Supplemental Data

Supplemental Table S1: Vinclozolin P60-150 generation animal body, testis weight and serum testosterone (T) levels.

Generation	Body Weight (gm)	Testis Weight (mg)	Testis/Body Ratio x10 ⁻³	T Levels (ng/ml)
F1 Control	513 <u>+</u> 12.63	3.95 ± 0.14	7.79 <u>+</u> 0.25	3.34 ± 0.39
F1 vinclozolin	517 <u>+</u> 12.00	3.67 ± 0.08	7.11 <u>+</u> 0.19	3.88 ± 0.40
F2 Control	416 <u>+</u> 25.13	3.54 ± 0.12	8.64 <u>+</u> 0.34	5.35 <u>+</u> 1.14
F2 vinclozolin	414 <u>+</u> 27.36	3.41 ± 0.09	8.47 <u>+</u> 0.46	4.78 <u>+</u> 1.14
F3 Control	551 <u>+</u> 14.96	4.18 <u>+</u> 0.09	7.60 <u>+</u> 0.17	3.65 ± 0.42
F3 vinclozolin	537 ± 15.80	3.66 ± 0.08	6.85 <u>+</u> 0.11	3.62 ± 0.30

The different generations data were collected at different times. Exact age matched controls were used within a generation, but ages varied between generations within the developmental period indicated.

Supplemental Table S2: Flutamide and $P60-150^+$ generation animal body, testis weight serum testosterone (T) levels.

	Body Weight	Testis Weight	Testis/Body Ratio	T Levels
Generation	(gm)	(mg)	x10 ⁻³	(ng/ml)
F1 Control	409 <u>+</u> 40.27	3.33 + 0.13	8.45 <u>+</u> 0.53	2.17 <u>+</u> 0.49
F1 Flutamide 5mg	405 <u>+</u> 32.37	3.21 <u>+</u> 0.18	8.02 <u>+</u> 0.26	7.35 <u>+</u> 1.36(*)
F1 Flutamide 20mg	445 <u>+</u> 34.37	3.38 <u>+</u> 0.16	7.76 <u>+</u> 0.35	2.90 <u>+</u> 0.36
F2 Control	480 <u>+</u> 25.40	3.66 <u>+</u> 0.07	7.89 <u>+</u> 0.43	3.75 <u>+</u> 0.45
F2 Flutamide 5mg	388 <u>+</u> 18.53	2.86 <u>+</u> 0.16	7.45 <u>+</u> 0.49	2.17 <u>+</u> 0.41
F2 Flutamide 20mg	495 <u>+</u> 26.12	3.75 <u>+</u> 0.11	7.71 <u>+</u> 0.25	2.83 <u>+</u> 0.38
F3 Control [⁺]	265 <u>+</u> 2.88	2.68 <u>+</u> 0.14	10.32 <u>+</u> 0.41	2.90 <u>+</u> 0.26
F3 Flutamide 5mg	402 <u>+</u> 28.39	3.52 <u>+</u> 0.15	8.81 <u>+</u> 0.24	1.14 <u>+</u> 0.34
F3 Flutamide 20mg	390 <u>+</u> 31.89	3.37 <u>+</u> 0.08	8.84 <u>+</u> 0.80	0.57 <u>+</u> 0.07

^(*) indicates significant difference compared to controls

The different generations data were collected at different times. Exact age matched controls were used within a generation, but ages varied between generations within the developmental period indicated. Note (+), the F3 control animals were collected at postnatal day 22, so were outside the P60-150 range for all other weights and T level data.

Supplemental Table S3: Vinclozolin P160-360 generation animal body, testis weight and serum testosterone (T) levels.

Vinclozolin animals

Generation/Dose	Body Weight (gm)	Testis Weight (mg)	Testis/Body Ratio x10 ⁻³	T Levels (ng/ml)
F1 Control	676 <u>+</u> 26.0	4.02 <u>+</u> 0.13	5.97 <u>+</u> 0.22	1.86 <u>+</u> 0.45
F1 Vinclozolin 100mg	635 <u>+</u> 17.35	3.79 <u>+</u> 0.11	5.99 <u>+</u> 0.16	1.74 <u>+</u> 0.24
F2 Control	589 <u>+</u> 18.27	3.99 <u>+</u> 0.08	6.88 <u>+</u> 0.22	1.02 <u>+</u> 0.07
F2 Vinclozolin 100mg	692 <u>+</u> 21.16	3.09 <u>+</u> 0.07	5.71 <u>+</u> 0.16	1.15 <u>+</u> 0.12
F3 Control	518 <u>+</u> 12.14	4.02 <u>+</u> 0.11	7.76 <u>+</u> 0.21	1.21 <u>+</u> 0.20
F3 Vinclozolin 100mg	601 <u>+</u> 20.0	4.00 <u>+</u> 0.05	6.75 <u>+</u> 0.20	1.47 <u>+</u> 0.22

The different generations data were collected at different times. Exact age matched controls were used within a generation, but ages varied between generations within the developmental period indicated.

Supplemental Table S4: Flutamide P160-360 generation animal body, testis weight and serum testosterone (T) levels.

Flutamide Animals

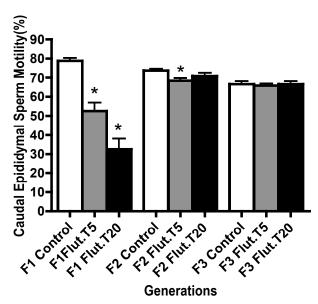
Generation/Dose	Body Weight (gm)	Testis Weight (mg)	Testis/Body Ra x10 ⁻³	tio T Levels (ng/ml)
F1 Control	325 <u>+</u> 9.27	3.10 <u>+</u> 0.10	9.53 <u>+</u> 0.20	1.28 <u>+</u> 0.39
F1 Flutamide 5mg	343 <u>+</u> 23.22	2.86 <u>+</u> 0.13	8.39 <u>+</u> 0.36	10.6 <u>+</u> 0.75 (*)
F1 Flutamide 20mg	356 <u>+</u> 12.12	3.00 <u>+</u> 0.04	8.46 <u>+</u> 0.40	2.92 <u>+</u> 0.65 (*)
F2 Control	544 <u>+</u> 19.53	3.69 <u>+</u> 0.08	6.87 <u>+</u> 0.23	1.55 <u>+</u> 0.20
F2 Flutamide 5mg	518 <u>+</u> 11.30	3.21 <u>+</u> 0.11	6.28 <u>+</u> 0.28	1.49 <u>+</u> 0.14
F2 Flutamide 20mg	565 <u>+</u> 23.92	3.96 <u>+</u> 0.06	7.14 <u>+</u> 0.23	1.42 <u>+</u> 0.37
F3 Control	576 <u>+</u> 9.34	3.79 <u>+</u> 0.05	6.60 <u>+</u> 0.17	0.622 <u>+</u> 0.03
F3 Flutamide 5mg	513 <u>+</u> 21.41	3.52 <u>+</u> 0.12	6.89 <u>+</u> 0.25	0.55 <u>+</u> 0.19 (*)
F3 Flutamide 20mg	584 <u>+</u> 10.21	3.98 <u>+</u> 0.09	6.83 <u>+</u> 0.18	0.80 <u>+</u> 0.13

^(*) denotes significant difference compared to controls

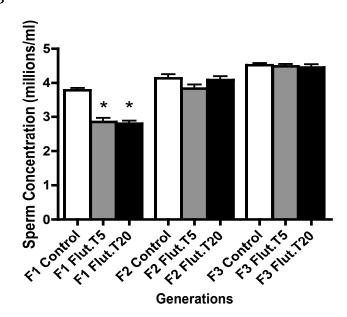
The different generations data were collected at different times. Exact age matched controls were used within a generation, but ages varied between generations within the developmental period indicated.

Supplemental Figure S1: Caudal epididymal sperm motility (A) and number (B) in P160-360 control and flutamide F1, F2 and F3 generation male rats. Statistically significant differences between control and treated generations are indicated by (*) for p < 0.05. The n value for each bar ranged between 10 and 25 animals.

A



В



Supplemental Figure S2: Photograph of low dose T5 (5mg/kg/day) Flutamide F2 generation animal with supernummary development (polymelia) of limbs. The multiple limbs are indicated with arrows.

