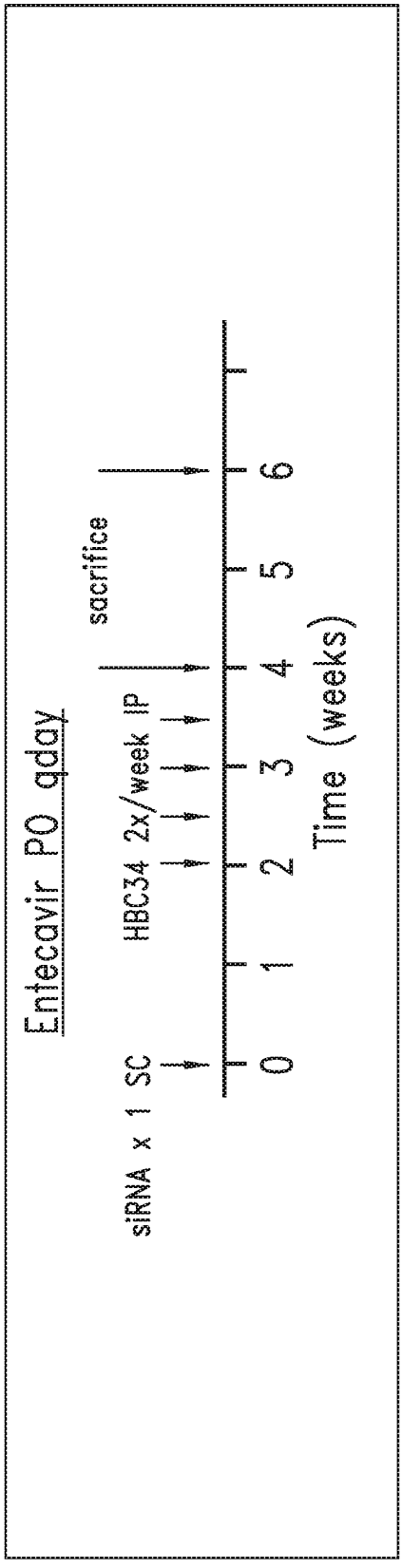

Entecavir PO qday
siRNA x 1 SC
HBC34 2x/week IP
sacrifice
0 1 2 3 4 5 6
Time (weeks)

FIG. 1

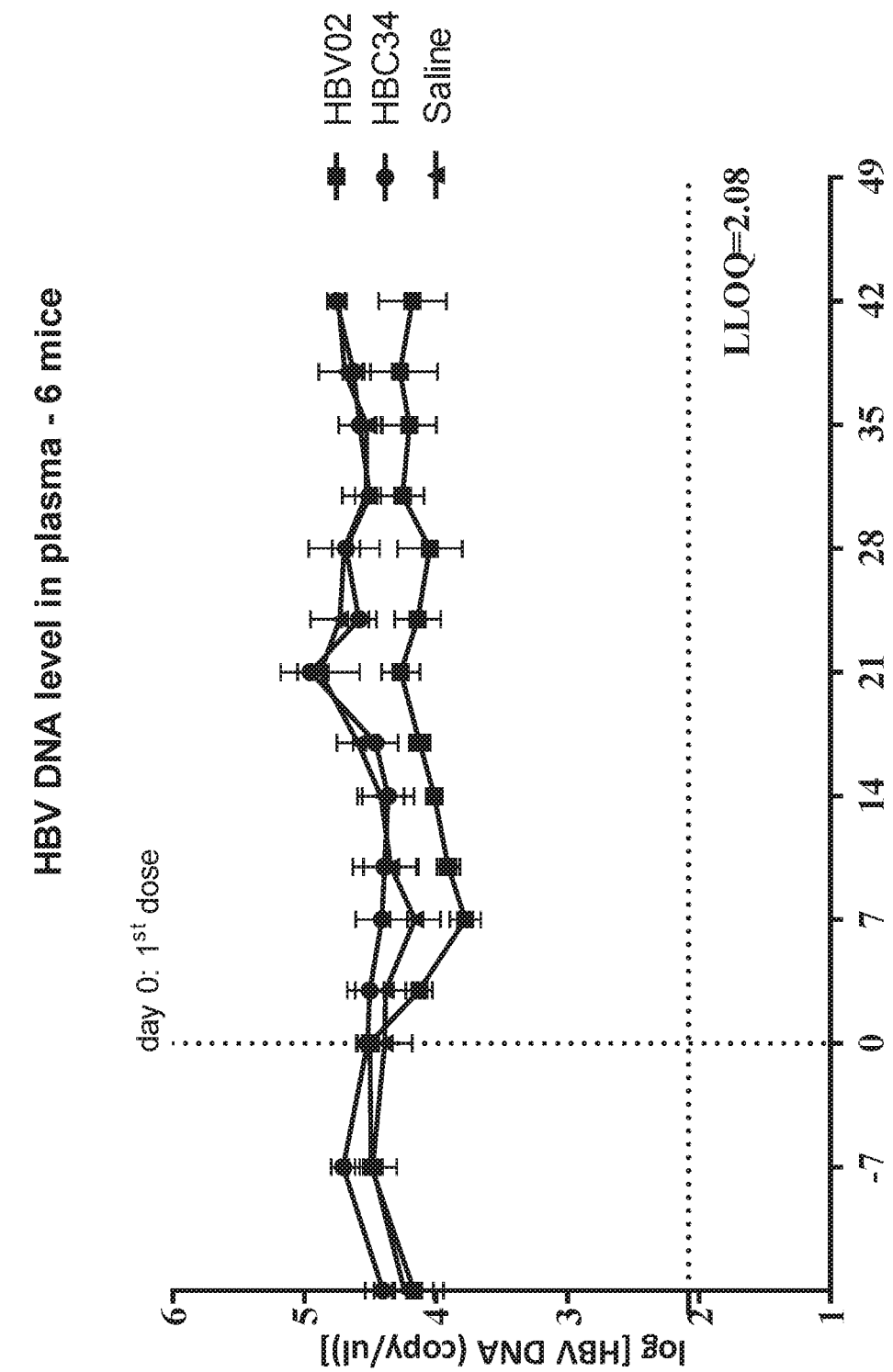

HBV DNA level in plasma - 6 mice
HBV02
HBC34
Saline
day 0: 1st dose
LLOQ=2.08
log [HBV DNA (copy/ul)]
day post-first dose
6
5
4
3
2
1
-7
0
7
14
21
28
35
42
49

FIG. 2A

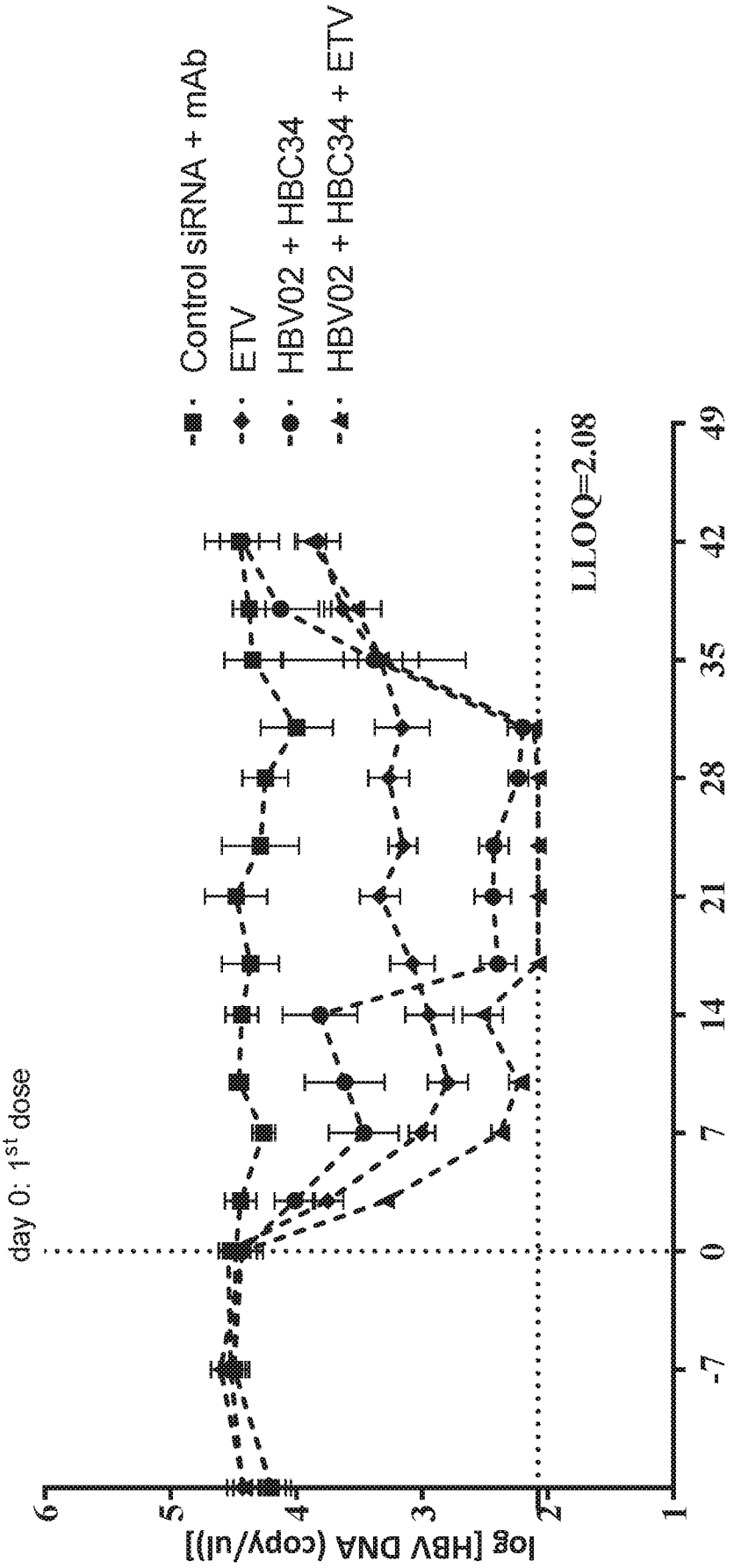

HBV DNA level in plasma - 6 mice
Control siRNA + mAb
ETV
HBV02 + HBC34
HBV02 + HBC34 + ETV
day 0: 1st dose
LLOQ=2.08
log [HBV DNA (copy/ul)]
day post-first dose
6
5
4
3
2
1
-7
0
7
14
21
28
35
42
49

FIG. 2B

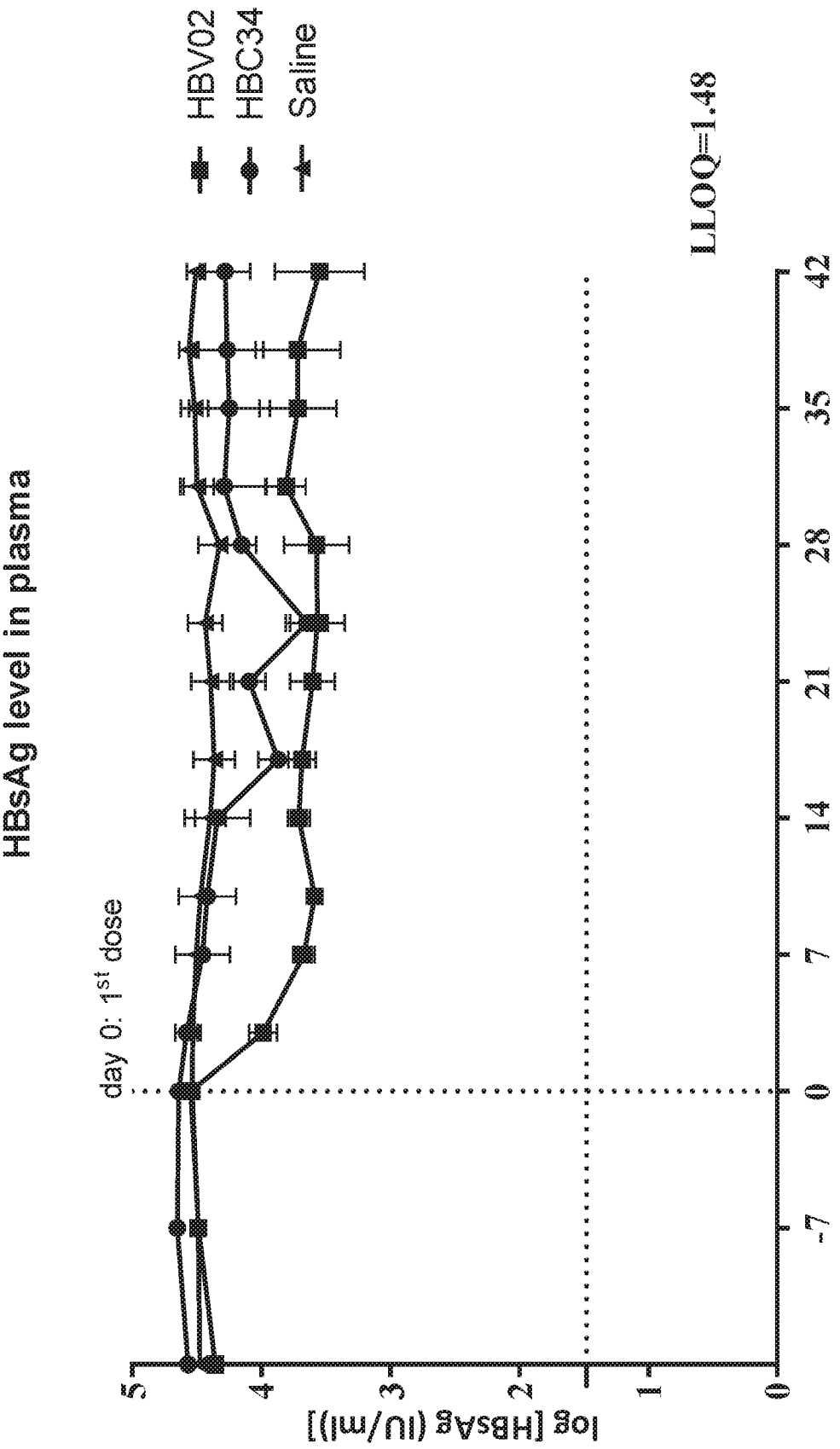

HBsAg level in plasma
HBV02
HBC34
Saline
LLOQ=1.48
day 0: 1st dose
log [HBsAg (IU/ml)]
day post-first dose
5
4
3
2
1
0
-7
0
7
14
21
28
35
42

FIG. 3A

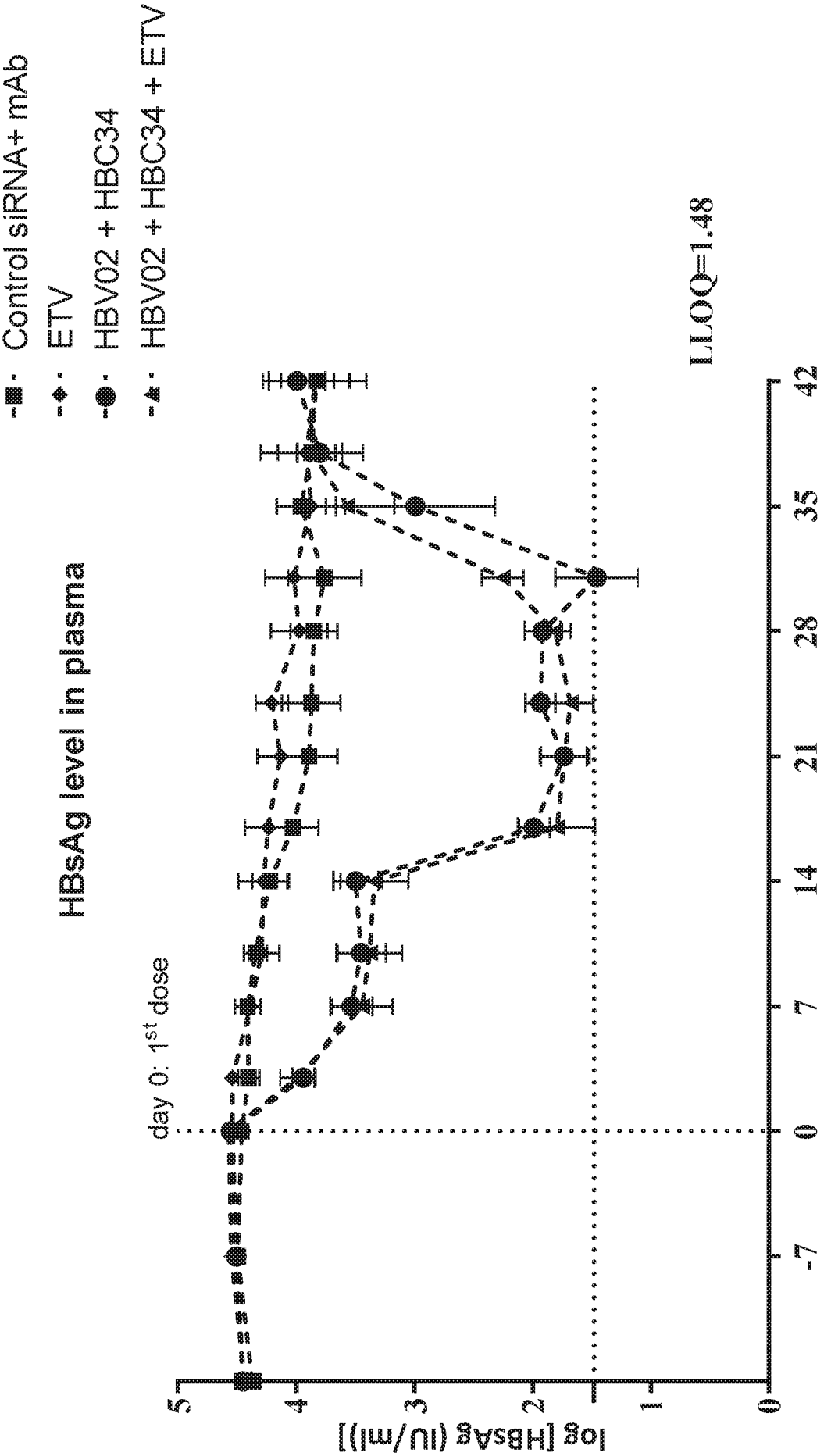

Control siRNA+ mAb
ETV
HBV02 + HBC34
HBV02 + HBC34 + ETV
HBsAg level in plasma
LLOQ=1.48
day 0: 1st dose
log [HBsAg (IU/ml)]
day post-first dose
5
4
3
2
1
0
-7
0
7
14
21
28
35
42

FIG. 3B

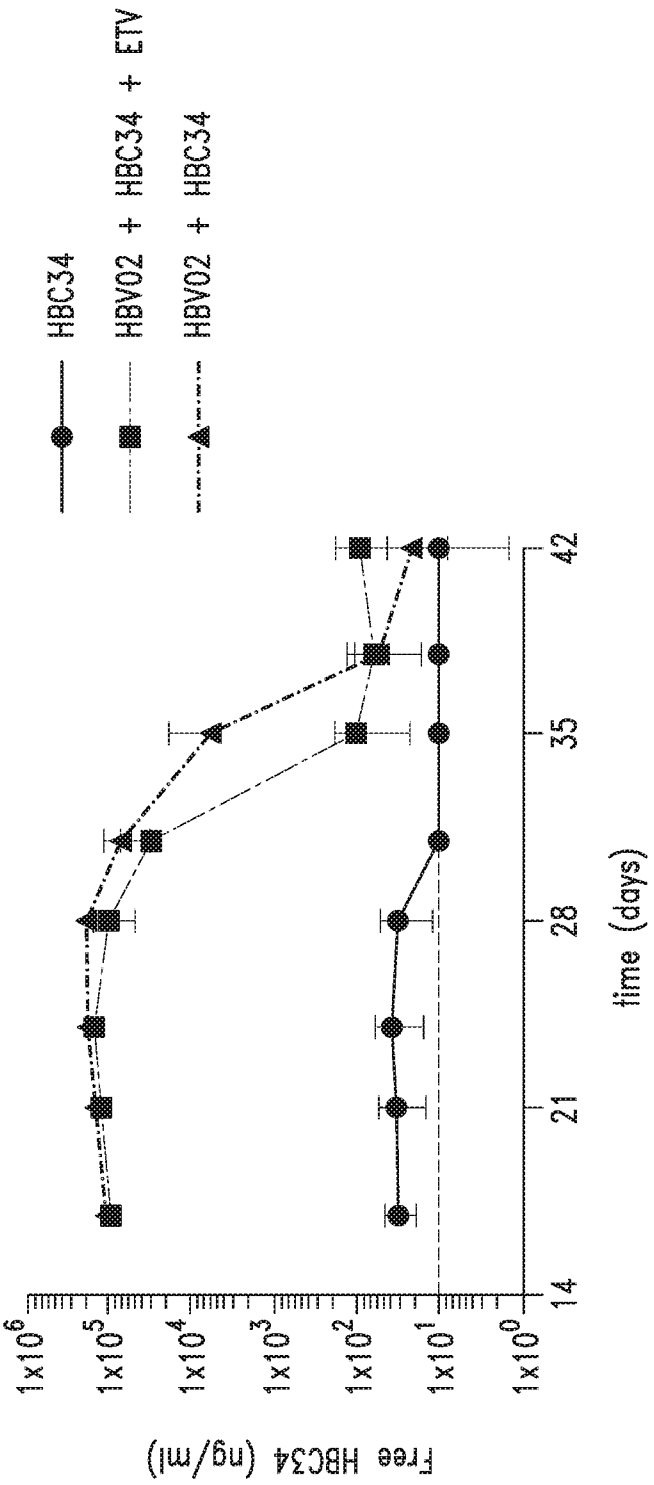

HBC34
HBV02 + HBC34 + ETV
HBV02 + HBC34
Free HBC34 (ng/ml)
1x10^6
1x10^5
1x10^4
1x10^3
1x10^2
1x10^1
1x10^0
14
21
28
35
42
time (days)

FIG. 4

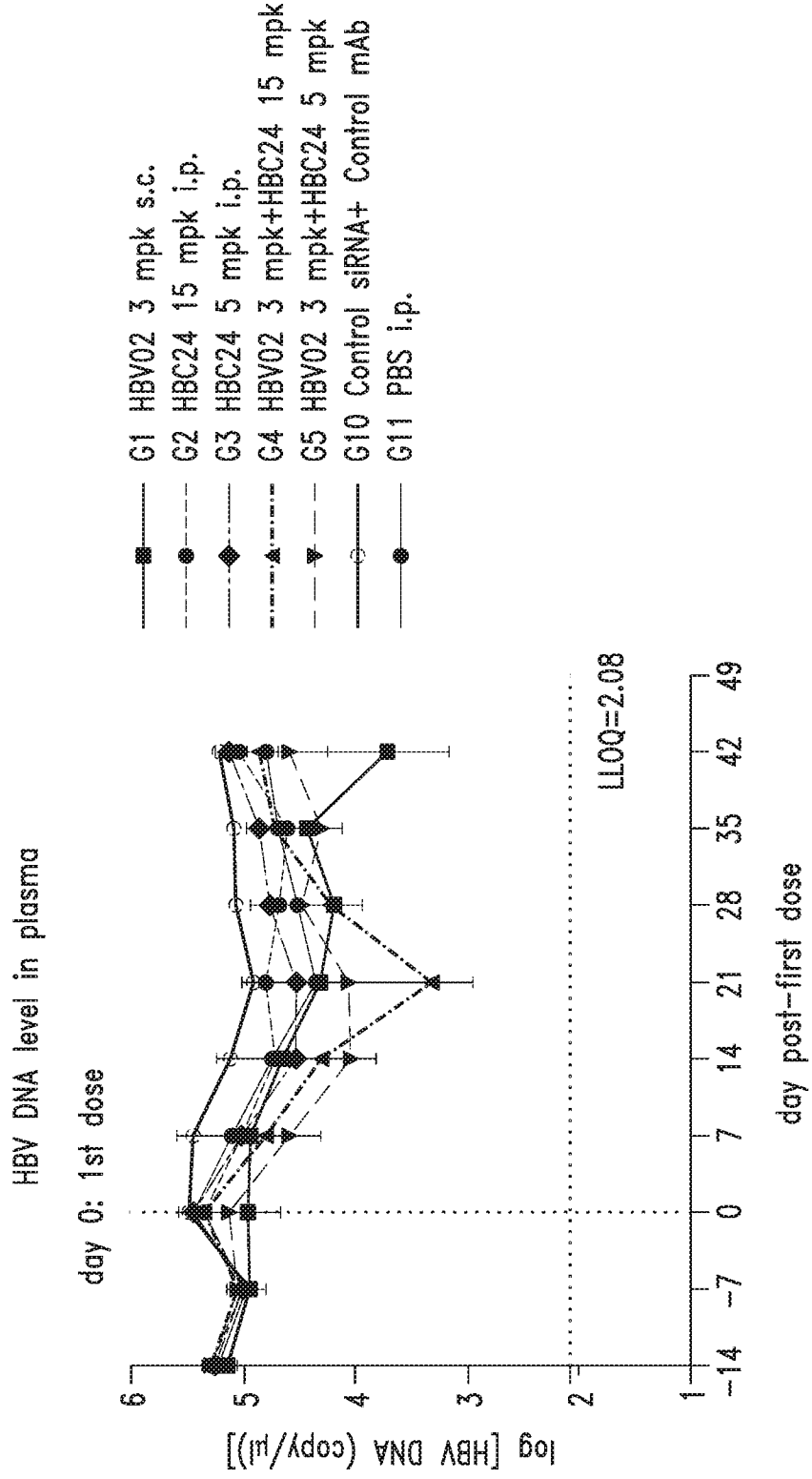

HBV DNA level in plasma
day 0: 1st dose
log [HBV DNA (copy/ml)]
6
5
4
3
2
1
LLOQ=2.08
-14
-7
0
7
14
21
28
35
42
49
day post-first dose
G1 HBV02 3 mpk s.c.
G2 HBC24 15 mpk i.p.
G3 HBC24 5 mpk i.p.
G4 HBV02 3 mpk+HBC24 15 mpk
G5 HBV02 3 mpk+HBC24 5 mpk
G10 Control siRNA+ Control mAb
G11 PBS i.p.

FIG. 5A

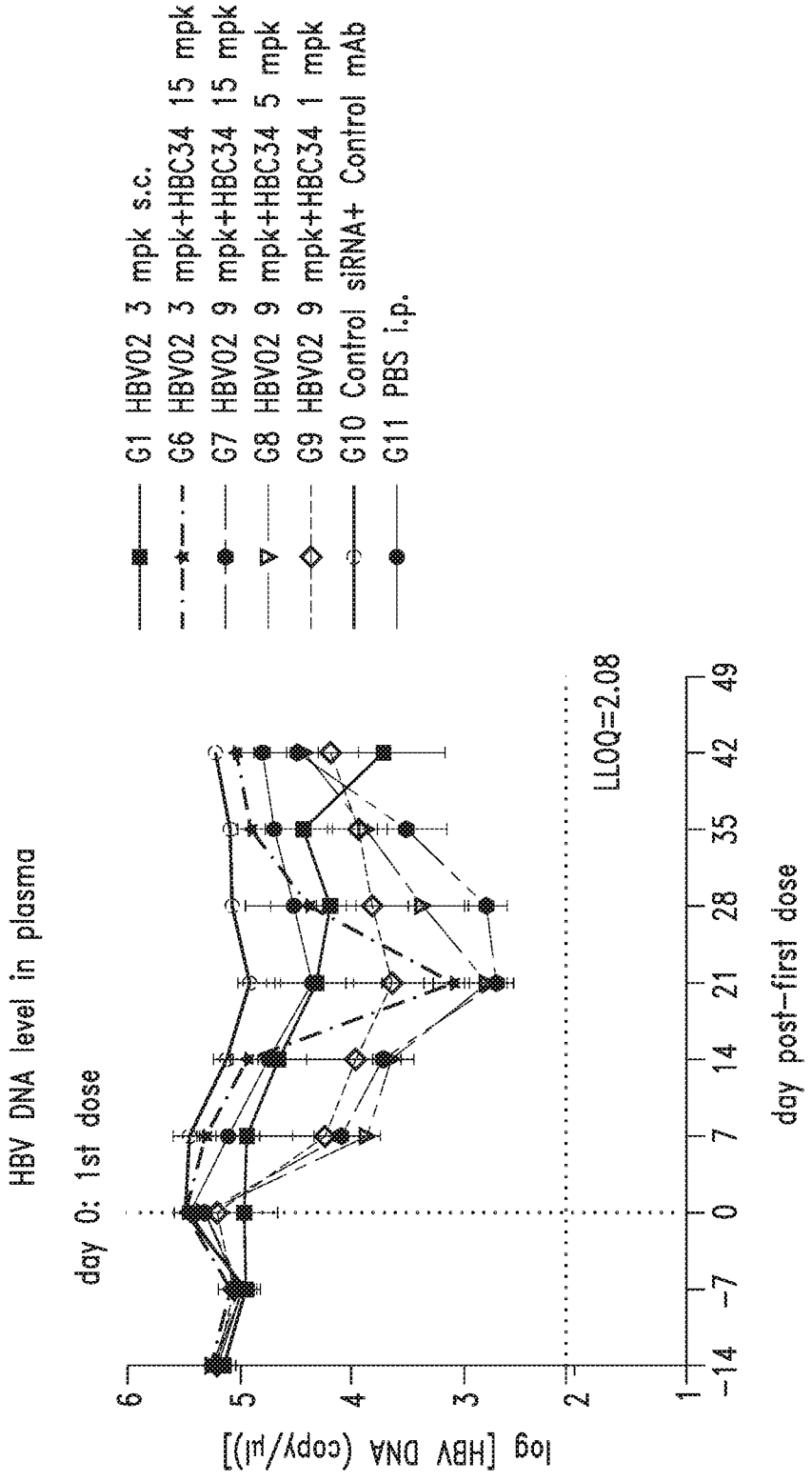

HBV DNA level in plasma
day 0: 1st dose
log [HBV DNA (copy/ul)]
6
5
4
3
2
1
LLOQ=2.08
-14
-7
0
7
14
21
28
35
42
49
day post-first dose
G1 HBV02 3 mpk s.c.
G6 HBV02 3 mpk+HBC34 15 mpk
G7 HBV02 9 mpk+HBC34 15 mpk
G8 HBV02 9 mpk+HBC34 5 mpk
G9 HBV02 9 mpk+HBC34 1 mpk
G10 Control siRNA+ Control mAb
G11 PBS i.p.

FIG. 5B

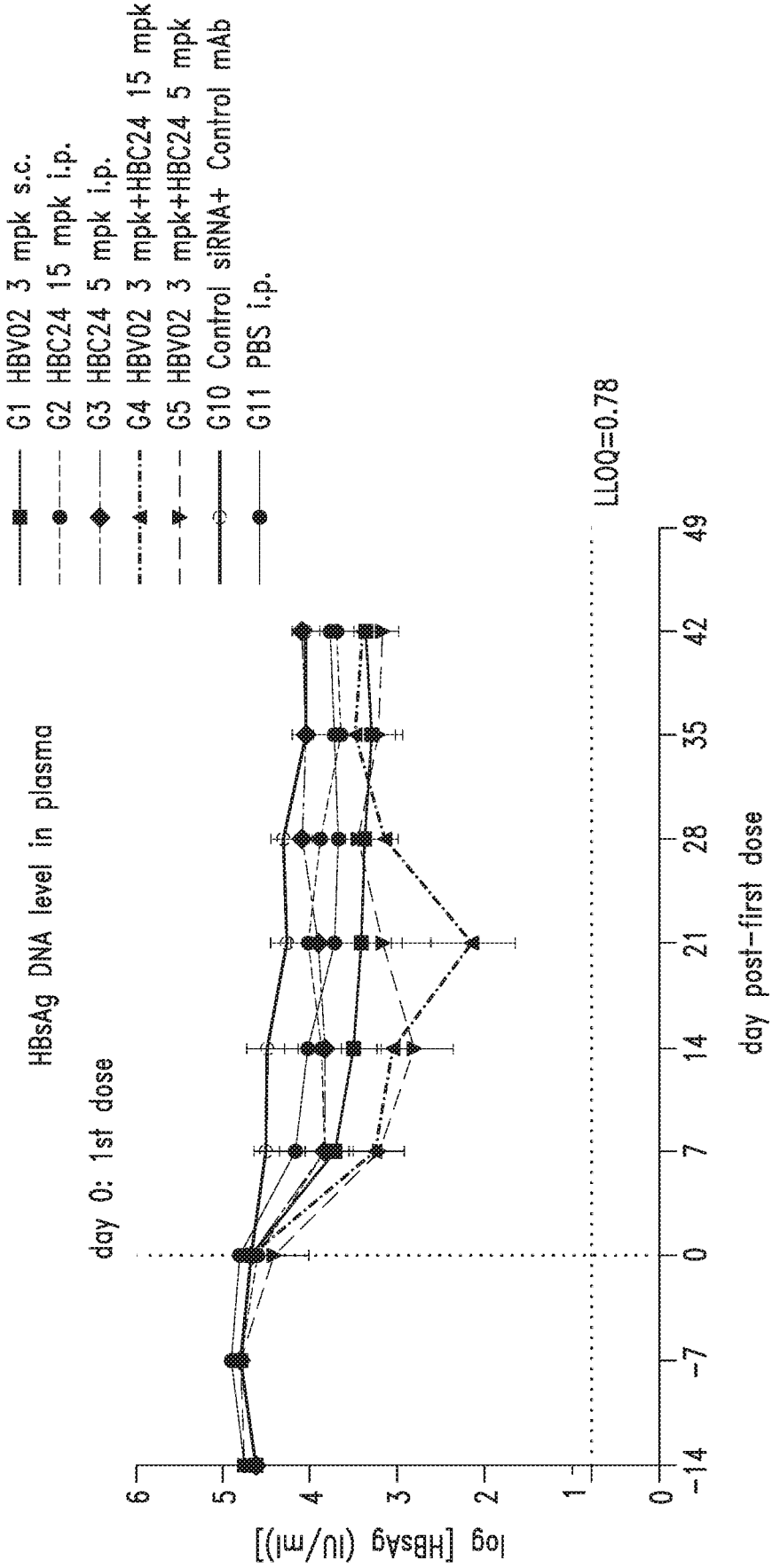

HBsAg DNA level in plasma
day 0: 1st dose
G1 HBV02 3 mpk s.c.
G2 HBC24 15 mpk i.p.
G3 HBC24 5 mpk i.p.
G4 HBV02 3 mpk+HBC24 15 mpk
G5 HBV02 3 mpk+HBC24 5 mpk
G10 Control siRNA+ Control mAb
G11 PBS i.p.
LLOQ=0.78
log [HBsAg (IU/ml)]
6
5
4
3
2
1
0
-14
-7
0
7
14
21
28
35
42
49
day post-first dose

FIG. 6A

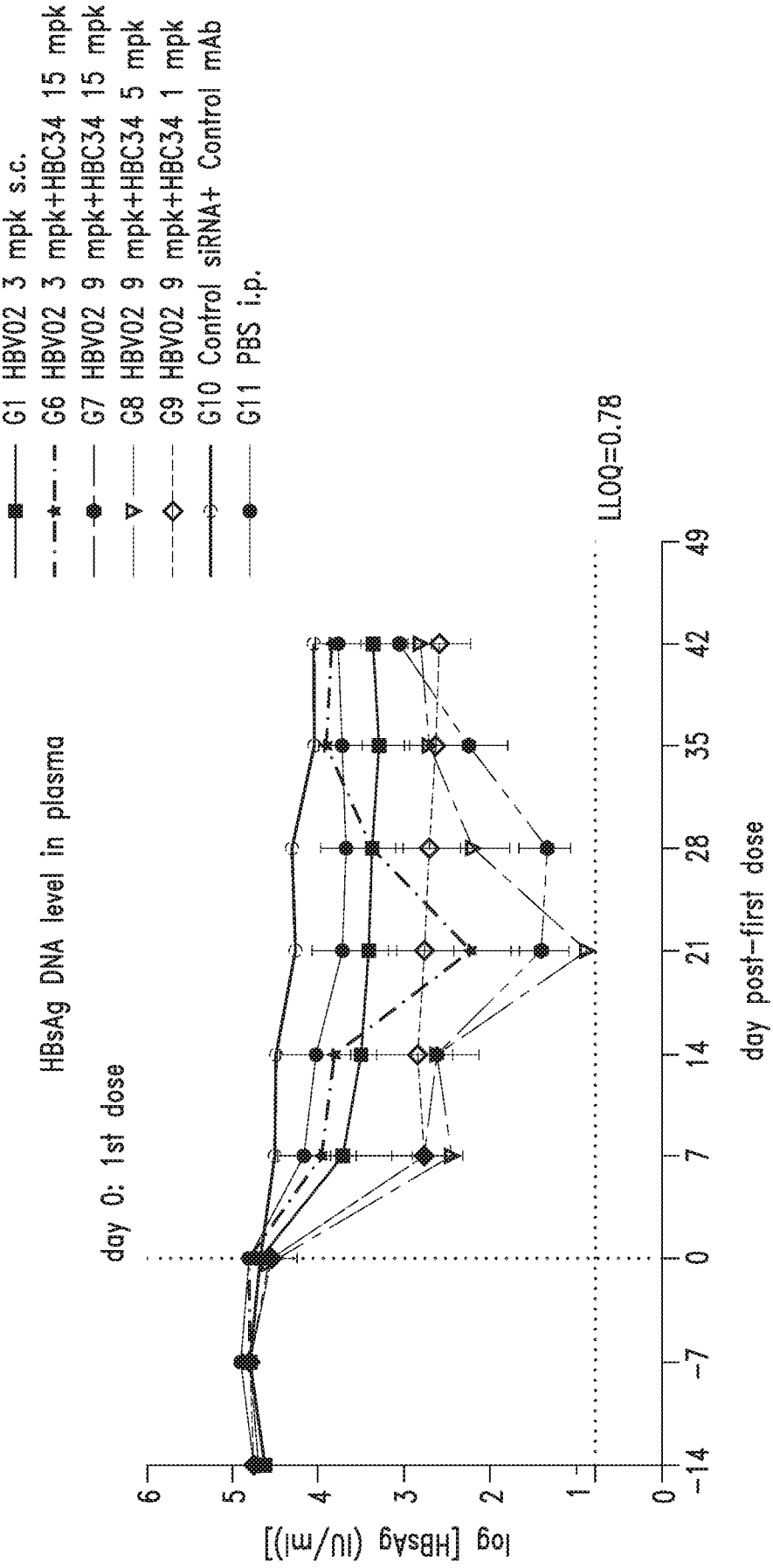

HBsAg DNA level in plasma
G1 HBV02 3 mpk s.c.
G6 HBV02 3 mpk+HBC34 15 mpk
G7 HBV02 9 mpk+HBC34 15 mpk
G8 HBV02 9 mpk+HBC34 5 mpk
G9 HBV02 9 mpk+HBC34 1 mpk
G10 Control siRNA+ Control mAb
G11 PBS i.p.
day 0: 1st dose
LLOQ=0.78
log [HBsAg (IU/ml)]
day post-first dose
6
5
4
3
2
1
0
-14
-7
0
7
14
21
28
35
42
49

FIG. 6B

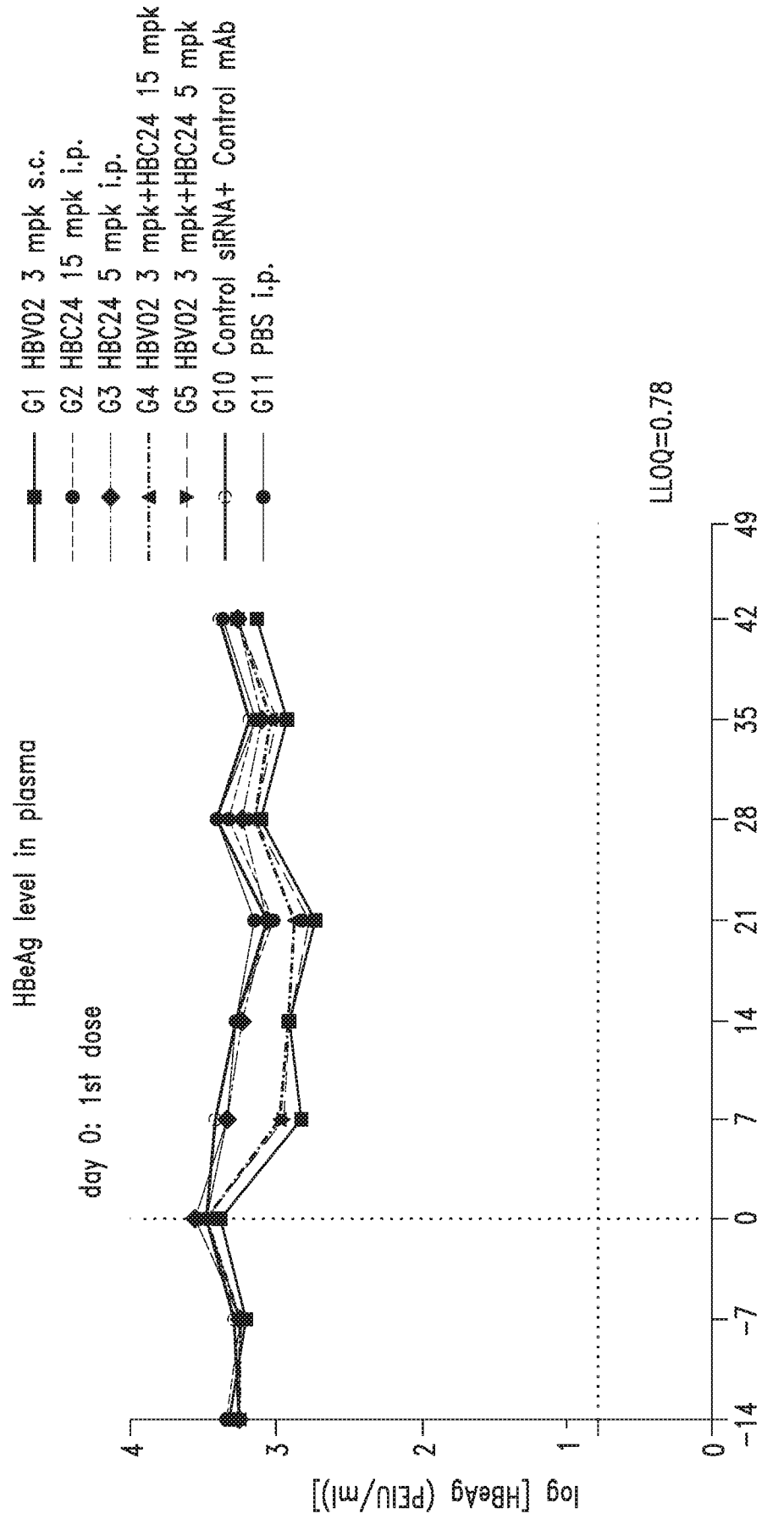

HBeAg level in plasma
day 0: 1st dose
G1 HBV02 3 mpk s.c.
G2 HBC24 15 mpk i.p.
G3 HBC24 5 mpk i.p.
G4 HBV02 3 mpk+HBC24 15 mpk
G5 HBV02 3 mpk+HBC24 5 mpk
G10 Control siRNA+ Control mAb
G11 PBS i.p.
LLOQ=0.78
log [HBeAg (PEIU/ml)]
day post-first dose
4
3
2
1
0
-14
-7
0
7
14
21
28
35
42
49

FIG. 7A

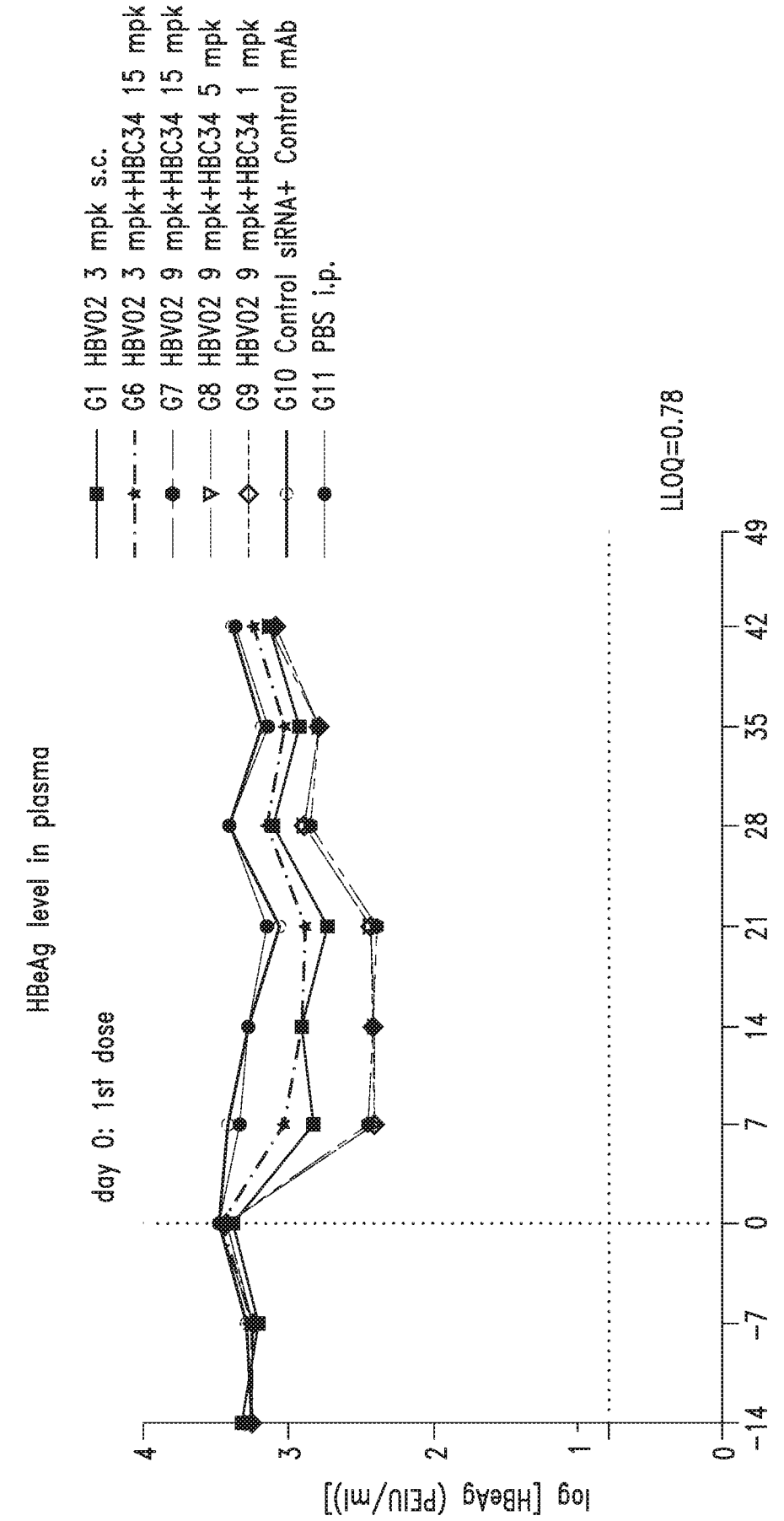

HBeAg level in plasma
day 0: 1st dose
G1 HBV02 3 mpk s.c.
G6 HBV02 3 mpk+HBC34 15 mpk
G7 HBV02 9 mpk+HBC34 15 mpk
G8 HBV02 9 mpk+HBC34 5 mpk
G9 HBV02 9 mpk+HBC34 1 mpk
G10 Control siRNA+ Control mAb
G11 PBS i.p.
LLOQ=0.78
log [HBeAg (PEIU/ml)]
day post-first dose
4
3
2
1
0
-14
-7
0
7
14
21
28
35
42
49

FIG. 7B

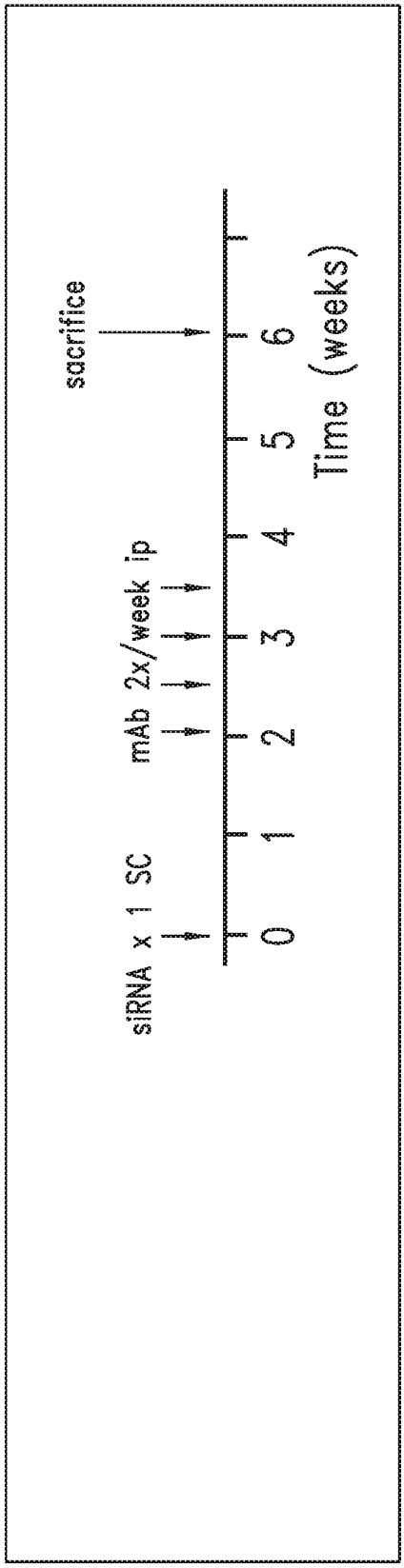


FIG. 8

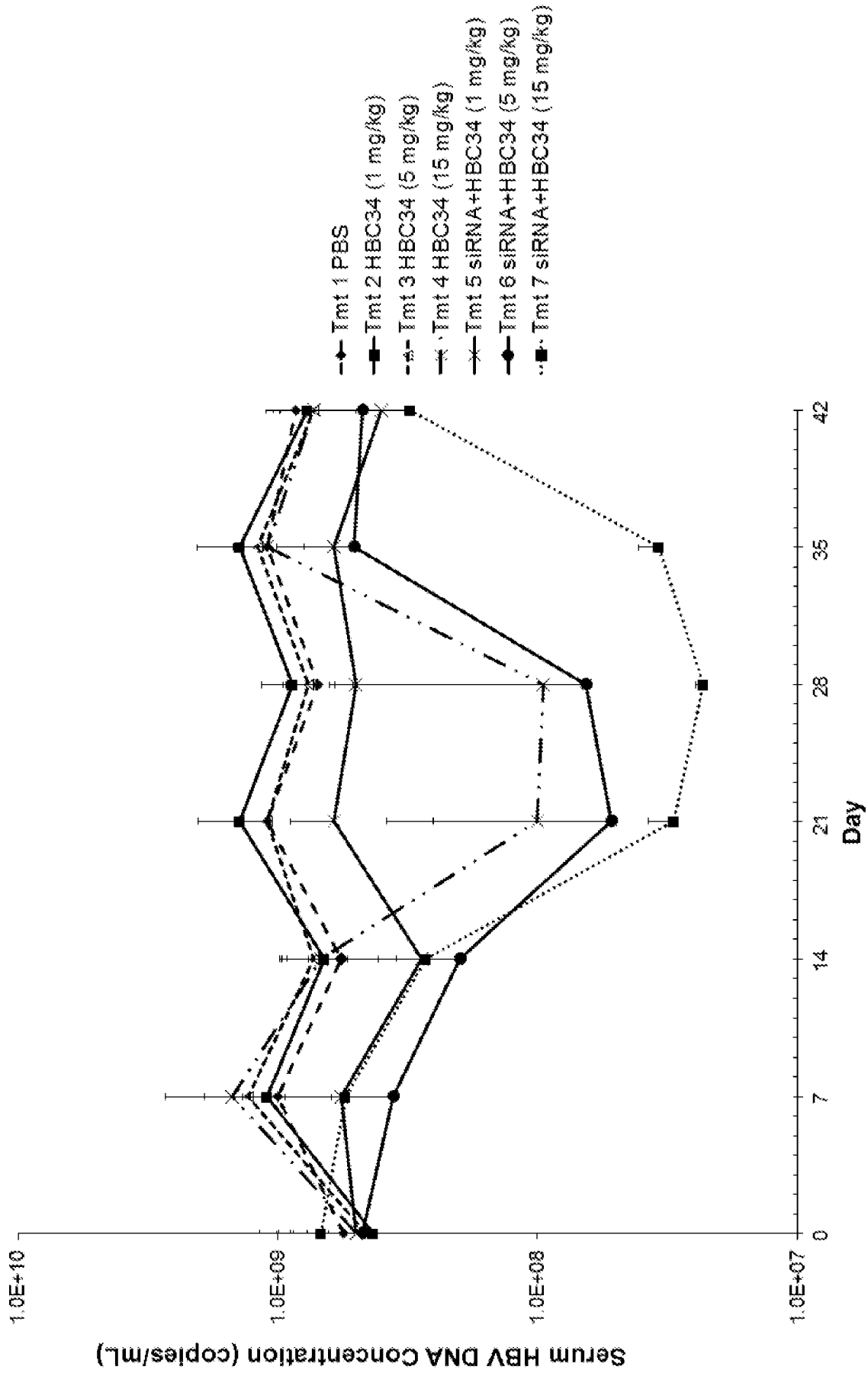

Tmt 1 PBS
Tmt 2 HBC34 (1 mg/kg)
Tmt 3 HBC34 (5 mg/kg)
Tmt 4 HBC34 (15 mg/kg)
Tmt 5 siRNA+HBC34 (1 mg/kg)
Tmt 6 siRNA+HBC34 (5 mg/kg)
Tmt 7 siRNA+HBC34 (15 mg/kg)
Serum HBV DNA Concentration (copies/mL)
1.0E+10
1.0E+09
1.0E+08
1.0E+07
0
7
14
21
28
35
42
Day

FIG. 9

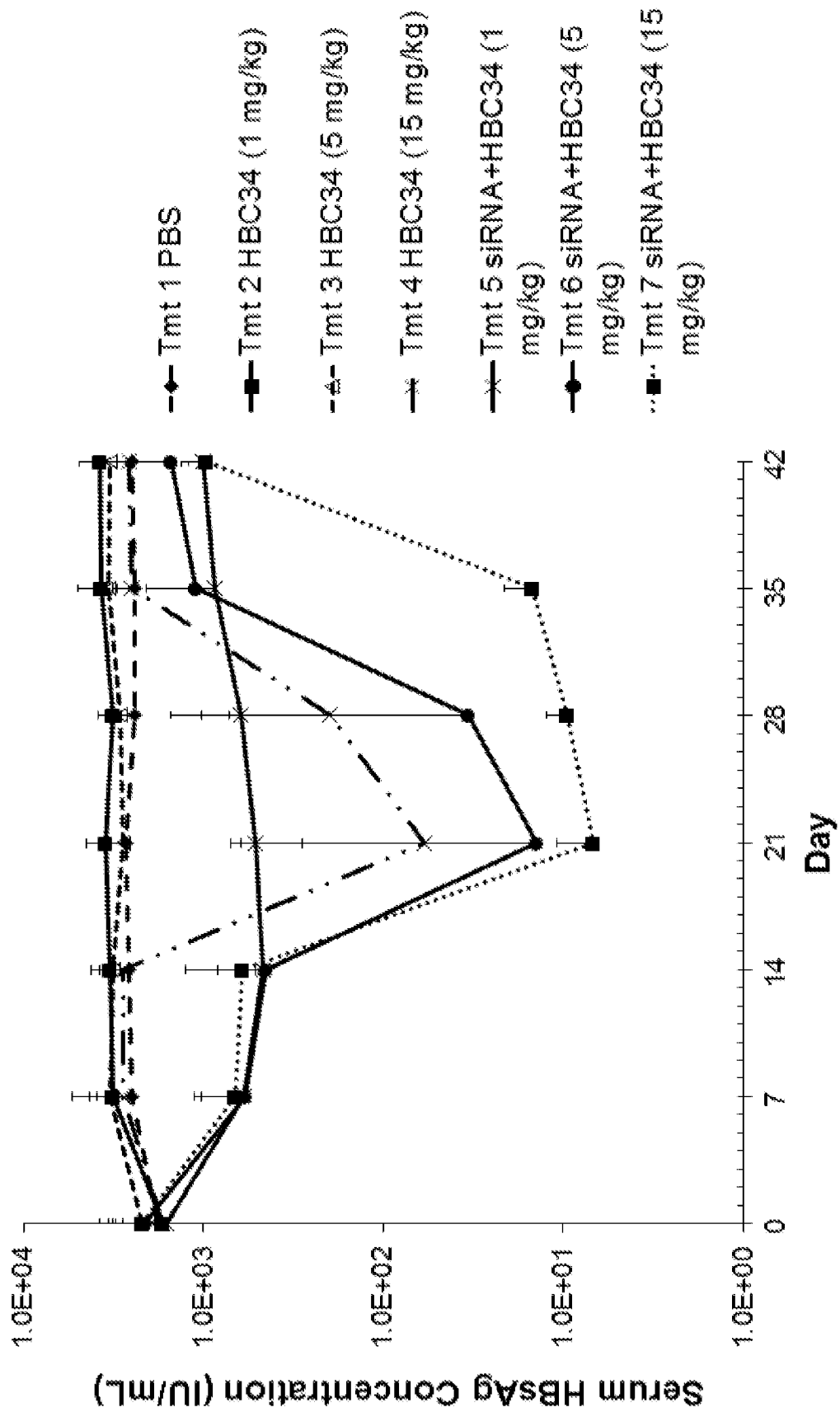

Tmt 1 PBS
Tmt 2 HBC34 (1 mg/kg)
Tmt 3 HBC34 (5 mg/kg)
Tmt 4 HBC34 (15 mg/kg)
Tmt 5 siRNA+HBC34 (1 mg/kg)
Tmt 6 siRNA+HBC34 (5 mg/kg)
Tmt 7 siRNA+HBC34 (15 mg/kg)
Serum HBsAg Concentration (IU/mL)
1.0E+04
1.0E+03
1.0E+02
1.0E+01
1.0E+00
0
7
14
21
28
35
42
Day

FIG. 10

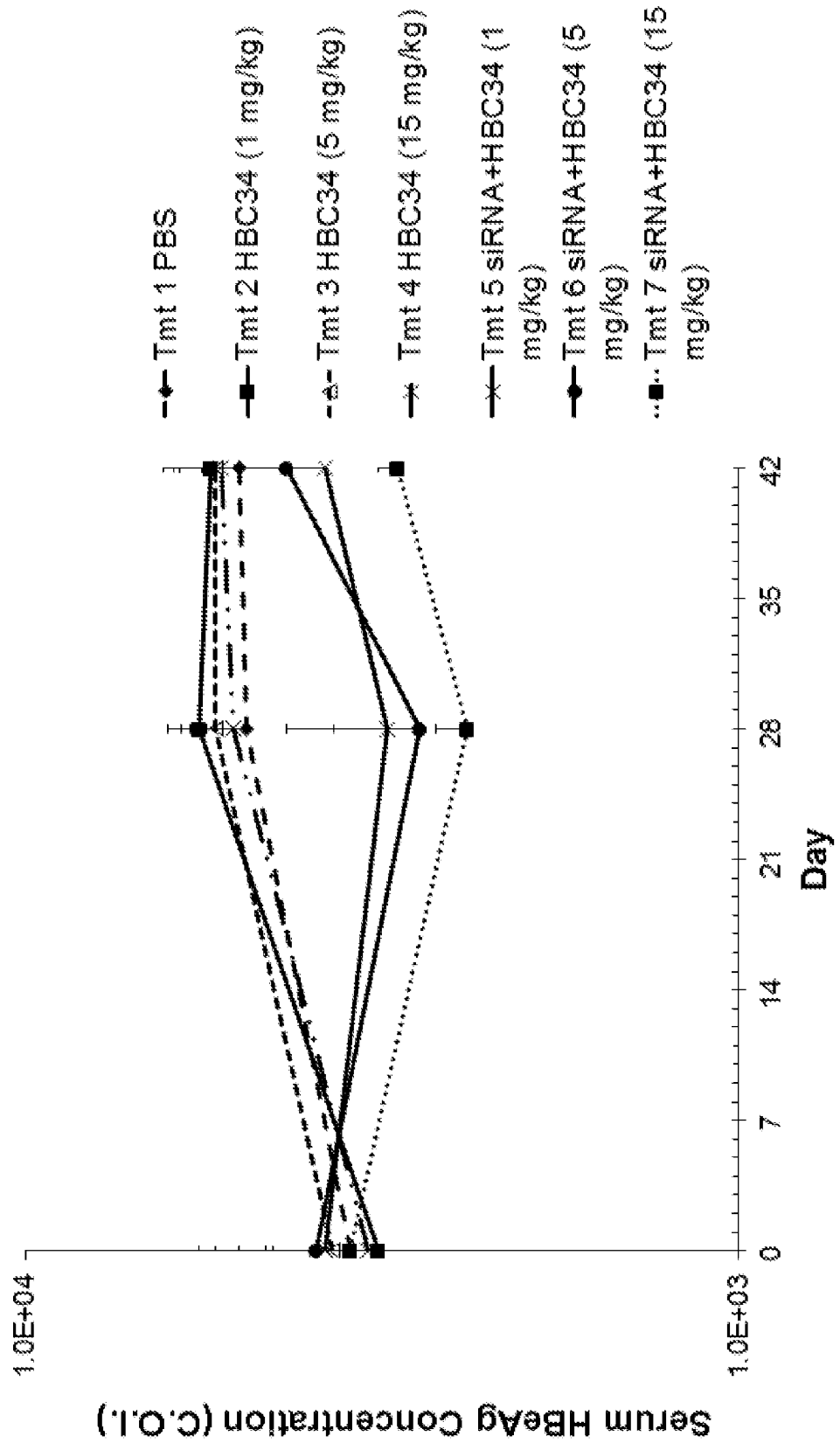

Tmt 1 PBS
Tmt 2 HBC34 (1 mg/kg)
Tmt 3 HBC34 (5 mg/kg)
Tmt 4 HBC34 (15 mg/kg)
Tmt 5 siRNA+HBC34 (1 mg/kg)
Tmt 6 siRNA+HBC34 (5 mg/kg)
Tmt 7 siRNA+HBC34 (15 mg/kg)
Serum HBeAg Concentration (C.O.I.)
1.0E+04
1.0E+03
0
7
14
21
28
35
42
Day

FIG. 11

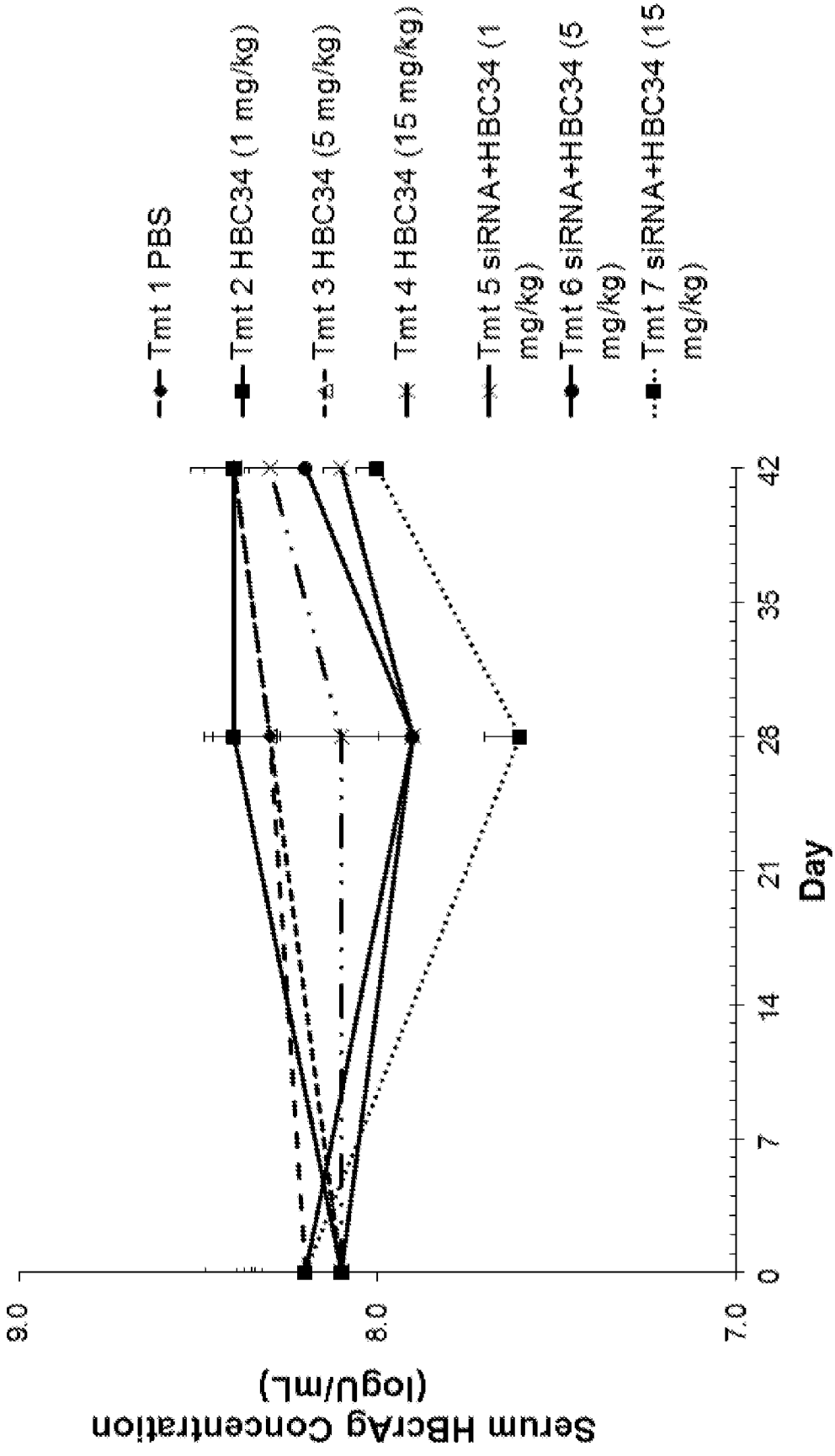

Tmt 1 PBS
Tmt 2 HBC34 (1 mg/kg)
Tmt 3 HBC34 (5 mg/kg)
Tmt 4 HBC34 (15 mg/kg)
Tmt 5 siRNA+HBC34 (1 mg/kg)
Tmt 6 siRNA+HBC34 (5 mg/kg)
Tmt 7 siRNA+HBC34 (15 mg/kg)
Serum HBcrAg Concentration (logU/mL)
9.0
8.0
7.0
0
7
14
21
28
35
42
Day

FIG. 12

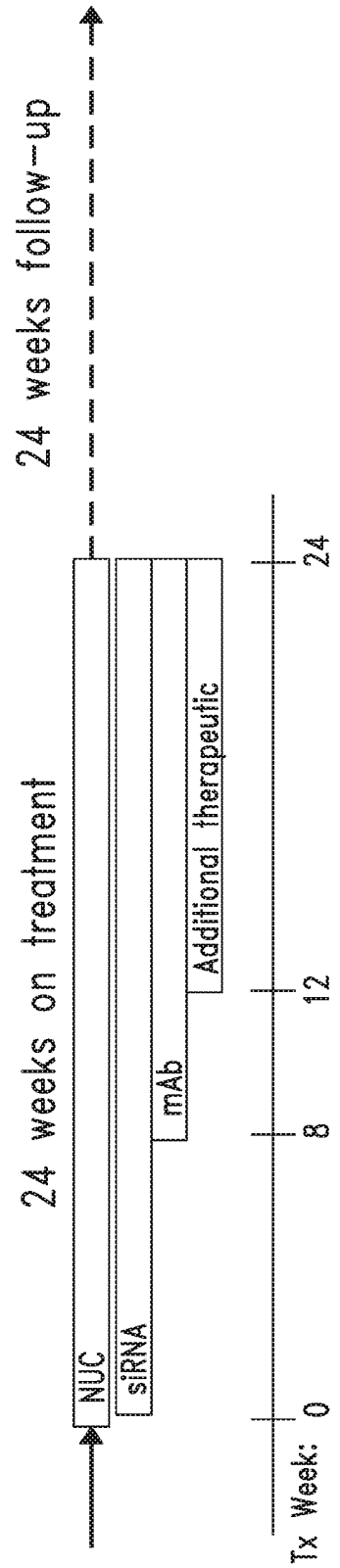

24 weeks follow-up
24 weeks on treatment
NUC
siRNA
mAb
Additional therapeutic
Tx Week: 0
8
12
24

FIG. 13