

SUMMARY

- 1) In the first part of this investigation, an adequate collection of solvent systems and colour reagents for the resolution and identification of different C-19 and C-21 steroids was selected.
- 2) 147 local fungal isolates, which belong to 11 species and 5 genera of order Mucorales, were tested for their ability of transformation of progesterone to its hydroxy derivatives.
- 3) Only six isolates achieved an oxidative splitting of the side chain of progesterone in position C-17 with the formation of  $\Delta^4$ -androsterone-3,17-dione  $\rightarrow$  testosterone  $\rightarrow$  testolactone as the final steroid product. These isolates belong to Circinella musca (3 isolates) and M. circinell-oides (3 isolates) both of family Mucoraceae.
- 4) In the remaining 141 isolates of Mucorales, hydroxylation at C-11 was achieved either at site 11  $\alpha$ -only or at both sites 11  $\alpha$ -and 11 $\beta$ . None of them could hydroxylate the steroid molecule at site 11 $\beta$ -only.
- 5) 11 $\beta$ -Hydroxylation frequently occurred in case of Absidia (Fam. Mucoraceae) and Cunninghamella (Fam. Choanephoraceae).
- 6) The isolates Rhizopus (70 isolates) could be divided into two groups:

a- The first group, comprising 22 isolates of R. oryzae which could transform progesterone to 11  $\alpha$ -hydroxyprogesterone as single and final end product.

b- The second group comprising one isolate of R. nodosus and 47 isolates of R. stoloniferum (nigricans) which could transform progesterone to 11  $\alpha$ -hydroxyprogesterone, which was further transformed to a second end product 6B, 11  $\alpha$ -dihydroxyprogesterone.

7) All isolates of Mucor griseo-cyanus (3 isolates) were characterised by a unique type of hydroxylation, leading to the production of 14  $\alpha$ -hydroxyprogesterone.

8) Mucor racemosus (18 isolates) and M. hiemalis (28 isolates) were similar in the aspect of the production of 11  $\alpha$ -hydroxy in addition to 11  $\alpha$ , 17  $\alpha$ -dihydroxyprogesterone, but were different in some aspects as all isolates of M. hiemalis gave rise to 21-hydroxyprogesterone whereas all isolates of M. racemosus could convert 11  $\alpha$ -hydroxyprogesterone into 11  $\alpha$ -allopregnane-3, 20-dione.

9) All isolates of C. elegans (6 isolates) and C. echinulata (13 isolates) both of family Choonephoraceae, gave 11  $\alpha$ -hydroxylation alone or combined with 11B-hydroxylation. The only difference was that all isolates belonging to C. elegans gave 6B, 11  $\alpha$ -dihydroxyprogesterone; a reaction

which was performed by some, but not all, isolates of C. echinulata.

10) C. echinulata (isolate No. 77) was the most active isolate, therefore was selected for further experiments.

a- The maximum production of  $17\alpha$ -hydroxyprogesterone was estimated: after 48 hours incubation; at pH.6.5; and in presence of 2 gm/ litre phosphate.

b- Acetate and fumarate were favourable for production  $17\alpha$ -hydroxyprogesterone.

c- Olive oil stimulated  $17\alpha$ -hydroxylation as, cotton seed oil stimulated  $11\alpha$ - and  $17\alpha$ -hydroxylation and castor oil promoted  $11\alpha$ -hydroxylation with no significant effect on  $17\alpha$ -hydroxylation.

d- Tween 80 supported the highest production of  $17\alpha$ -hydroxyprogesterone among all tweens used.

e- All water-soluble alcohols used, inhibited the production of both  $11\alpha$ - and  $17\alpha$ -hydroxylation, except t-butyl which stimulate  $17\alpha$ -hydroxylation.

f- By using C. echinulata (isolate No. 77) and a laboratory fermentor, 20 gm of added progesterone per 2 litres culture medium yielded  $17\alpha$ -hydroxyprogesterone

(40.40 %), 11  $\alpha$  -hydroxyprogesterone (23.65 %) and 11  $\alpha$  ,  
17  $\times$  -dihydroxyprogesterone (20.35 %).

g- Q. echinulata (isolate No. 77) is thus strongly recommen-  
ded for the technical production of 17  $\alpha$  -hydroxyprogesterone.