

2007 02 08
2008 03 21
2008 09 09
2009 05 12
2011 02 28
2011 05 17
2012 05 21
2014 11 21
2015 11 02
2016 05 11
2017 04 26
2018 05 22
2018 07 11
2019 02 26
2019 05 22
2019 05 28
2020 03 02
2020 05 15
2021 03 16

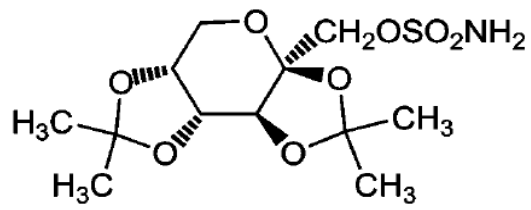
托吡酯片说明书

®

Topiramate Tablets

Tuobizhi Pian

2, 3, 4, 5- - (1-)-β-D-



C₁₂H₂₁NO₈S

339.36

25mg: 101

100mg: 101

2~16

(1) 25mg (2)100mg

25~50mg

1

1 2 25~50mg 100mg / 2
1
200mg 200mg
200~400mg 2 1600mg/

2~16

5~9mg/kg/ 2
25mg 1~3mg/kg/ 1 1 2
1~3mg/kg/ 2
30mg/kg/

2 1/3

17

25mg 1 1 2 25~50mg/
2

100mg/ 500mg/
1000mg/ 。

2~16

0.5~1mg/kg 1 1 2 0.5~1mg/kg/
2

100~400mg/
500mg/

(CL_{CR}<70ml/)

2

4111	3182	929	34	20	2847
<hr/>					
1					1%
	200~400mg/		5%		

/	200~400mg/ (N=354) %	600~1000mg/ (N=437) %	(N=382) %
	5.4	6.2	1.8
	5.1	8.7	3.7
	8.2	19.5	3.1
	4.5	9.4	1.6
	3.1	5.0	0.8
	3.1	11.7	3.4
	3.1	6.4	4.5
	2.8	3.2	1.8
	1.7	2.3	1.3
	1.7	2.1	0.5
	1.7	6.6	2.9
	1.7	3.2	1.0
	1.7	4.6	1.0
	17.8	17.4	8.4
	16.4	34.1	13.6
	8.2	17.2	3.7
	7.1	11.4	4.2
	6.2	11.7	6.8
	5.6		

1

1%

/	200~400mg/ (N=354) %	600~1000mg/ (N=437) %	(N=382) %
	6.8	15.1	8.4
	5.1	14.0	5.2
	3.7	3.9	2.1
	3.7	3.2	1.8
	3.1	3.2	1.3

2

2%

/	(N=104) %	(N=102) %
	19.2	12.7
	5.8	1.0
	8.7	6.9
	5.8	3.9
	2.9	2.0
	2.9	2.0
	15.4	6.9
	13.5	8.8
	10.6	2.0
	5.8	2.0
	4.8	2.9
	3.8	1.0
	4.8	1.0
	5.8	4.9
	6.7	5.9
	16.3	4.9
	11.5	8.8
	4.8	2.0
	9.6	1.0
2~16	5~9 mg/kg/	

3

1%

400mg/

5%

3

1%

/	50mg/ (N=257) %	400mg/ (N=153) %
	0.8	2.0
	3.5	12.4
	2.3	2.6
	4.3	8.5
	3.9	6.5
	2.3	4.6
	3.5	4.6
	0.8	2.6
	0.4	2.0
	1.6	2.0
	18.7	40.5
	1.2	7.2
	2.3	5.9
	4.3	5.2
	1.6	3.3
	1.6	2.6
	0.4	2.0
	1.2	2.0
	0.8	2.0
	0	2.0
	0	1.3
	0.4	1.3
	0	1.3
	0	1.3
	1.6	1.3
	1.2	2.0
	0	1.3
	5.4	6.5
	1.2	3.3

3

1%

/	50mg/ (N=257) %	400mg/ (N=153) %
	0.4	2.6
	0.8	2.6
	1.2	2.0
	0.4	2.0
	0	1.3
	0.4	3.9
	1.6	3.3
	0.4	3.3
	0.4	2.0
	0	1.3
	2.7	3.3
	1.9	2.0
	0.4	1.3
	0	2.6
	0.8	2.0
	0.8	2.0
	0.8	1.3
	15.2	14.4
	3.5	5.9
	3.1	3.3
	7.0	17.0
400mg/		

-

4

10~16

≥2%

400mg/

5%

4

2%

/	50mg/ (N=77) %	400mg/ (N=63) %
	1.3	4.8
	0	4.8
	1.3	4.8
	0	3.2
	3.9	15.9
	3.9	7.9
	0	3.2
	0	3.2
	3.9	9.5
	3.9	4.8
	0	6.3
	0	6.3
	0	4.8
	7.8	20.6
	0	3.2
10	400mg/	

5

1%

5

1%

--

$\geq 1/10$

$\geq 1/100$ $< 1/10$

$\geq 1/1000$ $< 1/100$

$\geq 1/10000$ $< 1/1000$

$< 1/10000$

1.

/

1

40

2.

3.

4.

400mg

6mg/kg/

4mEq/L

10mEq/L

/

/

Z

Z

			400mg/ <20mEq/L	32%	1%
50mg/			<17mEq/L	>5mEq/L	3%
	0%		≥16		50mg/
			14% 400mg/ <17mEq/L	25%	>5mEq/L
50mg/	1%	400mg/	6%		400mg/

Lennox-Gastaut
 2~16
 6mg/kg/ 67% 10%
 <17mEq/L >5mEq/L 11%
 0% 5
 5mg/kg/
 2
 25mg/kg/ -
 -5.9mEq/L <20mEq/L
 0% 5mg/kg/ 30% 15mg/kg/ 50% 25mg/kg/
 45% <17mEq/L ≥20mEq/L >5mEq/L
 0% 5mg/kg/ 4% 15mg/kg/ 5%
 25mg/kg/ 5%
 6~15
 50mg/ 9% 400mg/ 25%
 <17mEq/L
 >5mEq/L 50mg/ 1% 400mg/ 6%

5.

AED

AED

/

11 AED 199

AED

2

1.8 95% CI 1.2 2.7

12 27863

AED

0.43%

16029

0.24%

530

1

4

3 1 /

2 /

3

100~200mg/

200mg/

42% 400mg/

41% 600mg/

52% 800mg/

1000mg/

56%

14%

50mg/

19%

400mg/

26%

/

/

/

200mg/

1000mg/

400mg/

50mg/

400mg/

50mg/

9%

400mg/

15%

14%

/

/

/

50mg/ 400mg/

50mg/ 1 2% 400mg/ 7 12%
/

400mg/

7.

/

8.

9.

2796

10

0.0035

0.0005

0.003

0.005

10.

[

VPA]

/

1~24

8% 5mg/kg/

10%

15mg/kg/

0% 25mg/kg/

9%

≥50%

2

/

2

/

/

/ 1~24

0%	5mg/kg/	12%	15mg/kg/	7%	25mg/kg/	17%		
/						5mg/kg/	0%	
15mg/kg/	7%	25mg/kg/	8%					

11.

32/2086 1.5%

2 4

4/319 1.3%

284 1~24

7%

2

12.

35°C

13.

14.

15.

16.

6%

2%

0.4%

0.1%

3%

1%

2

BUN

17.

/

1.

D

/

AED

AED

<2500

SGA

10

SGA

SGA

3

SGA

2.

/

3.

5

10~20%

1~24

2

Lennox-Gastaut

1~24

20

5 15 25mg/

/ 1~24

/

/

/

/

12%

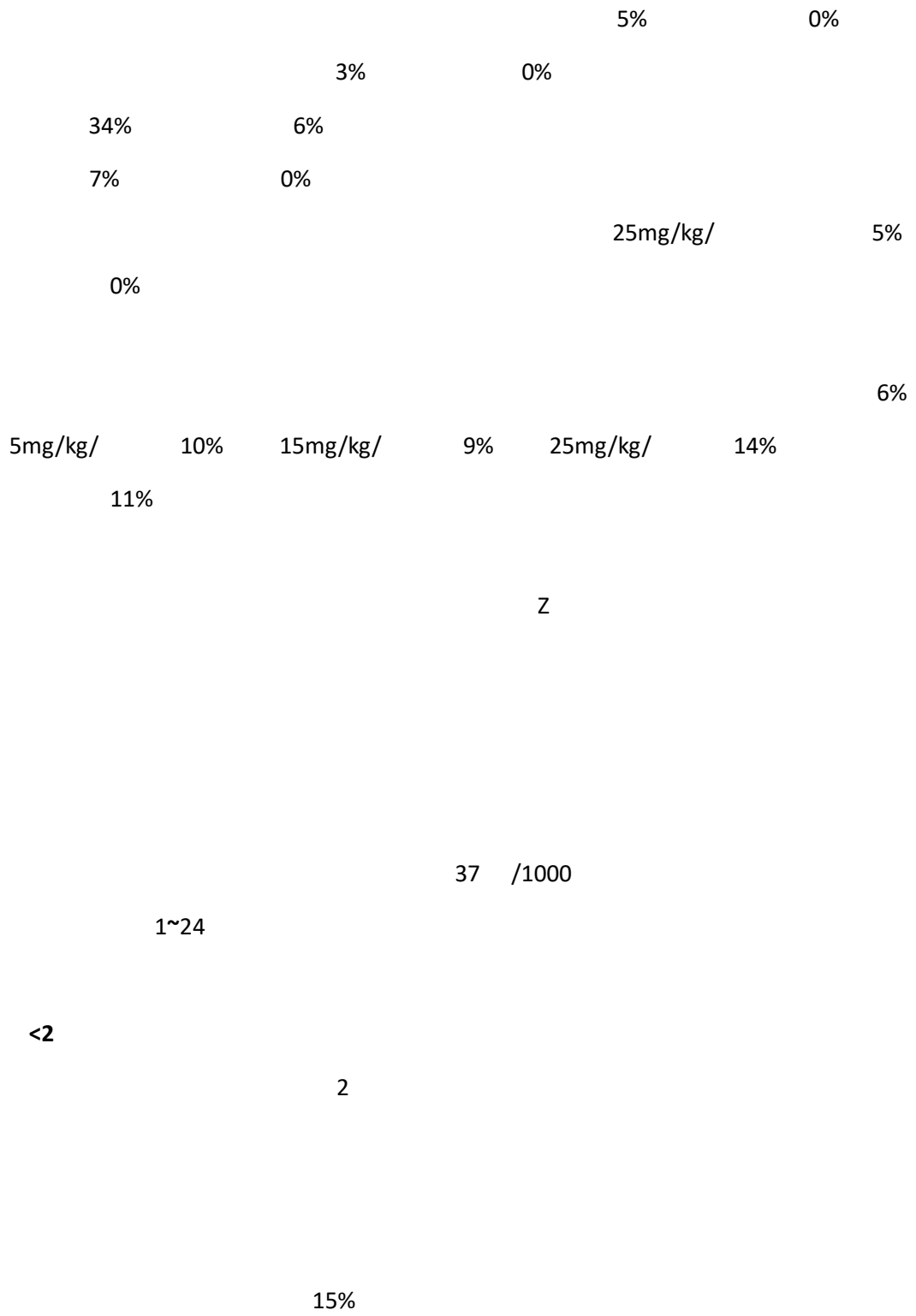
0%

40%

16%

3%

3% 7%



CYP2C19

100mg 400mg
327mg

9

	**	(48%)
CBZ		(40%)
		NS
		NS
**		
NS		

AUC 12%

(1mg /35μg

50~200mg/

AUC

200 400 800mg/

AUC

18% 21% 30%

50~800mg/ AUC 200~800mg/

50~200mg/ AUC

200mg/

AUC 18% 200mg/

600mg/ AUC 26%

100 250 400mg/

1~6mg/ AUC 250mg/ 400mg/

AUC 16% 33% 9-

9- AUC

25mg 2 96mg

C_{max} 27% AUC

29%

C_{max} AUC_{0-12h} 18% 25% CL/F

20% T_{max}

						AUC	15%	C _{max}
							C _{max}	AUC
13%	16%				C _{max}	AUC	60%	
				5mg/		150mg/	2	
						AUC ₂₄	25%	
	4-	-	-	M1	3-	-	M2	13%
15%								

35°C

	K		K
/		PT/INR	K
	INR		

10

	a%	a%
	C _{max} AUC 20%	NS
	AUC 31%	NS
4-	C _{max} 17% 2 50mg	C _{max} 9% 16% AUC 9% 17% 2 40mg 80mg
		NS
	AUC 25% AUC 18% N-	AUC 20%
	AUC 16% (50 mg /12h) ^b	
a%	C _{max} AUC	
NS	C _{max} AUC ≤15%	
b	AUC 14%	

96~110g

20~24

3~4

6
 2~16
 4
 EPMN-106 487 6~83 6
 50mg/ 400mg/ 6
 Kaplan-Meier 400mg/ 50mg/ p=0.0002 Log
 Rank
 p 0.046
 100mg/ Kaplan-Meier 6
 82.9% 71.4%
 p=0.005 75.7% 58.8% p=0.001
 0.516 95% 0.364~0.733)
 AED
 YI 15~63 48
 100mg/ 1000mg/
 54% 17%
 p=0.005

p=0.002

≤0.002

EPMN-104

6~85

252

25 50mg/

200 500mg/

54%

39%

p=0.022

3

p=0.015

EPMN-105

6~84

613

100mg/

200mg/

95%

0

YI EPMN-104

320

32

16

2~5

6

2

4~12

12

12

12

8

8

4

3

5

100mg

100 200mg/

119

25 50mg/

25 50mg/

200mg/

4 8 12

11

2~16

1

2

AED

8

AED

6

AED

25 50mg

25mg 150mg/

125 175 225 400mg/

6mg/kg

8

11 6

b

		a	mg/				
			200	400	600	800	1,000
YD		42	42	40	41		
		5.9	200	390	556		
		6.0	200	400	600		
YE		44			40	45	40
		9.7			544	739	796
		10.0			600	800	1,000
Y1		23		19			
		3.8		395			
		4.0		400			
Y2		30			28		
		5.7			522		
		6.0			600		
Y3		28				25	
		7.9				568	
		8.0				600	
119		90	157				
		8	200				
		8	200				

a				Y1 4 /	YD Y2 6 /
b	Y3 8 /	YE 10 /			

12

50%

12

			mg/					
			200	400	600	800	1,000	6mg/kg/ *
YD		45	45	45	46			
		11.6	27.2 ^a	47.5 ^b	44.7 ^c			
		18	24	44 ^d	46 ^d			
YE		47			48	48	47	
		1.7			40.8 ^c	41.0 ^c	36.0 ^c	
		9			40 ^c	41 ^c	36 ^d	
Y1		24		23				
		1.1		40.7 ^e				
		8		35 ^d				
Y2		30			30			
		-12.2			46.4 ^f			
		10			47 ^c			
Y3		28				28		
		-20.6				24.3 ^c		
		0				43 ^c		
119		91	168					
		20.0	44.2 ^c					
		24	45 ^c					
YP		45						41
		10.5						33.1 ^d
		20						39
		^a p=0.080	^b p 0.010	^c p 0.001				
		^d p 0.050	^e p=0.065	^f p 0.005				
*	YP	<9.3mg/kg/			6mg/kg			
	125 175 225	400mg/	mg/					

100mg¹⁴C-
C_{max} 1.5μg/ml
81%
4μg/ml
100~1200mg
50%
13~17%
0.80~0.55L/kg
100mg 2~3 T_{max}

2

21

100mg

$t_{1/2}$

$30.19 \pm 5.01h$

$31.07 \pm 4.67h$

100~400mg

2

$CL_{cr} < 70ml/$

26%

12

mg/kg

60 / /

36

WS₁- (X-070)-2006Z-2018

25mg: H20020555

100mg: H20020557

19

4 17F

19

710304

4008889988

029 82576616

<http://www.xian-janssen.com.cn>