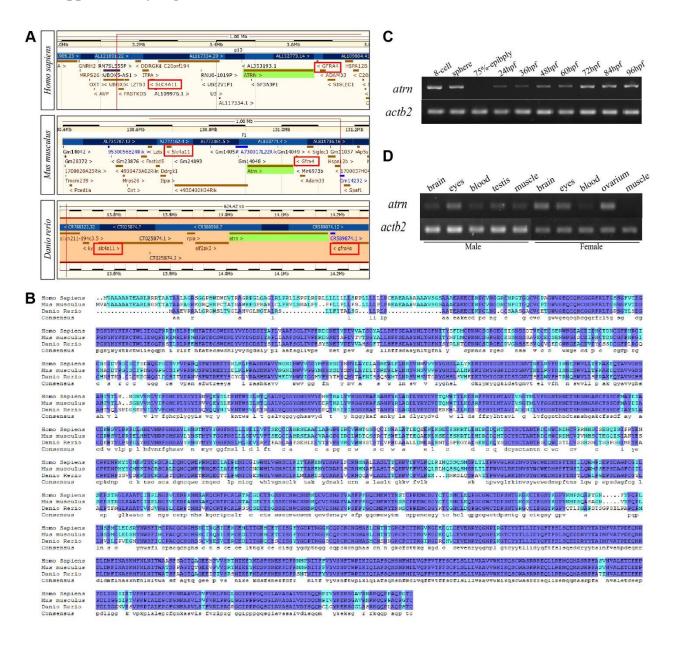


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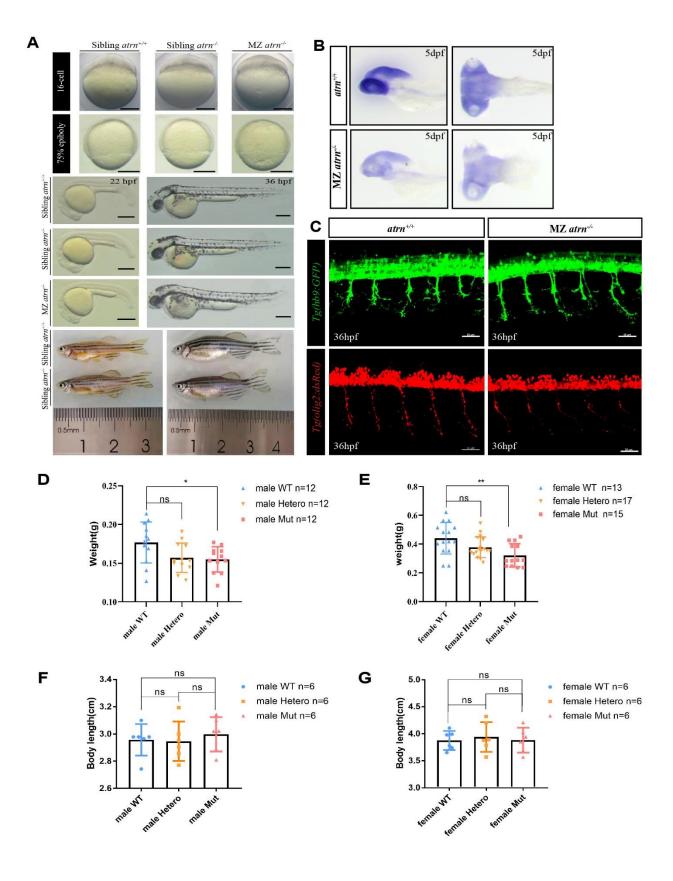
1 Supplementary Data

1.1 Supplementary Figures

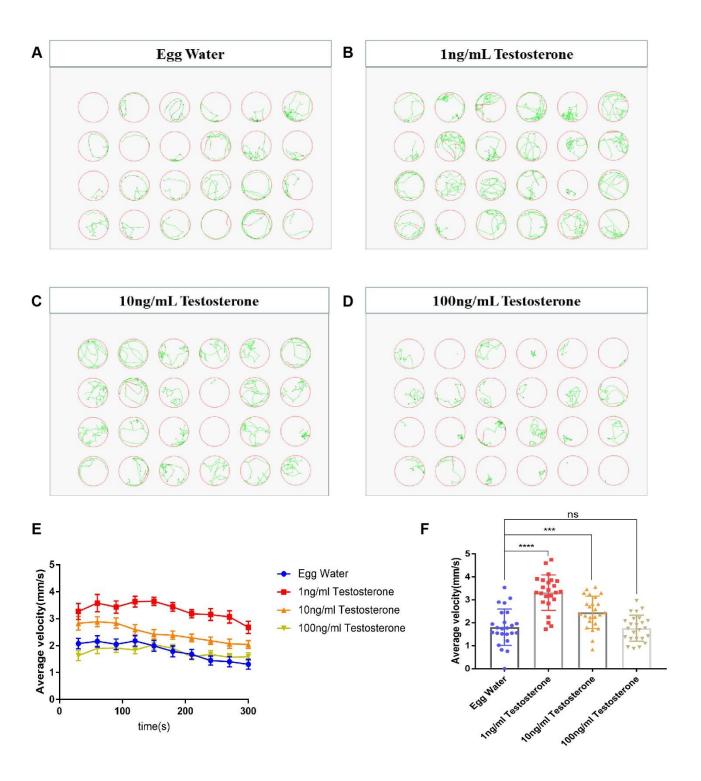


Supplementary Figure 1. Expression pattern of zebrafish ortholog gene atrn. (A) Syngeneic analysis of human, mice and zebrafish *ATRN* genes using the Ensembl genome browser (http://ensembl.org). In these three different species, there are two genes, *gfra4* and *slc4a11*, near the *ATRN* gene. (B) ATRN protein sequence alignment of human, mice and zebrafish attractin. (C) Semi-quantitative RT-

PCR detection of zebrafish *atrn* mRNA level at different development stages. (D) Semi-quantitative RT-PCR detection of zebrafish *atrn* mRNA level of adult zebrafish tissue.



Supplementary Figure 2. There are no obvious developmental defects in $atrn^{-/-}$. (A) The mutant can grow up normally with no obvious defects, and they can reproduce normally and lay eggs. (B) Reduced expression of atrn mRNA in $atrn^{+/+}$ and MZ $atrn^{-/-}$ larvae (5dpf) analyzed by WISH. (C) The development of motor neurons and oligodendrocytes was not affected in MZ $atrn^{-/-}$. (D) We measured the body weight of three genotype male adult zebrafishes, 12 in each group. The male mutant weigh much more than the wildtype and the heterozygous. (E) Identically, we measured 13 female $atrn^{+/-}$ zebrafishes, 17 female $atrn^{+/-}$ zebrafishes and 15 female $atrn^{-/-}$ zebrafishes, and the results as before in the male zebrafishes. (F-G) The length of different genotype adult zebrafish has no significantly difference. Data are shown as mean \pm SD; *, p < 0.05, **, p < 0.01, ***, p < 0.001.



Supplementary Figure 3. The average velocity of external testosterone treated larvae is faster than the control. Locus diagram of the control group (A), 1 ng/mL testosterone treated group (B), 10 ng/mL testosterone treated group (C) and 100 ng/mL testosterone treated group (D) at 5dpf. (E) The average velocity-time graph of testosterone treated larvae at 5dpf. (F) The average swimming speed of larvae during 5min period at 5dpf. Data are shown as mean \pm SD; ***, p < 0.001; ****, p < 0.001; ns, no significance (p > 0.05).

Table S1 Testosterone levels in men with mental disorders aged 20-49 years.

Gender		Male (20-49)						
	Low T	Medium T	High T	Tota l	χ2, df	p value		
Control	5 (8.3%)	51 (85%)	4 (6.7%)	60				
Epilepsy	3 (14.2%)	17 (81%)	1 (4.8%)	21	0.6798, 2	0.7118		
Anxiety disorders	9 (7.8%)	101 (87.8%)	5 (4.4%)	115	0.4612, 2	0.7941		
Schizophrenia	215 (26.1%)	589 (71.4%)	21 (2.5%)	825	11.87, 2	0.0027* *		
Obsessive- compulsive disorder	2 (6.1%)	31 (93.9%)	0	33	2.539, 2	0.281		
Insomnia	0	12 (100%)	0	12	2.057, 2	0.3575		
Bipolar disorder	44 (22.2%)	141 (71.2%)	13 (6.6%)	198	5.854, 2	0.0536		
Depression disorder	59 (11.4%)	442 (85.3%)	17 (3.3%)	518	2.15, 2	0.3413		
Undiagnosed mental disorder	134 (19.5%)	516 (75.2%)	36 (5.3%)	686	4.616, 2	0.0994		
Mania	14 (12.6%)	90 (81.1%)	7 (6.3%)	111	0.7223, 2	0.6969		

^{*,} *P* < 0.05; **, *P* < 0.01.

Table S2 Testosterone levels in women with mental disorders aged 20-49 years.

Gender		Female (20-49)						
	Low T	Medium T	High T	Tota l	χ2, df	p value		

Control	8 (6.6%)	104 (86%)	9 (7.4%)	121		
Epilepsy	1 (2.9%)	33 (94.3%)	1 (2.9%)	35	1.767, 2	0.4134
Anxiety disorders	16 (8.1%)	159 (80.3%)	23 (11.6%)	198	1.813, 2	0.4039
Schizophrenia	42 (4.0%)	785 (75%)	219 (20.9%)	1046	13.51, 2	0.0012**
Obsessive- compulsive disorder	4 (19.0%)	17 (81%)	0	21	4.889, 2	0.0868
Insomnia	1 (4.0%)	24 (96%)	0	25	2.327, 2	0.3123
Bipolar disorder	16 (3.8%)	314 (74.4%)	92 (21.8%)	422	13.75, 2	0.001**
Depression disorder	96 (10.0%)	754 (78.4%)	112 (11.6%)	962	3.748, 2	0.1535
Undiagnosed mental disorder	56 (5.8%)	704 (73.1%)	203 (21.1%)	963	12.72, 2	0.0017**
Mania	5 (3.8%)	103 (77.4%)	25 (18.8%)	133	7.677, 2	0.0215*

^{*,} *P* < 0.05; **, *P* < 0.01.

Table S3 Testosterone levels in men with mental disorders aged over $50\ years$.

Gender	Male (>50)					
	Low T	Medium T	High T	Tota l	χ2, df	p value
Control	7 (28.0%)	18 (72.0%)	0	25		
Epilepsy	1 (16.7%)	5 (83.3%)	0	6	0.3246, 1	0.5689

Anxiety disorders	2 (1.9%)	97 (90.7%)	8 (7.4%)	107	22.97, 2	<0.0001*** *
Schizophrenia	29 (12.6%)	193 (83.9%)	8 (3.5%)	230	5.04, 2	0.0805
Obsessive- compulsive disorder	0	1 (100%)	0	1	0.3832, 1	0.619
Insomnia	3 (9.1%)	26 (78.8%)	4 (12.1%)	33	6.067, 2	0.0482*
Bipolar disorder	11 (18.6%)	44 (74.6%)	4 (6.8%)	59	2.428, 2	0.297
Depression disorder	20 (5.7%)	303 (87.1%)	25 (7.2%)	348	18.37, 2	0.0001***
Mental disorder	23 (14.0%)	133 (81.1%)	8 (4.9%)	164	4.114, 2	0.1279
Mania	6 (16.7%)	26 (72.2%)	4 (11.1%)	36	3.667, 2	0.1598

^{*,} P < 0.05; ***, P < 0.001; ****, P < 0.0001.

Table S4 Testosterone levels in men with mental disorders aged over 50 years.

Gender	Female (>50)						
	Low T	Medium T	High T	Tota l	χ2, df	p value	
Control	6 (14.6%)	30 (73.2%)	5 (12.2%)	41			
Epilepsy	1 (25%)	3 (75%)	0	4	0.7412, 2	0.6903	
Anxiety disorders	21 (9.6%)	190 (87.2%)	7 (3.2%)	218	7.634, 2	0.022*	
Schizophrenia	33 (8.9%)	305 (82.4%)	32 (8.6%)	370	2.175, 2	0.3371	

Obsessive- compulsive disorder	0	7 (87.5%)	1 (12.5%)	8	1.353, 2	0.5083
Insomnia	6 (12.5%)	42 (87.5%)	0	48	6.49, 2	0.039*
Bipolar disorder	11 (10.7%)	89 (86.4%)	3 (2.9%)	103	5.559, 2	0.0621
Depression disorder	69 (9.8%)	596 (84.7%)	39 (5.5%)	704	4.413, 2	0.1101
Mental disorder	28 (12.2%)	186 (81.2%)	15 (6.6%)	229	1.938, 2	0.3795
Mania	2 (12.5%)	11 (68.8%)	3 (18.8%)	16	0.4209, 2	0.8102

^{*}*P* < 0.05.

 $Table \ S5 \ Testosterone \ levels \ of \ 14 \ ATRN-mutated \ schizophrenia \ patients \ were tested \ at \ different \ time \ points.$

Gende r	Age	Test Date	Testostero ne (ng/ml)	Testosterone (nmol/L)	Chang e
Female	49	2010/8/11	0.1600	0.5024	High
Male	40	2011/5/27	0.42	1.3188	Low
Female	43	2011/10/8	0.28	0.8792	High
Female	36	2013/10/22	0.162	0.50868	High
Female	57	2019/5/16	0.216	0.67824	High
Female	32	2013/10/23	0.389	1.22146	High
Female	34	2018/5/15	0.402	1.26228	High
Male	20	2013/3/6	3.68	11.5552	
Female	36	2014/2/24	0.362	1.13668	
	r Female Female Female Female Female Male	Female 49 Male 40 Female 43 Female 36 Female 57 Female 32 Female 34 Male 20	r Age Test Date Female 49 2010/8/11 Male 40 2011/5/27 Female 43 2011/10/8 Female 36 2013/10/22 Female 57 2019/5/16 Female 32 2013/10/23 Female 34 2018/5/15 Male 20 2013/3/6	r Age Test Date ne (ng/ml) Female 49 2010/8/11 0.1600 Male 40 2011/5/27 0.42 Female 43 2011/10/8 0.28 Female 36 2013/10/22 0.162 Female 57 2019/5/16 0.216 Female 32 2013/10/23 0.389 Female 34 2018/5/15 0.402 Male 20 2013/3/6 3.68	r Age Test Date ne (ng/ml) (nmol/L) Female 49 2010/8/11 0.1600 0.5024 Male 40 2011/5/27 0.42 1.3188 Female 43 2011/10/8 0.28 0.8792 Female 36 2013/10/22 0.162 0.50868 Female 57 2019/5/16 0.216 0.67824 Female 32 2013/10/23 0.389 1.22146 Female 34 2018/5/15 0.402 1.26228 Male 20 2013/3/6 3.68 11.5552

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10	Female	48	2016/10/21	0.239	0.75046	
11	Male	25	2019/1/24	4.4236	13.890104	
12	Male	17	2018/11/6	7.0288	22.070432	
13	Female	42	2011/10/13	0.89	2.7946	High
14	Male	28	2018/6/20	0.257	0.80698	Low

High: Testosterone level is higher than the normal range; Low: Testosterone level is lower than the normal range; --: Testosterone level is within the normal range.