Figure 1A

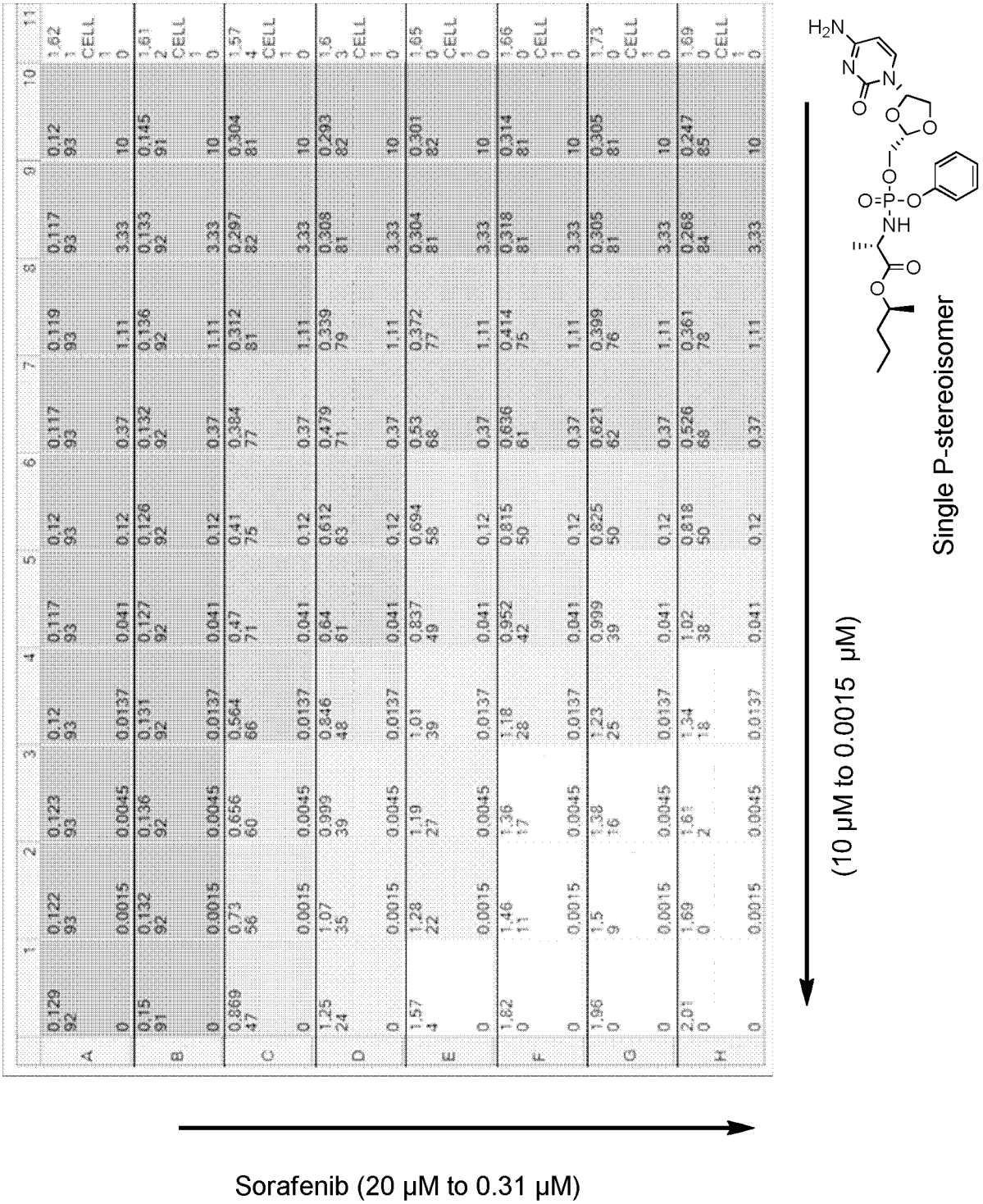

Sorafenib (20 µM to 0.31 µM)
(10 µM to 0.0015 µM)
Single P-stereoisomer

Figure 1B

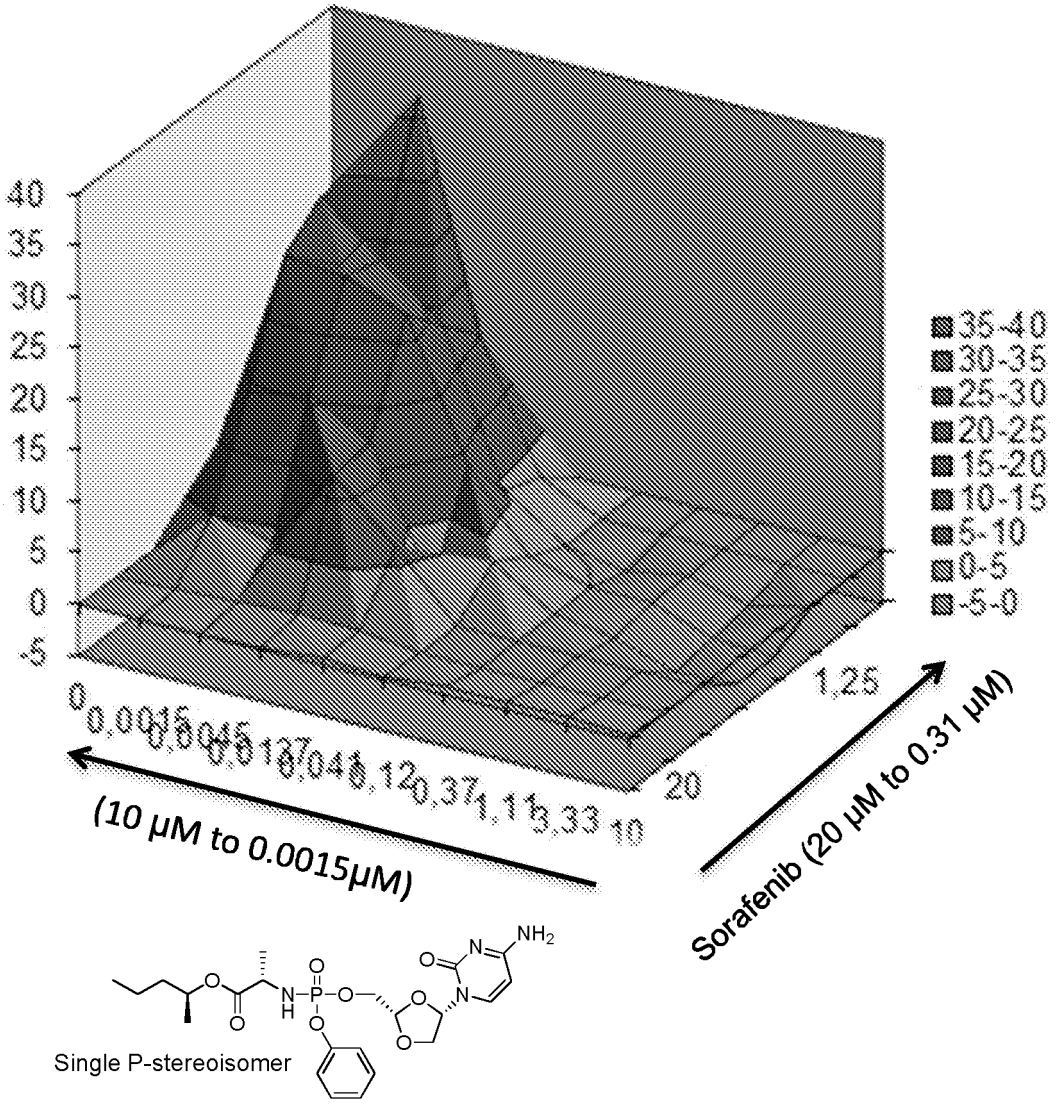

40
35
30
25
20
15
10
5
0
-5
0
0,0015
0,0045
0,014
0,041
0,12
0,37
1,11
3,33
10
20
1,25
(10 µM to 0.0015µM)
Sorafenib (20 µM to 0.31 µM)
35-40
30-35
25-30
20-25
15-20
10-15
5-10
0-5
-5-0
NH2
Single P-stereoisomer

Figure 2A

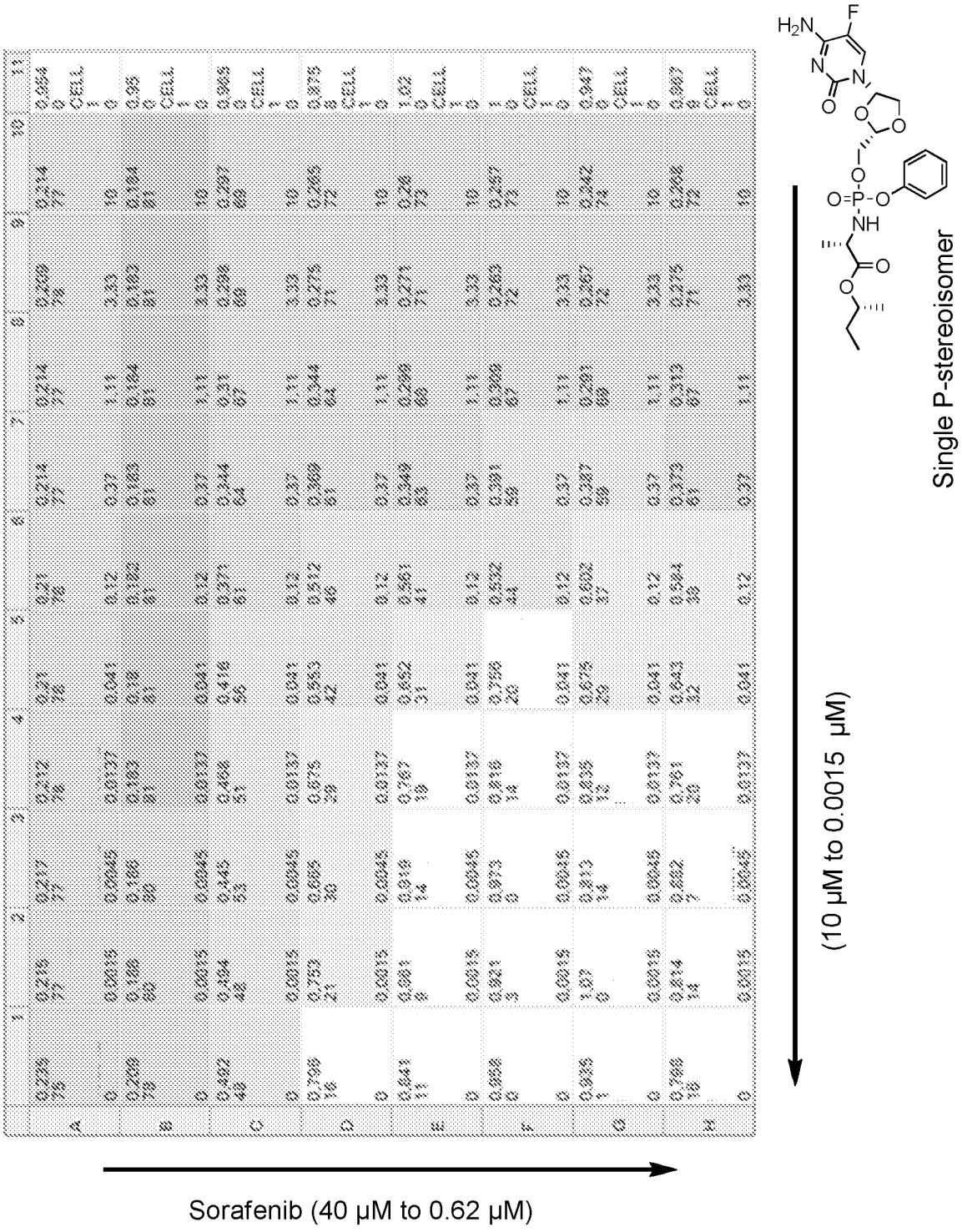

Sorafenib (40 µM to 0.62 µM)
(10 µM to 0.0015 µM)
Single P-stereoisomer

Figure 2B

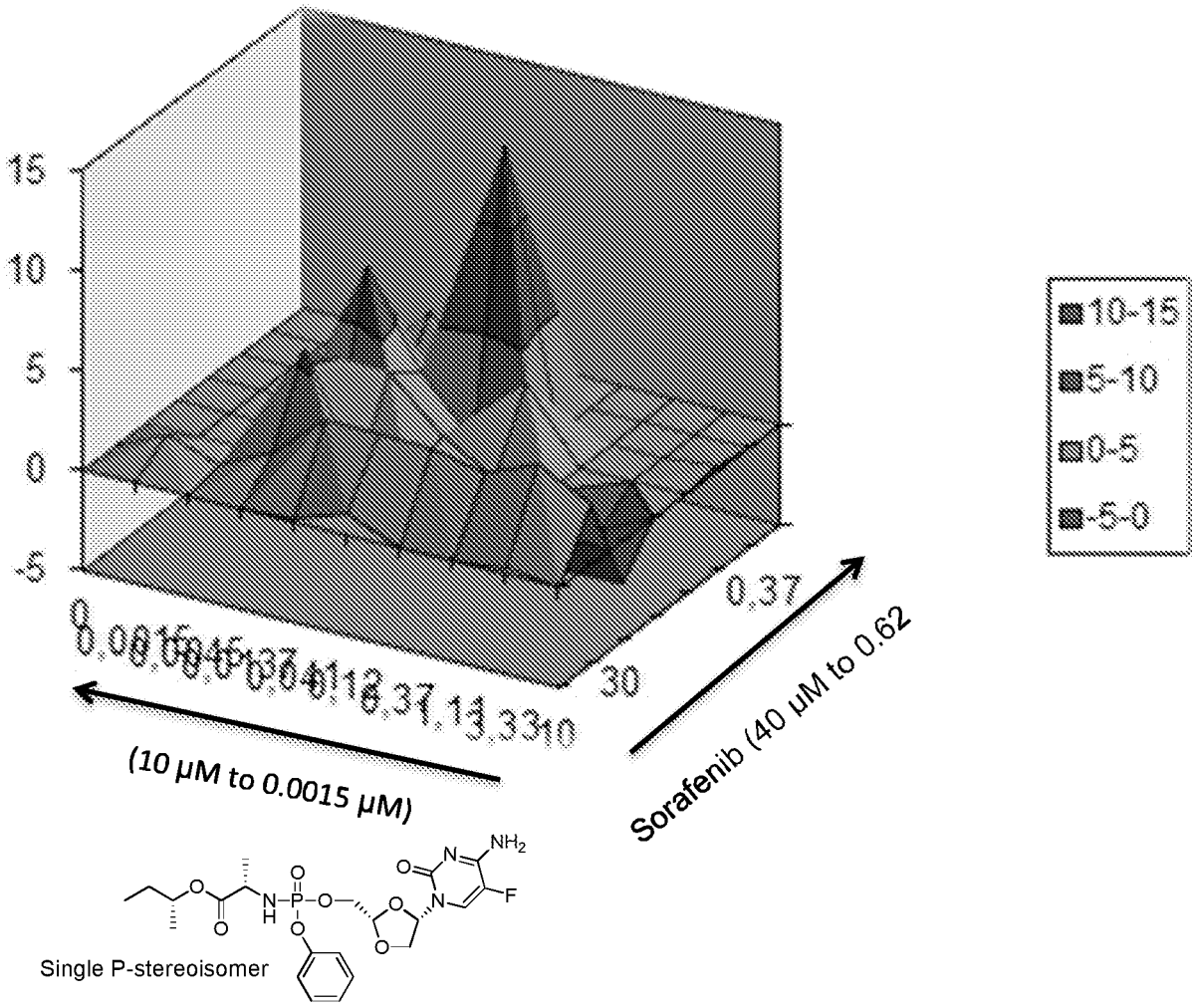

15
10
5
0
-5
10-15
5-10
0-5
-5-0
0.37
30
(10 µM to 0.0015 µM)
Sorafenib (40 µM to 0.62
Single P-stereoisomer

Figure 3A

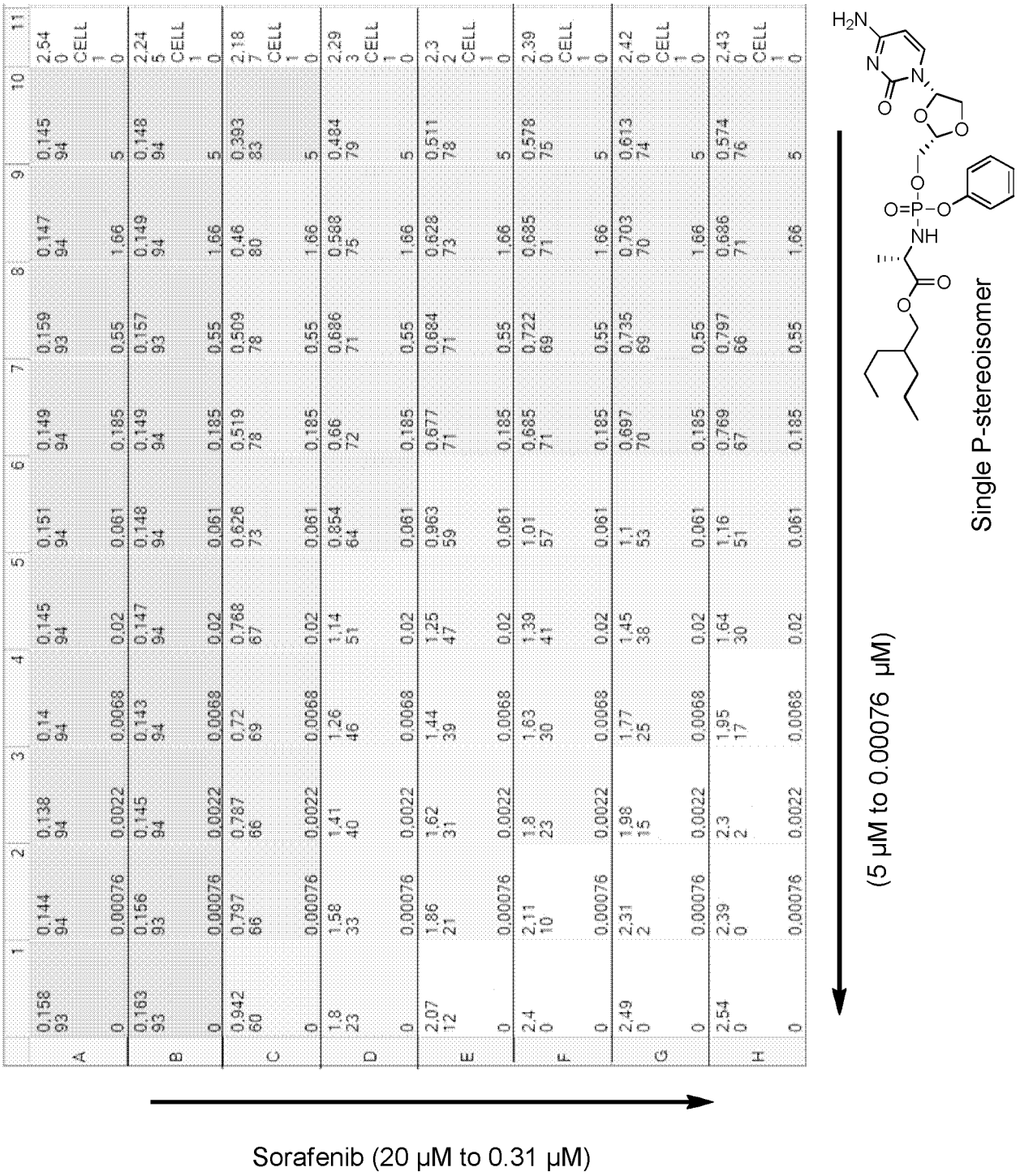

Sorafenib (20 µM to 0.31 µM)
(5 µM to 0.00076 µM)
Single P-stereoisomer

Figure 3B

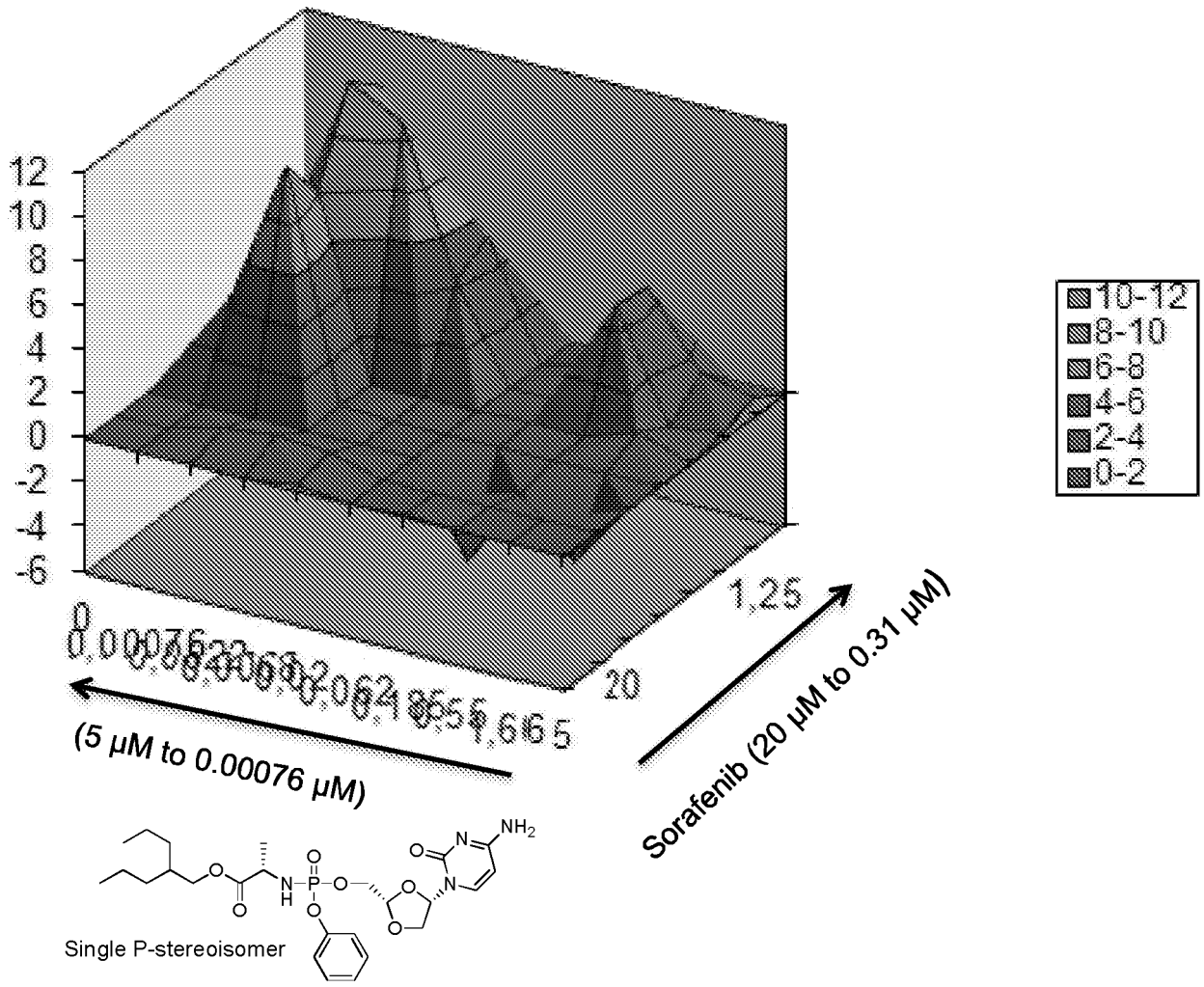
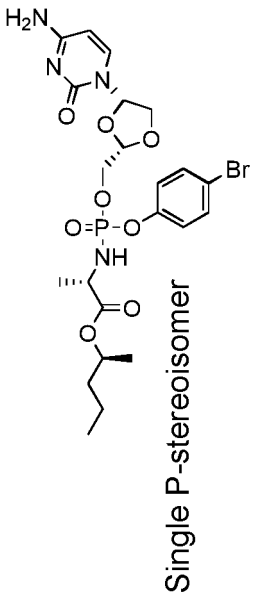

12
10
8
6
4
2
0
-2
-4
-6
10-12
8-10
6-8
4-6
2-4
0-2
0
1,25
20
(5 µM to 0.00076 µM)
Sorafenib (20 µM to 0.31 µM)
Single P-stereoisomer

Figure 4A

| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 10 | 11 |
|---|---|---|---|---|---|---|---|---|---|---|---|
| A | 0,146<br>94 | 0,15<br>93 | 0,15<br>93 | 0,143<br>94 | 0,16<br>93 | 0,146<br>94 | 0,145<br>94 | 0,151<br>93 | 0,153<br>93 | 0,153<br>93 | 2,57<br>0 |
| | 0 | 0,00076 | 0,0022 | 0,0068 | 0,02 | 0,061 | 0,185 | 0,55 | 1,66 | 5 | CELL<br>1<br>0 |
| B | 0,163<br>93 | 0,157<br>93 | 0,151<br>93 | 0,151<br>93 | 0,157<br>93 | 0,152<br>93 | 0,15<br>93 | 0,159<br>93 | 0,161<br>93 | 0,163<br>93 | 2,32<br>0 |
| | 0 | 0,00076 | 0,0022 | 0,0068 | 0,02 | 0,061 | 0,185 | 0,55 | 1,66 | 5 | CELL<br>1<br>0 |
| C | 0,867<br>62 | 1,01<br>56 | 0,856<br>63 | 0,665<br>71 | 0,579<br>75 | 0,659<br>71 | 0,599<br>74 | 0,681<br>70 | 0,541<br>77 | 0,527<br>77 | 2,18<br>5 |
| | 0 | 0,00076 | 0,0022 | 0,0068 | 0,02 | 0,061 | 0,185 | 0,55 | 1,66 | 5 | CELL<br>1<br>0 |
| D | 1,76<br>24 | 1,61<br>30 | 1,48<br>36 | 1,28<br>44 | 1,19<br>49 | 1,07<br>54 | 0,843<br>63 | 0,703<br>69 | 0,622<br>73 | 0,593<br>74 | 2,18<br>5 |
| | 0 | 0,00076 | 0,0022 | 0,0068 | 0,02 | 0,061 | 0,185 | 0,55 | 1,66 | 5 | CELL<br>1<br>0 |
| E | 2,08<br>10 | 1,95<br>15 | 1,73<br>25 | 1,52<br>34 | 1,38<br>40 | 1,23<br>47 | 0,91<br>60 | 0,744<br>68 | 0,644<br>72 | 0,624<br>73 | 2,22<br>3 |
| | 0 | 0,00076 | 0,0022 | 0,0068 | 0,02 | 0,061 | 0,185 | 0,55 | 1,66 | 5 | CELL<br>1<br>0 |
| F | 2,37<br>0 | 2,27<br>1 | 1,97<br>14 | 1,66<br>28 | 1,51<br>34 | 1,37<br>41 | 0,983<br>57 | 0,795<br>65 | 0,653<br>72 | 0,633<br>73 | 2,18<br>6 |
| | 0 | 0,00076 | 0,0022 | 0,0068 | 0,02 | 0,061 | 0,185 | 0,55 | 1,66 | 5 | CELL<br>1<br>0 |
| G | 2,49<br>0 | 2,4<br>0 | 2,3<br>0 | 1,89<br>18 | 1,62<br>30 | 1,48<br>36 | 1,03<br>55 | 0,829<br>64 | 0,684<br>70 | 0,635<br>72 | 2,25<br>2 |
| | 0 | 0,00076 | 0,0022 | 0,0068 | 0,02 | 0,061 | 0,185 | 0,55 | 1,66 | 5 | CELL<br>1<br>0 |
| H | 2,5<br>0 | 2,46<br>0 | 2,44<br>0 | 2,19<br>5 | 1,93<br>16 | 1,57<br>32 | 1,07<br>53 | 0,797<br>65 | 0,684<br>70 | 0,669<br>71 | 2,52<br>0 |
| | 0 | 0,00076 | 0,0022 | 0,0068 | 0,02 | 0,061 | 0,185 | 0,55 | 1,66 | 5 | CELL<br>1<br>0 |

Sorafenib (20 µM to 0.31 µM)

(5 µM to 0.00076 µM)




Single P-stereoisomer

Figure 4B

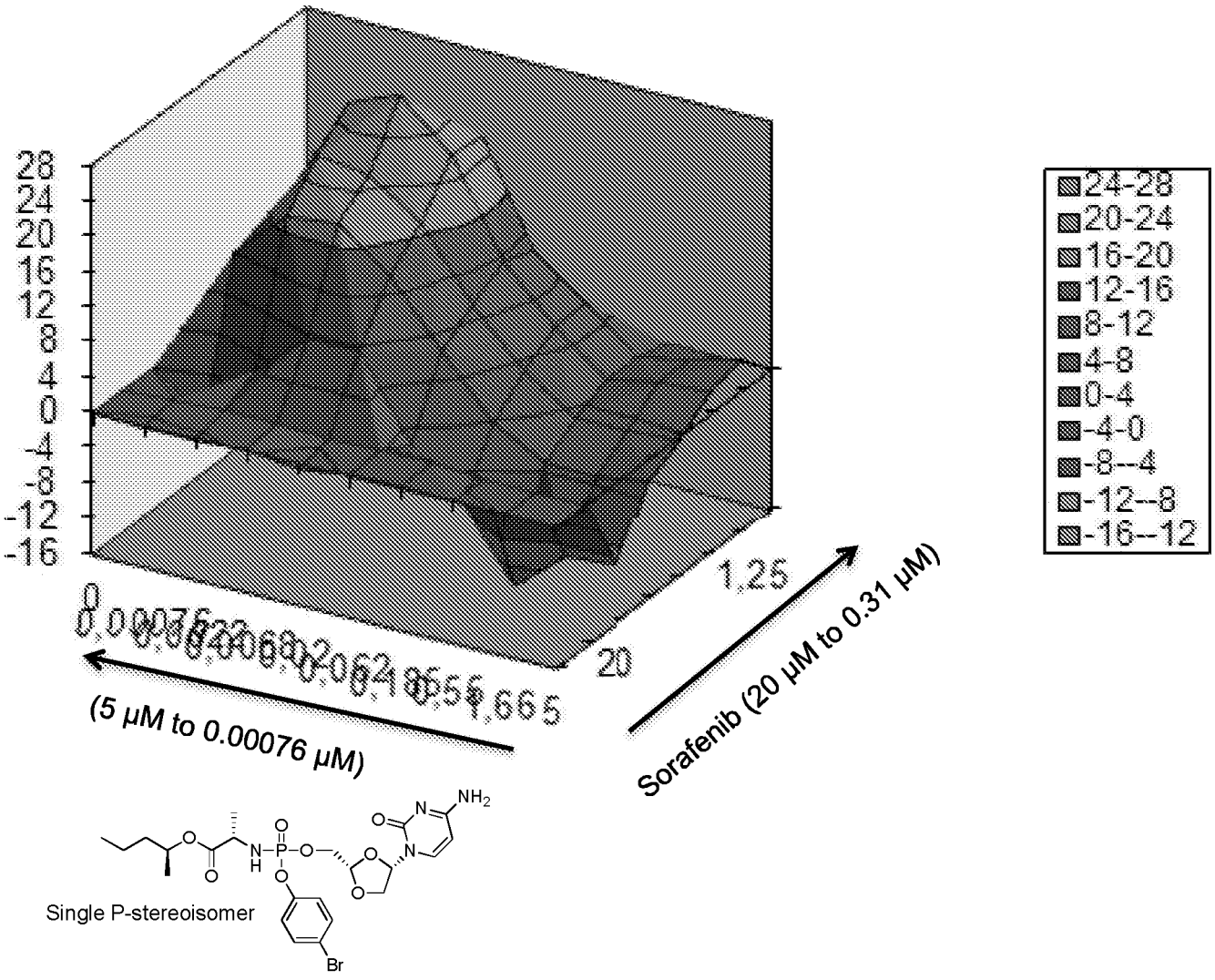

28
24
20
16
12
8
4
0
-4
-8
-12
-16
0
1,25
20
(5 µM to 0.00076 µM)
Sorafenib (20 µM to 0.31 µM)
24-28
20-24
16-20
12-16
8-12
4-8
0-4
-4-0
-8--4
-12--8
-16--12
NH2
Single P-stereoisomer
Br

Figure 5A

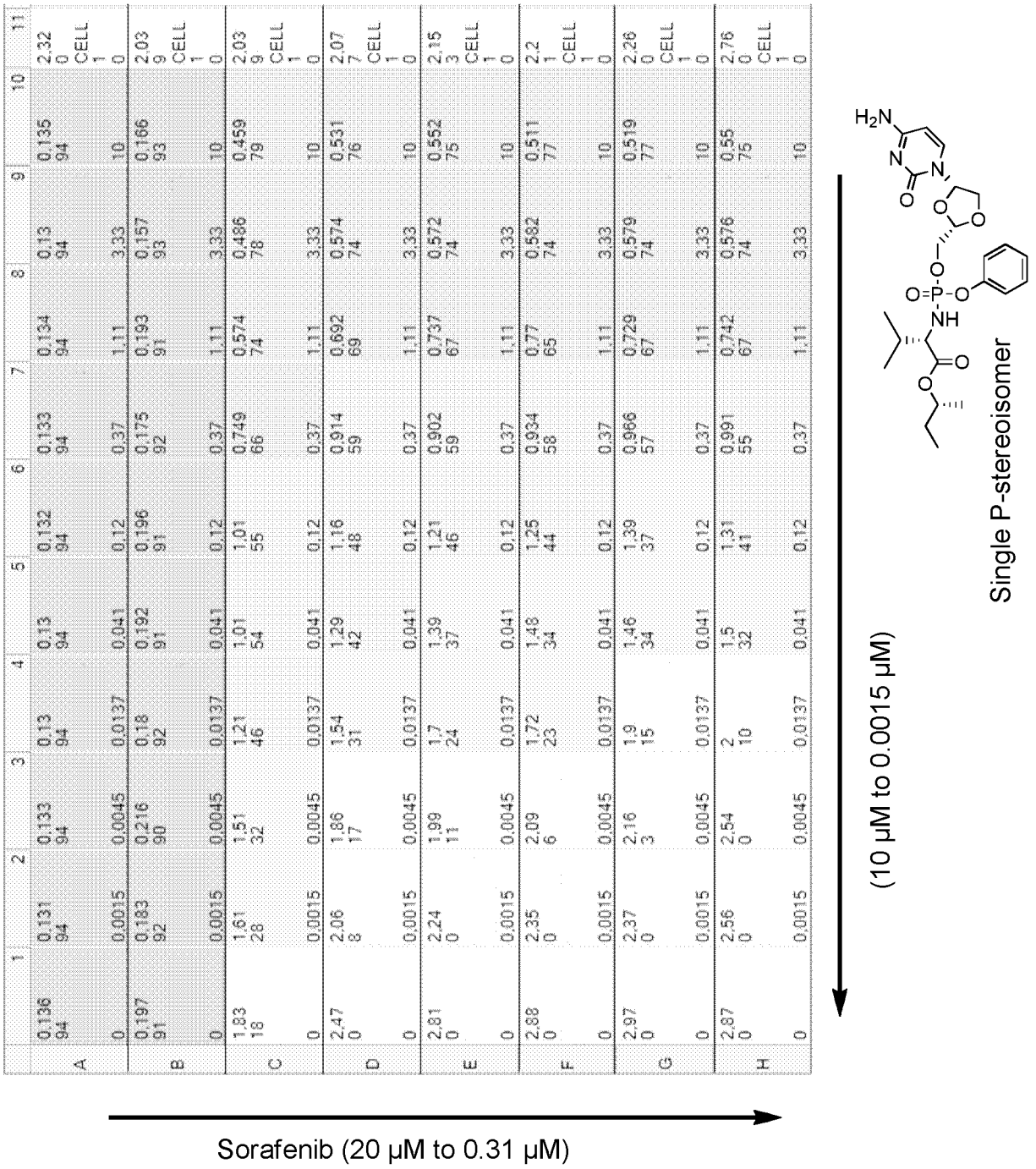
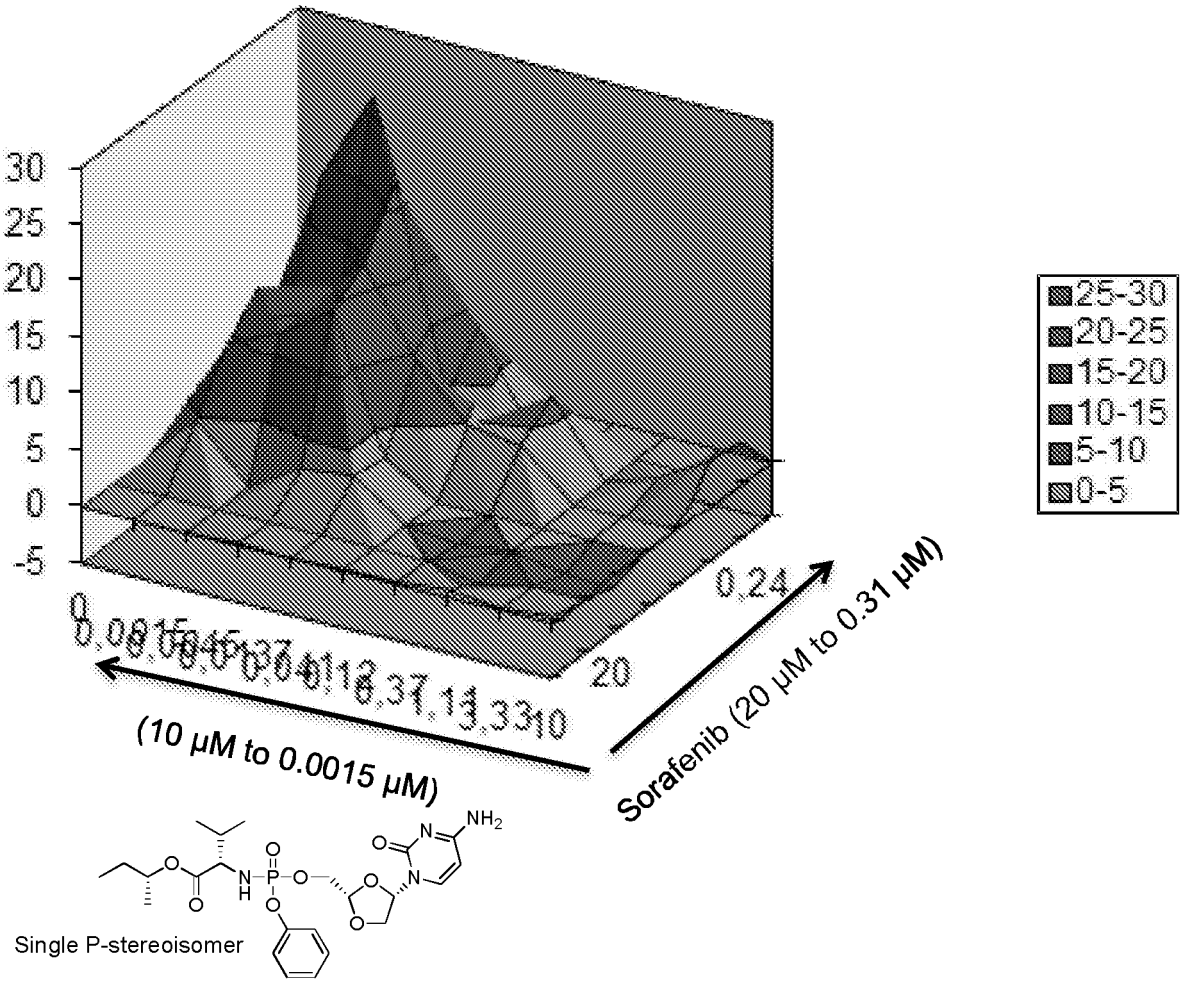

1 2 3 4 5 6 7 8 9 10 11
A 0.136 94 0.131 94 0.133 94 0.13 94 0.13 94 0.132 94 0.133 94 0.134 94 0.13 94 0.135 94 2.32 0 CELL 1 0
0 0.0015 0.0045 0.0137 0.041 0.12 0.37 1.11 3.33 10
B 0.197 91 0.183 92 0.216 90 0.18 92 0.192 91 0.196 91 0.175 92 0.193 91 0.157 93 0.166 93 2.03 9 CELL 1 0
0 0.0015 0.0045 0.0137 0.041 0.12 0.37 1.11 3.33 10
C 1.83 18 1.61 28 1.51 32 1.21 46 1.01 54 1.01 55 0.749 66 0.574 74 0.486 78 0.459 79 2.03 9 CELL 1 0
0 0.0015 0.0045 0.0137 0.041 0.12 0.37 1.11 3.33 10
D 2.47 0 2.06 8 1.86 17 1.54 31 1.29 42 1.16 48 0.914 59 0.692 69 0.574 74 0.531 76 2.07 7 CELL 1 0
0 0.0015 0.0045 0.0137 0.041 0.12 0.37 1.11 3.33 10
E 2.81 0 2.24 0 1.99 11 1.7 24 1.39 37 1.21 46 0.902 59 0.737 67 0.572 74 0.552 75 2.15 3 CELL 1 0
0 0.0015 0.0045 0.0137 0.041 0.12 0.37 1.11 3.33 10
F 2.88 0 2.35 0 2.09 6 1.72 23 1.48 34 1.25 44 0.934 58 0.77 65 0.582 74 0.511 77 2.2 1 CELL 1 0
0 0.0015 0.0045 0.0137 0.041 0.12 0.37 1.11 3.33 10
G 2.97 0 2.37 0 2.16 3 1.9 15 1.46 34 1.39 37 0.966 57 0.729 67 0.579 74 0.519 77 2.26 0 CELL 1 0
0 0.0015 0.0045 0.0137 0.041 0.12 0.37 1.11 3.33 10
H 2.87 0 2.56 0 2.54 0 2 10 1.5 32 1.31 41 0.991 55 0.742 67 0.576 74 0.55 75 2.76 0 CELL 1 0
0 0.0015 0.0045 0.0137 0.041 0.12 0.37 1.11 3.33 10
Sorafenib (20 µM to 0.31 µM)
(10 µM to 0.0015 µM)
Single P-stereoisomer

Figure 5B


30
25
20
15
10
5
0
-5
0
0,24
20
(10 µM to 0.0015 µM)
Sorafenib (20 µM to 0.31 µM)
25-30
20-25
15-20
10-15
5-10
0-5
Single P-stereoisomer