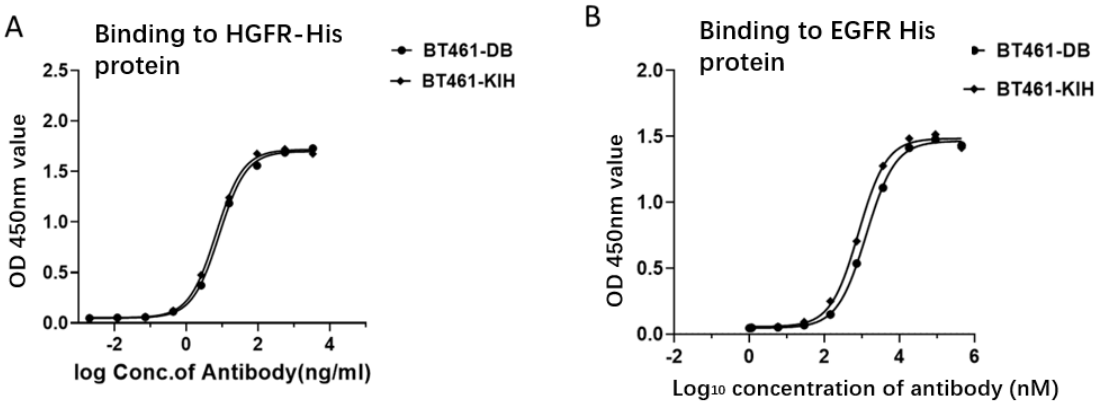


FIG. 1

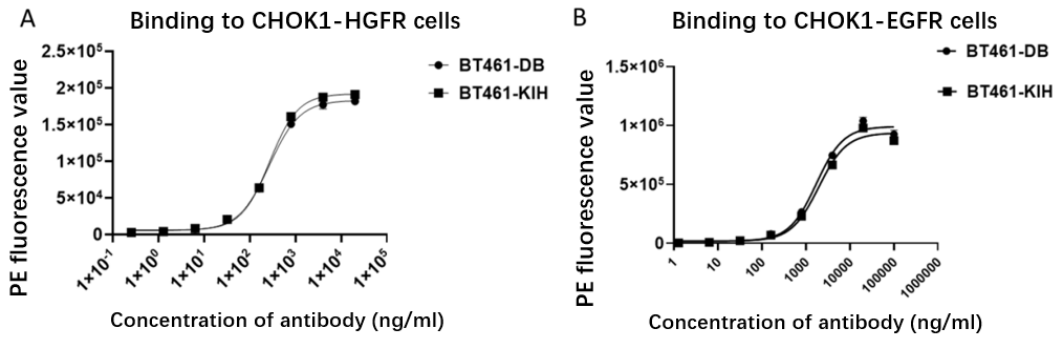
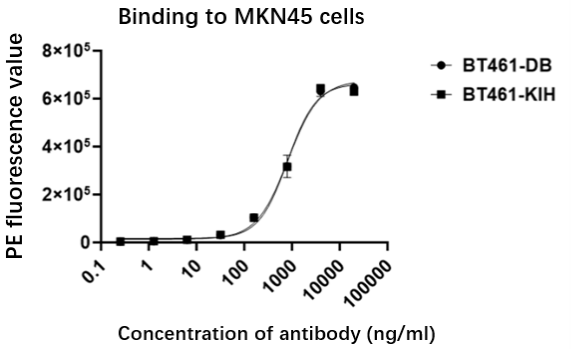


FIG. 2



	EC50（ng/mL）
BT461-DB	829.2
BT461-KIH	809.5

FIG. 3

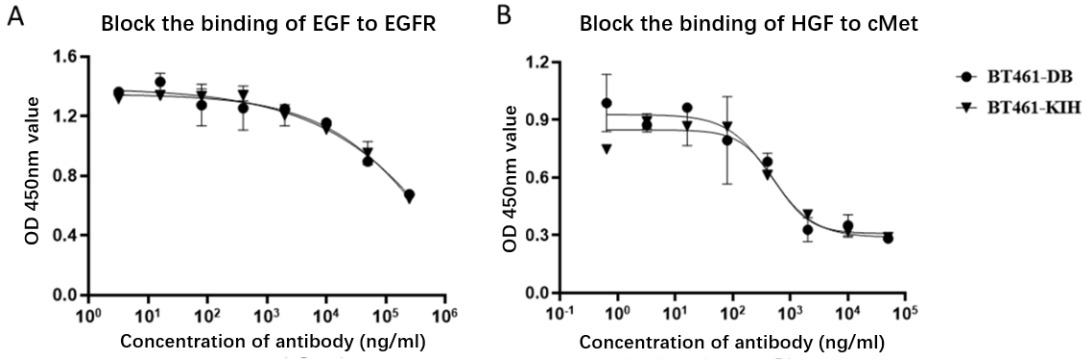


FIG. 4

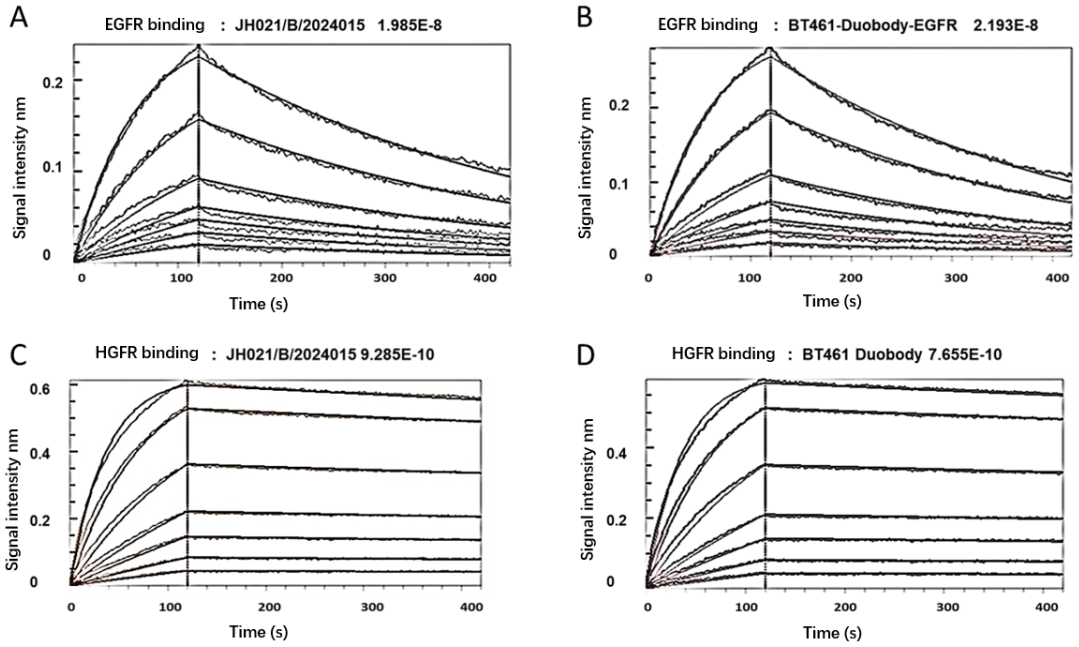

A
EGFR binding : JH021/B/2024015 1.985E-8
B
EGFR binding : BT461-Duobody-EGFR 2.193E-8
C
HGFR binding : JH021/B/2024015 9.285E-10
D
HGFR binding : BT461 Duobody 7.655E-10
Signal intensity nm
Time (s)

FIG. 5

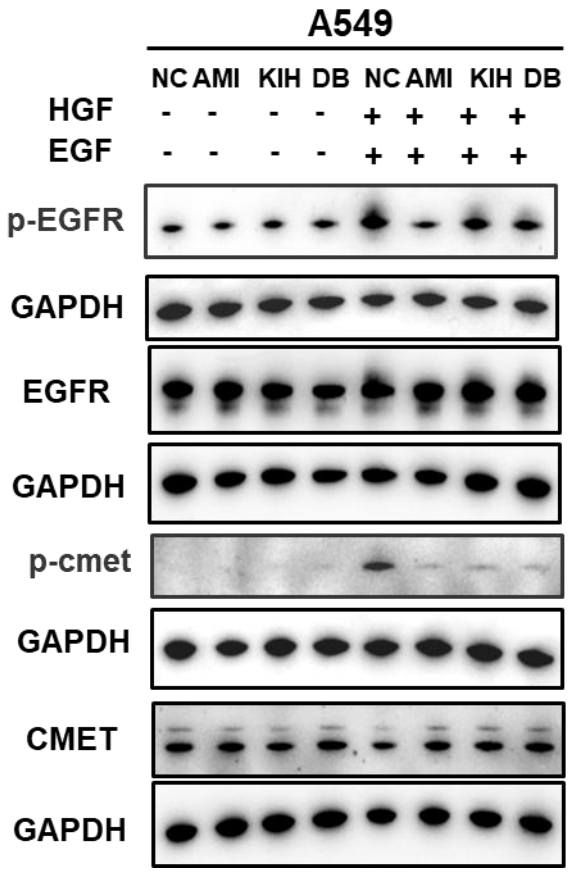
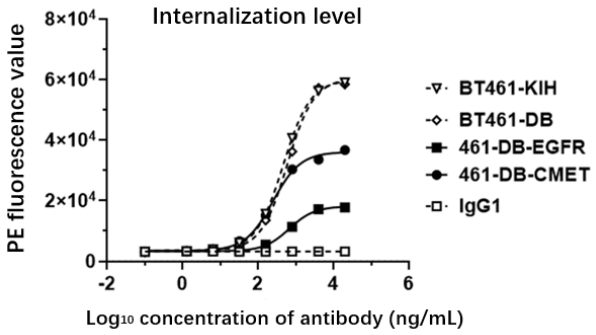

A549
NC AMI KIH DB NC AMI KIH DB
HGF - - - - + + + +
EGF - - - - + + + +
p-EGFR
GAPDH
EGFR
GAPDH
p-cmet
GAPDH
CMET
GAPDH

FIG. 6



	EC50(ng/mL)	Top 10^3
IgG1	~	~
461-50L	460.9	59.9
461-Duobody	593.8	60.3
461-DB-EGFR	671.4	18.1
461-DB-CMET	246.5	36.1

FIG. 7

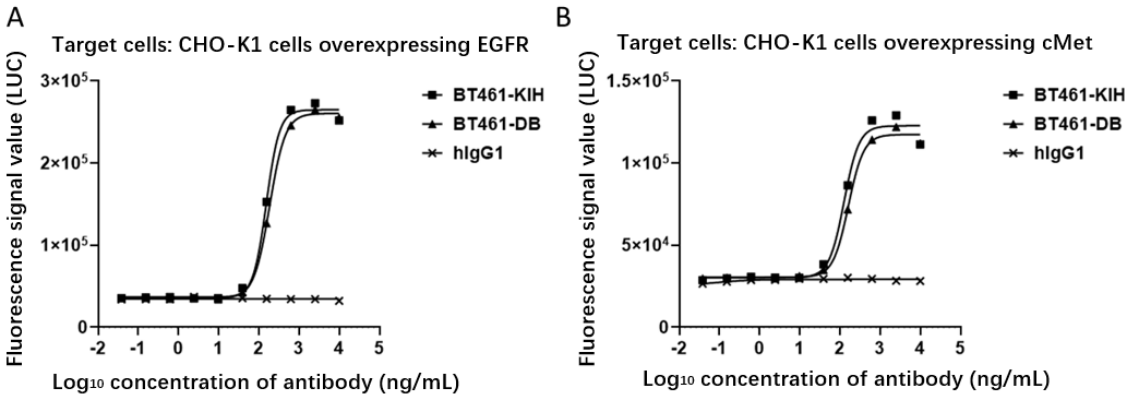
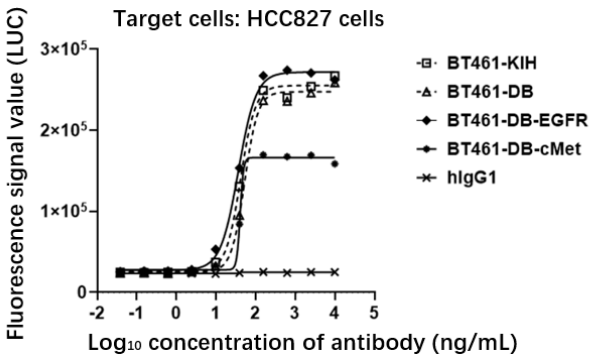


FIG. 8



	EC50 (ng/mL)	TOP(10^3)
BT461-KIH	41.67	255.3
BT461-DB	52.58	247.6
BT461-DB-EGFR	36.49	271.8
BT461-DB-cMet	40.51	166.3
hIgG1	NA	24.8

FIG. 9

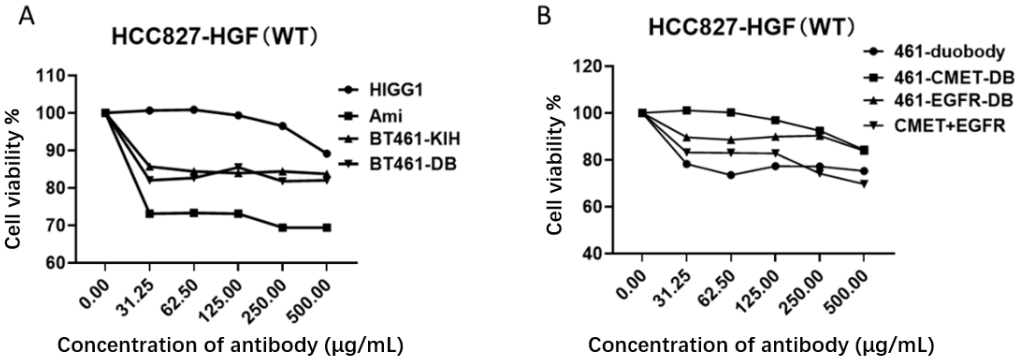


FIG. 10

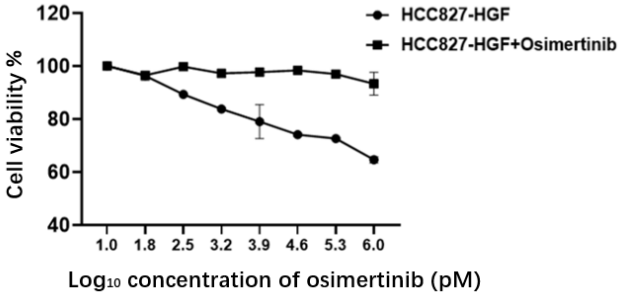


FIG. 11

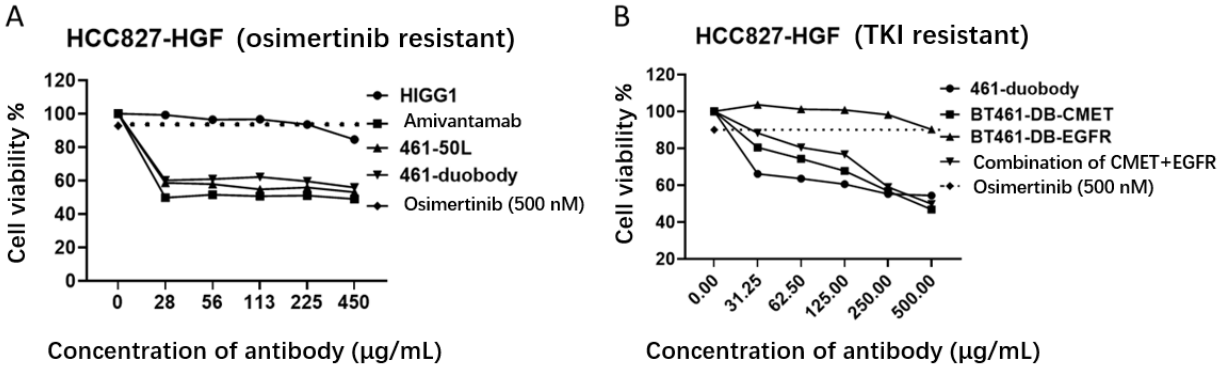
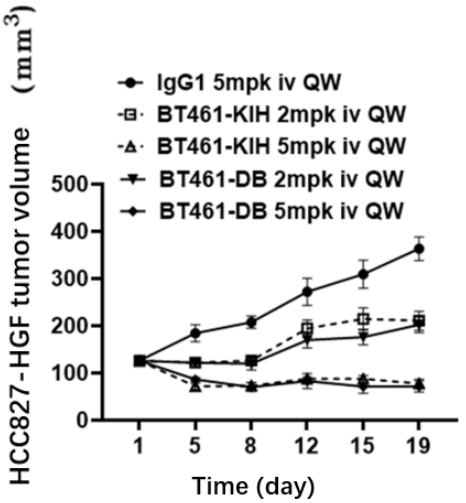


FIG. 12

A B



TGI of tumor model HCC827 (EGFR exon 19 deletion mutation)	
IgG1 5mpk iv QW	-
BT461-KIH 2mpk iv QW	64.2%
BT461-KIH 5mpk iv QW	120.4%
BT461-DB 2mpk iv QW	67.7%
BT461-DB 5mpk iv QW	122.9%

FIG. 13

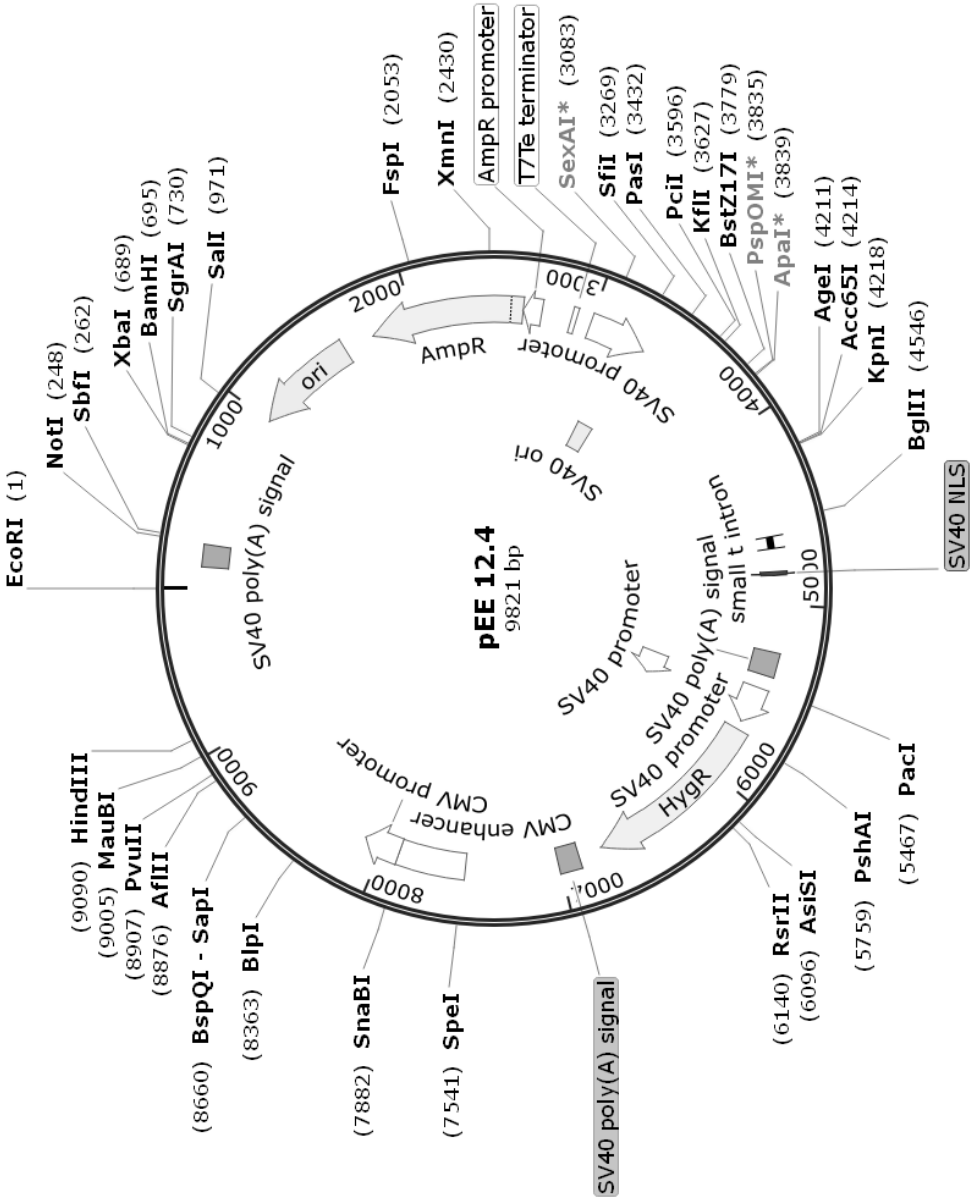

pEE 12.4
9821 bp
EcoRI (1)
NotI (248)
SbfI (262)
XbaI (689)
BamHI (695)
SgrAI (730)
SalI (971)
FspI (2053)
XmnI (2430)
AmpR promoter
T7Te terminator
SexAI* (3083)
SfiI (3269)
PasI (3432)
PciI (3596)
KflI (3627)
BstZ17I (3779)
PspOMI* (3835)
ApaI* (3839)
AgeI (4211)
Acc65I (4214)
KpnI (4218)
BglII (4546)
SV40 NLS
PacI (5467)
PshAI (5759)
RsrII (6140)
AsiSI (6096)
SV40 poly(A) signal
SpeI (7541)
SnaBI (7882)
BlpI (8363)
BspQI - SapI (8660)
AflII (8876)
PvuII (8907)
MauBI (9005)
HindIII (9090)
SV40 poly(A) signal
ori
AmpR
SV40 promoter
SV40 ori
SV40 poly(A) signal
small t intron
SV40 promoter
HygR
CMV enhancer
CMV promoter
1000
2000
3000
4000
5000
6000
7000
8000
9000

FIG. 14

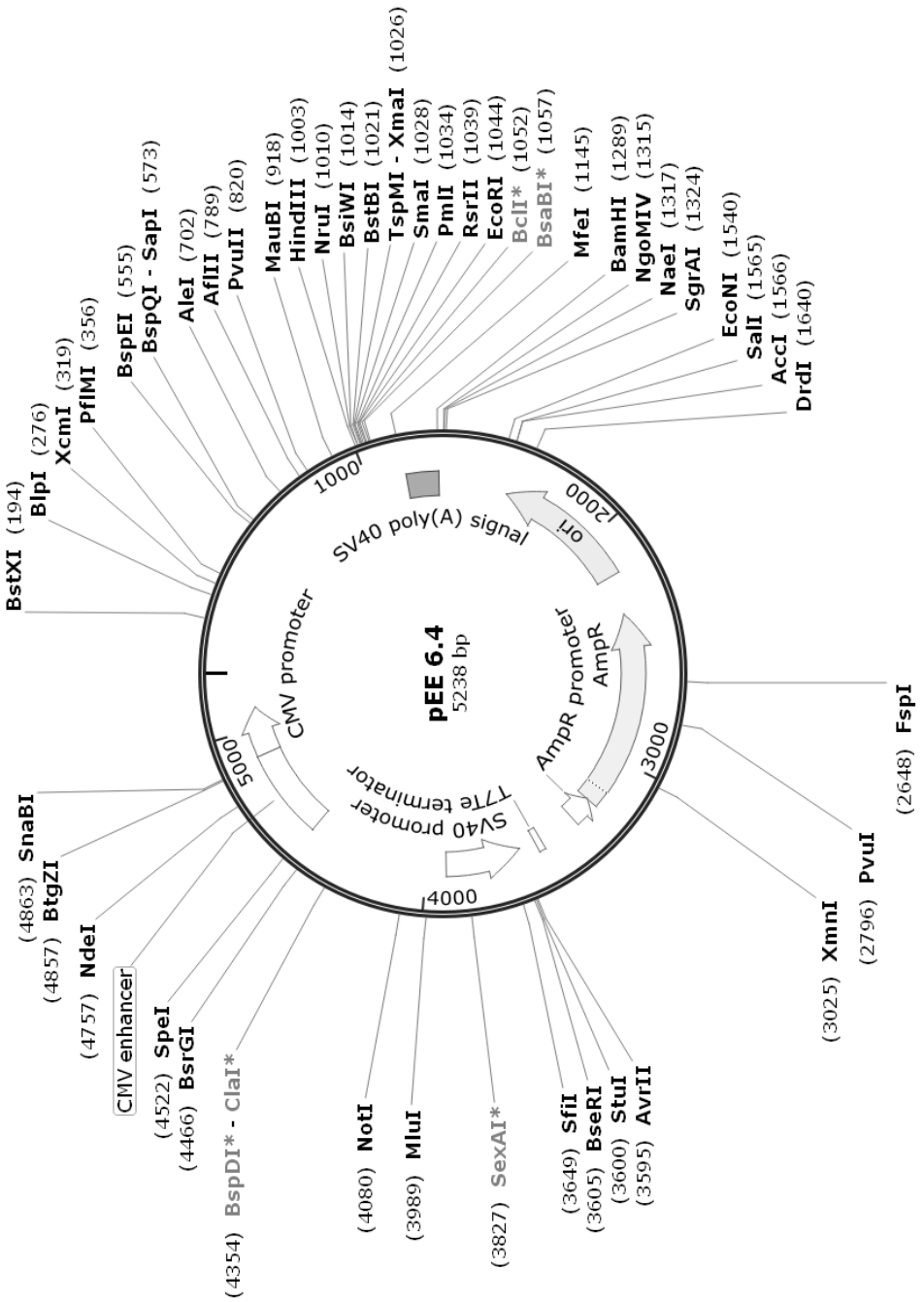

pEE 6.4
5238 bp
1000
2000
3000
4000
5000
SV40 poly(A) signal
ori
AmpR promoter
AmpR
SV40 promoter
T7Te terminator
CMV promoter
CMV enhancer
BstXI (194)
BlpI (276)
XcmI (319)
PflMI (356)
BspEI (555)
BspQI - SapI (573)
AleI (702)
AflII (789)
PvuII (820)
MauBI (918)
HindIII (1003)
NruI (1010)
BsiWI (1014)
BstBI (1021)
TspMI - XmaI (1026)
SmaI (1028)
PmlI (1034)
RsrII (1039)
EcoRI (1044)
BclI* (1052)
BsaBI* (1057)
MfeI (1145)
BamHI (1289)
NgoMIV (1315)
NaeI (1317)
SgrAI (1324)
EcoNI (1540)
SalI (1565)
AccI (1566)
DrdI (1640)
(2648) FspI
(2796) PvuI
(3025) XmnI
(3595) AvrII
(3600) StuI
(3605) BseRI
(3649) SfiI
(3827) SexAI*
(3989) MluI
(4080) NotI
(4354) BspDI* - ClaI*
(4466) BsrGI
(4522) SpeI
(4757) NdeI
(4857) BtgZI
(4863) SnaBI

FIG. 15