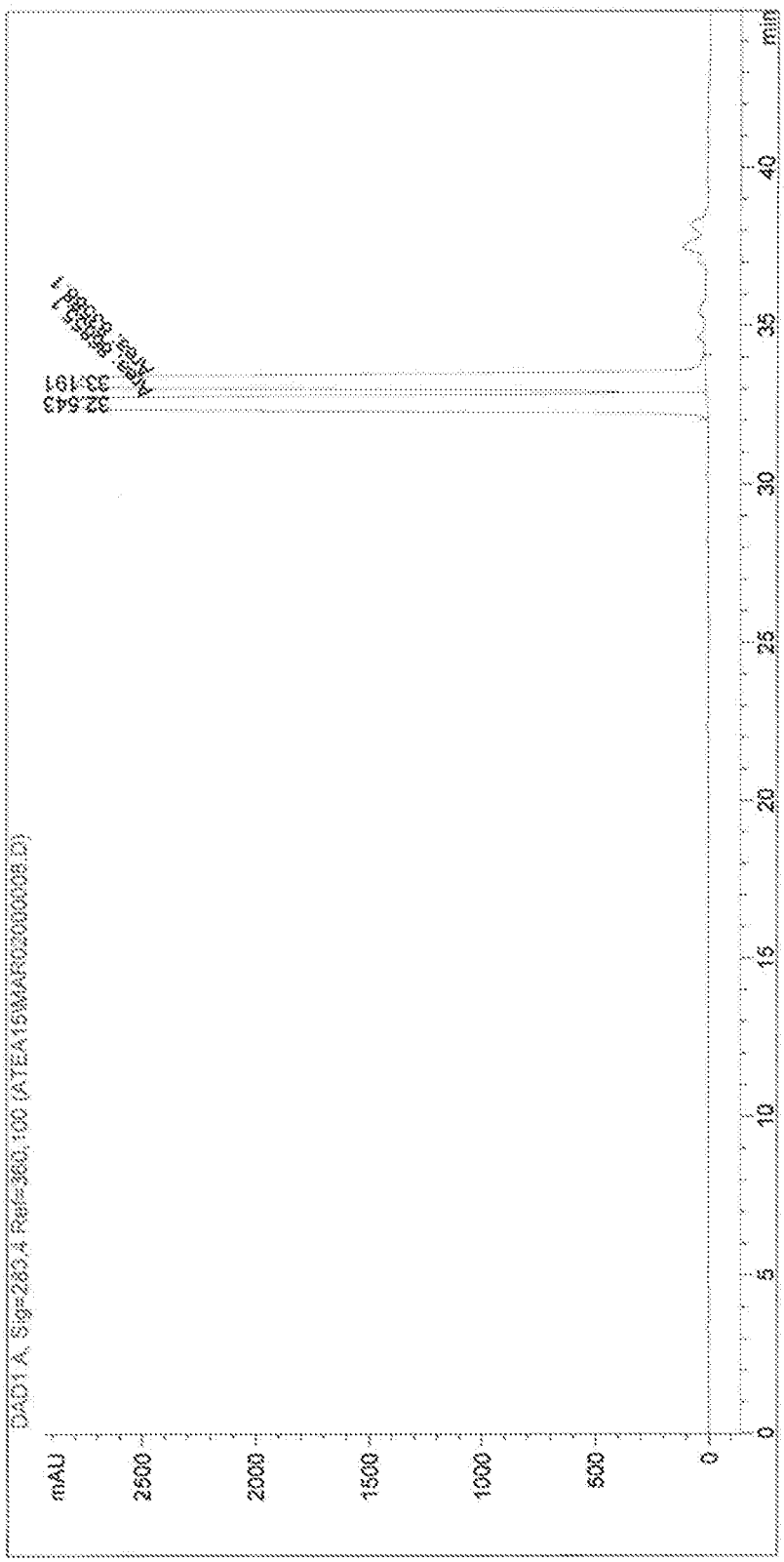

DAD1 A, Sig=280,4 Ref=360,100 (ATEA15MAR2015000009.D)
mAU
2500
2000
1500
1000
500
0
0
5
10
15
20
25
30
35
40
min

FIG. 1

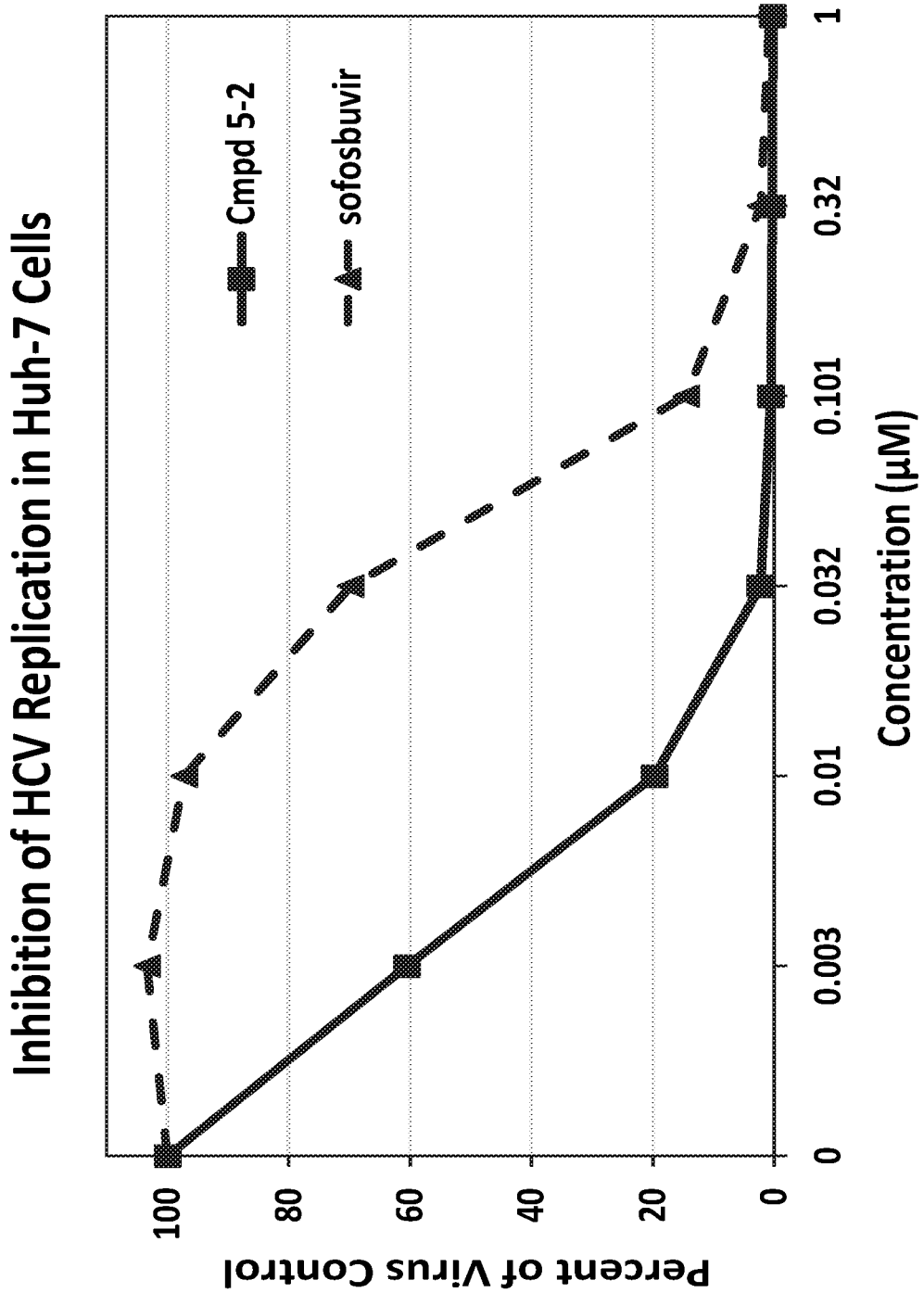

Inhibition of HCV Replication in Huh-7 Cells
Cmpd 5-2
sofosbuvir
Percent of Virus Control
100
80
60
40
20
0
0
0.003
0.01
0.032
0.101
0.32
1
Concentration (μM)

FIG. 2

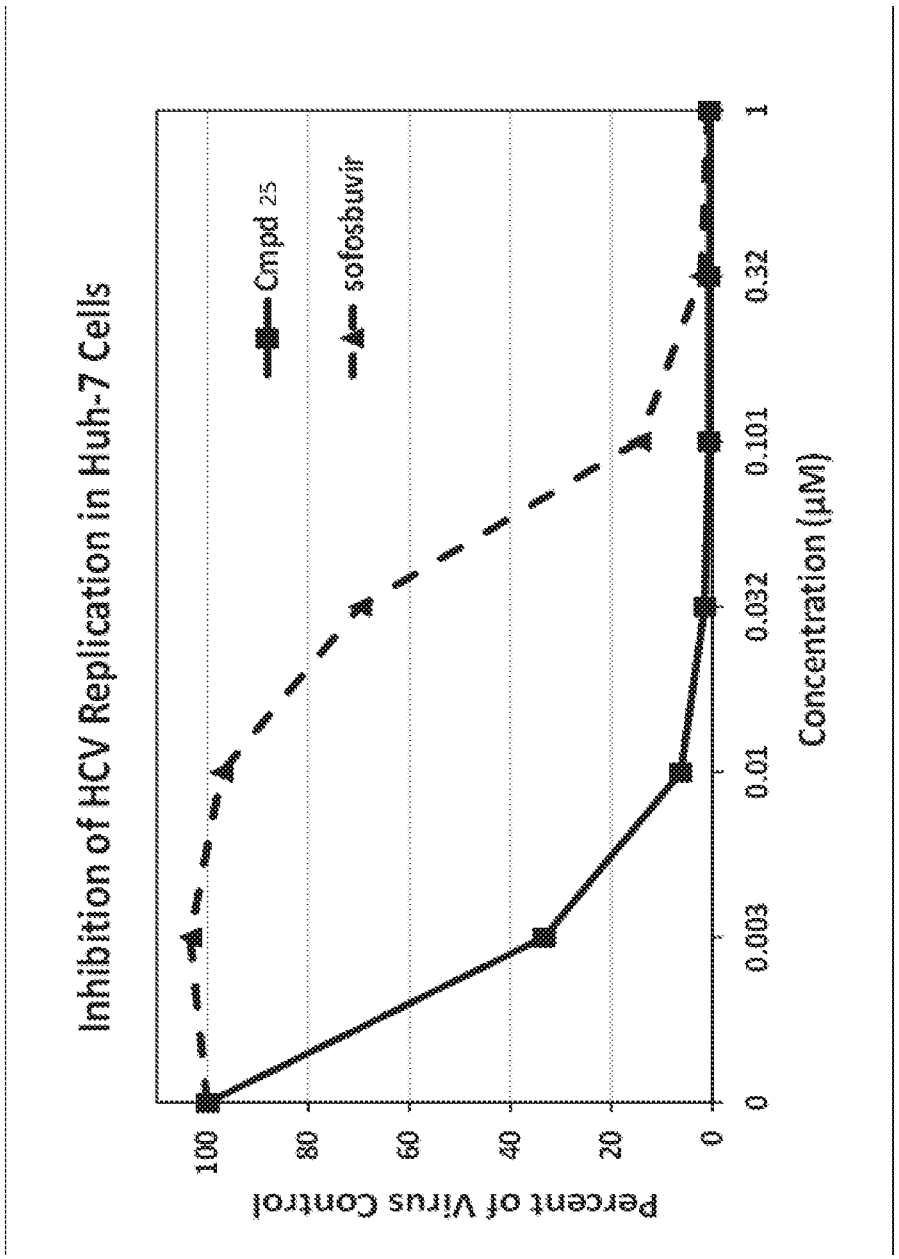

Inhibition of HCV Replication in Huh-7 Cells
Cmpd 25
sofosbuvir
Percent of Virus Control
100
80
60
40
20
0
0
0.003
0.01
0.032
0.101
0.32
1
Concentration (μM)

FIG. 3

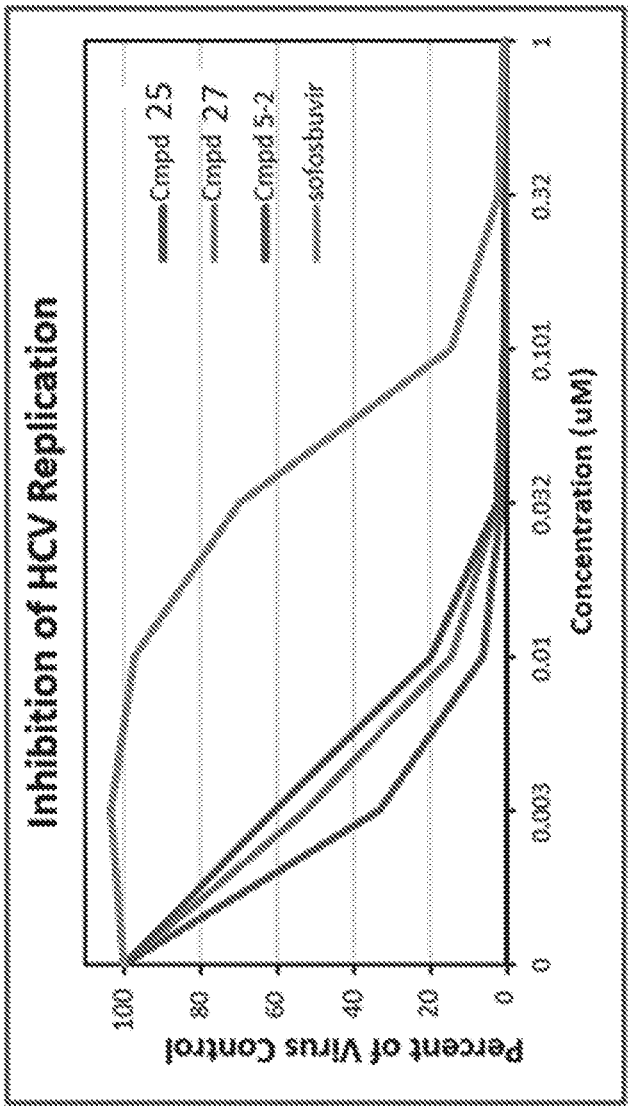

Inhibition of HCV Replication
Cmpd 25
Cmpd 27
Cmpd 5-2
sofosbuvir
Percent of Virus Control
100
80
60
40
20
0
0
0.003
0.01
0.032
0.101
0.32
1
Concentration (uM)

FIG. 4

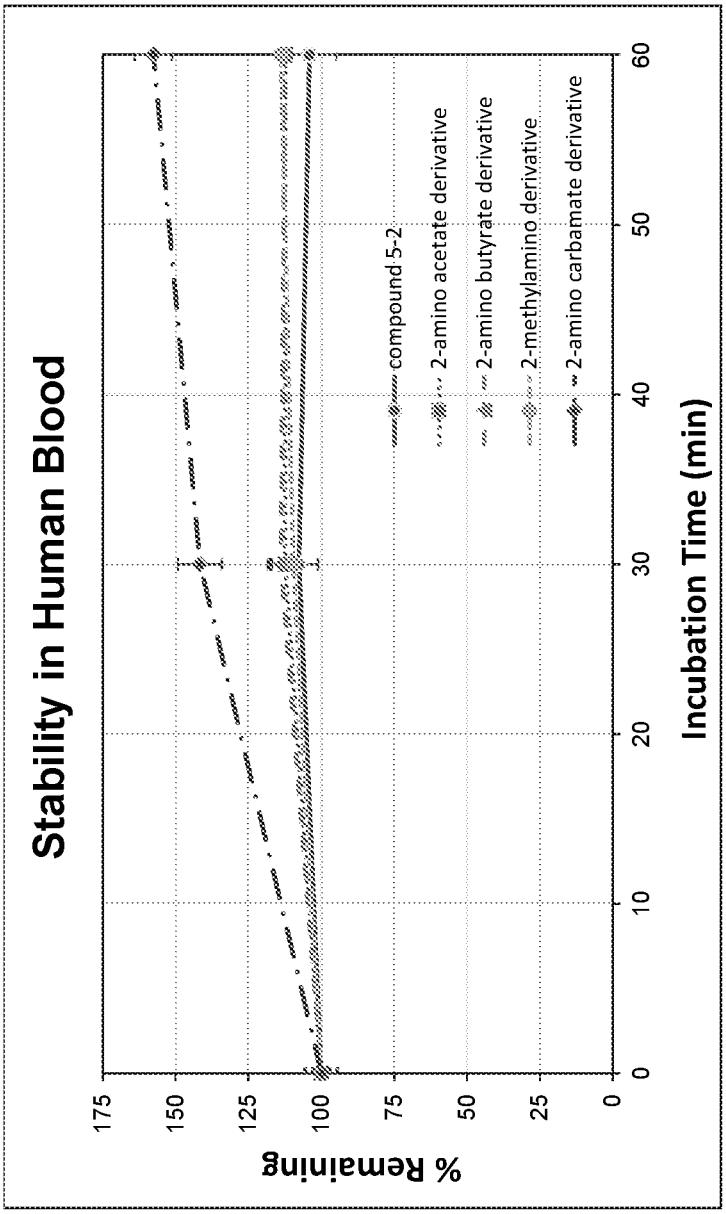

Stability in Human Blood
% Remaining
175
150
125
100
75
50
25
0
0
10
20
30
40
50
60
Incubation Time (min)
compound 5-2
2-amino acetate derivative
2-amino butyrate derivative
2-methylamino derivative
2-amino carbamate derivative

FIG. 5

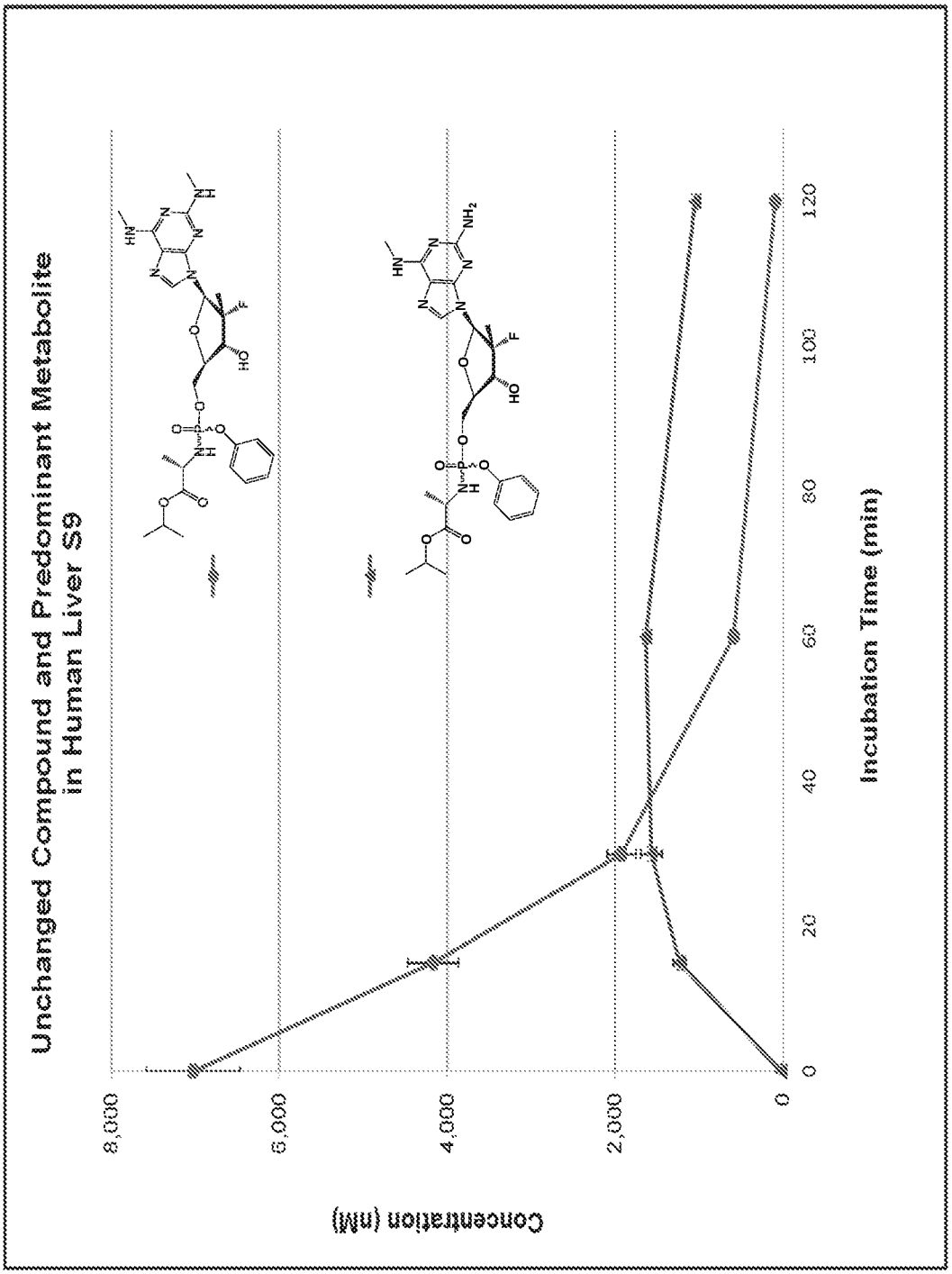

Unchanged Compound and Predominant Metabolite in Human Liver S9
Concentration (nM)
8,000
6,000
4,000
2,000
0
0
20
40
60
80
100
120
Incubation Time (min)

FIG. 6

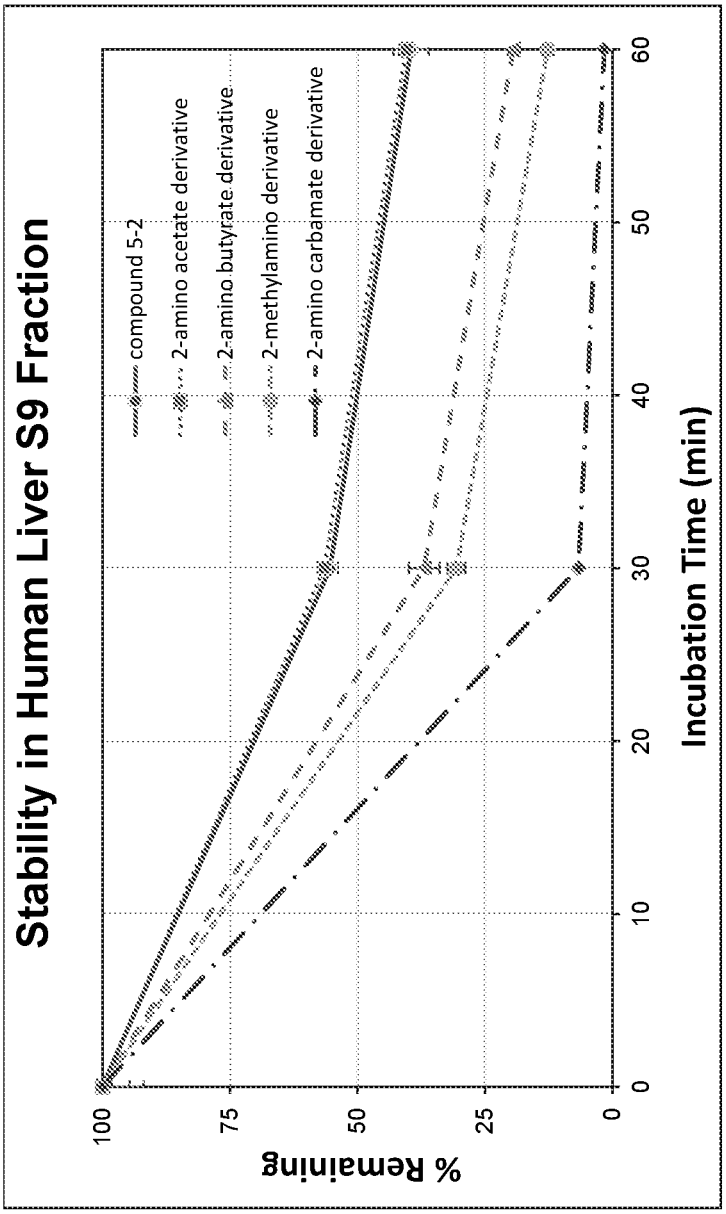

Stability in Human Liver S9 Fraction
compound 5-2
2-amino acetate derivative
2-amino butyrate derivative
2-methylamino derivative
2-amino carbamate derivative
% Remaining
100
75
50
25
0
0
10
20
30
40
50
60
Incubation Time (min)

FIG. 7

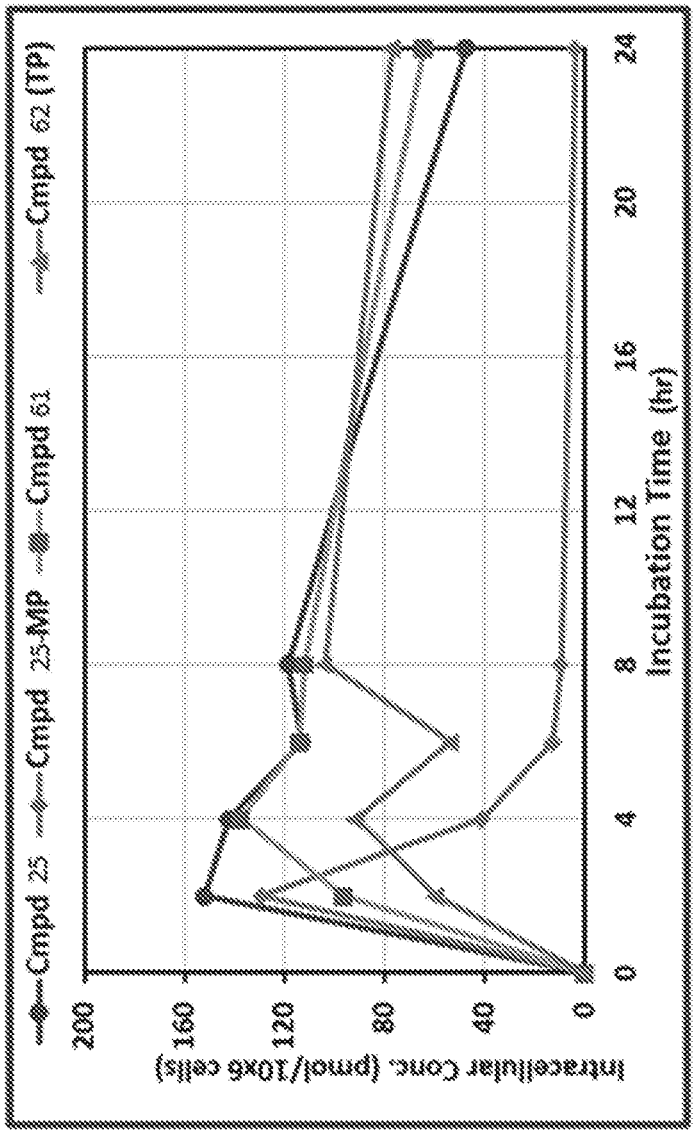

Cmpd 25
Cmpd 25-MP
Cmpd 61
Cmpd 62 (TP)
Intracellular Conc. (pmol/10x6 cells)
200
160
120
80
40
0
0
4
8
12
16
20
24
Incubation Time (hr)

FIG. 8

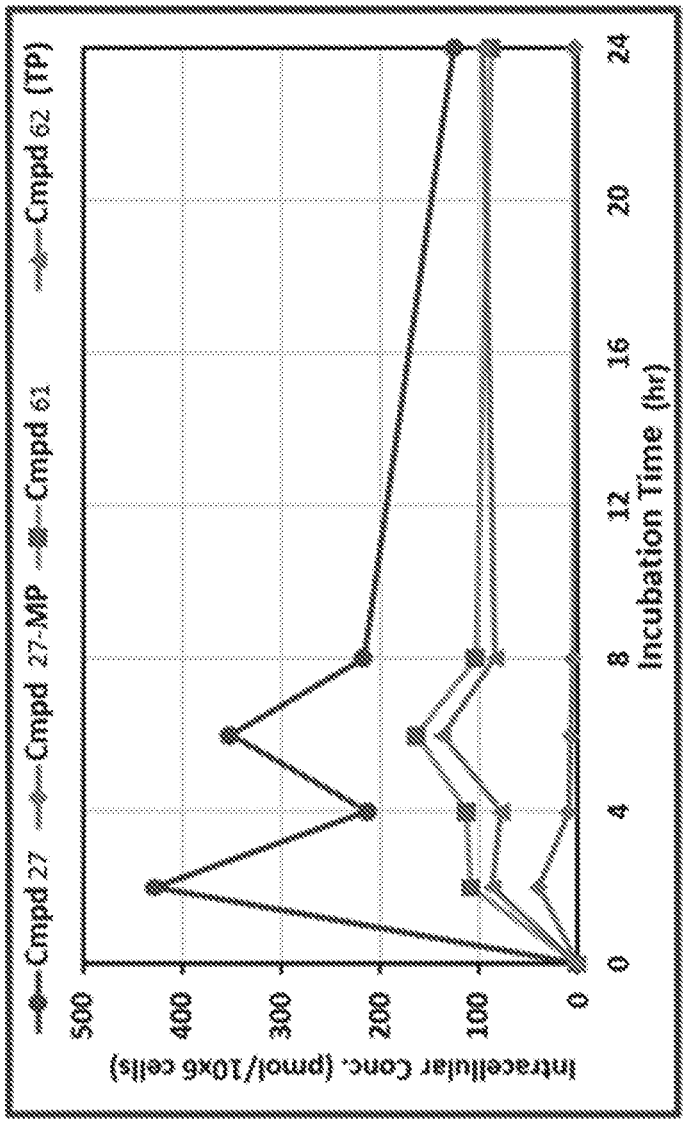

Cmpd 27
Cmpd 27-MP
Cmpd 61
Cmpd 62 (TP)
Intracellular Conc. (pmol/10x6 cells)
500
400
300
200
100
0
0
4
8
12
16
20
24
Incubation Time (hr)

FIG. 9

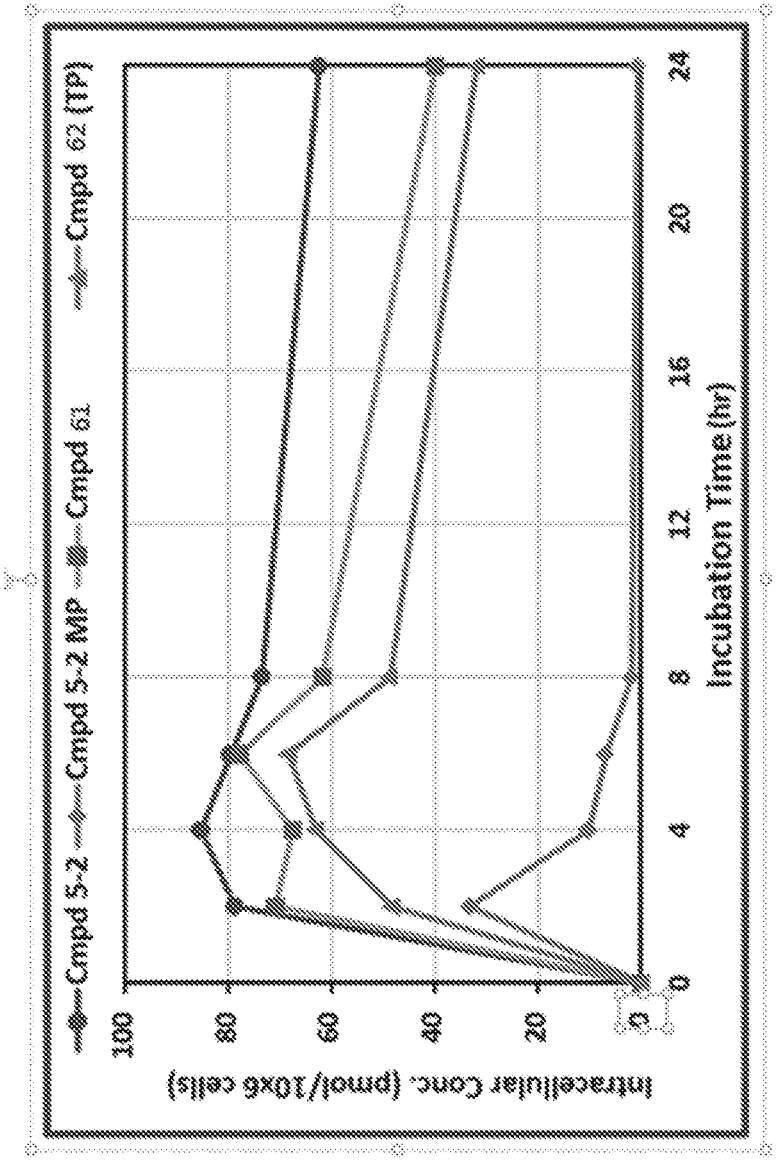

Cmpd 5-2
Cmpd 5-2 MP
Cmpd 61
Cmpd 62 (TP)
Intracellular Conc. (pmol/10x6 cells)
100
80
60
40
20
0
0
4
8
12
16
20
24
Incubation Time (hr)

FIG. 10

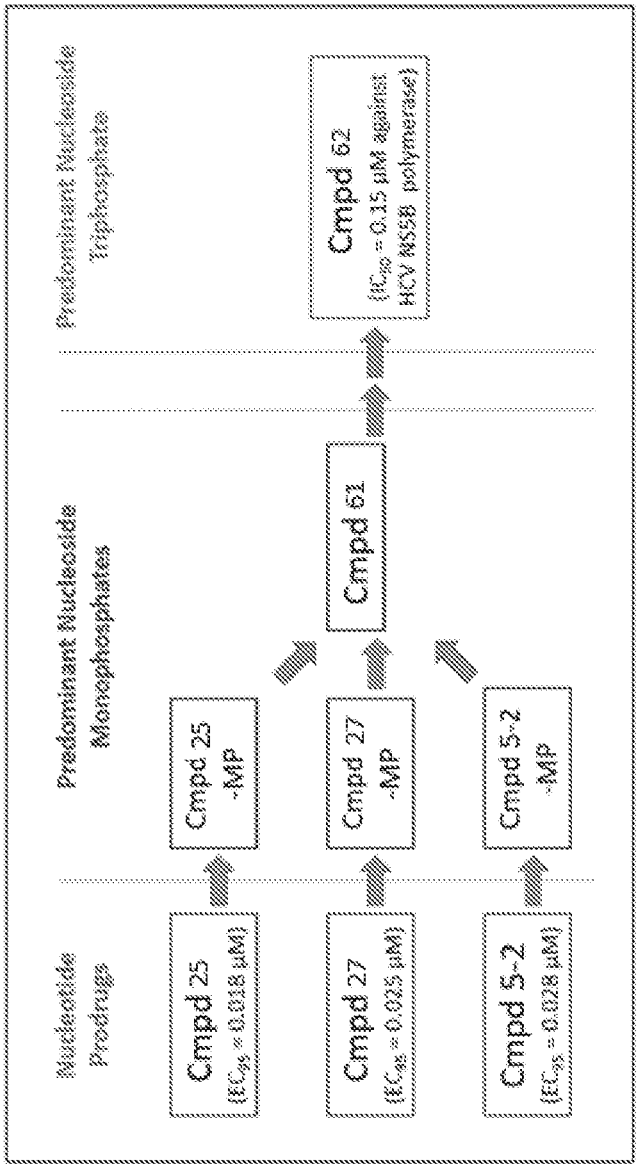

Nucleotide Prodrugs
Predominant Nucleoside Monophosphates
Predominant Nucleoside Triphosphate
Cmpd 25 (EC_{95} = 0.018 μM)
Cmpd 27 (EC_{95} = 0.025 μM)
Cmpd 5-2 (EC_{95} = 0.028 μM)
Cmpd 25 -MP
Cmpd 27 -MP
Cmpd 5-2 -MP
Cmpd 61
Cmpd 62 (IC_{50} = 0.15 μM against HCV NS5B polymerase)

FIG. 11