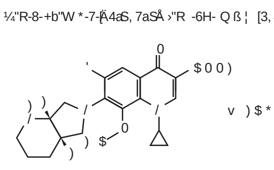
Moxifloxacin Hydrochloride and Sodium Chloride Injection

Yansuan Moxishaxing Lühuana Zhusheye

[illegible]



$C_{21}H_{24}FN_3O_4\cdot HCl$

437.9

【性状】 [illegible]

【适应症】 [illegible]

【规格】 250ml: [illegible] $C_{21}H_{24}FN_3O_4$ [illegible] 2.0g

【用法用量】 [illegible]

	0.4g	
U W4ö9ü WUk0Ö&¾	0.4g	10
' W _"D1Ñ&¾ U W Œ	0.4g	5
/n j9ç Ç W6ê&¾	0.4g	7-14
MŽ = r W,Þ6Ô ¼,Þ6Ô4ô4÷ O	0.4g	7
= r W,Þ6Ô ¼,Þ6Ô4ô4÷ O	0.4g	7-21
= r W7©7„ µ O	0.4g	5-14
UP+Û	0.4g	10-14

2、老年患者 [illegible]

3、肝功能损害患者 [illegible] Child Pugh A、B [illegible]

[illegible] 30ml/min/1.73m² [illegible]

4、给药说明 [illegible]

5、药物相容性 [illegible] 1:10 [illegible]

0.9%"_ FJÐ# 4#â — 9ü# 4+X"d

1M "_ FJÐ# 4#â — 10%:':43 # 4#â

5%:':43 # 4#â — £G¨Ç l# 4#â

6、种族差异 [illegible]

【不良反应】

1、严重和其他重要的不良反应 [illegible]

2、临床试验的经验 [illegible] 14981 [illegible]

表2：盐酸莫西沙星（口服与静脉）与对照药物的临床试验中常见（≥1%）不良反应

		% (N=14981)
[illegible]	C[>p	1
[illegible]	¦ó	7
	7©#+	6
	… @	
	ï0	
	7©G +ì,	
	#¸ F =8Ÿ	1
[illegible]	'	1
42« ð	Bg lEœ"XG¦ wQ	1
BR ¼:U «2«+î+õ	~Jî>p+÷	1
/Ž4ÿ3+5 +î+õ	d,	4
	d …	3
2î/Ž+î+õ	a-P	

表3：盐酸莫西沙星（口服与静脉）与对照药物的临床试验中不常见（0.1%~<1%）不良反应 (N=14981)

[illegible]

3、实验室检查参数变化 [illegible]

4、上市后的经验 [illegible]

表4：上市后报告的药物不良反应

[illegible]

【禁忌】 [illegible]

【注意事项】

1、肌腱炎和肌腱断裂、周围神经病变和中枢神经系统的影响 [illegible]

2、肌腱炎和肌腱断裂 [illegible]

3、重症肌无力加重 [illegible]

4、QT间期延长 [illegible]

5、过敏反应 [illegible]

6、其他严重且有时致命的反应 [illegible]

7、中枢神经系统的不良反应 [illegible]

8、周围神经病变 [illegible]

9、精神病的不良反应 [illegible]

10、艰难梭菌相关性腹泻 [illegible]

11、动物的关节病 [illegible]

12、对血糖的干扰 [illegible]

13、光敏感性/光毒性 [illegible]

14、细菌耐药性的发生 [illegible]

15、对驾驶或操作机械能力的影响 [illegible]

16、主动脉瘤和主动脉夹层的风险 [illegible]

【孕妇及哺乳期妇女用药】 [illegible]

【儿童用药】 [illegible]

【老年用药】 [illegible]

【药物相互作用】

1、抗酸药、硫糖铝、复合维生素和其他含有多价阳离子的产品 [illegible]

2、华法林 [illegible]

3、抗糖尿病药物 [illegible]

4、非甾体类抗炎药物（NSAIDs） [illegible]

5、延长QT间期的药物 [illegible]

【药物过量】 [illegible]

【临床试验】

1、慢性支气管炎急性发作 [illegible] 89%(222/250) [illegible] 89%(224/251)

表5：按病原菌分类的临床可评价患者在随访时的临床治愈率（慢性支气管炎急性发作）

病原菌	盐酸莫西沙星(n)	克拉霉素(n)
6ê&¾K.*39ü	16/16(100%)	20/23(87%)
#q O >p v9ü	33/37(89%)	36/41(88%)
Ÿ#q O >p v9ü	16/16(100%)	14/14(100%)
' 9Û ù9ü	29/34(85%)	24/24(100%)
H Tô8¢:':4*39ü	15/16(94%)	6/8 (75%)
6ê&¾ {M' _9ü	18/20(90%)	10/11(91%)

[illegible]

2、社区获得性肺炎 [illegible] 95%(184/194) [illegible] 95%(178/188) [illegible] 86%(157/182) [illegible] 89%(161/180) [illegible] 93%(241/258) [illegible] (2.9% 13.2%) [illegible] 84%(216/258) [illegible] (74%, 208/280) [illegible] 95% [illegible] (2.6% 16.3%) [illegible]

表6：按病原菌分类的临床治愈率（社区获得性肺炎，汇总数据）

+õ Ï9ü	- G¨9Û?¯"É O	
6ê&¾K.*39ü	80/85	(94%)
H Tô8¢:':4*39ü	17/20	(85%)
6ê&¾ {M' _9ü	11/12	(92%)
#q O >p v9ü	56/61	(92%)
6ê&¾>" Ï ƒ	119/128	(93%)
6ê&¾ _ Ï ƒ	73/76	(96%)
' 9Û ù9ü	11/12	(92%)

3、由耐多药肺炎链球菌（MDRSP）引起的社区获得性肺炎 [illegible] MDRSP [illegible] 37 [illegible] MDRSP [illegible] 35 [illegible] 95% [illegible]

[illegible] MDRSP [illegible] MIC 1 2 mcg/ml [illegible]

表7：采用盐酸莫西沙星治疗MDRSP CAP患者的临床治愈率和细菌学治愈率（按耐药表型分析）

敏感性筛选	临床治愈率		细菌学治愈率	
	n/N[a]	%	n/N[b]	%
6@M,M93P	21/21	100%c	21/21	100%c
6@1\ ¼ d '9ü3P	25/26	96%c	25/26	96%c
6@ W)ß µ72	22/23	96%	22/23	96%
6@+b"W8ô ¦ /*7*+b q A	28/30	93%	28/30	93%
6@)ß3P	17/18	94%	17/18	94%

a) n= [illegible] N= [illegible] MDRSP [illegible] 37 [illegible]

b) n= [illegible] N= [illegible] MDRSP [illegible] 37 [illegible]

c) [illegible]

d) [illegible]

[illegible]

表8：耐多药肺炎链球菌所致社区获得性肺炎的临床治愈率和细菌清除率

耐多药肺炎链球菌	临床治愈率	细菌清除率
6 @ /ý Ç9ü9Ÿ	12/13(92.3%)	12/13(92.3%)
6 @ /ý Ç9ü9Ÿ	10/11(90.9%)a	10/11(90.9%)a
6@4 /ý Ç9ü9Ÿ	6/6 (100%)	6/6 (100%)
6 @ /ý Ç9ü9Ÿ	7/7 (100%)a	7/7 (100%)a
d 96@ J9Ÿ6ê&¾K.*39ü,´9ü>p+÷	9/9 (100%)	9/9 (100%)

a) [illegible]

4、急性细菌性鼻窦炎 [illegible] 400mg [illegible] 250mg [illegible] 90% [illegible] 89% [illegible] 97% (29/30) [illegible] 83% (15/18) [illegible] 80% (24/30) [illegible]

5 5 .复杂性皮肤和皮肤组织感染

X5¾ -F >|,´ 0N©L¿ j Ã ü-" Ã)'—,´ d ºB P¼ È)- G¨9Û?¯"É O Ä = È 400mg!ÿ 0!Q È F 5 5 9Ÿ 7 Y Å> d '"X8ô- G¨- Ä = È 500mg!ÿ 9!Q È F 5 5 9Ÿ 7 Y Å,+Ç xF >| ¶" E³ Ä B .D0¦] È9 30%,´ Ó65"ë+ÇMŽ = r W7C6ï Ã8%"ë+Ç+ÆEÂ%"ë+Ç=2 0Í4÷&¾ Å0%"ë+Ç7C+á+õZ6%"ë+Ç ¦ ,Þ6Ô O Ä - G¨9Û?¯"É O4ô ¼)'—4ô] 6 [9 17% ¼14%,´ Ó65 í>|Eµ Ù0; ¿ Ä 7 0 ¼ E #q F$5 KÃG¨9Û?¯"É O4ô ¼ d '"X8ô- G¨- 4ô] È Aô ' Ó65,´ d º"ë 8)· 6 [j 89%(108/122) ¼91%(110/121)Ä

6 6 .复杂性皮肤和皮肤组织感染

= r W,Þ6Ô ¼,Þ6Ô5 ´ O Ä cSSSI Å >| TN©L¿ j Ã #k W)'—B P¼ ÃN© j ü-"B P¼ È k?± X G5¾ ` jF >| È)- G¨9Û?¯"É O IV/PO ¿C_5 9ŸÄ400mg!ÿ 0!Q È F 5 5 9Ÿ 7-14 Y Å> IV/PO)'— ñ β- µG 7* /β- µG 7*G¦ Á f r+X ¾"ë+Ç cSSSI Ó65,´+Ç xF >| ¶" E³ Ä B .D0¦ ¡4ã • 617 = Ó65 È¦] 335 = Ó65F2 8F >|+Ç x 6 À Ä 0N© j -Lu 0 n.D0¦ È)- G¨9Û?¯"É OÄ 400mg!ÿ 0!Q ÈF 5 5 9Ÿ 7-21 Y Å >)'— ñ β- µG 7* /β- µG 7*G¦ Á f r IV/PO ¿C_5 9Ÿ+X ¾"ë+ÇcSSSI Ó65,´ õ åF >| ¶" E³ Ä B .D0¦ ¡4ã • 804 = Ó65 È¦] 632 = Ó65F2 8F >|+Ç x 6 À Ä F Ë.D0¦] È - G¨9Û?¯"É O4ô ¼)'—4ô] 6 [9 55% ¼53%,´ Ó65F >| { _ 7 0 Ã E#q F$5 K ÈB Eµ Ù ý Œ ´ @ \F2 Ä+÷"ë+Ç p õM0,´ 0G 6 Ä "ë 8)·L¿Aú Ý2« »6< F È þ O W$³+Ñ Ó65,´ 61% ` ¦ ì " Ó65,´ 90% Ä Ë"ë 8)· > )'—4ô?ò `,´"ë 8)·-(l Ä Aô ' Ó65,´ k ƒ"ë 8)· ú 9+õ Ï9ü 62«,´ d º"ë 8)· ²> ˜ 表 9 ¼表 10 p/j Ä

表 9：复杂性皮肤和皮肤组织感染患者的临床治愈率

研究	盐酸莫西沙星 n/N(%)	对照药 n/N(%)	95% 置信区间 *
G5¾	125/162(77.2%)	141/173(815%)	(-14.4%，2%)
-Lu	254/315(80.6%)	268/317(845%)	(-9.4%，2.2%)

9Û?¯"É O >)'—9Ÿ {L$,´"ë 8)· 2 Ä9Û?¯"É O ")'—9Ÿ Å

表 10：复杂性皮肤和皮肤组织感染患者按病原菌分类的临床治愈率

病原菌	盐酸莫西沙星 n/N(%)	对照药 n/N(%)
H Tô8¢:':4*39üÄ)+b"W?¯ Ç • O,´ 6/ë Z Å a	106/129(822%)	120/137(876%)
W6Ð v9ü	31/38(816%)	28/33(84.8%)
6ê&¾ {M' _9ü	11/12(917%)	7/10(70%)
Ld"Ï6Ð v9ü	9/11(81.8%)	4/7 (57.1%)

D +b"W?¯ Ç • O W X G5¾.D0¦]#{ Ê

7.复杂性腹腔内感染

= r W7©7„ µ O F >| ¶ TN©L¿ j Ã #k W)'—B P¼Ã0N© j ü-"B P¼ È k?± X G5¾ ` jF >| È)- G¨9Û?¯"É O IV/PO ¿C_5 9Ÿ Ä400mg!ÿ 0!Q ÈF 5 5 9Ÿ 5-14 Y Å > ù ù?¯ Ç A $ –M‰795 9Ÿ > =Lo9Û?¯ Ç { ù5$G¨+X ¾"ë+Ç = r W7©7„ µ O ÄclAI ÅÓ65Ä 5 7©7Ì&¾ Ã7C6ï Ãd0¯„'LA n&¾ ú6Ð0¯„ Å ,´+Ç xF >| ¶" E³ Ä B .D0¦ ¡4ã • 681 = Ó65 È¦] 379 = Ó65>Û?ö j d º Aô ' Ó65 Ä 0N© j -Lu 0 n.D0¦ È)- G¨9Û?¯"É OÄ 400mg!ÿ 0!Q È F 5 5 9Ÿ 5-14 Y Å> d ' " ®M‰795 9 Ÿ +b.• AM‰795 9Ÿ > =Lo9Û?¯ Ç / { ù5$G¨+X ¾"ë+ÇcIAI Ó65,´ õ åF >| ¶" E³ Ä B .D0¦ ¡4ã • 595 = Ó65 È¦] 511 = Ó65>Û?ö j d º Aô ' Ó65 Ä d º Aô ',´ ê5Ô+a4ÿ { _žAú +O = r W O Ã8# A5 9Ÿ 5 Y ú"ë 8 ðP¼Aï?ö & Ó65F >|1\ 25-50 YL¿AïAô ` ,´ B 65 Ä d º Aô ' B 65,´ k ƒ d º"ë 8)· ²> ˜ 表 11 p/j Ä

表 11：复杂性腹腔内感染患者的临床治愈率

研究	盐酸莫西沙星 n(N%)	对照药 n(N%)	95% 置信区间 *
G5¾ kAÑ Å	146/183(79.8%)	153/196(781%)	(-7.4%，9.3%)
7C6ï	40/57(70.2%)	49/63(778%)a	NAb
MŽ7C6ï	106/126(84.1%)	104/133(782%)	NA
-LuÄ kAÑ Å	199/246(80.9%)	218/265(823%)	(-8.9 %，4.2%)
7C6ï	73/93(78.5%)	86/99(869%)	NA
MŽ7C6ï	126/153(82.4%)	132/166(795%)	NA

* - G¨9Û?¯"É O >)'—9Ÿ {L$,´"ë 8)· 2 Ä- G¨9Û?¯"É O ")'—9Ÿ Å

D ÂL" T = X 0 M 48 ? & µM0 >| { _,´ Ó65

b) NA– =F2+X

8.微生物学

+a ¾ V*6 – ¼ >| W éM',´ Ï È # X6êUP+Û Ó65] 0 …- G¨9Û?¯"É O,´ 9 x W.D0¦ Ä!" È F ZF2 Ä+÷,´ © ö _ * ¾ Ø (™ j Q » 0 …,´ 0N© 9 x W.D0¦ ú @ ê > Ø(™,´ _ 1 W9Ÿ Ø Ë – ž Ä

Ó 96êUP+Û,´MŽ#b5/)d j Ø(™ Q » È 0 … ¶ 0N©L¿ j Ã-"# Ã ¹ r)'—.D0¦ Ä 20 MŽ#b5/)dÄ 10 Lô)d ¼ 1 Lü)d Åh • £ w rGÿÄ ± SD Å j 100±50 LD50Ä93 $ j 92-127 LD50 ÅUP+Û6f D "? v9üÄ CO929ü Z ÅDM. r ÄB .D0¦] È 9Û?¯"É O)UP+Û6f D "? v9ü9ü Z,´ 0 ? Á9ü#ƒ ÖÄ MIC j 0.06 mcg/mL Ä15 '8# A 4 Z ? & È 6 9'— ê ƒ5 9Ÿ é x5 ¸9Û?¯"É O F ¹ r ÈF >| 10 Y"ë+Ç ÄX 0 û.D0¦5 9Ÿ } È p 9.D0¦ Ø(™ w ' È +O9ü>p+÷ ÄUP+Û6f D "? v9üÃë+Ç > 83-139 ? & OL$Ä £ w 115±19 ? & ÅÈ Õ ¹ r,´ 100% Ø(™!« ¾+î+õ ÄX"ë+Ç5 • >,´ 3 Y µ È 9Û?¯"É O"ë+Ç4ô,´10/10 100% Ø(™ ý'f ^#k Å> ¹ r4ô-(" È 9Û?¯"É O4ô,´!« Ñ)· n:‡E³ ~ Ä+O ˆ)· 2 Ö 100%Ëü × ðP¼Ä2î.žšž jL$ j [66.3% 100%] p<0.0001ÄÄ

X6êUP+Û,´MŽ#b5/)d Q »] È> ¹ r4ô-(" È 9Û?¯"É O4ô Ø(™+O ˆ)· i ´ § 95 AÑ – n:‡ W È 9Û?¯"É O,´ £ w>p#v#ƒ ÖEî ` FCµE÷ ¶ = ¼M‰795 9Ÿ é x,´ @ ê,´9Ÿ(™>p#v#ƒ Ö ÅM‰79%$400mg 9Û?¯"É O,´ @ ê £ w Ä ±SD Å #ƒ ÖÄ CmaxÅ ¼ k äMbGÿ ÄAUC Å6 [j 3.9±0.9 mcg/mL ¼39.3±8.6 mcg•h/mLÄG÷+X ê ƒ5 9Ÿ é x Q ê ƒ5 9Ÿ Ä rGÿ j 400mg ÅMŽ#b5/)d È X5 9Ÿ 0 Y > 9Û?¯"É O,´ £ wÅ ±SD Å#ƒ Ö ¼ AUC_{0-24} 6 [j 4.4±1.5 mcg/mL ¼22±8.0 mcg·h/mLÄ

Ï 药理毒理 Ð

药理作用

(1) 作用机制

9Û?¯"É O _ § 9 ¯Ba#k W ¼ p9ü Œ+X,´ 8- +b"W *"O éB*Gž2« Ç9ü9ŸÄ9Û?¯"É O X ƒ F n/j *)M™ Lc W9ü Ã M™ Ld W9ü Ã¼ "W9ü ÃG¨9ü ¼MŽ ¨ » Þ+O(™ ² _ Ï ƒ Ã>" Ï ƒ ¼ Ë 9ü § 9 ¯Ba Ç9ü#k W Ä

p9ü Œ+X j f j ¢ • 2 ´G¦ û ¼ý Ä • 2 ´G¦ _ × f DNA • ¼ X DNA = f Ã = ¼Eœ …] £K^,´G¦ Ä

9Û?¯"É O>˜)à j#ƒ Ö ÍC† W,´ p9ü#k W Ä 0 ~ p9ü#ƒ Ö ¼ 0 ~ Á9ü#ƒ Ö * \ 08$ Ä

9Û?¯"É O)β- µG 7*2« ¼ W)ß µGŸ2«6@9Ÿ,´4ö9ü Ö 9 xFÄIE÷ O ,´ ÎP¼ Ø(™ Q »Añ Î È 9Û?¯"É O ƒ µ#k WQ Ä

(2) 耐药性

,8$)M,M93P2« Ãd '9ü3P2« Ã"X *3 9'2« Ã W)ß µGŸ2« ¼)ß3P2«6@9Ÿ,´6@9Ÿ j f = ¡ ý9Û?¯"É O,´ Ç9ü#k W Ä9Û?¯"É O ¼ F Ë Ç9ü9ŸL$ Ô ù6@9Ÿ W8# ú Z)àCX2Â û „´6@9Ÿ W *)à Ä

9Û?¯"É O,´ 8- +b"W *G 6 > 8- "RG 6-(" § 9)M™ Lc W9üQ #k W ¼6@9Ÿ0± ,´ ~F9 W Ä Î W,´ 7 } ¼"^ r)ß *7- Lk!' k Ø F Â È B k Ø F Â j"O éB*Gž2«9Ÿ(™,´6@9Ÿ j f Ä

ƒ FB P¼ n/j4ÿE÷ J!• 2 }7-5C ' ` *)à )9Û?¯"É O,´6@9Ÿ W Ä k { ¦6@9Ÿ)· ¸ ~ (10-7-10-10)Ä¿ G ` 64ö9ü äMb X ~ ¾ 9Û?¯"É OMIC #ƒ Ö & 7- ¯ MIC l 9 AGÿ,´ Î Ð Ä

> ¦ éB*Gž2«9Ÿ(™ {L$ ˆ X Ô ù6@9Ÿ Ä v _ È 0 Ë) ¦ éB*Gž2«9Ÿ(™6@9Ÿ,´M™ Lc W9ü ¼ ¼"W9ü)9Û?¯"É O • O Ä

(3) 对其它类药物的协同作用

X TN© o65 =- G¨9Û?¯"É O,´.D0¦]?ò ` ; G F Ö W6Ð ó <9ü Ã 8í ' v9ü Ž ÃžFJ v9ü Ã 6Ð*39ü ÃM' _"?9ü Ž ¼ ¼"W9ü ² ü!— v9ü Ã -O v9ü ¼#¸ FK.*39ü1y Å´ ÿ A È F Ë F X T ˜ µ ' =!" h È Z)à8 Lî Ý9ü" 3P Ä

(4) 体外试验敏感数据

• O	] Ö • O	6@9Ÿ
M™ Lc W9ü		
LdFƒ Ð ç9ü		
6ê&¾K.*39ü È5 JGý6@9Ÿ6ê&¾K.*39ü9ü[ØDRSP] È5 ". ,´M,M93P6@9Ÿ9ü Z ÄPRSPÅú) ;F Ç+O3P] T/ý F $ J Ç+O3P6@9Ÿ9ü Z ÖM,M93P (MIC 12µg/mL Ã 1\ ¼ d '9ü3PÄ ² d ' {EË ÅÃW)ß µGŸ2« Ã)ß3P2« Ãb"W8ô ¦ / /*7*+b q A Ä		
A 4ô F7CK.*39ü*		
2£"?K.*39ü		
E« »K.*39ü		
£K.*39ü		
Œ £K.*39ü		
í &¾K.*39ü *		
O ×K.*39ü *		
H Tô8¢:':4*39üÄ 5 +b"W?¯ Ç • O9ü Z Å*	H Tô8¢:':4*39ü Ä+b"W?¯ Ç/ "W"O "É O6@9Ÿ9ü Z Å	
0 "?:':4*39ü		
>˜,Þ:':4*39ü Ä 5 +b"W?¯ Ç • O9ü Z Å	>˜,Þ:':4*39üÄ+b "W?¯ Ç "W"O"É O6@9Ÿ9ü Z Å	
$æ>p:':4*39ü		
ê:':4*39ü		
7€+O:':4*39ü		
Q /:':4*39ü		
,- ¹ (æ v9ü		
2Ú6Ð*39ü(õ) 7 M93P Ã ¶ WM93P • O9ü Z)		

*/** "4ÿAñ Î7-) • O9ü ,8$,´ " © ö,´ d ºF2 Ä+÷ 9 x

9Û?¯"É O õ)+a MecA * û „´+b"W?¯ Ç6@9Ÿ:':4*39ü,´ ƒ F MIC l X ¦ • O93 $ µ Ä !" È ² Ì)àF 2«9ü Z = Ø 9€¯+X9Û?¯"É O Ä

• O	] Ö • O	6@9Ÿ
M™ Ld W9ü		
#q O >p v9ü (5 ×+O ¼ = ×+O £- µG 7*G¦,´9ü Z)*		
Ÿ#q O >p v9ü *		
' 9Û ù9 ü 5 ×+O ¼ = ×+O £- µG 7*G¦,´9ü Z)*		
,® ã Š ç(©9ü		
W6Ð ó <9ü*		
6ê&¾ {M' _9ü*		
×G¨ {M' _9ü		
×"D6Ð v9ü		
@ 6Ð v9ü		
Ld"Ï6Ð v9ü *		
]L$6Ð v9ü		
LZ >6Ð v9ü		
	K 5/ w …7 9ü	
	9— y w …7 9ü	
	#;:¡ _ {M= D ç9ü	
	TÖ8í « …7 9ü	
	w 2 ' v9ü *	
žFJ ' v9ü		
™"? ™ ï9ü		
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) ¾ Ë 4ö9ü,´9ç Ç W6@9Ÿ)· 7- L¿ `*6 ¼ &L$6< i Ä pG 6@9Ÿ _ 7- *)à ,´ È(© [_"ë +Ç UGÿO & ÄF ƒ F • OB P¼5 Ì +X • 7 , T Ý Þ+O(™ _ V)9Û?¯"É O • O Ä

… rGÿM‰79 ¼ =5 ¸ 0.4g- G¨9Û?¯"É O,´ PK/PD' E³ Ä

M0 •L' Ó65,´ AUC/MIC90 l W ¾ 5 D C_{max}/MIC_{90} j 8-10 j d º"ë 8,´ O K l Ä L Aú Ó65,´ ò lFJ hE³ ~ È AUC/MIC_{90} W ¾30-40Ä

;>˜ j … rGÿM‰79 ¼ =5 ¸ 0.4g- G¨9Û?¯"É O,´ PK/PD,´AÑ1Ç l" E³ Ö

5 9Ÿ é ?	M‰79		=	
ò] })	AUIC[h]	C_{max}/MIC_{90} a)	AUIC[h]	C_{max}/MIC_{90}
MIC_{90} 0.125 mg/L	313	32.5	279	23.6
MIC_{90} 0.25 mg/L	156	16.2	140	11.8
MIC_{90} 0.5 mg/L	78	8.1	70	5.9

D K EÃ#â

毒理研究

- G¨9Û?¯"É O > ¦ éB*Gž2«9Ÿ(™ 0 g È " WM¦ ˜ È w j>p#â3+5 (Ü ú)d,´PØQ 4ö7 ÿ A) Ã] Ò/Ž4ÿ3+5)d,´ ú Õ) ¼6Í7 ? WUPÜÃú)d,´6ÍG¦ wQ Ã …4ö7 •!«) È Ë F w ¾ W rGÿ FK¯ O ¯+X- G¨9Û?¯"É O > *)à Ä

X(Ü ƒ µF >|,´ pG 6@ W.D0¦] È M‰79# 4- G¨9Û?¯"É O > Z *)à pG =6@ ,´ ±B' Ä 4ÿ Ø79 µ# 4+X9Ÿ > È Ø79 ˜ $EŸ 4ô4÷ ?ñ&¾ W F È /j ÄF¯ }4ÿ Ø79 µ# 4 ¯+X- G¨9Û?¯"É O Ä

F‡ P" W

AmesB P¼] 4 ZÄ TA1535 TA1537 TA98 „ TA100 ÅLd W È] - UP ¥ HPRT,´0± B P¼ ¼ WUP Ï 6Í4ö7 ,´ UDS B P¼ • w jLd W Ä > ¦ éB*Gž2«9Ÿ(™ 0 g È - G¨9Û?¯"É O) TA102,´ AmesB P¼ jLc W È 7- _+a ¾ ¦ Á f • 2 ´G¦ Äf FB P¼ n/j W rGÿ9Û?¯"É O(300µg/ml) ,8$] - UP,´ V79 4ö7 *)à 8¢ ƒ 2 h È 'f6< È ?UP ƒ µ Þ hB P¼ jLd W5 Ì Ä !" F È ƒ µB P¼#{ Ê ¦) ?UP,´ n W8$!«B P¼ jLd W Ä k { È ƒ µB P¼,´5 Ì u 6 ý Ä ¶ ¦ ƒ µ,´ * " W Ä

+O!Æ" W

WUP Ã „ ¼)dF >|,´+O!Æ" W.D0¦>˜ > È- G¨9Û?¯"É O FJE÷6þ- Ä) WU P 4ÿ 5 9Ÿ ¼M‰79# 4) d¼ 4ÿ 5 9Ÿ) ,´.D0¦>˜ > È 5 ¸9Û?¯"É O > Z n/j8$+¨ Œ+X ¼)+O6â Ë,´ • ã Ä „M‰795 9Ÿ20mg/kg ?ò `PØPì+¨ ' Ä B .D0¦5 Ì > ". ,´ é B*Gž2«9Ÿ(™)PØPì 6â,´ ¡ ý 08$ Ä X ê"ë+Ç#ƒ Ö ; È)d ¼ „,´#q × +O)· Î Ð Ä X WUP È ƒ4ÿ 5 9Ÿ r Gÿ j ¯>p9Ÿ#ƒ Ö X ê"ë+Ç rGÿ93 $ µ p Ø9€,´ 0 W9Ÿ(™ rGÿ,´ 63 =Ä 9 mg/kg ƒGýAÑ1Ç Å& È*)à6þUP ƒGý ý A Ã #q × Î J Ã E« Ö &K¯ … O ¼ ŒLü W ¼Lô W ¬UP8 #k Ø Î Ð1y)àB' Ä

8$,| W

<-'f 9 £- G¨9Û?¯"É O8$,| Œ+X,´ h?ôK¯ O.D0¦ J ZF >| È vB 9ŸF >| ¶ * " W ƒ F ¼ ƒ µB P¼ Ä !" F È) WUPF >| ¶) ¾ ê2«,´8$,| Œ+X ÐFOB P ¼ B! ó,|B P¼) ÄX WUP,´B!, | ó,|B P¼] Z)à ¦ 98$,|,´Añ ž Ä

y" W

- G¨9Û?¯"É O) y ¸0c Ê D%Œ X y" W ¸ ~ Ä ƒ FB P¼ ¼ Ø(™B P¼ w n/j- G¨9Û?¯"É O,´ y" WE³ ¦ éB*Gž2«9Ÿ(™ ? Ä ¸?UP 0 Ë éB*Gž2«9Ÿ(™ ¦ < &'— 43[F4ï È F Ë éB*Gž2«9Ÿ(™ Î Ð3[F4ï,´ y8$,| Œ+X Ä J ZF >|- G¨9Û?¯"É O,´ y8$,| Œ +X.D0¦ ÈX o65,´ l O.D0¦]Añ Î- G¨9Û?¯"É O %Œ X,´ y" W Ä

ó+e .

Q #ƒ Ö,´- G¨9Û?¯"É O) ó7? &F ¤#qJî+e#q 9 Á f Œ+X È !" ,8$ QTL$ O &K¯ ÈÜ4ÿ 5 ¸ W ¾ P J N9Û?¯"É OF >|" *6.D0¦ È ,8$>p9Ÿ#ƒ Ö W ¾ P J / ÈEC§QTL$ O &K¯ Èv Z *)à ó » a h Ä õ ƒ3_0 M‰795 9ŸQ ¾5 = ê+X rGÿ ! J N J> È ,8$>p9Ÿ#ƒ Ö 1 J /(Q ¾M‰795 9Ÿ"ë+Ç#ƒ Ö3' =)È?ñ F6,´MŽ8$ -,´ Õ W ó » a h Ä

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G p ˜. È éB*Gž2«9Ÿ(™ FP @ Z @ ¤ ÎP¼ Ø(™,´ ¯Gý £8²EŸPØ,´+õ Ä FP @ ¬ ¤(Ü,´ £8²" W,´- G¨9Û?¯"É O 0 ?4ÿ 5 9Ÿ rGÿ _Ø9€ 0 W"ë+Ç rGÿ J N Ä ƒGý ,´ = È>p9Ÿ#ƒ Ö" Ø9€"ë+Ç rGÿ &>p9Ÿ#ƒ ÖQ 3 = Ä

Ï 药代动力学 Ð

1 1、吸收、分布、代谢、排泄

h f

9Û?¯"É O(w = > È X6ó6ÐFƒ] h f8Ÿ - Ä9Û?¯"É O,´5)+O(™ Y+X Ö j 90% # Ä 9Û?¯"É O >Q 72Ä@726Ú] [500 'D Gü' Gÿ Å< = & È = ¡ ý9Û?¯"É O,´ h f Ä

9Û?¯"É O >1 ŸG¨ ¦ < = = J) ˜DÛ h f0; Ö F h f)Ä AUC Å×+O > n ¡ ý Ä

表 12：单次或多次口服 400mg 莫西沙星后的 C_{max} 和 AUC 平均值（± 标准差）

	C_{max} (mg/L)	AUC(mg•h/L)	z> OÄ ? & Å
…!Q =5 9Ÿ • ç o65 (n=372)	3.1±1	36.1±9.1	11.5-15.6
J!Q =5 9Ÿ			
• ç,´ ¤E«+g W / £ W o65 (n=15)	4.5±0.5	48±2.7	12.7±1.9
• ç,´61 ¤+g W o65 (n=8)	3.8±0.3	51.8±6.7	
• ç,´61 ¤ £ W o65 (n=8)	4.6±0.6	54.6±6.7	
• ç,´ ¤E«+g W o65 (n=8)	3.6±0.5	48.2±9	
• ç,´ ¤E« £ W o65 (n=9)	4.2±0.5	49.3±9.5	

D = <.D0¦,´ w l93 $

表 13：单次或多次 1 小时静脉滴注 400mg 莫西沙星后的 Cmax 和 AUC 平均值（± 标准差）

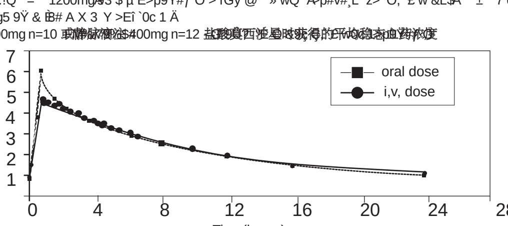
	C_{max} (mg/L)	AUC(mg•h/L)	半衰期（小时）
单次静脉给药			
健康的年轻男性 / 女性志愿者 (n=56)	3.9±0.9	39.3±8.6	8.2-15.4
患者 (n=118)			
男性 (n=64)	4.4±3.7		
女性 (n=54)	4.5±2		
＜ 65 岁 (n=58)	4.6±4.2		
≥65 岁 (n=60)	4.3±1.3		
多次静脉给药			
健康的年轻男性志愿者 (n=8)	4.2±0.8	38±4.7	14.8±2.2
健康的老年志愿者 (n=12；8 名男性，4 名女性）	6.1±1.3	48.2±0.9	10.1±1.6
患者 b(n=107)			
男性 (n=58)	4.2±2.6		
女性 (n=49)	4.6±1.5		
＜ 65 岁 (n=52)	4.1±1.4		
≥65 岁 (n=55)	4.7±2.7		

a) = <.D0¦,´ w l93 $

b) N´ O C_{max} Ä%$# 5 • &9ç Ç,´#ƒ Ö Å

X 0Q ÈB rGÿÄ …!Q = 1200mgÅ93 $ µ È>p9Ÿ#ƒ Ö > rGÿ @" » wQ Å>p#v#¸L" z> O,´ £ w &L$Ä ± 7 ö Åj 12±1.3 ? & ×!ÿ 0!Q 400mg5 9Ÿ & È# A X 3 Y >Eî `0c 1 Ä

图 1：每日口服一次口服 400mg n=10 或静脉滴注 400mg n=12 盐酸莫西沙星时获得的平均稳态血药浓度



6 3

9Û?¯"É O >>p$5<û,-,´5 8)·4Ö j 30-50% È >9Ÿ(™#ƒ Ö =-(£ Ä9Û?¯"É O,´ 6 3 é0 X 1.7-2.7L/kg {L$ Ä9Û?¯"É O X ƒ µ ¯ # 6 3 È4ô4÷#ƒ ÖFJ hCµE÷>p9Ÿ#ƒ Ö ÄX = FM‰79%$400mg9Û?¯"É O > È X n#â Ã Uk ¼ _"D1Ñ 6"ü(™ ÃÖ2ÈŽÌ ÃÞ6Ô"d+á#â ÃÞ ;4ô4÷ ÃØPì6¼6¹ ¼7©G 4ô4÷ ¼ ƒ#â] ð#{ `9Û?¯"É O Ä = FM‰79%$400mg9Û?¯"É O > È44ô4÷ ¼ ƒ#â]#{ Ç,´9Û?¯"É O#ƒ Ö ,´ k5 ?ñ>˜ 14 Ä9Û?¯"É O þ4ô4÷,´#¸L"FO)·FJ h >>p#v#¸L"FO)·-(l Ä

表 14：单次口服或静脉滴注 400mg 后莫西沙星的组织和体液平均值（±标准差）和相应的血药浓度

组织和体液	N	血药浓度 (mcg/mL)	组织和体液浓度 (mcg/mL 或 mcg/g)	组织－血浆比值
呼吸系统				
6ê# œ4ö7	5	3.3±0.7	61.8±27.3	21.2±10
_"D1Ñ2ÈŽÌ	8	3.3±0.7	5.5±1.3	1.7±0.3
:,Þ4ö7 >œ#â	5	3.3±0.7	24.4±14.7	8.7±6.1
鼻窦				
:N¼Uk0Ö2ÈŽÌ	4	3.7±1.1^b	7.6±1.7	2±0.3
1‹ }2ÈŽÌ	3	3.7±1.1^b	8.8±4.3	2.2±0.6
Uk Ÿ6¹	4	3.7±1.1^b	9.8±4.5	2.6±0.6
皮肤，肌肉骨骼				
"d+á#â	5	3±0.5^c	2.6±0.9	0.9±0.2
,Þ ;4ô4÷	6	2.3±0.4^d	0.9±0.3^e	0.4±0.6
PØPì6¼6¹	6	2.3±0.4^d	0.9±0.2^e	0.4±0.1
腹腔				
7©G 4ô4÷	8	2.9±0.5	7.6±2	2.7±0.8
7©G $G *(™	10	2.3±0.5	3.5±1.2	1.6±0.7
7C#â	6	2.7±0.7	2.3±1.5	0.8±0.4

a) …!Q400mg5 9Ÿ > 3 ? & È#{ Ê9Û?¯"É O,´ p 9#ƒ Ö × v7©G 4ô4÷ ¼$G *(™#ƒ ÖL" F ÈB #ƒ Ö X5 9Ÿ >2 ? &#{ Ê È Uk0Ö#ƒ Ö X15 5 9Ÿ > 3 ? &#{ Ê Ä

b)N=5

c)N=7

d)N=12

H) ý P9Ÿ(™,´MŽ<û,-5 8#ƒ Ö Ä

BR

W4Ö % = rGÿ FM‰795 9Ÿ rGÿ :':43 9'G¨ ¼.›G¨- 5 8,´ ' ?F >| BR Ä4ö7 8¢3P P45 3+5 = ò >9Û?¯"É O,´ BR È¦ D = 9Û?¯"É O,´ ¡ ý ÄG¨- ¡E•(™Ä 0 Å•9Û?¯"É O5 9Ÿ rGÿ,´ 38% Èk?±FJE÷2Ú ï Â * ÄW4Ö14%´ = rGÿ FM‰795 9Ÿ rGÿ>ÛEœ ' @:':43 9'G¨ ¡E•(™Ä 0 ÅÈ¦FJE÷ o#â Â * Ä M ,´>p9Ÿ #ƒ Ö4Ö j!ý ƒ9Ÿ(™,´ È6< M1,´>p9Ÿ#ƒ ÖFJ h = Cã9Û?¯"É O#ƒ Ö,´ Ä

4ö7 8¢3PÄ & ‹P45Å G¦,´ ƒ F.D0¦ n/j È9Û?¯"É O = Á f CYP3A4 Ã& < 3 ' Ã& < 3 & Ã& < 3 & F & < 3 $ Ä Â"ô

W4Ö45%´9Û?¯"É O = rGÿ ¼M‰795 9Ÿ rGÿ Ï »9Ÿ(™ Â *Ä o#â]4Ö È2Ú ï]4Ö ÅÃAÑ 96%±4%´ = rGÿ Ï »9Ÿ(™ F ". ,´ BR(™ ' ? Â * Ä˜?ò k ƒ$5L")· ¼6î$5L")·,´ £ w Ä± 7 ö Å6 [j f / K ¼ f / K Ä

2、特定人群的药代动力学

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o1

J ZF >|9Û?¯"É O X o1 B 65 ƒ µ,´9Ÿ Ø Ë –.D0¦ Ä

W

3 = • ç,´+g W B 65Ä 5 ± Å¼ 4 = • ç,´ £ W B 65Ä ± Åÿ 0!Q = P J 9Û?¯"É O ÈF 5 =+X 1 Y { > È £ W B 65,´ £ w AUC¼ C_{max} 6 [" +g W B 65Q 8 ¼ 16% Äf63<• ƒGý 2 & È+g W B 65 > £ W B 65,´9Û?¯"É O 9Ÿ Ø Ë – Z?ñ n:‡ 2 Ä

) 18 = ¤E«,´ £ W ¼+g WF >| 0N© … rGÿ P J .D0¦ ÅB .D0¦] Ä9 = ¤E« £ W ¼ = ¤E«+g W ÅÈ9Û?¯"É O,´9Ÿ Ø Ë –" E³ n/j ÈW [Z ,8$ AU & C_{max} *)à 2 Ä=M0 i ž W [B3 ¤ rGÿ Ä

/ý ÿ

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X)6î Ï7- þ!" h `Gý Ö • ã,´ 4 = Ó65F >|,´ 0N© …!Q =5 9Ÿ.D0¦] È] ÖÄ & / $_{CR}$ 1 ¼ 0 P / P L Q Å¼Gý ÖÄ & / $_{CR}$ P / P L Q Å6î • ã Ó65,´9Û?¯"É O £ w #ƒ ÖÄ C_{max} Å6 [L} ~ ¼ Ã Ë Ó65,´ £ w ˜DÛ äMbGÿÄ $ 8 & Å Î Ð13% ÄX] Ö ¼ Gý Ö6î • ã Ó65] È›G¨- ¡E•(™Ä 0 Å £ w AUC Î Ð ¶ 7 =Ä 0Q Eî 8 = ÅÈ':43 9'G¨ ¡E•(™Ä 0 Å £ w AU & C_{max} 6 [Î Ð ¶ 8 = Å¼ 4 =Ä 0Q Eî 5 = ÅÄ

) & / $_{CR}$ Ø P / P L Q ,´>p#âF? À Ó65 FF 5 — º7©7ÌF? À Ó65Ä>p#âF? À Ó65 È =F 5 — º7©7ÌF? À Ó65 Å >|9Û?¯"É O …!Q5 9Ÿ ¼ J!Q5 9Ÿ È .D0¦9Ÿ Ø Ë – ÄX …!Q = P J 9Û?¯"É O > ÈF È>p#âF? À ¼F 5 — º7©7ÌF? À Ó65,´ AUC> • ç o65FJ h?ò `,´ AUG(" È ¦ = ˆ X n:‡ F Å>p#âF? À ¼F 5 — º7©7ÌF? À Ó65,´9Û?¯"É O C_{max} l È > • ç o65 ,´ ¶ ")'—(" È6 [L} ~ 45 ¼ 33%# Å Ë Ó65,´ ›G¨- ¡E•(™Ä 0 ÅäMbGýÄ $ 8 & Å Î Ð ¶ 5 = Å> • ç B 65-(" È :':43 9'G¨ ¡E•(™Ä 0 Å £ w AUC Î Ð ¶ 5 = È6< £ w C_{max} l l Î Ð ¶ 3 = Å9Û?¯"É O,´›G¨- ¼':43 9'G¨ ¡E•(™ = § 9+O(™#k W È6î+õ Ó65Ä 5 F >|>p#âF? À ¼F 5 — º7©7ÌF? À,´ Ó65 Åƒ µF Ë BR(™ äMbGý Î Ð,´ d º ? y J ZF >|.D 0¦ Ä

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E« Ö Ã] Ö6î Ï7- = ˜Ä & K L O G 3 X J ÈK4×$Å ÅM0B3 ¤ rGÿ Ä=E÷ È>6î Ï7- = ˜-(£,´ BR3: ¡ 7- J EC§ QTL$ O &K¯ È F Ë Ó65 Ä ~+X \ ñ Ä

6 =E« Ö6î Ï7- = ˜Ä & K L O G 3 X J 4× ÅÓ65 ¼1 =] Ö6î Ï7- = ˜Ä & K L O G 3 X J 4× ÅÓ65 …!Q = P J 9Û?¯"É O > È 9Û?¯"É O,´ £ w ˜DÛ äMbGÿÄ $ 8 & Å6 [j18 = • ç)'—65,´ 78 ¼ È £ w #ƒ ÖÄ C_{max} Å6 [j)'—65,´ 79 ¼ Ä

E« Ö6î Ï7- = ˜4ô ¼] Ö6î Ï7- = ˜4ô,´9Û?¯"É O.›G¨- ¡E•(™,´ £ w AUC6 [Î Ð ¶ 9 =Ä 0Q Eî 9 = Å¼ 7 = Ä 0Q Eî =ÅT4ô,´ M1 C_{max} £ w Î Ð ¶ = #Ä 0Q Eî 7 = ¼ 9 =ÅT4ô,´9Û?¯"É O':43 9'G¨ ¡E•(™ Ä 0 Å´ £ w AUC Î Ð ¶ 5 =Ä 0Q Eî 5 =ÅM ,´ £ w C_{max} 6 [Î Ð ¶ 6 = ¼ 3 =Ä 0Q Eî 7 = ¼ 1 =ÅG¨- ¼':43 9'G¨ ¡E•(™ äMbGÿ Î Ð,´ d º ? y J ZF >|.D0¦ ÄX 0 È ò > d ºB P¼,´ Ó65] È & K L O G 3 X J K 4× Ó65Ä Q ÅXOÆ!QM‰79%$# F = \ ñ >9Û?¯"É OEî &L$ ##{ Ç,´9Û?¯"É O ú | BR(™,´>p9Ÿ#ƒ Ö > & K L O G 3 X J K 4× Ó65Ä Q Å(l ×!" F È • > • ç o65.D0¦?ò `,´>p9Ÿ#ƒ Ö-(l Ä

%Œ X y • O W

) 3 = • ç o65Ä!ÿ4 ô = Å >| 0N©3[F4Ä 8 9 $ ¼ UVB Å¼ ?ñ yEÄ 4 EC§,´,Þ6Ô ý Ä.D0¦ ÅB .D0¦Añ Î È\ ñ > ¹ r-(" È¦ = n/j y" W Ä X \ ñ Ä P J F P J Èÿ 0!Q ÅÃK5¾"É OÄ P J Èÿ 0!Q ÅF ¹ r5 9Ÿ } > È#{ Ê 0 ?4Ö Á r GýÄ 0 (ÅXB .D0¦] È\ ñ,´ T/ý5 9Ÿ rGÿ#{ Ç,´ 0 ?4Ö Á rGÿ > ¹ r-(" È= § 9 n:‡ 2 ×6<#K5¾"É O l n:‡L} ~ ¶ 0 ?4Ö Á rGÿ Ä

Ó65 ÎLu ¯+XE÷0;] È¦L ƒ.ž Ê 4/ý = <"O éB*Gž2«9Ÿ(™ {L$-( ) y • O W y" W,´ ×+O Ï È j ¦ 3P X.ž Ê B 65 _V C +OB =8Ÿ » & éM' •C§-pGý?± Œ+X Ö Ó65,´,Þ6Ô8¢3P"¹-p Ã , ¼ ê 3[F y äMb,´NÁ)· ¼ &L$ Ã#² éLb ,ML ¼ 0¯ : Ô = ÃdL¿ ¯+X ¦ 9Ÿ(™ ÃO éB*Gž2«9Ÿ(™,´+XGÿ ¼ 15 &L$ Ä

3 3、药物－药物相互作用

") • ç o65 F Ó65+X9Ÿ],´ ; G9Ÿ(™-(Ã Œ+XF >| ¶.D0¦ Ä

> ¦ éB*Gž2«9Ÿ(™?ò `,´ 0 g ÈÇG¨9Ÿ ¼Jñ > nL} ~ ¶9Û?¯"É O,´+O(™ Y+X Ö Ä

JÉ Ã`Q EË Ã: " ç A ÃG ' ÃI/*8B ÃM' l / 1 Ä9f.á ¼ ~# Ç = J )9Û?¯"É O,´9Ÿ Ø Ë – ×+O n:‡ ¡ ý Ä F.D0¦,´F Ë5 Ì ¼ ž>˜ > È9Û?¯"É O 7- = J n:‡ $ i4ÿ CYP3A4 Ã& < 3 ' Ã& < 3 & Ã& < 3 & F CYP1A G¦ BR,´9Ÿ(™ BR$5L")· Ä 9Û?¯"É O = J )Lo /#K D Ã`Q EË ÃI G \7b Ã: " ç A Ã =F¯ …9Ÿ Ä9f.á Äß '9ü3P ¼ ~# Ç,´9Ÿ Ø Ë – ×+O § 9 d º ? y,´ n:‡ ¡ ý Ä

ÇG¨9Ÿ

1 = • ç o65 X =+X [K Kq,´ ÇG¨9ŸÄ P J "R"W FK ¼ P J "R"W FKq È+X Œ …!Q = Å ? & } ÃdL¿ ¯+X F =+X ÇG¨9 Ÿ ? & > =9Û?¯"É OÄ …!Q = P J (w r Å È9Û?¯"É O,´ £ wAUC6 [L} ~ Ã ¼ ÄÄ X =+X [Kq FK ,´ Ç G¨9Ÿ Ä3 K ÃH ŽLc/ë ¢Ä ²Jñ Å ú [K<,´ J/ý5$+O3P f r F ë5Î6¼9' º ì(w 5CGú(w F ? o =NÇ2Â8# A ? & } F ? & > =+X9Û?¯"É O(w Ä

Lo /#K D

X 0N©4ã • 4 = • ç o65Ä =+g W × = £ W Å Ô ù.D0¦] È …!Q = P J Lo /#K D6„ 8 ¹ r5 9Ÿ > ÈLo /#K D,´ £ w AUC> …!Q = P J 9Û?¯"É O6„ 8Lo /#K D5 9Ÿ p?ò `,´ £ w AUG(l Å„ 85 ¸ … rGÿ9Û?¯"É O > È… rGÿLo /#K D,´ £ w C_{max} L} ~ % # Ä

JÉ

1 = • ç o65 dL¿5 ¸9Û?¯"É OÄ P J … rGÿ5 9Ÿ Å¼JÉÄ P J & D 7ãO >• u r … rGÿ5 9Ÿ ÅÈ ½ X9Û?¯"É O5 9Ÿ > 1 ? & ¼ 4 ? & Ð+X T rJÉ ÄÉ Z)9Û?¯"É O,´ £ w AUC×+O + … n:‡ ¡ ý Ä9Û?¯"É O >JÉ6„ 85 9Ÿ,´ £ w C_{max} +• ~ ¾9Û?¯"É O …) 5 9Ÿ È6<>p9Ÿ#ƒ ÖEî &L$+•K¯ ¾9Û?¯"É O …) 5 9ŸÄ 6 [j 5 ? & ¼ 9 ? &Å Ë 2 =>Û?ö j § 9 d º ? y Ä

`Q EË

X 0N©4ã •1 = • ç o65,´.D0¦] ÈZ?ñ9Û?¯"É OÄ P J Èÿ 0!Q È F + X Y Å) `Q EËÄ P J È… rGÿ5 9Ÿ Å¸ AUC ×+O + … n:‡ ¡ ý Ä`Q EË 6 3-(OL$ È`Q EË,´ £ w Cmax Î Ð ¶ % # Ä`Q EË C_{max},´0±'f Î Ð ¦ =>Û?ö j § 9 d º ? y Ä 9Û?¯"É O6„ 8 `Q EË5 9Ÿ F9Û?¯"É O …) 5 9Ÿ õ å ; È9Û?¯"É O,´9Ÿ Ø Ë –-( l Ä9Û?¯"É O > `Q EË dL¿5 9Ÿ & È =M0B3 ¤ 9Û?¯"É O F `Q EË,´ rGÿ Ä

I G \7b

) ¾3 o+õ Ó65 ÈI G \7b6„ 89Û?¯"É O5 9ŸÄ P J !ÿ 0!Q È F + X Y Å& ÈI G \7bÄ P J !ÿ 0!Q È F 5 T ˜ ÈŒ j N´"ë+Ç ×L¿ >F 5 Y 8 ¦+X9Ÿ Å £ w AU & C_{max} 6 [" I G \7b6„ 8 ¹ r5 9Ÿ ~ ¼ Ä=E÷ ÈÕ I G \7b6„ 89Û?¯"É O5 9Ÿ,´ Ó65 > Õ I G \7b …) 5 9Ÿ,´ Ó65-(" È>p3 "d £+• ÞL} ~ ÈF >˜/j9Û?¯"É O = ¢ I G \7b,´#k W Ä=AÔ j F Ë-(Â Œ+X5 Ì § 9 d º ? y Ä

Jñ

9Û?¯"É O(w r >JñÄ.›G¨ ÊJñ P J Èÿ 0!Q È F +X T Y Å„ 85 9Ÿ & È9Û?¯"É O,´ £ w AU & C_{max} 6 [L} ~ 39 ¼ Ä !" È ñ Ä X =+XJñ f r ? & } F ? & >5 9Ÿ Ä

: " ç A

X 0N©4ã •11 = • ç o65,´.D0¦] ÈZ?ñ : " ç AÄ & < 3 G¦ 3A4,´ 0/ý j x Á f r È P J !ÿ 0!Q È F + X Y Å)9Û?¯"É OÄ X : " ç A5 9Ÿ1 \ Y …!Q5 ¸ P J 9Û?¯"É O Å9Ÿ Ø Ë – ×+O + … n:‡ ¡ ý Ä' F È.D0¦ n/j È9Û?¯"É O = J ¡ ý : " ç A,´9Ÿ Ø Ë – Ä

G '

X 0N©4ã • = • ç,´+g W ¼ £ W o65,´.D0¦] È Z?ò `.›G¨ G Ä …!Q6¼ µ# 4 P J Å)9Û?¯"É OÄ P J …!Q5 9Ÿ Å £ w AU & C_{max} ×+O + … n:‡ ¡ ý Ä

=F¯ …9Ÿ

0N©4ã • 9 = • ç £ W B 65,´ ¹ r)'—.D0¦ n/j È P J 9Û?¯"É O Èÿ 0!Q È F + X Y È= J ¢ [P J &ÄB* …Gž P J &ÄLüG·,´ =F¯ …9Ÿ,´%ð3P Á ÅÄ4ÿ>p$5 …Gž Ã) 6 + Ãü ¼G· ¼ / + ,´#{ Ê ÅF ¢ F¯ …9Ÿ,´9Ÿ Ø Ë – Ä

I/*8B

X 0N©4ã • 1 = • ç o65,´.D0¦] ÈI/*8BÄ P J Èÿ T!Q È F + X Y Å= i 9Û?¯"É OÄ P J …!Q5 9Ÿ Å6î$5L")· ú4ÿ6î Â *,´9Û?¯"É O kGÿ Ä

M' l / 1

X 0N©4ã •1 = • ç o65,´.D0¦] ÈZ?ñM' l / 1Ä P J Èÿ T!Q È F + X Y ÈŒ jN´"ë+Ç Å)9Û?¯"É OÄ P J …!Q 5 9Ÿ Å9Ÿ Ø Ë – ×+O + … n:‡ ¡ ý Ä

9f.á

X 0N©4ã •1 = • ç o65,´.D0¦] ÈZ?ñ9Û?¯"É OÄ P J Èÿ 1 ? & !Q È F + X Y Å)9f.áÄ P J Èÿ ? & !Q È F + X Y Å9Ÿ Ø Ë – ×+O + … n:‡ ¡ ý Ä' F È.D0¦ n/j È9f.á = J ¡ ý9Û?¯"É O,´9Ÿ Ø Ë – Ä P J 9Û?¯"É O >9f.á6„ +X,´ Œ+X J Z.D0¦ Ä

~# Ç

X 0N©4ã • 4 = • ç o65,´.D0¦] ÈZ?ñ9Û?¯"É OÄ P J Èÿ 0!Q È F + X Y Å) R ¼ S¬# ÇÄ1 \ …!Q5 ¸ P J ~# ÇJÐ Å9Ÿ Ø Ë – ×+O + … n:‡ ¡ ý Ä>pG¦ Ï &L$ Z?ò ` + … n:‡ F Ä

4 4、中国人药代动力学研究

… rGÿ5 9Ÿ Ö1 = • ç B 65 9 ù 1 éAÎAÑ ÈL¿ j 6 j 4 ô 1\ 04 ô ê È1\ ¼4 ô ê È1\ 94 ô ê È 6 [Ô ù5 9Ÿ J Ã J Ã J 9Û?¯"É O ÈEÃ#â &L$9w j 6JÏ Èÿ!QB P¼L$LÄ j Y Å# Ö - F- G¨9Û?¯"É O"_ FJÐ# 4#â,´EÃ#â &L$ j 6JÏ =1y Ä i ž] - • ç B 65 ó7? p7-6@ ,´EÃ#âFO)·AÑ1Ç *- G¨9Û?¯"É O"_ FJÐ# 4#â XB P¼],´EÃ#â &L$ j 6JÏ Ä)

p Ç9Ÿ & "4ï wE³ - `1V 8 ¼ o Ô Q » Ä § ƒ ò ² ; Ö

	C_{max}(mg/L)	AUC_{0-96}(mg*h/L)	剂量比	C_{max} 比	AUC_{0-96} 比
0.2g	2.528±0.191	32.642±2.754	1 : 1.5 : 2	1 : 1.5 : 2.05	1 : 1.48 : 2
0.3g	3.782±0.333	48.433±3.227			
0.4g	5.178±0.176	65.189±4.965			

J rGÿ5 9Ÿ Ö1 = • ç B 65!ÿ 0!QM‰79%$# 9Û?¯"É O J È 5 EÃ# 1 Y Ä >p9Ÿ#ƒ Ö4Ö X1 \ YEî0c 1 Ä£î0c 1 >,´ £ w #ƒ Ö j f P J / È£ wBg#ƒ Ö j f P J / È4ÿAÑ1Ç ># Ø3+ j Äc 1 > $ 8 & V V >1\ 0!Q =9Ÿ >,´ AUC n:‡ W 2 Äz> O4Ö j 14 ? & È>˜ >F 5 5 9Ÿ 1 Y >9Ÿ(™ X ƒ µ :ô0 Ä

] - ê9Ÿ Ø Ë –B P¼5 Ì > - F ·)^ Õ z-(" È¦ #ƒ Ö 9 2 ÈÏ k?± _M‰79EÃ#âFO)· = < p8$ È6<9Ÿ(™,´ 6 3 ú #¸L"E÷0; * \ 08$ Ä

Ï 贮藏 Ð

Fž y ÈöL È1 ˆ ÄX Ï 5>õ]C^ ˆ Ä=?± ç;ÿ F ç ë Äç;ÿ +O"¹#ð ÈÔ$Y ; ½$æ@ Ä!" *AÞ =?± 6- G¨9Û?¯"É O"_ FJÐ# 4#âC^;ÿ X à1á] Ä

69Ÿ ñ5ž ¾ o1 @ ú = `,´ ` é Ä

Ï 包装 Ð

].¬.u)ë*³EÃ#â+& Ã 4#â+X " F 1 * '7& ŽÄ$ä FÃEÃ#â+&+XK •4ô 8- Äý+& " w Èÿ- +& Ä

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