
F1
F3
C21
C3
C9
C20
N3
C16
C15
C10
C8
C7
C4
C2
F2
C19
N2
N1
C1
N4
C14
C17
C18
C11
C13
O2
C12
O3
C5
C6
Cl1
O1

图 1

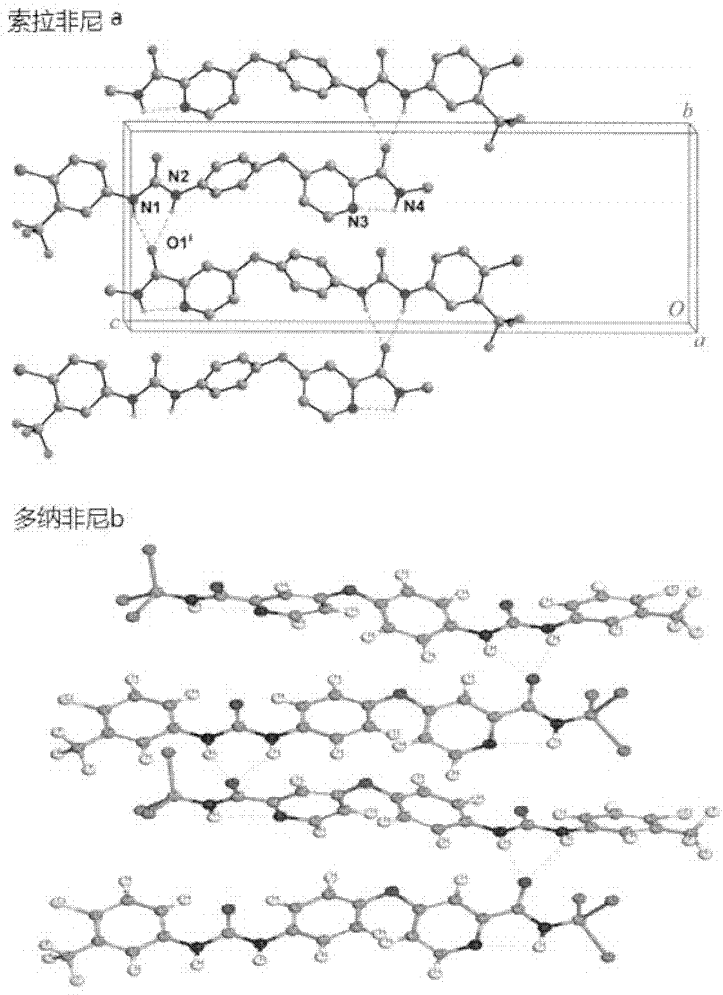

索拉非尼 a
b
N2
N1
N3
N4
O1ⁱ
c
O
a
多纳非尼b

图 2

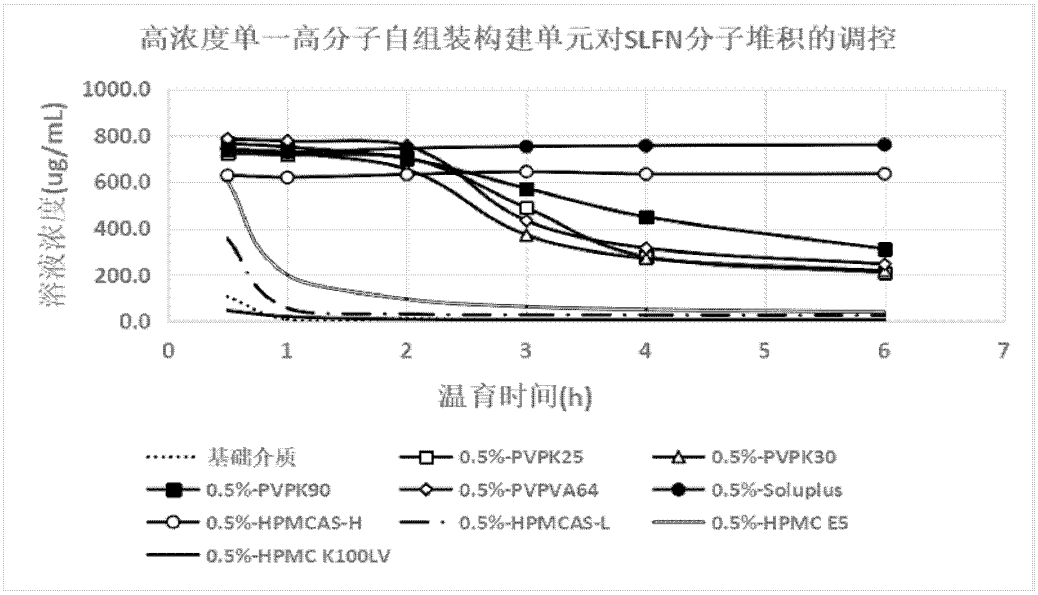

高浓度单一高分子自组装构建单元对SLFN分子堆积的调控
溶液浓度(ug/mL)
1000.0
800.0
600.0
400.0
200.0
0.0
0
1
2
3
4
5
6
7
温育时间(h)
基础介质
0.5%-PVPK25
0.5%-PVPK30
0.5%-PVPK90
0.5%-PVPVA64
0.5%-Soluplus
0.5%-HPMCAS-H
0.5%-HPMCAS-L
0.5%-HPMC E5
0.5%-HPMC K100LV

图 3

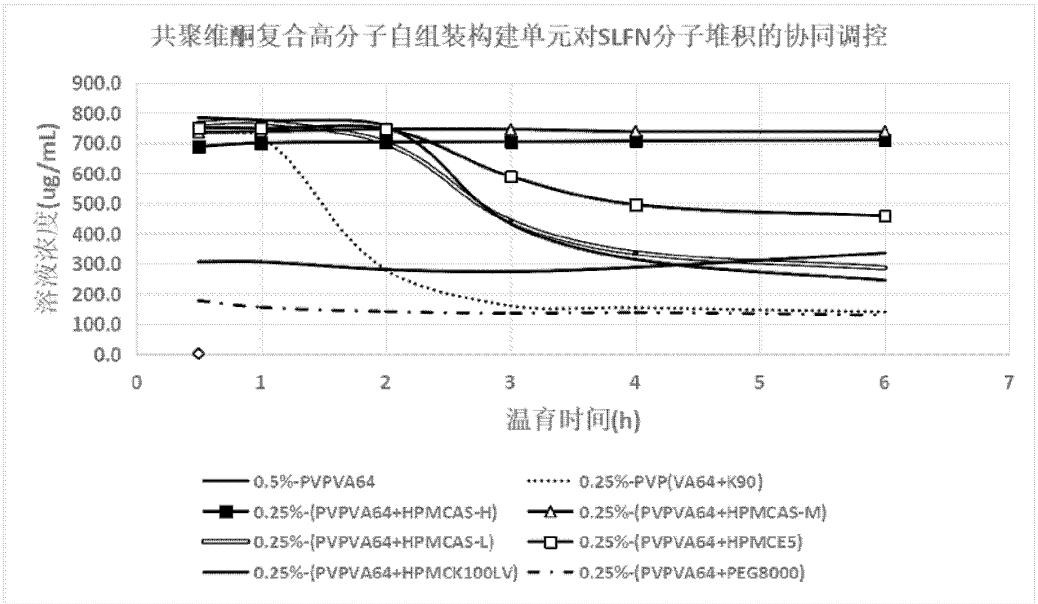

共聚维酮复合高分子自组装构建单元对SLFN分子堆积的协同调控
溶液浓度(ug/mL)
900.0
800.0
700.0
600.0
500.0
400.0
300.0
200.0
100.0
0.0
0
1
2
3
4
5
6
7
温育时间(h)
0.5%-PVPVA64
0.25%-PVP(VA64+K90)
0.25%-(PVPVA64+HPMCAS-H)
0.25%-(PVPVA64+HPMCAS-M)
0.25%-(PVPVA64+HPMCAS-L)
0.25%-(PVPVA64+HPMCE5)
0.25%-(PVPVA64+HPMCK100LV)
0.25%-(PVPVA64+PEG8000)

图 4

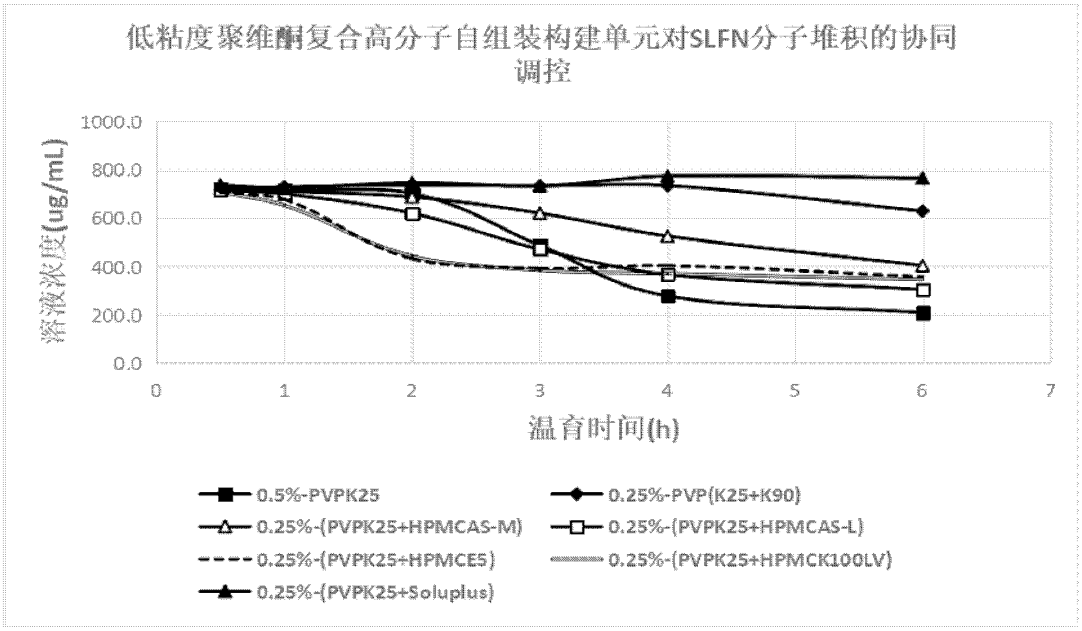

低粘度聚维酮复合高分子自组装构建单元对SLFN分子堆积的协同调控
溶液浓度(ug/mL)
1000.0
800.0
600.0
400.0
200.0
0.0
0
1
2
3
4
5
6
7
温育时间(h)
0.5%-PVPK25
0.25%-PVP(K25+K90)
0.25%-(PVPK25+HPMCAS-M)
0.25%-(PVPK25+HPMCAS-L)
0.25%-(PVPK25+HPMCE5)
0.25%-(PVPK25+HPMCK100LV)
0.25%-(PVPK25+Soluplus)

图 5

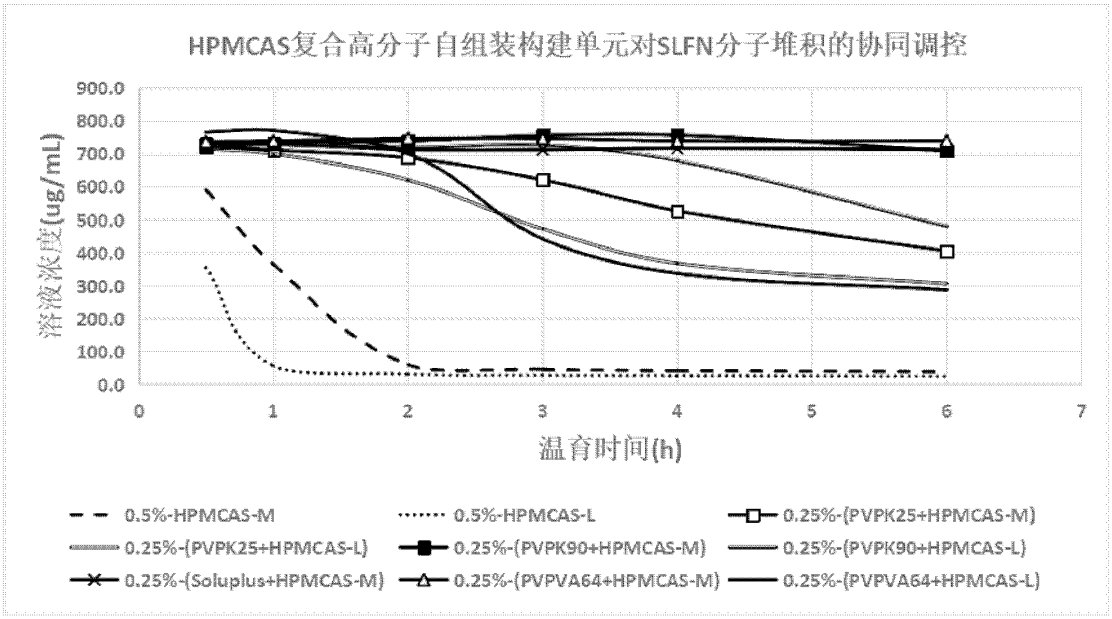

HPMCAS复合高分子自组装构建单元对SLFN分子堆积的协同调控
溶液浓度(ug/mL)
900.0
800.0
700.0
600.0
500.0
400.0
300.0
200.0
100.0
0.0
0
1
2
3
4
5
6
7
温育时间(h)
0.5%-HPMCAS-M
0.5%-HPMCAS-L
0.25%-(PVPK25+HPMCAS-M)
0.25%-(PVPK25+HPMCAS-L)
0.25%-(PVPK90+HPMCAS-M)
0.25%-(PVPK90+HPMCAS-L)
0.25%-(Soluplus+HPMCAS-M)
0.25%-(PVPVA64+HPMCAS-M)
0.25%-(PVPVA64+HPMCAS-L)

图 6

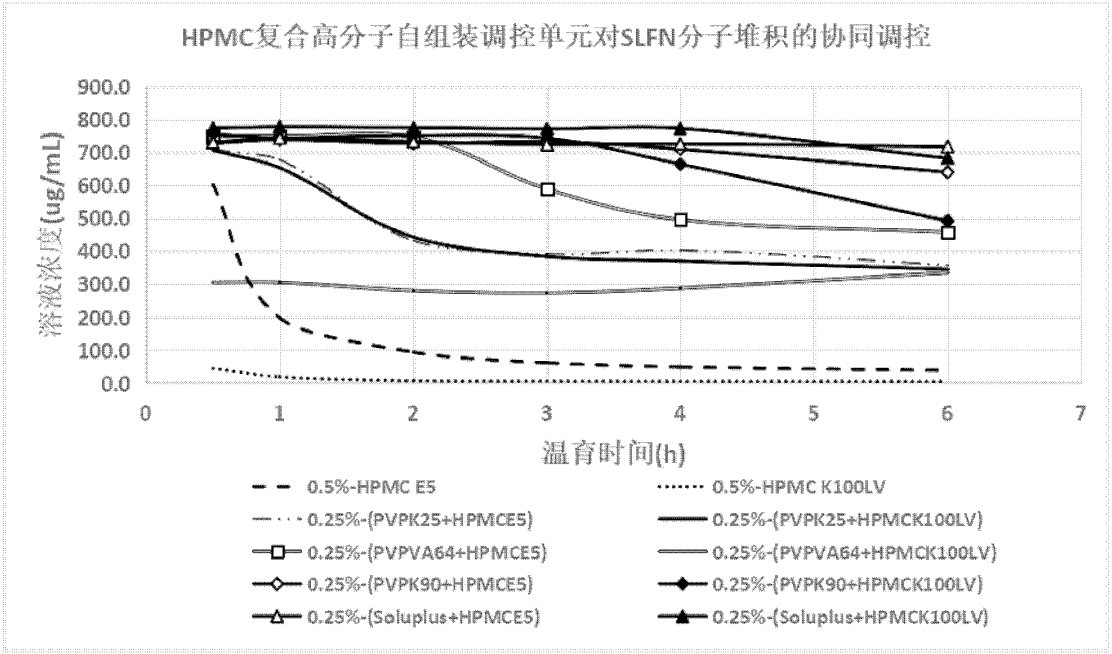

HPMC复合高分子自组装调控单元对SLFN分子堆积的协同调控
溶液浓度(ug/mL)
900.0
800.0
700.0
600.0
500.0
400.0
300.0
200.0
100.0
0.0
0
1
2
3
4
5
6
7
温育时间(h)
0.5%-HPMC E5
0.5%-HPMC K100LV
0.25%-(PVPK25+HPMCE5)
0.25%-(PVPK25+HPMCK100LV)
0.25%-(PVPVA64+HPMCE5)
0.25%-(PVPVA64+HPMCK100LV)
0.25%-(PVPK90+HPMCE5)
0.25%-(PVPK90+HPMCK100LV)
0.25%-(Soluplus+HPMCE5)
0.25%-(Soluplus+HPMCK100LV)

图 7

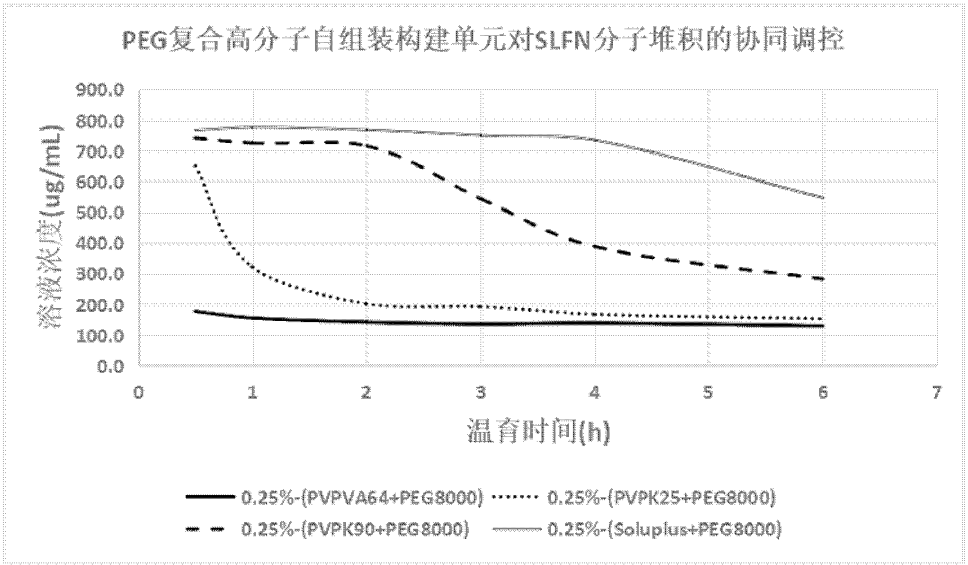

PEG复合高分子自组装构建单元对SLFN分子堆积的协同调控
溶液浓度(ug/mL)
900.0
800.0
700.0
600.0
500.0
400.0
300.0
200.0
100.0
0.0
0
1
2
3
4
5
6
7
温育时间(h)
0.25%-(PVPVA64+PEG8000)
0.25%-(PVPK25+PEG8000)
0.25%-(PVPK90+PEG8000)
0.25%-(Soluplus+PEG8000)

图 8

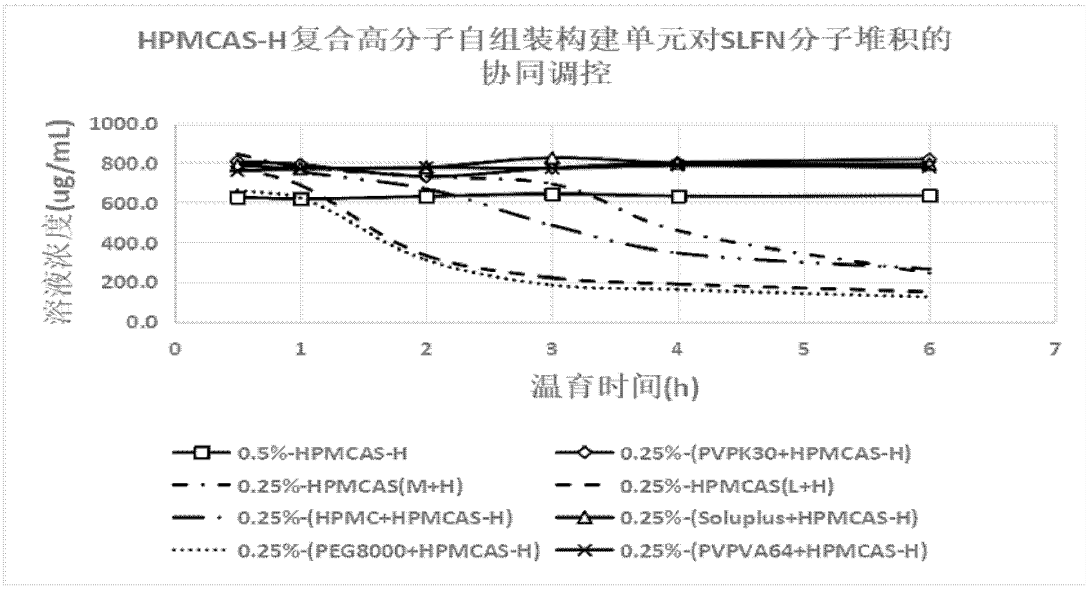

HPMCAS-H复合高分子自组装构建单元对SLFN分子堆积的协同调控
溶液浓度(ug/mL)
1000.0
800.0
600.0
400.0
200.0
0.0
0
1
2
3
4
5
6
7
温育时间(h)
0.5%-HPMCAS-H
0.25%-(PVPK30+HPMCAS-H)
0.25%-HPMCAS(M+H)
0.25%-HPMCAS(L+H)
0.25%-(HPMC+HPMCAS-H)
0.25%-(Soluplus+HPMCAS-H)
0.25%-(PEG8000+HPMCAS-H)
0.25%-(PVPVA64+HPMCAS-H)

图 9

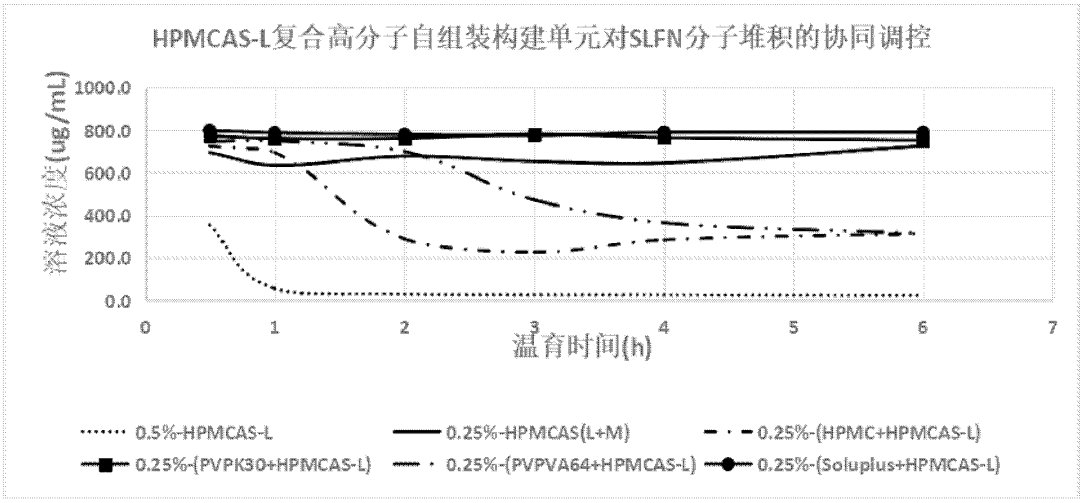

HPMCAS-L复合高分子自组装构建单元对SLFN分子堆积的协同调控
溶液浓度(ug/mL)
1000.0
800.0
600.0
400.0
200.0
0.0
0
1
2
3
4
5
6
7
温育时间(h)
0.5%-HPMCAS-L
0.25%-HPMCAS(L+M)
0.25%-(HPMC+HPMCAS-L)
0.25%-(PVPK30+HPMCAS-L)
0.25%-(PVPVA64+HPMCAS-L)
0.25%-(Soluplus+HPMCAS-L)

图 10

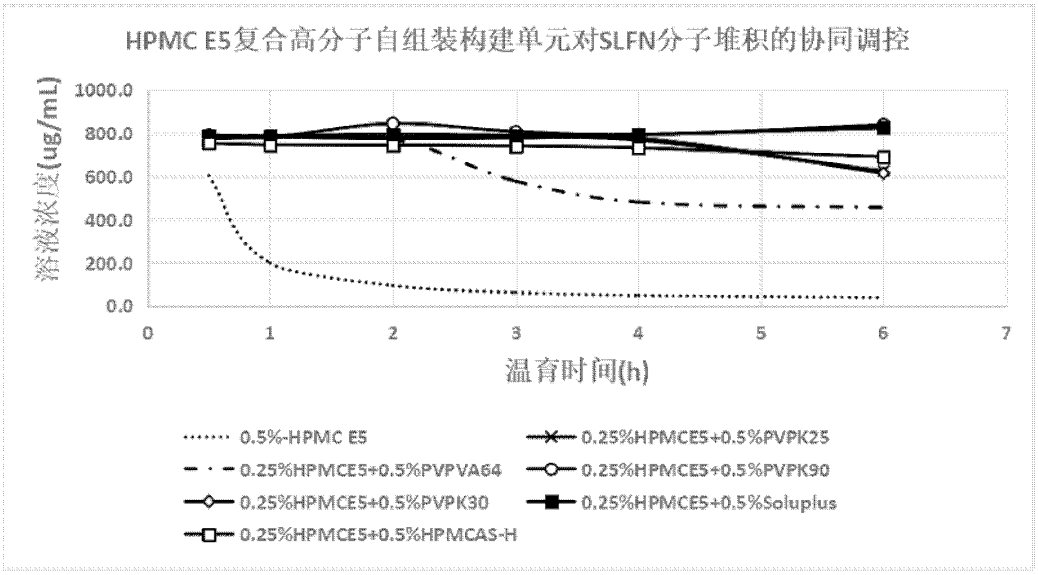

HPMC E5复合高分子自组装构建单元对SLFN分子堆积的协同调控
溶液浓度(ug/mL)
1000.0
800.0
600.0
400.0
200.0
0.0
0
1
2
3
4
5
6
7
温育时间(h)
0.5%-HPMC E5
0.25%HPMCE5+0.5%PVPK25
0.25%HPMCE5+0.5%PVPVA64
0.25%HPMCE5+0.5%PVPK90
0.25%HPMCE5+0.5%PVPK30
0.25%HPMCE5+0.5%Soluplus
0.25%HPMCE5+0.5%HPMCAS-H

图 11

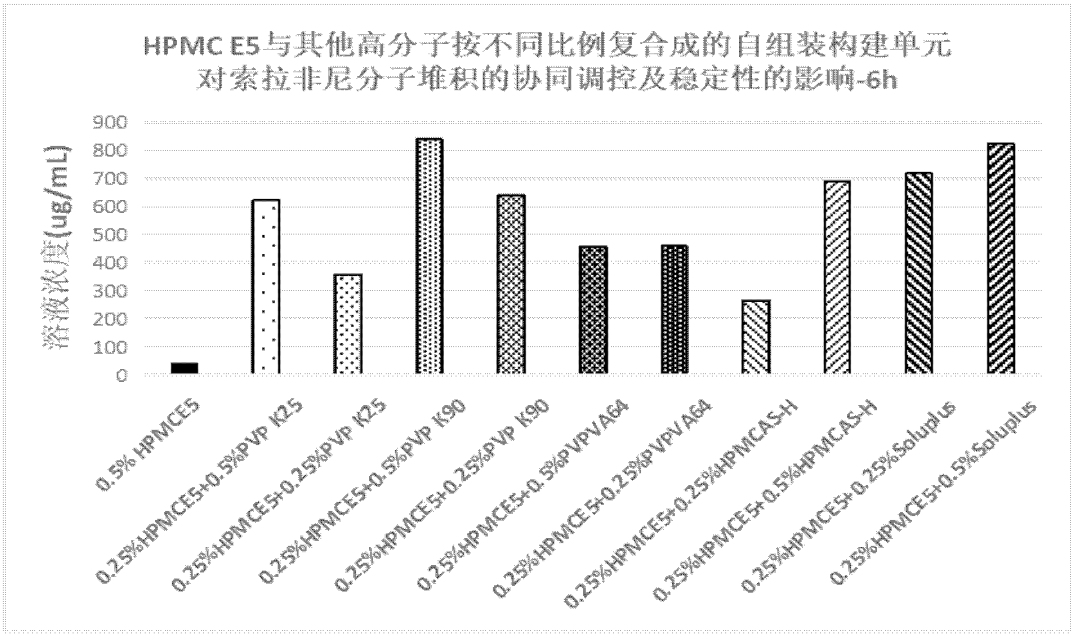

HPMC E5与其他高分子按不同比例复合成的自组装构建单元
对索拉非尼分子堆积的协同调控及稳定性的影响-6h
溶液浓度(ug/mL)
900
800
700
600
500
400
300
200
100
0
0.5% HPMCE5
0.25%HPMCE5+0.5%PVP K25
0.25%HPMCE5+0.25%PVP K25
0.25%HPMCE5+0.5%PVP K90
0.25%HPMCE5+0.25%PVP K90
0.25%HPMCE5+0.5%PVPVA64
0.25%HPMCE5+0.25%PVPVA64
0.25%HPMCE5+0.25%HPMCAS-H
0.25%HPMCE5+0.5%HPMCAS-H
0.25%HPMCE5+0.25%Soluplus
0.25%HPMCE5+0.5%Soluplus

图 12

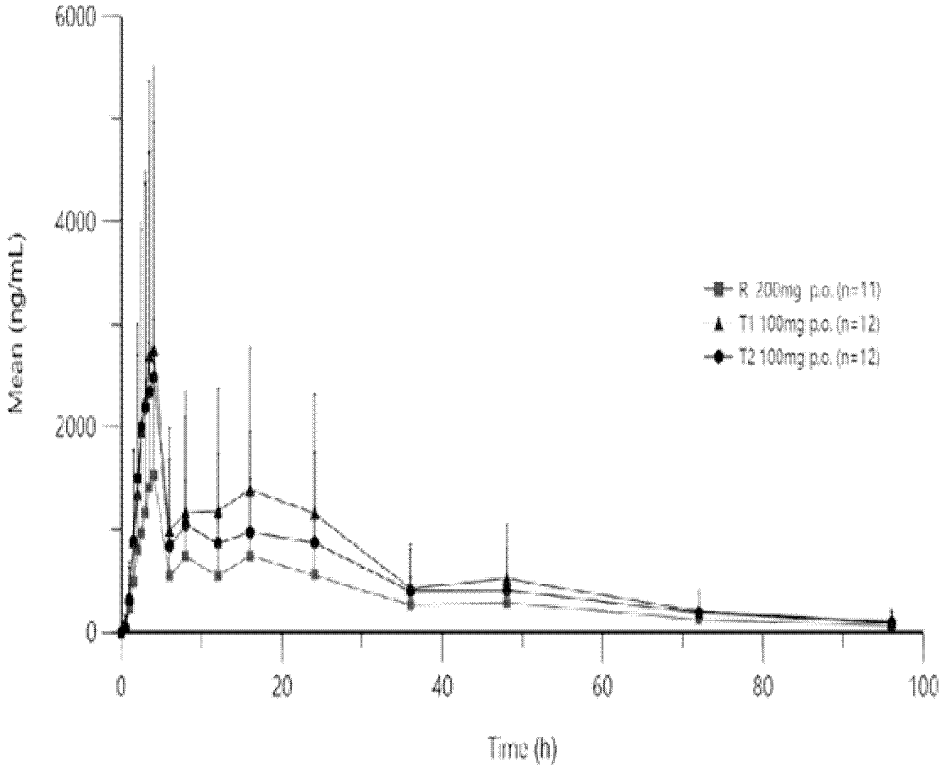

6000
4000
2000
0
Mean (ng/mL)
0
20
40
60
80
100
Time (h)
R 200mg p.o. (n=11)
T1 100mg p.o. (n=12)
T2 100mg p.o. (n=12)

图 13

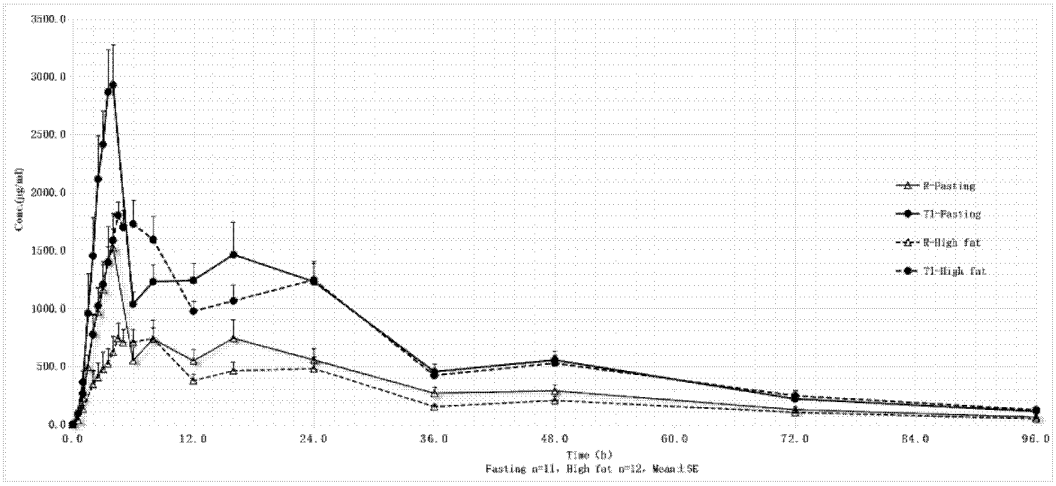

3500.0
3000.0
2500.0
2000.0
1500.0
1000.0
500.0
0.0
Conc.(μg/ml)
0.0
12.0
24.0
36.0
48.0
60.0
72.0
84.0
96.0
Time (h)
Fasting n=11, High fat n=12, Mean±SE
R-Fasting
T1-Fasting
R-High fat
T1-High fat

图 14