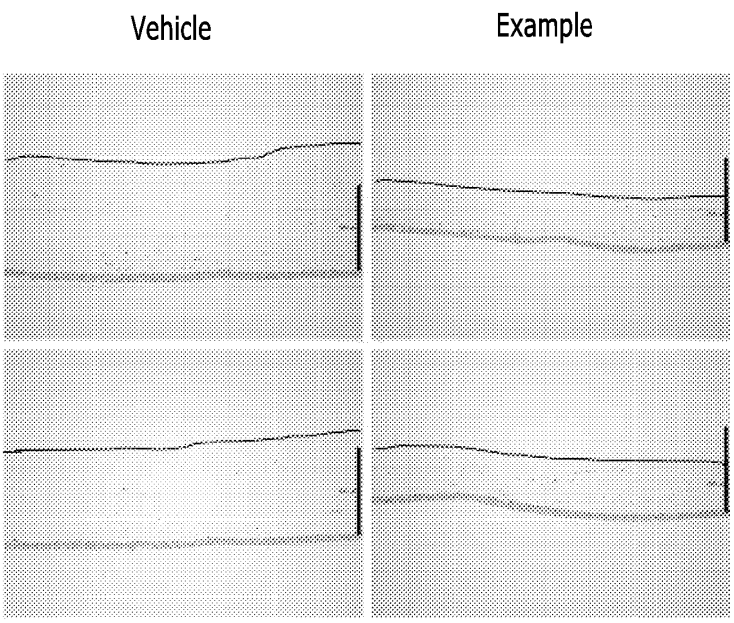
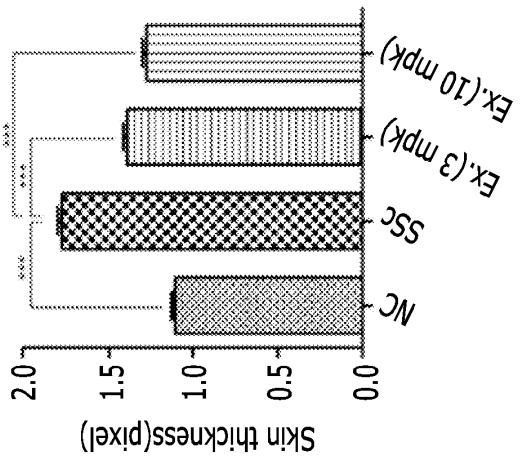
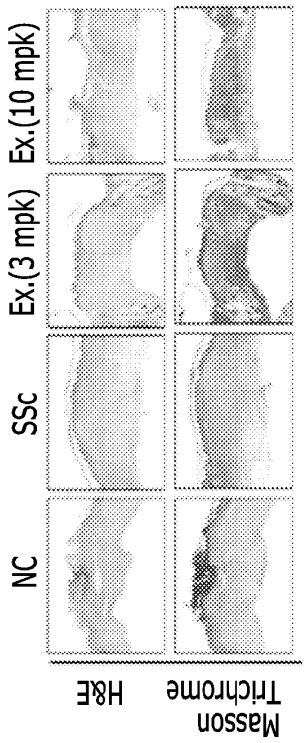
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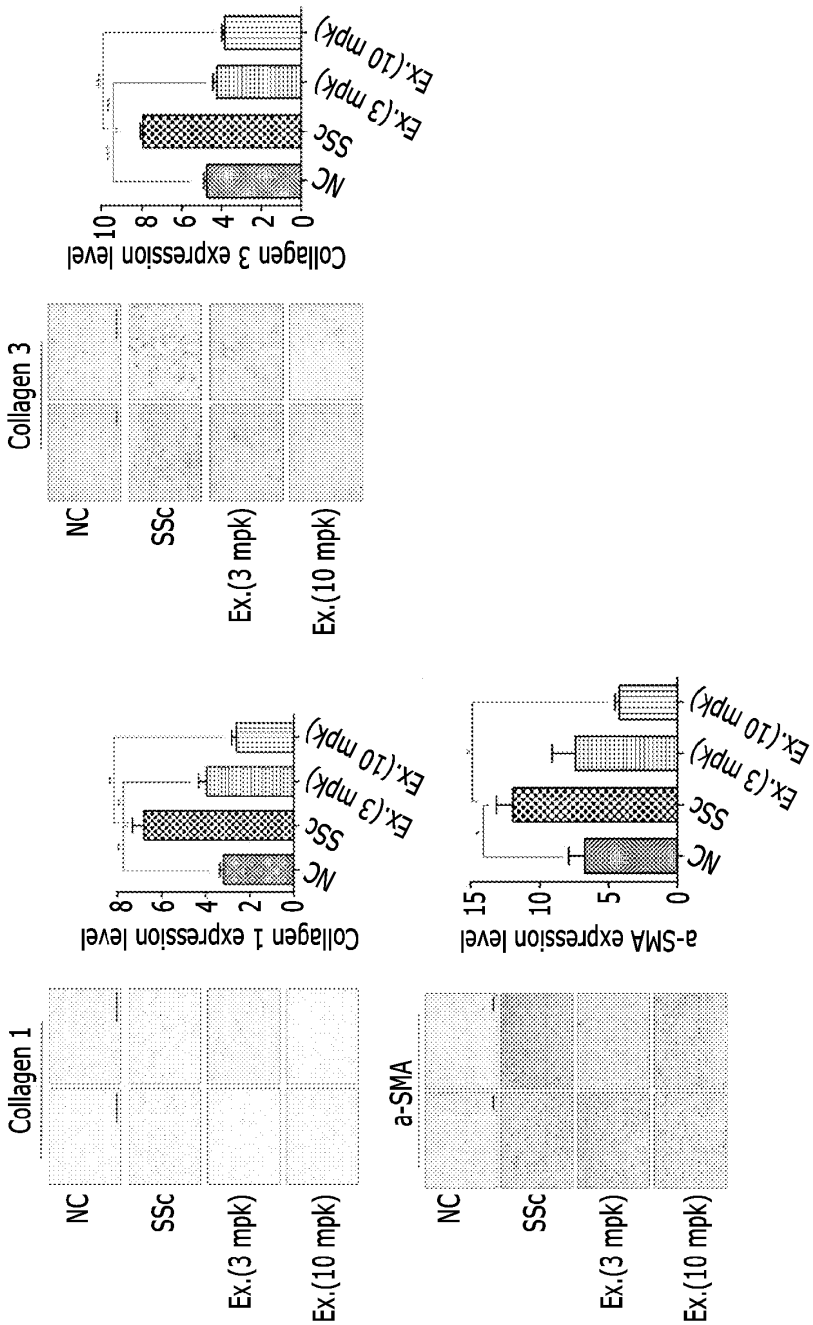


[도2]

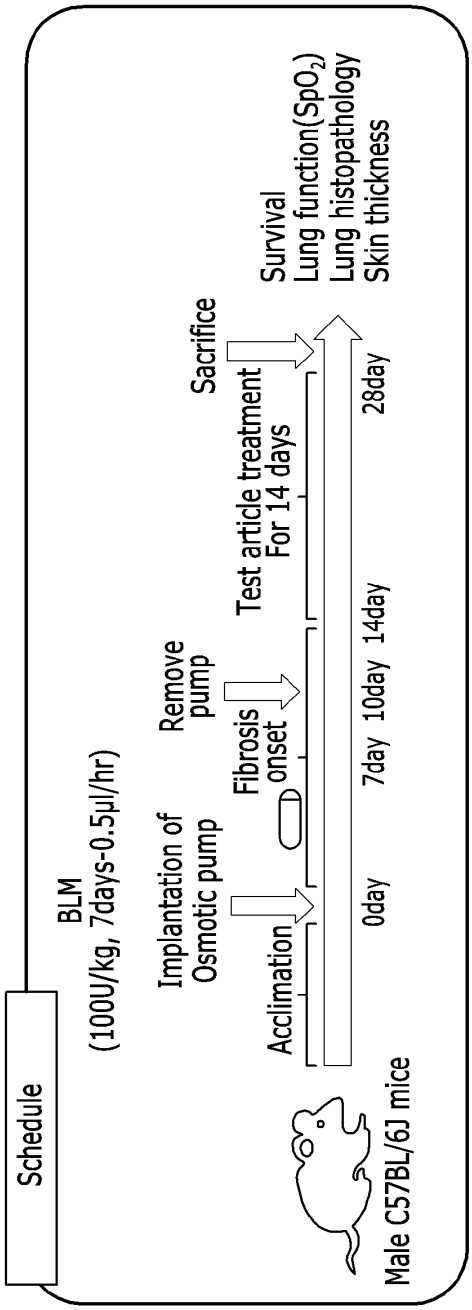

Skin thickness(pixel)
2.0
1.5
1.0
0.5
0.0
NC
SSc
Ex.(3 mpk)
Ex.(10 mpk)
***
***
***


NC
SSc
Ex.(3 mpk)
Ex.(10 mpk)
H&E
Masson Trichrome

[도3]


Collagen 3 expression level
0
2
4
6
8
10
NC
SSc
Ex.(3 mpk)
Ex.(10 mpk)
Collagen 3
NC
SSc
Ex.(3 mpk)
Ex.(10 mpk)
Collagen 1 expression level
0
2
4
6
8
NC
SSc
Ex.(3 mpk)
Ex.(10 mpk)
a-SMA expression level
0
5
10
15
NC
SSc
Ex.(3 mpk)
Ex.(10 mpk)
Collagen 1
NC
SSc
Ex.(3 mpk)
Ex.(10 mpk)
a-SMA
NC
SSc
Ex.(3 mpk)
Ex.(10 mpk)

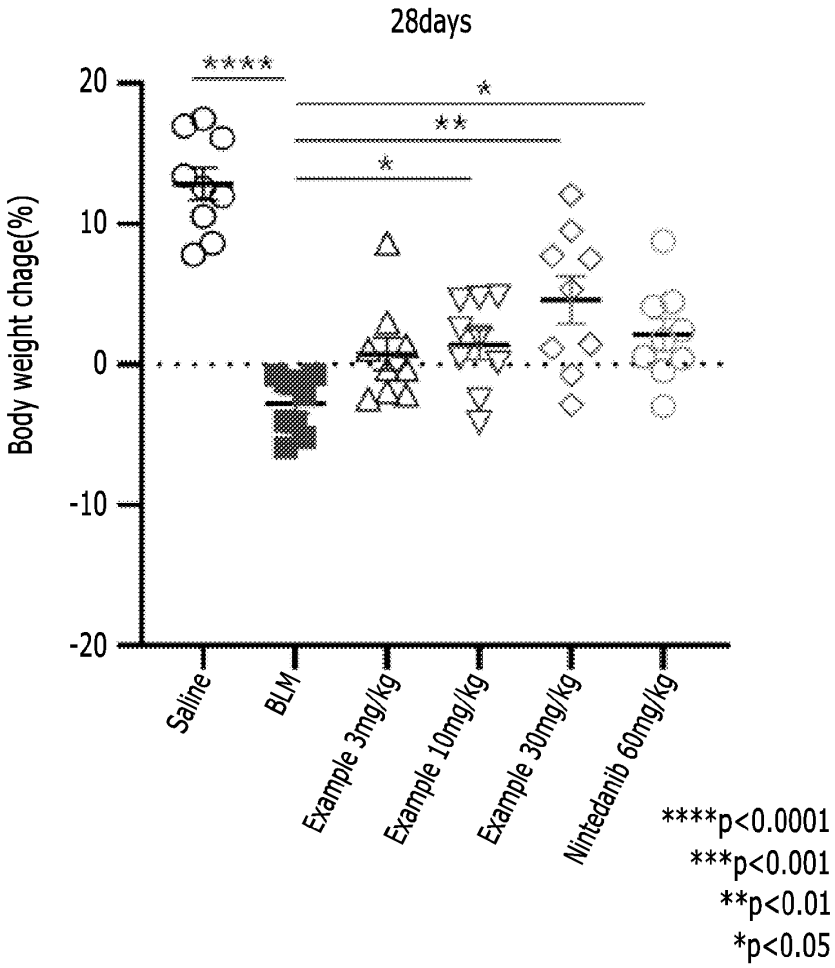
[도4]



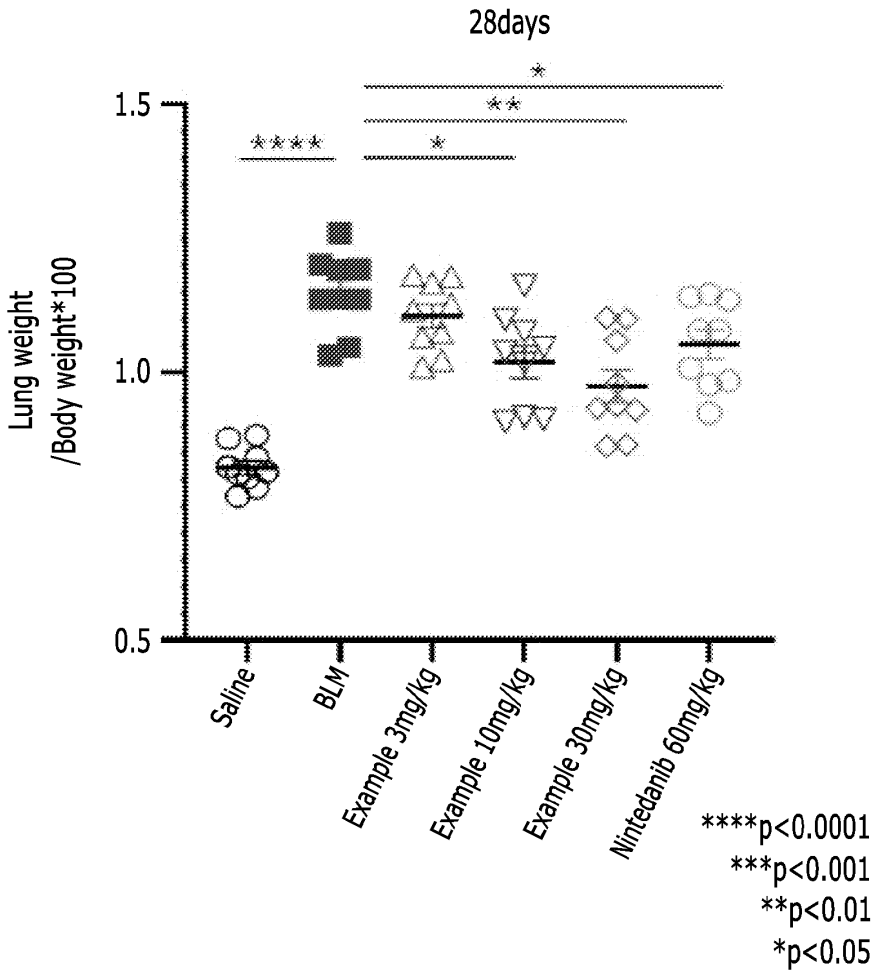
• Dosing : Example = 3 - 30 mg/kg, QD, once a day (N=9)
Nintedanib 60 mg/kg, QD, once a day

• Inducing agent : Bleomycin (BLM)

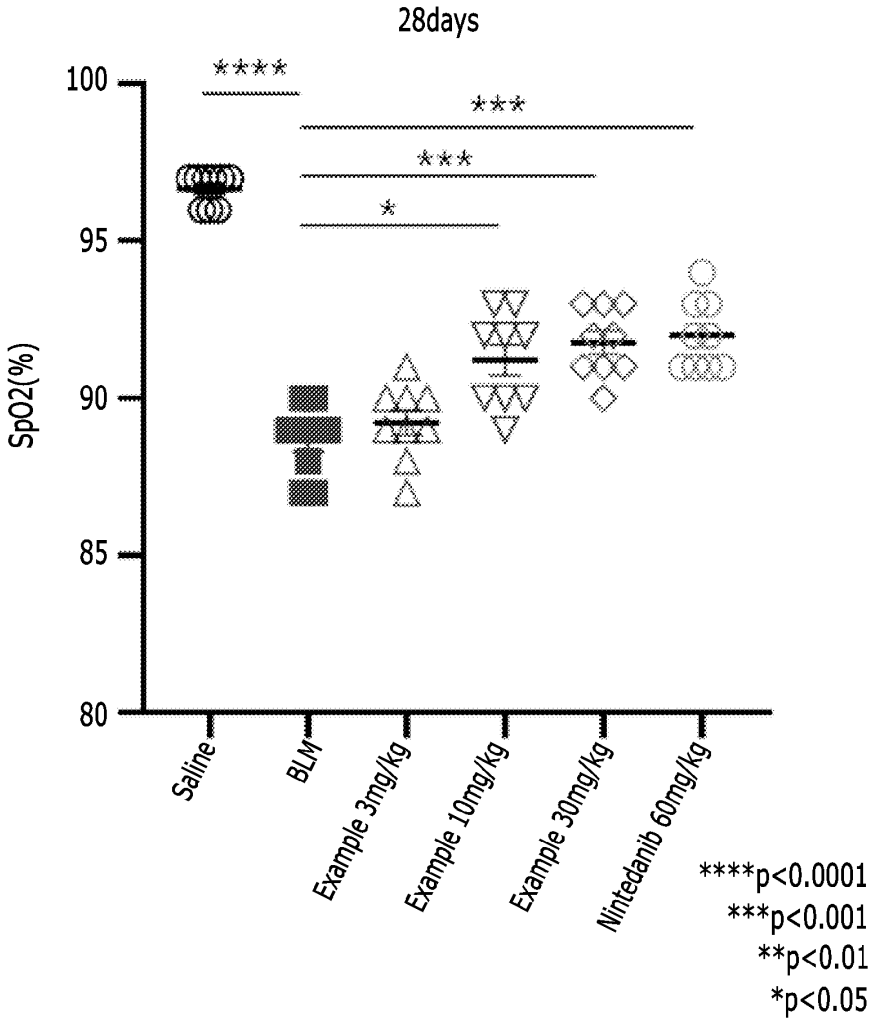
[도5]


28days
****
*
**
*
20
10
0
-10
-20
Body weight chage(%)
Saline
BLM
Example 3mg/kg
Example 10mg/kg
Example 30mg/kg
Nintedanib 60mg/kg
****p<0.0001
***p<0.001
**p<0.01
*p<0.05

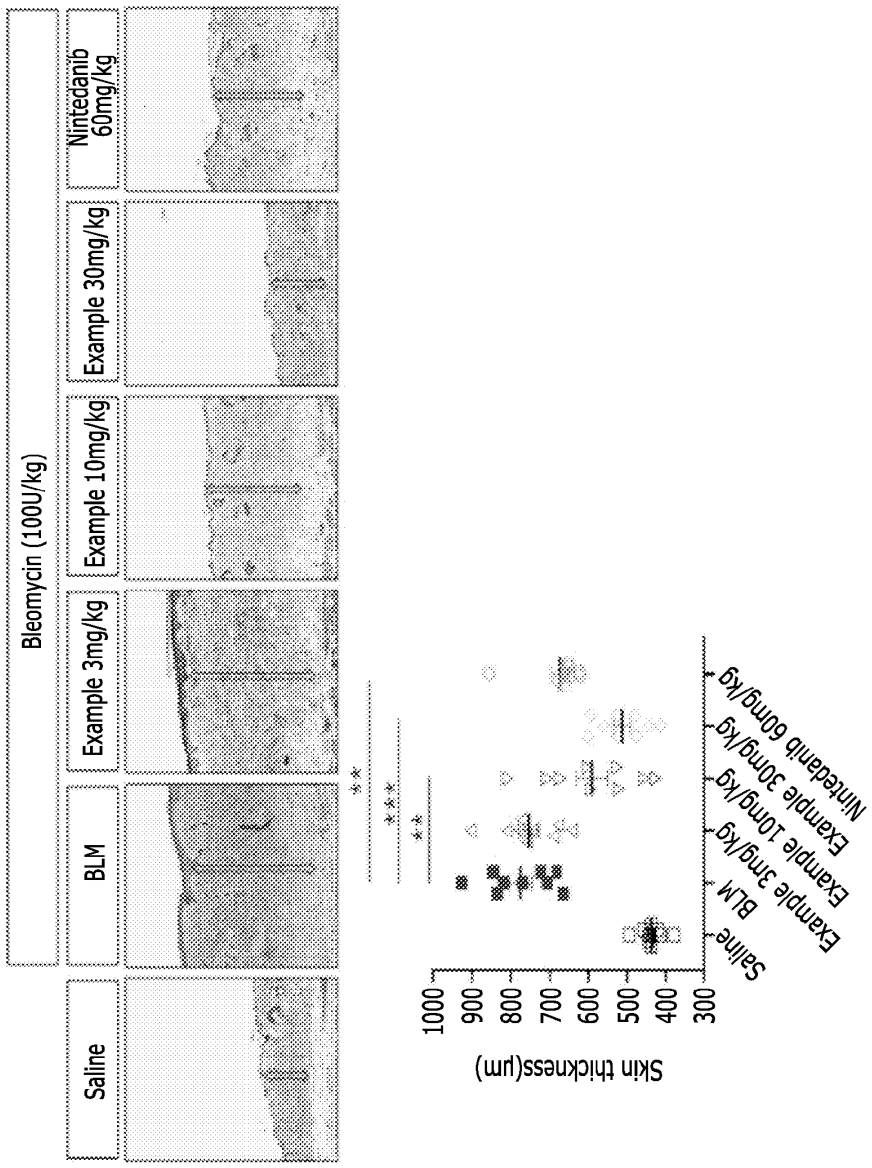
[도6]


28days
Lung weight /Body weight*100
1.5
1.0
0.5
****
*
**
*
Saline
BLM
Example 3mg/kg
Example 10mg/kg
Example 30mg/kg
Nintedanib 60mg/kg
****p<0.0001
***p<0.001
**p<0.01
*p<0.05

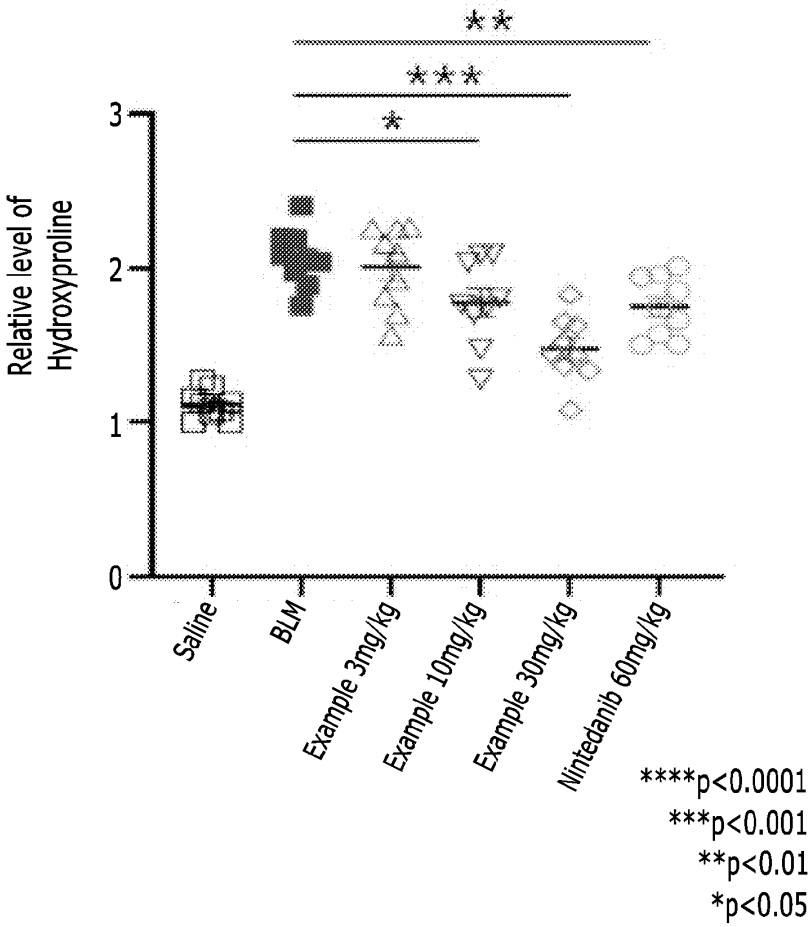
[도7]


28days
SpO2(%)
100
95
90
85
80
****
***
***
*
Saline
BLM
Example 3mg/kg
Example 10mg/kg
Example 30mg/kg
Nintedanib 60mg/kg
****p<0.0001
***p<0.001
**p<0.01
*p<0.05

[도8]


Bleomycin (100U/kg)
Saline
BLM
Example 3mg/kg
Example 10mg/kg
Example 30mg/kg
Nintedanib 60mg/kg
Skin thickness(μm)
1000
900
800
700
600
500
400
300
**
***
**
Saline
BLM
Example 3mg/kg
Example 10mg/kg
Example 30mg/kg
Nintedanib 60mg/kg

[도9]


Relative level of Hydroxyproline
3
2
1
0
Saline
BLM
Example 3mg/kg
Example 10mg/kg
Example 30mg/kg
Nintedanib 60mg/kg
****p<0.0001
***p<0.001
**p<0.01
*p<0.05