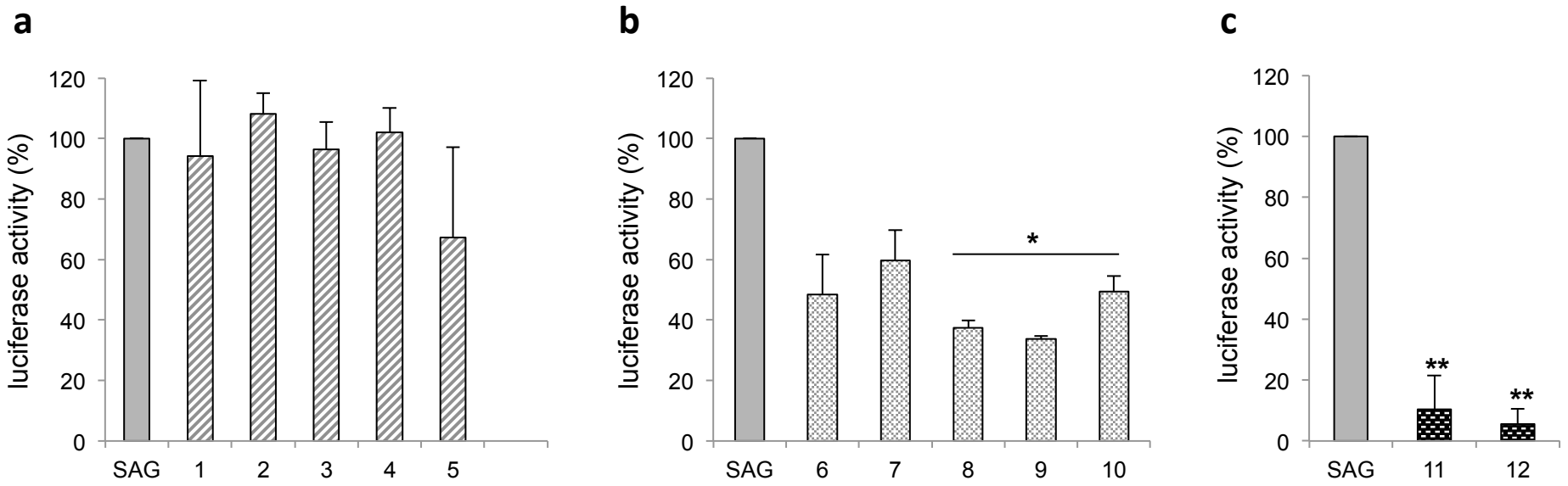
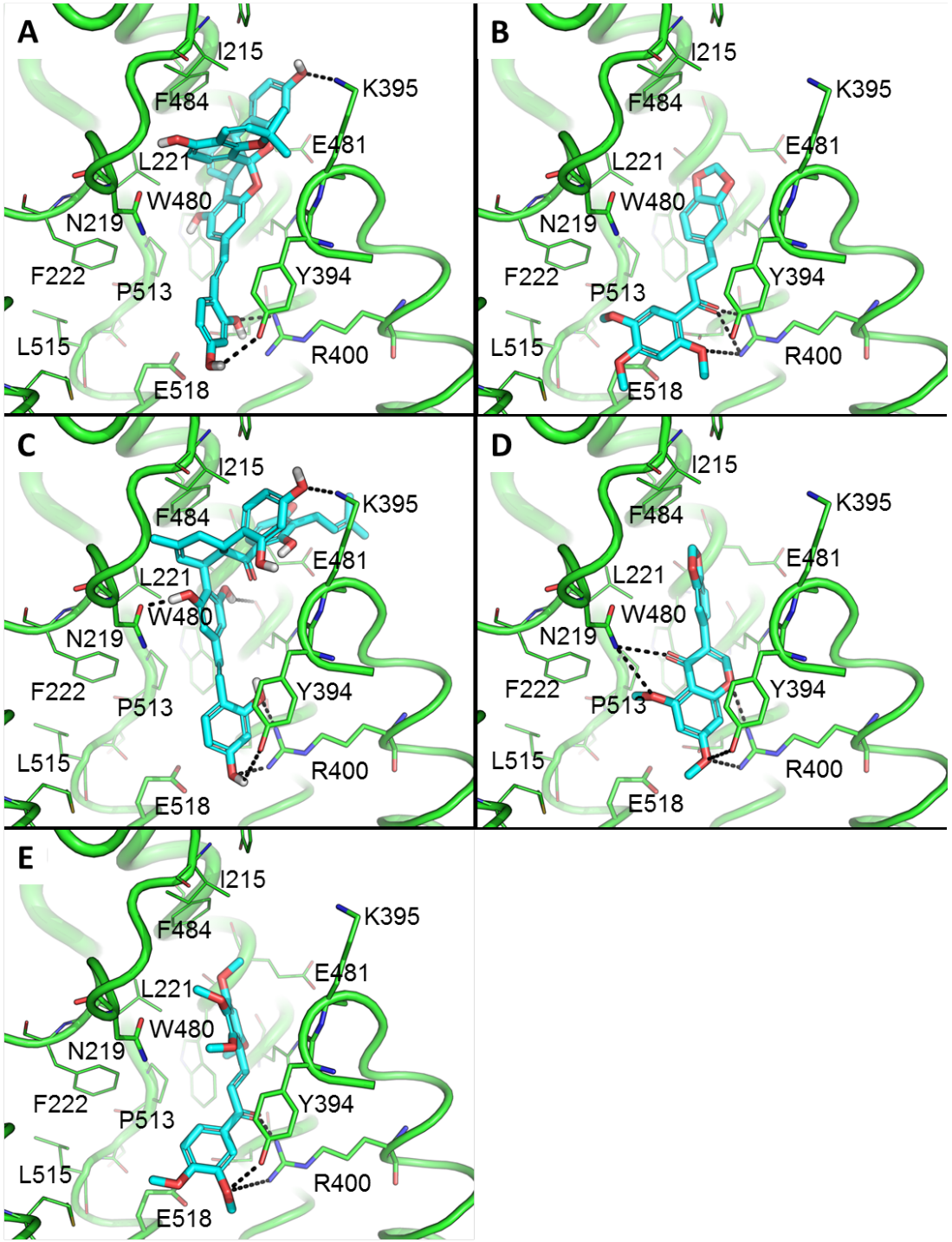
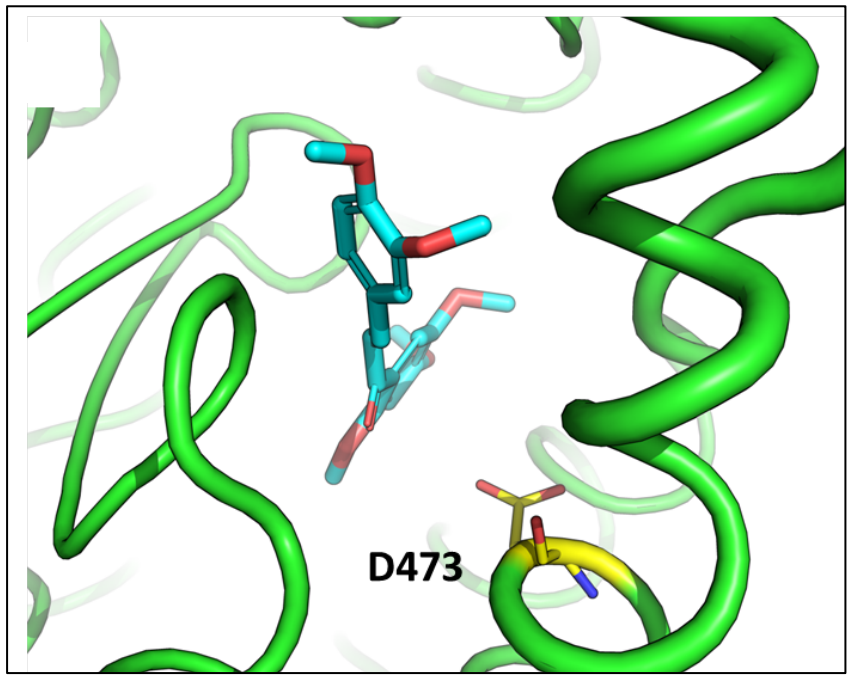
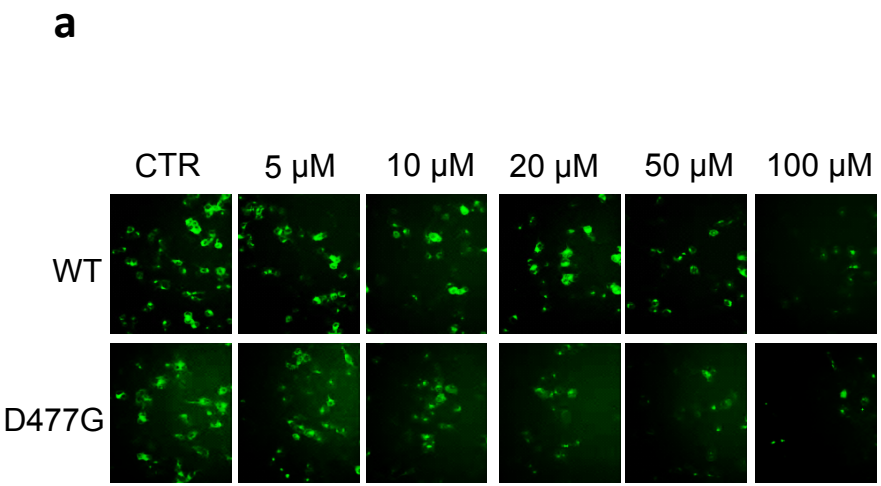

a
b
c
luciferase activity (%)
120
100
80
60
40
20
0
SAG
1
2
3
4
5
6
7
8
9
10
11
12
*
**
**

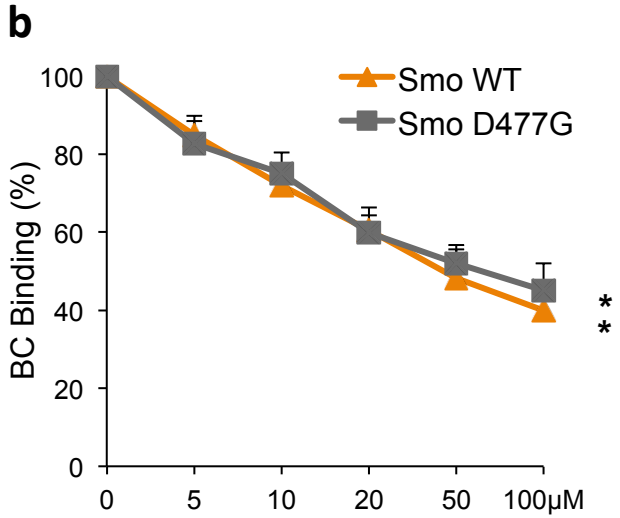
S1

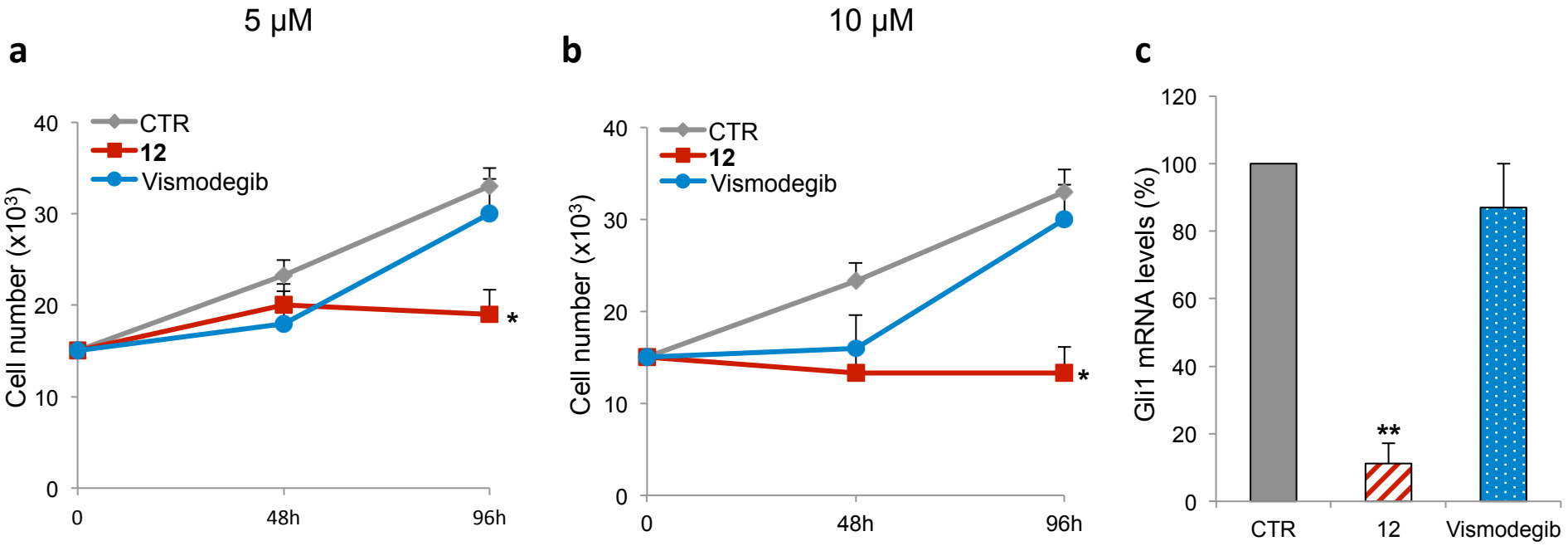

A
I215
K395
F484
E481
L221
W480
N219
F222
P513
Y394
L515
R400
E518
B
I215
K395
F484
E481
L221
W480
N219
F222
P513
Y394
L515
R400
E518
C
I215
K395
F484
E481
L221
W480
N219
F222
P513
Y394
L515
R400
E518
D
I215
K395
F484
E481
L221
W480
N219
F222
P513
Y394
L515
R400
E518
E
I215
K395
F484
E481
L221
W480
N219
F222
P513
Y394
L515
R400
E518

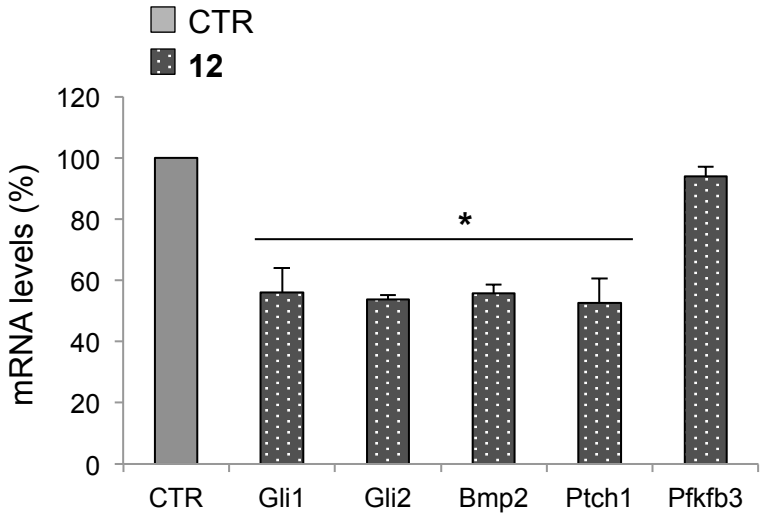
S2


D473


a
CTR
5 μM
10 μM
20 μM
50 μM
100 μM
WT
D477G


b
Smo WT
Smo D477G
BC Binding (%)
100
80
60
40
20
0
0
5
10
20
50
100μM
**


a
5 μM
CTR
12
Vismodegib
Cell number (x10³)
40
30
20
10
0
0
48h
96h
*
b
10 μM
CTR
12
Vismodegib
Cell number (x10³)
40
30
20
10
0
0
48h
96h
*
c
Gli1 mRNA levels (%)
120
100
80
60
40
20
0
**
CTR
12
Vismodegib


CTR
12
120
100
80
60
40
20
0
mRNA levels (%)
*
CTR
Gli1
Gli2
Bmp2
Ptch1
Pfkfb3