

2021 10 26

2021 12 30

合源® 思源清®

/ ADHD

ADHD

/

12

2200

11 ADHD 1

6 18

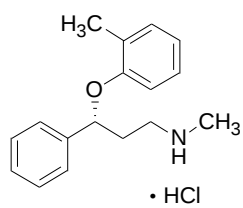


851

Atomoxetine Hydrochloride Capsules

Yansuan Tuomoxiting Jiaonang

- -N- -3- -3- O-



C₁₇H₂₁NO·HCl

291.82

/ ADHD

25mg(C₁₇H₂₁NO)

70 -

0.5mg/kg

3

1.2mg/kg

1.2mg/kg/

1.4mg/kg 100mg

70 -

40mg

3

80mg

2 4

100mg

70kg

100mg

/

ADHD

6-15

1.2-1.8mg/kg/

120mg

150mg

-

HI

ADHD

HI

Child-Pugh ClassB

HI

50%

HI

Child-Pugh ClassC

25%

CYP2D6

CYP2D6

Poor metabolizers,PM

_____ -	CYP2D6			CYP2D6 PM
70				0.5mg/kg/ 4
				1.2mg/kg/
	CYP2D6			70
		40mg/	4	
		80mg/		
	/			
ADHD	DSM-IV	7		-
			2	

I

	2	1
	3	0
	16	4
	3	1
	19	15
c	11	4
	5	2
	2	1

a 2%

2%

c	2	0	1	1

a

b Breslow-Day

c Breslow-Day 0.05 P 0.1

2%

CYP2D6 PM

PM

CYP2D6 EM

PM

11%,EM

6%

PM

7%

EM

4%

PM

7%

EM

4%

1

PM

7%,EM

EM

4%

PM

5%

EM

1%

PM

4%

EM

2%

PM

3%

EM

1%

PM

3%

EM

1%

PM

3%

EM

1%

PM

2%

EM

1%

PM

2%

EM

1%

PM

4%

EM

2%

1

11.3%

61/541

3.0%

12/405

0.9%

N=5

0.1%

N=5

0.6%

N=3

0.6%

N=3

0.4%

N=2

0.4%

N=2

0.4%

N=2

0.4%

N=2

0.4%

N=2

0.4%

N=2

0.1%

1/748

0/43

0.1%

1/705

3

2%

5%

2

3

ADHD

5%

10%

≥20 /

≥15 20mmHg

3:

25

a		
/	N=1697	N=1560

	3	1
	20	5
	26	6
	8	3
b	7	4
	4	2
	4	2
	10	6
	3	0
	2	1
	5	3
	2	1

f	8	1
g	3	2
f / f	4	1
	4	1
	3	0

a 2%

2%

b

c

d

e

f

g

STRATTERA N=943 N=869

STRATTERA N=754 N=691

	2%	CYP2D6		PM		PM			
CYP2D6	EM			PM	4%	EM	1%		PM
35%	EM	17%	PM	11%	EM	7%	PM	5%, EM	2%
	PM	23%	EM	15%	PM	5%	EM	1%	PM
11%		PM	7%, EM	3%		PM	5%	EM	3%
PM	3%	EM	1%	PM	6%, EM	1%		PM	21%,
EM	9%		PM	6%, EM	2%	PM	15%	EM	7%
PM	3%	EM	1%						

_____ -

4

2%

_____-QT

_____-

_____-

_____-

_____-

_____-

_____-

_____-

_____-



ADHD

MAOI

MAOI

MAOI

MAOI

MAOI

15-20mmHg 20 /

/ ADHD

6 18

12 ADHD 11 1 2200

1357

851

0.4% 5/1357

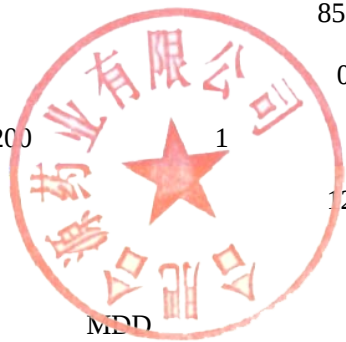
2200

1

12

ADHD

MDD



6000

1

“ ”

120

[20 × ULN]

2 × ULN

40

12

:

	≥15mmHg		≥20mmHg		≥20bpm ^a			
	b				b			
	%	%	%	%	%	%	%	%
DBP ≥15mmHg	21.5	14.1	9.3	4.8	12.6	8.7	4.8	3.5
SBP ≥20mmHg	12.5	8.7	4.9	3.3	12.4	7.8	4.2	3.2
HR ≥20bpm	23.4	11.5	12.2	3.8	22.4	8.3	10.2	2.0

^a :bpm= / :DBP=

HR= mmHg=

SBP=

^b

0% 0/934

0.3% 5/1597

EM

5.0 /

PM

9.4 /

EM/PM

PM

EM

11

/ 7.5 / PM

0.5% 2/402

1.5% 8/540

EM/PM

PM

EM 4.21 2.13mmHg ,

EM

PM:2.75

EM:2.40mmHg PM

0.2% 12/5596

0.8% 46/5596

1.8% 6/340

0.5% 1/207

0.2% 1939

4

1056

ADHD

:

ADHD

ADHD

21/1308 1.6%

9/806 1.1%

1.33[95% 0.67-2.64-

6/1697 0.35%

4/1560 0.26%

1.38[95% 0.39-4.38-

]

ADHD

1.7% 9/540

5.6%

30/540

0% 0/402

0.5% 2/402

2

4

9-12

3

17.9

0.5

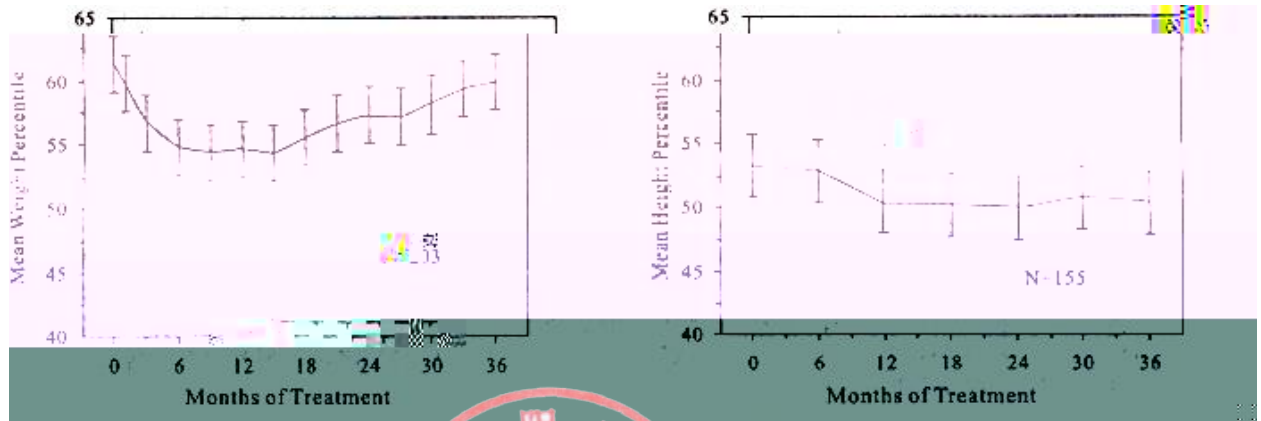
12

3

19.4

0.4

1



1:

3

≤

8

≤9

8

≤13

9

≤14

1.2

13

14

EMs PMs

2

2.4

1.1

0.2

0.4

9

0.4

0.9

1.5

1.1

0.5mg/kg/

1.2mg/kg/

1.8mg/kg/

1.3% 7.1% 19.3% 29.1%

3.5%

CYP2D6

CYP2D6

EM

PM

AUC 10 5 7% PM CYP2D6
PM PM CYP2D6 PM

CYP2D6 CYP2D6 PM
CYP2D6 4- CYP2D6
CYP2D6 PM

- Child-Pugh Class B 2 Child-Pugh Class
C 4 EM AUC

- EM
65%
ADHD
-
-
-
- PM

Tourette ADHD 18
7-17 148 DSM-IV ADHD 80%
Tourette Touretter 116 :29
0.5-1.5mg/kg/ 1.3mg/kg/

YGTSS
148 103 69.6% 76 38 50.0%
72 45 62.5% 12 CGI-S≥4
“ ” CGI-S

- ADHD - ADHD
12 8-17 176 ADHD
DSM-IV 2

		0.8mg/kg/	,	1.2mg/kg/	1.30mg/kg/
± 0.29mg/kg/					PARS
		158	26	16%	
16				18-65	442
ADHD	DSM-IV		23%		2
		40mg/	100mg/	83mg/	± 19.5mg/
			LSAS		
	413	149	36.1%		

_____ -



_____ -

“ ”

_____ -

_____ -

4

_____ -

_____ -

_____ -

_____ -

24

24

_____ -

2-4% 15-20%



6

MAOI

EM CYP2D6

PM

 β_2

60mg	5
------	---

21

80mg.

5

P450

200-800mg

~~P450~~

CYP1A2 CYP3A CYP2D6

CYP3A -

60mg	12
------	----

CYP3A4	5mg	AUC	15%
--------	-----	-----	-----

CYP2D6 -

40 60mg 13

CYP2D6 50mg

18 / 24

,

pH

pH /

1400mg

QT



_____ - 6 18 4
ADHD DSM-IV

/ ADHD ADHD Rating Scale-IV-Parent Version ADHDRS
/

DSM-IV ADHD intent-to-treat ADHDRS

1 8 18 N=297 8

0.5 1.2 1.8mg/kg/

ADHDRS ADHD

1.8mg/kg/ 1.2mg/kg/

0.5mg/kg/

2	6	16	N=171	6	
		1.5mg/kg/ ADHD		1.3mg/kg/	ADHDRS
2	7	13	3 N=147	4 N=144	9
2.0mg/kg/ ADHD				1.6mg/kg/	ADHDRS
	12	12	17		
-		6-15			ADHD
1.2-1.8mg/kg/	10			4	DSM-IV ADHD
				N=292	N=124
		CGI-ADHD-S	≤2		ADHDRS-IV-Parent
25%				8	
N=81	N=82				CGI-ADHD-S
	2	ADHDRS-IV-Parent			90%
2	DSM-IV	ADHD	18		
		ADHD			
	30			CAARS	ADHD
18	ADHD	CAARS	/		
		intent-to-treat			
2	10			5 N=280	6 N=256
					60
120mg/					95mg/

CAARS

ADHD

<42 ≥42

/ ADHD

Ames

CHO

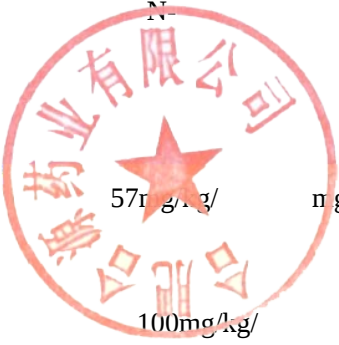
DNA

UDS

CHO

Ames

UDS



57mg/kg/

mg/m²

6

100mg/kg/

3

1

30mg/kg/

mg/m²

100mg/kg

23

AUC

AUC

3.3

0.4

50mg/kg/

mg/m²

6

2

10

2

1

25mg/kg

13mg/kg

2

10

40mg/kg/

mg/m²

5

20mg/kg/

150mg/kg/

mg/m²

17

2 47 458mg/kg/

8 5 mg/m²

AUC AUC 1.8 0.2

39 26 mg/m²

10

1 10 50mg/kg/ mg/m² 0.2 2

8 10 50mg/kg

10 50mg/kg 50mg/kg 50mg/kg

15 10 50mg/kg 50mg/kg 30 50mg/kg

60

CYP2D6 P450 2D6

PM 7% 2% CYP2D6 PM

1% [EM] 10

AUC 5 24 CYP2D6

400 mg/kg

Cmax AUC

EM PM 63% 94%

1-2 Cmax

AUC Cmax 37% Tmax 3

Cmax 9%



0.85L/kg

98%

_____ - CYP2D6 EM

PM EM PM

AUC 10 C_{ss} C_{max} 5 CYP2D6 PM

CYP2D6

CYP2D6 CYP2D6

4-

4-

EM 1% PM 0.1% 4- CYP2D6

PM 4- P450 N-

CYP2C19 P450

EM 5% PM 45%

EM 0.35L/hr/kg 5.2

PM 0.03L/hr/kg 21.6 EM

PM AUC 10 C_{ss} C_{max} 5 EM 4-

N- 6 8 PM N-

34 40

4- -O- 80%

17%

3%

/ 7 / 1 / 14 / 1 / 14 / 2 /

24

YBH12472021

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