

2007 02 20
2008 01 14
2009 07 01
2009 08 12
2010 03 25
2011 01 05
2011 01 10
2011 08 23
2011 12 12
2012 08 24
2013 03 14
2014 02 05
2015 01 05
2015 05 07
2017 09 25
2018 04 19
2018 10 25
2019 11 15
2020 10 30
2021 02 10
2021 05 06

~

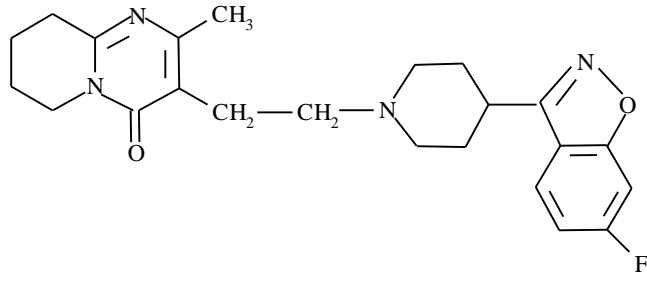
®

Risperidone for Depot Suspension

Zhusheyong Lipeitong Weiqiu

3-[2-[4-(6- -1 2- -3-)-1-]]-6 7 8 9- -2- -4H-
[1 2-] -4-

C₂₃H₂₇FN₄O₂



410.49

7525 DL JN1 - d, 1 - 于 - - 于

40mPa.s

20

1 25mg 2 37.5mg 3 50mg

2

1

1

于

2

于

两 25mg

2 1

37.5mg

50mg

2 50mg

3

4 1

3

两 25mg

2 1

3

1

2

0.5mg

2

2

1mg

2mg

2mg

2

1

25mg

18

12

11% 22/202

13% 13/98

2392

2392

332

12

332

202

25

50mg

12

4

于

两

-

12

2%

ADR

1

1

12

2%

| / | 25 mg (n=99) % | 50 mg (n=103) % | (n=98) % |
|---|----------------------|-----------------------|-------------|
| | 2 | 0 | 1 |
| | | | |

| | | | |
|---|----|----|----|
| | 15 | 21 | 12 |
| * | 8 | 15 | 9 |
| | 7 | 11 | 6 |
| * | 4 | 11 | 6 |
| | 4 | 4 | 0 |
| | 0 | 3 | 0 |
| | 2 | 2 | 3 |
| | 2 | 1 | 0 |
| | 2 | 0 | 0 |
| | | | |
| | 2 | 3 | 0 |
| | | | |
| | 4 | 2 | 3 |
| | 2 | 0 | 0 |
| | | | |
| | 5 | 7 | 1 |
| | 0 | 7 | 1 |
| | 6 | 6 | 0 |
| | 3 | 4 | 5 |
| | 1 | 3 | 0 |
| | 4 | 1 | 0 |
| | | | |
| | 2 | 2 | 0 |
| | 2 | 0 | 0 |
| | | | |
| | 6 | 2 | 1 |
| | | | |
| | 3 | 6 | 0 |
| | 0 | 3 | 0 |
| | 2 | 3 | 1 |
| | 4 | 1 | 0 |
| | 2 | 1 | 0 |
| | | | |
| | 5 | 4 | 2 |
| | 4 | 1 | 1 |
| * | | | |

3 25mg 50mg 75mg 12

 EPS 1 EPS 2

ESRS

1 25mg EPS 50mg EPS

| | | | | | | | | | |
|------|------|------|-----|------|------|------|---------|------|------|
| | | | | ESRS | | | | 50mg | |
| 25mg | | | | 0 | 25mg | -1 | 50mg | 0 | |
| | | | | | | | | 25mg | 50mg |
| | | | | | | | | | 2% |
| | 12 | | | 9% | | 6% | | | 7% |
| | | | | | | | | | |
| | ECG | | | | | | | | |
| | 12 | | | | | 25mg | 50mg | | 202 |
| | | 98 | | | | | QTc | | |
| | | | | | | ECG | QT | QTc | |
| PR | | | | | | | | | 1 |
| | | | | | | | 8~16mg/ | | |
| | | | 4~6 | / | | | | | |
| | | | | | | 0 | ~100 | | |
| | | | | | 16.7 | 12.6 | 25mg | 12.0 | 9.0 |
| | 18.2 | 11.8 | 6 | 10 | | | | 1% | 25mg |
| | | | | | | | | 50mg | 50mg |
| | | | | | | | | | |
| | | | | | | | | | |
| | 2a | | | | | | | | |
| | | | | | | | | | 2% |
| | 2a | | | | | | | | |
| | | | | | | | | | 2% |

| | |
|---|-----|
| / | |
| | * |
| | * * |

2a

2%

/

| | |
|---|---|
| / | |
| | |
| | |
| | |
| | |
| | |
| | * |
| * | 二 |

2b

2%

/

2b:

2%

/

| | |
|---|----|
| / | |
| | |
| | |
| | |
| | |
| | * |
| | |
| | |
| | QT |
| | |
| | - |
| | |
| | |
| | |
| | |
| * | |

2c

25mg

50mg

/

2c

25mg 50mg

/

| | |
|---|--------|
| / | |
| | |
| | |
| | |
| | |
| | |
| | |
| | |
| | * |
| | * 二 |
| | |
| | |
| | |
| | 血 |
| | |
| | |
| | |
| | |
| | * |
| | |
| * | 二 严 |

两 /

3

1/10

1/100 <1/10

1/1000 <1/100

1/10000 <1/1000

<1/10000

事

17

事

事

两 4.0%

3.1%

事

两 86

67~100

事

两 7.3%

89

75~97

3.1%

84

70~96

4.1%

80

67~90

4

2

事

事 义

事

----- (CAE)

85

73~97

事

<1/10000

/

<1×10⁹/L

VTE

VTE

VTE

| | | | | | |
|------------|------------|------------|------------|-----------|------------|
| | TD | | 严 | | TD |
| | TD | | TD | | |
| | | | TD | 两 | 义 |
| | | | | TD | |
| | TD | | | TD | |
| TD | | | TD | | |
| | | | TD | | |
| | | 1 | | 2 | |
| | | | 两 | | |
| | TD | | | | TD |
| | | | | | |
| | EPS | | | | |
| | | | 两 | | |
| | | | | | 义 |
| | | -- | | | |
| 两 | | | | | |
| | | | | | |
| | | NMS | | | |
| | | | - | | NMS |
| | | | | 严 | |
| | | CPK | | 两 | |
| | | | | | |
| NMS | | | | | |
| | | | | | |
| | | | EPS | | |
| | | 严 | | | |
| NMS | | 1 | | | 2 |
| | | 3 | | | NMS |

NMS
NMS

NMS

事

IFIS

-

IFIS

-

-

两 0.8%

12/1499

1

2

两 0.3% 5/1499

事

事

D₂

GnRH

-

义

TTP

1300

1 28

TTP

TTP

义

两

30mL/min/1.73 m²

9-

25 mg

QT

QT

QT

5%

1

Claims

1.02-1.56

=1.26 95% CI

义

/

二

9-

义

9-

18

57

65

12

2

1

QT

QT

QT

P- ^{QT} CYP2D6 CYP3A4 9-
P-gp CYP2D6 / CYP3A4 / P-gp

CYP2D6

CYP2D6

CYP2D6

CYP2D6

CYP3A4 / P-gp

CYP3A4 / P-gp

CYP3A4 / P-gp

CYP3A4 / P-gp

CYP3A4 / P-gp

CYP3A4 / P-gp

- CYP3A4
- CYP3A4 P-gp
- CYP2D6 CYP3A4
- CYP3A4 P-gp
-
-
- CYP3A4 P-gp 两 2~8mg/ 200mg/
70%
- CYP3A4 P-gp 200mg/
9-
-
- CYP2D6 CYP3A4

- CYP2D6 两 CYP3A4

β-

- -
- CYP3A4 P-gp
-
- 事
- H₂- CYP2D6 CYP3A4

SSRI

- CYP2D6
- CYP2D6
20mg/
-
- CYP2D6 CYP3A4
100mg/
100mg/

8

两 20~300mg

事

两

240mg

36mg

QT

QRS

360mg

两

QT

事

QT

QT

/ 于

DSM-IV

两 12

25mg

50mg

两 12

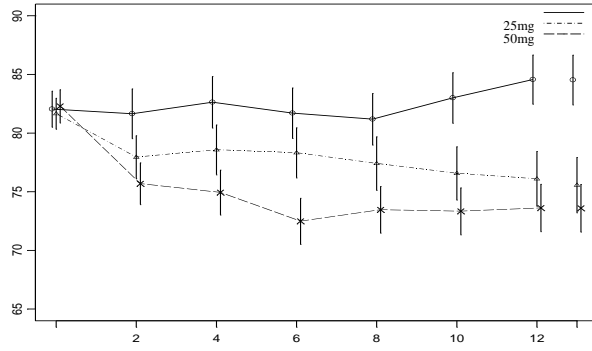
两 50

两 DSM-IV

1

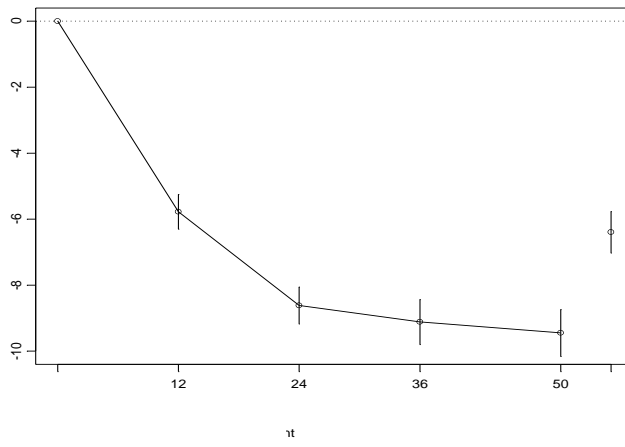
PANSS

LOCF



2 50

PANSS



5HT₂ D₂ 1 2 H₁
 5HT_{1C} 5HT_{1D} 5HT_{1A}

D₁ σ M 1 2

两

D₂ 5HT₂ D₂ 5HT₂

Ames DNA

两 Ames 两

Wistar
 0.16~5mg/kg [mg/m² 两
 10mg/ 0.16~4.8] 于 两
 于 两 MRHD
 两 0.31~5mg/kg mg/m² 两 Beagle
 MRHD
 1.0~16.0

SD Wistar - 两
 0.63 10mg/kg 0.31 5mg/kg mg/m² 两 MRHD 0.64~9.6
 0.64~9.6

两 0.16~5mg/kg mg/m² 两 MRHD 0.16~4.8
 4 事 事
 事 2.5mg/kg
 mg/m² 两 MRHD 2.4 于
 两 于 事 两
 1~4
 5mg/kg mg/m² 两 MRHD 4.8

0.63 2.5 10mg/kg mg/kg 两
 MRHD 10mg/ 3.2 14.4 60.8 mg/m² 两 MRHD
 0.32 1.2 4.8 两 0.64 2.4 9.6 两 18 25

5 40mg/kg [mg/m²
 MRHD 50mg 1 8 AUC MRHD AUC 0.3
 2] 2 1 24

D₂

| | | | | | | | | |
|---------|--------|-------|------------|------|--------------|---------|-----|----------|
| | 5 | 6 | | 24 | | | | |
| | | | | | 6 | 1.5 | | 两 |
| | 40 | | | 0.31 | 1.25 | 5mg/kg/ | | |
| | | 两 | 0.31mg/kg/ | | | | | |
| 12 | | | | | | | | |
| | | 12~50 | | | | | | |
| | | 两 | 0.63mg/kg/ | | 1.25 mg/kg / | | | 两 |
| | | | | | | | | |
| | CYP2D6 | 两 | | | 9- | | | 9- |
| | | | | | 两 N- | | | |
| | | | | | | | | |
| | 两 | 3 | | 严 | 3 | | <1% | 7 |
| | | | 3 | | | | 4~6 | |
| | | | 4~6 | | | 7~8 | | |
| 25~50mg | | 9- | | 严 | | | | |
| | | | | | | | | |
| | | 两 | 1~2L/kg | | | | 1- | |
| | 两 | 90% | | 9- | 两 | 77% | 9- | |
| | | | | | 100mcg/mL | | | 10mcg/mL |

10mcg/mL 9- 10ng/mL
50ng/mL

CYP2D6 两 9- 产
N- 产 9-

70% 14% 1mg ¹⁴C- 1 两 84% 3
70% 14%

两 3~6 3~6
CYP2D6 9-

两 5.0L/h 13.7L/h 两 3.2L/h 3.3L/h
25mg 50mg

9.9~19.2ng/ml 17.9~45.5ng/ml 2 25 50mg
12 两 7~8

30% 60%
35%

III

PANSS

ESRS

2~8°C

1

- 1

-

1

- 1

2

1

2

36

25°C

24

25°C

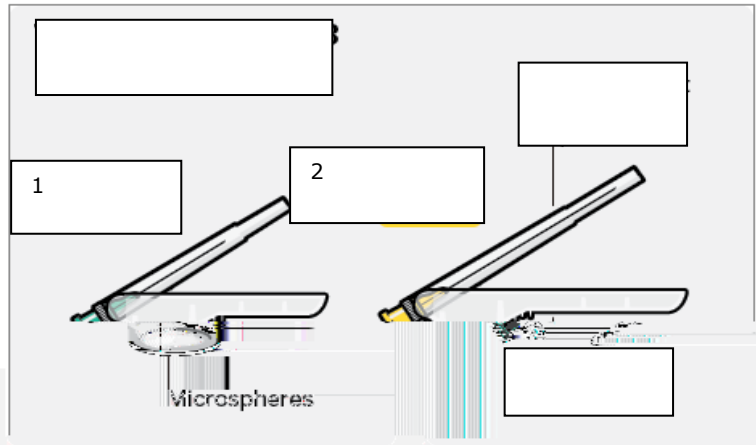
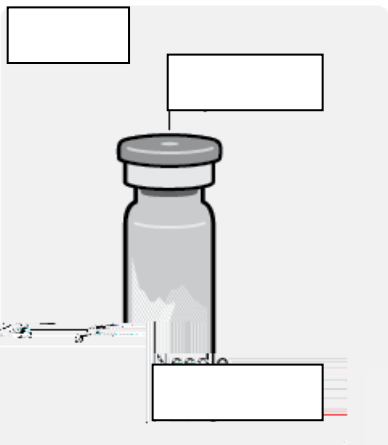
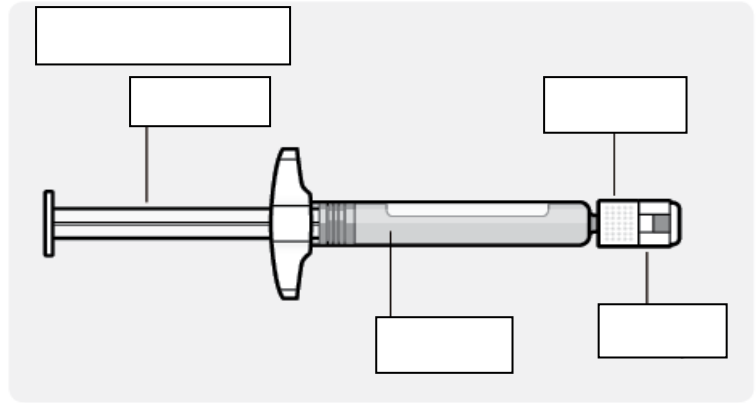
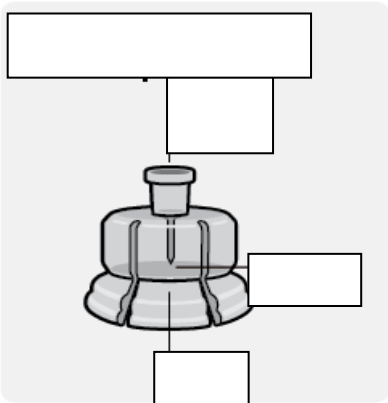
6

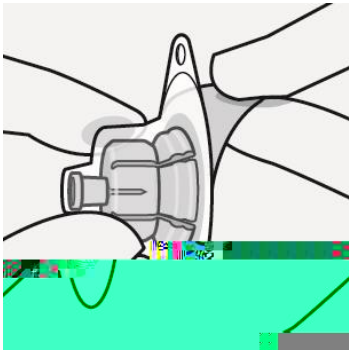
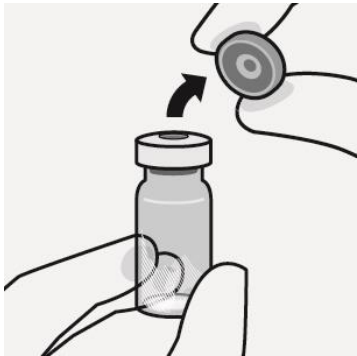
JX20190185

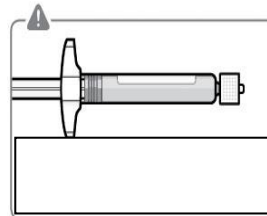
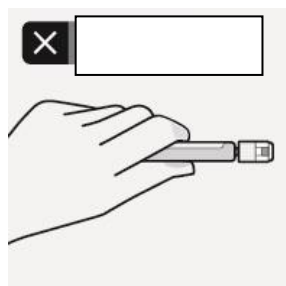
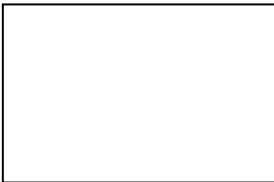
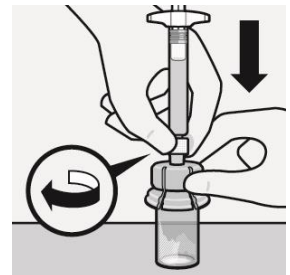
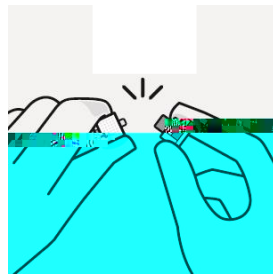
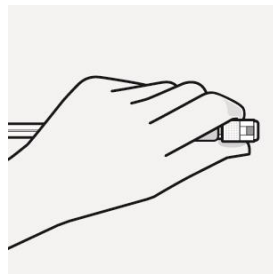
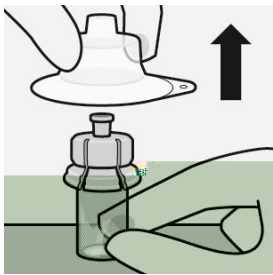
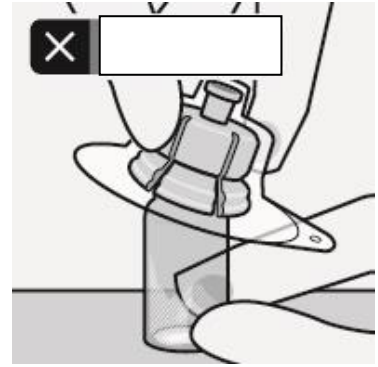
-
-
-
-
-
-

两 ®

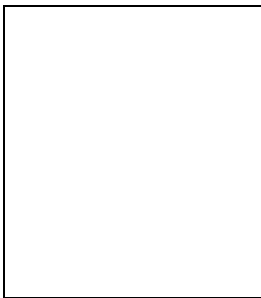
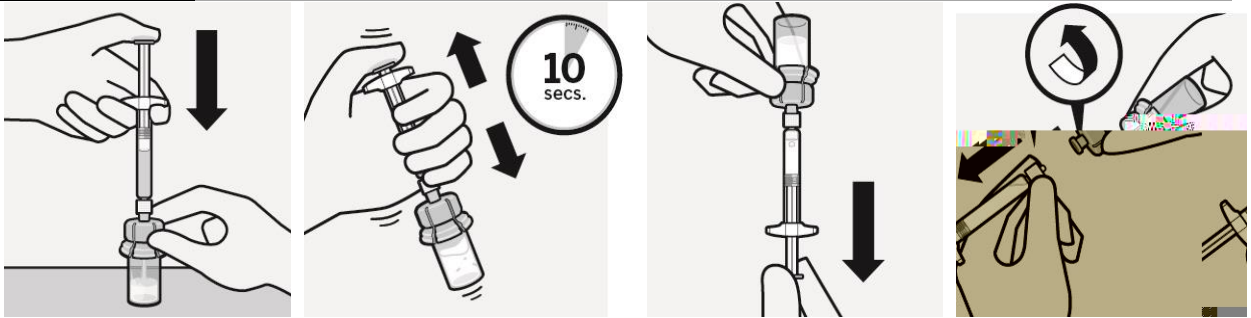
两







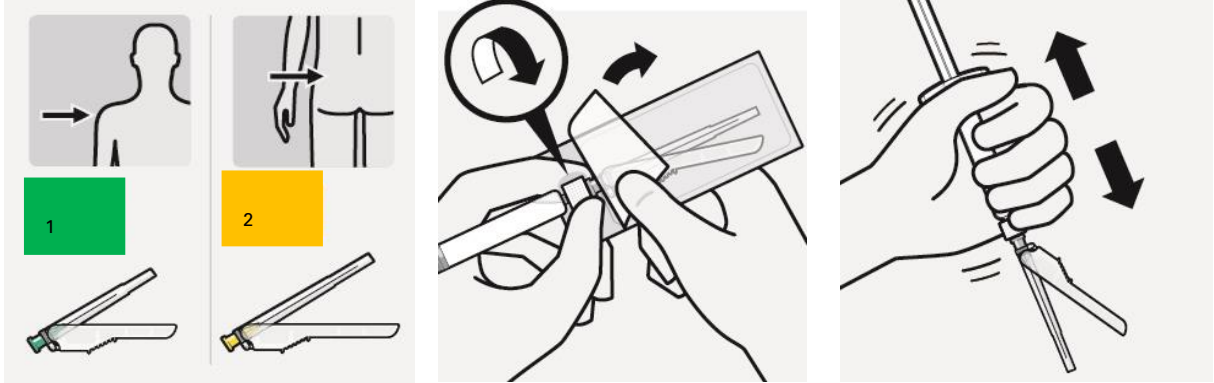
2



10

两

3

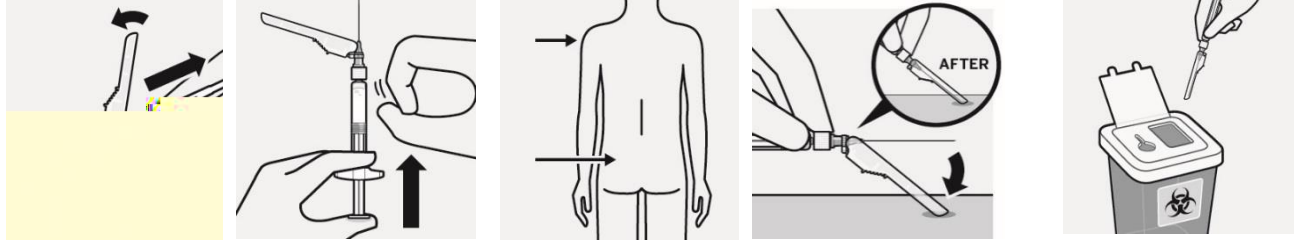


1

2



4



IM

45°

1/4

义