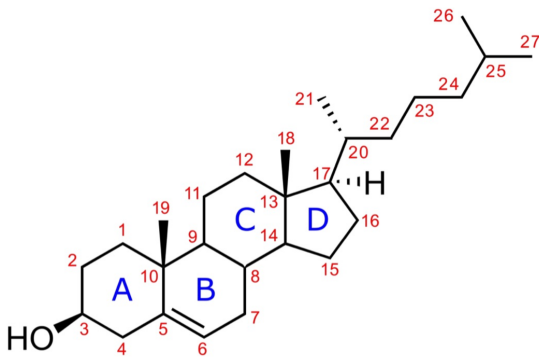


* Numbering system of Cholesterol



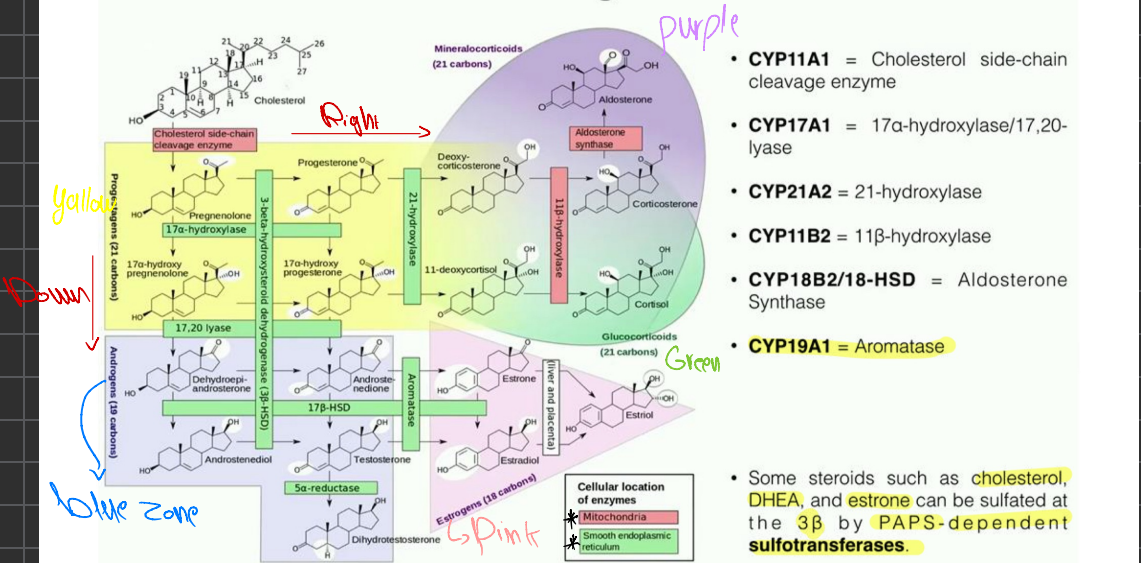
1) the site of OH (C₃)

2) where the tail begins (22-27)

3) 4 rings (3 six membered rings and 1 five membered ring)
 a) the site of Double bond.

* based on this system the enzymes took their names

Steroidogenesis



- **CYP11A1** = Cholesterol side-chain cleavage enzyme
- **CYP17A1** = 17 α -hydroxylase/17,20-lyase
- **CYP21A2** = 21-hydroxylase
- **CYP11B2** = 11 β -hydroxylase
- **CYP18B2/18-HSD** = Aldosterone Synthase
- **CYP19A1** = Aromatase

• Some steroids such as **cholesterol**, **DHEA**, and **estrone** can be sulfated at the 3 β by **PAPS-dependent sulfotransferases**.

* The Zones * The type of enzymes * The directions

- The 1st step (remove the tail) ~> the tail is carboxaldehyde and pregnenolone is ketone (the parent steroid).

- 17 α hydroxylase ~> hydroxylase C 17

- 3 beta hydroxyl steroid dehydrogenase (Pregnenolone \rightarrow progesteron)

oxidation of C₃

isomerization of double bond
5=6 \Rightarrow 4=5

- Progesteron can act independently

* Convert Progesteron to Cortisol \Rightarrow 1) 17 α hydroxylase

hydroxylate C₁₇ \leftarrow 2) 21 hydroxylase produces

- a) deoxycorticosteron
- b) 11-deoxycortisol

must be transported to mitochondria \leftarrow
by simple diffusion.

- hydroxylate C₁₁
- mitochondrial enzyme

- 3) 11 β hydroxylase produces
- a) corticosteron
 - b) cortisol (glucocorticoid)

* Aldosteron synthesis

Corticosteron

Aldosteron synthase

Aldosteron

- 1) hydroxylation of C₁₈
- 2) oxidation of C₁₈

end with aldehyde

* Cortisol and aldosteron are **Deadend**.

act independently on their receptors

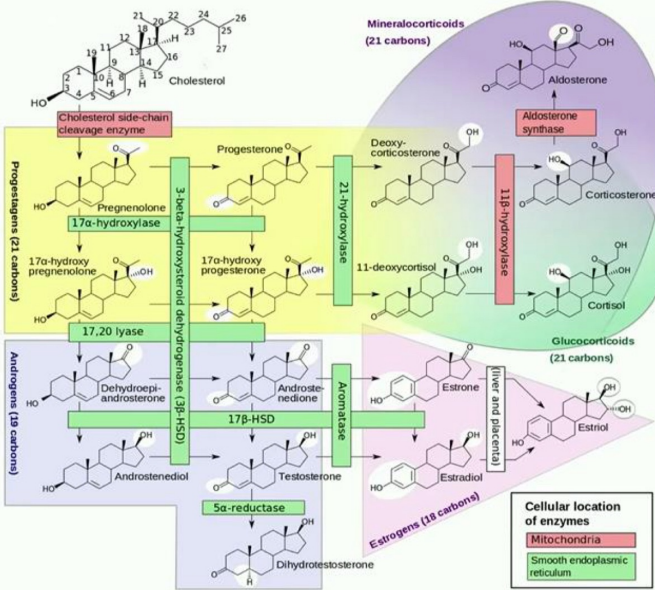
metabolize in liver.

* Development of sex hormones.

We need two progesteron (17 α hydroxyl progesteron and 17 α hydroxyl pregnenolone).

2) 17-20 lyase use the OH group on C₁₇ to remove the keton group attached to the C₁₇ (this enzyme has activity like 17 α -hydroxylase) to produce DHEA and androstenedione the 1st two androgens.

Steroidogenesis



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* In progesterone \rightarrow keton tail on C₁₇, in androgens \rightarrow No !!

3) 3 β HSD which will oxidize 1) DHEA into androstenedione
2) androstenediol into testosterone.

4) 17 β HSD 1) androstenedione for testosterone
2) DHEA for androstenediol.

5) 5 α reductase testosterone reduction \rightarrow Dihydrotestosterone

\checkmark H on C₅ \rightarrow use NADPH \rightarrow remaining of (=) \rightarrow most potent androgen !!
 \hookrightarrow this process increase the potency from 20-100 folds

* **Estrogen synthesis**

- Aromatase (hydroxylation of the C₃, convert A ring to Aromatic)

1) androstenedione

→ estrone (less active estrogen)

2) testosterone

→ estradiol (primary estrogen)

↳ both metabolized in liver and placenta to be estrone

* estradiol $\xrightarrow{16\text{-hydroxylase}}$ estrone

* estrone $\xrightarrow{1) 17\text{BHS}, 2) 16\text{-hydroxylase}}$ estrone

* 16-hydroxylase \rightsquigarrow hydroxylase C16

* 17BHS \rightarrow reduce ketone group into OH group.

-By Luqman Ahmad

022 endo



Good luck ✨