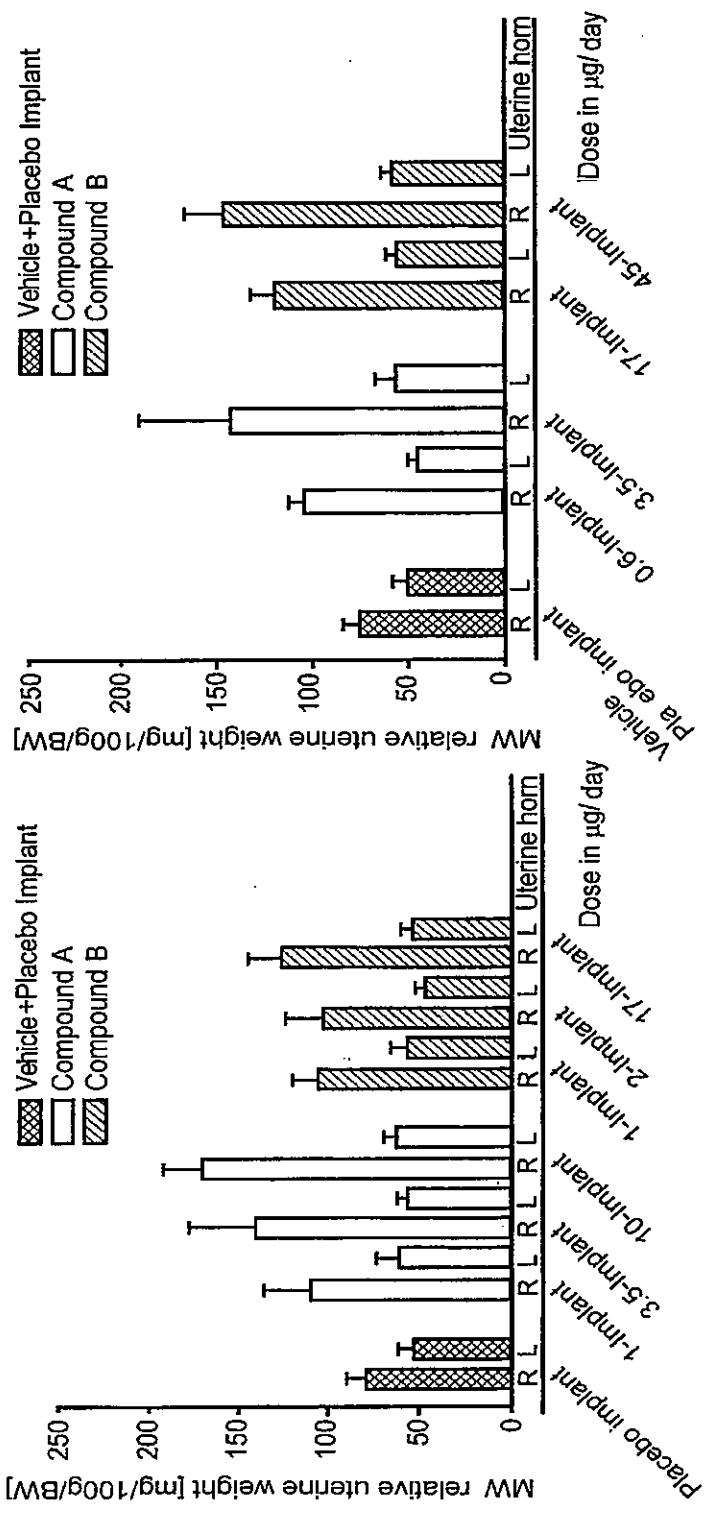


Local progestin administration induces dose-dependent weight gain in the IUS-carrying right uterine horn (R). The left uterine horn (L) does not carry an IUS and does not show any weight gain.

Fig. 1

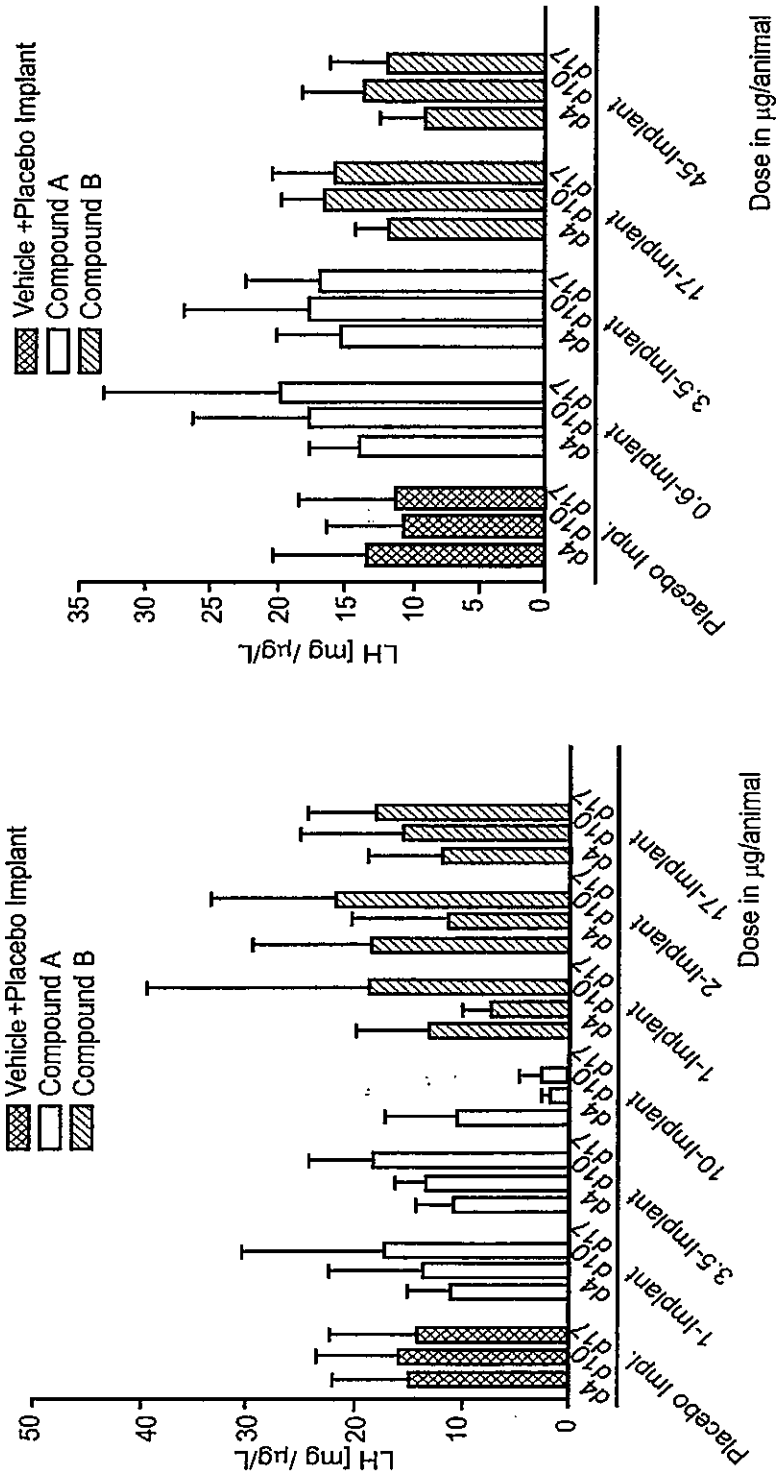
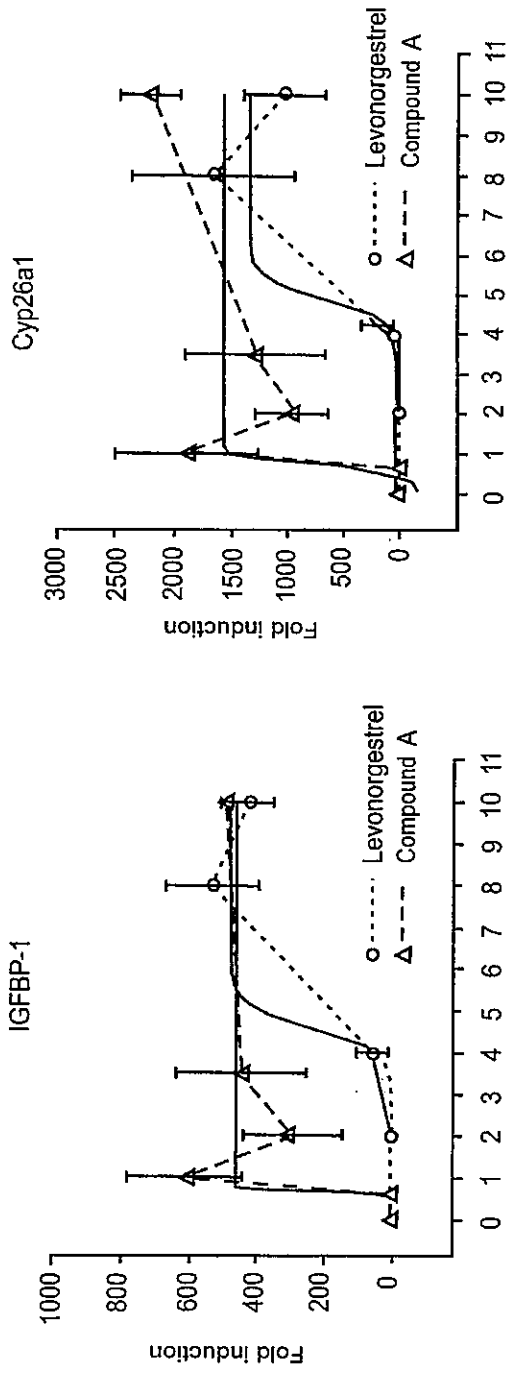


Fig. 2

No decrease in LH levels occurs with compound (A) in the dose range of up to 3.5 µg.
No decrease in LH levels occurs with compound (B) in the dose range of up to 45 µg.



Compound (A) exhibits a distinct increase in gene expression in the dose range from 1 µg/day upwards.
Levonorgestrel exhibits a distinct increase in gene expression in the dose range from 8 µg/day upwards.

Fig. 3

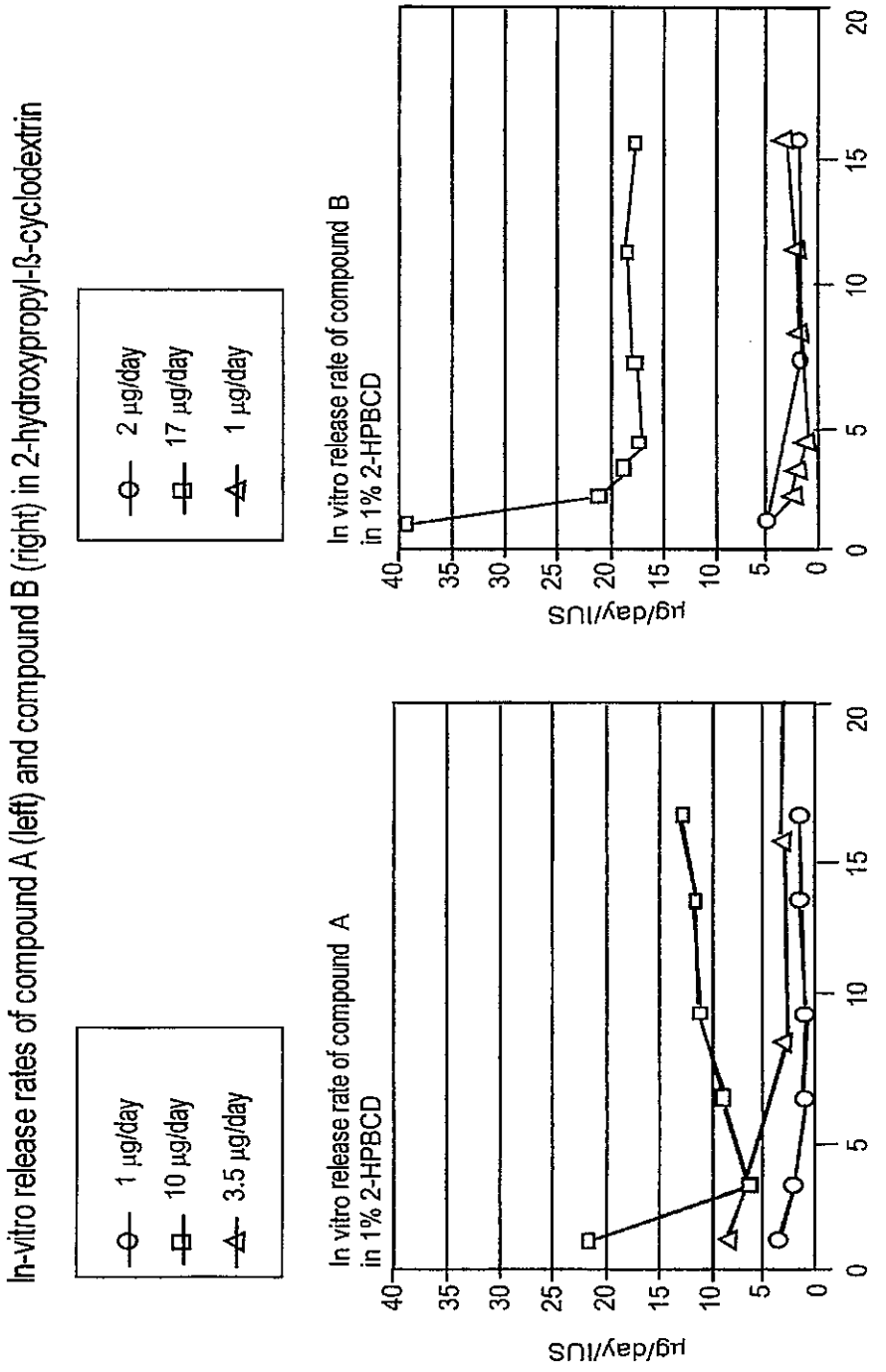

In-vitro release rates of compound A (left) and compound B (right) in 2-hydroxypropyl-ß-cyclodextrin
1 µg/day
10 µg/day
3.5 µg/day
2 µg/day
17 µg/day
1 µg/day
In vitro release rate of compound A
in 1% 2-HPBCD
In vitro release rate of compound B
in 1% 2-HPBCD
µg/day/IUS
0
5
10
15
20
25
30
35
40

Fig. 4