N *H*

!

	/ /	
	1 / /	
	2 / /	• 2 2 • 3 2
	3 / /	
	QT QTc 500 ms 60 ms	3 * • 480ms 3 QTc 55mg • 480ms 3 QTc 480ms - 110mg - 55mg
	QTc	
	LVEF 10% 50%	3 * 3 LVEF • 110mg • 55mg
CK	CK 5 2	# 2
	5 CK 10 3 #	3 * 3 2 5 • 110mg • 55mg
	CK 10 4 #	3 * 3 2 5 • 55mg •

	$^{\#}$ 3 CK	3 * 3 2 • 110mg • 55mg
	3	3 * 3 2 • 110mg • 55mg
55mg 3		

"　　!!　　!

	110mg N=697	
	%	3 %
[a]	21.8	0.6
	10.8	0.3
[b]	10.6	0.3
	10.2	0.1
[c]	11.5	0.3
	34.0	6.5
	20.4	0.7
	19.8	1.7
	16.5	0.6
	12.1	0.6
	10.3	0.7

•

□! " □ " !

	N=106		N=107	
	%	3 %	%	3 %
a	26.4	1.9	7.5	0
	17.9	0.9	7.5	0
b	14.2	1.9	0.9	0
	11.3	0	4.7	0.9
	60.4	6.6	14.0	0
	27.4	0	17.8	0
	26.4	0	25.2	0.9
	15.1	0	14.0	0
	14.2	0	4.7	0
	13.2	0	9.3	0
QT	12.3	1.9	3.7	1.9
	12.3	0.9	3.7	0

-

- □! " □ !

	N=94		N=53	
	%	3 %	%	3 %
[a]	14.9	0	5.7	0
	45.7	6.4	7.5	0
	24.5	0	7.5	0
[d]	23.4	1.1	18.9	0
	14.9	1.1	1.9	0
	13.8	0	3.8	0

-

! " ! !

	N=214		N=215	
	%	3 %	%	3 %
[a]	26.6	0	49.3	0
	15.4	1.4	34.4	0.9
[b]	12.6	0	11.2	0
	17.8	0	13.5	0.9
	11.2	0	8.4	0
[c]	19.2	0.5	19.5	0
	19.2	0.9	8.8	0
[d]	15.4	0	6.5	0

e	43.0	3.7	62.3	14.4
	35.5	7.0	8.8	0.5
	22.9	2.3	12.6	0
	21.0	0.9	7.4	0.9
	12.6	1.9	9.3	0
	11.7	0	6.5	0
QT	10.7	0.9	8.8	1.9

HS-10296-03-01

NCI-CTCAEen-8NCI

	%	3 %
[a]	21.6	0.7
	13.1	0
[b]	11.0	0
	10.2	0.4
[c]	12.7	0.4
[d]	11.7	0.4
	11.3	0.7
	20.1	6.0
[e]	19.4	1.8
	11.7	0

a.

b.

c.

d.

e. -

f.

110 mg 697 237 34.0%

1 113 16.2% 2 79 11.3% 3 34 4.9% 4
11 1.6% 5 33 4.7% 17 2.4%
1 0.1% 1 0.1%

64
44
2 2 1 2
21 3.0% 5 3 7 2
9 1 5 2 1 14 2.0%
3

QT

110mg 697 54 7.7% QT
1 36 5.2% 2 13 1.9% 3 5 0.7% 4
5 4 0.6% 1 0.1%
QT 44.5 QT
38.5

110mg 697 8 LVEF
10% 50% 1 4

/

110mg 697 2 2

HS-10296-304 147 7 4.3%
1~2

110mg 697 35 5.0%
1 27 3.9% 2 8 1.1%
3

42 93

110mg 697 16.5% 12.1%

10.3% 3.3%

1~2 3 1%

1%

---

t_{max}

C_{max} 318.50 ng/mL

ng/mL

mg

- **EGFR II-IIIB NSCLC-III**

HS-10296-302

HS-10296-302 III

EGFR II-IIIB AJCC 8

NSCLC 110 mg 110 mg

EGFR 19 L858R

II IIIA N2 IIIA/B N2+

IRC DFS

DFS IRC 2 DFS 3 DFS 5 DFS OS 5 OS

214 1 1

107 107

27.6

59.2 30~75

24.05 kg/m^2 15.8~37.3 kg/m^2 44.9% 55.1% 97.2%

72.0% 28.0% 100.0% IB 0.9% II 44.9% IIIA 49.5%

IIIB 4.7% 19 47.7% L858R 52.3% 1 T790M

1 EGFR 20 ECOG PS

1 63.6% 0 36.4% 94.4% 6.9%IB-IIA 93.1%IIB-IIIB

EGFR II-IIIB NSCLC

106 104

83% mDFS mDFS

19.4 HR 0.166 95% CI 0.094, 0.294 p 0.0001 IRC

DFS Kaplan-Meier OS

DFS

EGFR ECOG PS

CNS

6 5.7% 17 16.3%

HS-10296-302 IRC

	(N=106)	**(N=104)**
(DFS)		
n (%)		
(95% CI) [ ]		

HR (95% CI)

24.16kg/m^2 19.1~33.0 kg/m^2 45.7% 54.3% 97.8% 69.6% 30.4% 90.2% IIIA 23.9% IIIB/IIIC 76.1% 19 41.3% L858R 58.7% 75.0% 25.0% ECOG PS 1 81.5% 0 18.5%

PFS HR=0.200 PFS mPFS 30.4 3.8 BIRC 8 PFS Kaplan-Meier 2 OS OS mOS 12.7% 12.0% 14.0% mOS mOS 36.99 HR=0.797 95%CI 0.306 2.074

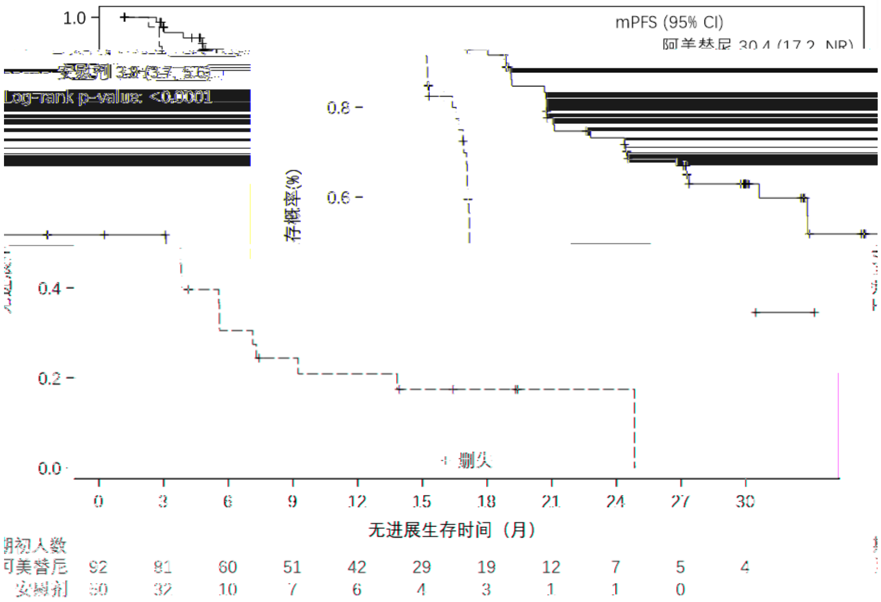
PFS TNM IIIA IIIB IIIC EGFR Ex19del L858R <65 65 ECOG PS 0 1

! "

	N=92	N=50
PFS		
n %		
95% CI [ ]		
HR 95% CI		
p		
ORR		
% 95% CI		
95% CI		
p		
DCR		
% 95% CI		
95% CI		
p		

HR= CI= NA= BIRC RECIST v1.1

mPFS 30.36 3.78 HR=0.150 95% CI 0.080 0.284 p 0.0001 BIRC



2 ! "

- **EGFR** **NSCLC-III**

HS-10296-03-01

HS-10296-03-01

1

1 EGFR 19 L858R

1.1 RECIST v1.1 PFS

OS ORR DoR DCR

DepOR 429 214

215 20.5 20.7

58.1 32~78

23.13 kg/m^2 16.0~37.0 kg/m^2 37.4% 62.6% 96.7%

72.9% 27.1% 98.1% IV 94.4% IIIB 5.6% 19

65.4% L858R 34.6% ECOG PS 1 74.8%

0 23.8% 26.2%

PFS

PFS mPFS 19.3 9.9 HR=0.463

PFS Kaplan-Meier

OS OS mOS 24

70.4% vs. 58.8%

PFS

EGFR 19 L858R

65 65 ECOG PS 0 1

EGFR 19

mPFS 20.8 12.3 HR=0.389 p 0.0001 L858R

mPFS 13.4 8.3 HR=0.599 p=0.0102

mPFS 15.3 8.2 HR=0.376 p 0.0001

mPFS 19.3 12.6 HR=0.507 p 0.0001

□! " □ □!

	N=214	**N=215**
PFS		
n %		
95% CI [ ]		
HR 95% CI		
p		
DoR		
95% CI [ ]		
HR 95% CI		
p		
ORR		
% 95% CI	73.8 (	72.1 (
95% CI		
p		

HR= CI= NA= RECIST v1.1

BIRC

mPFS 17.9 9.7 HR=0.500 95% CI 0.390 0.641 p 0.0001

ORR (95% CI)	68.9 (62.6,74.6)
DCR (95% CI)	93.4 (89.6,96.2)
mPFS (95% CI) ( )[1]	12.3 (9.6,13.8)
mDoR (95% CI) ( )[2]	12.4 (11.3,NA)

IRC RECIST v1.1

PR SD 5 1

48.0% 2

4

CNS II 91

23 9.4% IRC IRC

RECIST v1.1 CNS ORR 60.9% (95%CI 38.5% 80.3%)

4.3% CR CNS DCR 91.3% (95%CI 72.0% 98.9%)

CNS mPFS 10.8 47.8% CNS mDoR 11.3