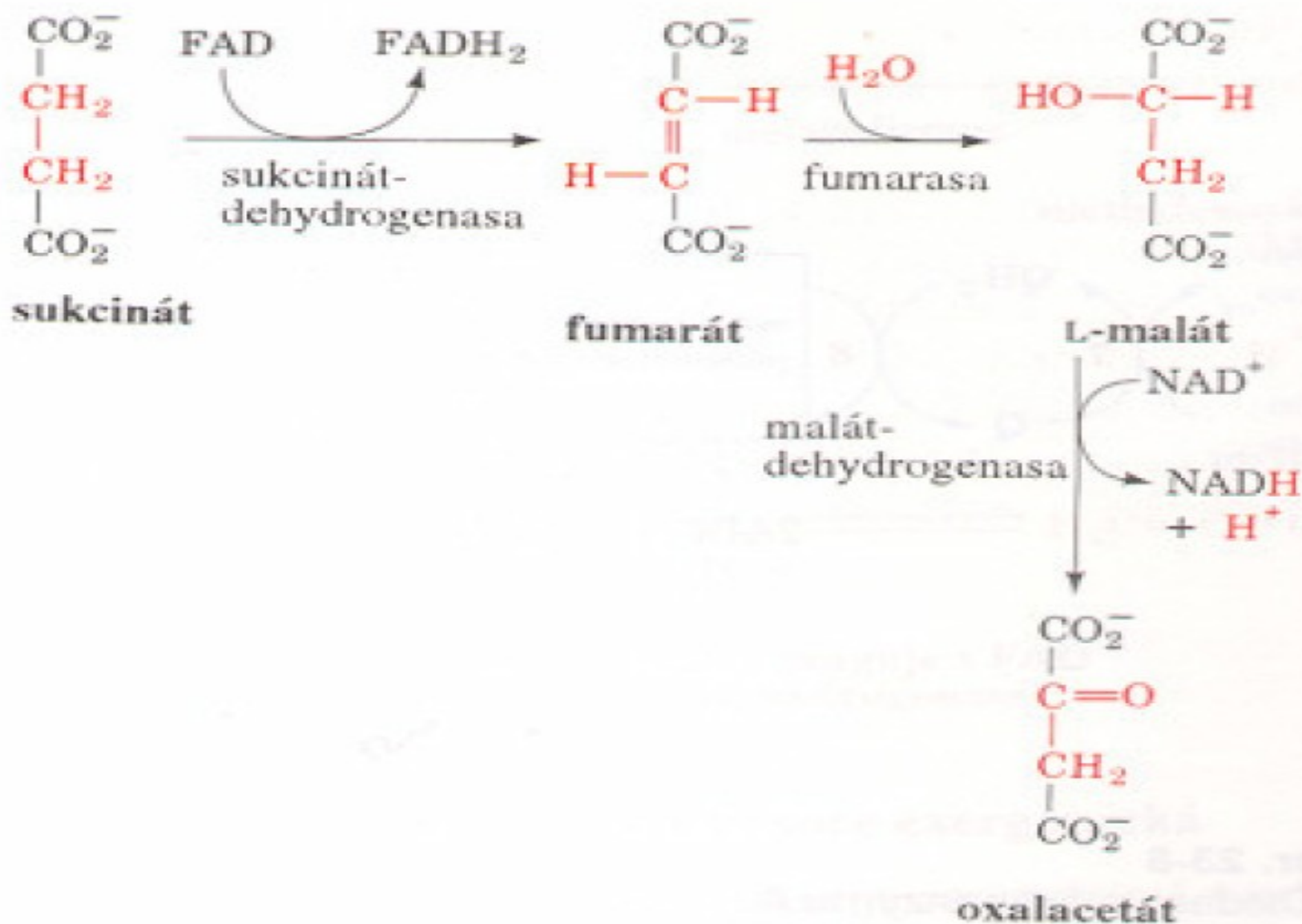
acylkarnitin

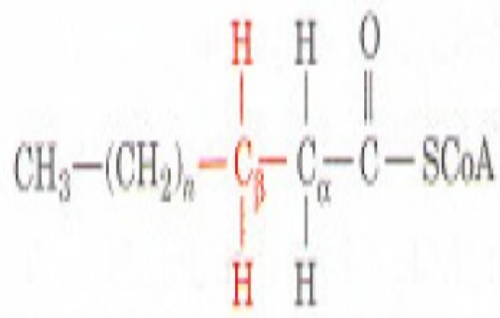
B. Transport $RCOSCoA$ - karnitinový člunek



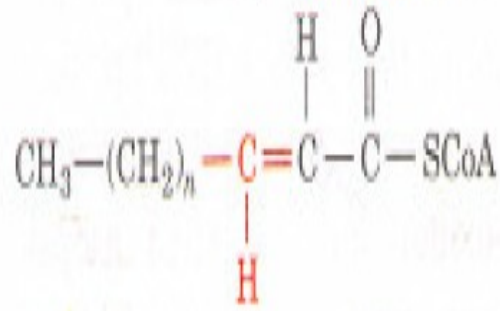
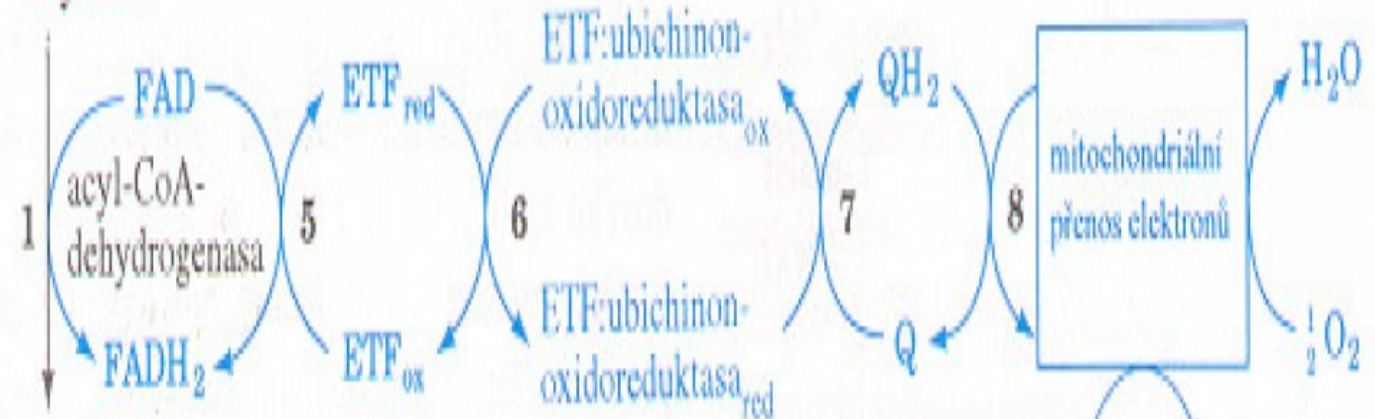
C. β oxidace







acyl-CoA

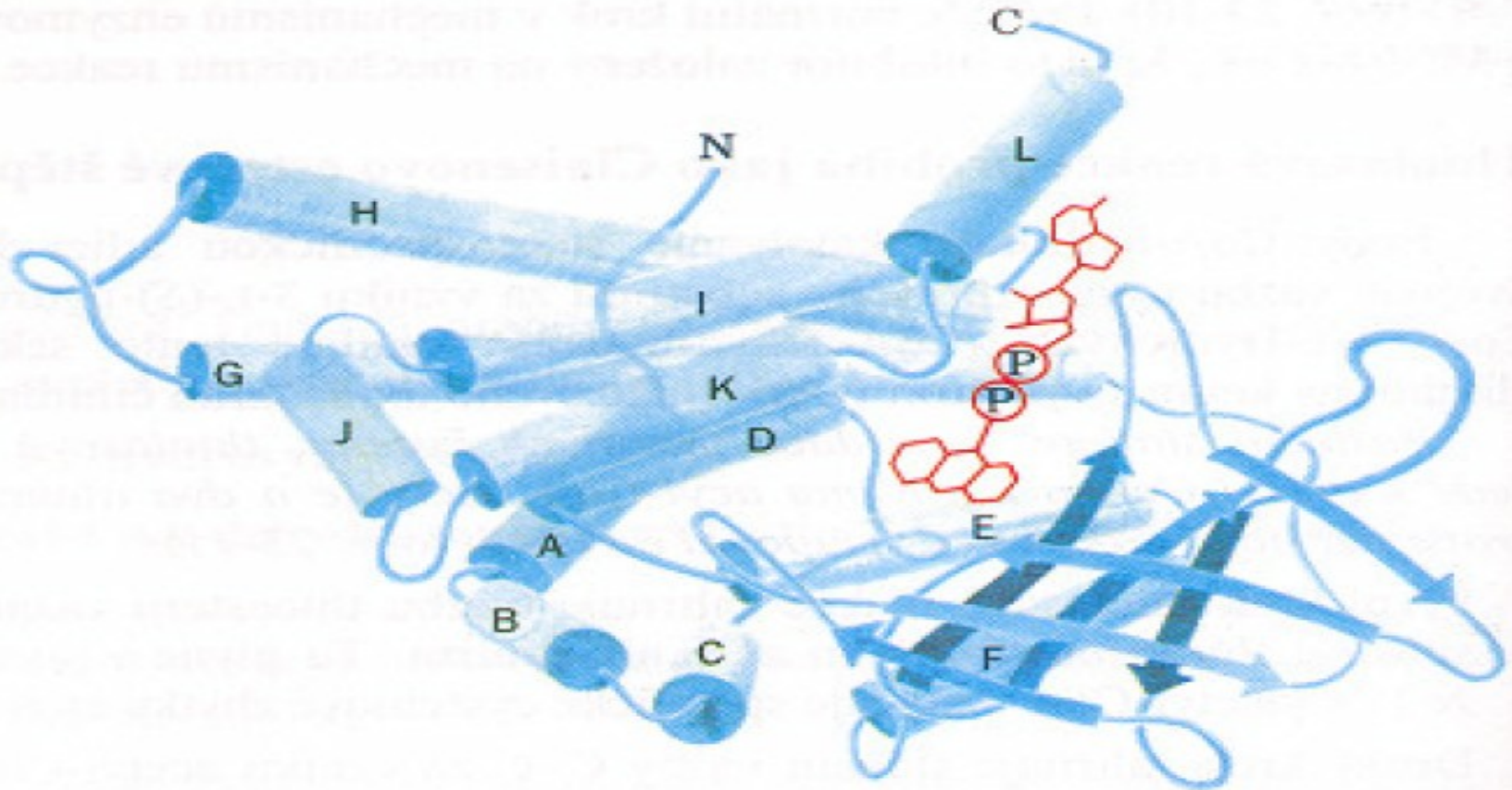


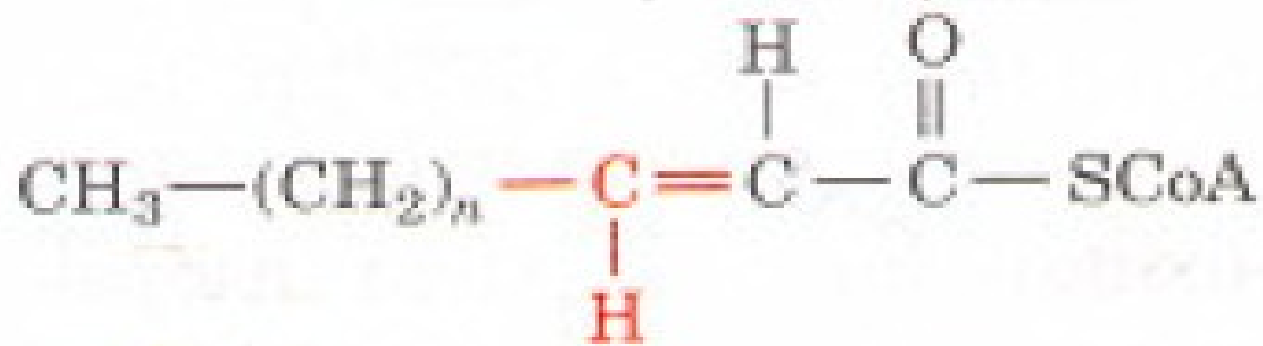
2ADP + 2P_i → 2ATP

Acyl-CoA DH

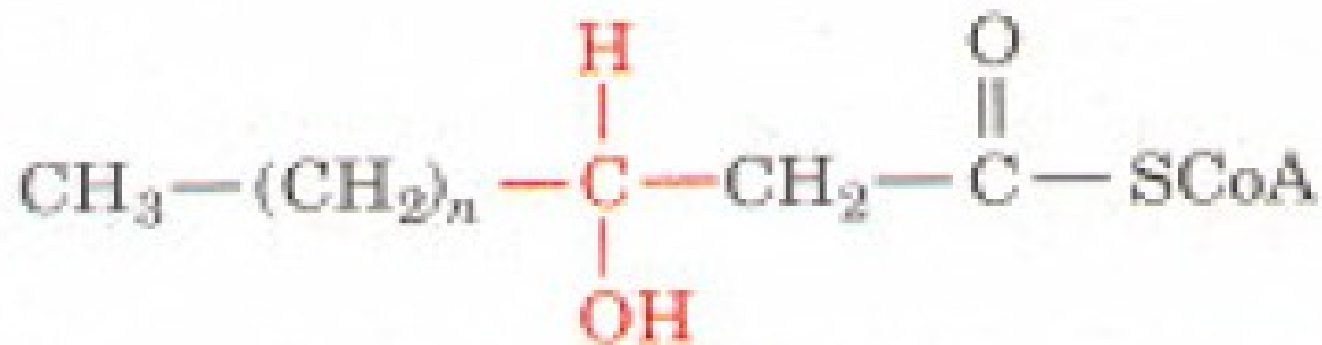
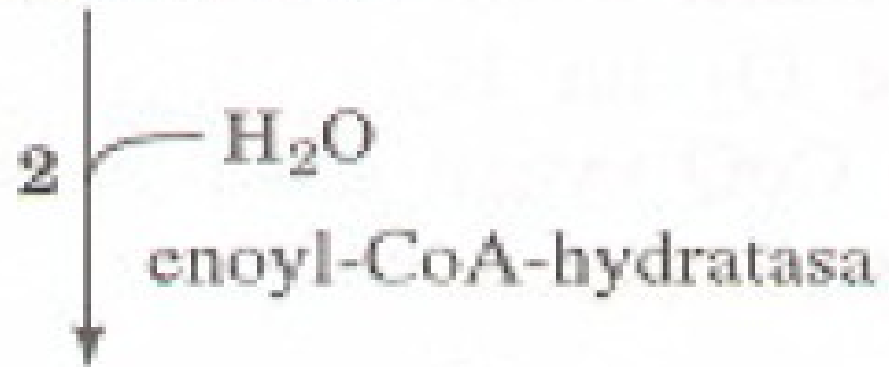
nedostatek

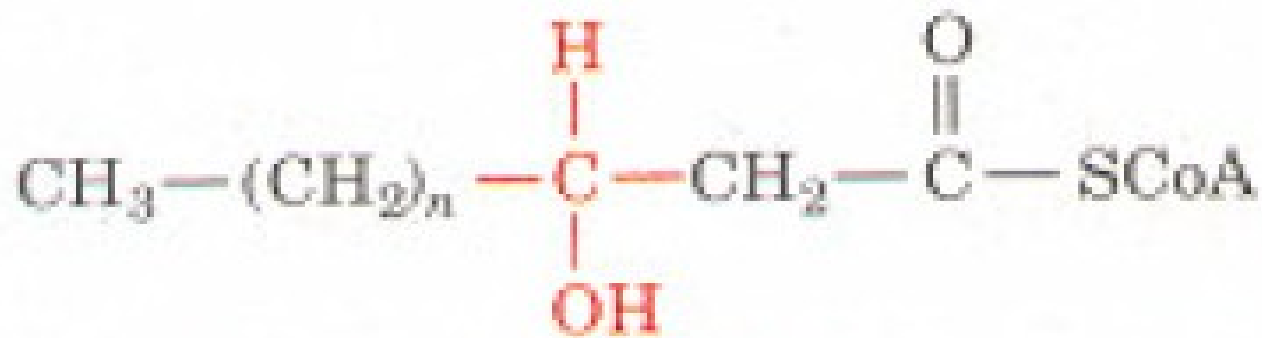
syndrom náhlého dětského úmrtí (sudden infant death syndrome, SIDS)



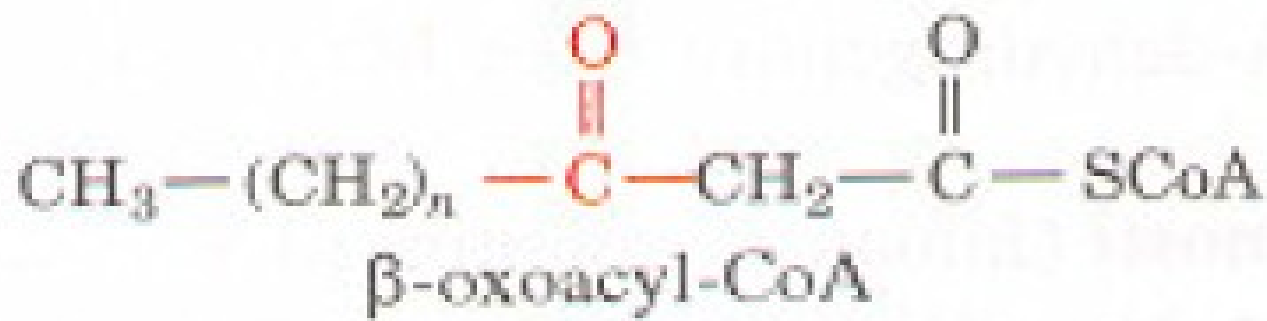
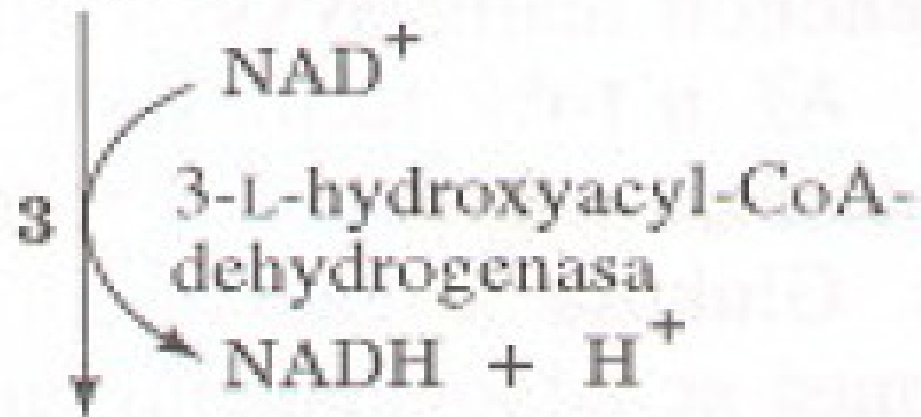


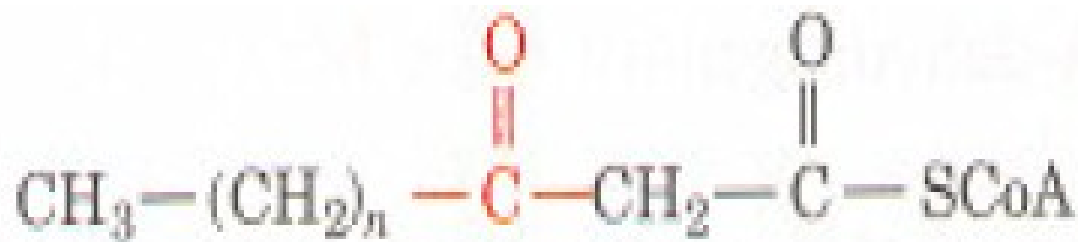
trans- Δ^2 -enoyl-CoA





3-L-hydroxyacyl-CoA



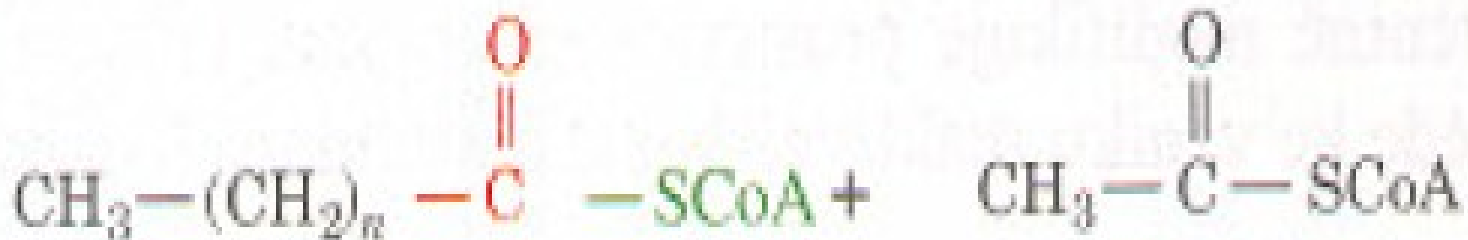


β -oxoacyl-CoA



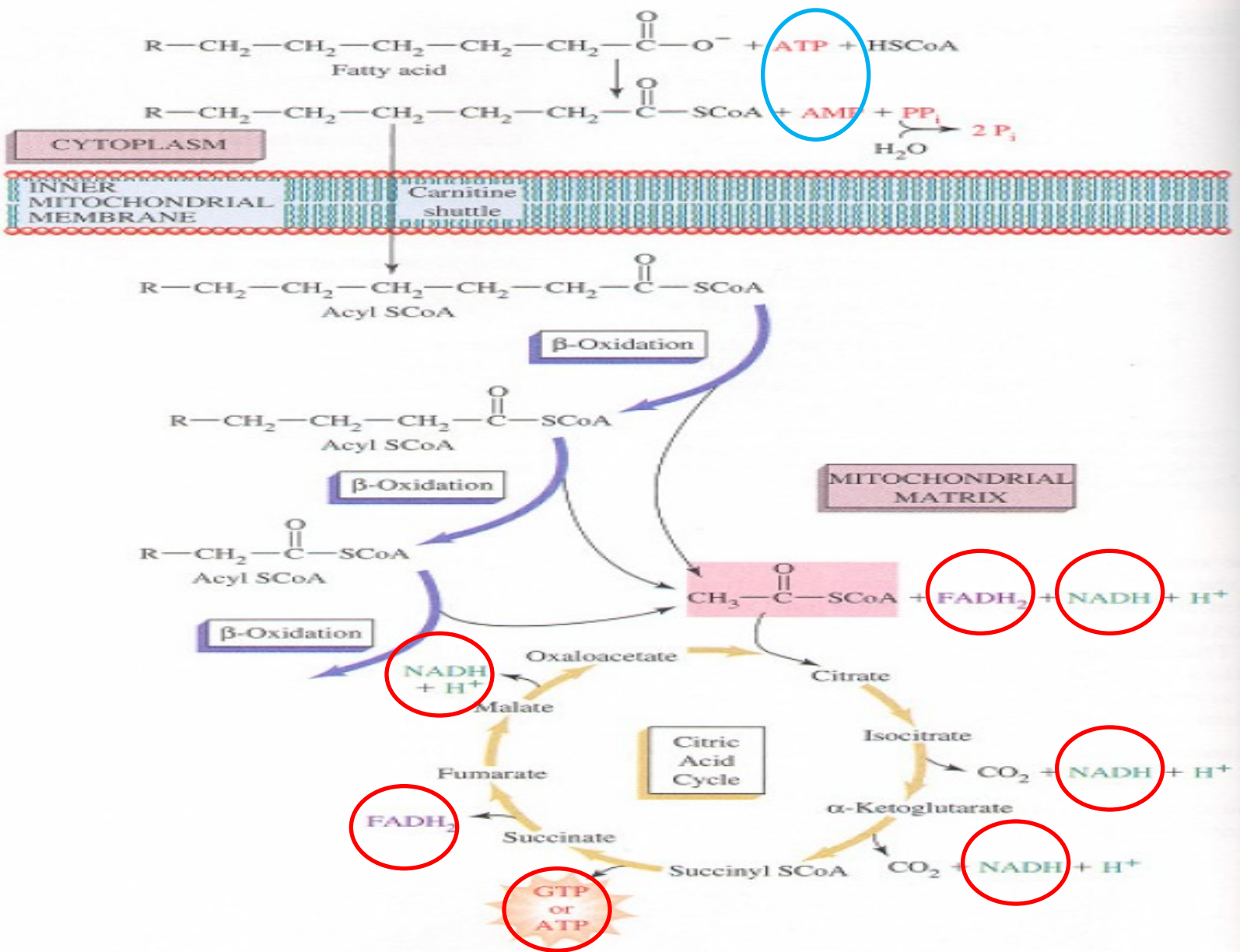
CoASH

β -oxoacyl-CoA-thiolasa



acyl-CoA
(kratší o 2C)

acetyl-CoA

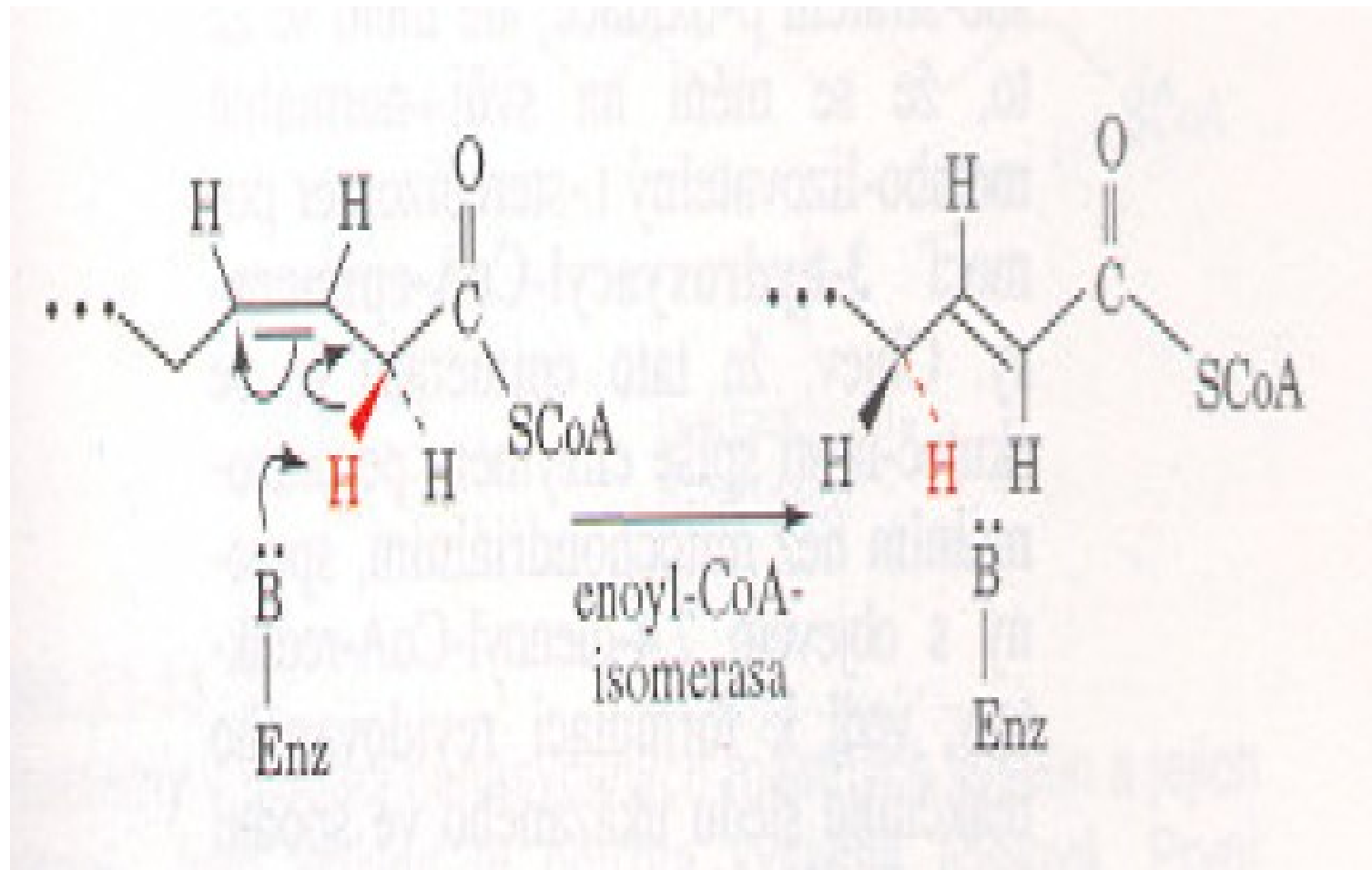


Bilance β oxidace :

1. cyklus - 1 FADH₂ (2 ATP) + 1 NADH (3 ATP) - 5 ATP
acetylCoA (citrátový cyklus) - 12 ATP

na C₁₆ - 7 x β oxidace + 8 x citrátový cyklus - aktivace
(7 x 5) + (8 x 12) - 2 ATP = 129 ATP

Odbourávání nenasycených mastných kyselin





three cycles of β -oxidation



enoyl-CoA isomerase



complete β -oxidation cycle



acyl-CoA dehydrogenase



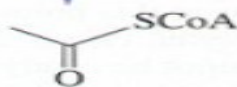
$\text{NADPH} + \text{H}^+ \rightarrow \text{NADP}^+$ (catalyzed by 2,4-dienoyl-CoA reductase)

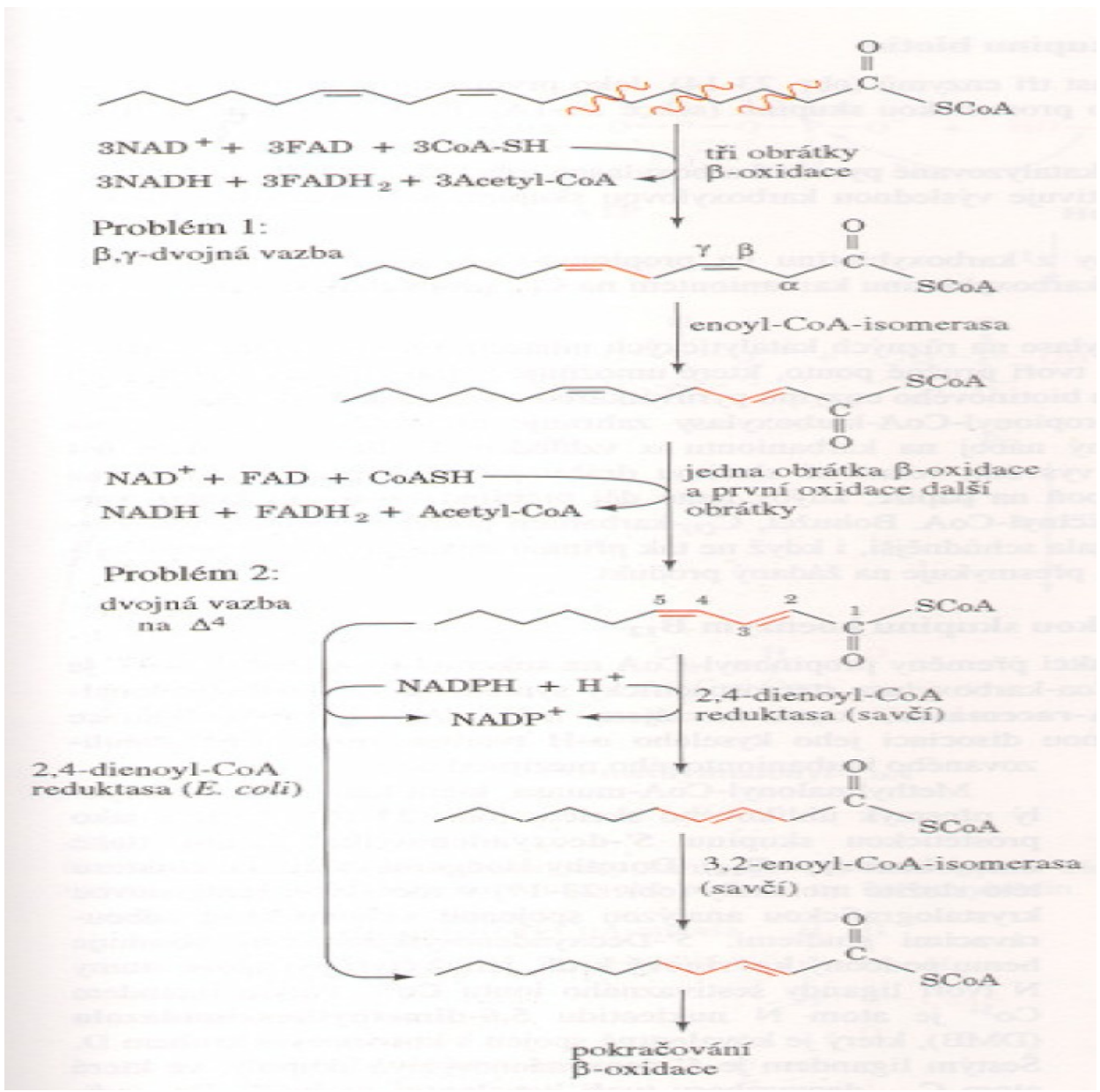


enoyl-CoA isomerase

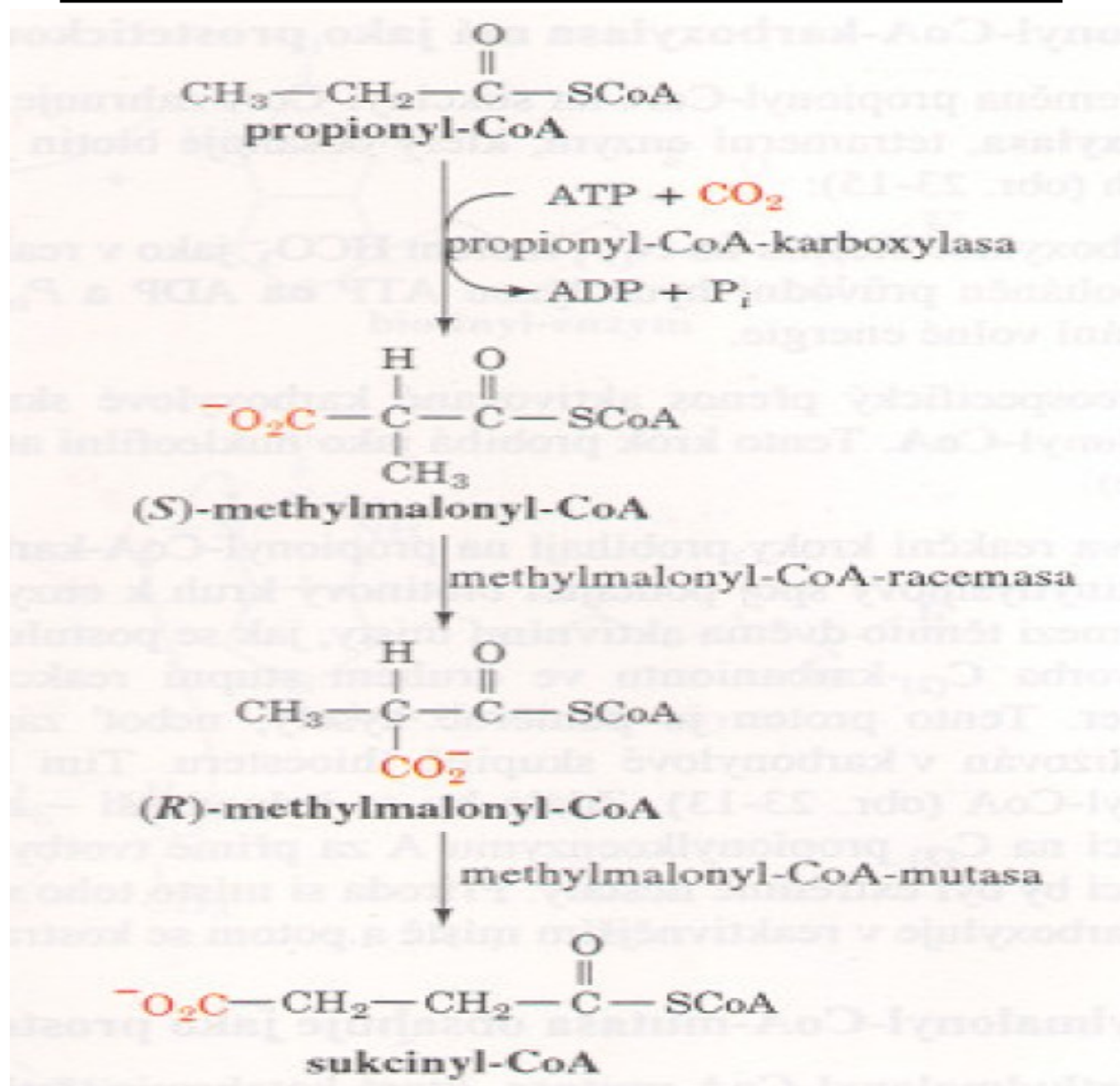


three cycles of β -oxidation

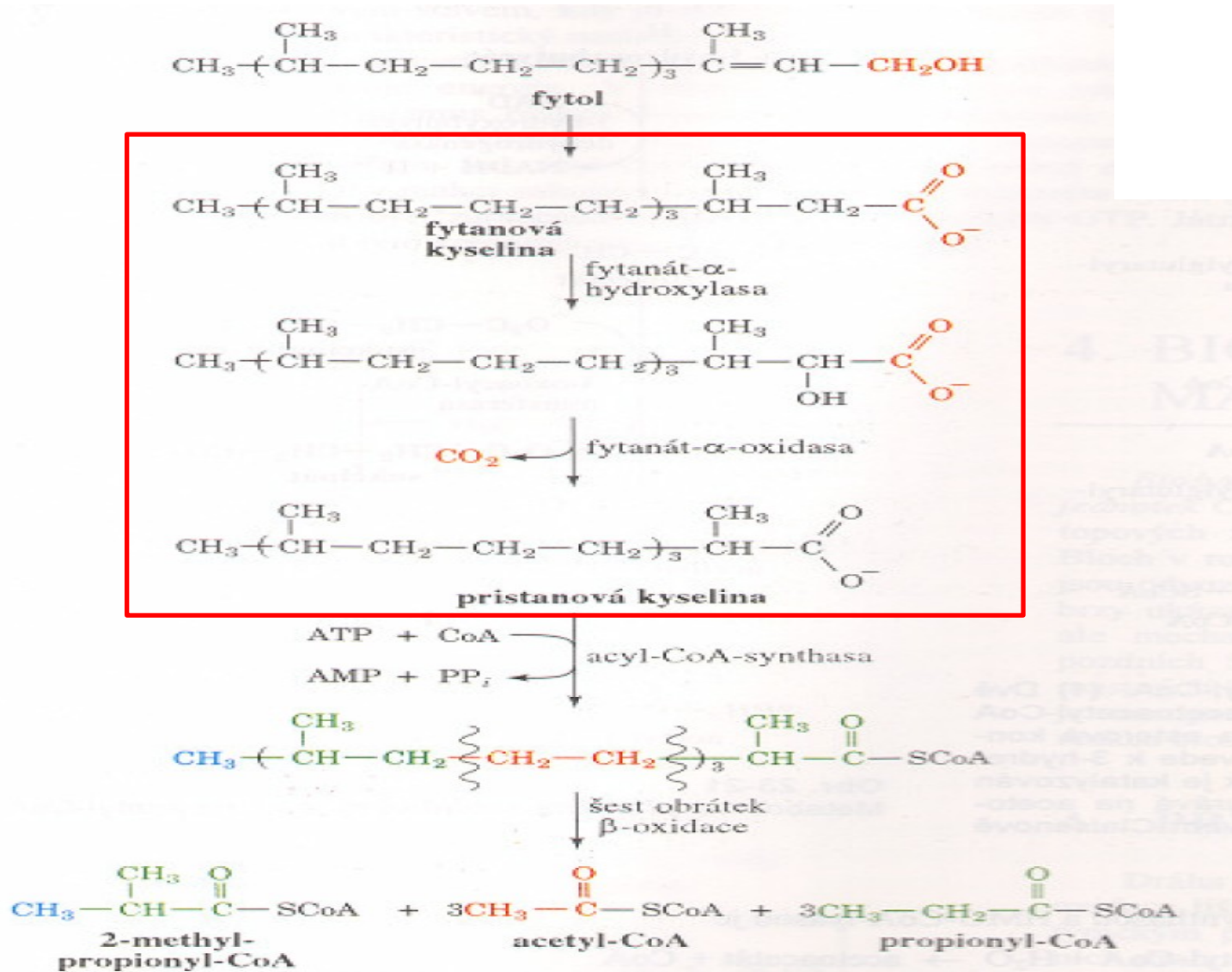


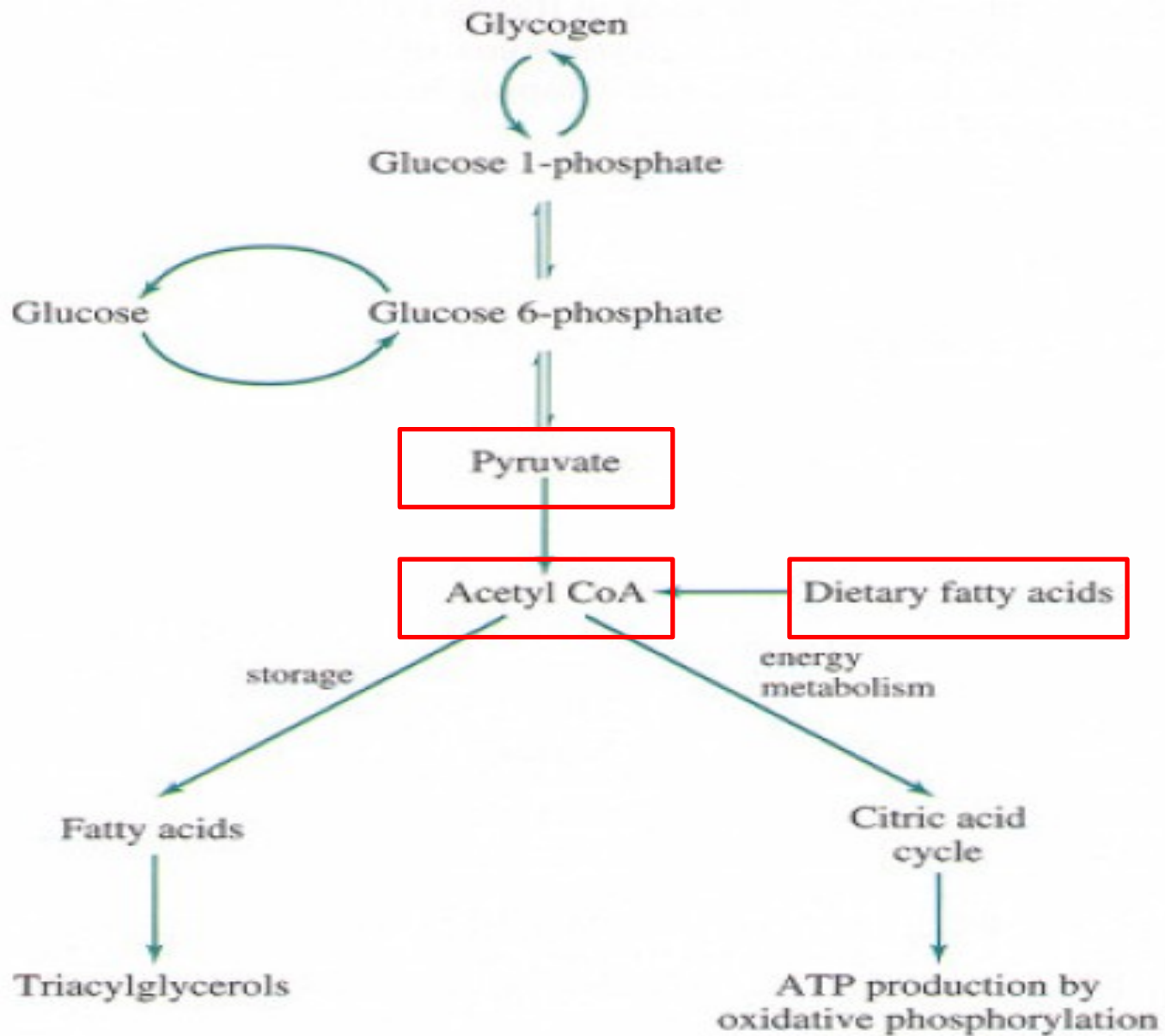


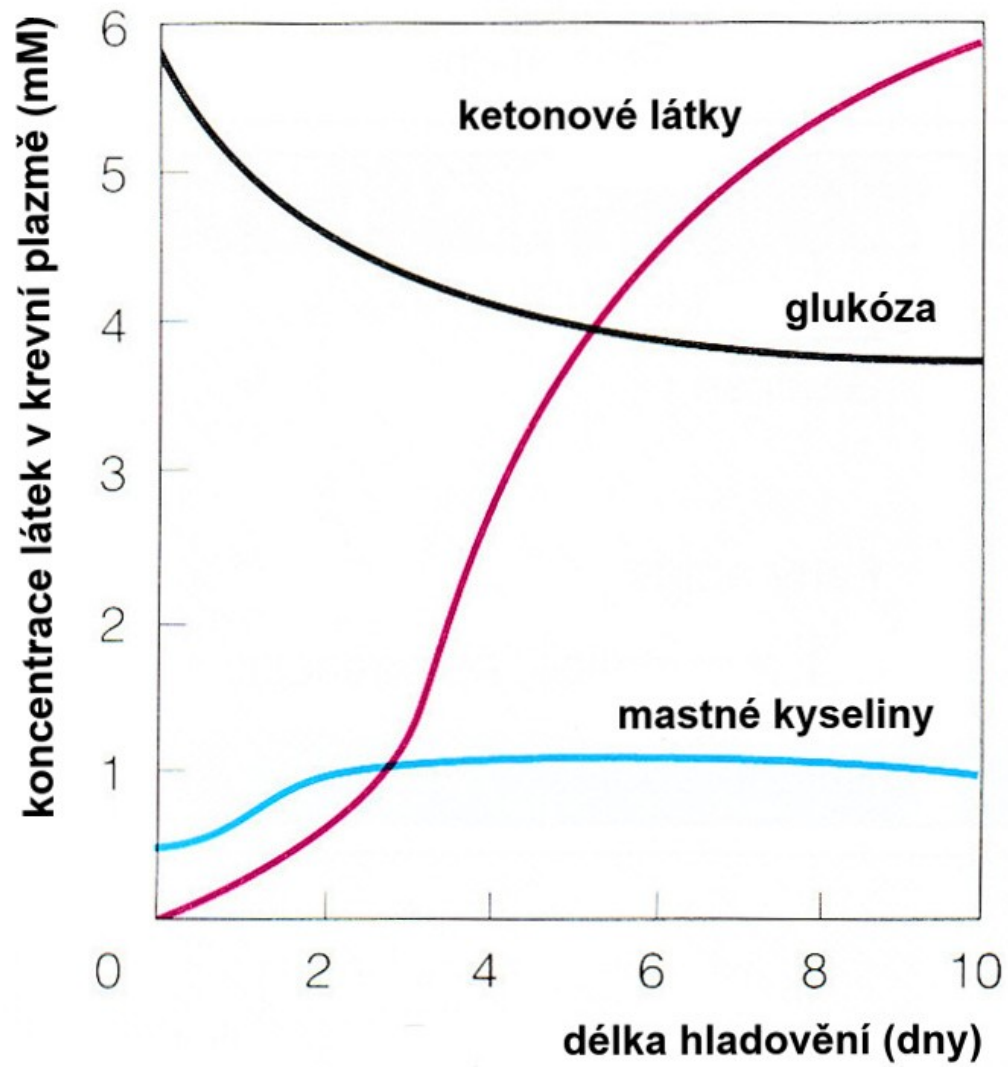
Odbourávání mastných kyselin s lichým počtem C atomů



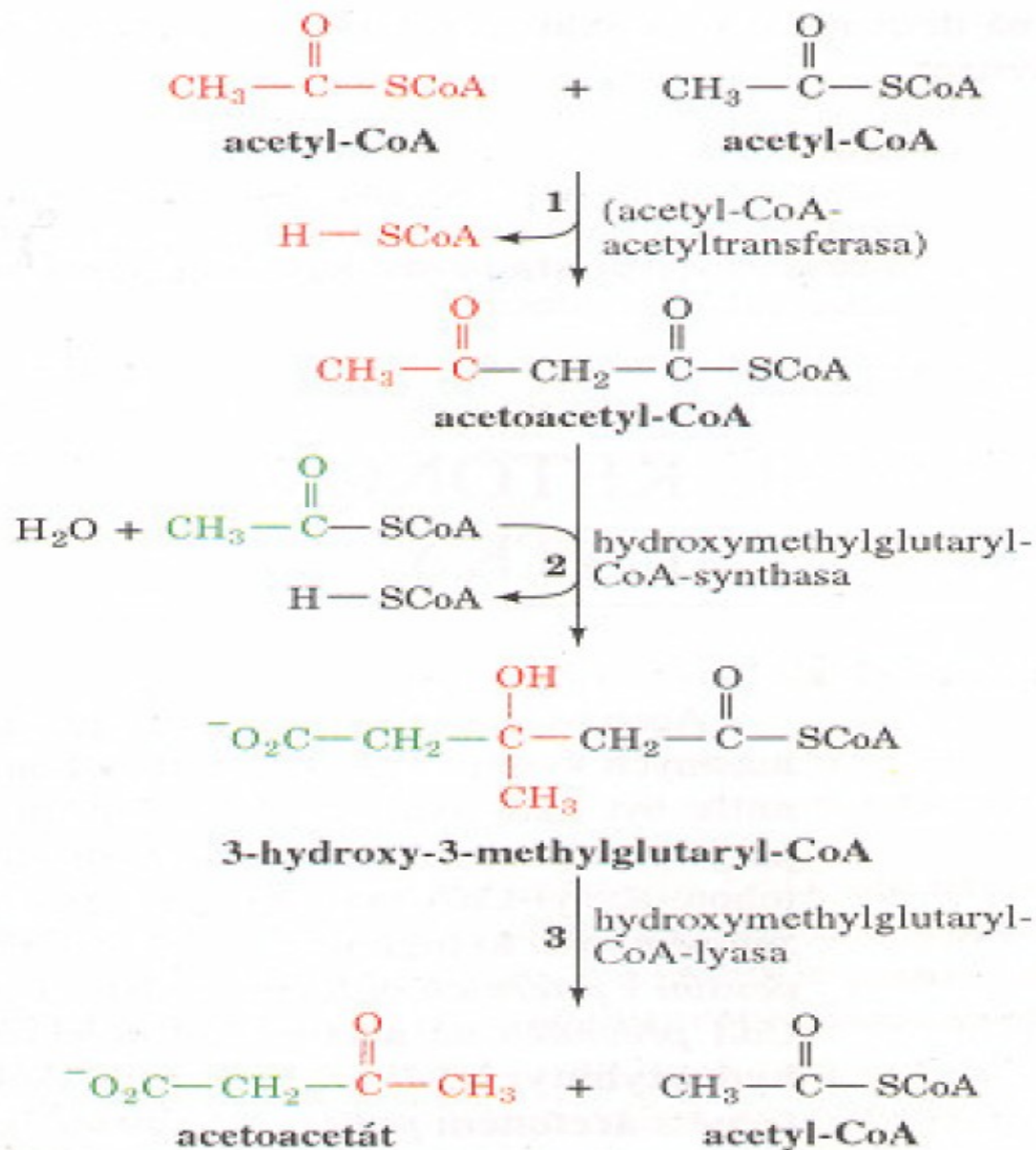
α -oxidace



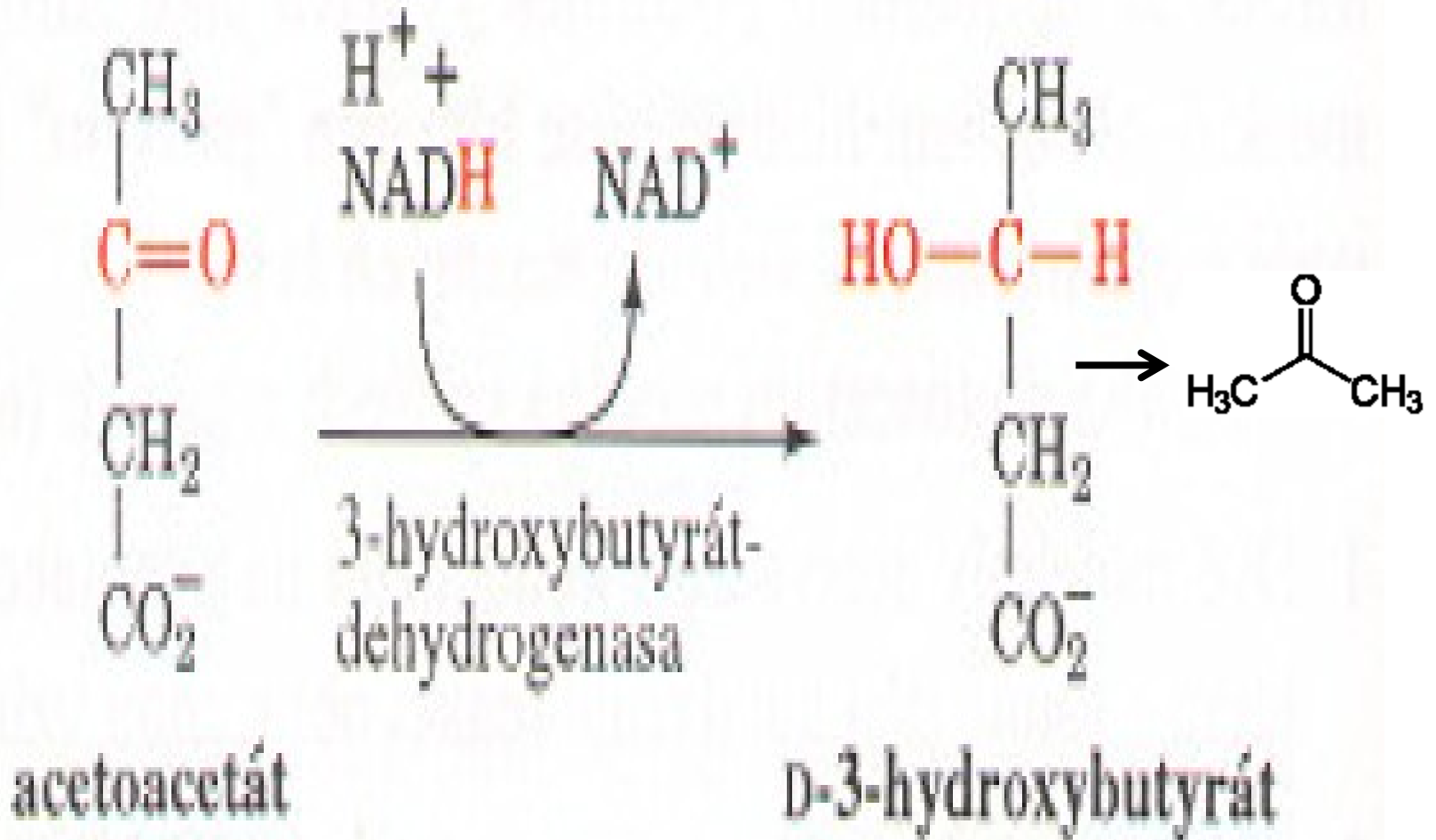




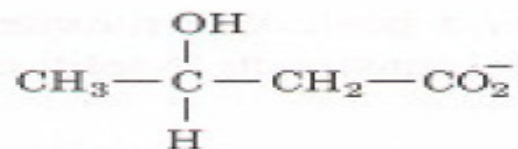
Ketonové látky



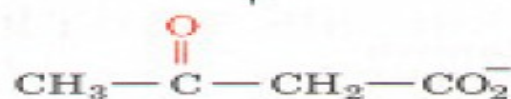
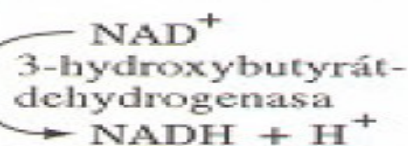
Ketonové látky



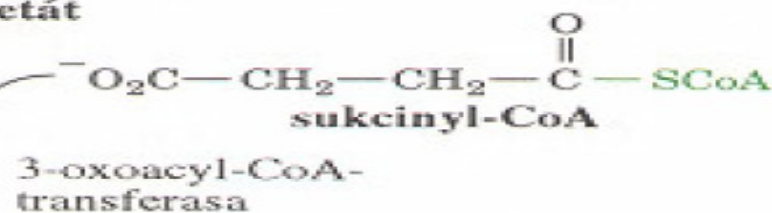
Ketonové látky



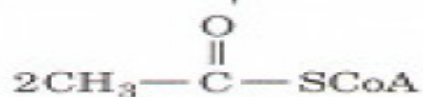
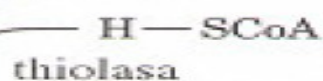
D-3-hydroxybutyrát



acetoacetát

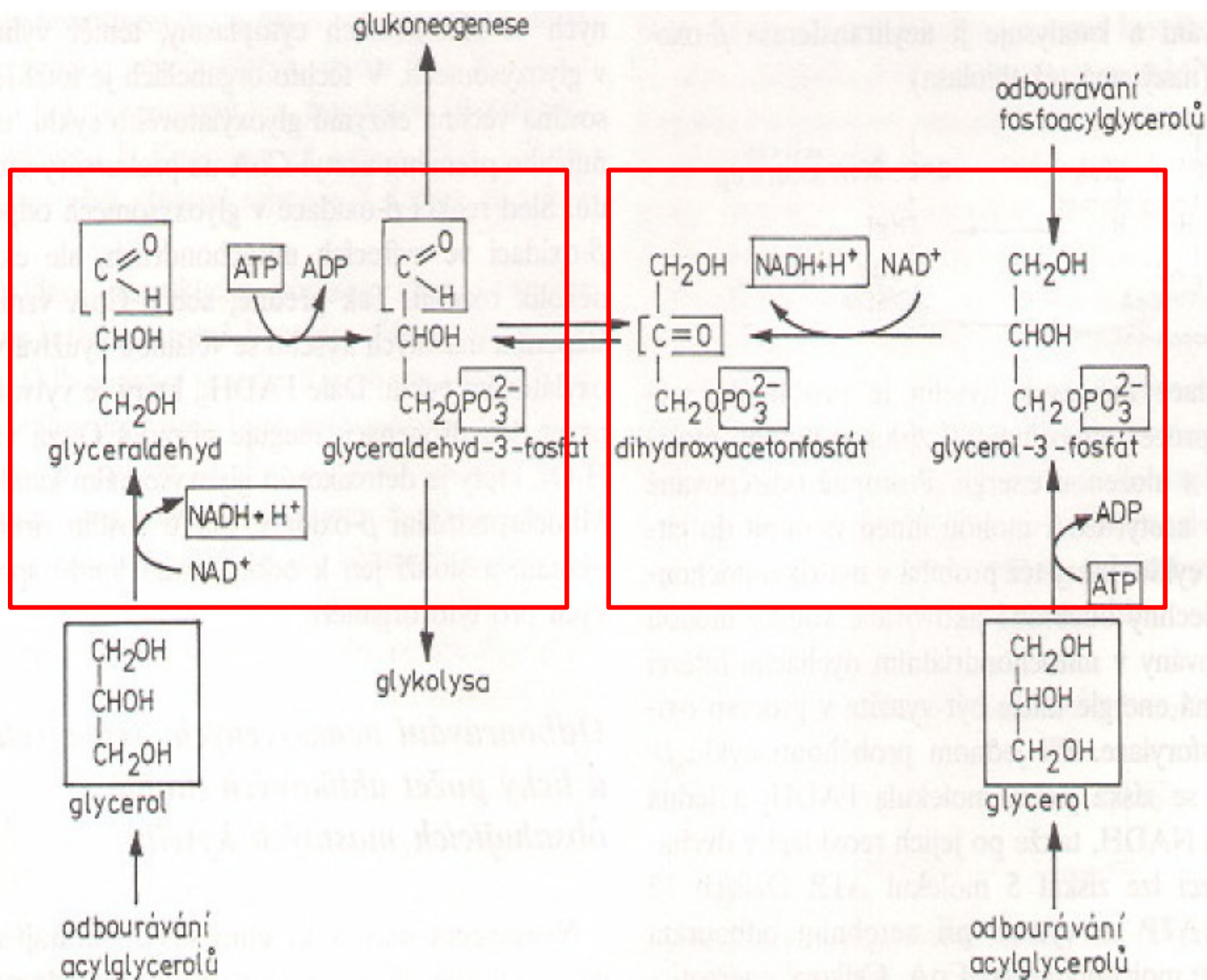


acetoacetyl-CoA

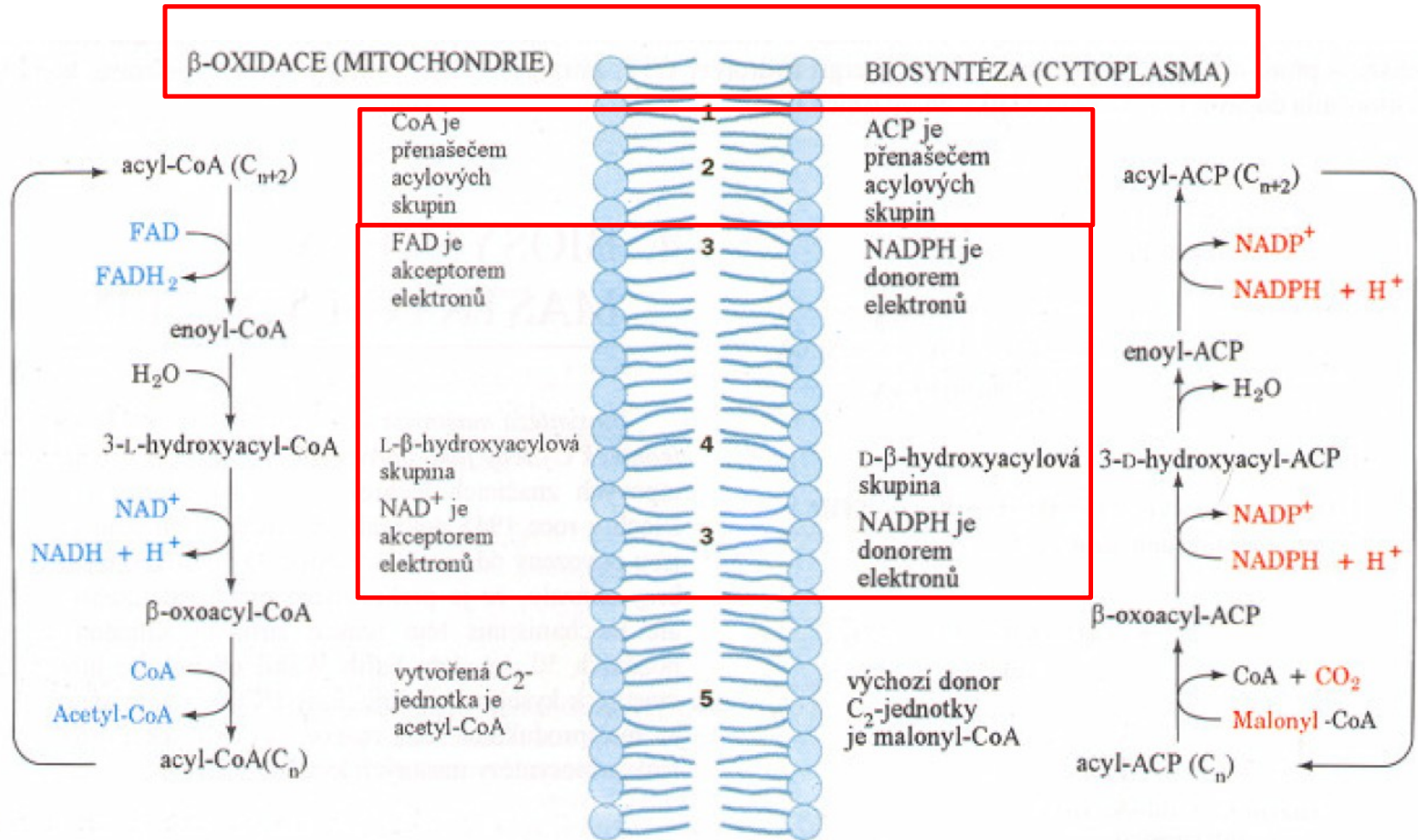


acetyl-CoA

Metabolismus glycerolu

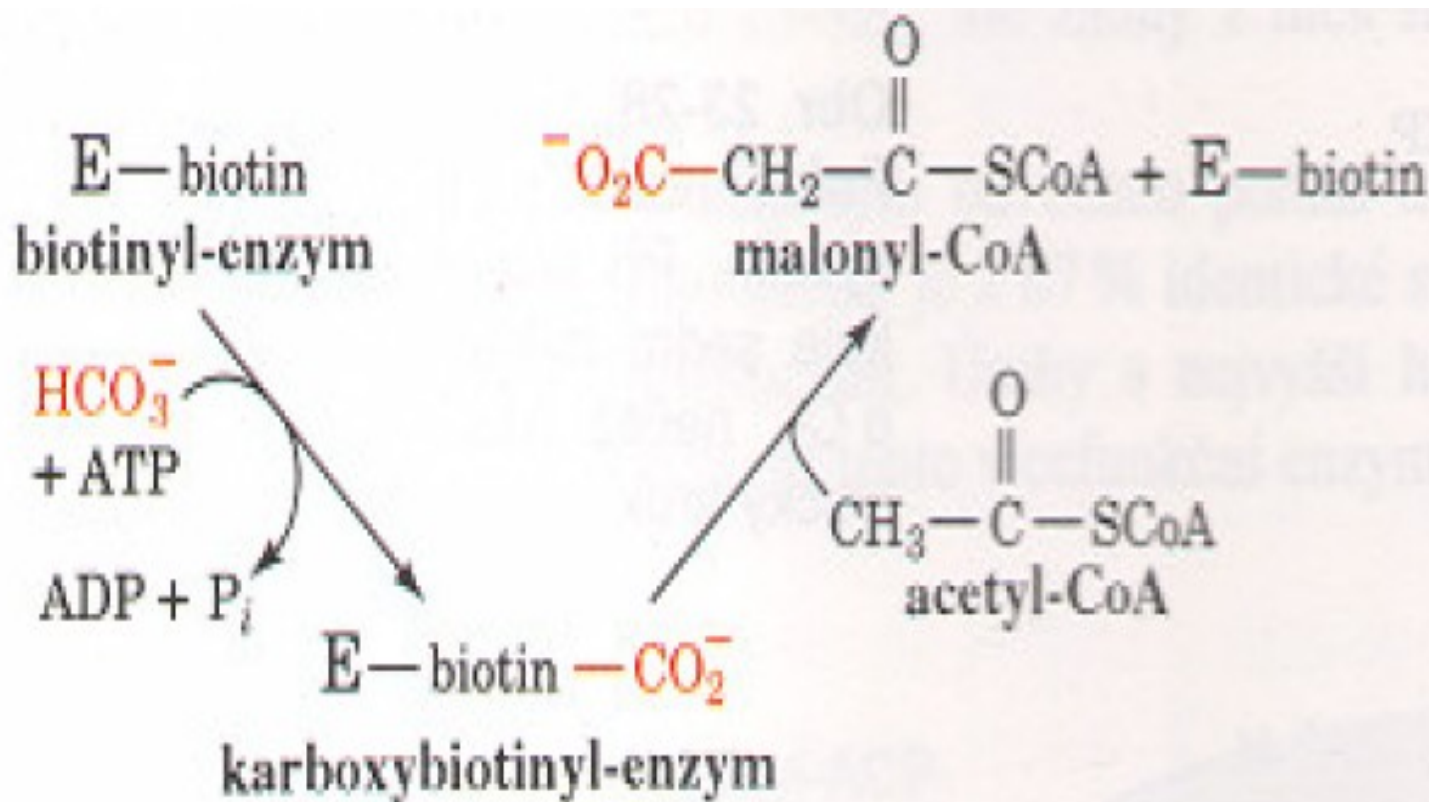


Metabolismus versus biosyntéza MK

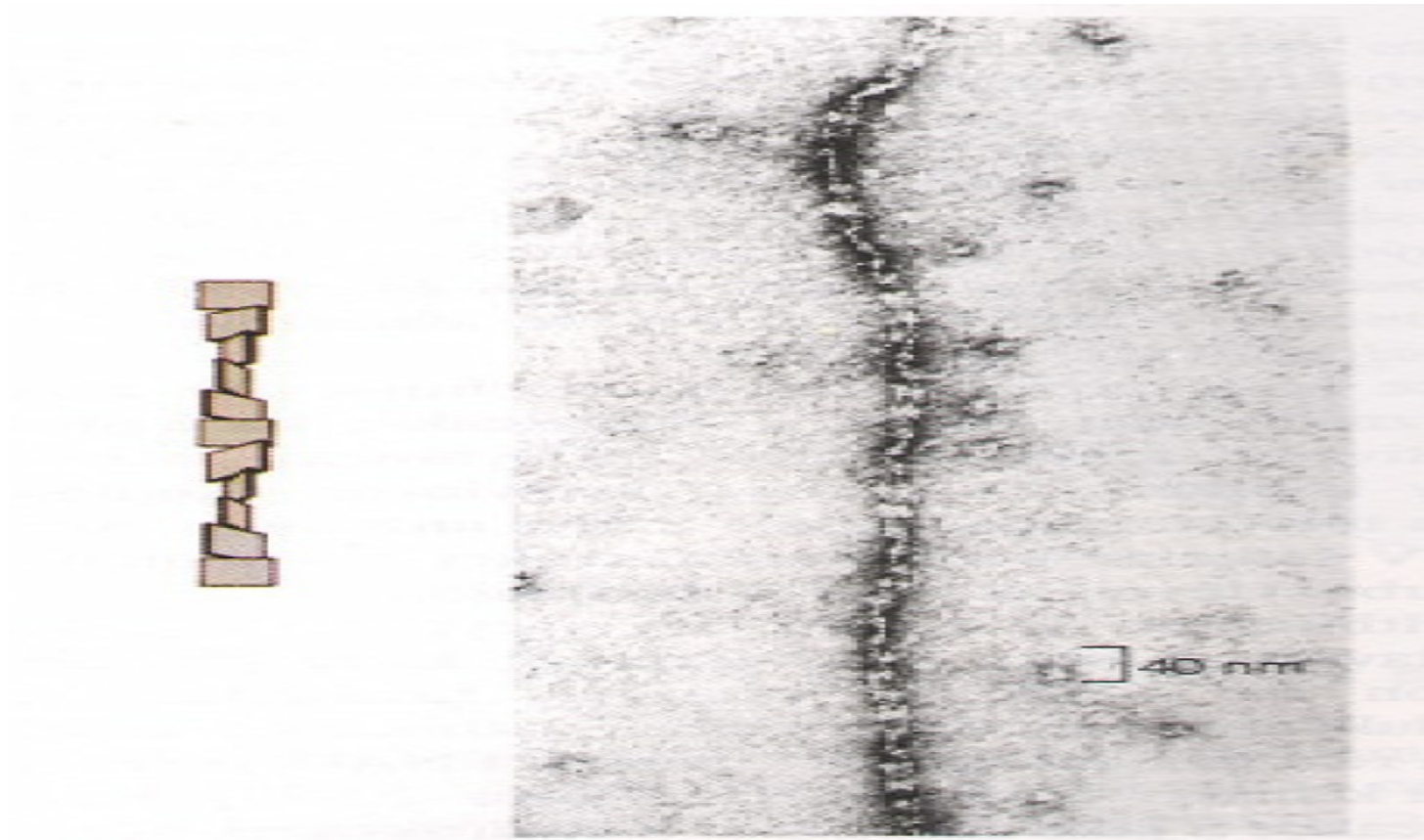


Biosyntéza mastných kyselin

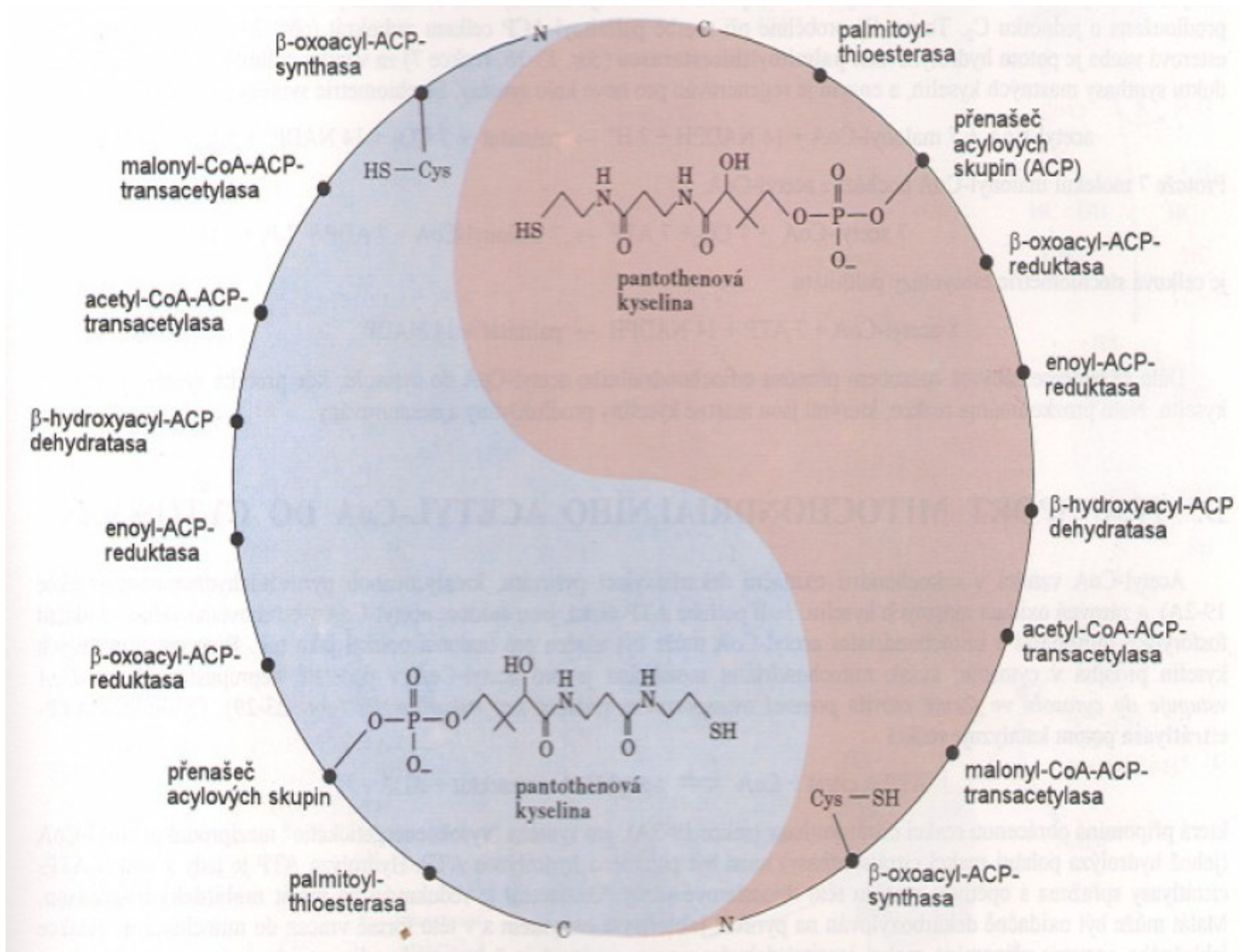
A. Syntéza malonylCoA



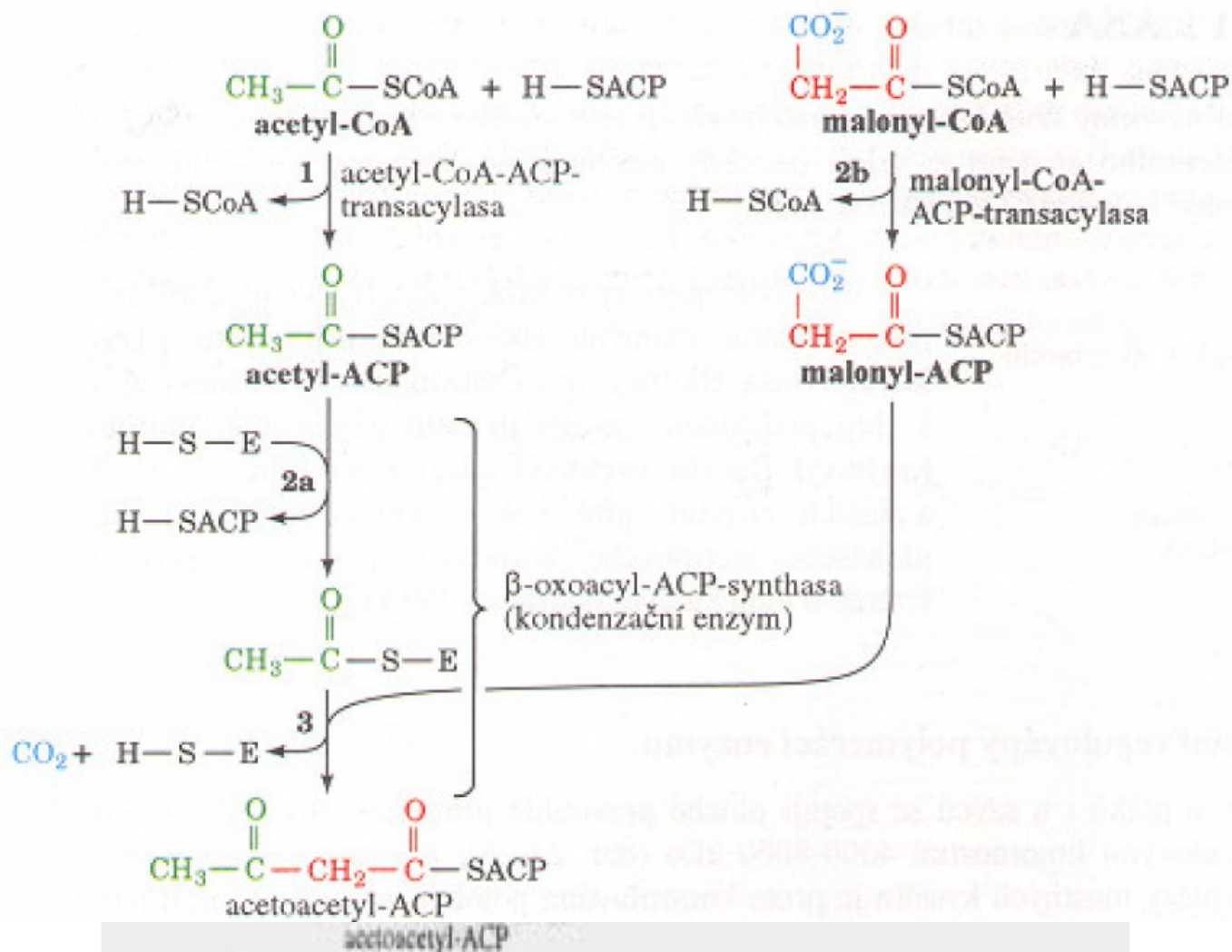
Regulace Acetyl-CoA karboxylasy

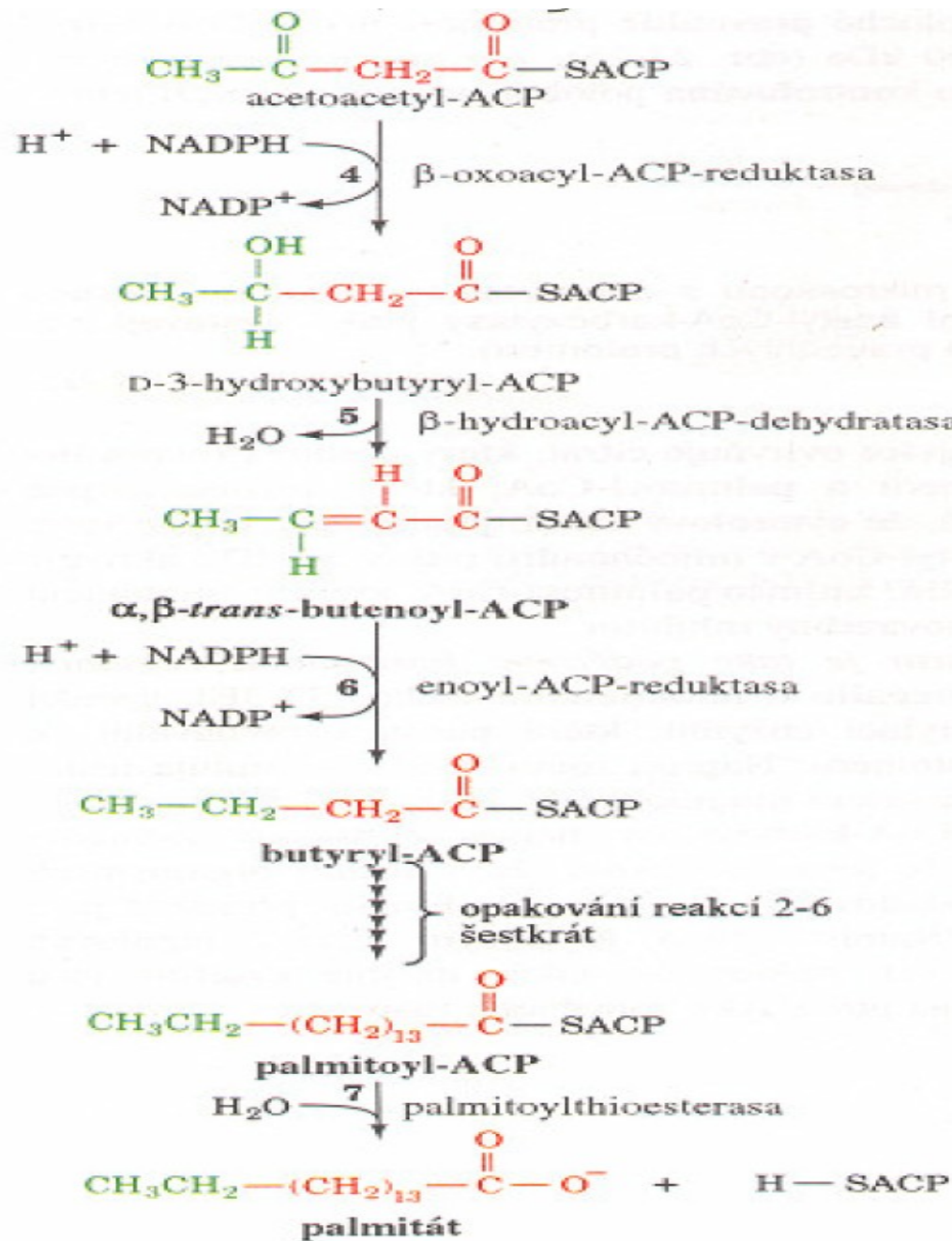


Synthasa mastných kyselin

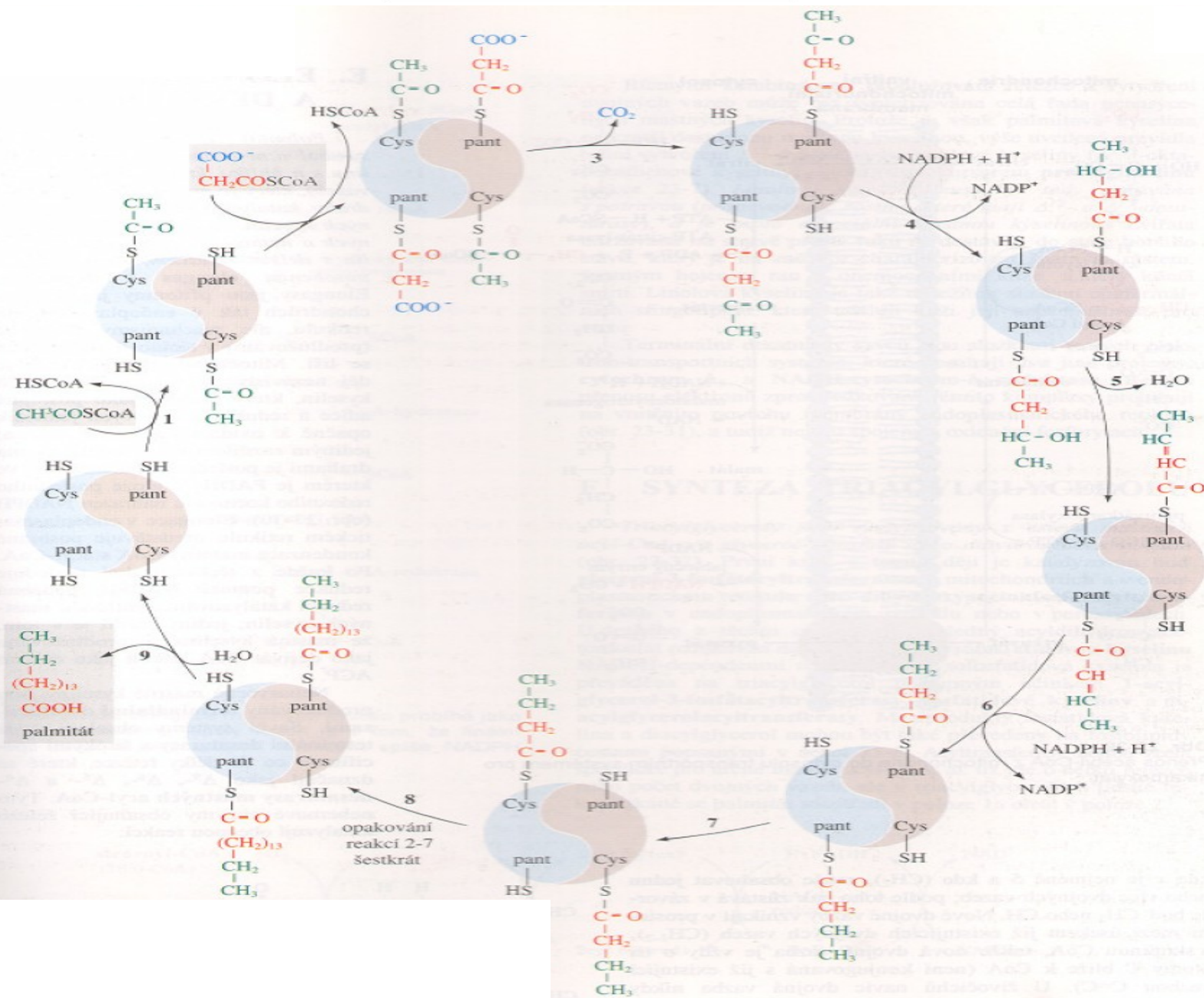


B. Syntéza palmitové kyseliny





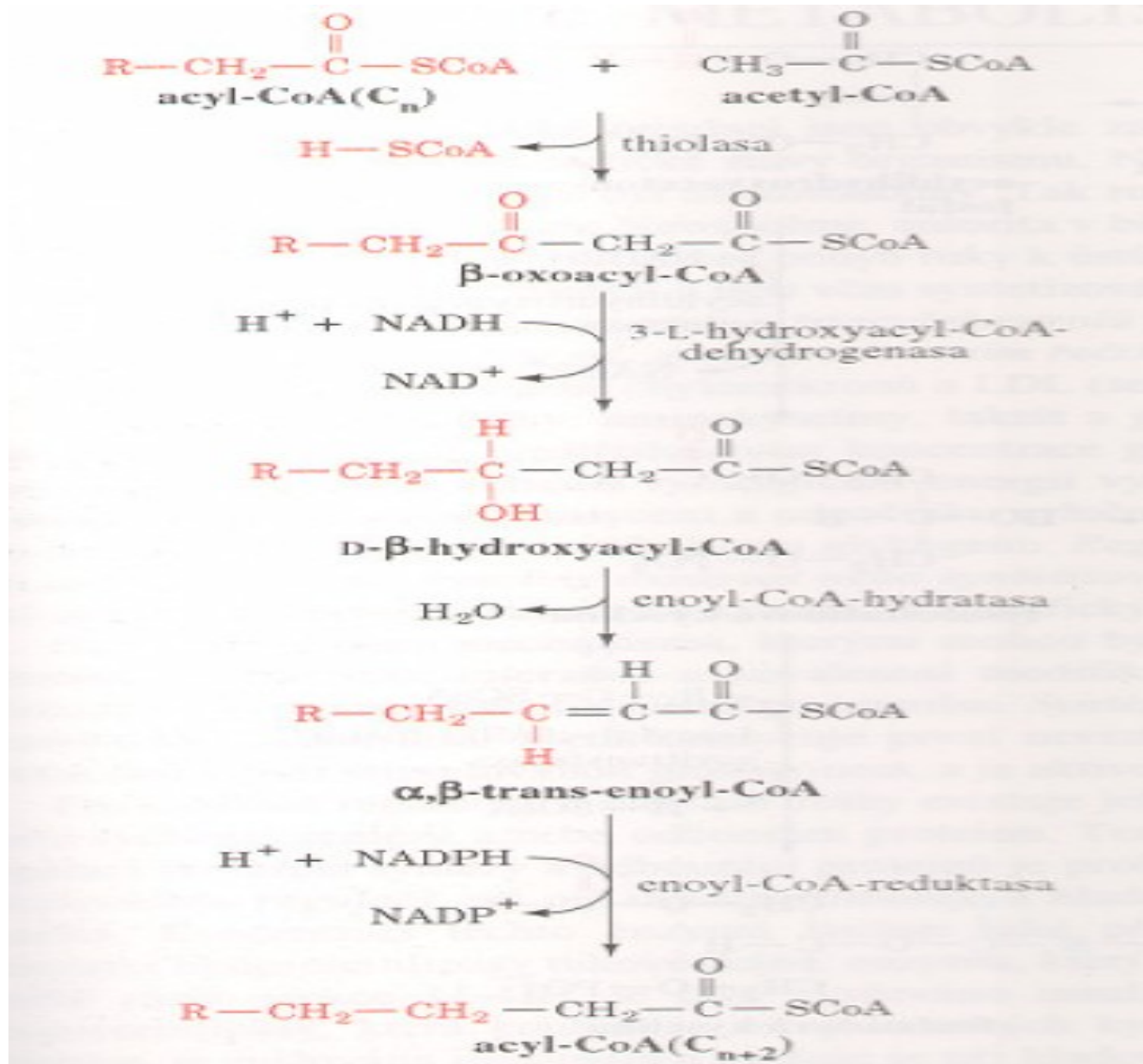
Synthesa mastných kyselin



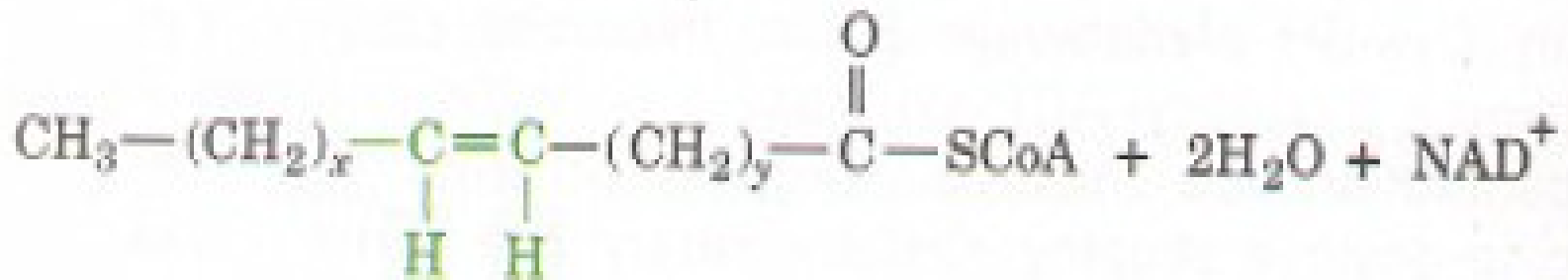
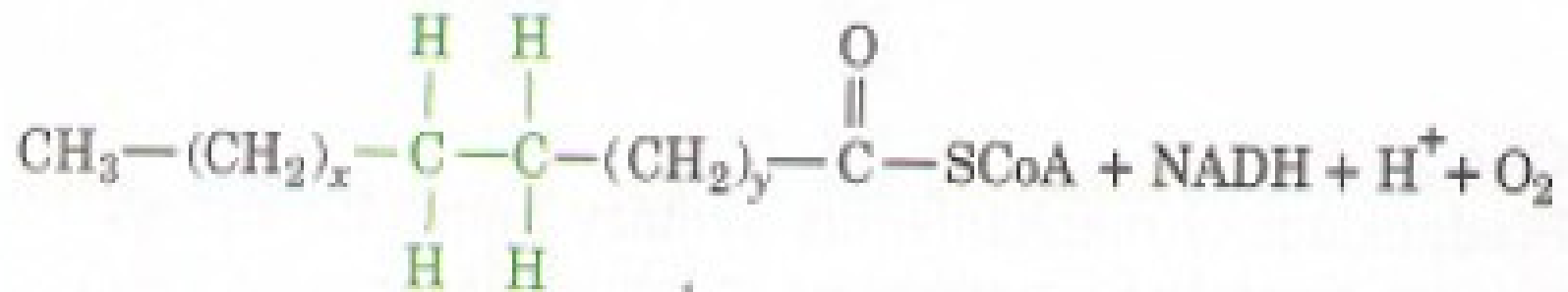
C. Další přeměny palmitové kyseliny

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

Elongace



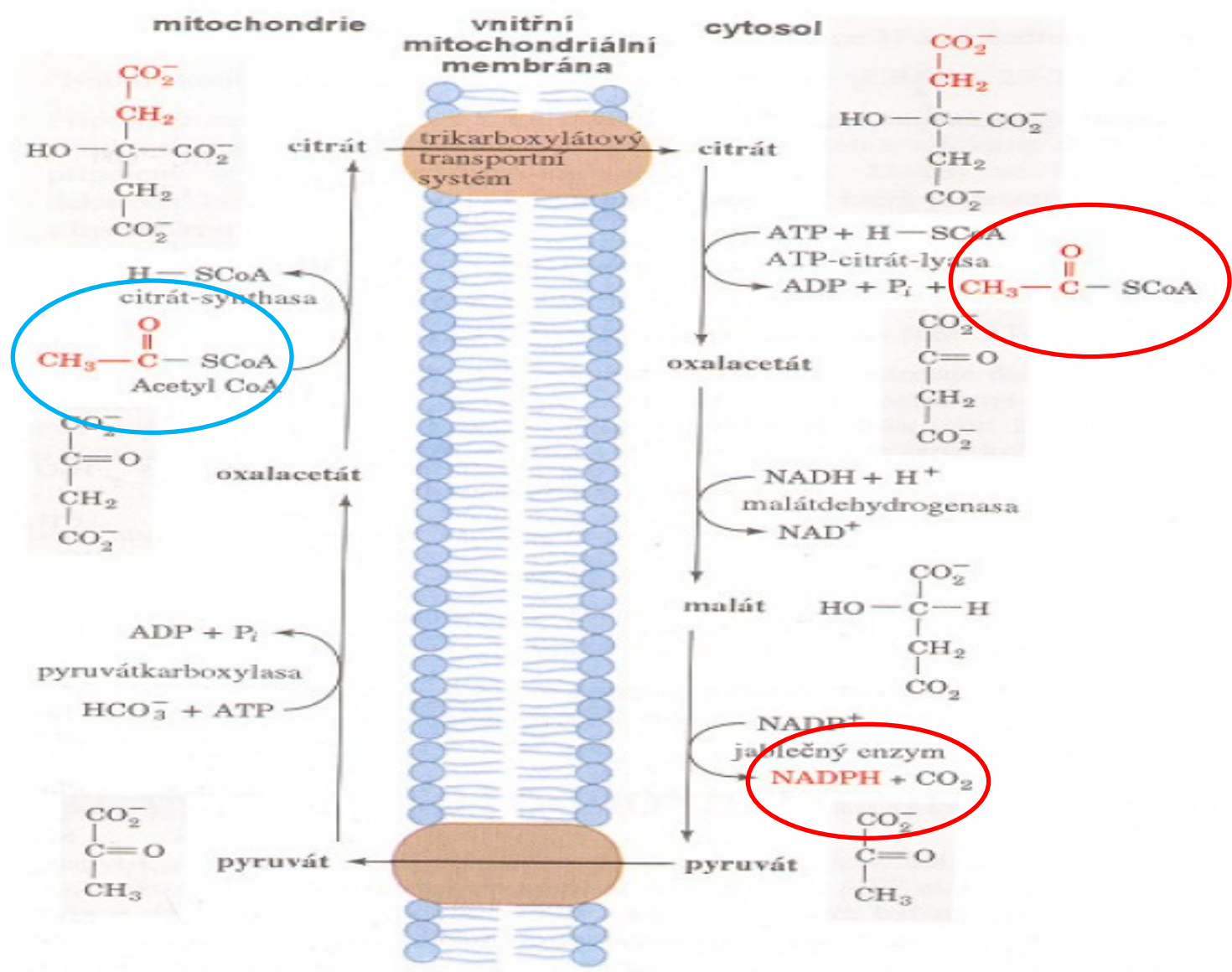
Desaturase



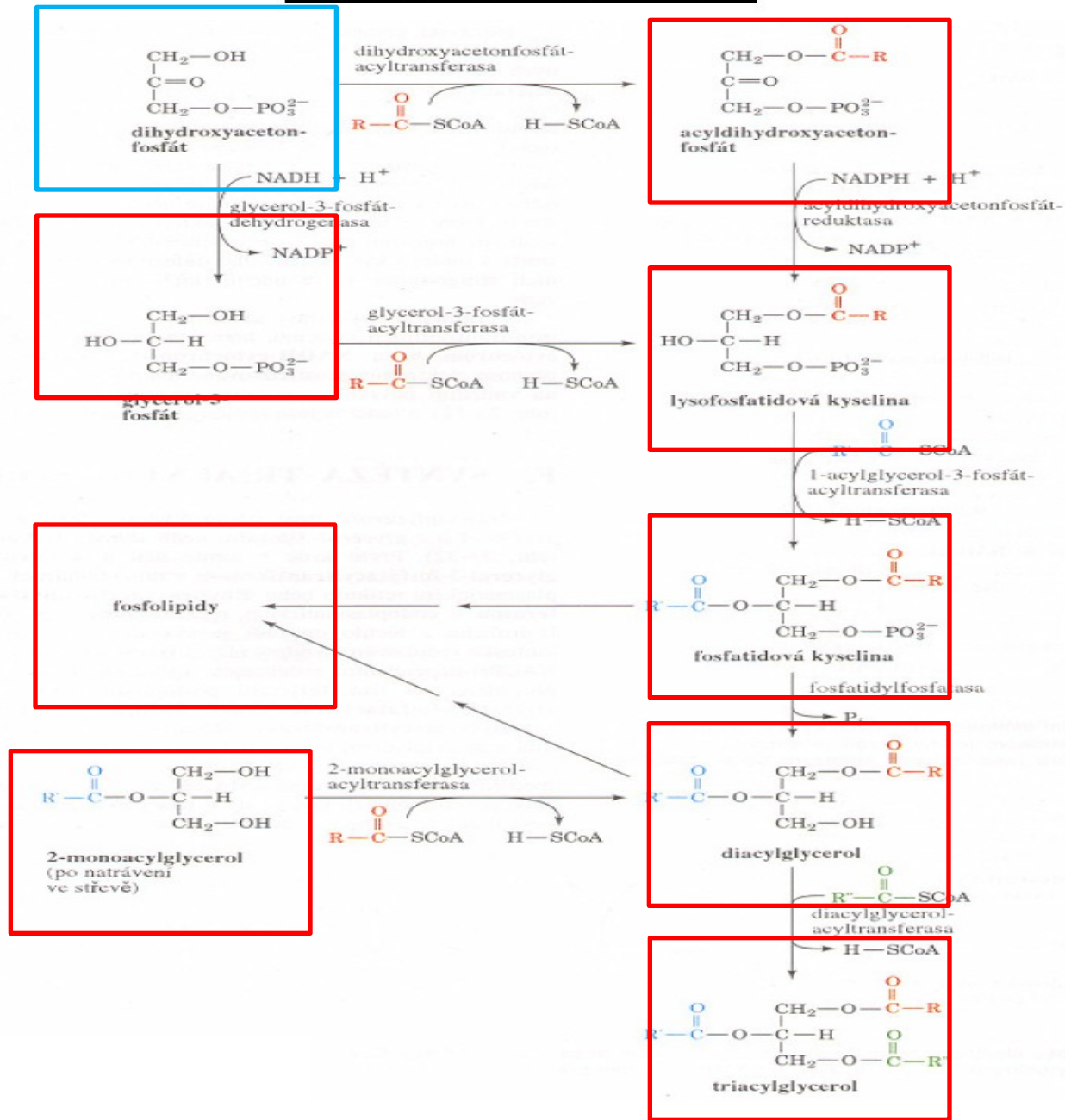
Bilance biosyntézy mastných kyselin :

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

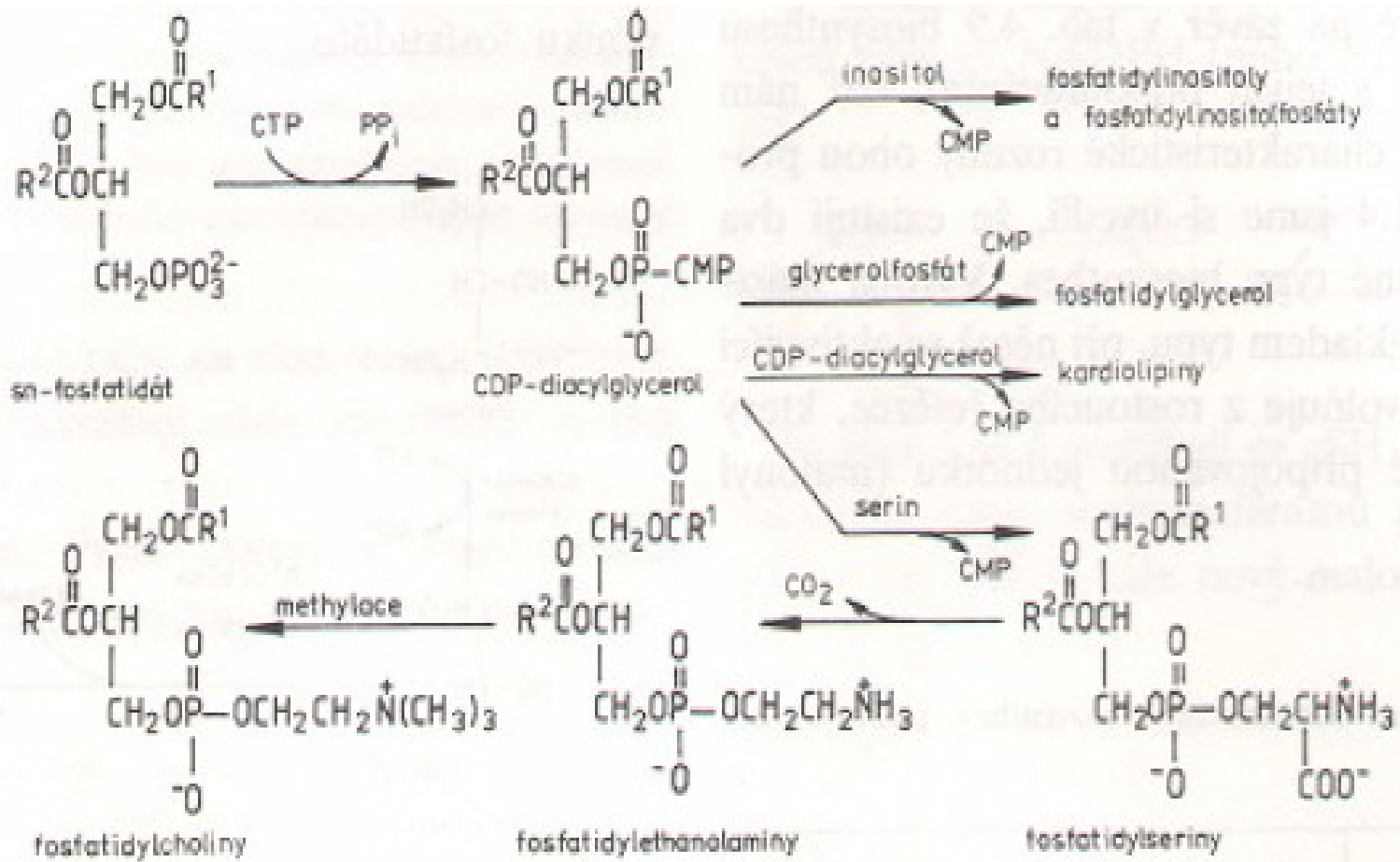
Přenos AcetCoA vně mitochondrie



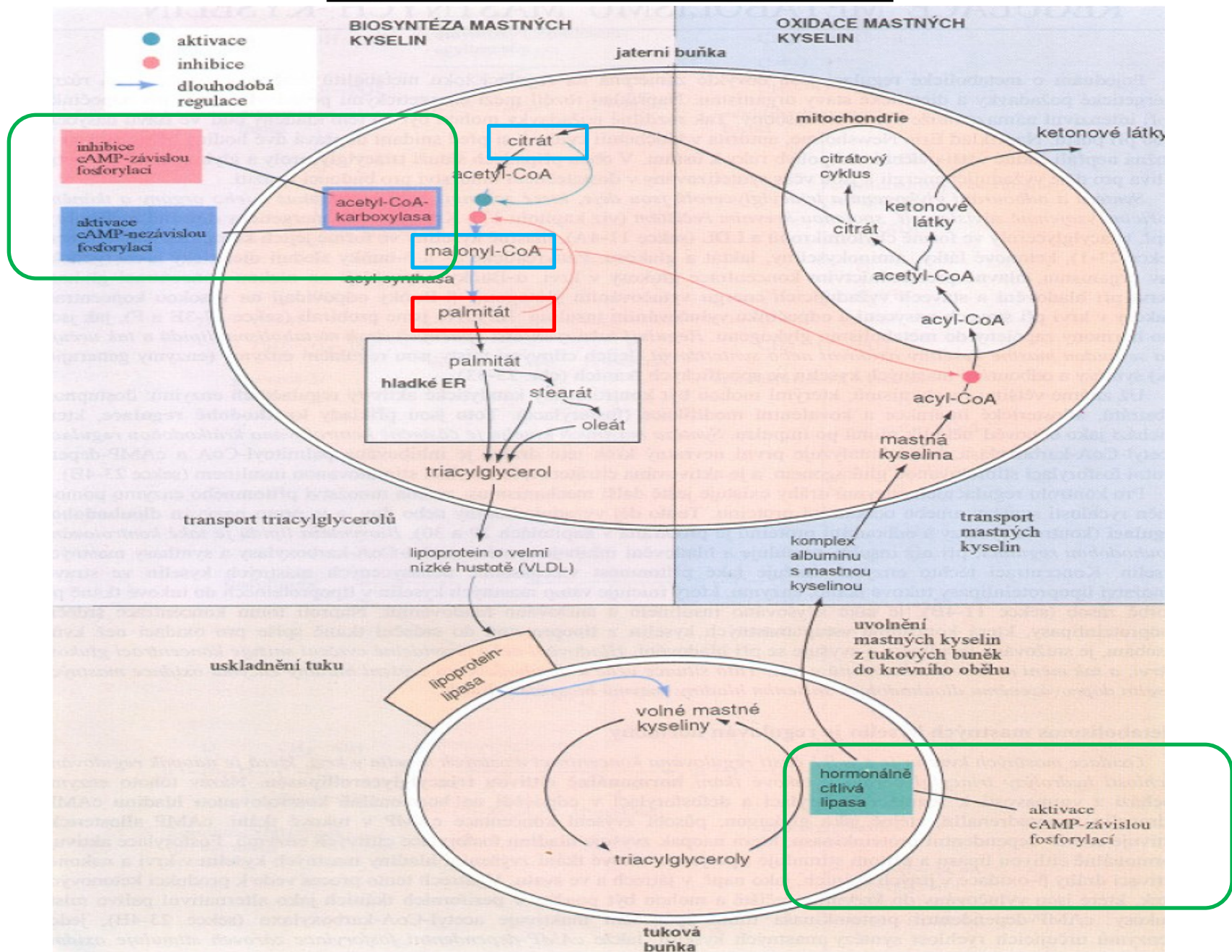
Biosyntéza triacylglycerolů



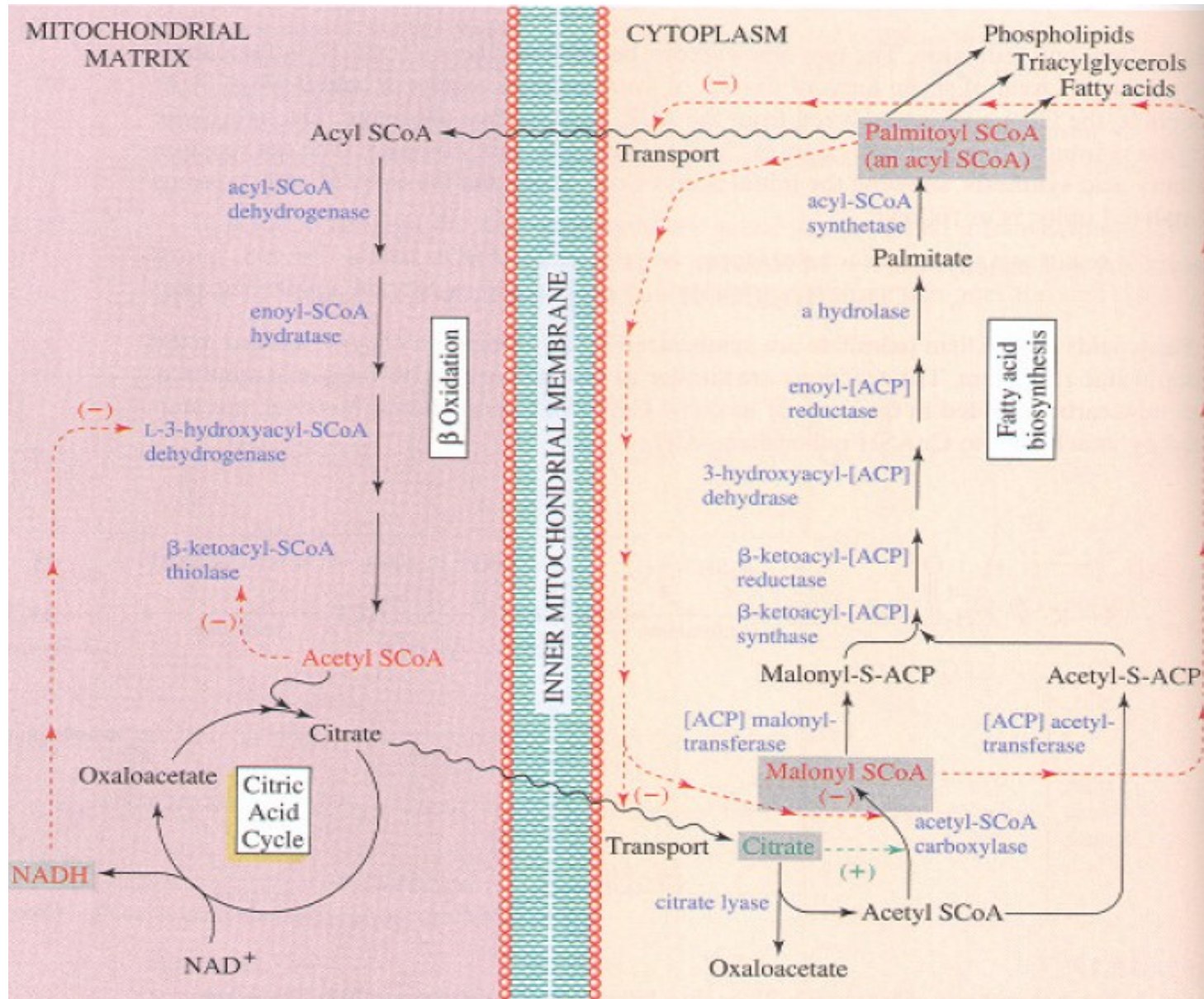
Biosyntéza fosfolipidů



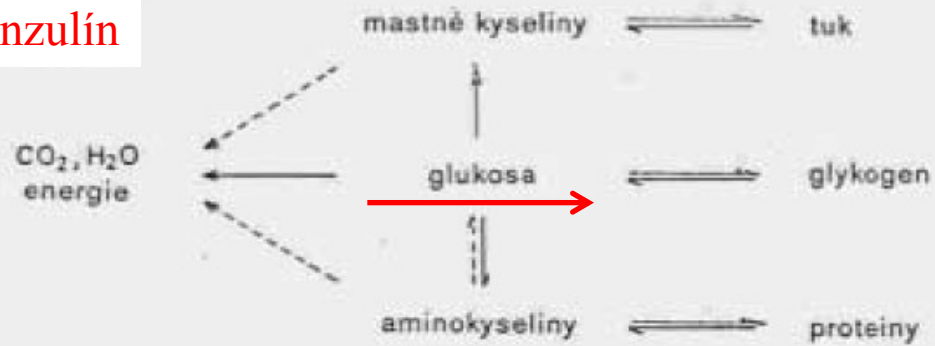
Regulace metabolismu triacylglycerolů



Regulate metabolismu triacylglycerolů



inzulín



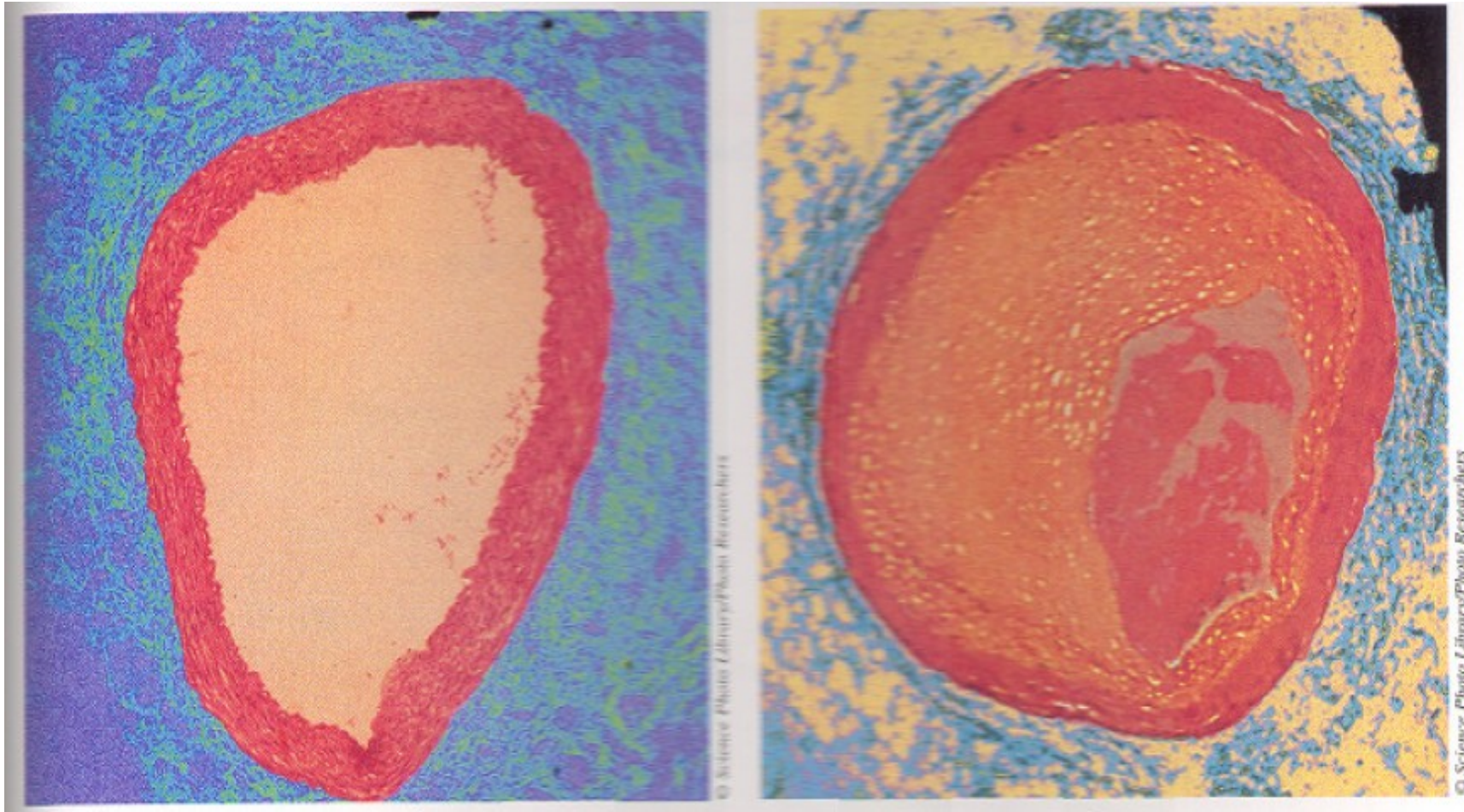
glukagon



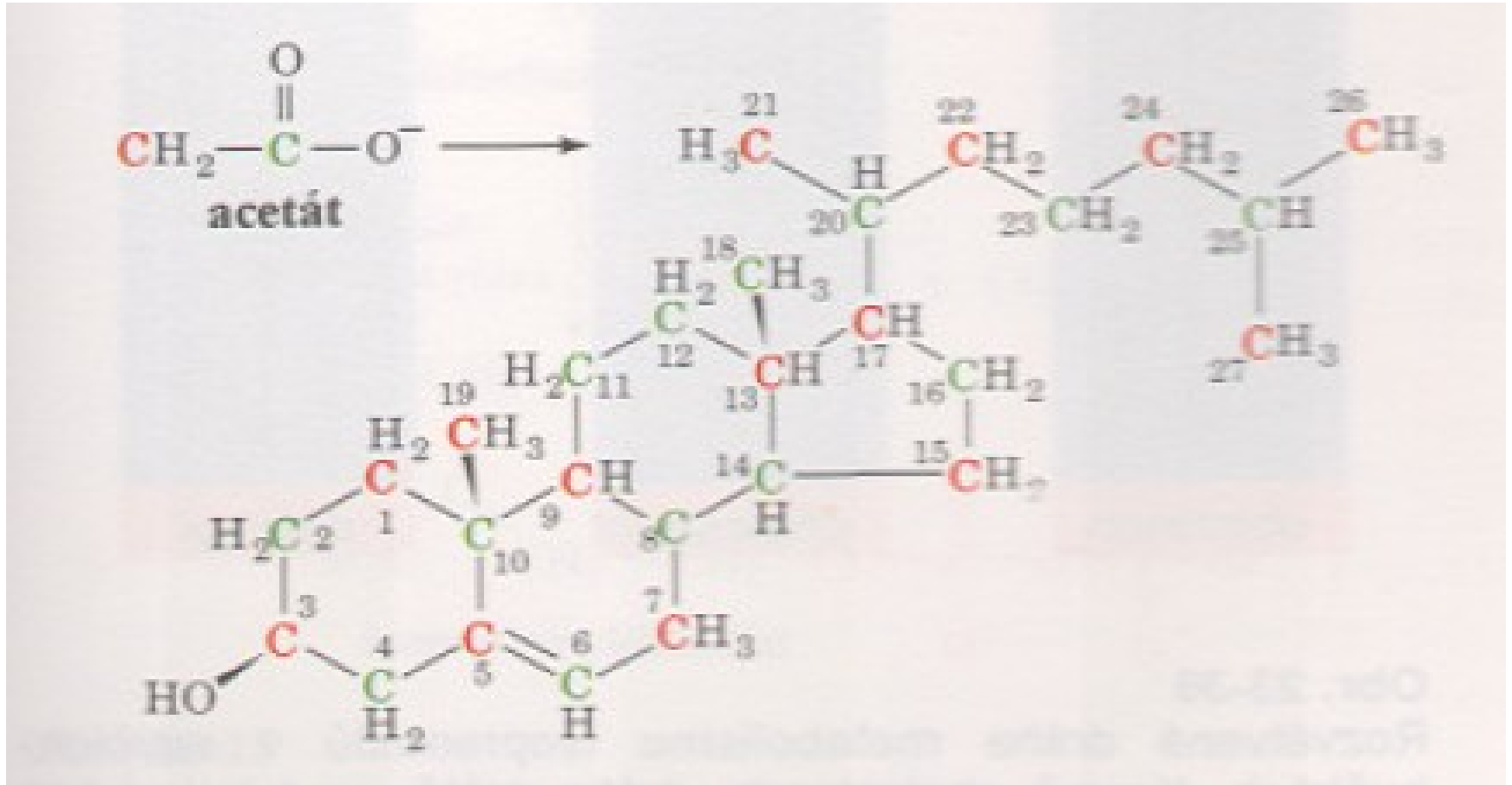
adrenalin



Cholesterol

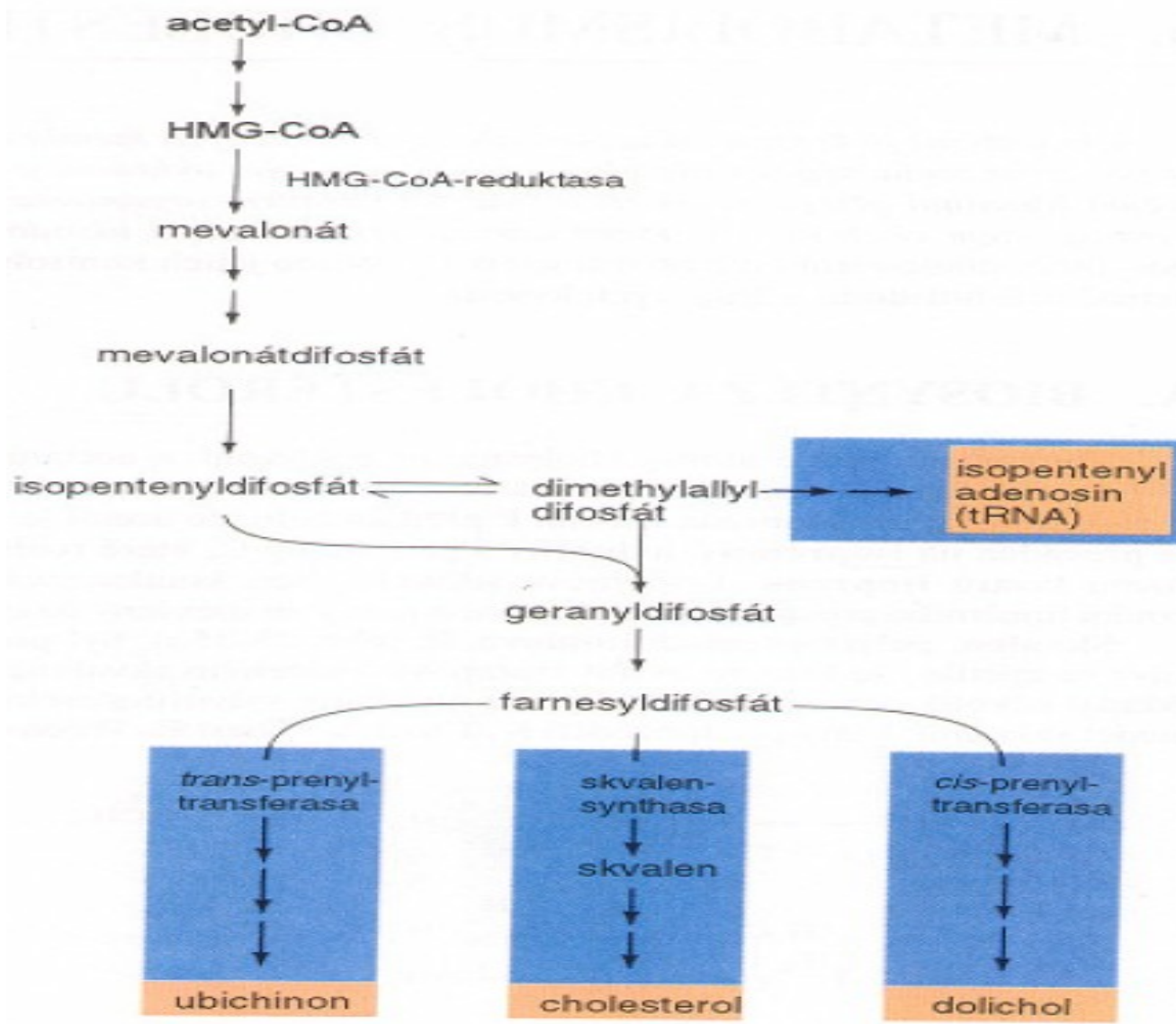


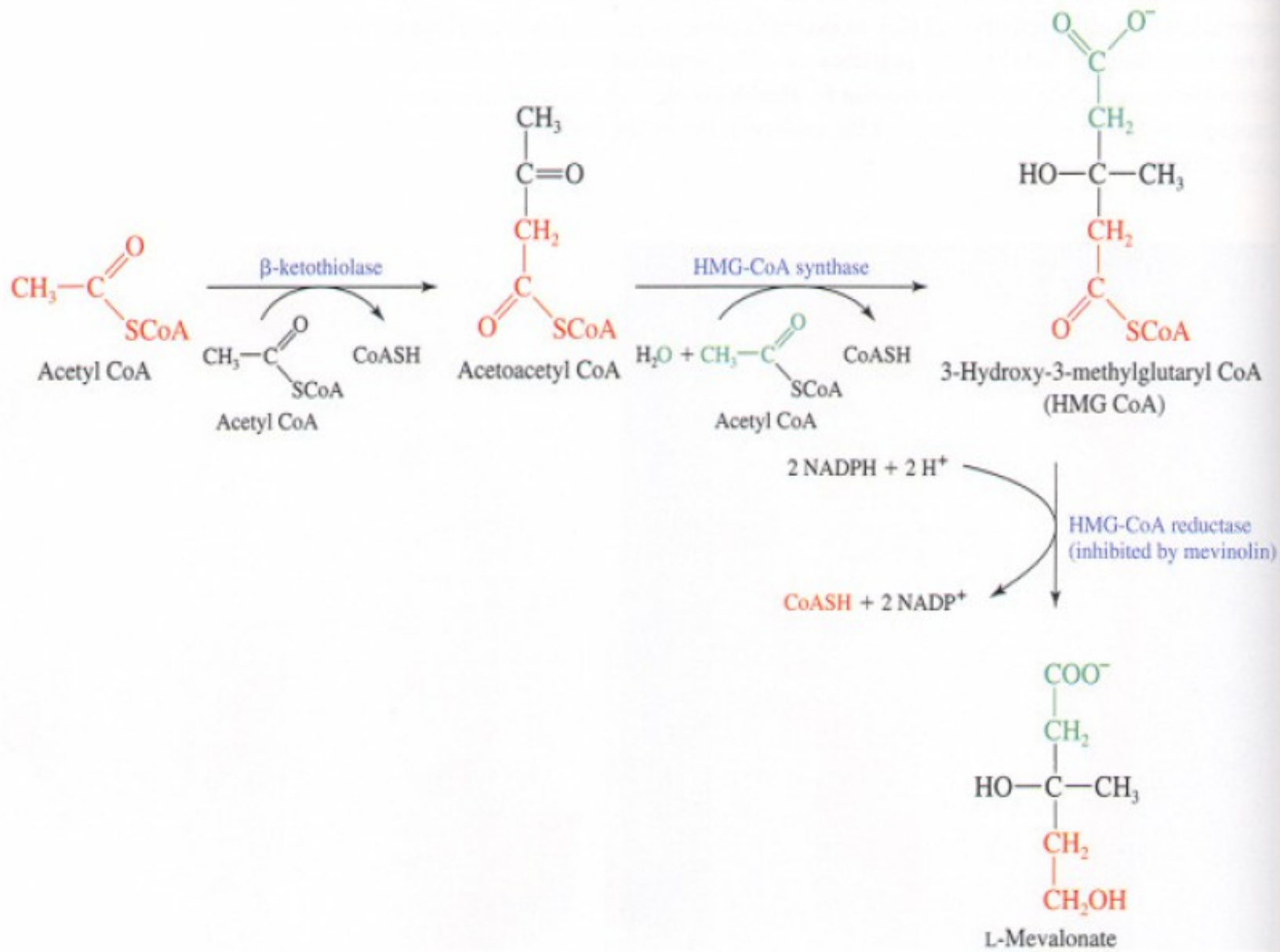
Biosyntéza cholesterolu

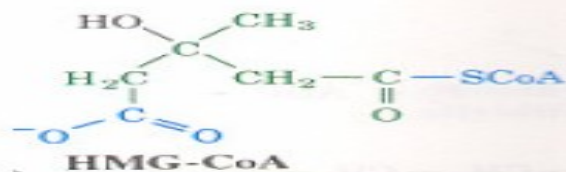


acetát → isoprenoidní intermediát → skvalen
→ produkt cyklizace → cholesterol

Biosyntéza cholesterolu



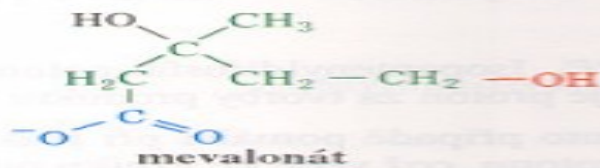




HMG-CoA-reduktasa

1

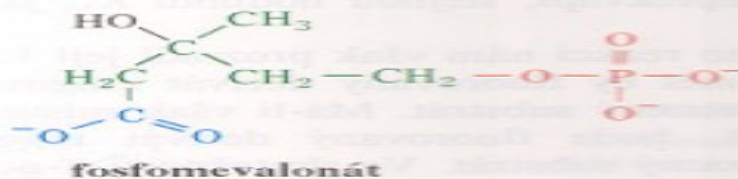
2NADPH → 2NADP⁺ + CoA



mevalonát-5-fosfo-transferasa

2

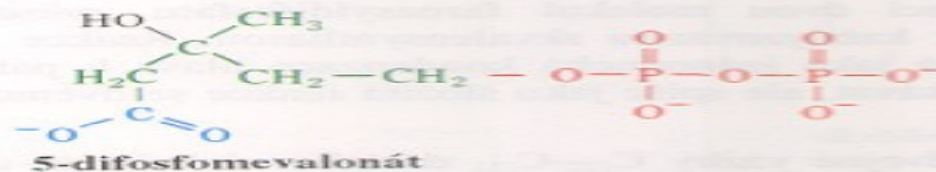
ATP → ADP



fosfomevalonát-kinasa

3

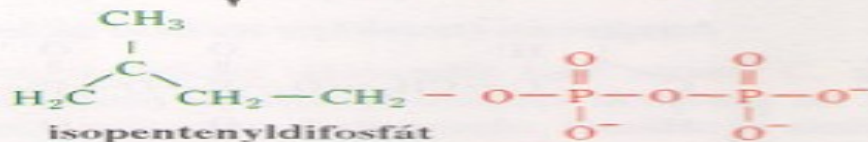
ATP → ADP

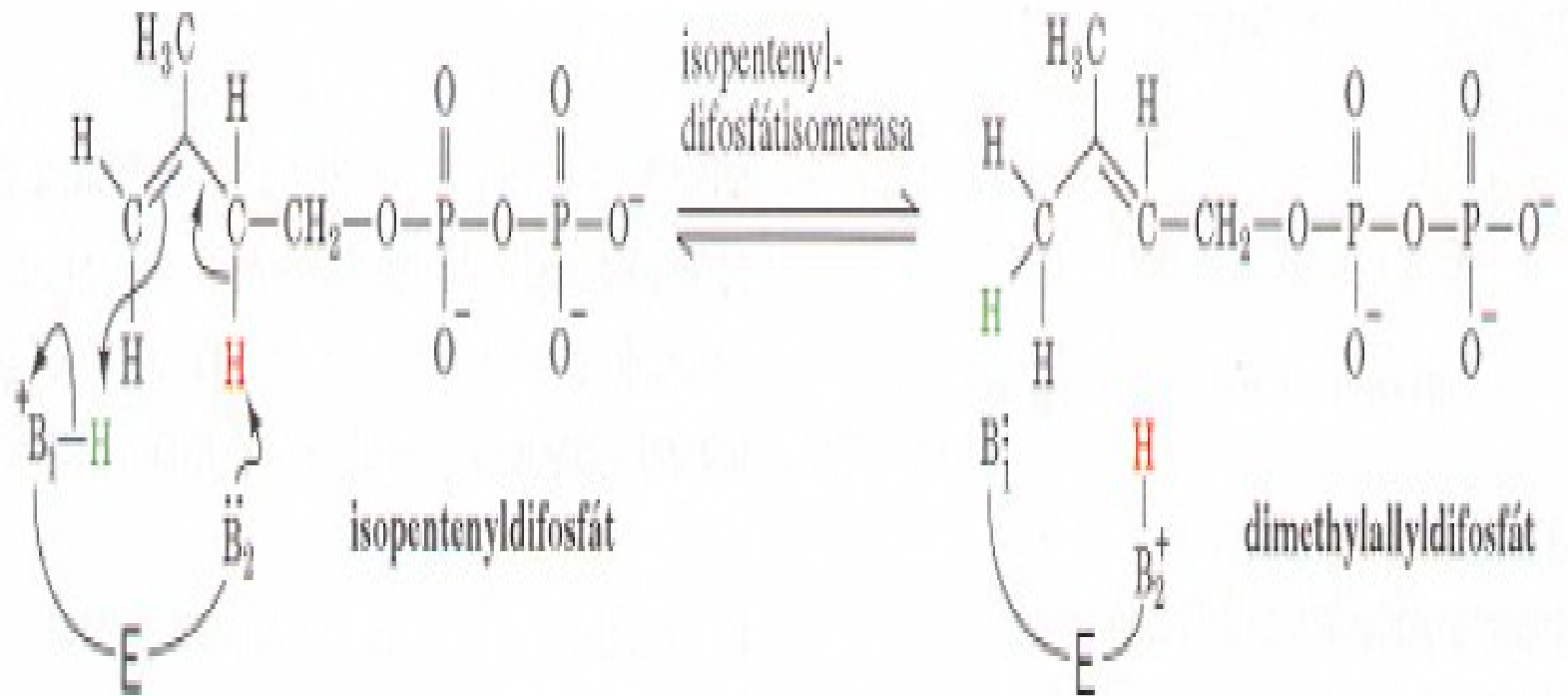


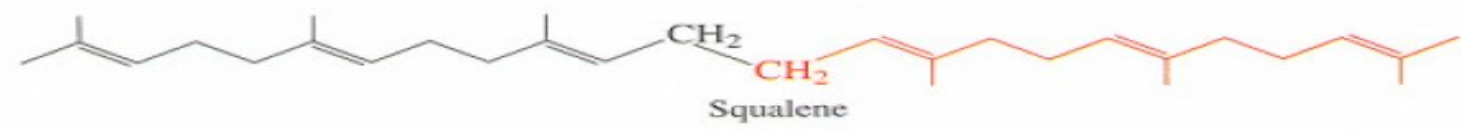
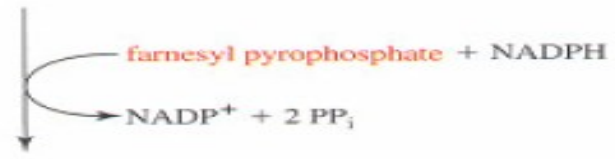
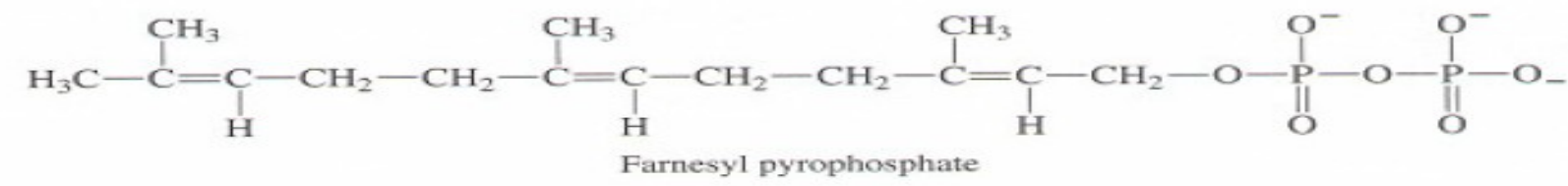
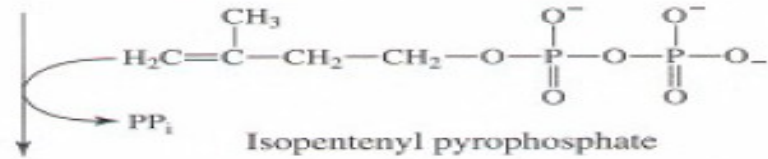
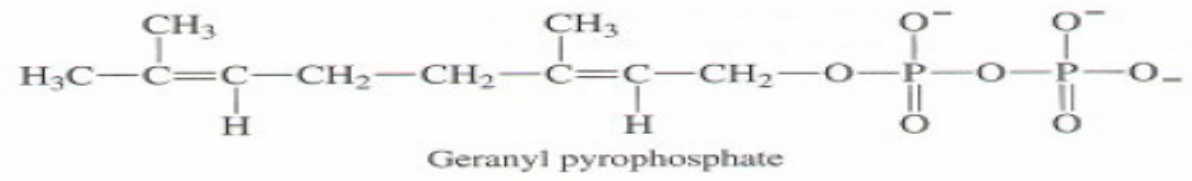
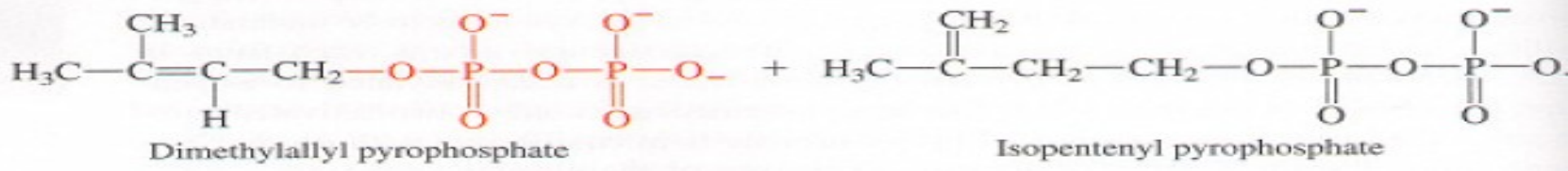
difosfo-mevalonát-dekarboxylasa

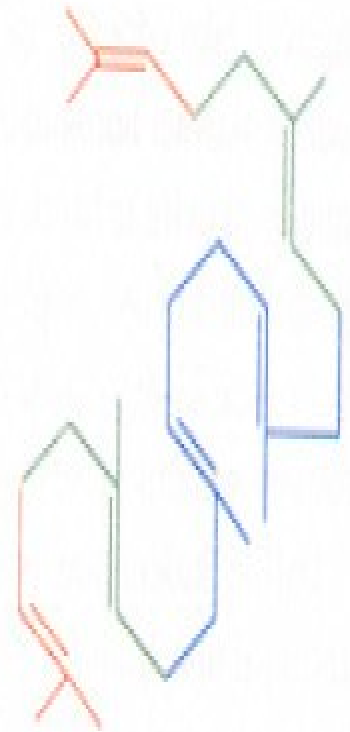
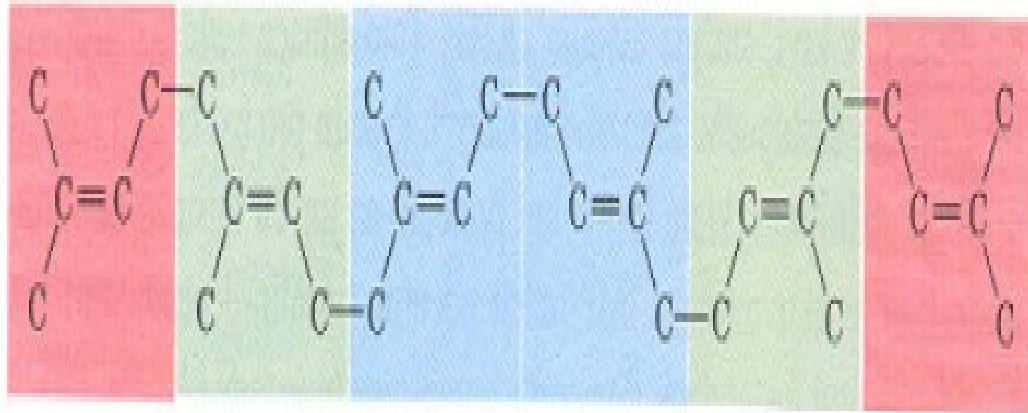
4

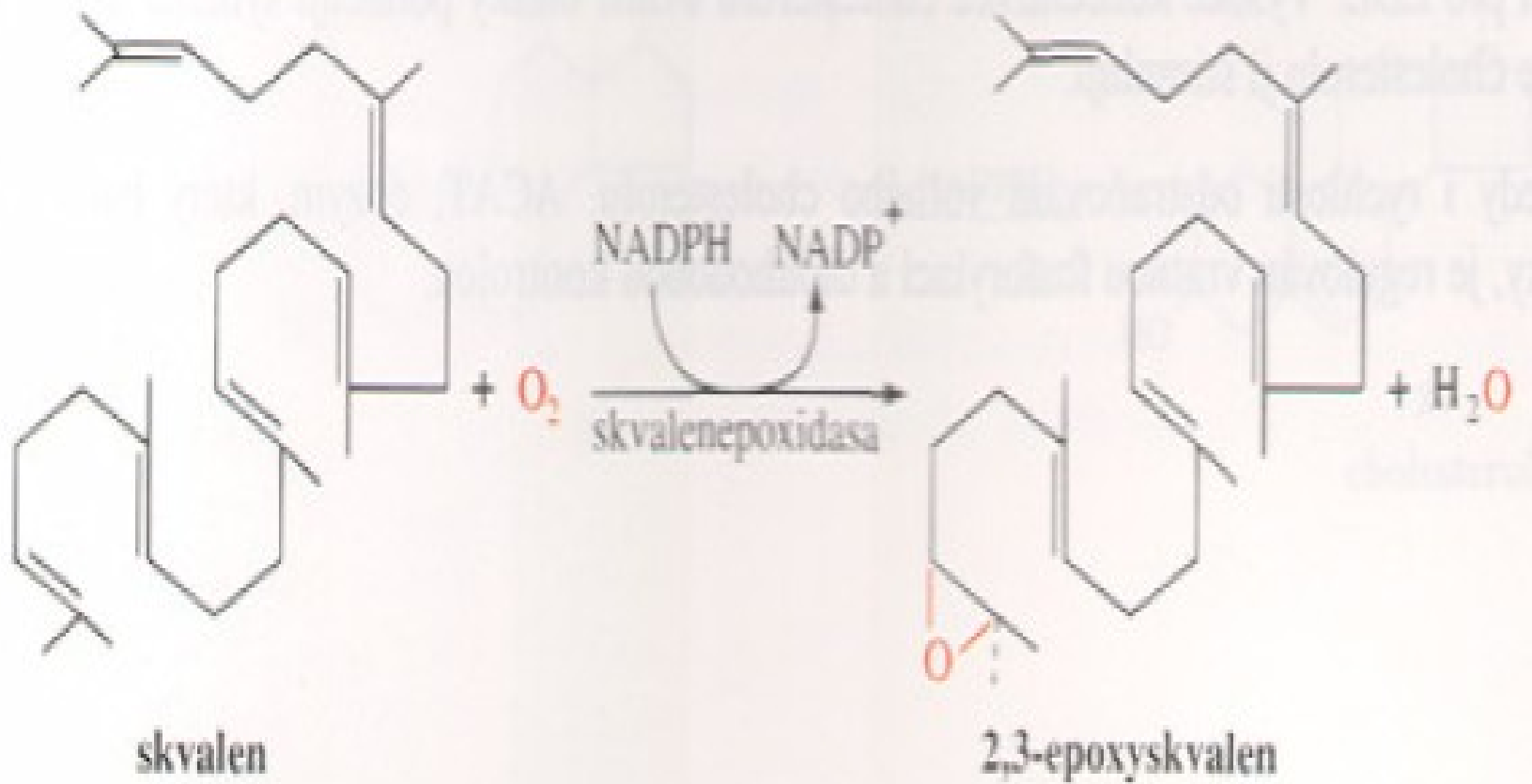
ATP → ADP + P_i + CO₂

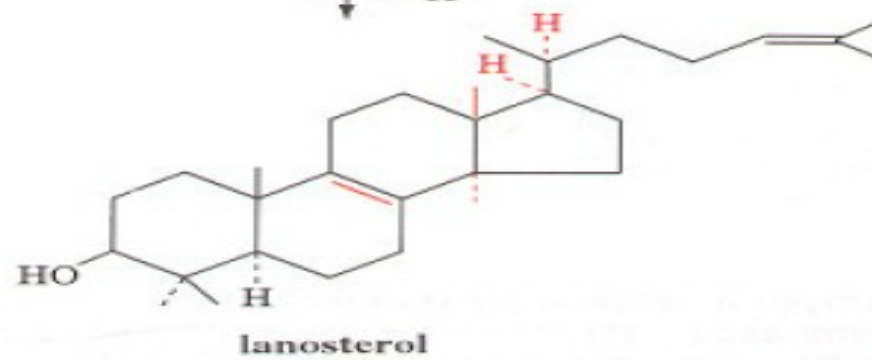
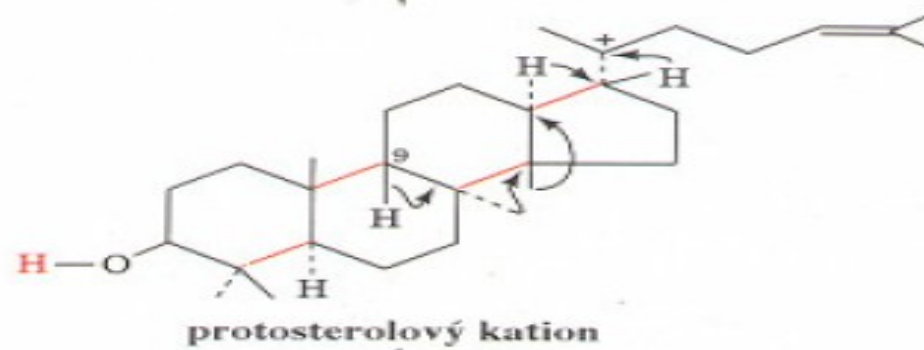
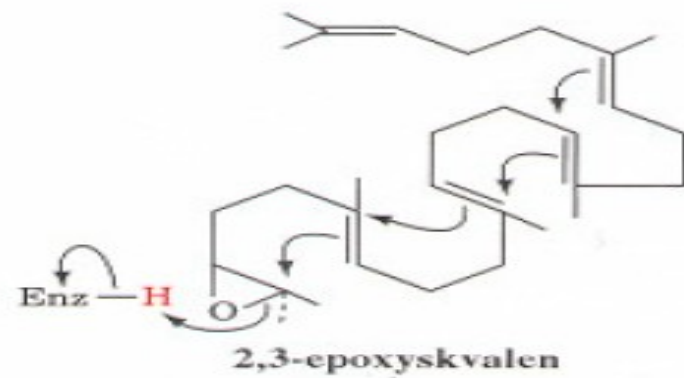


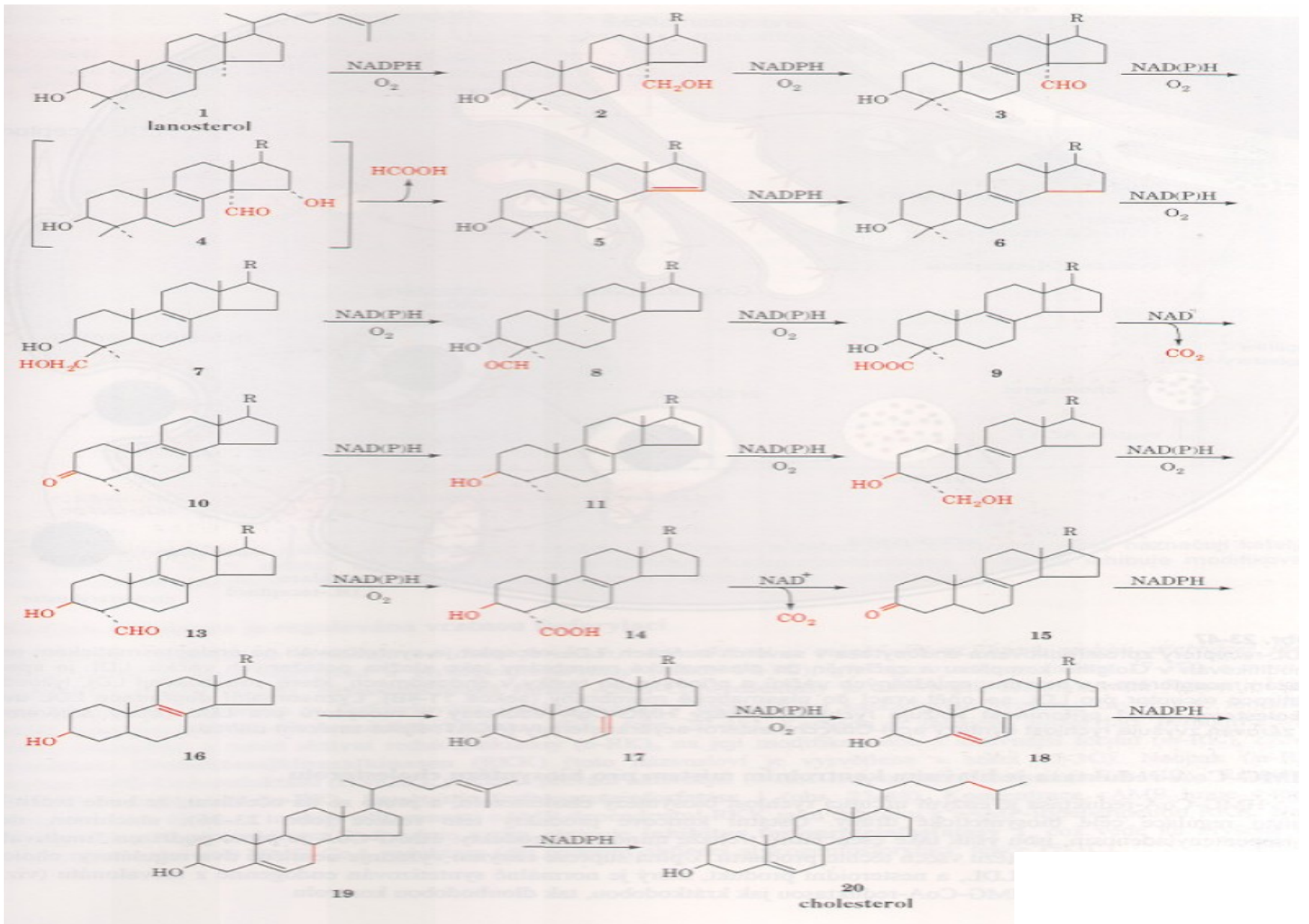












Metabolismus bílkovin

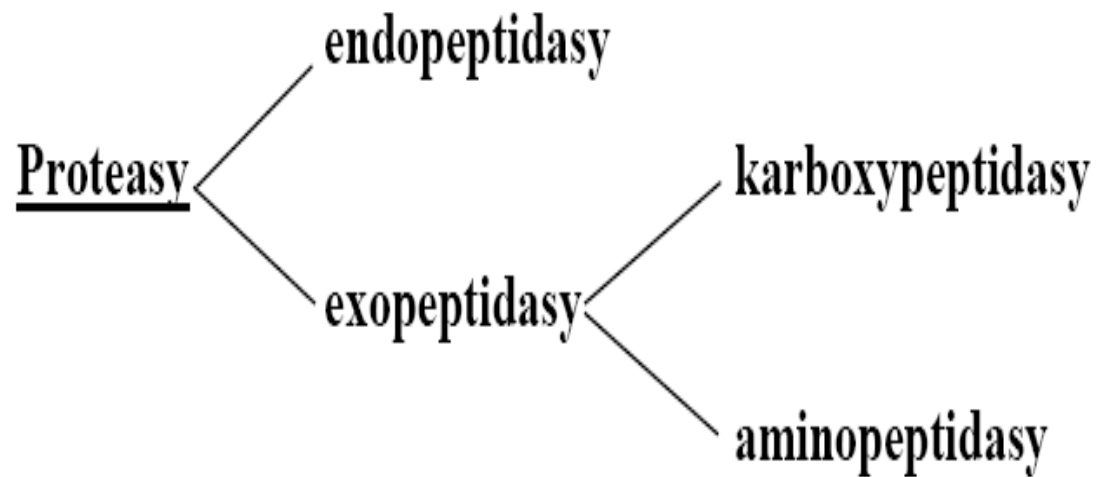
- Tuky, sacharidy – zásobárna energie, mohou se vzájemně zastupovat. Bílkoviny – tvorba tělních bílkovin, jsou zdrojem N pro heterotrofy
- V organismu neexistuje skladiště bílkovin
- U sacharidů (glykolýza, pentozový cyklus) a lipidů (β oxidace) jednotný metabolismus, AMK individuální metabolismus

Metabolismus bílkovin

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

METABOLISMUS BÍLKOVIN

Proteolýza



Proteasy - serinové

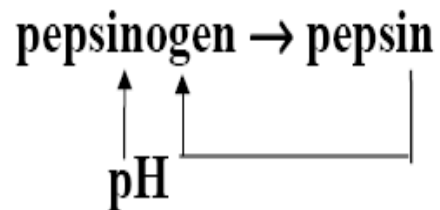
cysteinové

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

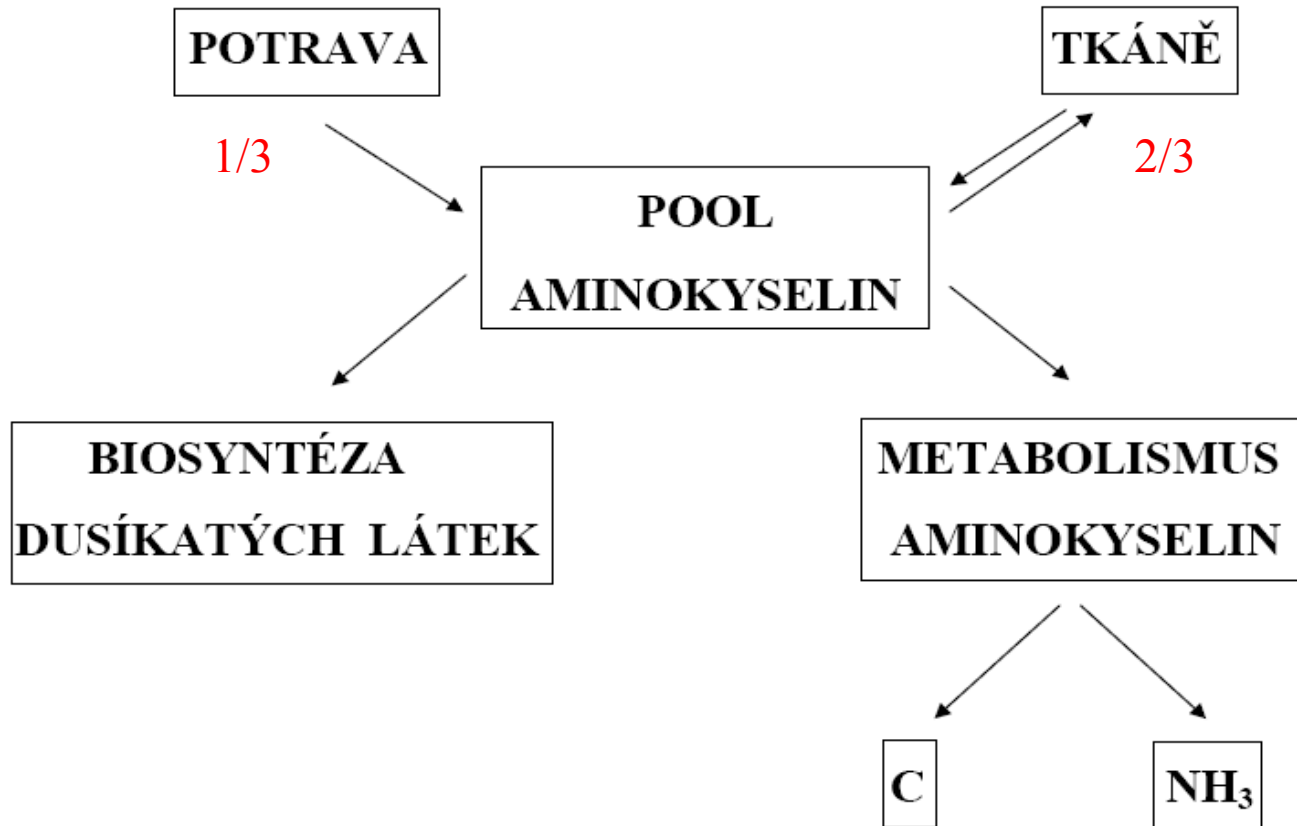
- aminopeptidasy
- dipeptidasy

4. Buněčné proteasy

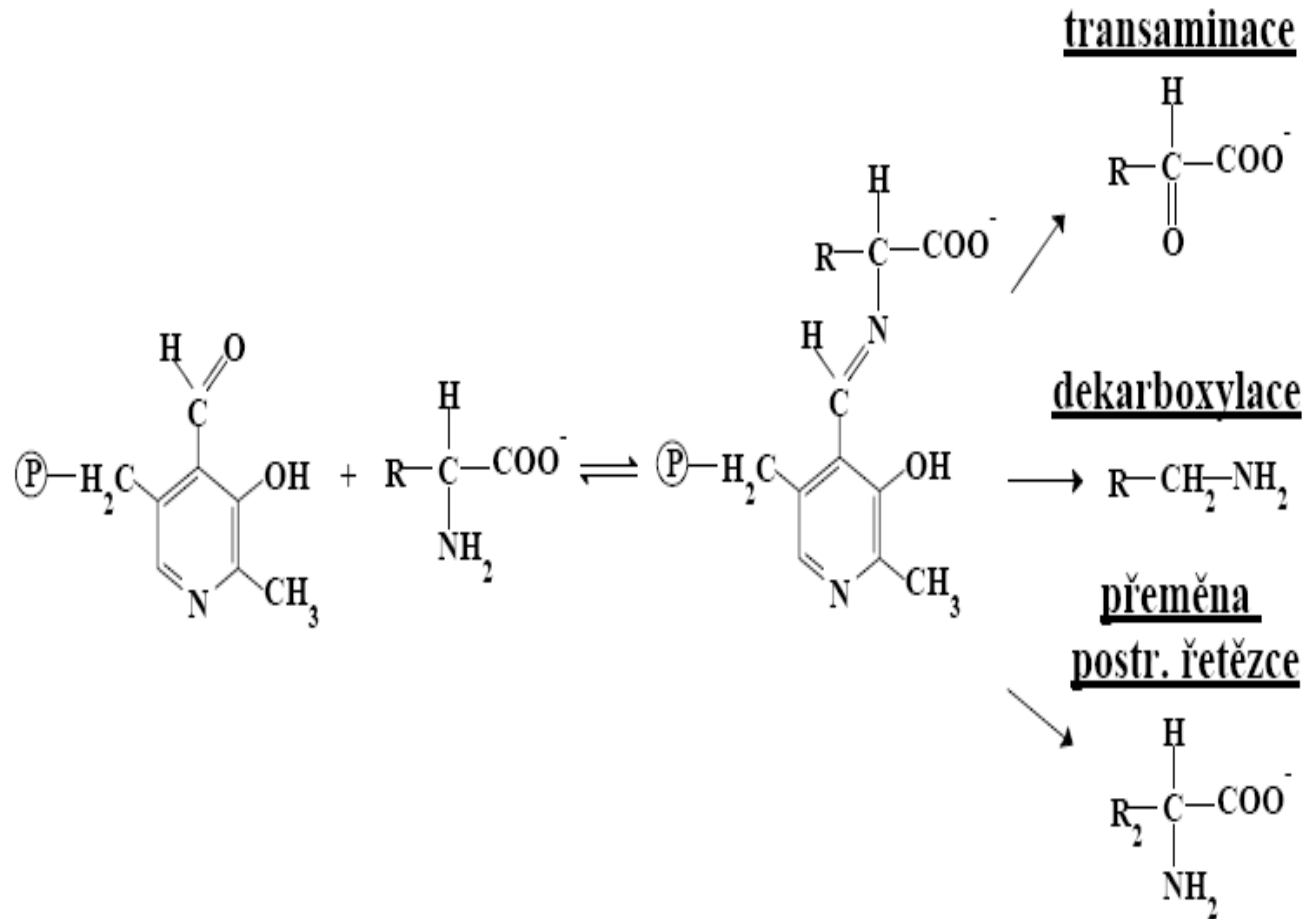
- živočichové - kathepsiny B, D, L, H, M, S a T
- rostliny - papain
- bakterie - subtilisin, pronasa

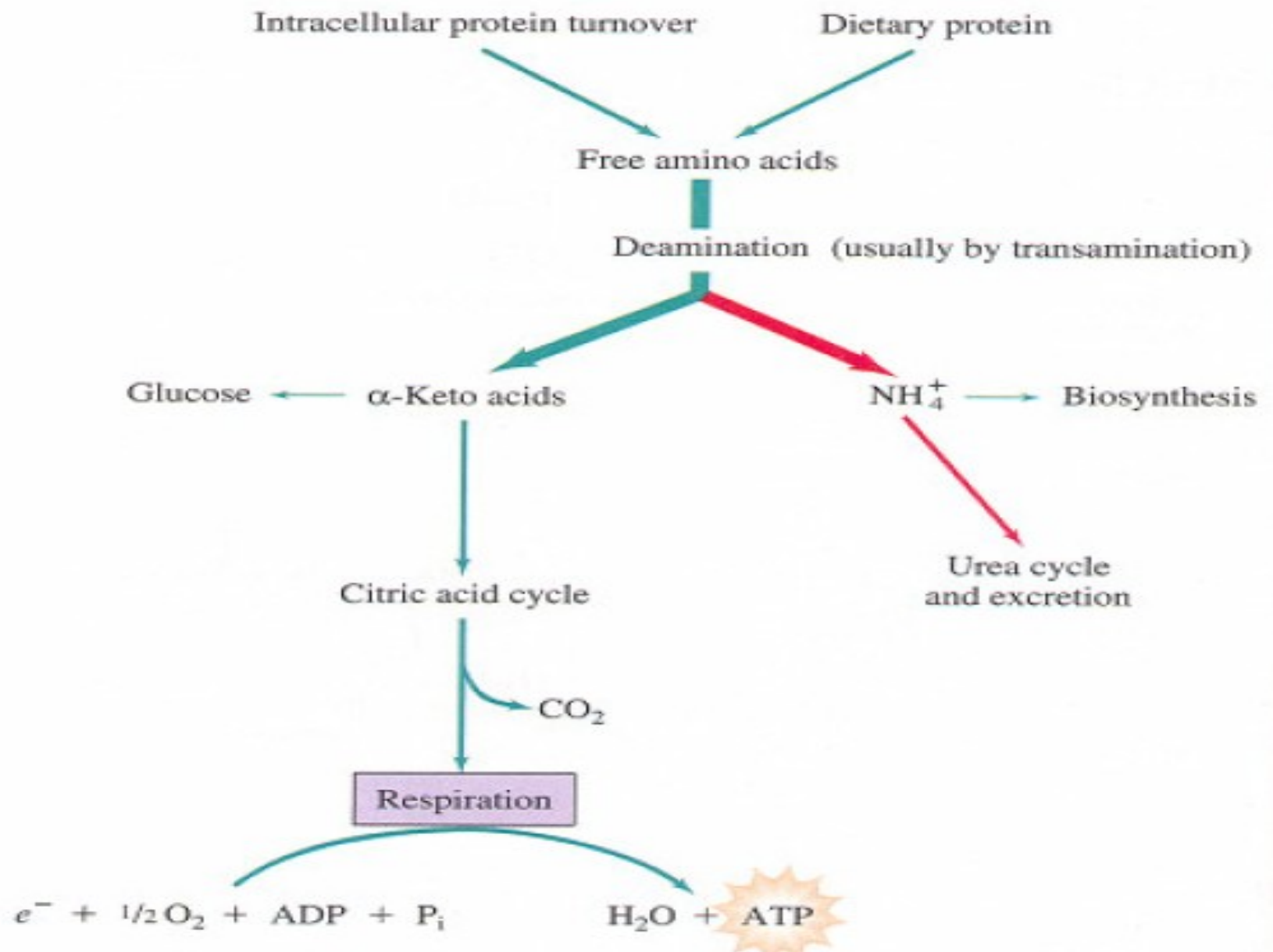
- Proteasy s jinou funkcí
- enterokinasa - aktivace zymogenů
- trombin - srážení krve

Hotovost - pool aminokyselin

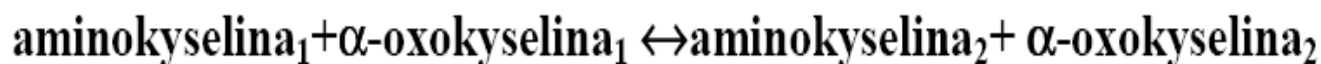


METABOLISMUS AMINOKYSELIN



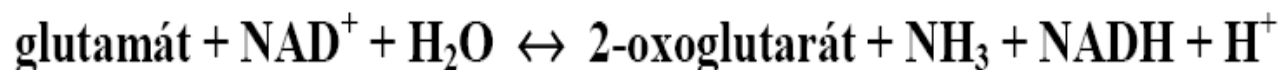


Transaminace

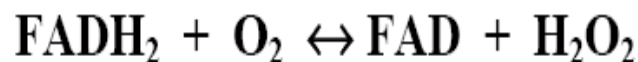
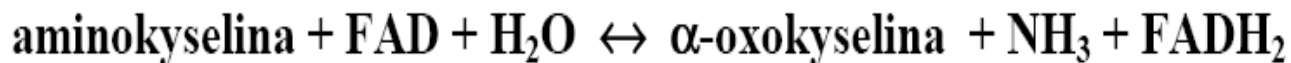


Oxidační deaminace

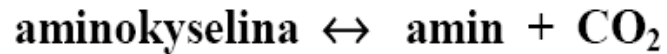
savci



vejcorodí

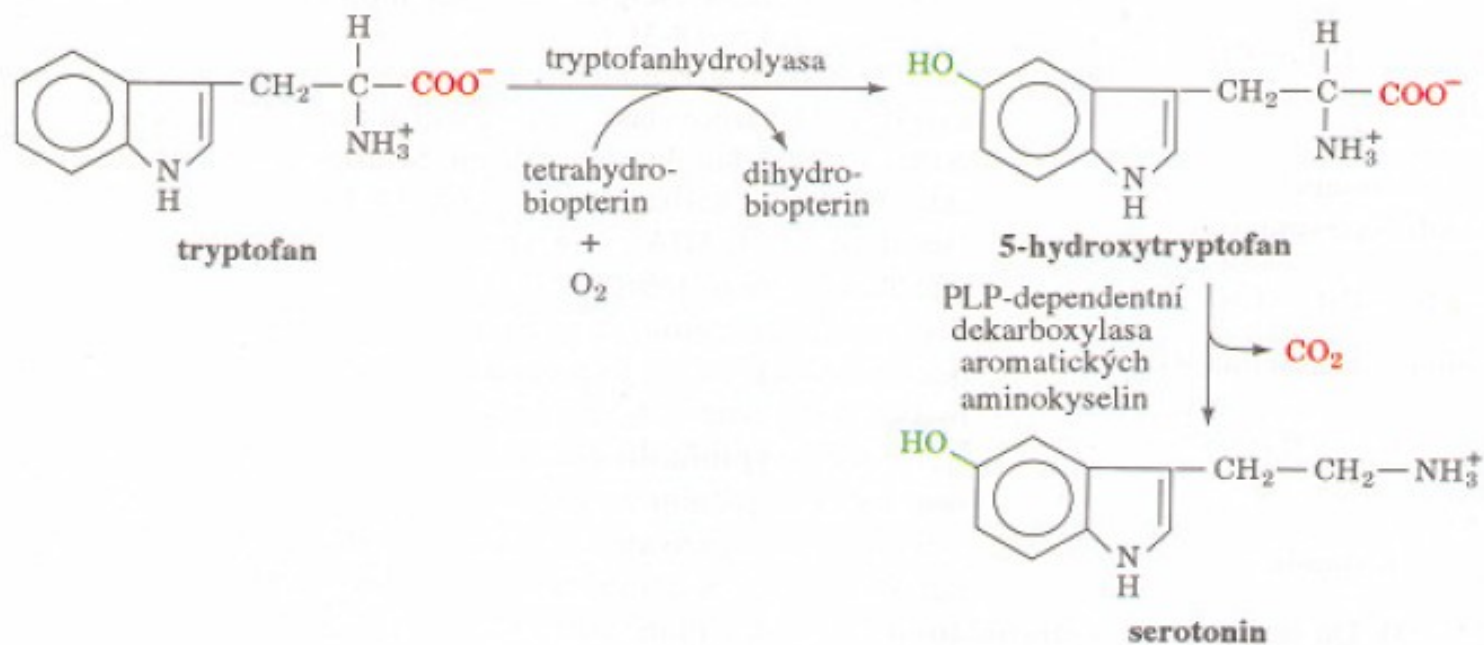
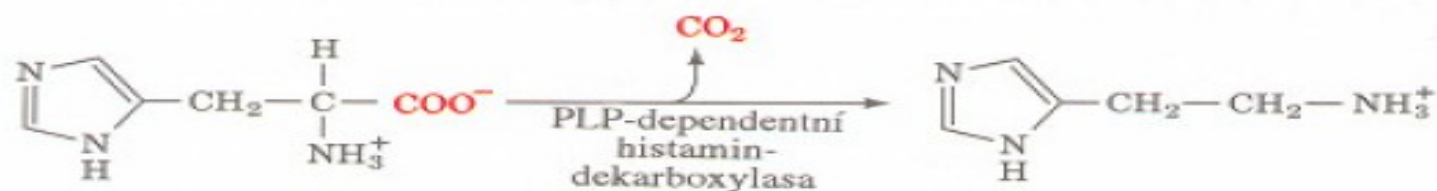
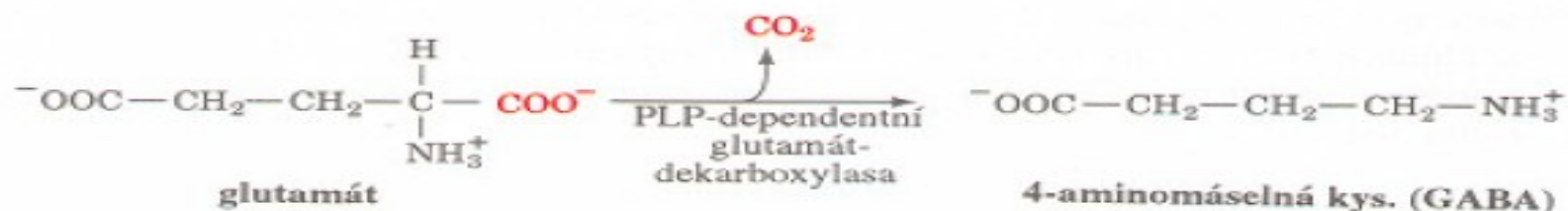


Dekarboxylace



Biogenní aminy

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



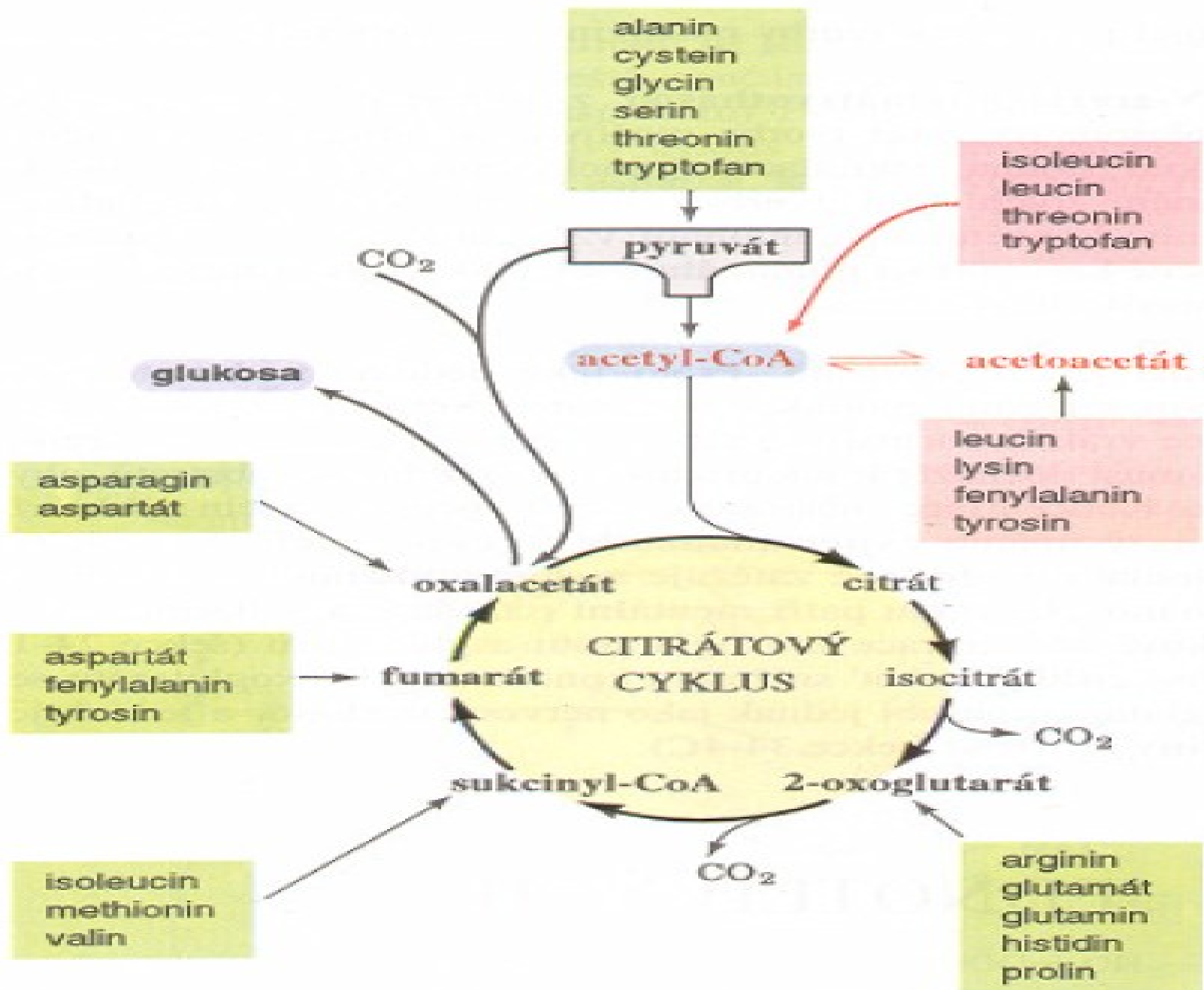
Degradace uhíkových koster aminokyselin

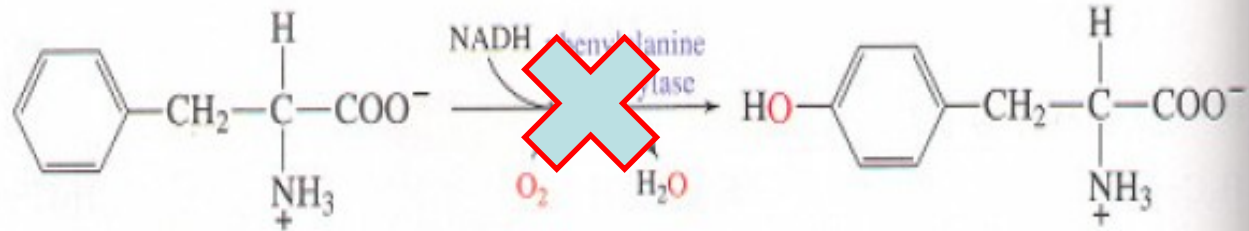
1. Glukogenní aminokyseliny - prekurzory sacharidů

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



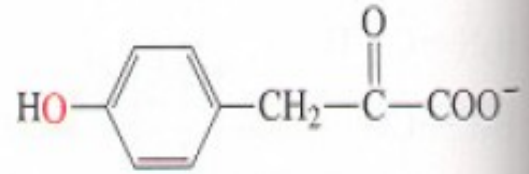


Phenylalanine

Fenylketonurie

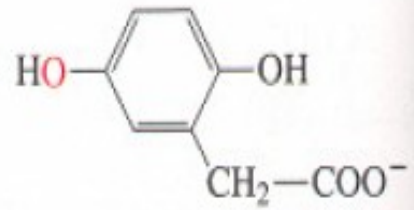
Tyrosine

aminotransferase



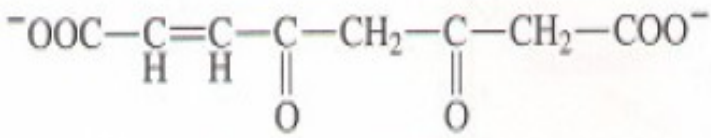
p-Hydroxyphenylpyruvate

↓



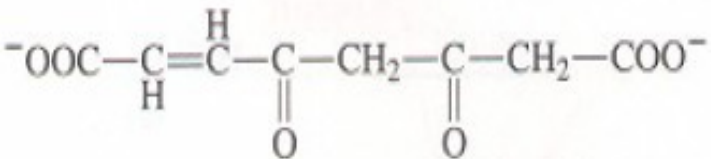
Homogentisate

homogentisate oxidase



4-Maleylacetoacetate

↑

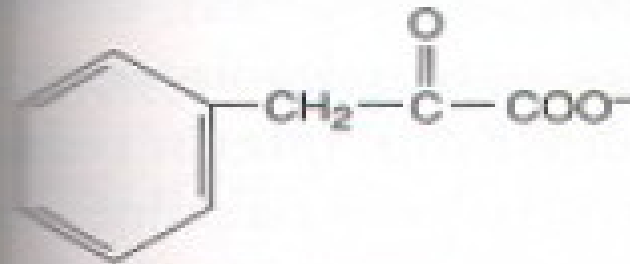


4-Fumarylacetoacetate

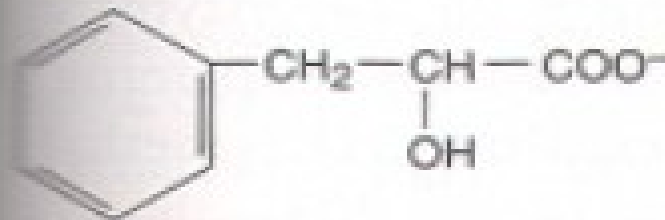
Fumarate + Acetoacetate

←

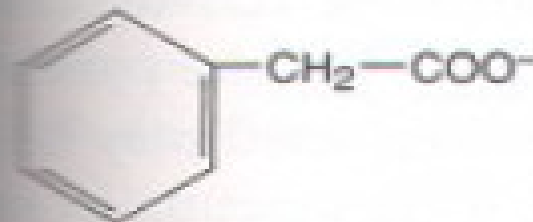
Fenylketonurie



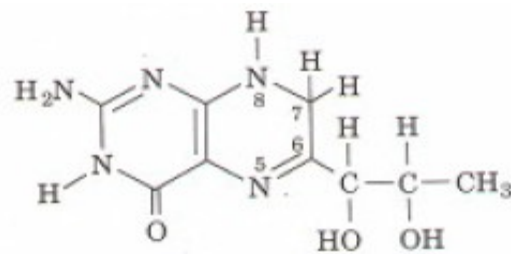
Phenylpyruvate



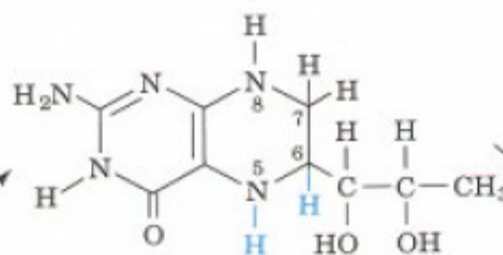
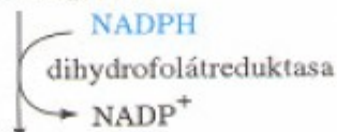
Phenyllactate



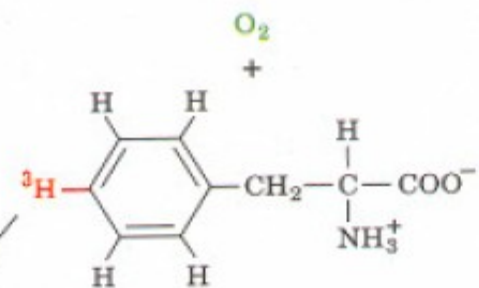
Phenylacetate



7,8-dihydrobiopterin

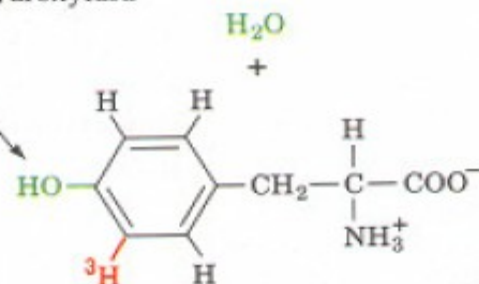


5,6,7,8,-tetrahydrobiopterin

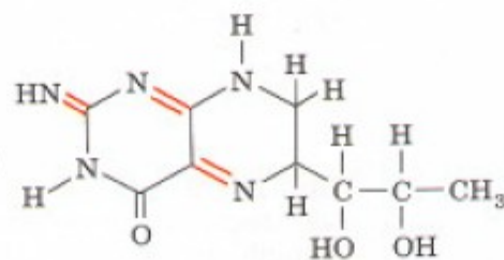


fenylalanin

fenylalanin-hydroxylasa



tyrosin



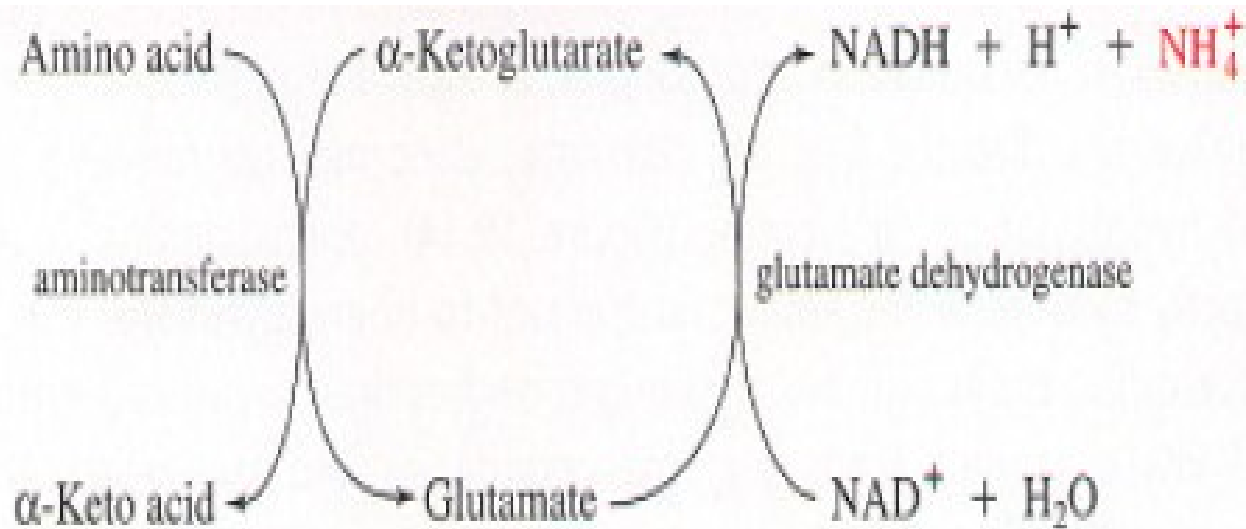
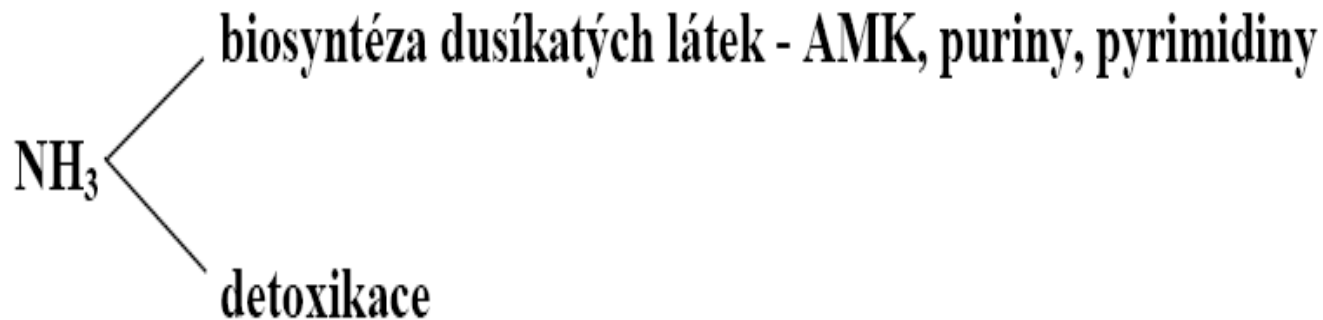
7,8-dihydrobiopterin
(chinoidní forma)

dihydropteridinreduktasa

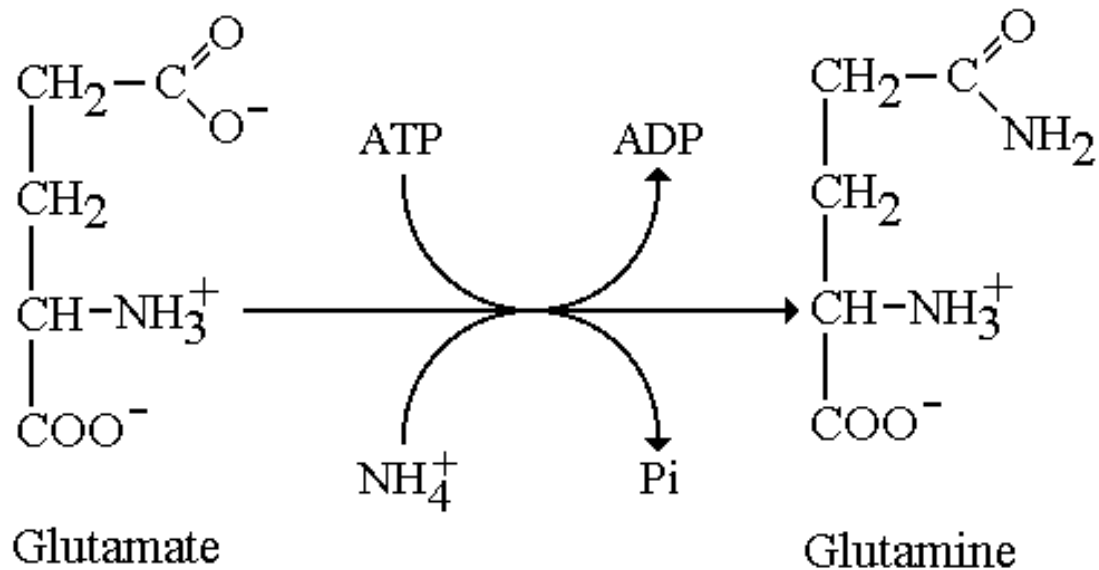
NAD⁺

NADH

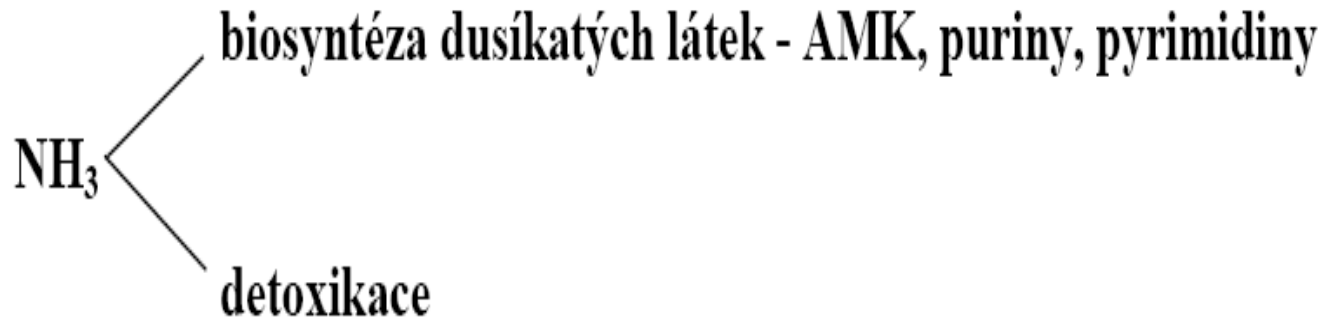
Metabolismus amoniaku



Glutamin synthethasa



Metabolismus amoniaku



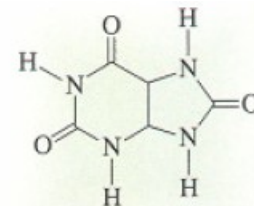
Živočichové - amonotelní - NH_3 - vodní živočichové

- urikotelní - k.močová - vejcorodí

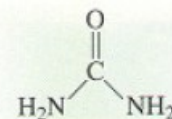
- ureotelní - močovina - placentálové

Rostliny - nevylučují NH_3

NH_4^+
(c) Ammonium ion



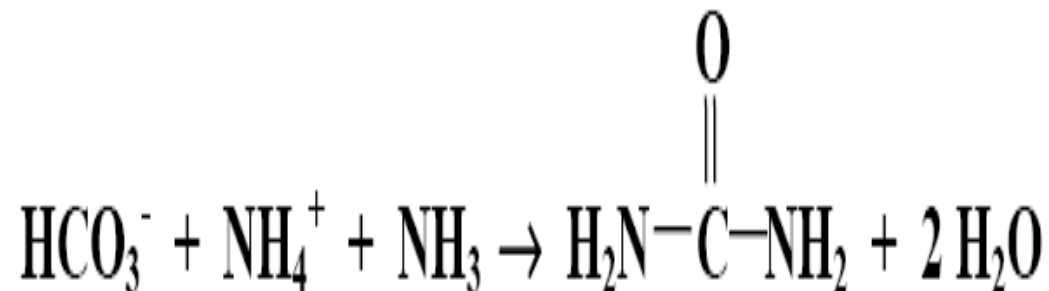
(b) Uric acid

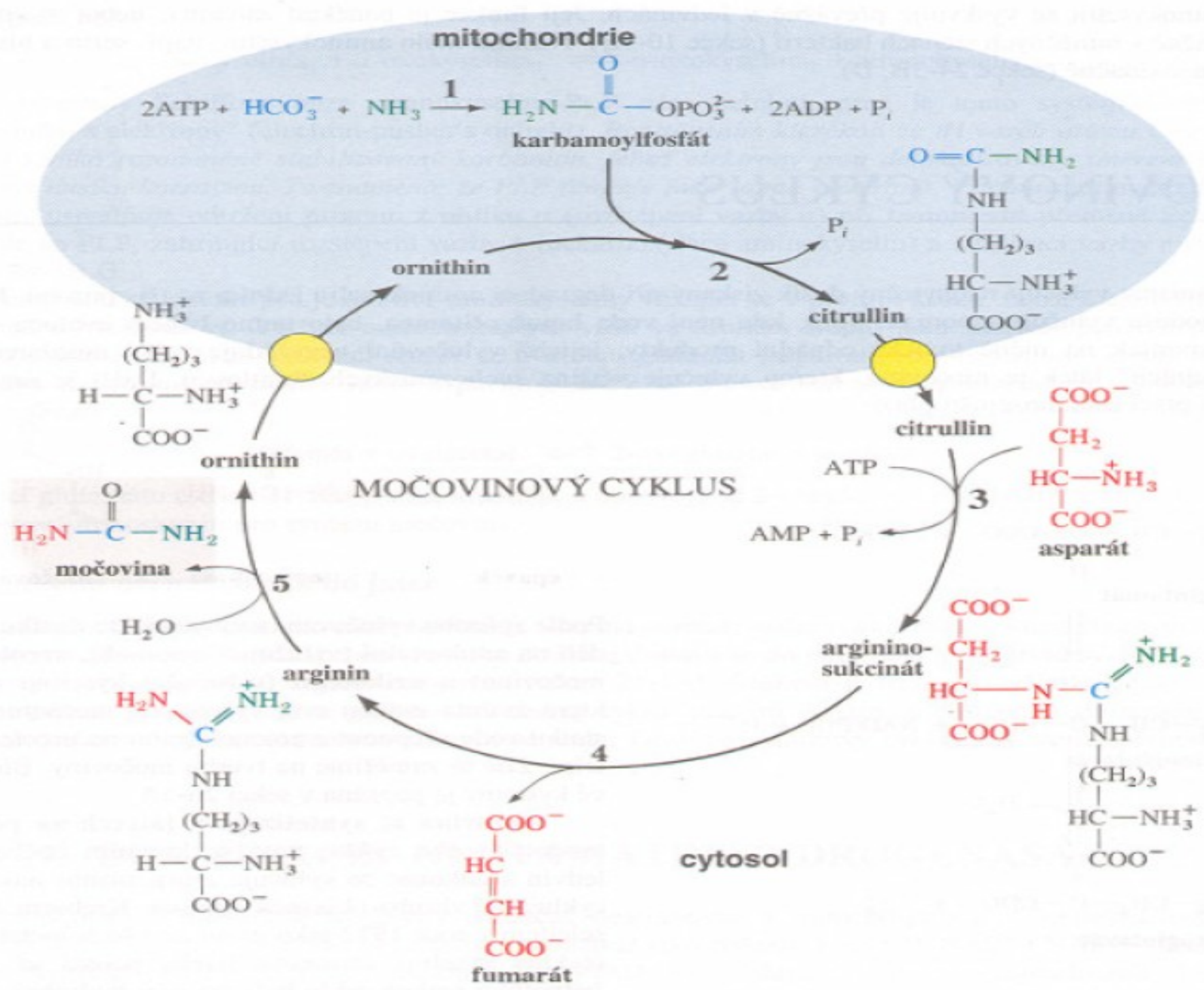


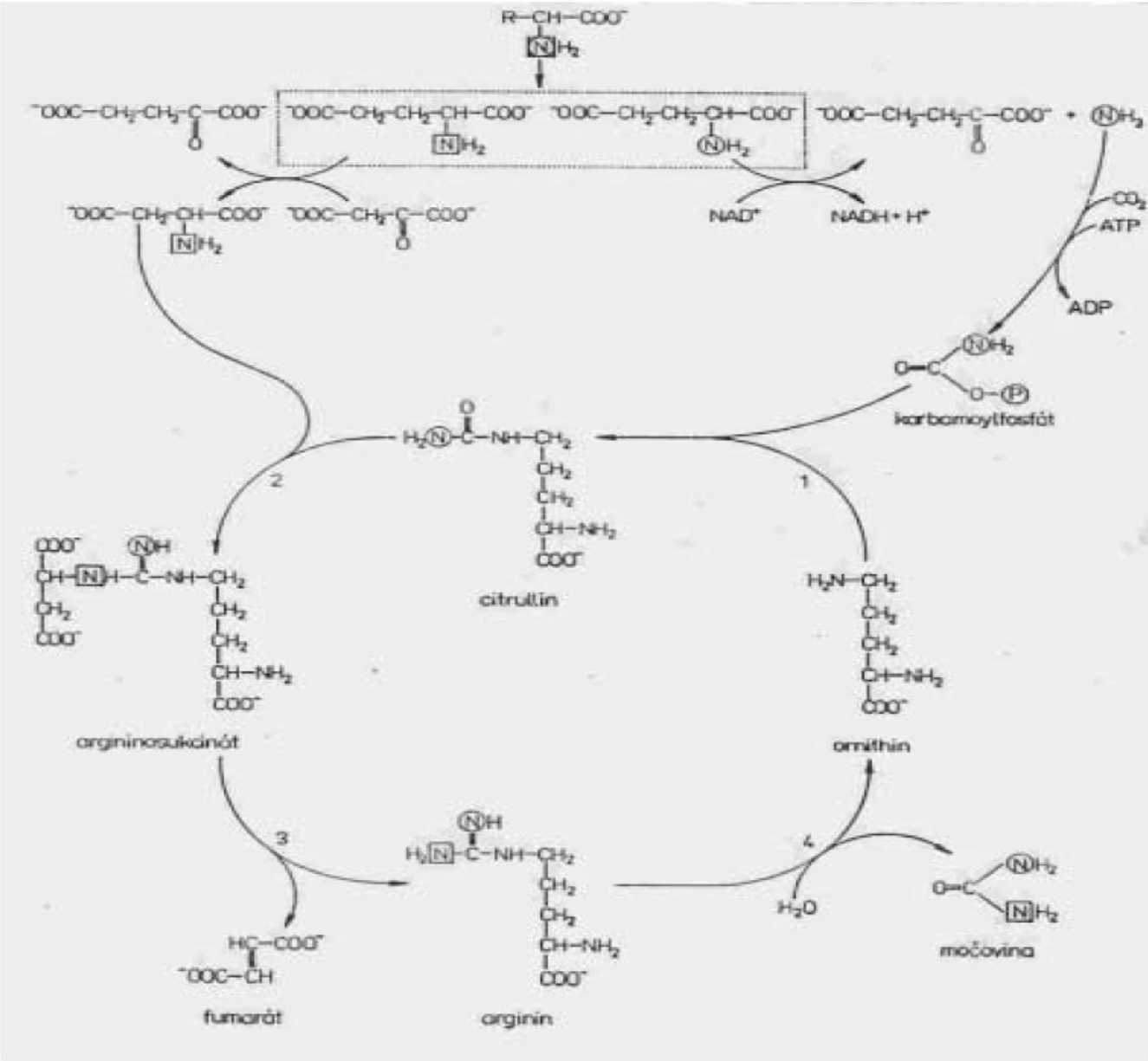
(a) Urea

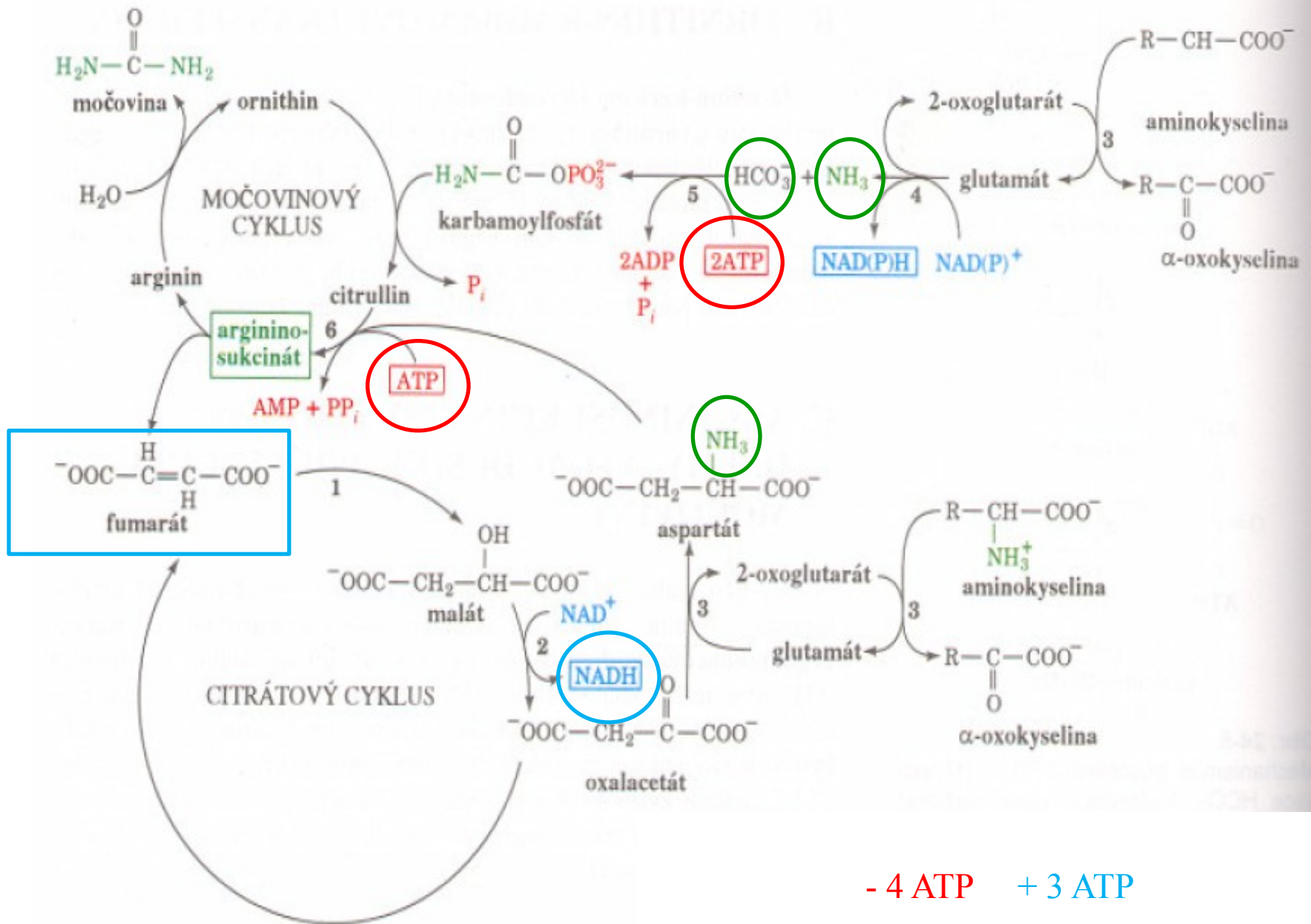
Tvorba močoviny - ornitinový cyklus

H. KREBS, K. HENSELEIT - 1932

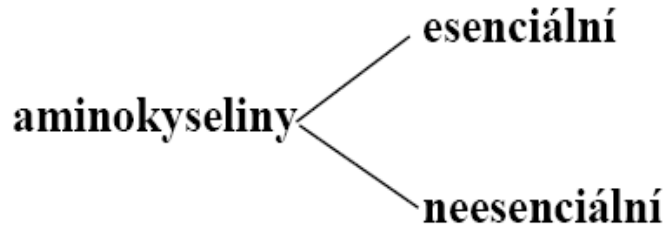






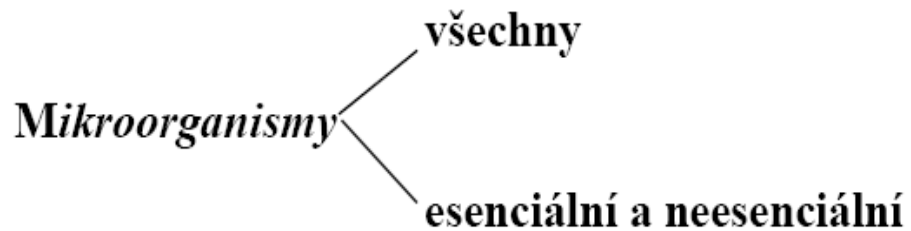


Biosyntéza aminokyselin



Rostliny - všechny

NH_2 , NH_4^+ , NO_3^- , NO_2^- , N_2



NH_2 , NH_4^+ , NO_3^- , NO_2^- , N_2

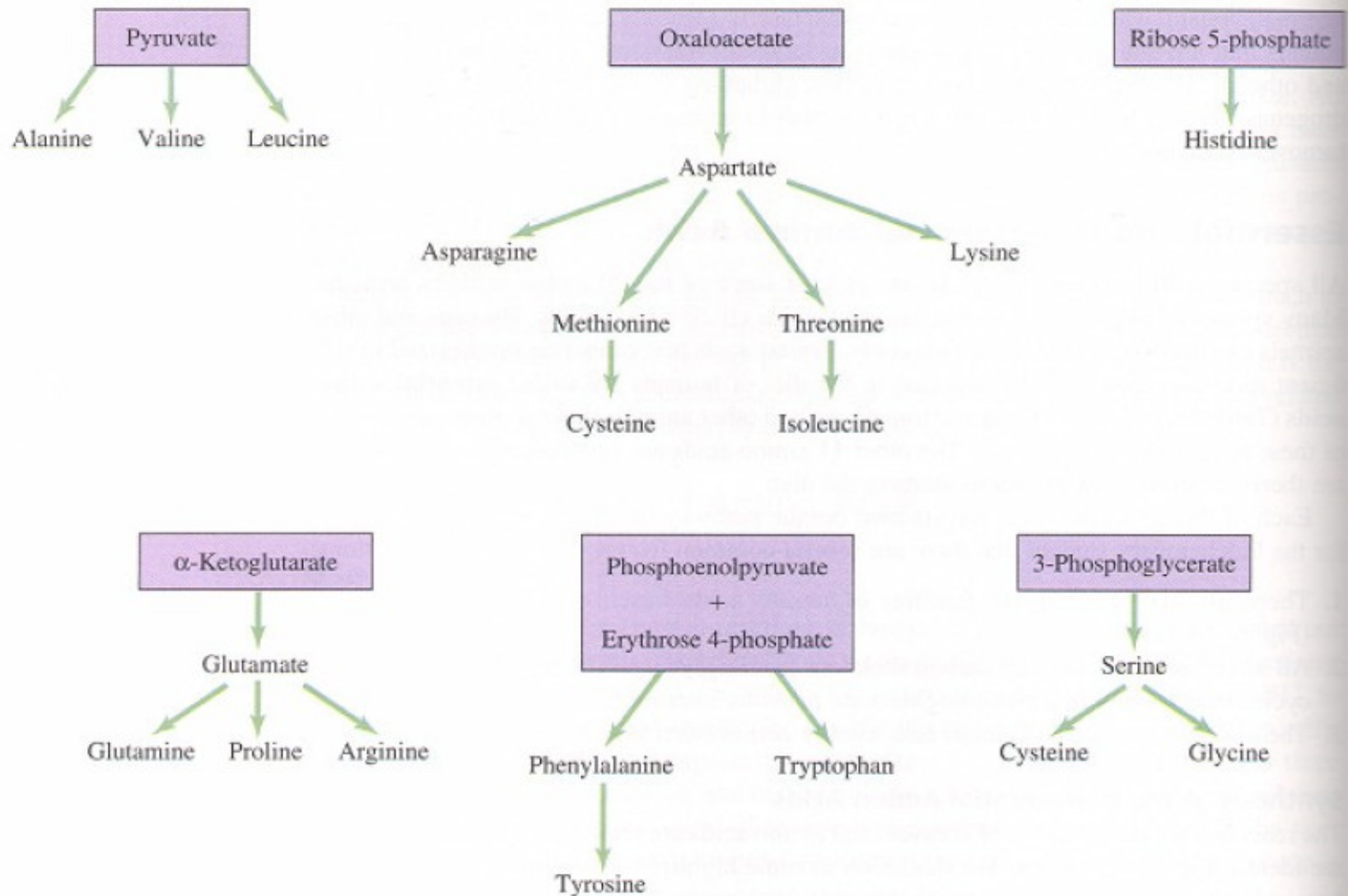
Živočichové - esenciální a neesenciální

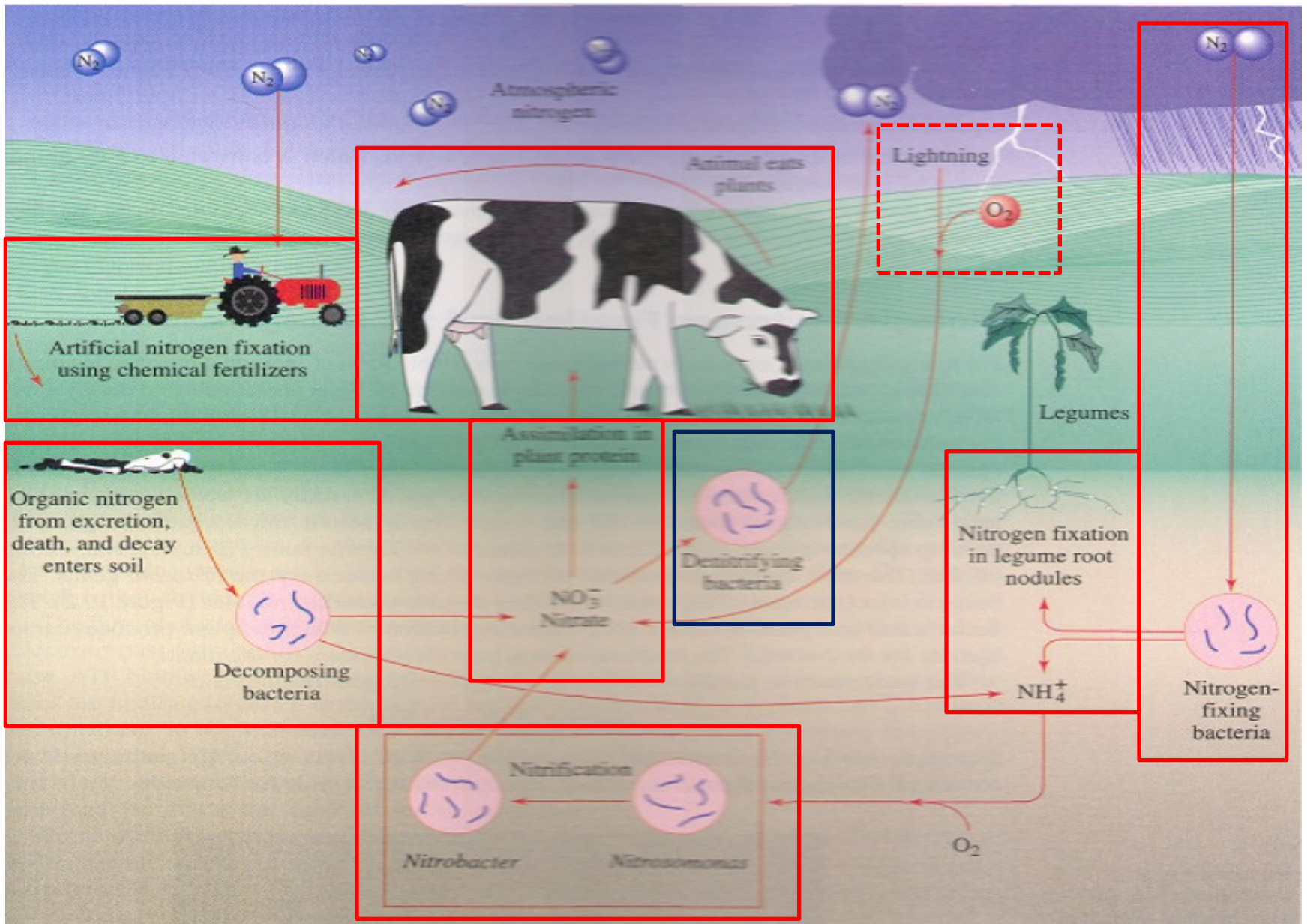
NH_2

Biosyntéza aminokyselin

- **Uhlíkové kostry pocházejí z meziproduktů glykolýzy, pentózového a citrátového cyklu**
- **Biosyntéza vychází ze společných prekurzorů a probíhá přes společné meziprodukty – jen 6 drah**
- **Další AMK se tvoří přestavbou jiných AMK**
- **Neesenciální AMK jsou syntetizovány jednodušší cestou**
- **Biosyntéza esenciálních AMK je komplikovanější**

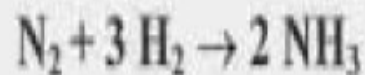
Biosyntéza aminokyselin



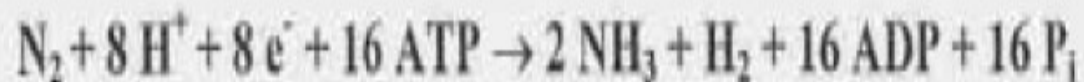


Fixace N₂

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



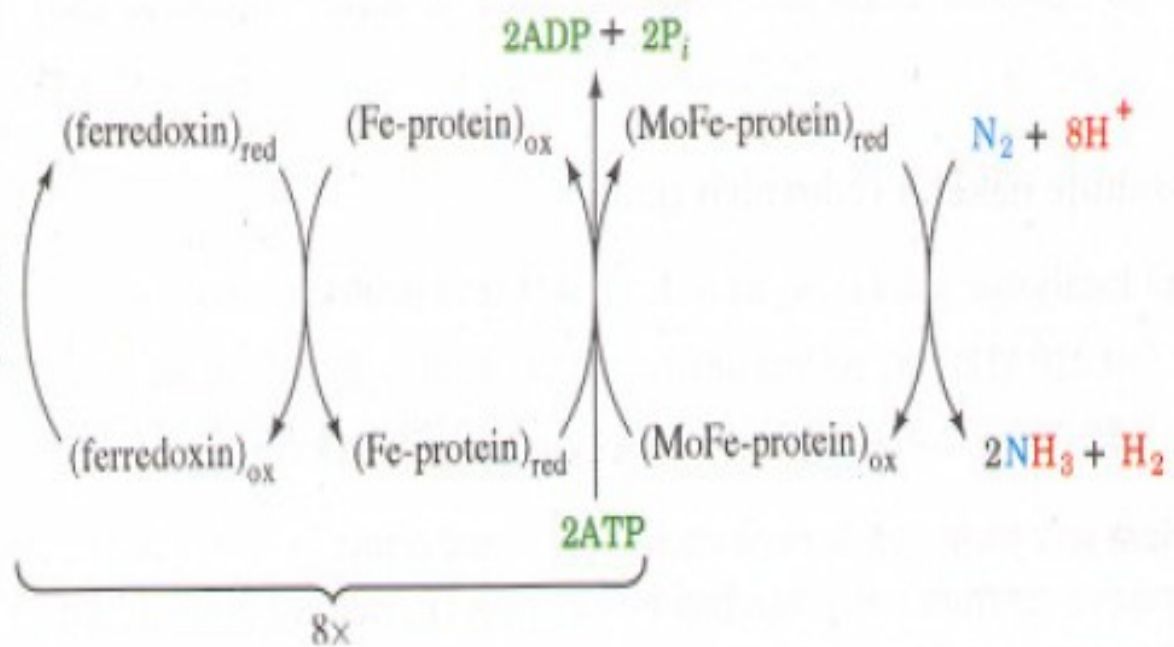
Biosyntéza – sinice, bakterie - *Rhizobium*, *Azotobacter*

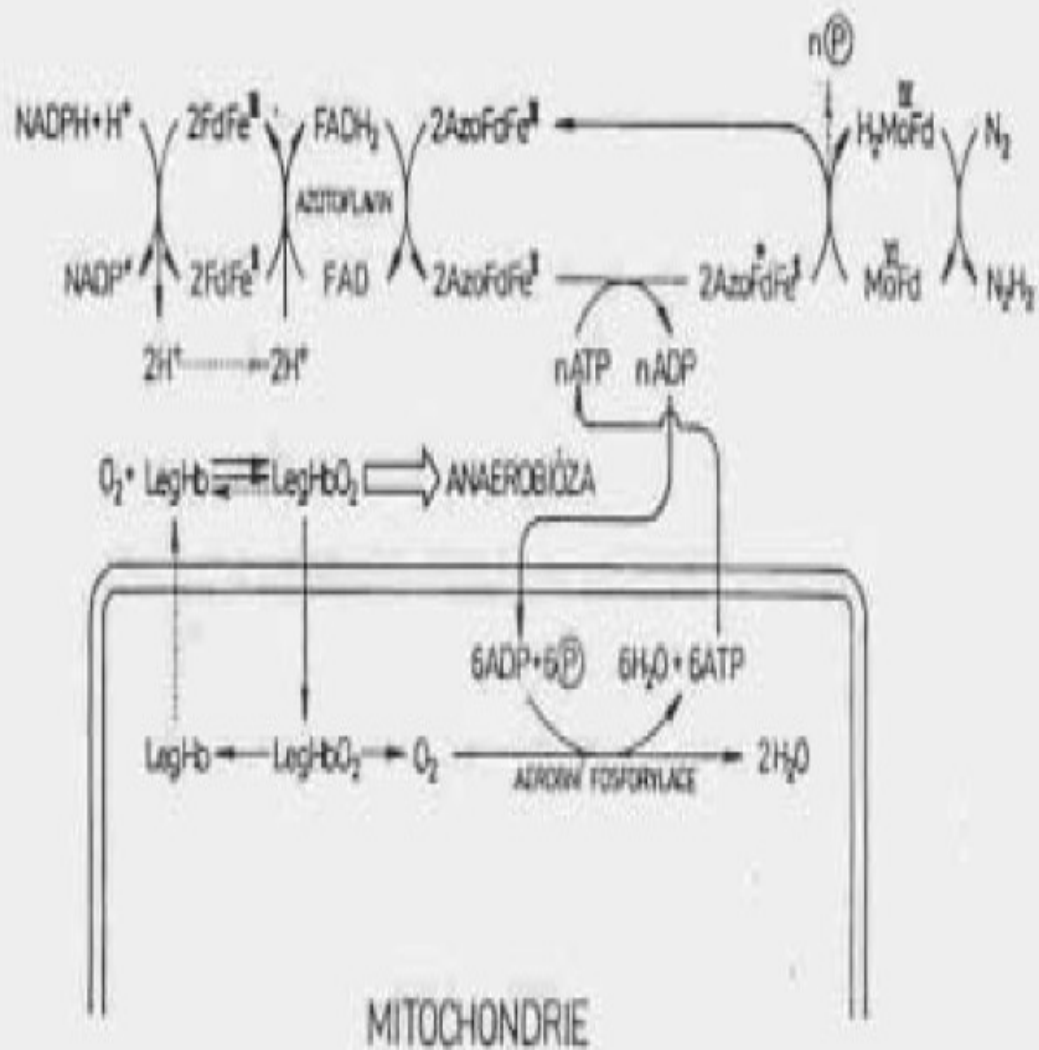


Nitrogenasa : 1. protein Fe
2. protein MoFe

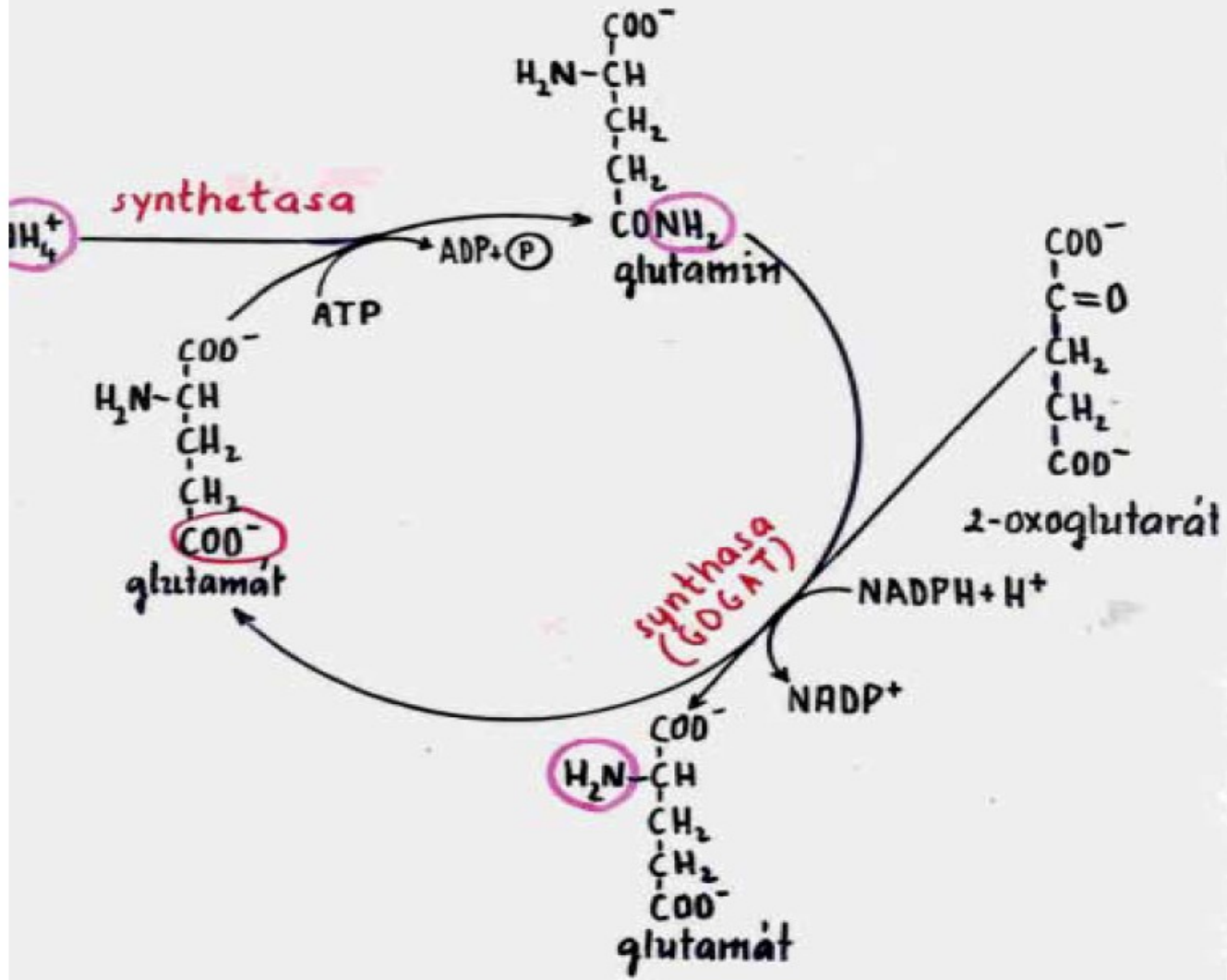


fotosyntéza
nebo oxidační
transport elektronů





INKORPORACE NH₃ U PROKARYOT



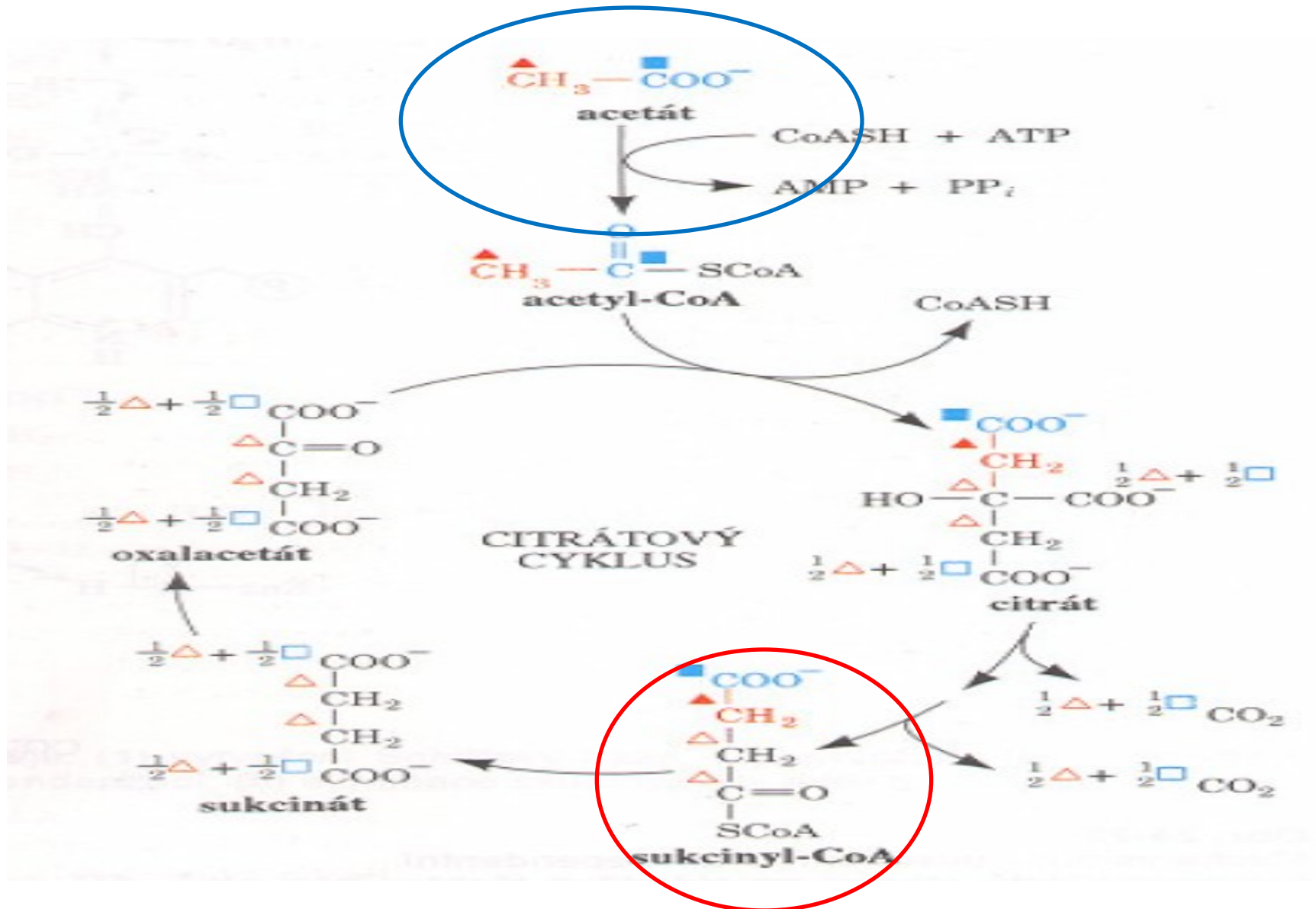
Eukaryota - GluDH

AMK jako prekurzory

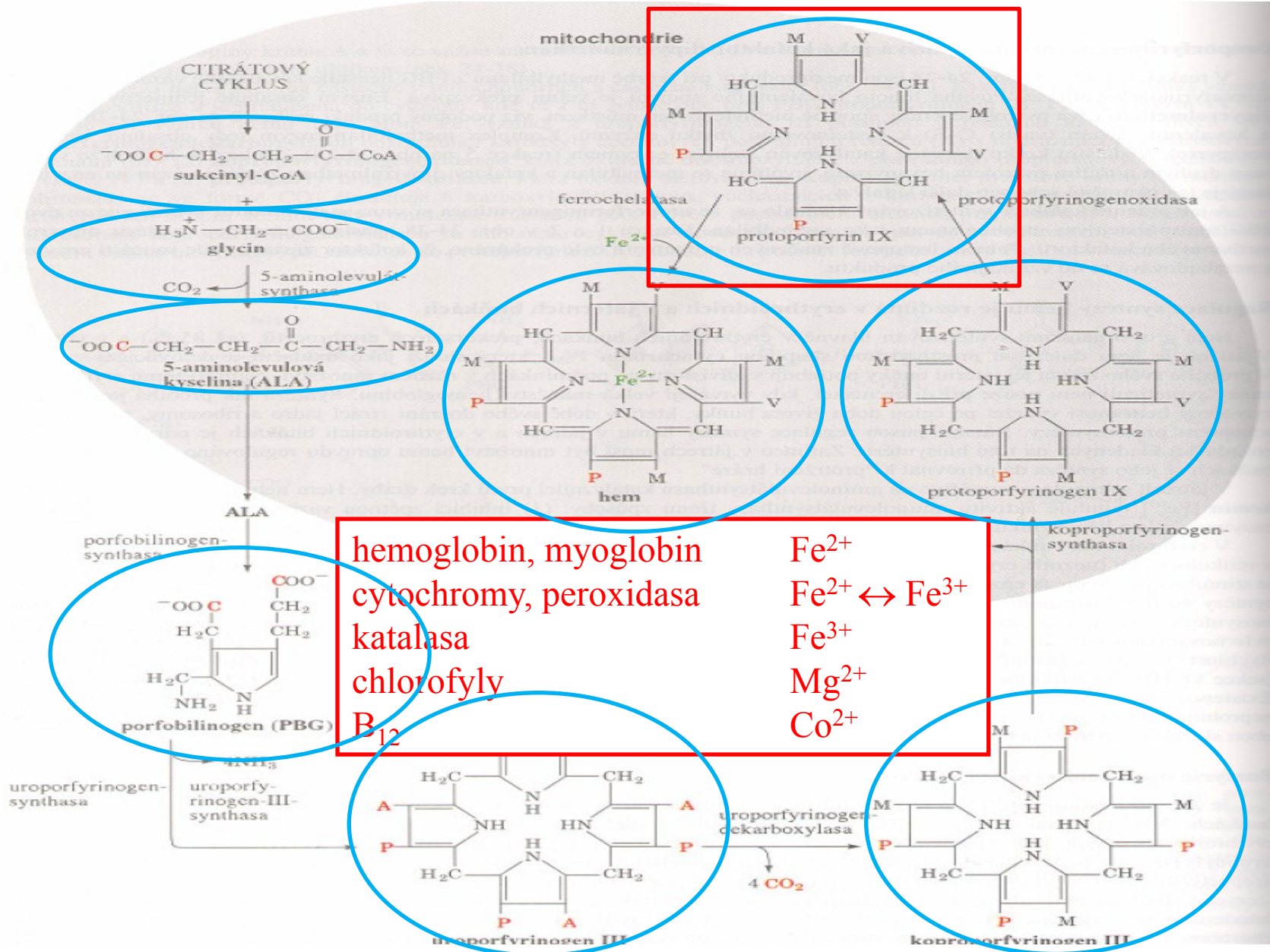
Bioactive Product	Biological Function	Amino Acid Precursor(s)
Alkaloids	Nitrogen bases in plants	Ornithine, Asp, Lys, Tyr, Trp, Phe, His
γ -Aminobutyric acid (GABA)	Inhibitory neurotransmitter	Glu
Auxin	Plant growth hormone	Trp
Catecholamines	Neurotransmitters, hormones	Tyr, Phe
Glutathione	Redox tripeptide	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
Porphyrin	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



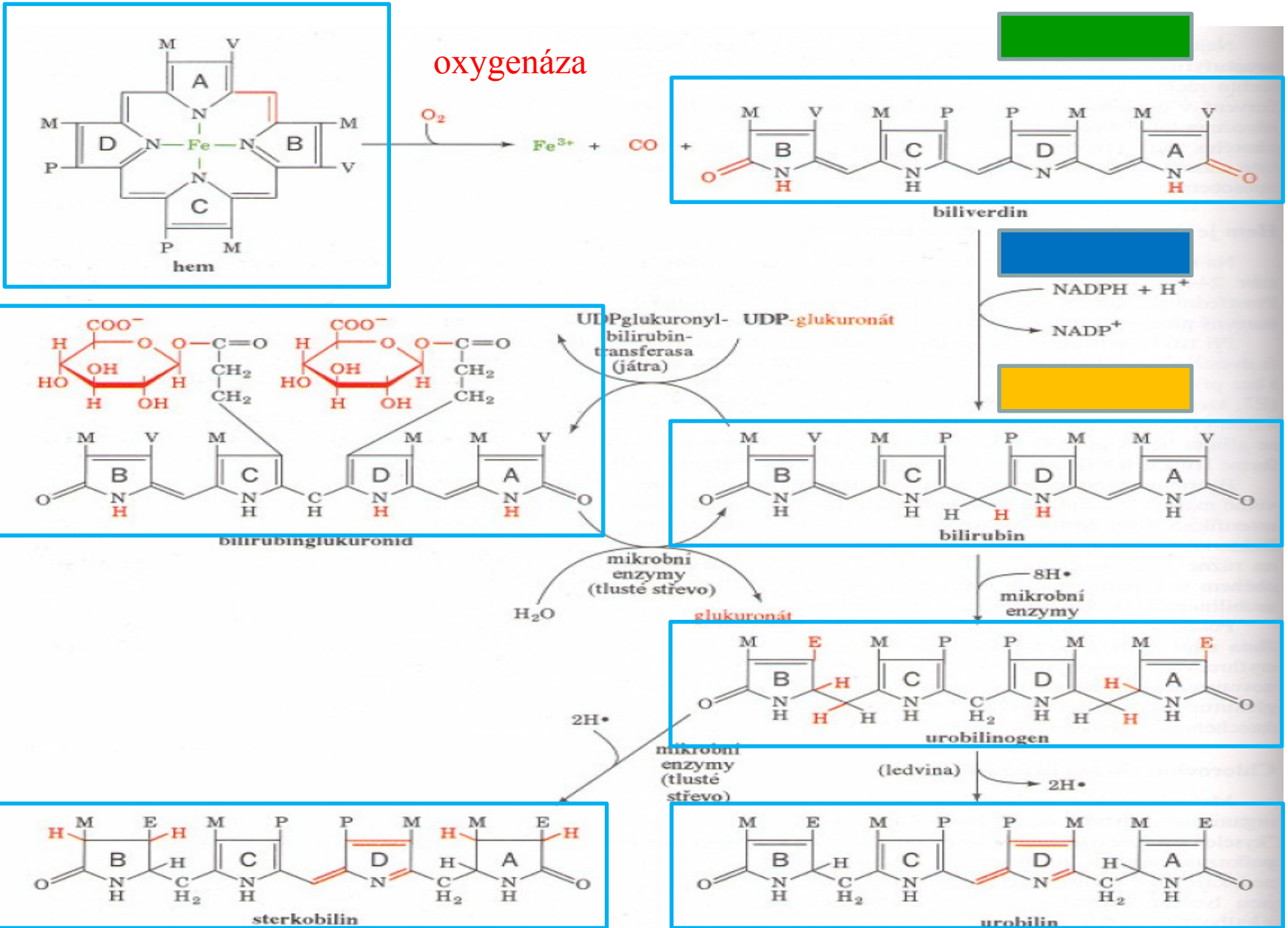
Hem



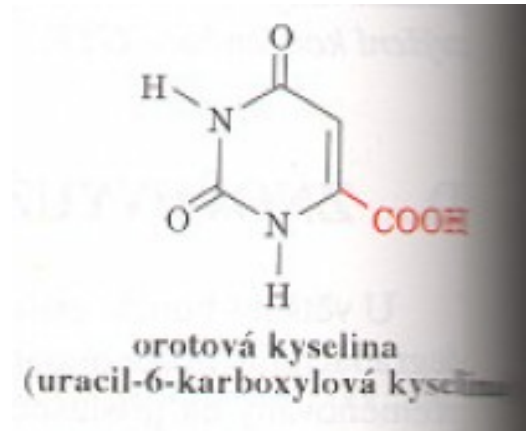
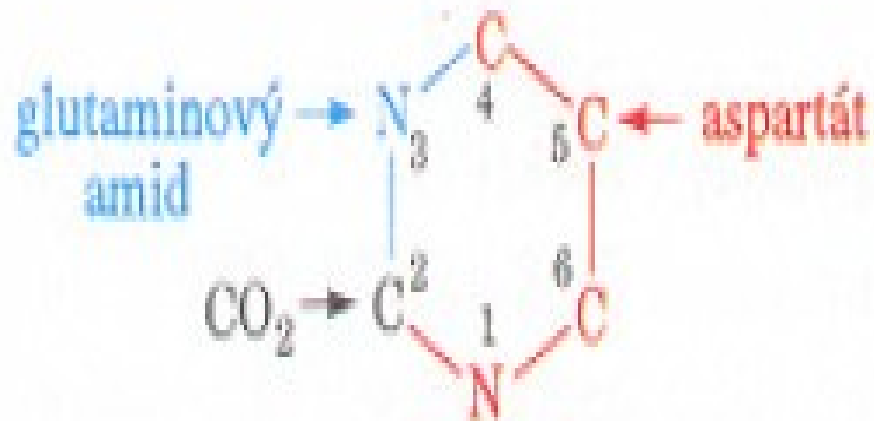
Biosyntéza hemu



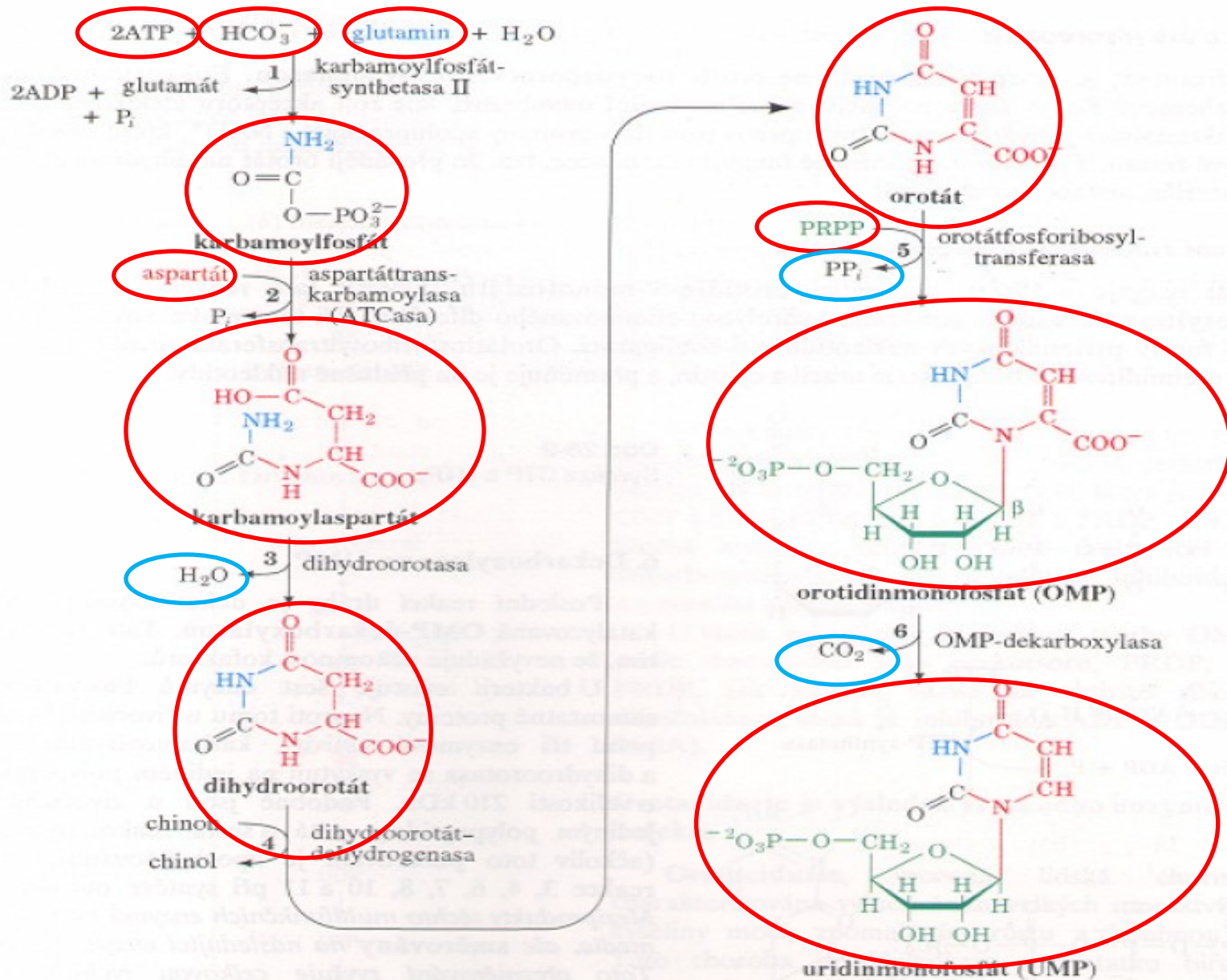
Degradace hemu



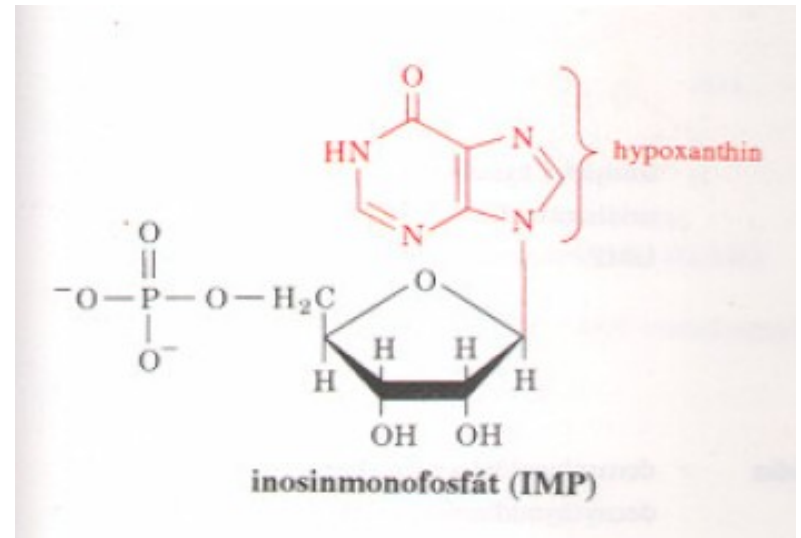
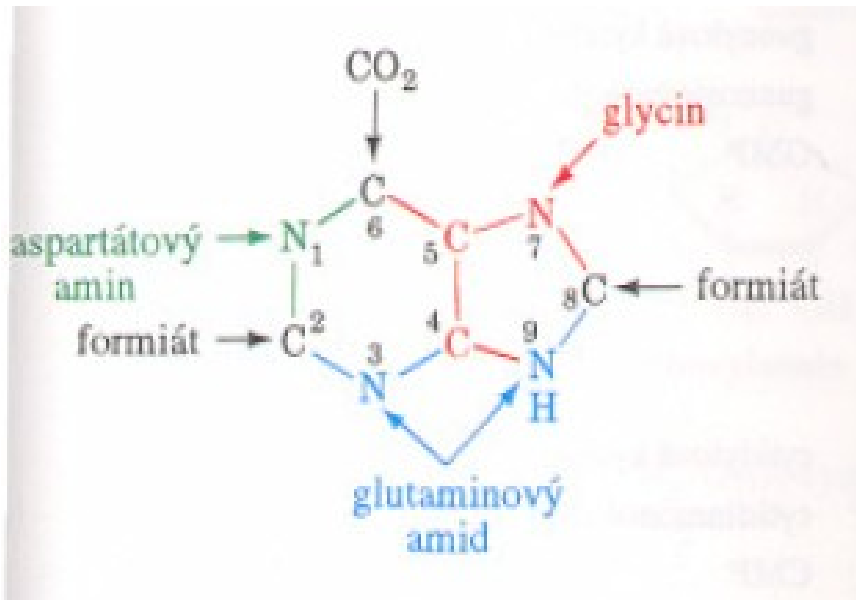
Biosyntéza pyrimidinových bází



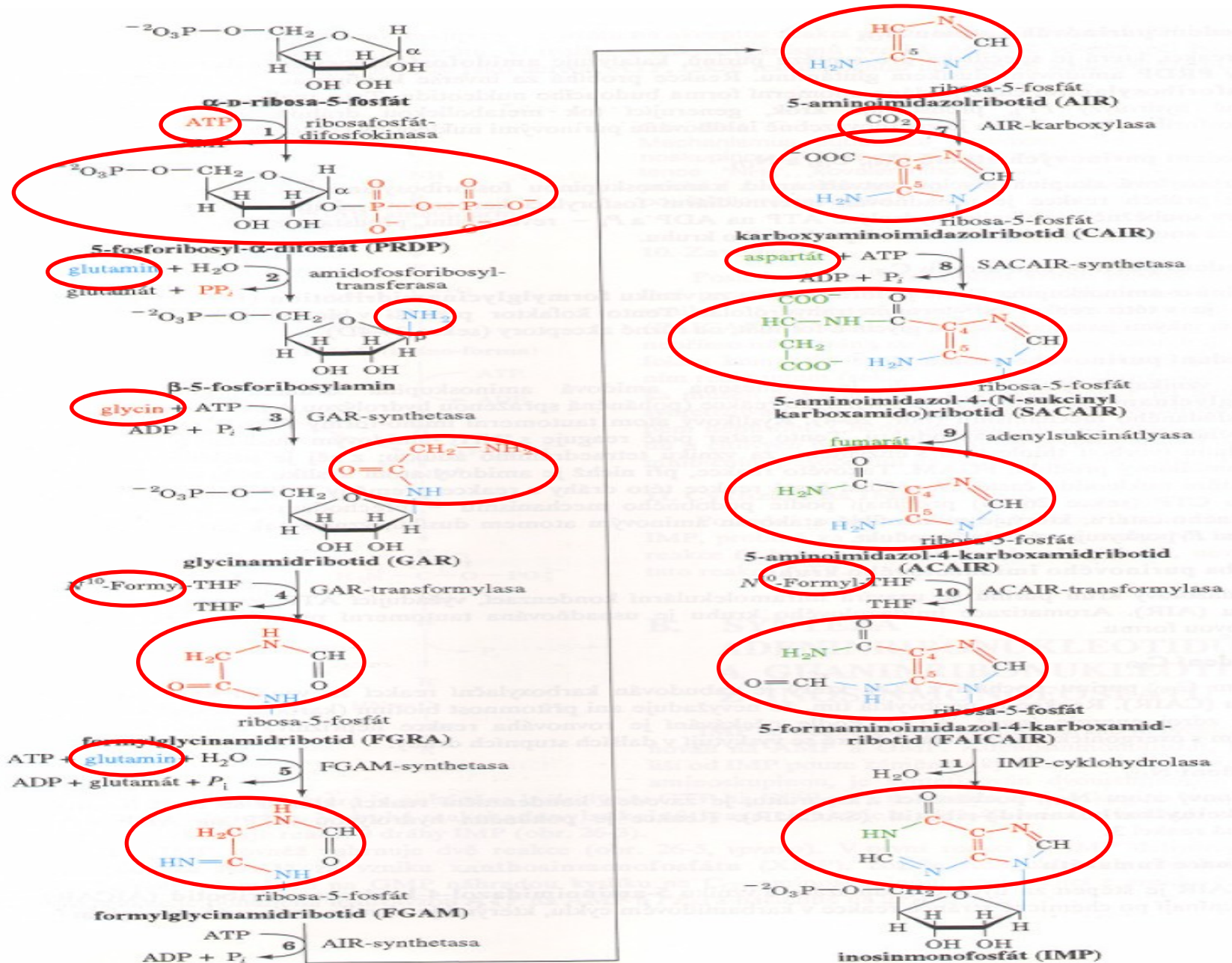
Biosyntéza pyrimidinových bází



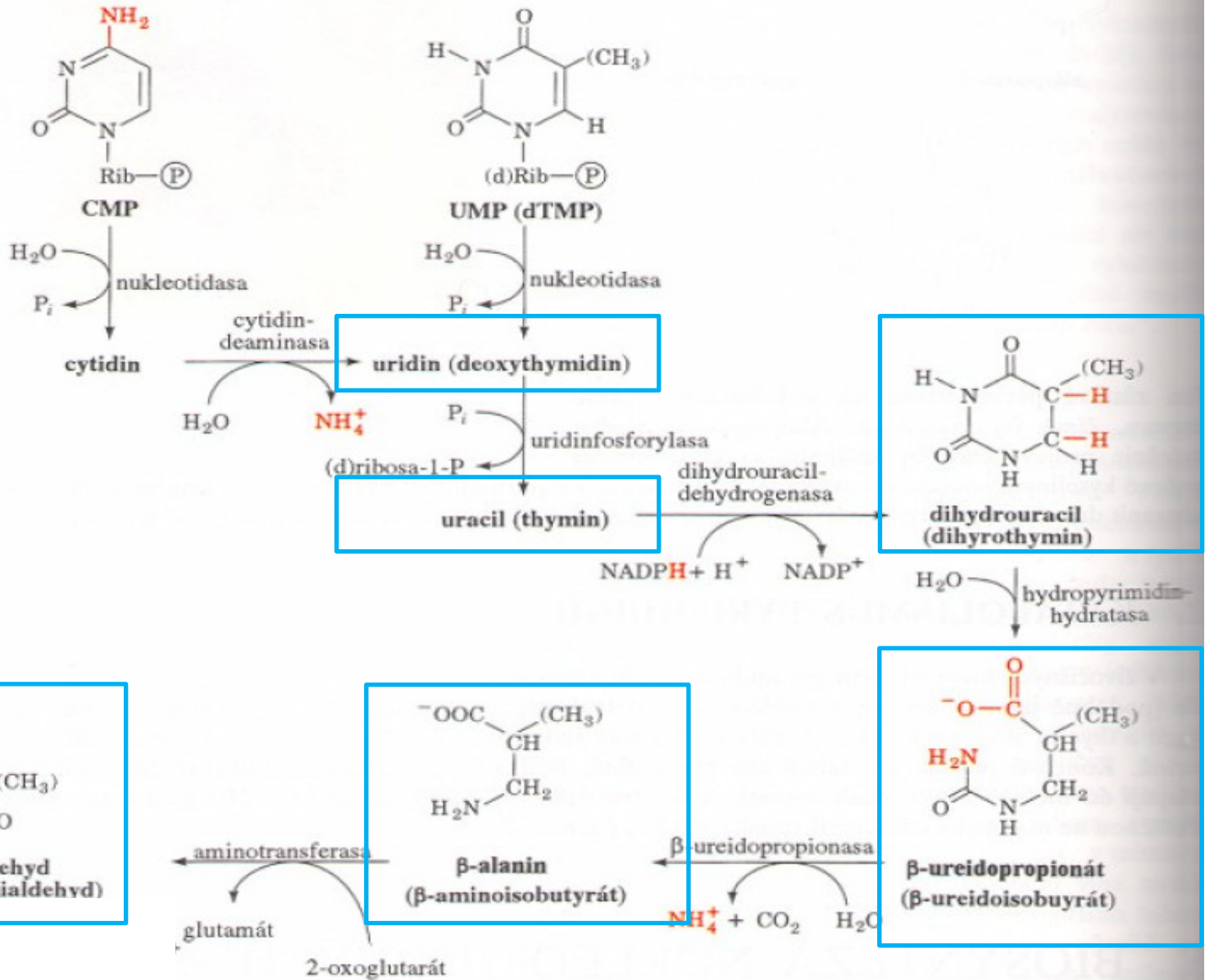
Biosyntéza purinových bází



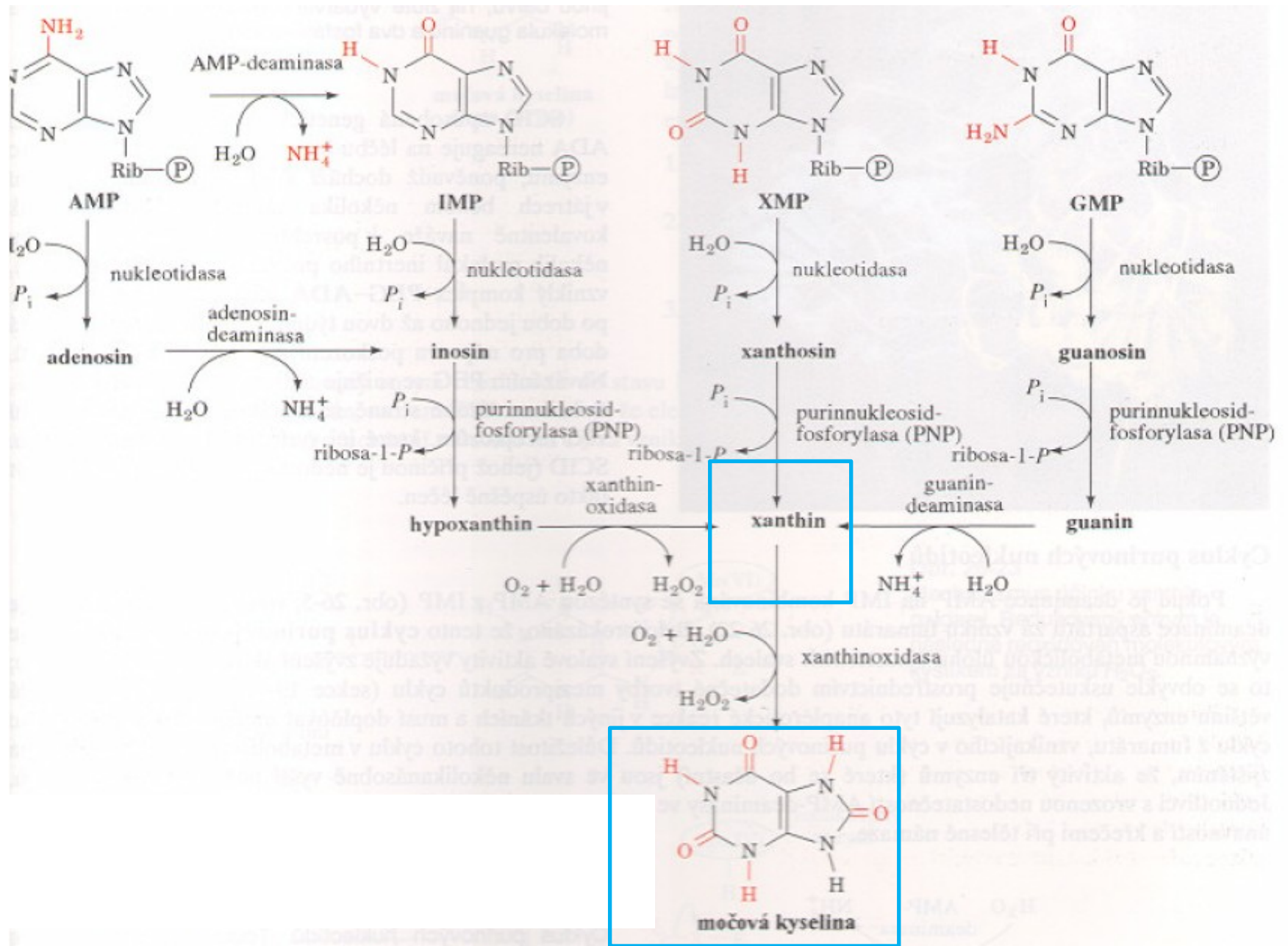
Biosyntéza purinových bází



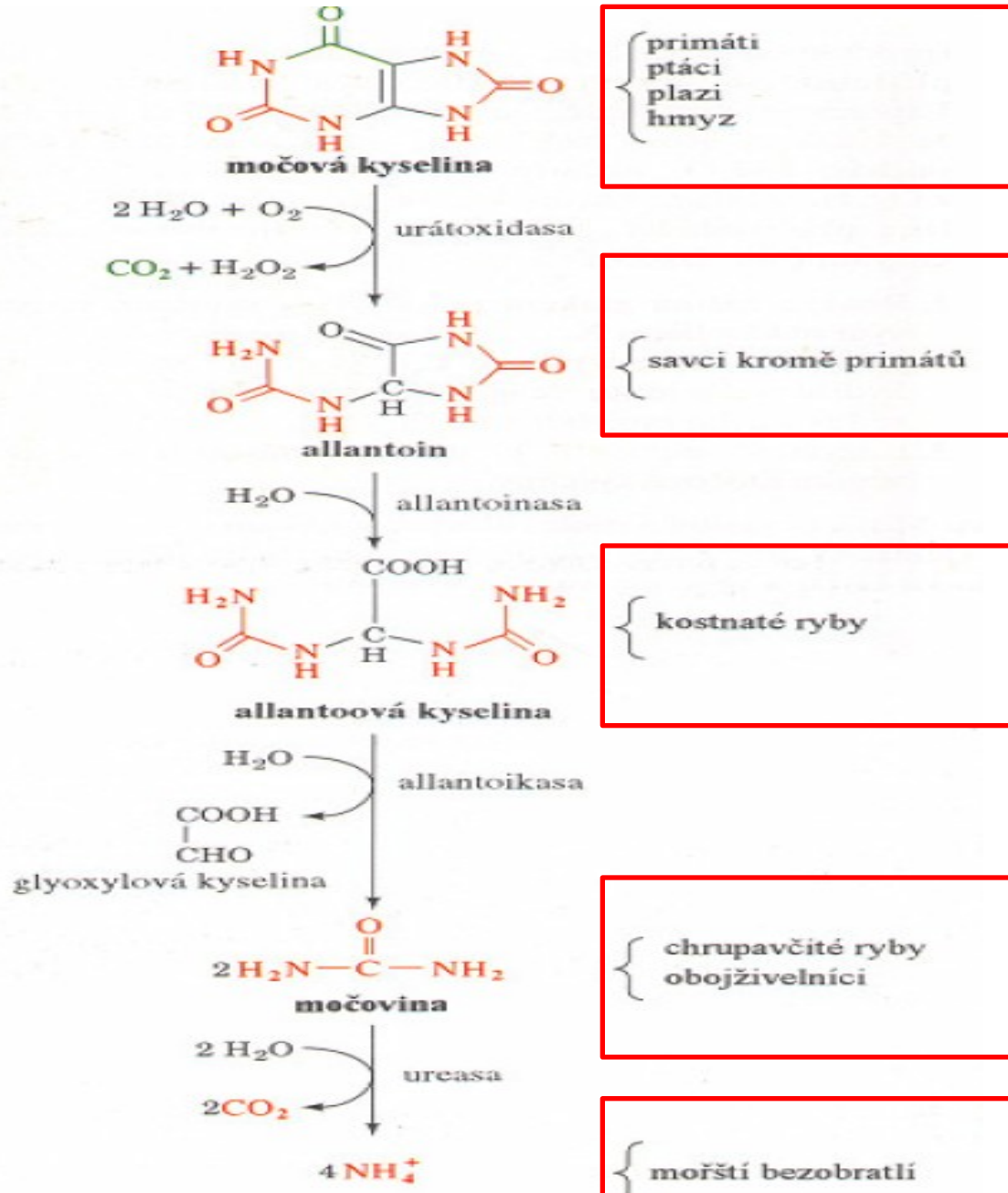
Degradace pyrimidinových bází



Degradace purinových bází



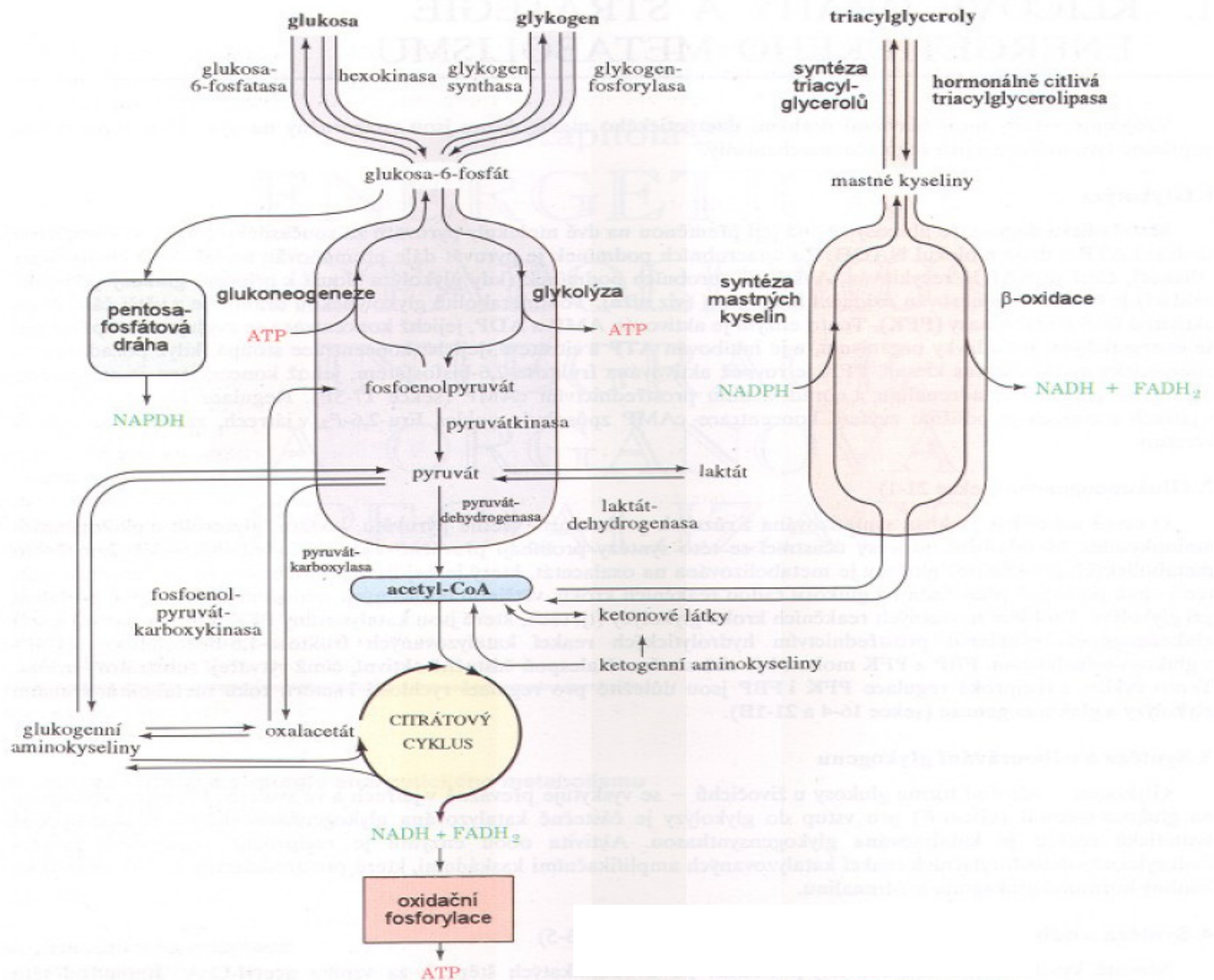
Degradace purinových bází



Metabolismus NK

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

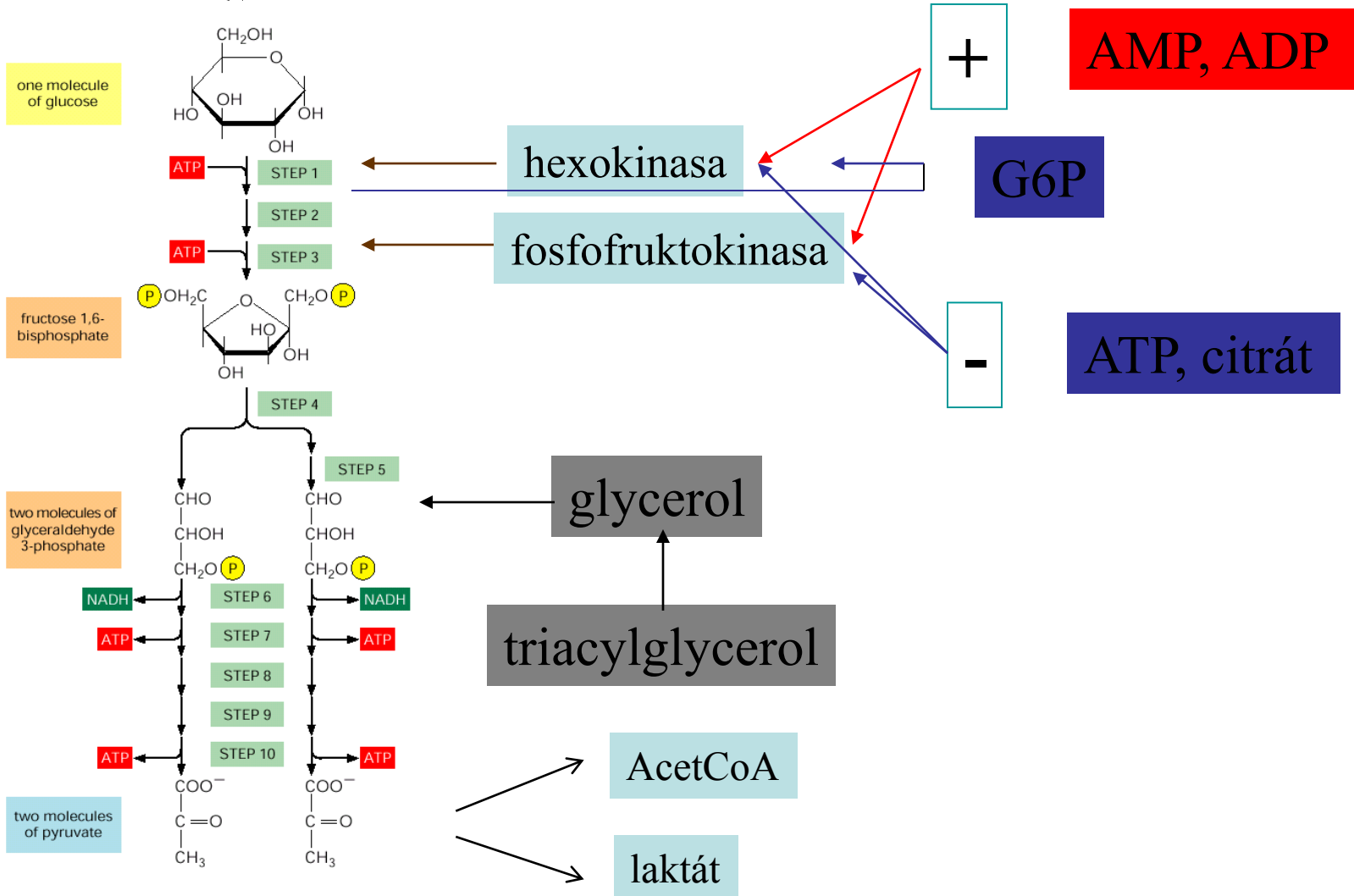
Hlavní dráhy energetického metabolismu



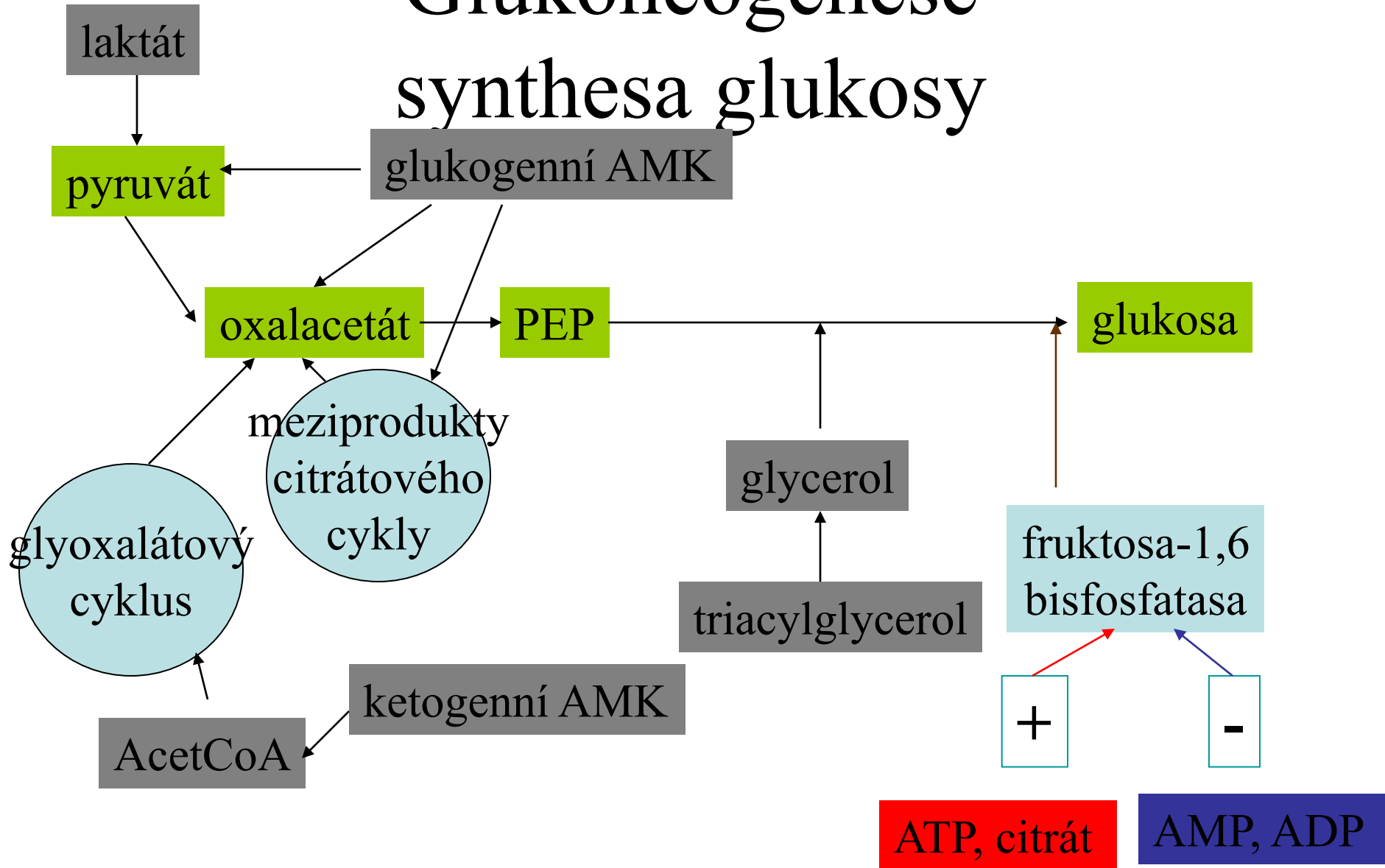
Glykolyza zisk energie

glykogen

glukosa monosacharidy

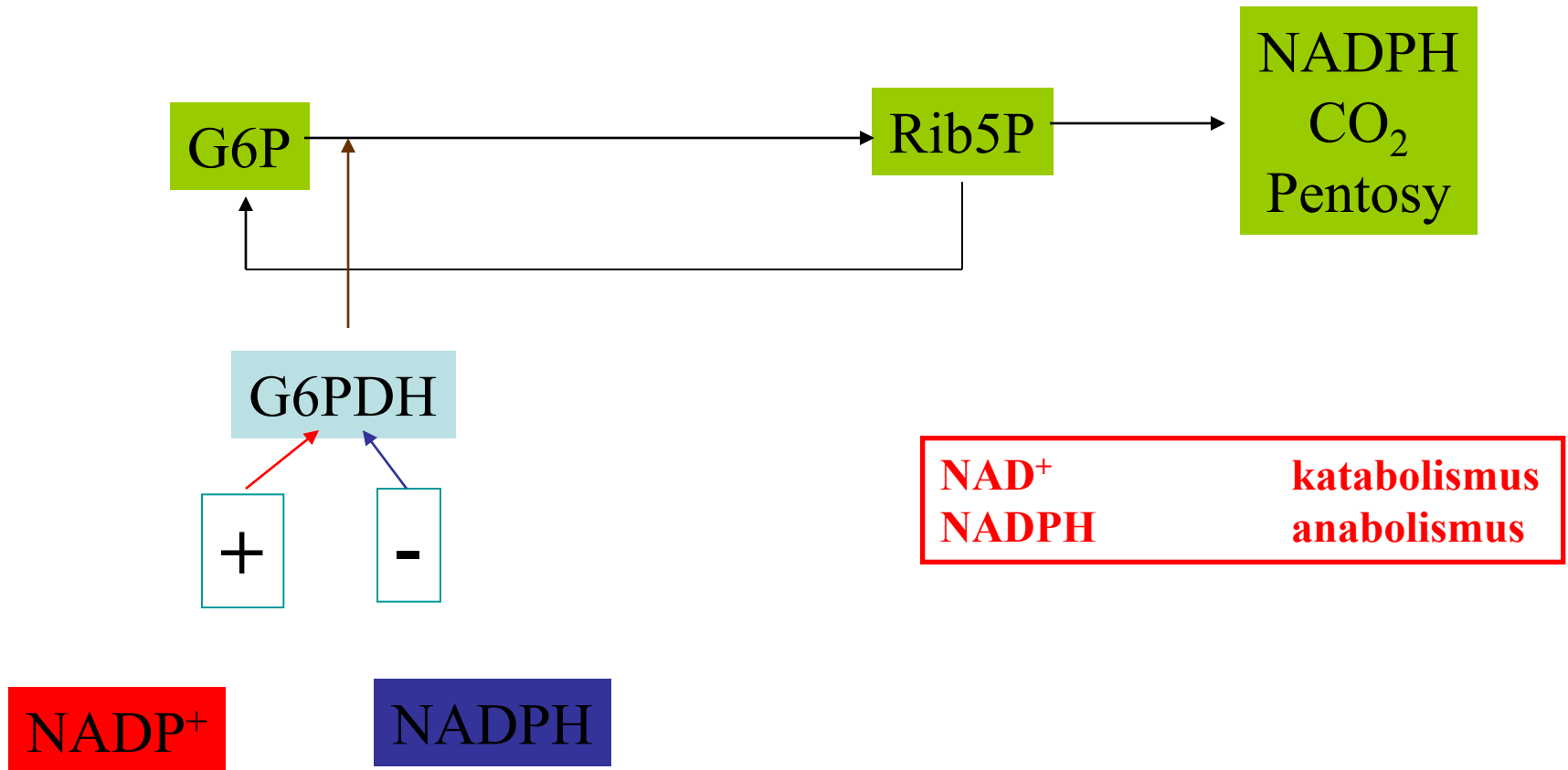


Glukoneogeneze synthesa glukosy

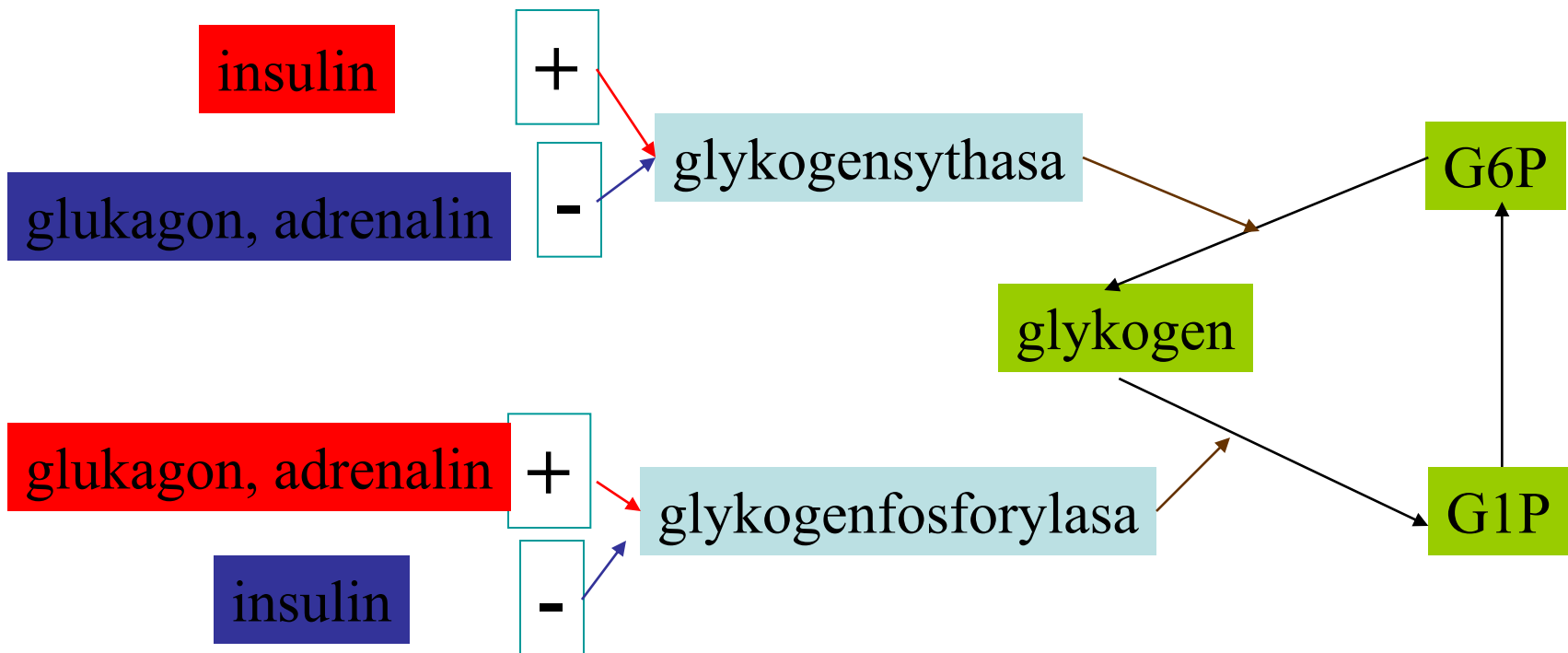


Pentosový cyklus

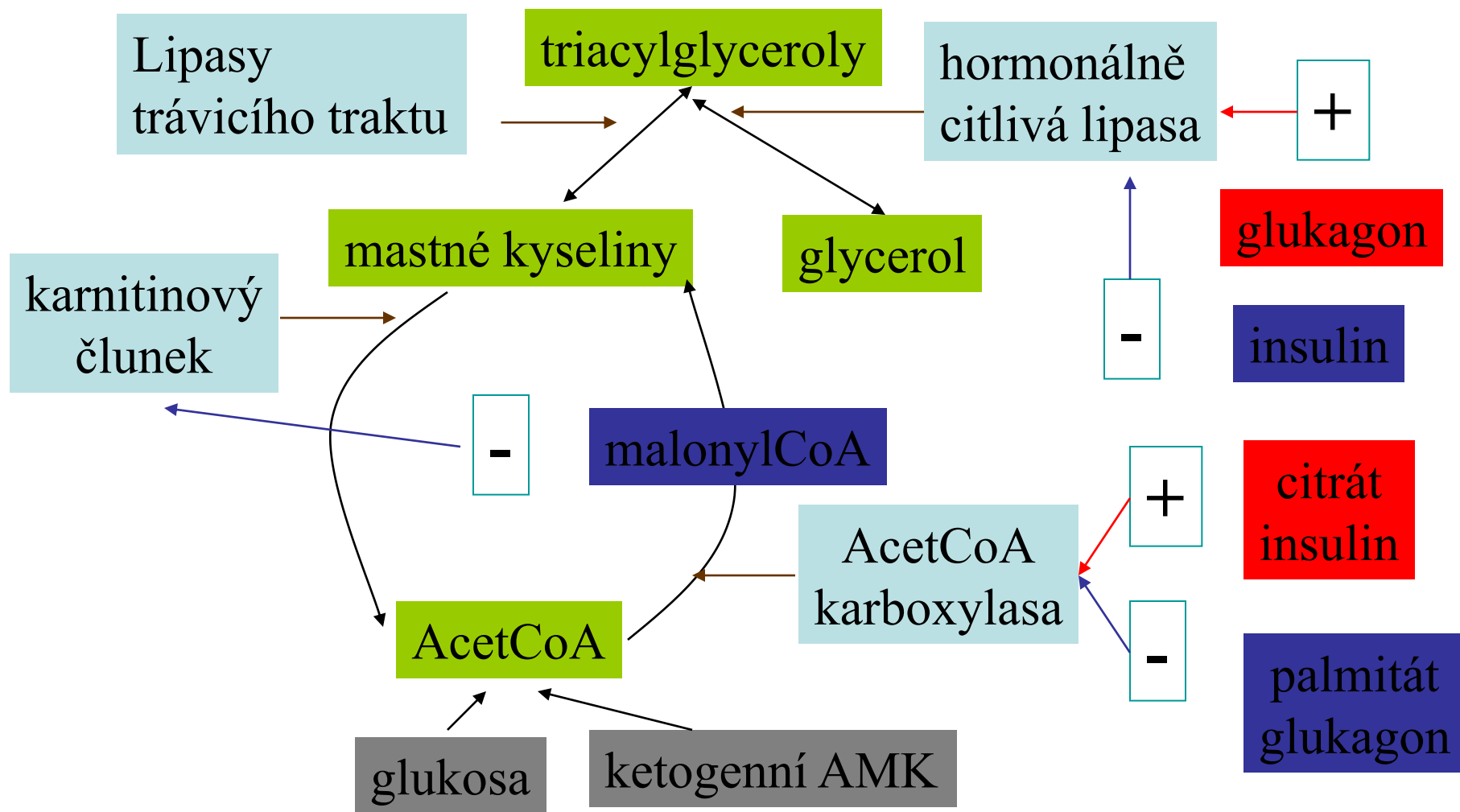
NADPH pro biosynthesu



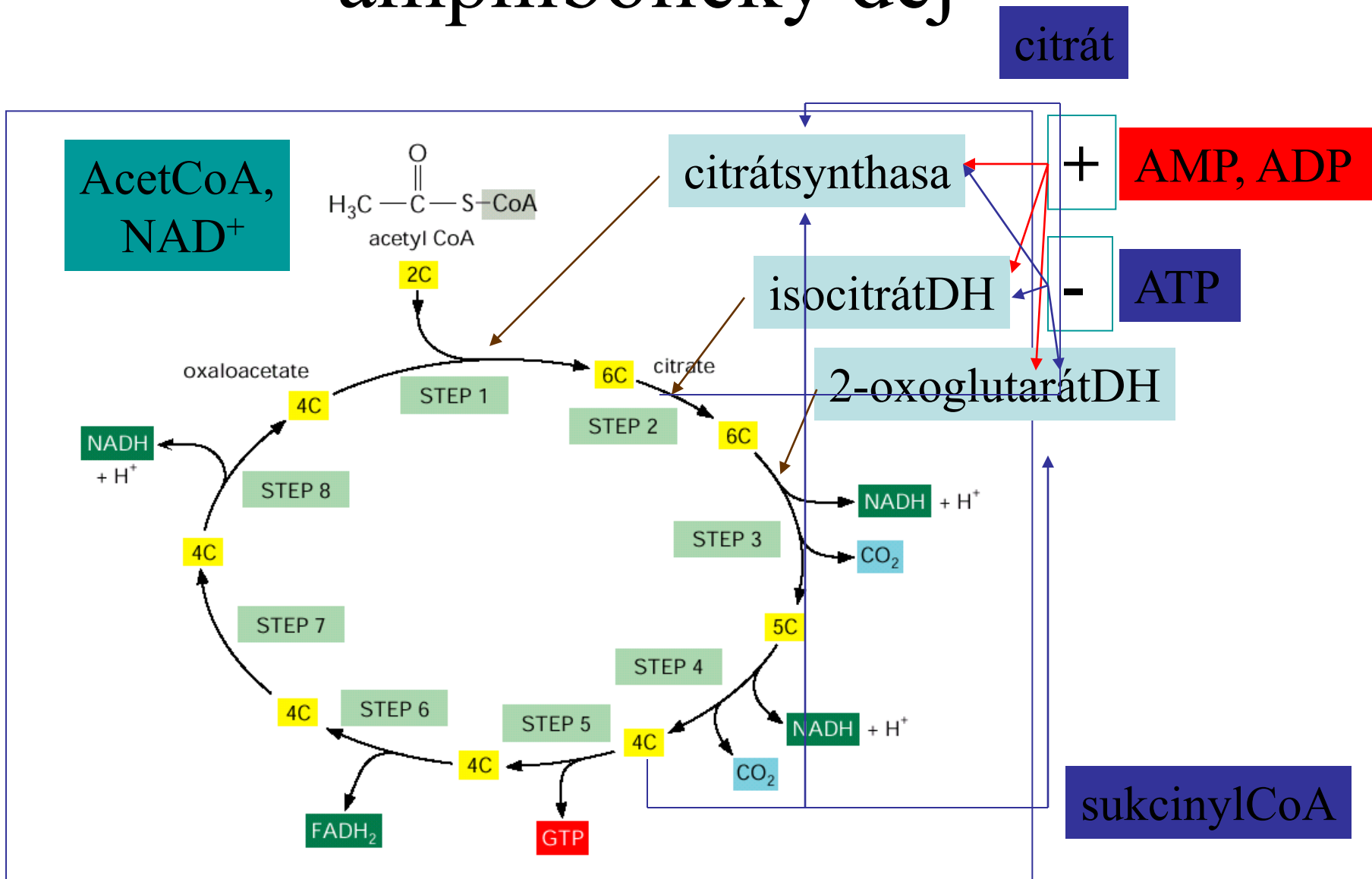
Synthesa a odbourávání glykogenu



Metabolismus triacylglycerolů

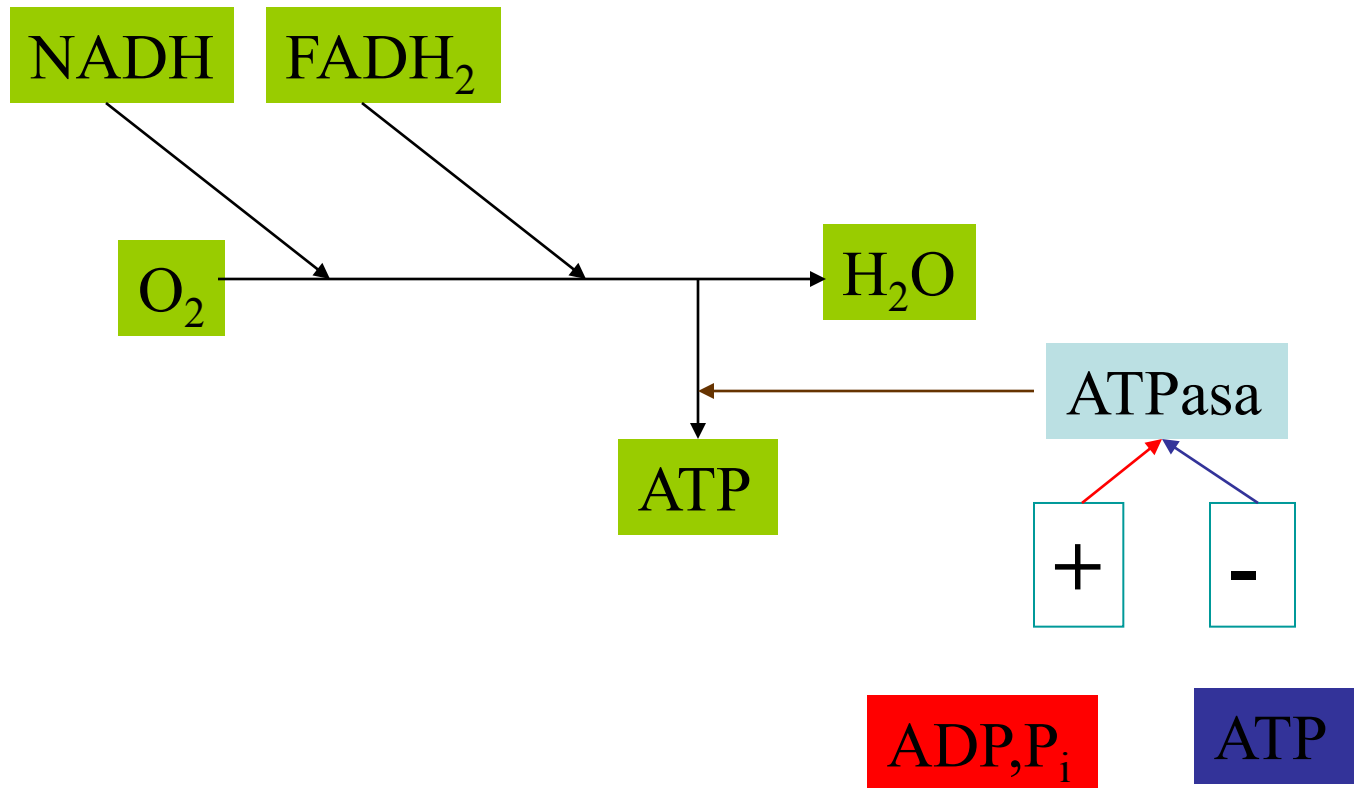


Citrátový cyklus amphibolický děj



Oxidační fosforylace

zisk ATP



Orgánová specializace

Hnědá tuková tkáň
versus
Bílá tuková tkáň

