

SUPERINTENDENCIA DE INDUSTRIA Y COMERCIO



No. 06-129487- -00012-0000

Fecha: 2011-04-06 16:44:29 Dep. 2020 DIR.NUEVASCR
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JEFE DE DIVISIÓN DE NUEVAS CREACIONES

SUPERINTENDENCIA DE INDUSTRIA Y COMERCIO

E.

S.

D.

REF: Expediente No. 06 129 487 – solicitud de patente de invención a favor de GENENTECH INC.

Respuesta al oficio No. 15384 del 26 de noviembre de 2010

JUAN PABLO CADENA SARMIENTO, portador de la tarjeta profesional No. 57.492 del C.S.J., obrando como apoderado de la sociedad GENENTECH INC., según lo acredita el documento de poder presentado ante esa División me permito dar respuesta a las observaciones de carácter técnico y jurídico, relacionadas con la patentabilidad, para la concesión de la patente de acuerdo con la Decisión 486 según el concepto técnico No. 3694 notificado por fijación en lista de fecha 26 de noviembre de 2010 y cuyo término fue prorrogado el 17 de febrero de 2011.

En respuesta a los comentarios realizados por el Señor Examinador, adjuntamos un nuevo capítulo reivindicatorio de 10 reivindicaciones, que subsana las objeciones a la presente solicitud sin ampliar la materia inicialmente divulgada y con sustento adecuado en la descripción. Es preciso tener en cuenta que este nuevo capítulo representa una restricción sustancial a la materia anteriormente reclamada y que cada uno de los compuestos que abarca el alcance de las nuevas reivindicaciones se encuentra debidamente soportado en la descripción.

Al diseñar este nuevo capítulo reivindicatorio se tuvo en cuenta cada uno de los radicales mencionados por el Señor Examinador así como los soportados en la descripción. Sin embargo, se ha realizado una generalización razonable

del anillo A para brindar adecuada protección al desarrollo alcanzado por los inventores ya que los mismos se encuentran en el alcance de la descripción y además comparten la misma estructura molecular, actividad y afinidad por el dominio BIR3 de proteínas X-IAP siendo efectivos en el tratamiento del cáncer. En este sentido, consideramos que la amplitud de las posibilidades del anillo A está totalmente justificada y solicitamos entonces su aceptación.

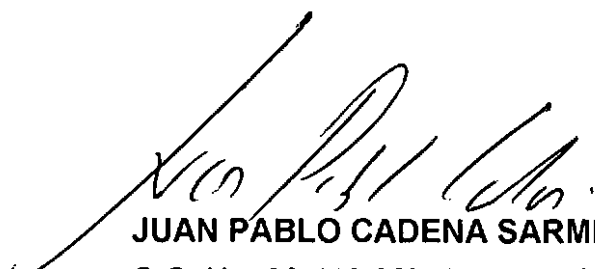
Para dar mayor soporte a lo anterior, adjuntamos a este memorial un listado de 287 compuestos con sus respectivas estructuras moleculares, los cuales incluyen las diferentes definiciones del anillo A en la nueva reivindicación 1, demostrando de esta forma la equivalencia que existe entre ellas. Además, se observa que todos los compuestos listados prueban afinidad por el dominio BIR3 de proteínas X-IAP, presentan valores de K_i que hace a cada uno de ellos un potente inhibidor de estas proteínas y por ende un potencial fármaco efectivo en el tratamiento del cáncer quedando en total sustento la nueva limitación del anillo A. Esta afinidad fue determinada al calcular la constante de afinidad nanomolar K_i a partir de los valores IC_{50} , mediante el método de polarización de fluorescencia descrito en la página 137 de la presente solicitud.

De lo anterior, es claro que al compartir esta actividad, afinidad y estructura molecular todos los compuestos probados por el solicitante justifican el mantenimiento de la amplitud reivindicada para el anillo A.

Adicionalmente, consideramos que, al no existir documento alguno en el arte previo que se acerque o pueda afectar la patentabilidad de la presente solicitud, todos los requisitos de patentabilidad se cumplen adecuadamente. Más aún, enfatizamos que, de acuerdo con los datos presentados, todos los compuestos desarrollados por los inventores constituyen un aporte sustancial al estado de la técnica tanto en el presente como para futuros desarrollos en la búsqueda de agentes que logren combatir el cáncer. En este sentido, la protección sobre este desarrollo es vital por lo que solicitamos al Señor Examinador el retiro de sus objeciones.

En vista de lo anteriormente expuesto, respetuosamente solicitamos el retiro de las objeciones a la presente solicitud y la concesión del privilegio solicitado, ya que en opinión del solicitante cumple con todos los requisitos establecidos en la decisión 486.

Del señor Director,

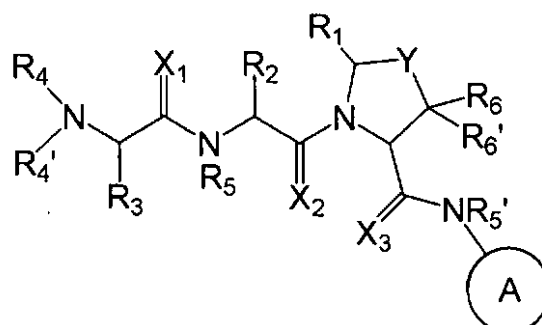


JUAN PABLO CADENA SARMIENTO
C.C. No. 80.410.337 de Usaquén
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Calle 70A No. 4-41

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REIVINDICACIONES

1. Un compuesto de fórmula I:



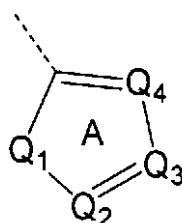
I

en donde

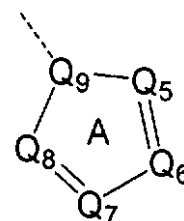
X₁, X₂ y X₃ son O;

Y es CH₂ o S;

A tiene la fórmula IIa o IIb



IIa

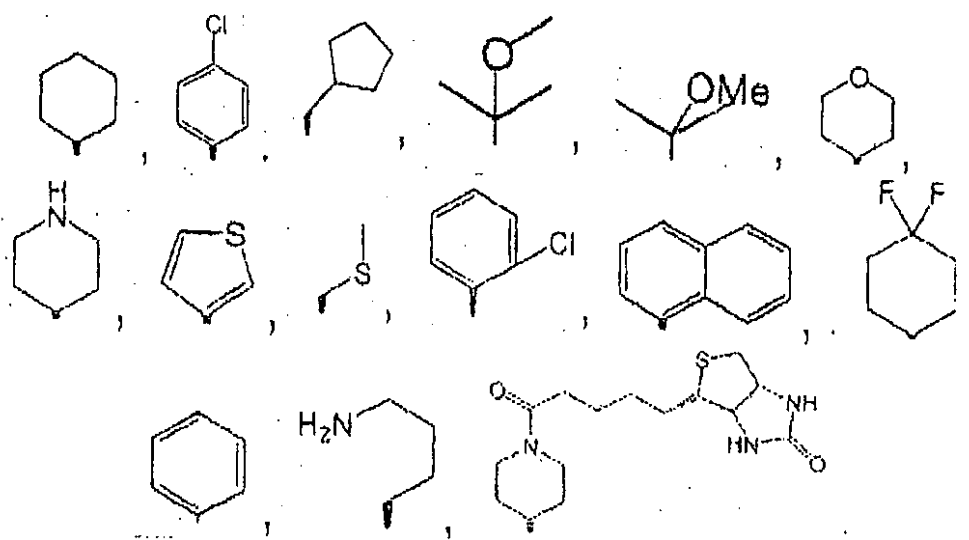


IIb

En donde Q₁ es NR₈, O o S; Q₂, Q₃, Q₄, Q₅, Q₆, Q₇, y Q₈ son independientemente CR₉ o N; en donde R₉ es H, amino, hidroxilo, mercapto, halógeno, carboxil, amidino, guanidino, alquil, alcoxi, arilo, ariloxi, acil, aciloxi, acilamino, cicloalquilo o un heterociclo; en donde cada sustitución alquilo, alcoxi, arilo, ariloxi, acil, aciloxi, acilamino, cicloalquilo y heterociclo es opcionalmente sustituida con hidroxilo, halógeno, mercapto, carboxilo, alquilo, haloalquilo, amino, nitro, cicloalquilo, arilo o un heterociclo; R₈ es H, alquilo, acil, arilo, cicloalquilo o un heterociclo; en donde cada alquilo, arilo, cicloalquilo y heterociclo es opcionalmente sustituido con hidroxilo, halógeno, mercapto, carboxilo, alquil, haloalquilo, amino, nitro, cicloalquilo, arilo o un heterociclo; y Q₉ es CH o N

R₁ es H;

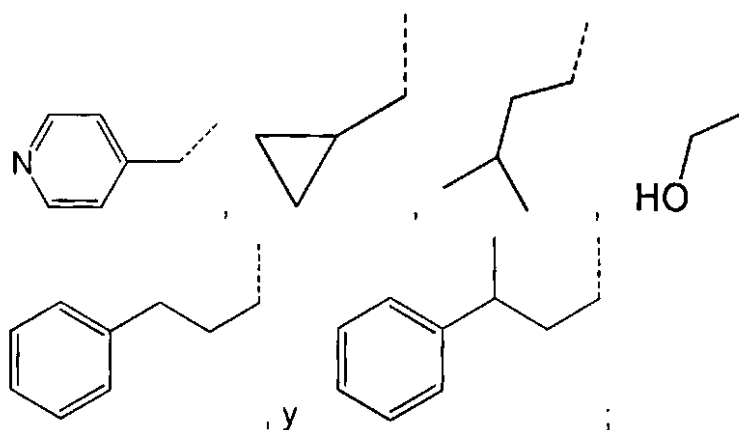
R₂ es isopropilo, CF₃,



R_3 es CH_3 o CH_2CH_3 ;

R_4 es H o CH_3 ;

R_4' es H, CH_3 ,



R_5 , y R_5' son cada uno H;

R_6 , y R_6' son cada uno independientemente H o CH_3 ;

y sales de los mismos.

2. El compuesto de acuerdo con la reivindicación 1, caracterizado porque

X_1 , X_2 , y X_3 son cada uno O;

Y es CH_2 ;

R_1 es H;

R_2 es ciclohexil, tetrahidropiran-4-il o fenilo;

R_3 es metilo;

R_4 is metilo;

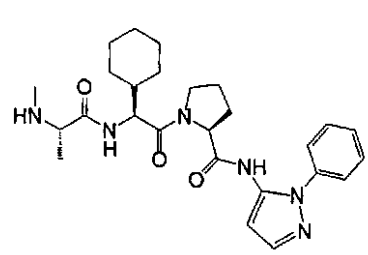
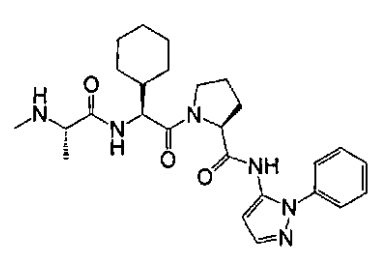
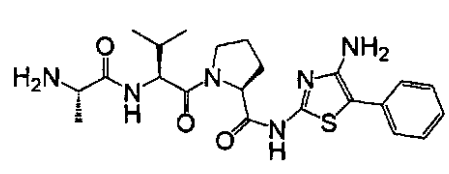
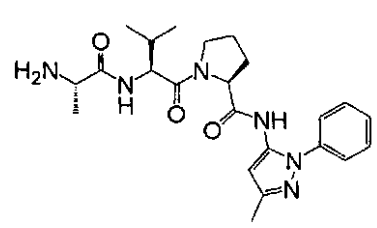
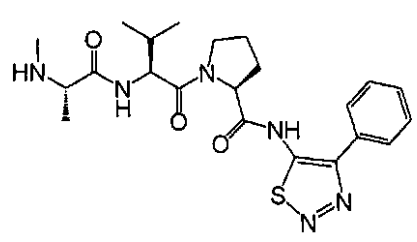
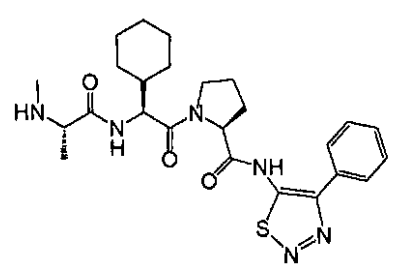
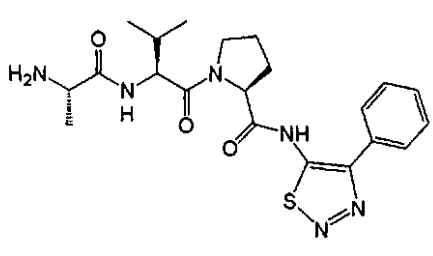
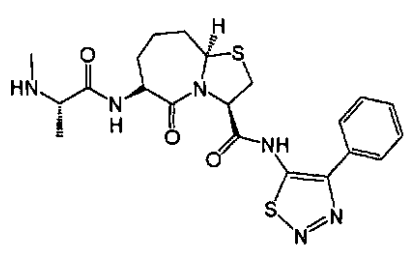
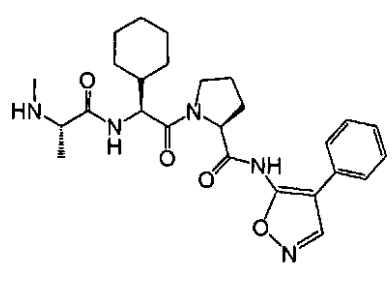
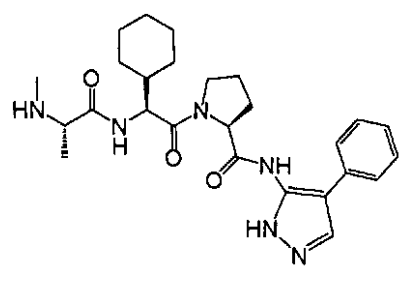
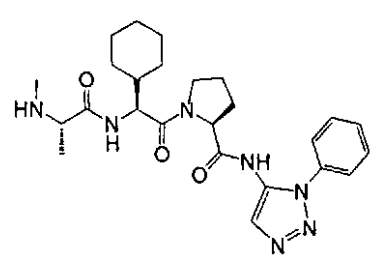
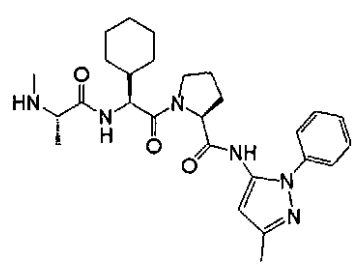
R_4' es H;

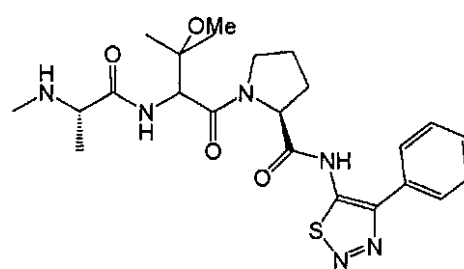
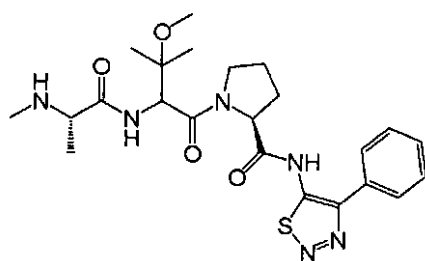
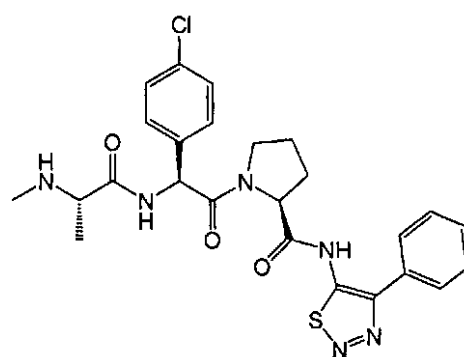
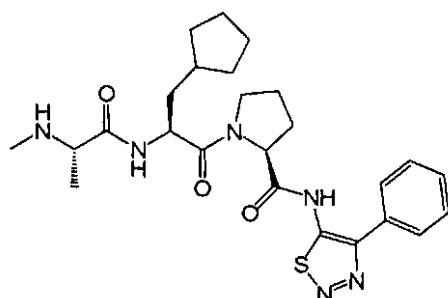
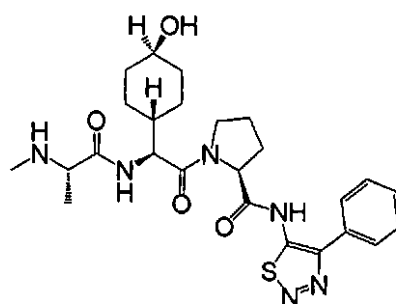
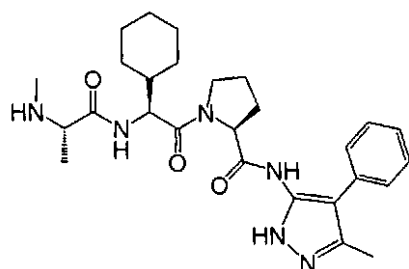
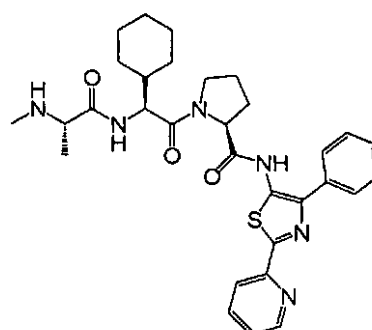
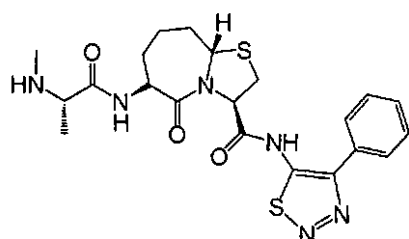
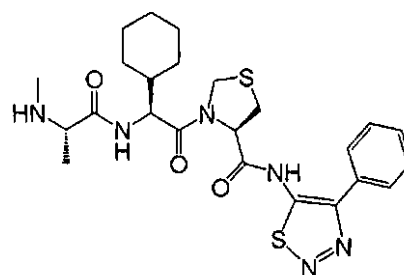
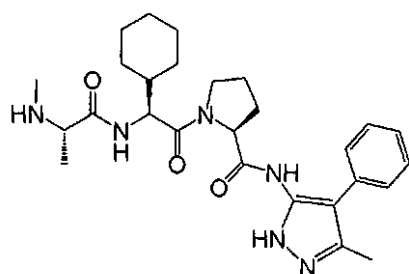
R_5 y R_5' son ambos H; y

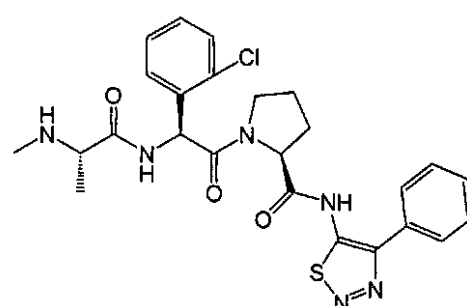
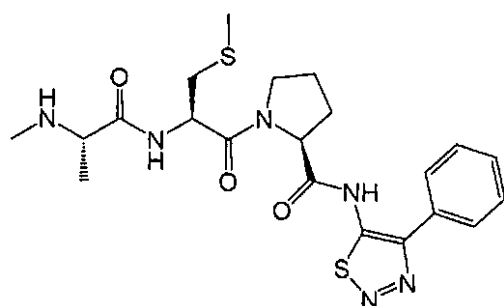
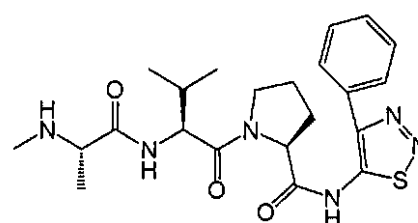
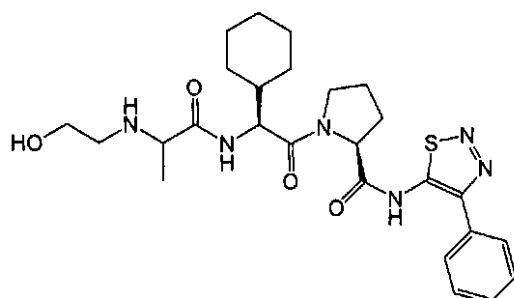
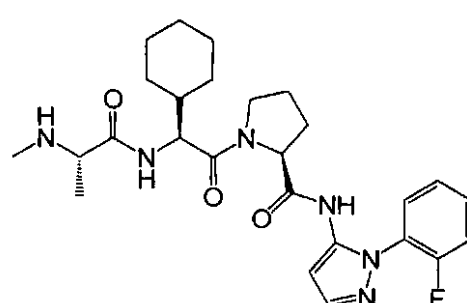
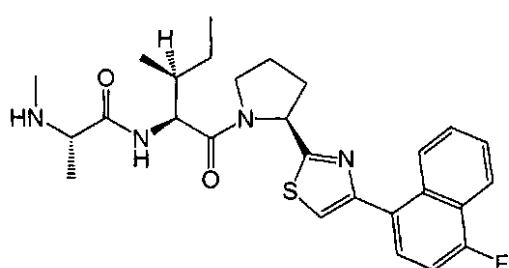
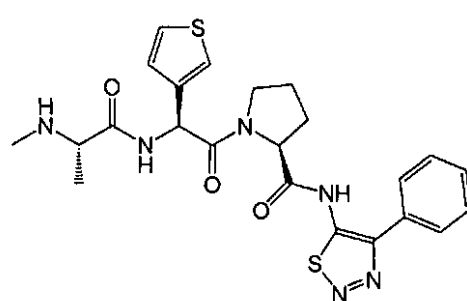
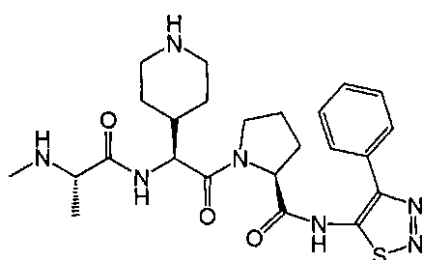
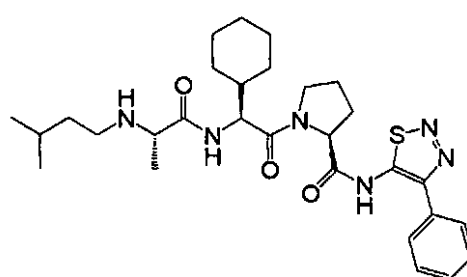
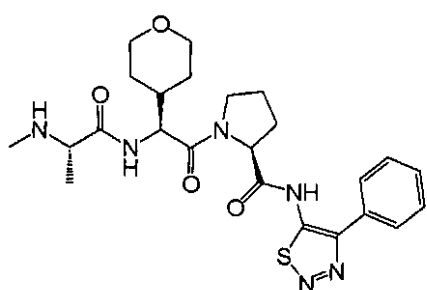
R_6 y R_6' son ambos H.

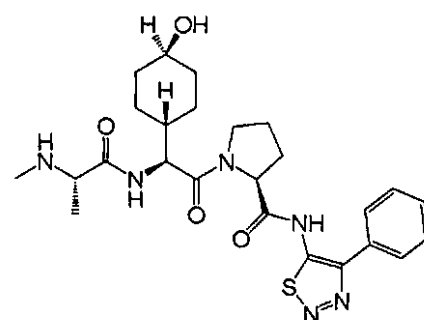
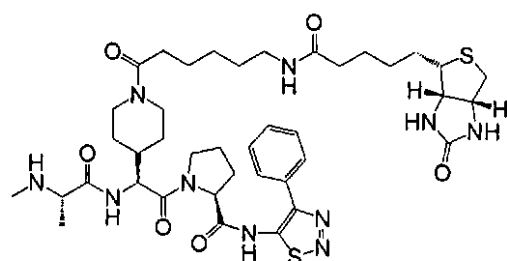
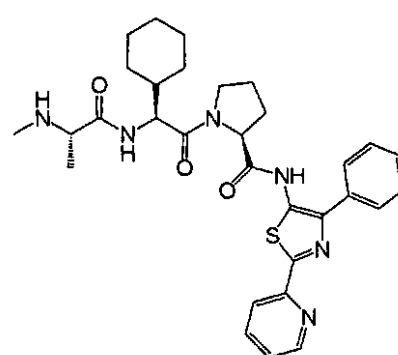
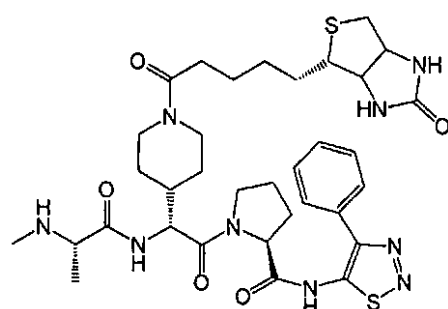
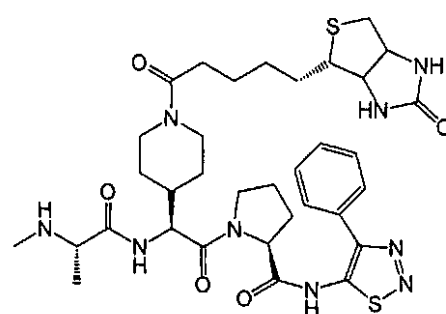
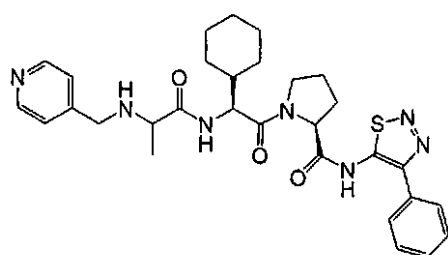
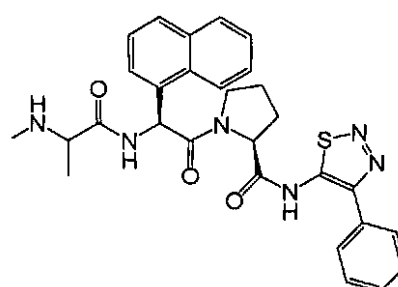
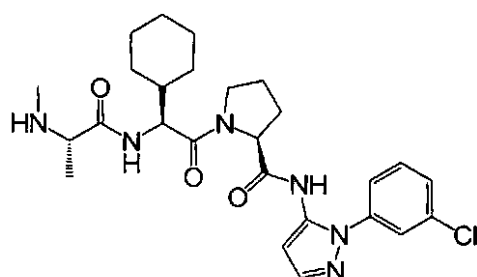
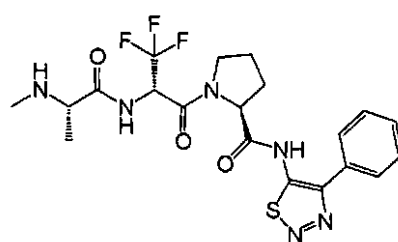
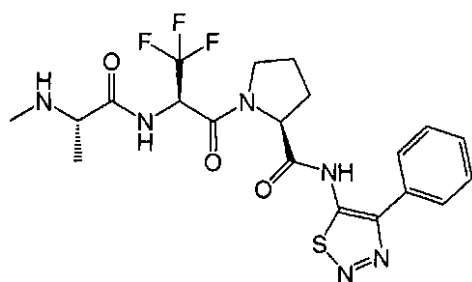
3. El compuesto de acuerdo con la reivindicación 2, caracterizado porque el anillo A es un tiofeno opcionalmente sustituido con alquilo, arilo, aralquilo, cicloalquilo, cicloalquilalquilo, un heterociclo o un heterociclo-alquil opcionalmente sustituido con halógeno hidroxilo, mercapto, carboxilo, alquilo, haloalquilo, amino, nitro, arilo o heteroarilo.
4. El compuesto de acuerdo con la reivindicación 2, caracterizado porque el anillo A es pirazol opcionalmente sustituido con alquilo, arilo, aralquilo, cicloalquilo, cicloalquilalquilo, un heterociclo o un heterociclo-alquilo opcionalmente sustituido con halógeno hidroxilo, mercapto, carboxilo, alquilo, haloalquilo, amino, nitro, arilo o heteroarilo.
5. El compuesto de acuerdo con la reivindicación 2, caracterizado porque el anillo A es oxazol opcionalmente sustituido con alquilo, arilo, aralquilo, cicloalquilo, cicloalquilalquilo, un heterociclo o un heterociclo-alquilo opcionalmente sustituido con halógeno hidroxilo, mercapto, carboxilo, alquilo, haloalquilo, amino, nitro, arilo o heteroarilo.
6. El compuesto de acuerdo con la reivindicación 2, caracterizado porque el anillo A es isoxazol opcionalmente sustituido con alquilo, arilo, aralquilo, cicloalquilo, cicloalquilalquil, un heterociclo o un heterociclo-alquilo opcionalmente sustituido con halógeno hidroxilo, mercapto, carboxilo, alquilo, haloalquilo, amino, nitro, arilo o heteroarilo.
7. El compuesto de acuerdo con la reivindicación 2, caracterizado porque el anillo A es tiazol opcionalmente sustituido con alquilo, arilo, aralquilo, cicloalquilo, cicloalquilalquilo, un heterociclo o un heterociclo-alquilo opcionalmente sustituido con halógeno hidroxilo, mercapto, carboxilo, alquilo, haloalquilo, amino, nitro, arilo o heteroarilo.
8. El compuesto de acuerdo con la reivindicación 2, caracterizado porque el anillo A es isotiazol opcionalmente sustituido con alquilo, arilo, aralquilo, cicloalquilo, cicloalquilalquilo, un heterociclo o un heterociclo-alquilo opcionalmente sustituido con halógeno hidroxilo, mercapto, carboxilo, alquilo, haloalquilo, amino, nitro, arilo o heteroarilo.
9. El compuesto de acuerdo con la reivindicación 2, caracterizado porque el anillo A es thiadiazol opcionalmente sustituido con alquilo, arilo, aralquilo, cicloalquilo, cicloalquilalquilo, un heterociclo o un heterociclo-alquilo opcionalmente sustituido con halógeno hidroxilo, mercapto, carboxilo, alquilo, haloalquilo, amino, nitro, arilo o heteroarilo.

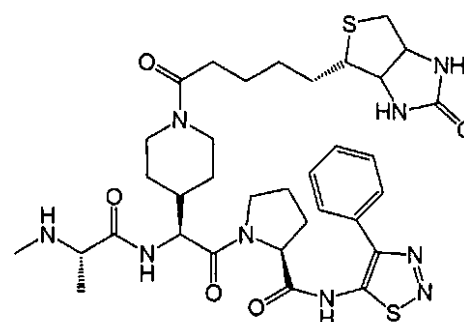
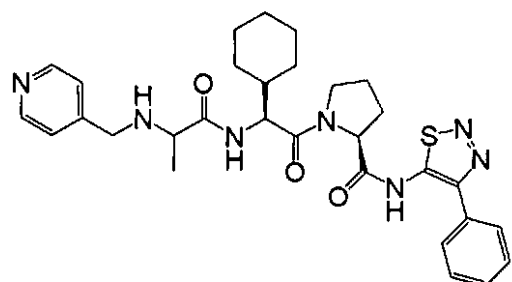
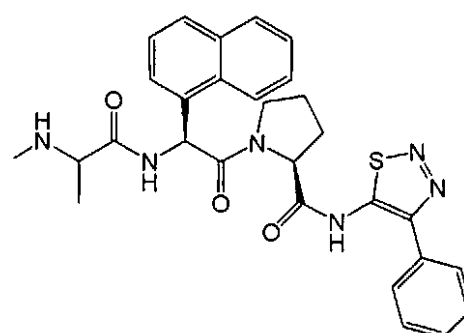
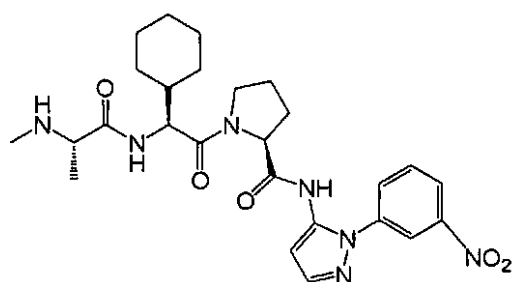
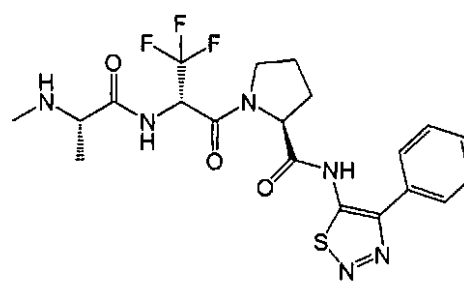
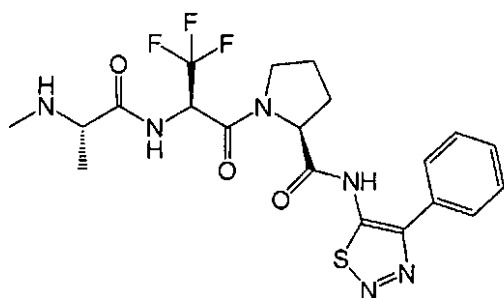
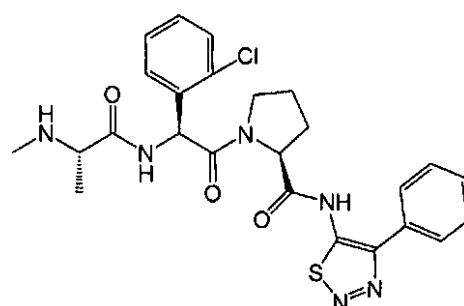
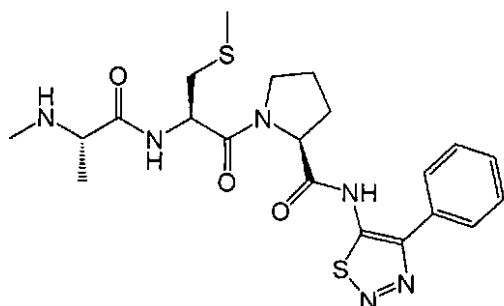
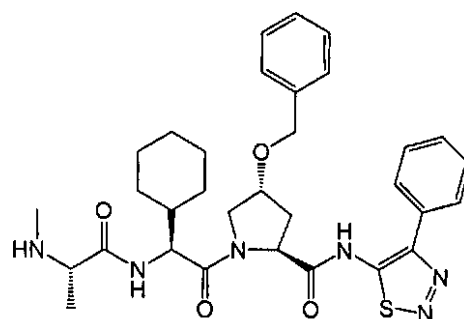
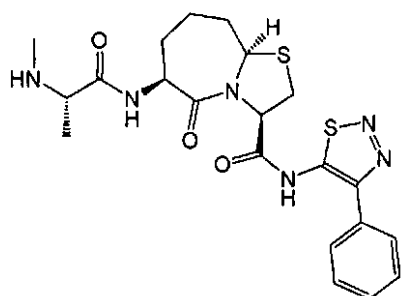
10. Un compuesto de acuerdo con la reivindicación 1 seleccionado del grupo que consiste de:

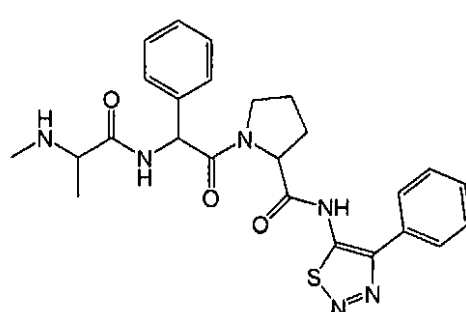
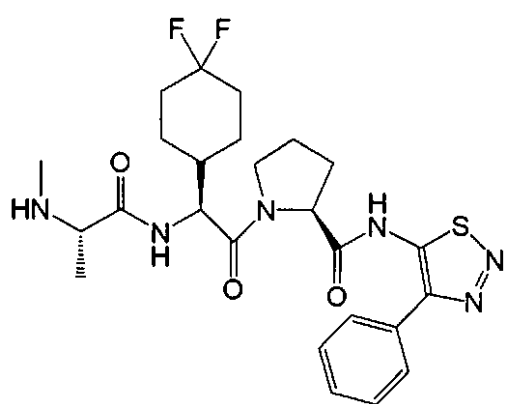
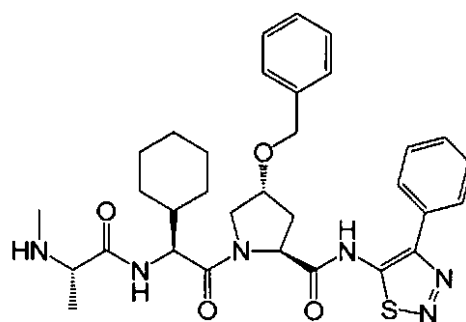
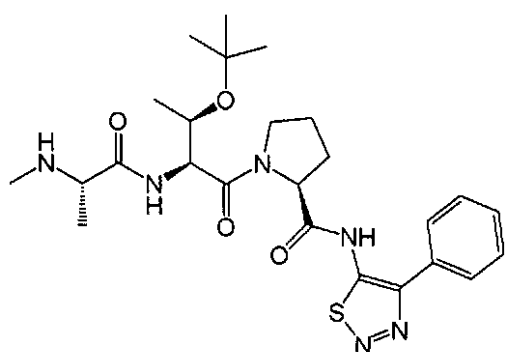
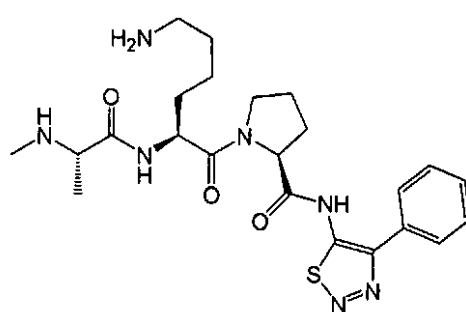
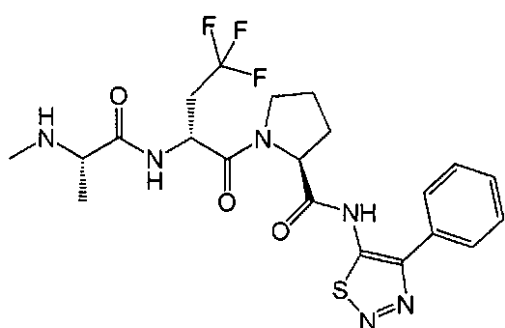
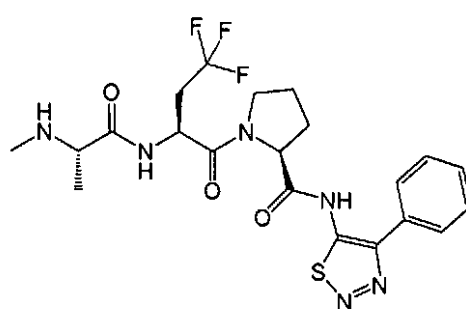
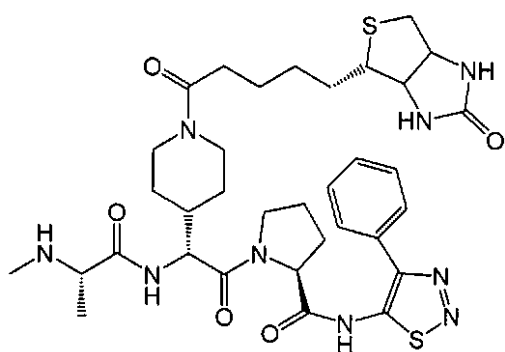


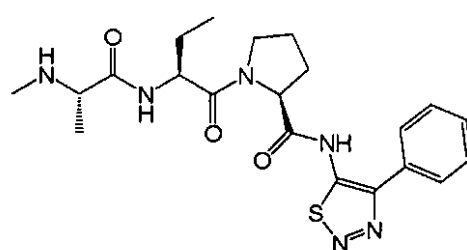
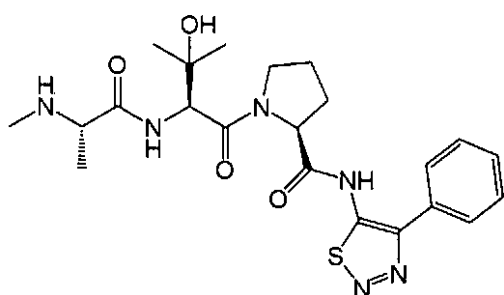
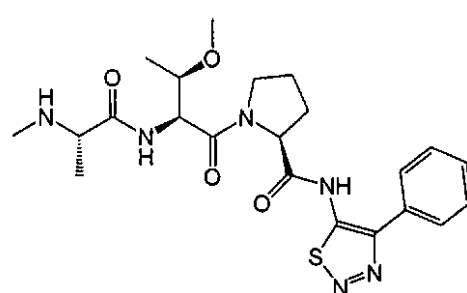
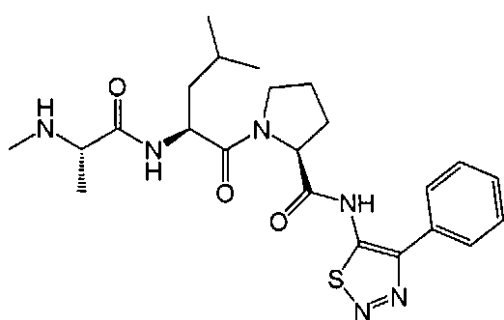
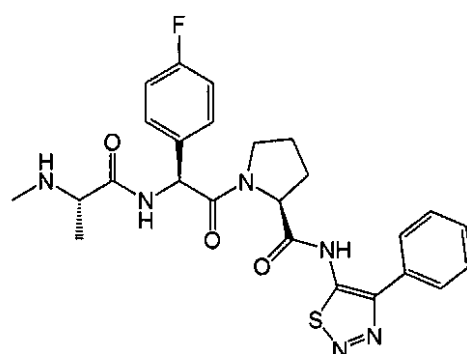
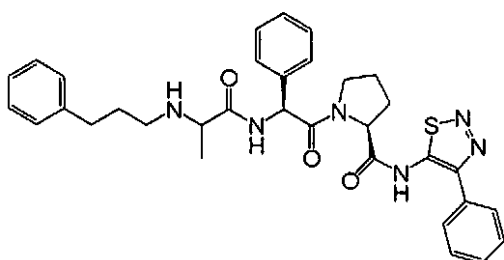
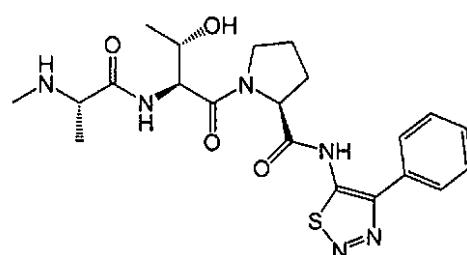
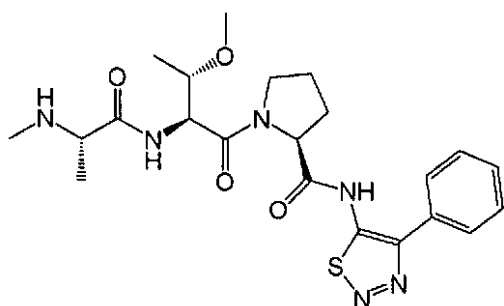
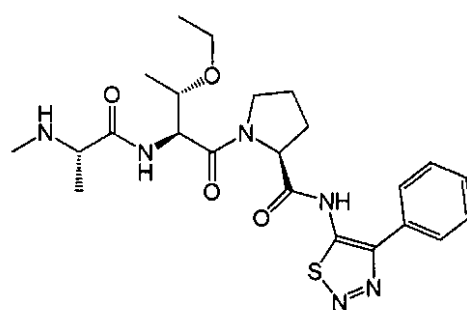
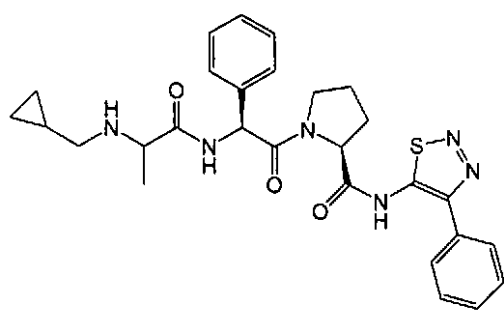


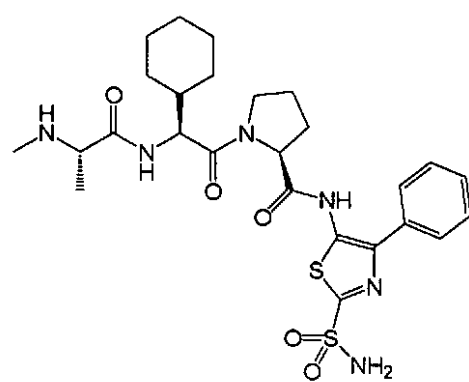
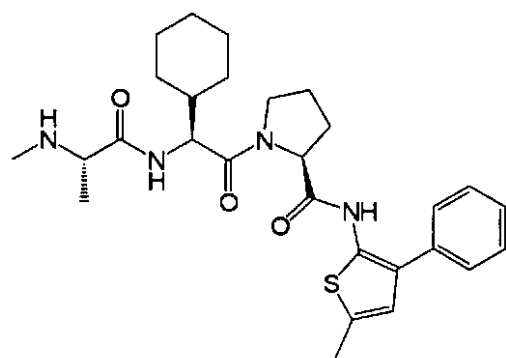
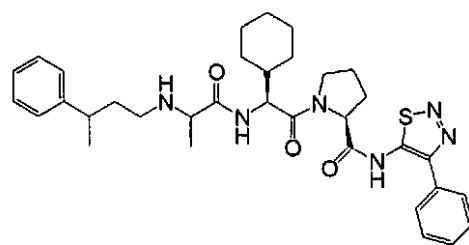
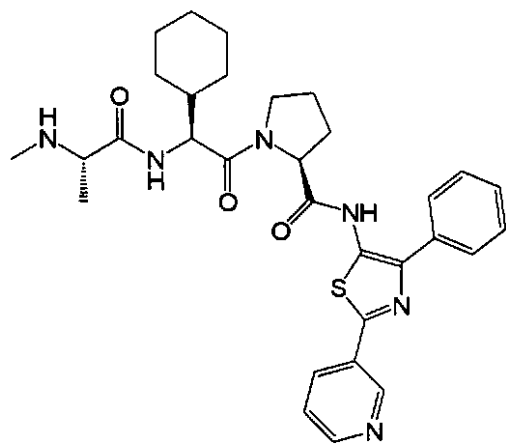
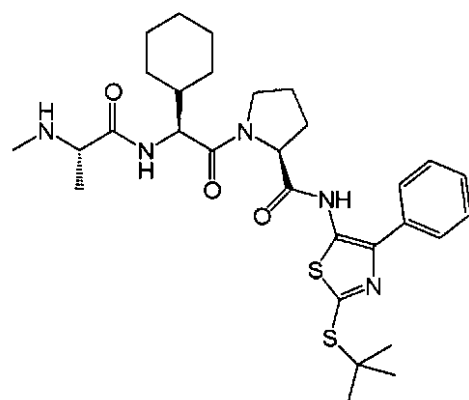
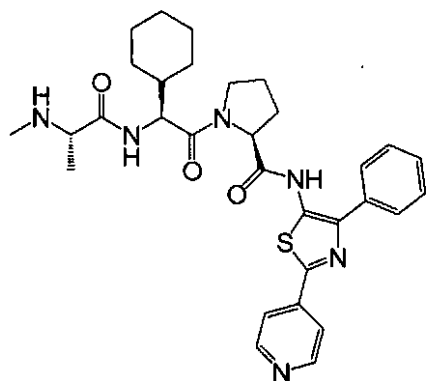
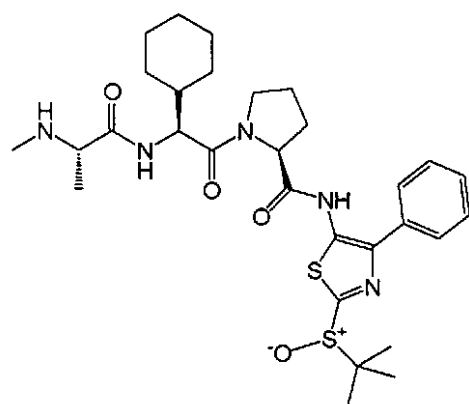
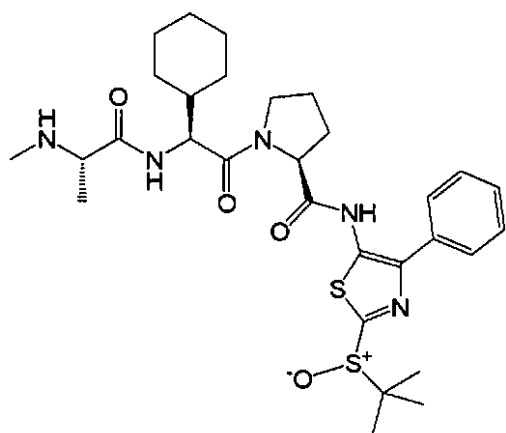




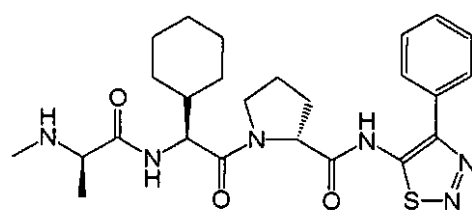
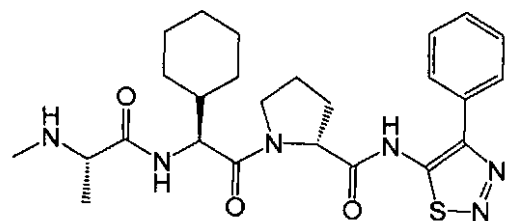
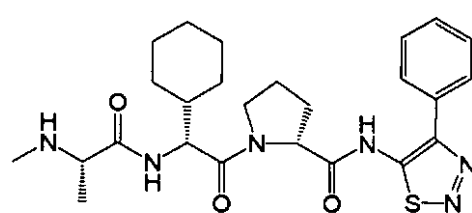
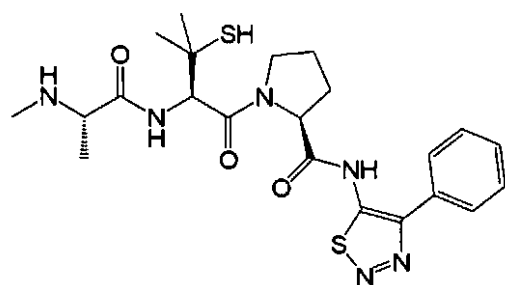
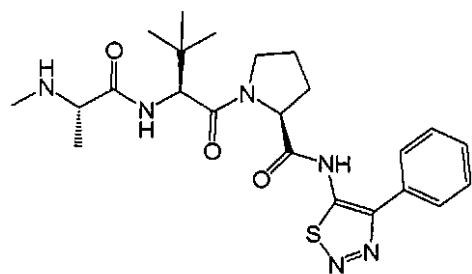
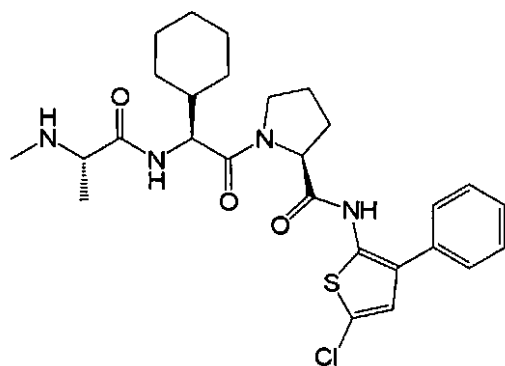
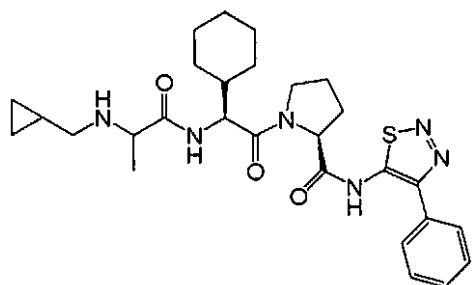
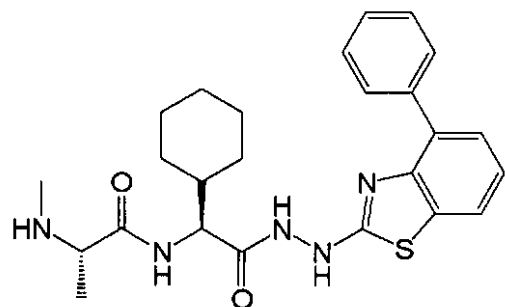
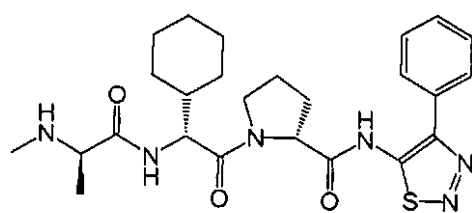
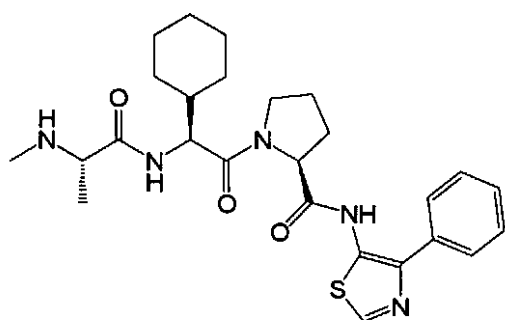


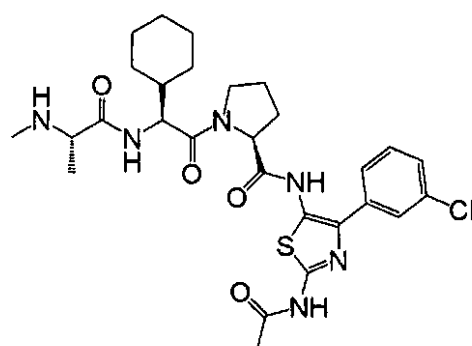
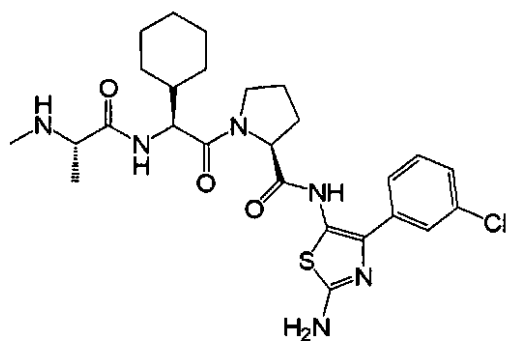
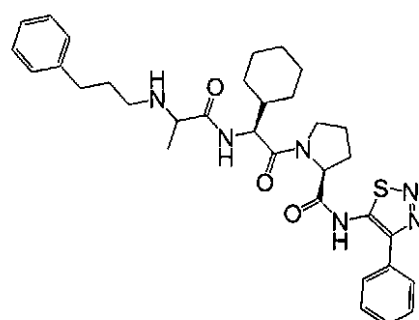
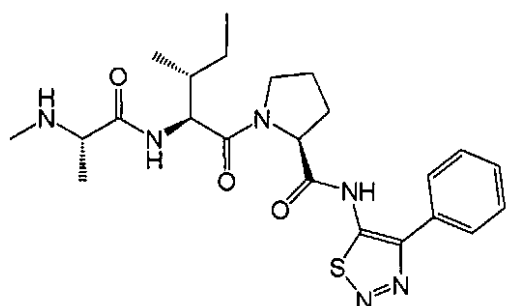
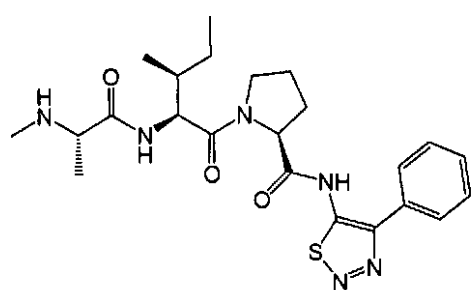
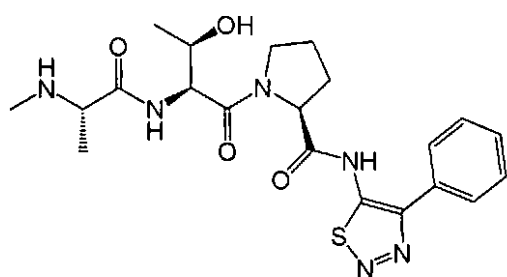
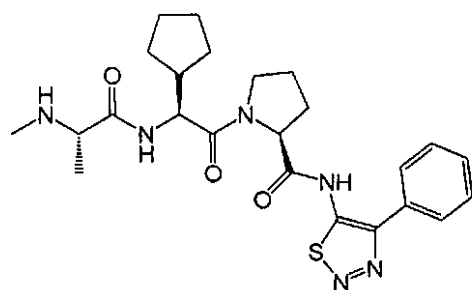
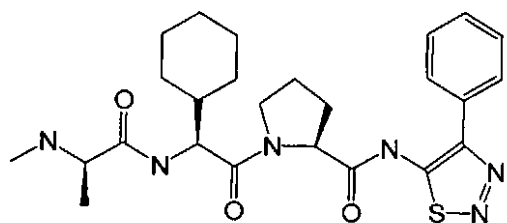
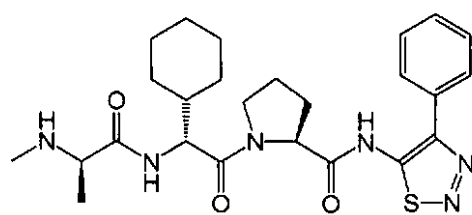
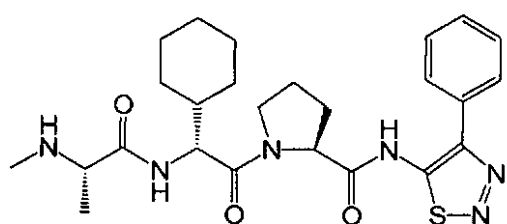


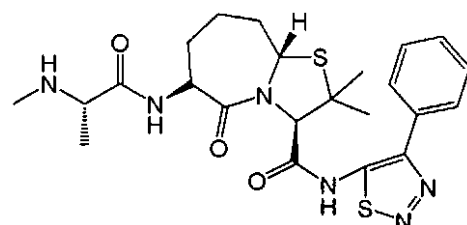
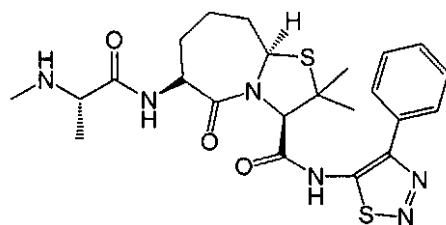
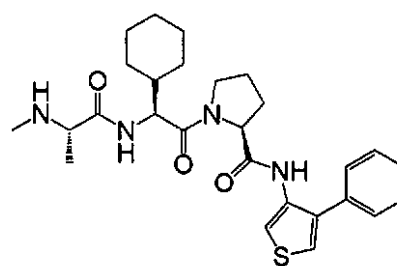
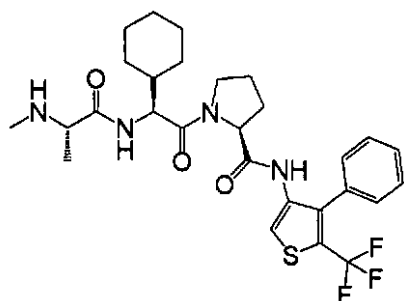
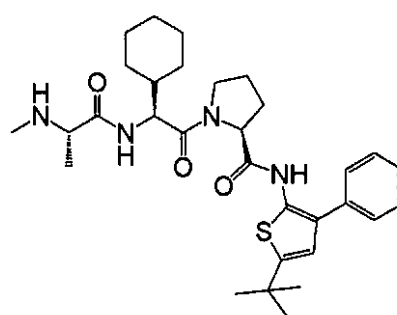
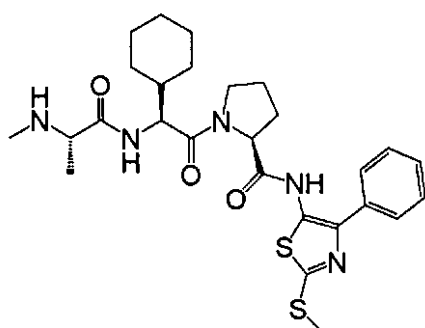
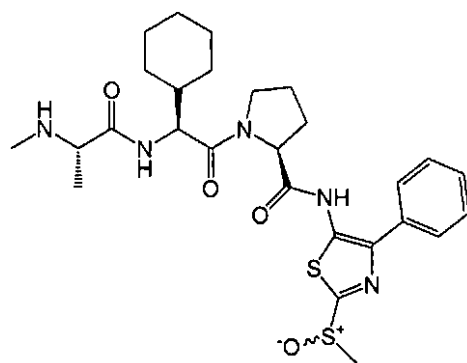
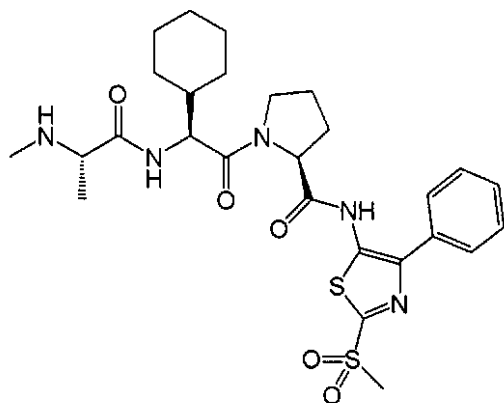
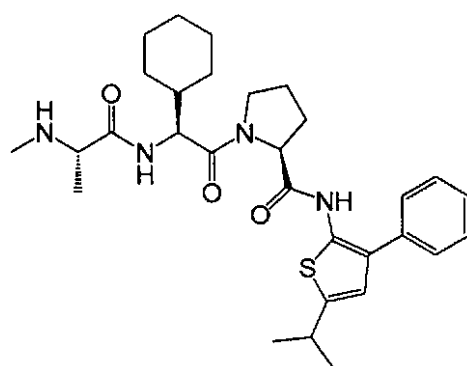
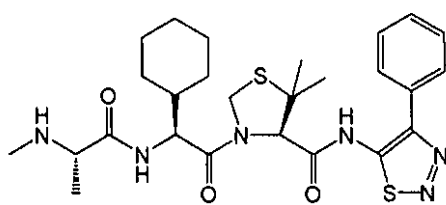


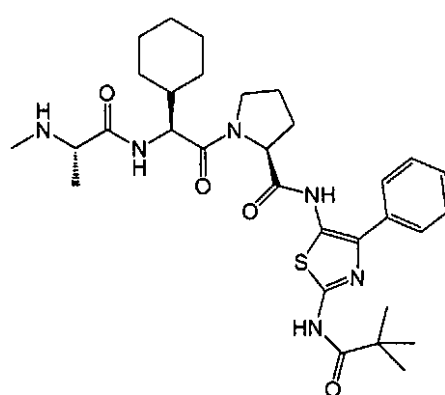
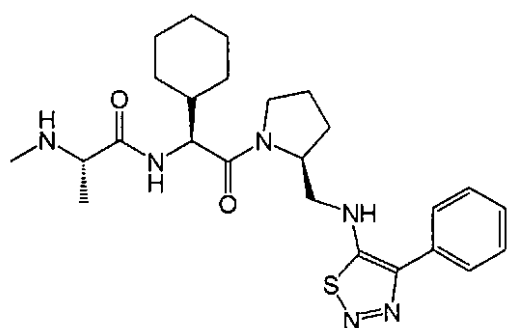
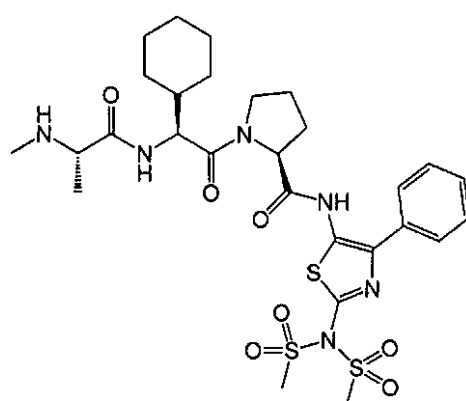
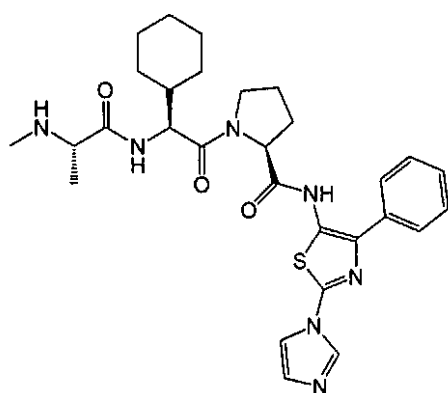
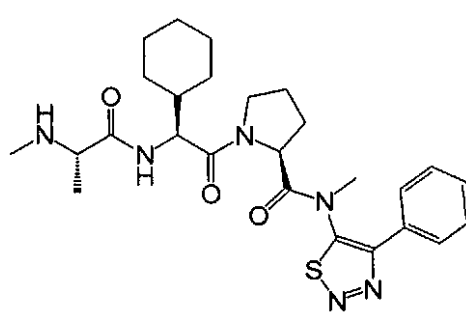
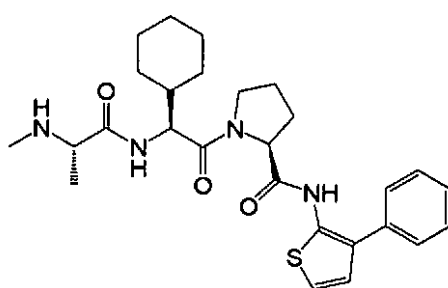
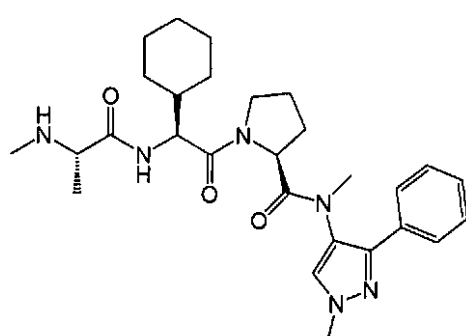
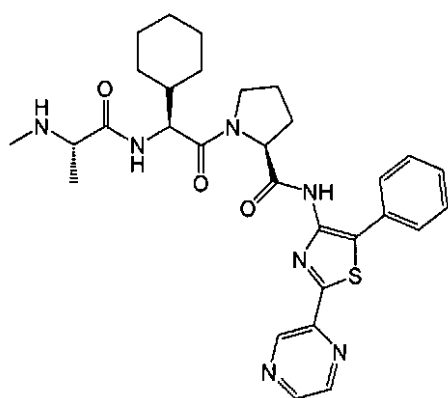


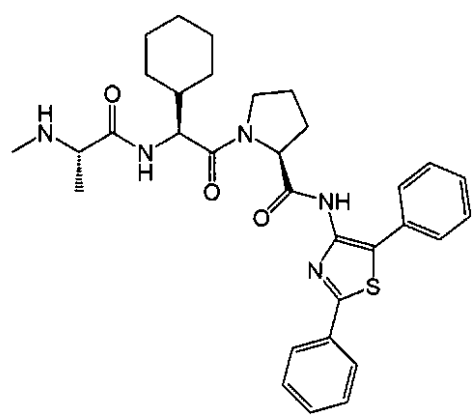
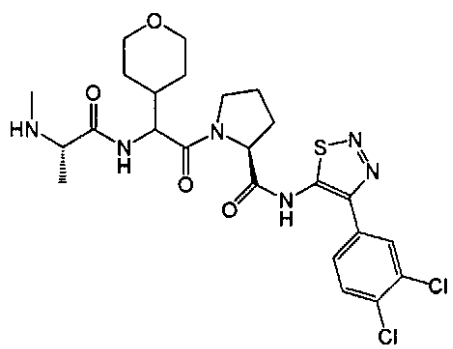
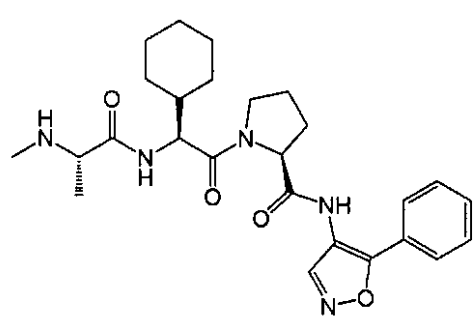
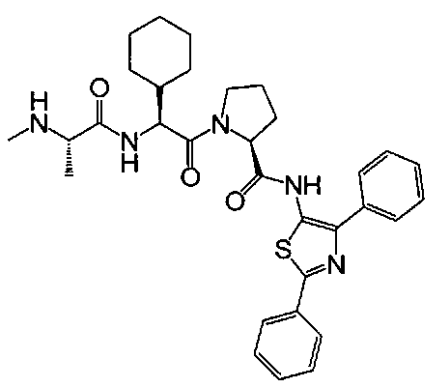
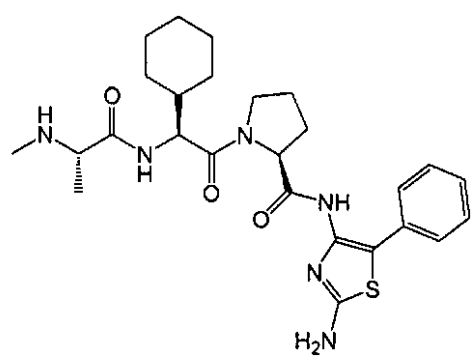
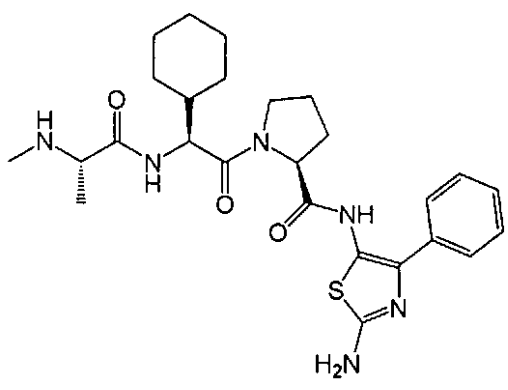
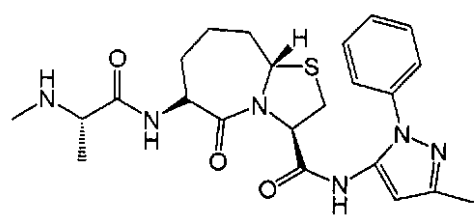
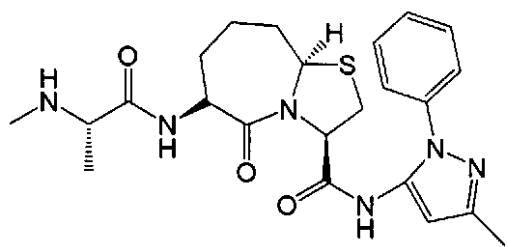
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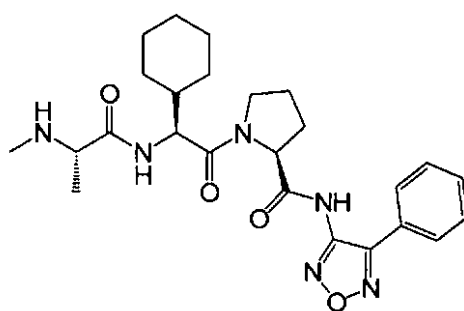
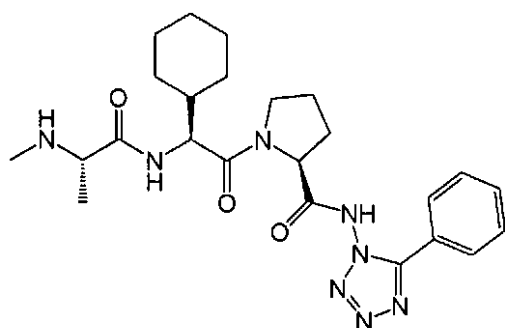
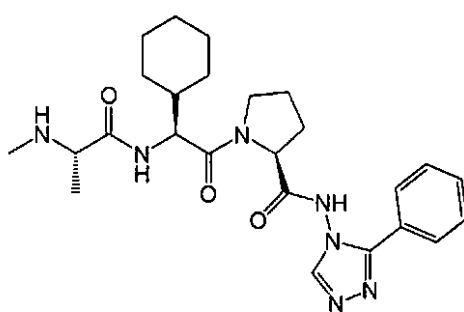
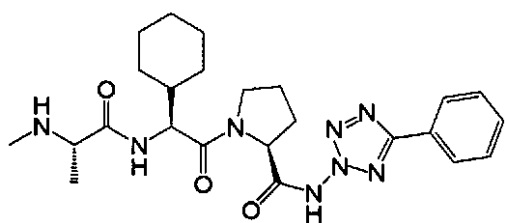
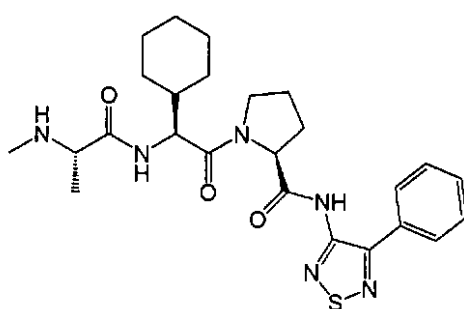
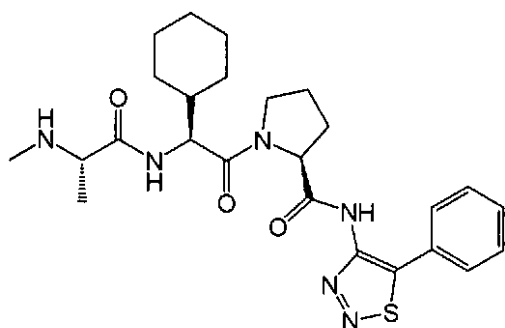
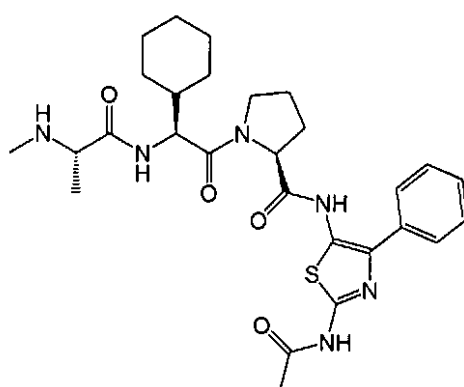
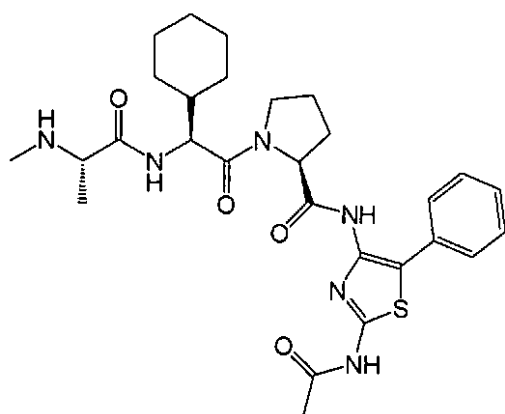


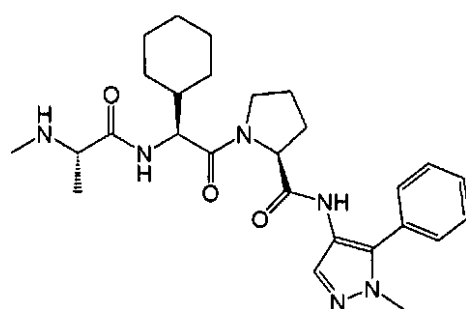
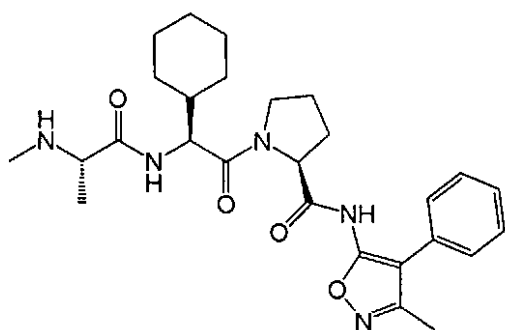
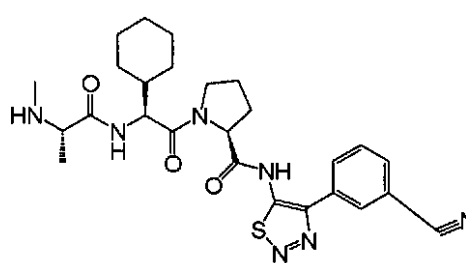
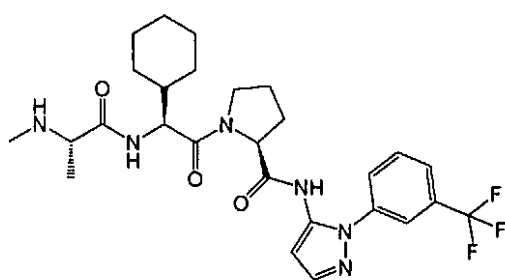
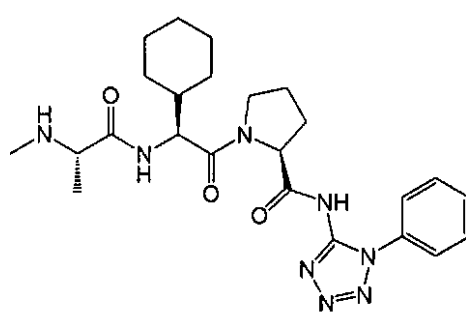
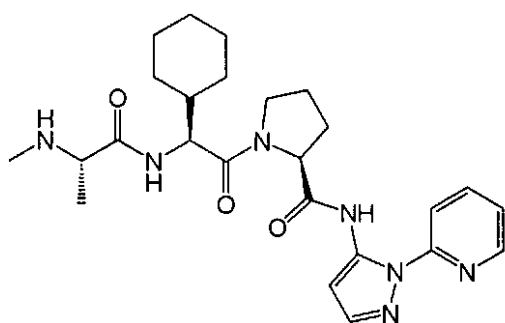
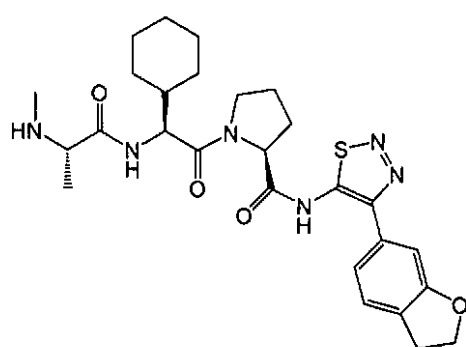
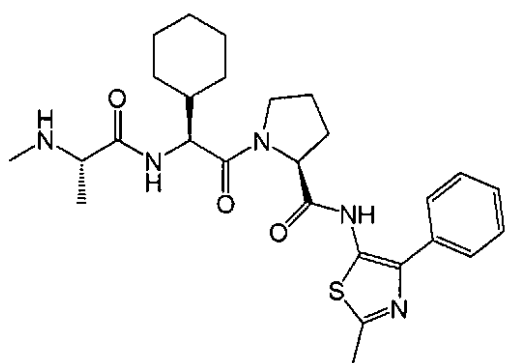


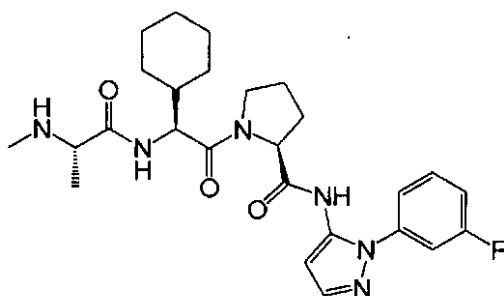
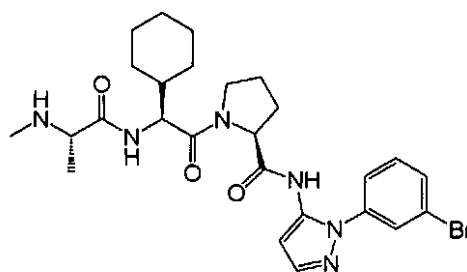
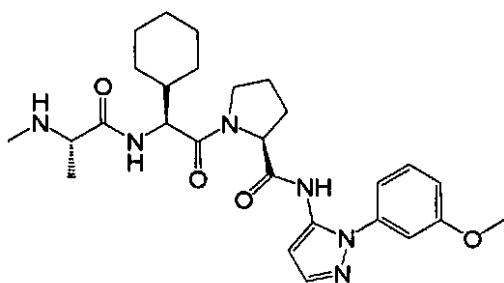
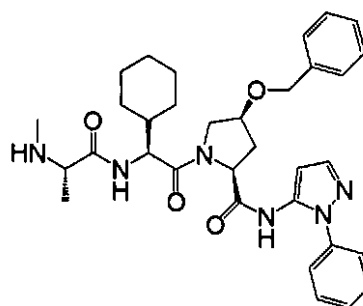
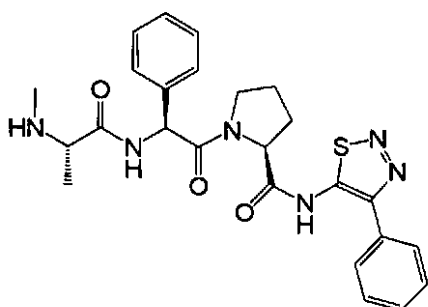
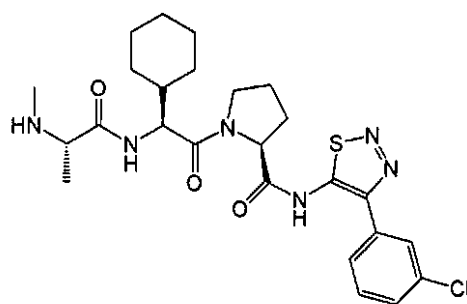
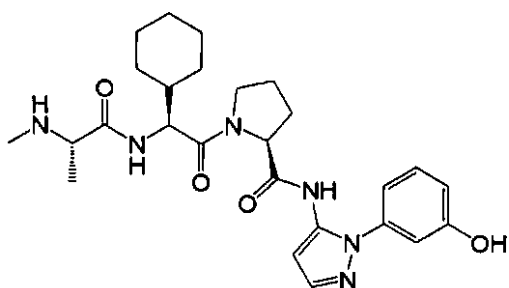
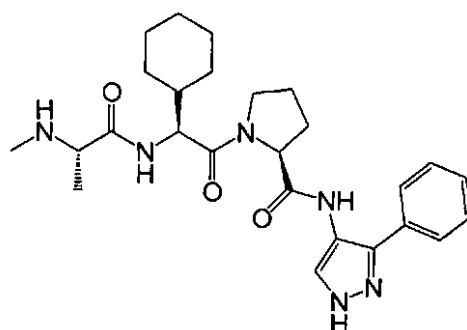
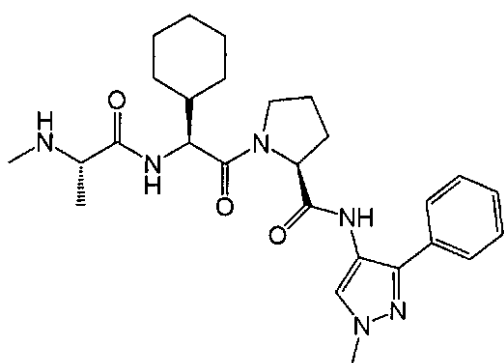




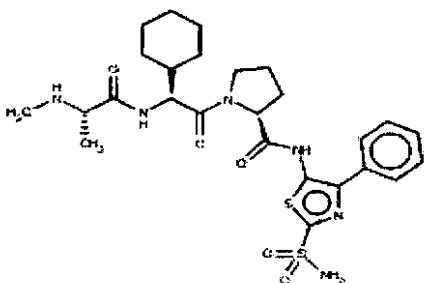
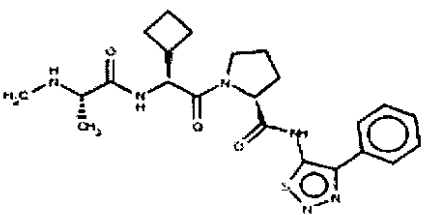
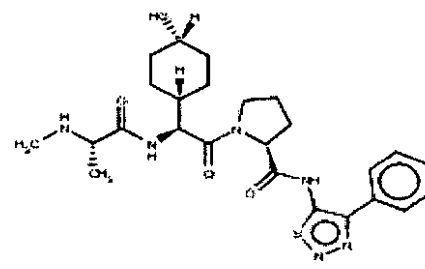
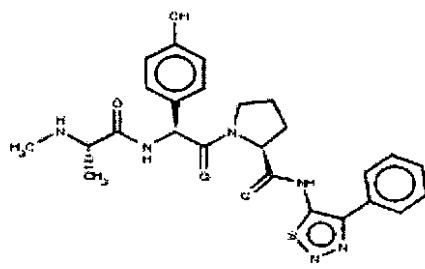
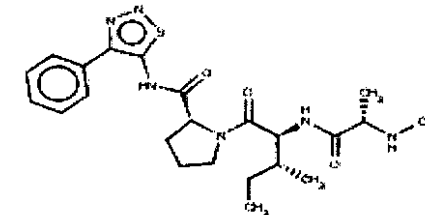
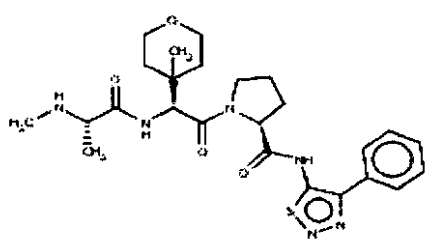


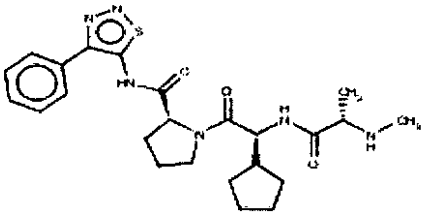
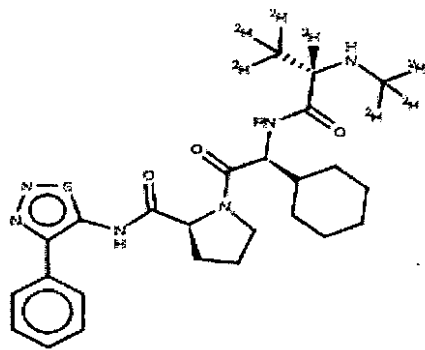
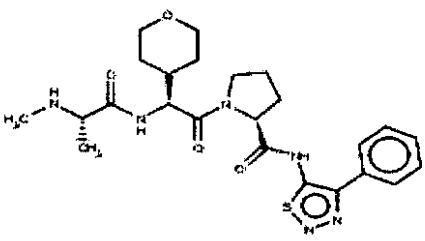
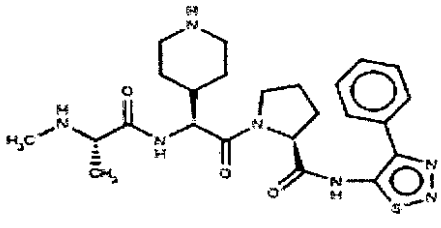
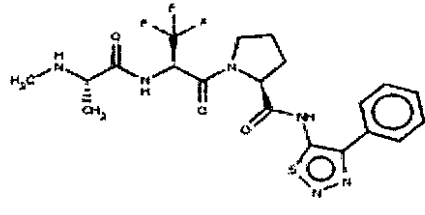
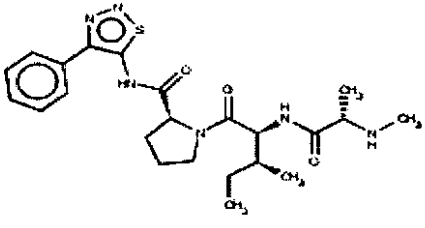


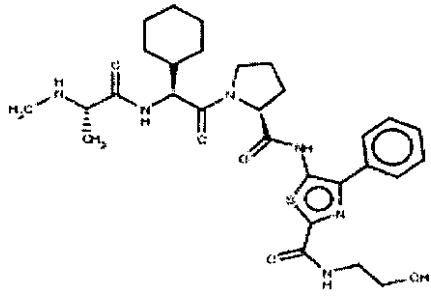
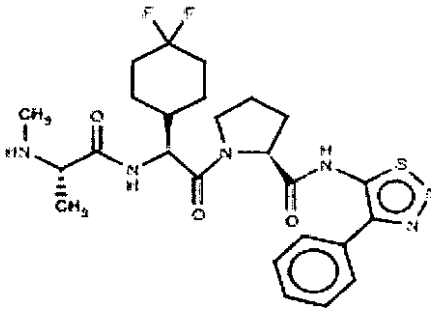
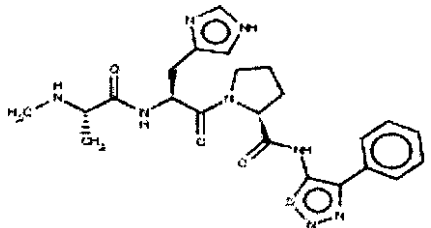
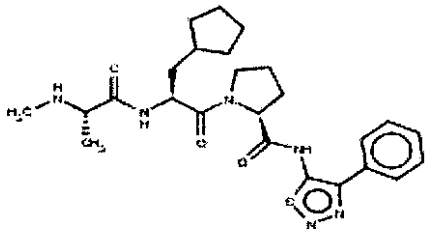
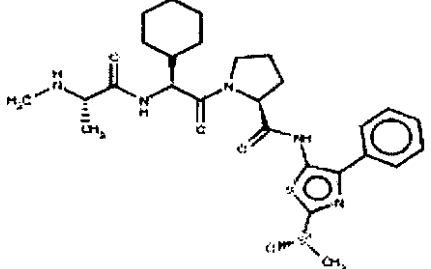


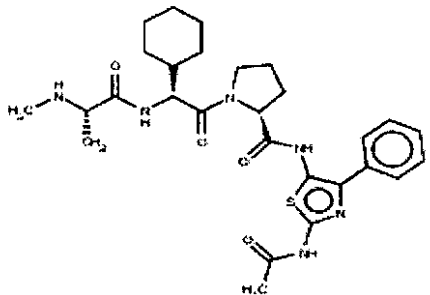
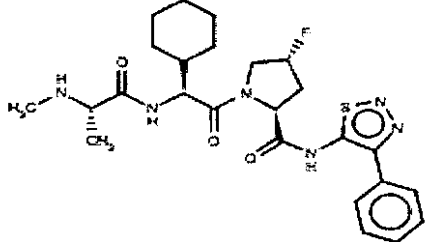
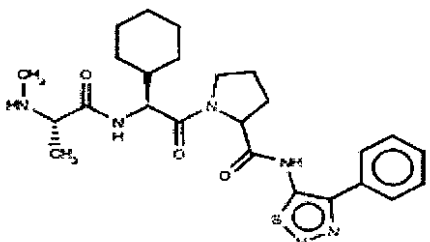
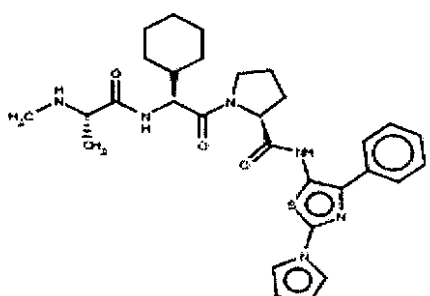
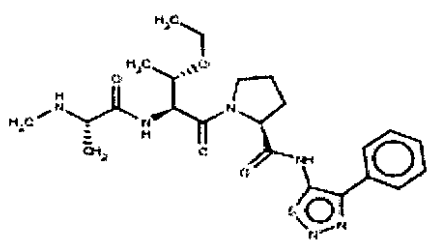


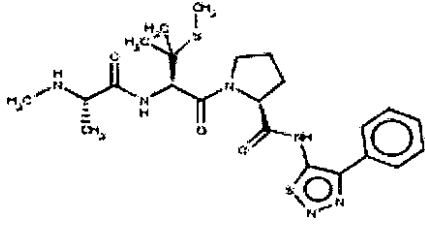
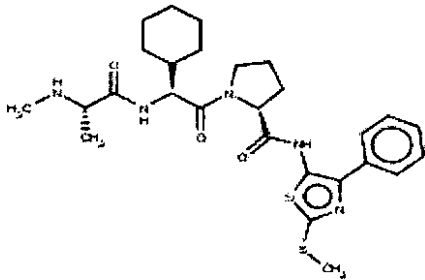
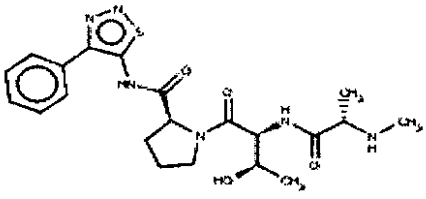
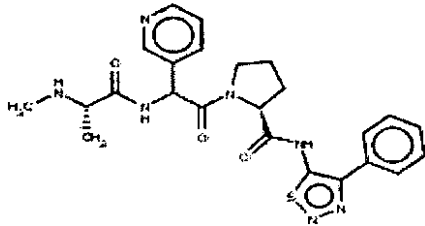
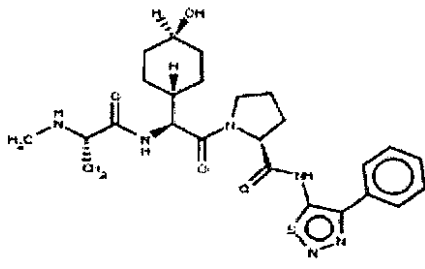
Afinidad por el dominio BIR3 de proteínas X-IAP

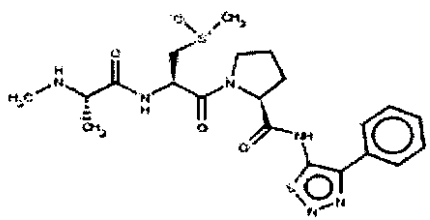
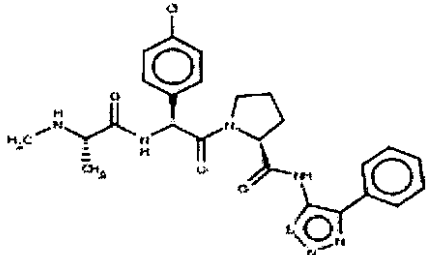
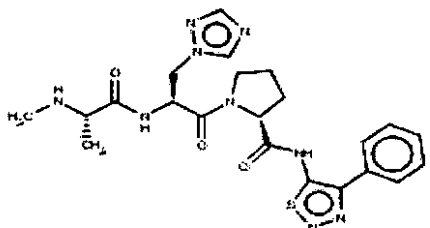
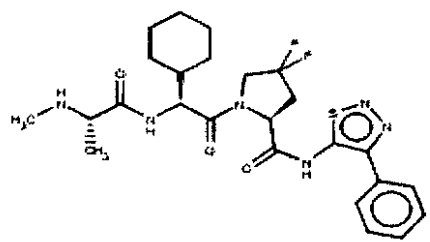
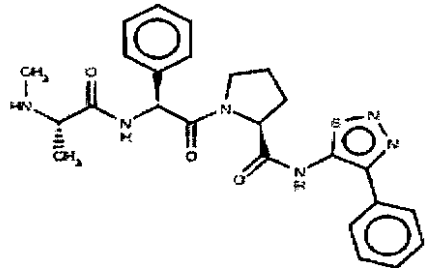
Item	Estructura del Compuesto	BIR3 FP (Ki) [uM]
1		0,0193
2		0,0202
3		0,0226
4		0,024
5		0,025
6		0,0254

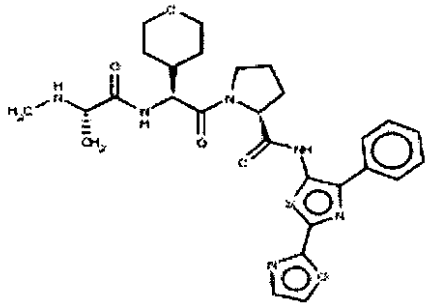
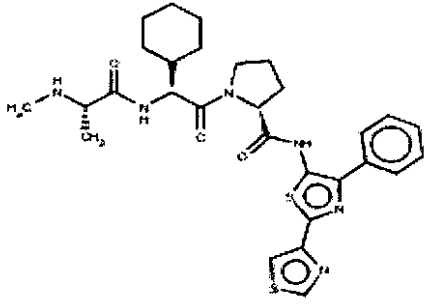
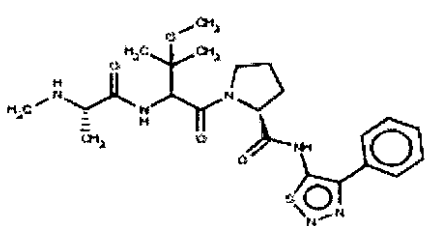
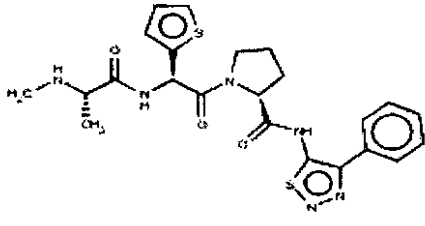
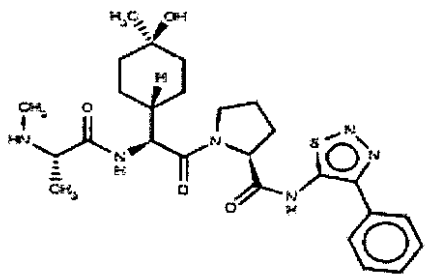
7		0,0255
8		0,0261
9		0,0269
10		0,0277
11		0,029
12		0,0292

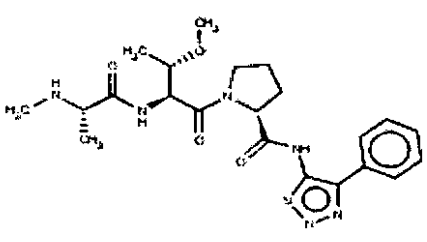
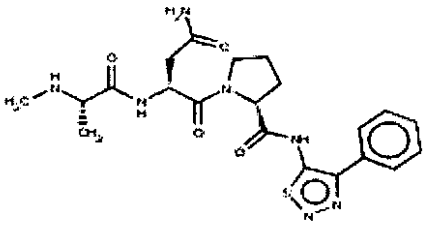
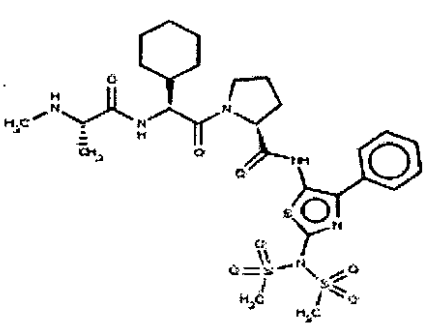
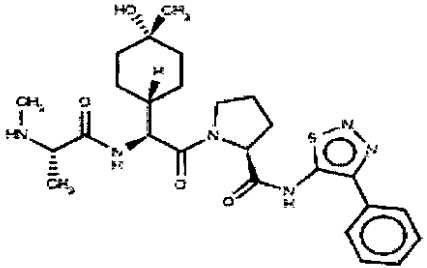
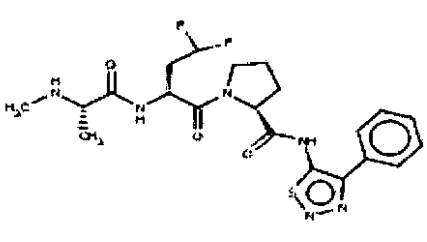
13	 <chem>NC(=O)[C@H](O)C(=O)N[C@@H](C1CCCCC1)C(=O)N2CCCC2C(=O)Nc3nc(C(=O)NCCO)c4ccccc34</chem>	0,0293
14	 <chem>CC[C@H](N)C(=O)N[C@@H](C1CCCC(F)(F)C1)C(=O)N2CCCC2C(=O)Nc3nc4ccccc4n3</chem>	0,0294
15	 <chem>NC(=O)[C@H](O)C(=O)N[C@@H](Cc1c[nH]cn1)C(=O)N2CCCC2C(=O)Nc3nc4ccccc34</chem>	0,0302
16	 <chem>NC(=O)[C@H](O)C(=O)N[C@@H](Cc1ccccc1)C(=O)N2CCCC2C(=O)Nc3nc4ccccc34</chem>	0,0305
17	 <chem>NC(=O)[C@H](O)C(=O)N[C@@H](C1CCCCC1)C(=O)N2CCCC2C(=O)Nc3nc(C(=O)N(C)C)c4ccccc34</chem>	0,0309

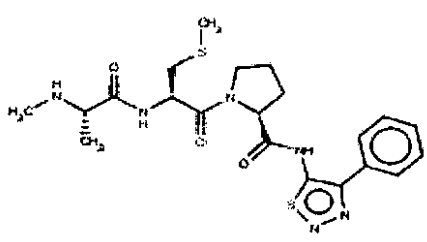
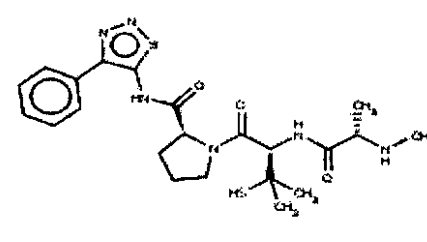
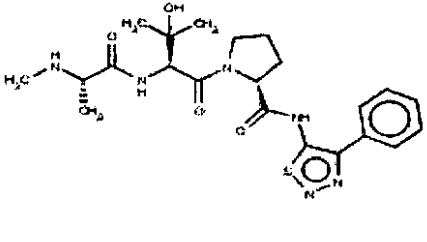
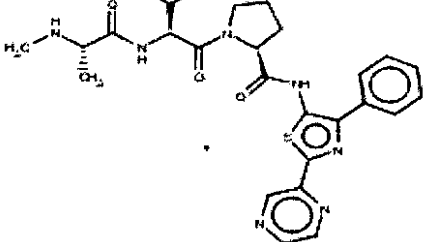
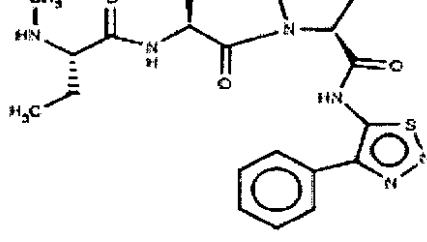
18		0,0312
19		0,0316
20		0,0318
21		0,0318
22		0,0321

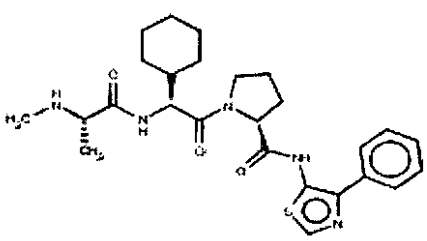
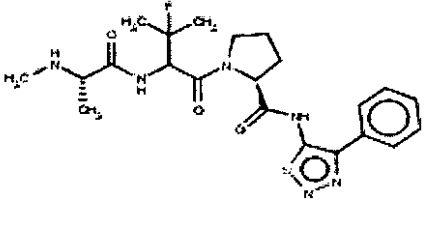
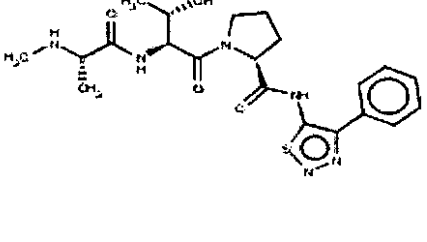
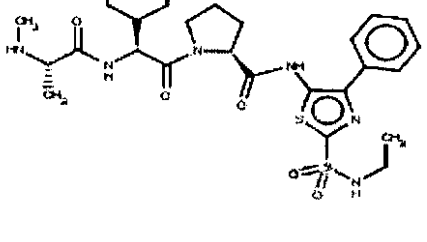
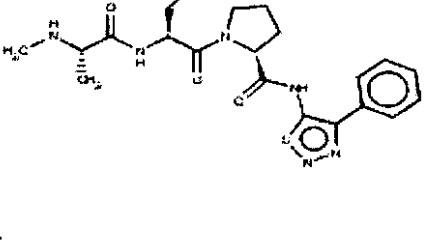
23		0,0323
24		0,0327
25		0,033
26		0,0332
27		0,0334

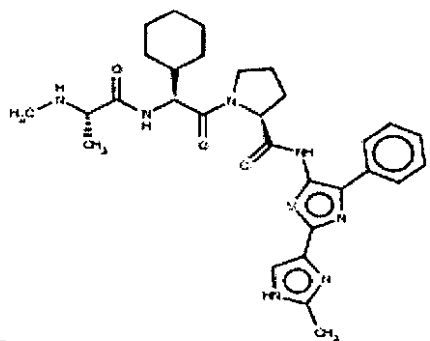
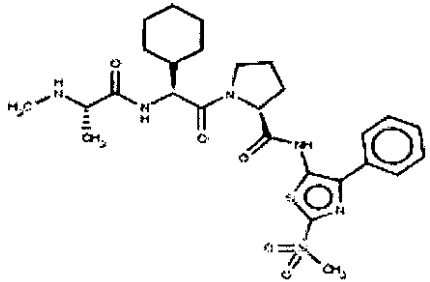
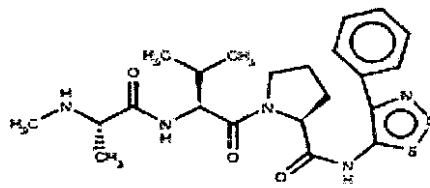
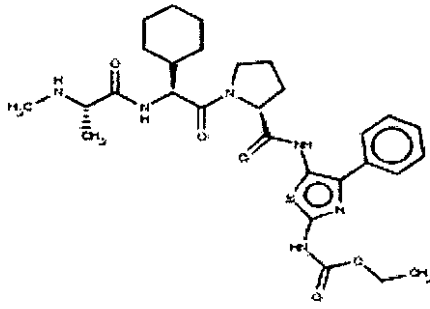
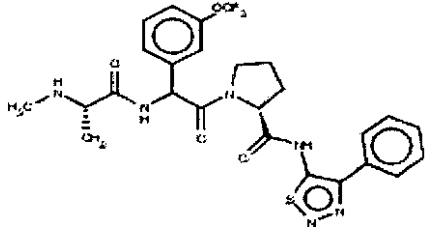
28		0,0336
29		0,0338
30		0,0344
31		0,0344
32		0,0353

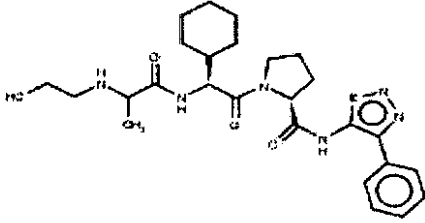
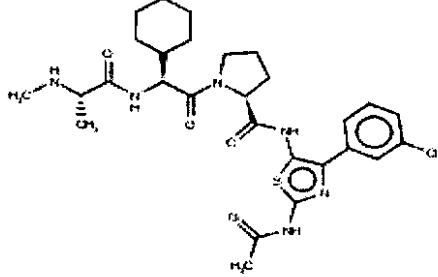
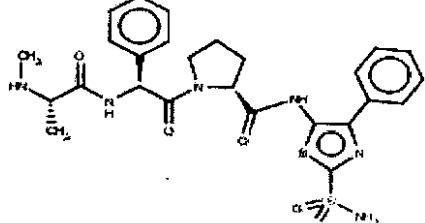
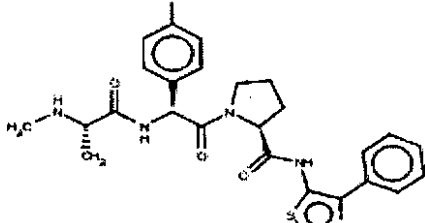
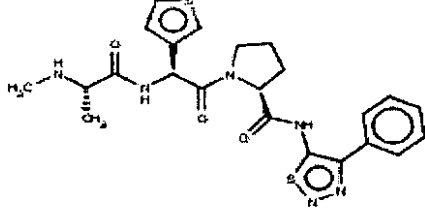
33		0,036
34		0,0361
35		0,0365
36		0,0373
37		0,0373

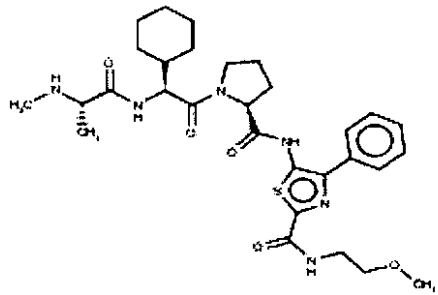
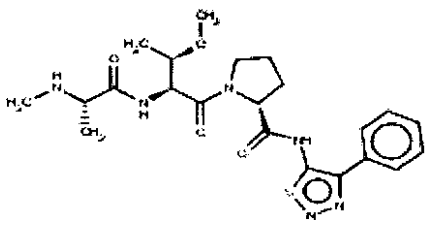
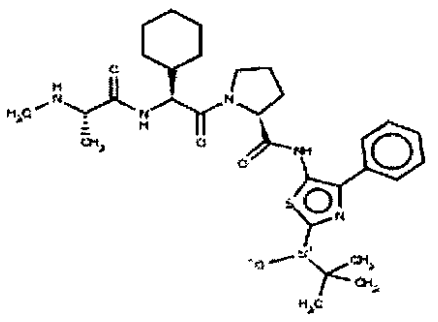
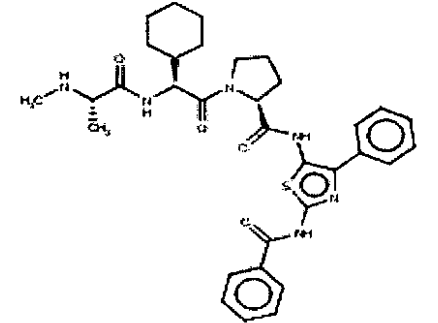
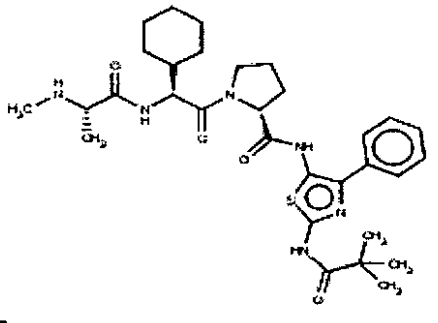
38		0,0373
39		0,0374
40		0,0375
41		0,0381
42		0,0381

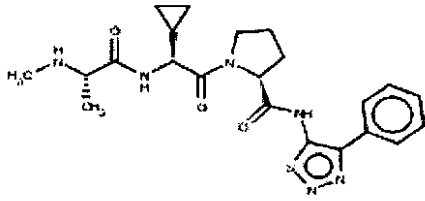
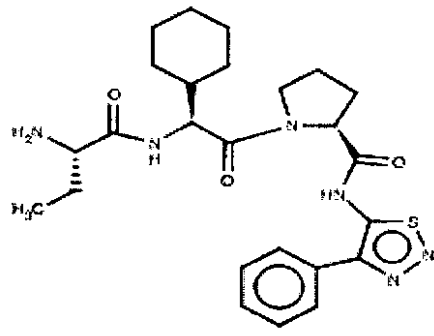
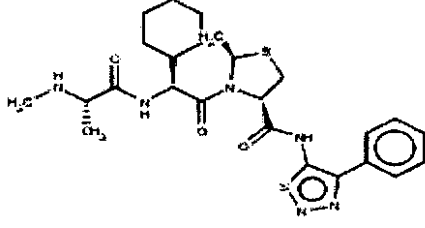
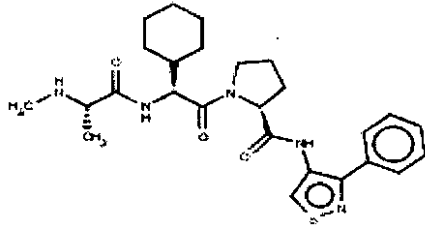
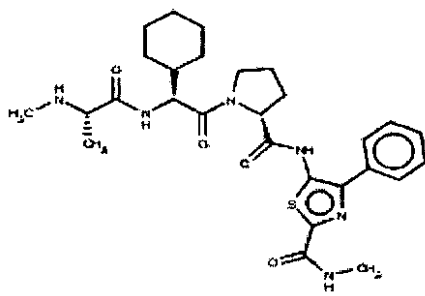
43		0,0387
44		0,039
45		0,0392
46		0,0397
47		0,0405

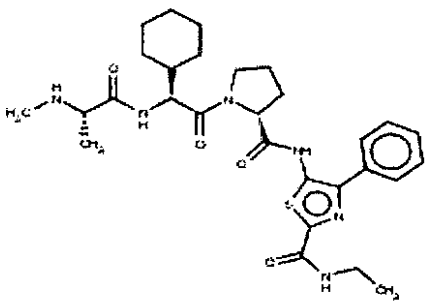
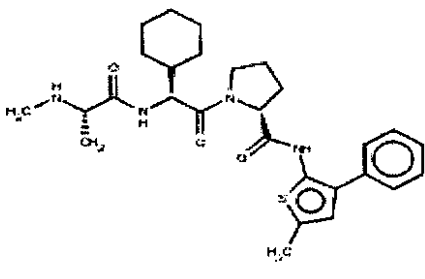
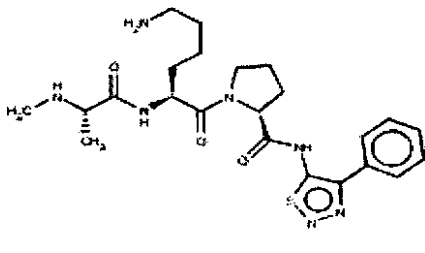
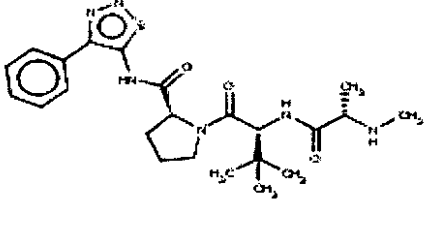
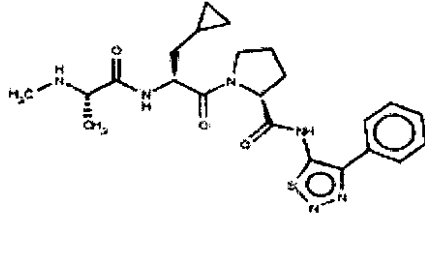
48		0,0405
49		0,0408
50		0,0411
51		0,0412
52		0,0413

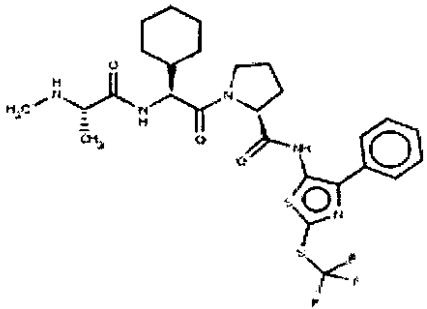
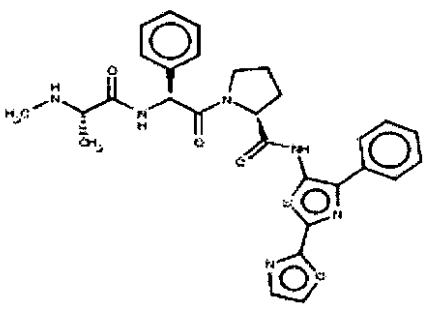
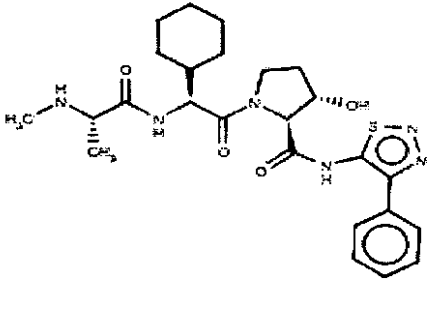
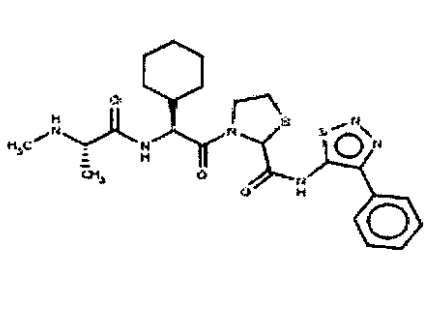
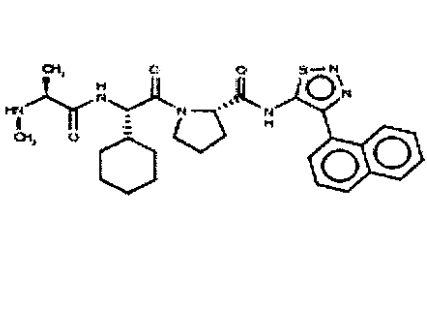
53		0,0416
54		0,0423
55		0,0425
56		0,0427
57		0,0429

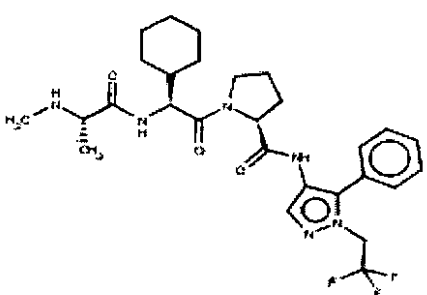
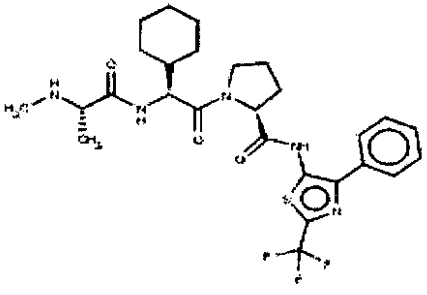
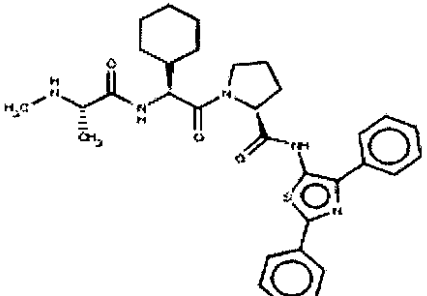
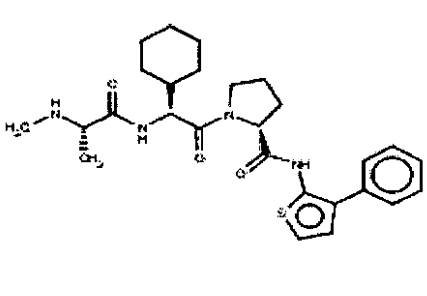
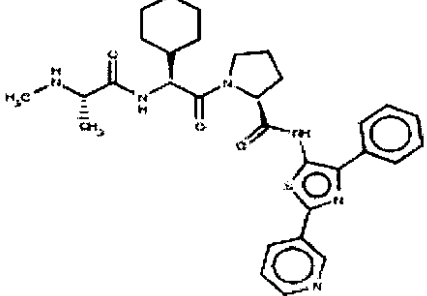
58		0,0431
59		0,0435
60		0,0446
61		0,0446
62		0,0447

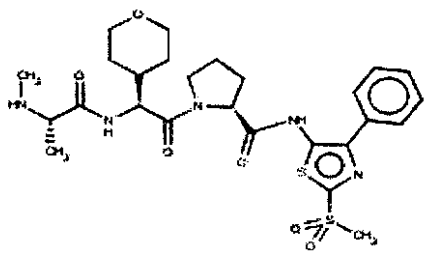
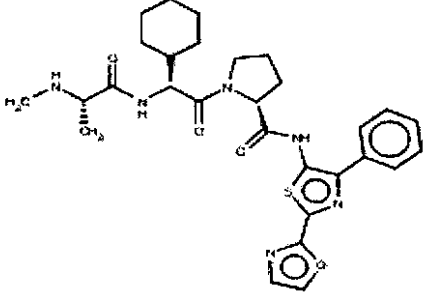
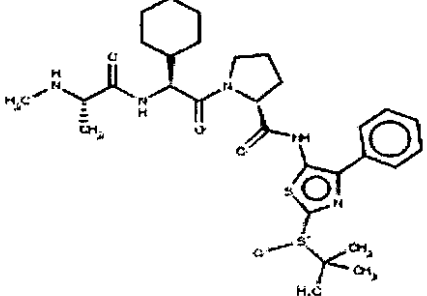
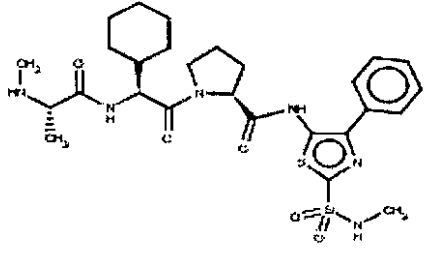
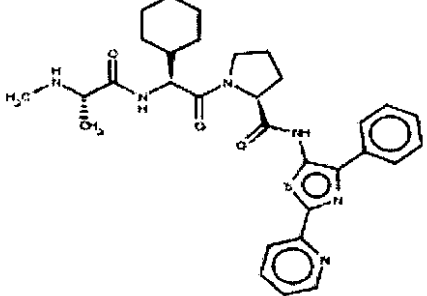
63		0,0453
64		0,0457
65		0,0458
66		0,0465
67		0,0474

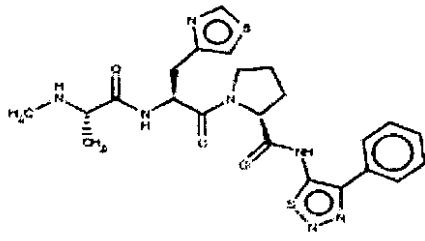
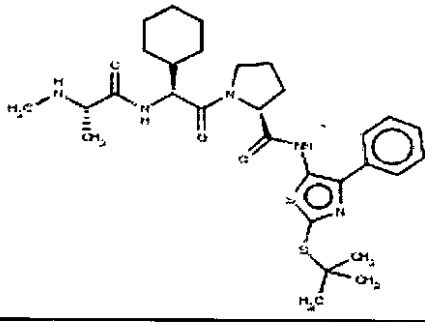
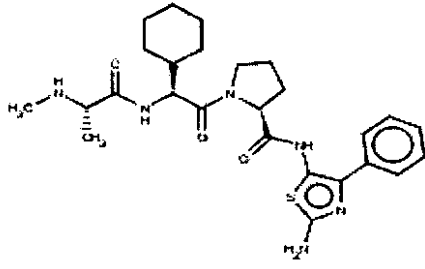
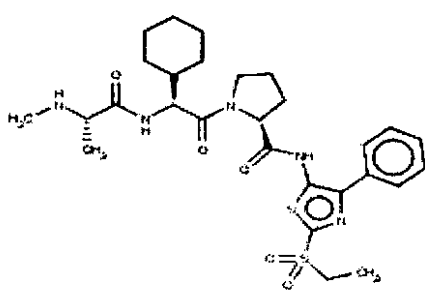
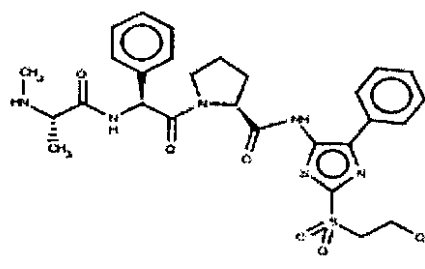
68		0,0474
69		0,048
70		0,0482
71		0,0484
72		0,0486

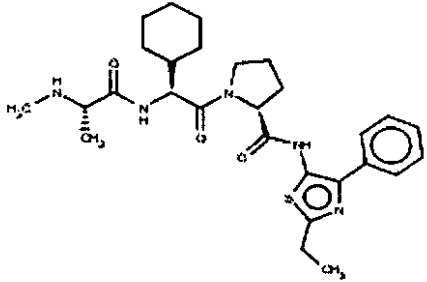
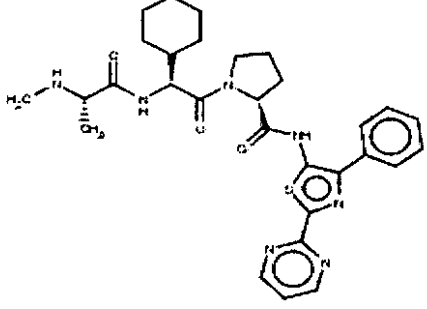
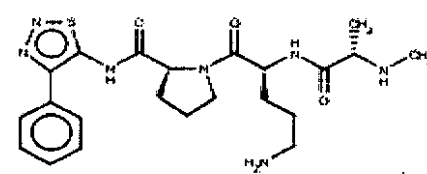
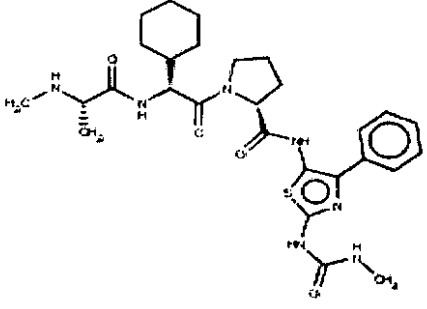
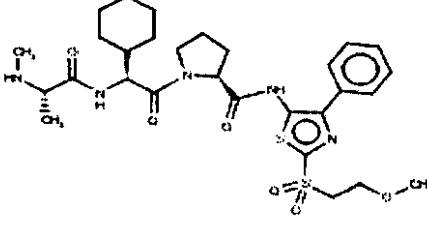
73		0,0488
74		0,0488
75		0,0489
76		0,0496
77		0,0498

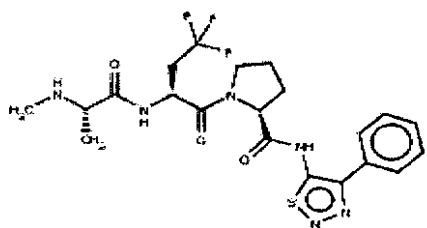
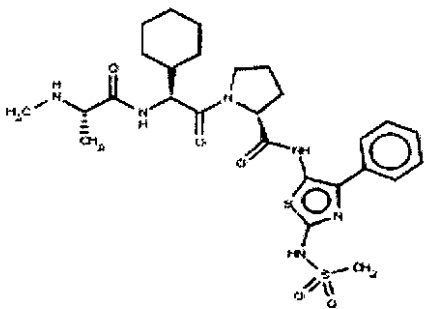
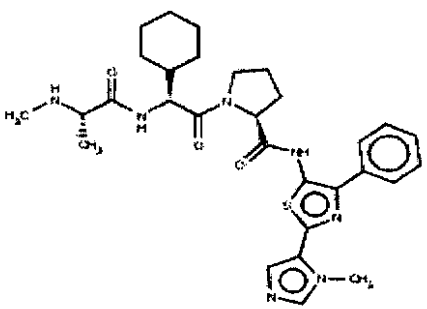
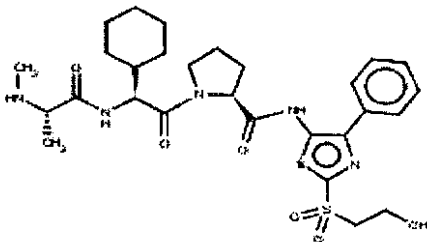
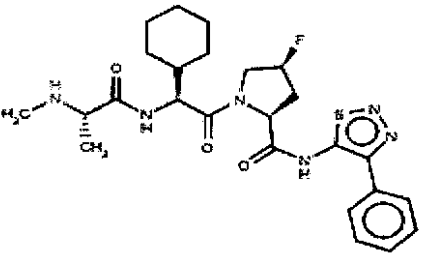
78		0,0499
79		0,0508
80		0,0513
81		0,0523
82		0,0524

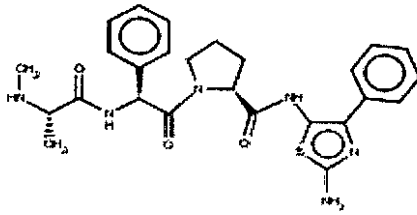
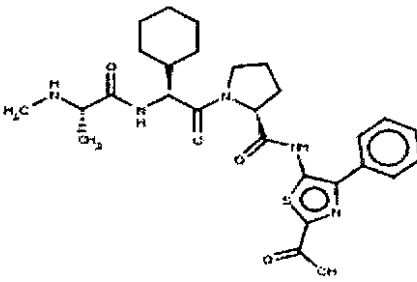
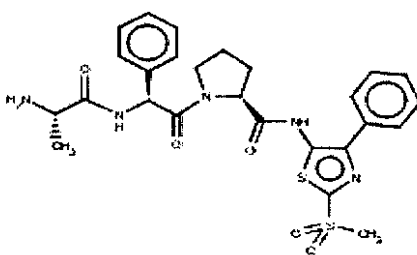
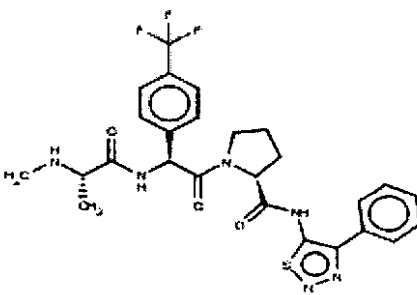
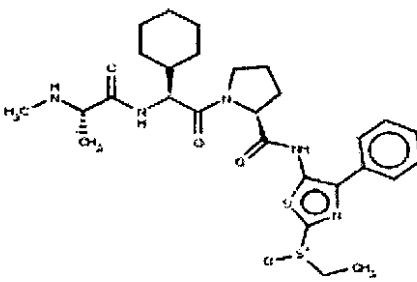
83		0,0531
84		0,0531
85		0,0532
86		0,0532
87		0,0534

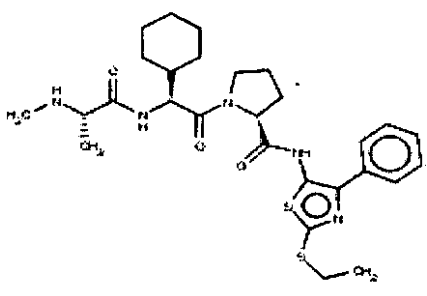
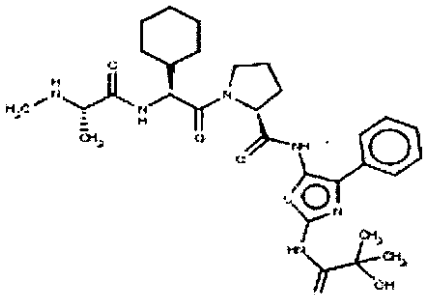
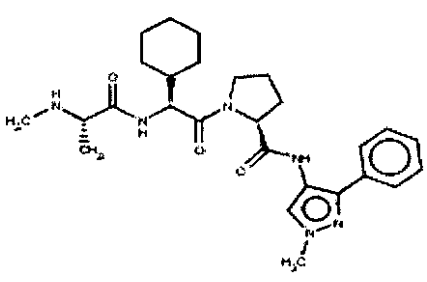
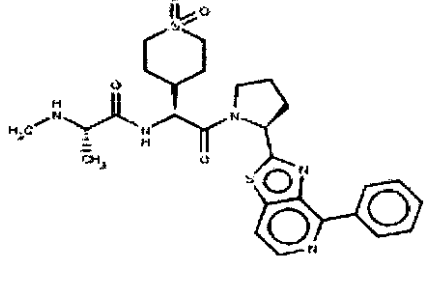
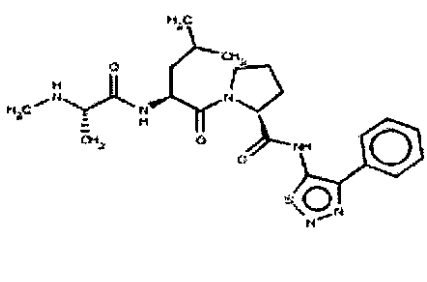
88		0,0541
89		0,0544
90		0,0551
91		0,0553
92		0,0556

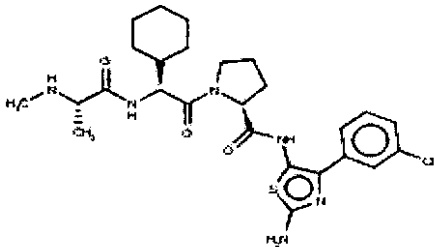
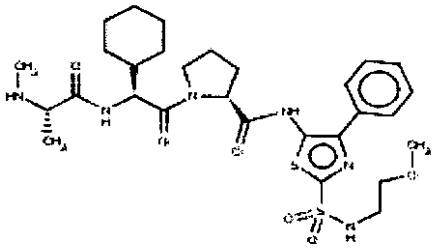
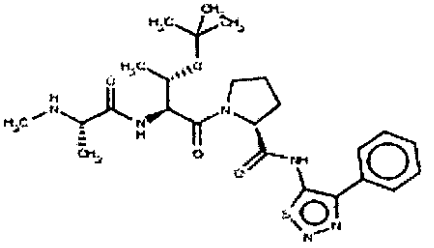
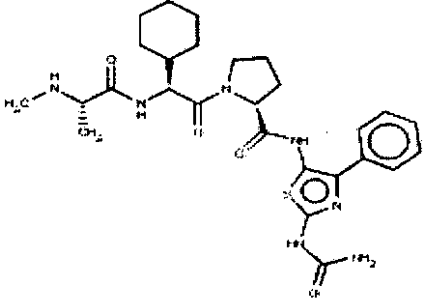
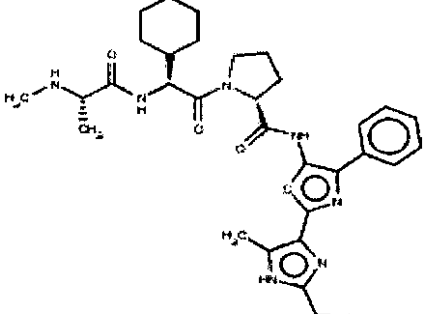
93		0,0568
94		0,0568
95		0,0577
96		0,0578
97		0,0581

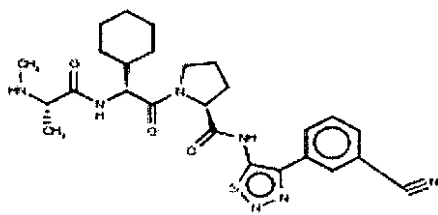
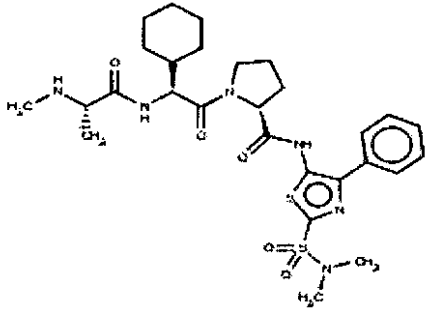
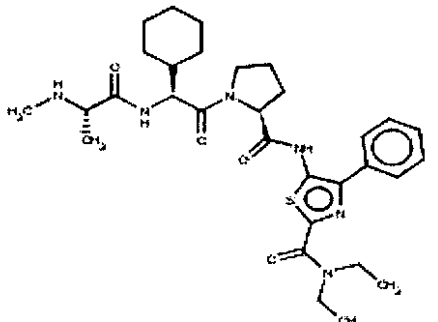
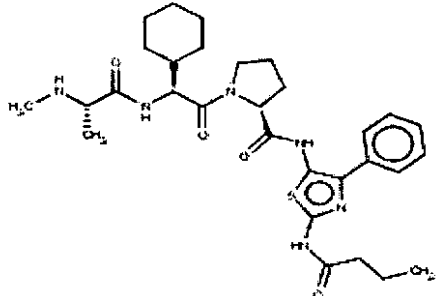
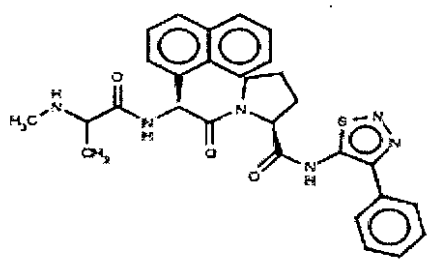
98		0,0581
99		0,0584
100		0,0588
101		0,0599
102		0,0604

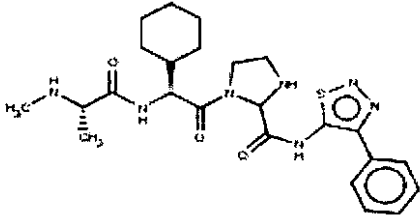
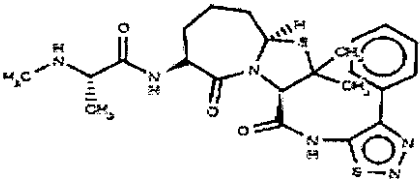
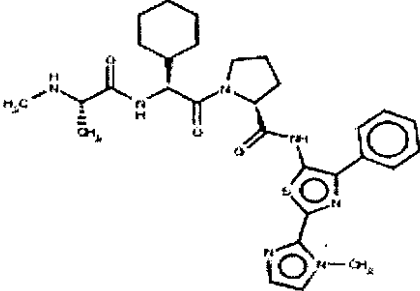
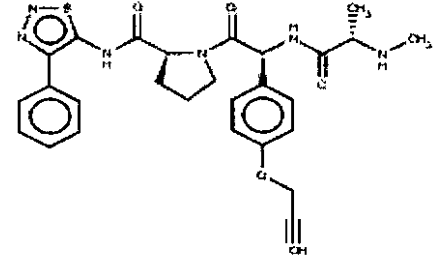
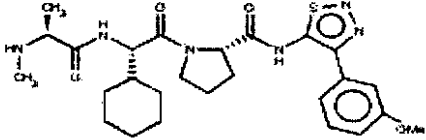
103		0,061
104		0,0611
105		0,0617
106		0,063
107		0,0637

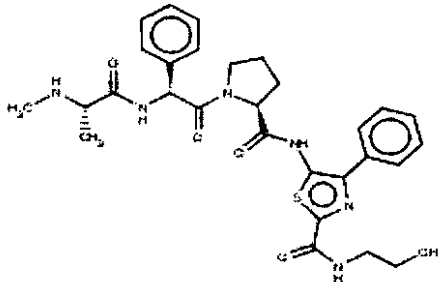
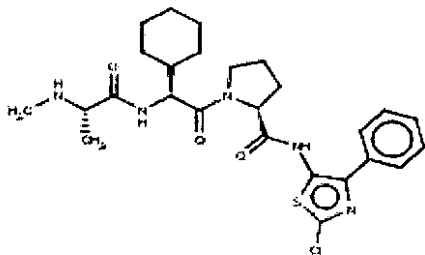
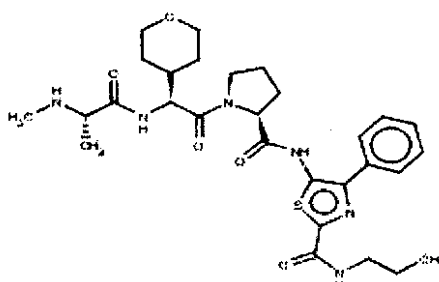
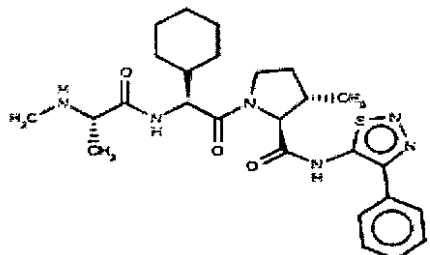
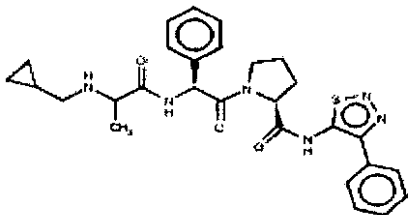
108		0,0641
109		0,0646
110		0,0649
111		0,0658
112		0,066

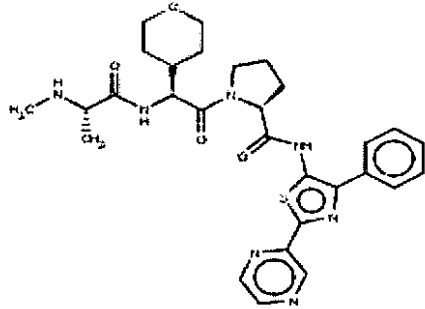
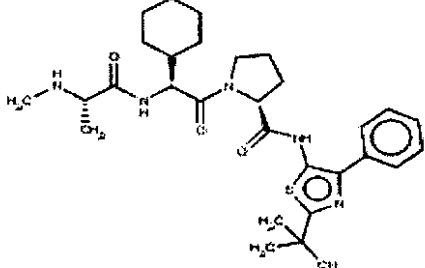
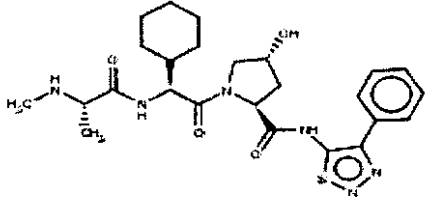
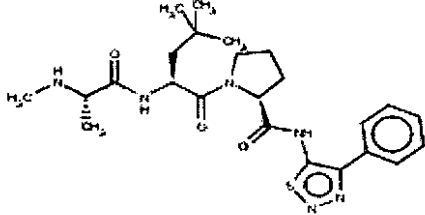
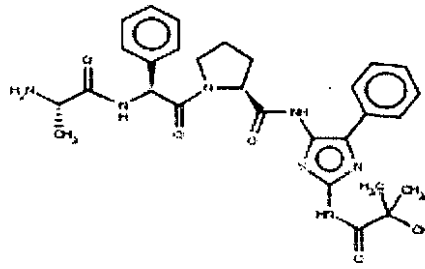
113		0,0661
114		0,0661
115		0,0667
116		0,0675
117		0,068

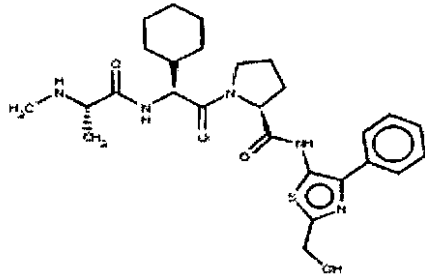
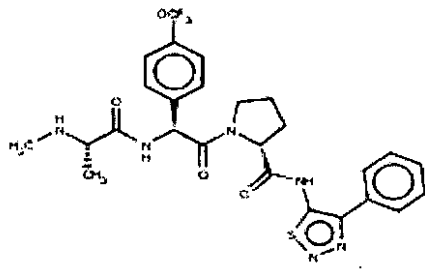
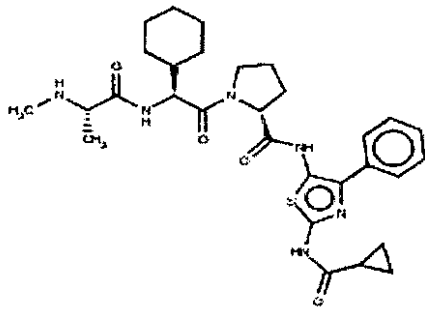
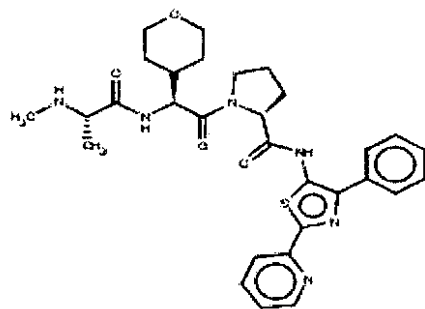
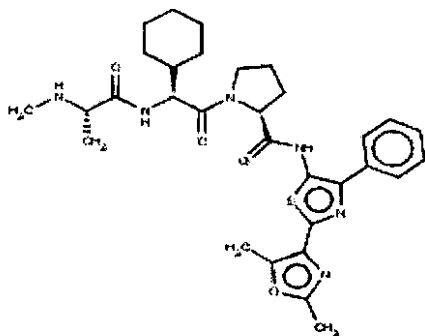
118		0,0683
119		0,069
120		0,0691
121		0,0692
122		0,0703

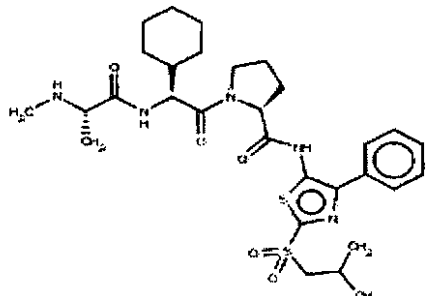
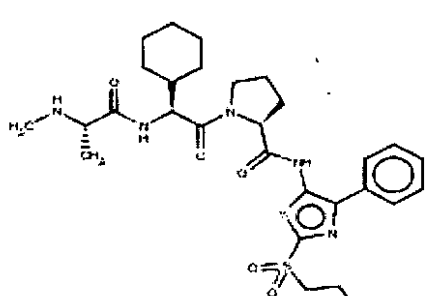
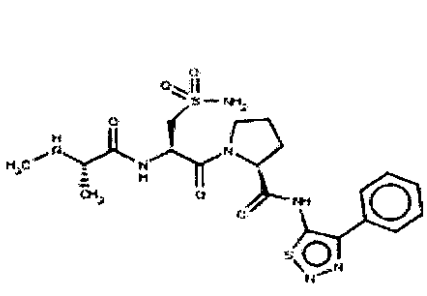
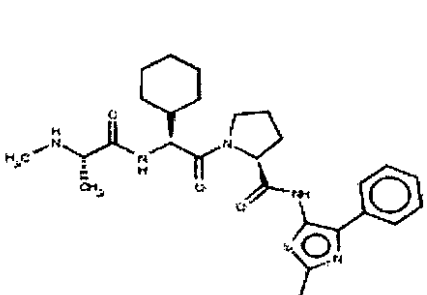
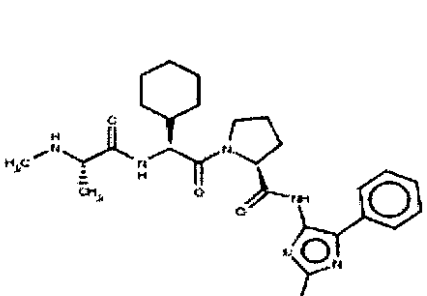
123		0,0705
124		0,0712
125		0,0714
126		0,0715
127		0,0716

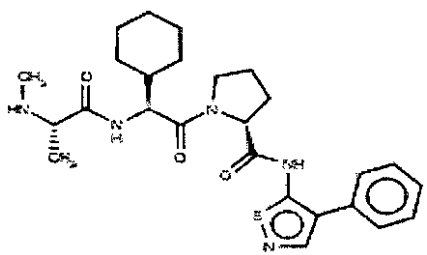
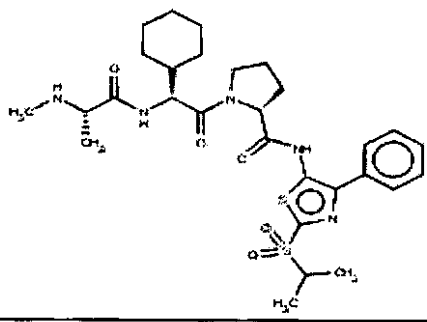
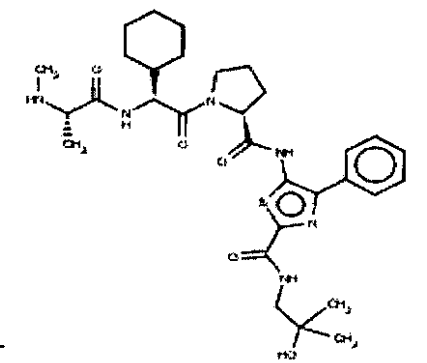
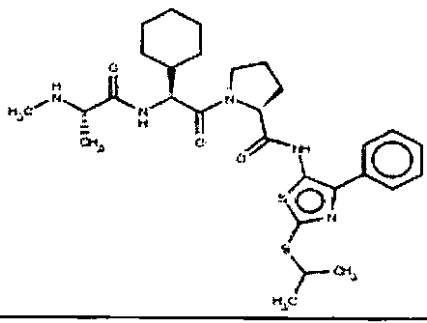
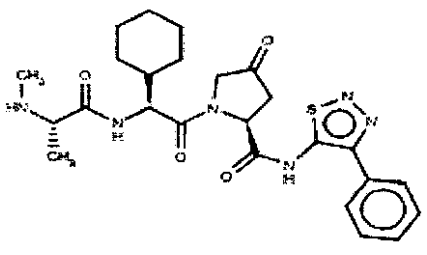
128		0,0722
129		0,0734
130		0,0742
131		0,0746
132		0,0751

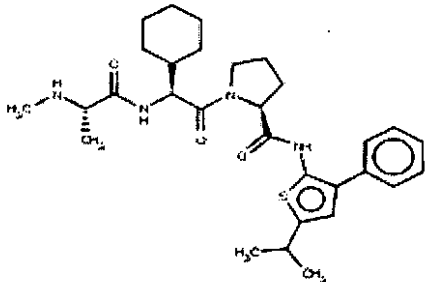
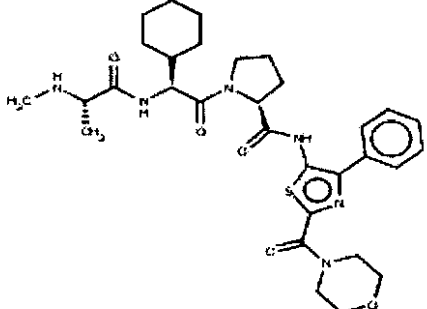
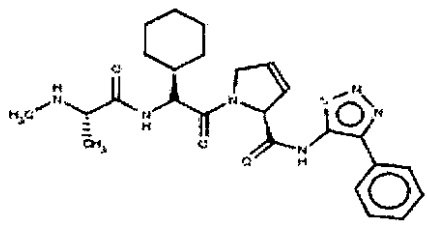
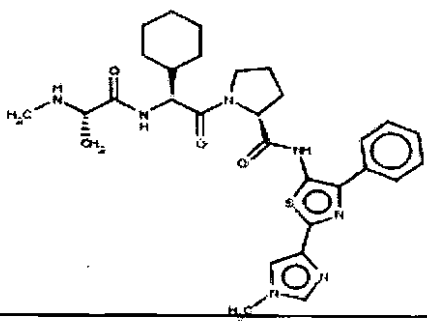
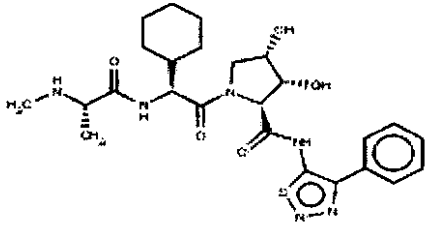
133		0,0753
134		0,076
135		0,0761
136		0,0769
137		0,0775

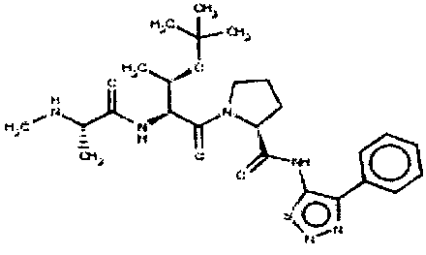
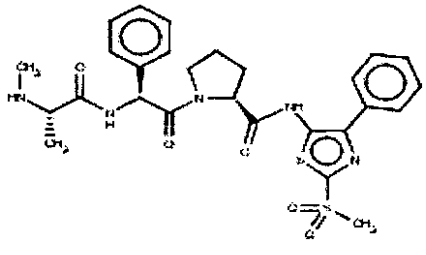
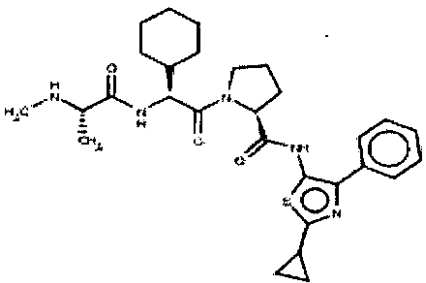
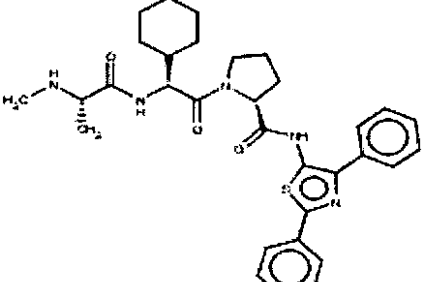
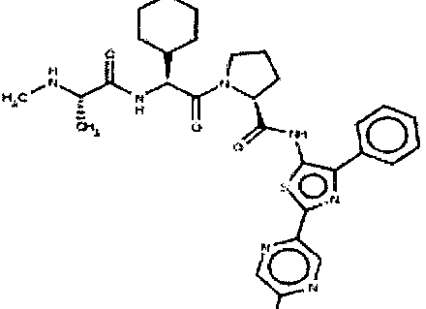
138		0,0777
139		0,0783
140		0,0785
141		0,0789
142		0,0794

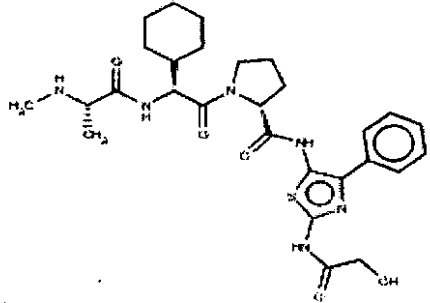
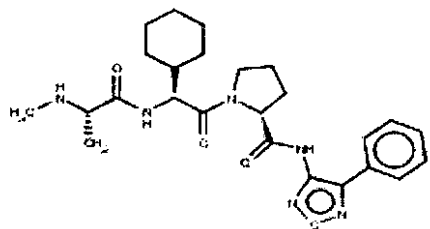
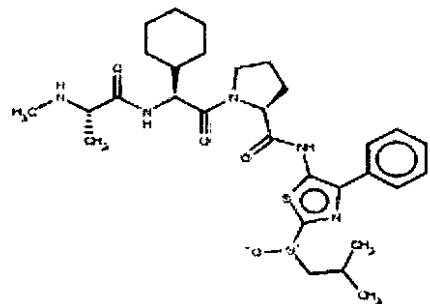
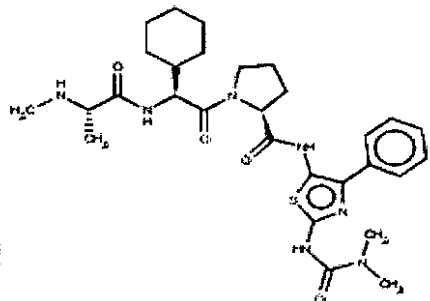
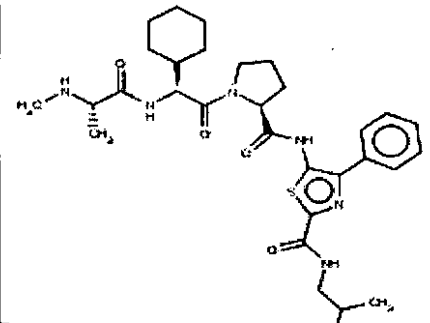
143		0,0802
144		0,0808
145		0,0808
146		0,0814
147		0,0814

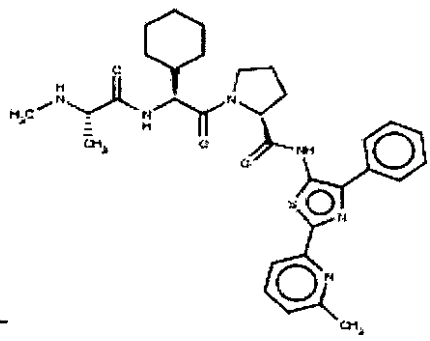
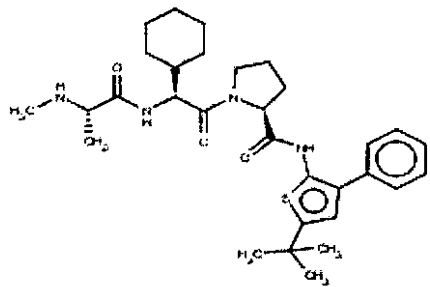
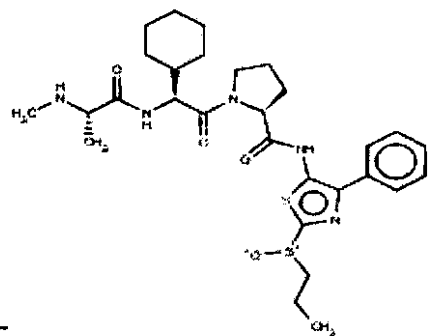
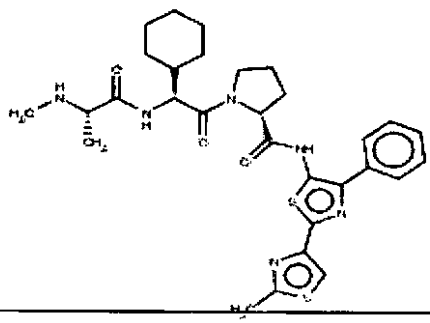
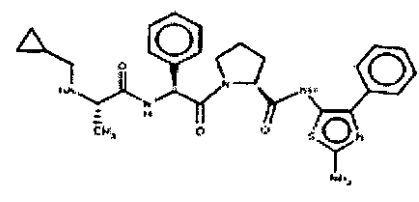
148	 <chem>CCNC(=O)C(C1CCCCC1)NC(=O)C2CCCC2C(=O)Nc3nc(cc3c4ccccc4)S(=O)(=O)CC5C</chem>	0,0825
149	 <chem>CCNC(=O)C(C1CCCCC1)NC(=O)C2CCCC2C(=O)Nc3nc(cc3c4ccccc4)S(=O)(=O)CC5CC</chem>	0,0832
150	 <chem>CCNC(=O)C(C1CCCCC1)NC(=O)C2CCCC2C(=O)Nc3nc(cc3c4ccccc4)S(=O)(=O)N</chem>	0,0834
151	 <chem>CCNC(=O)C(C1CCCCC1)NC(=O)C2CCCC2C(=O)Nc3nc(cc3c4ccccc4)c5cc[nH]5</chem>	0,0838
152	 <chem>CCNC(=O)C(C1CCCCC1)NC(=O)C2CCCC2C(=O)Nc3nc(cc3c4ccccc4)c5cc6nn[nH]6c5</chem>	0,0839

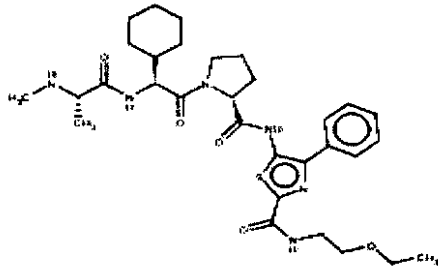
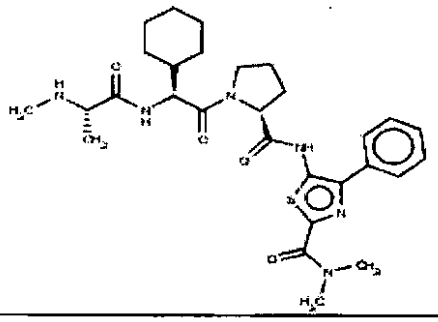
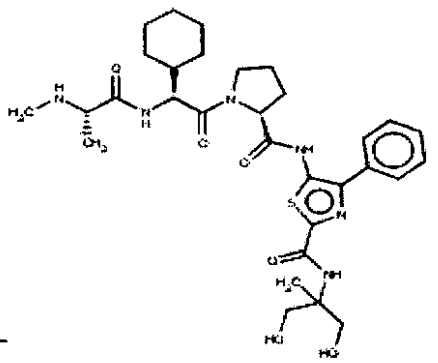
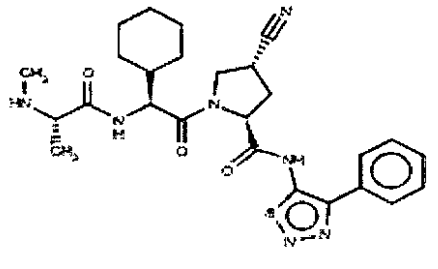
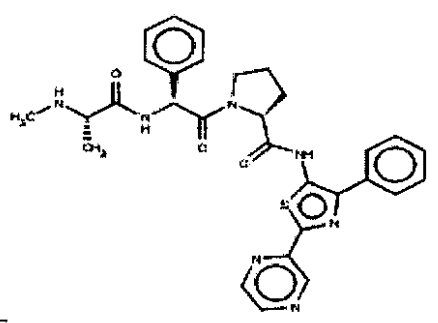
158		0,0852
159		0,0857
160		0,0863
161		0,0868
162		0,087

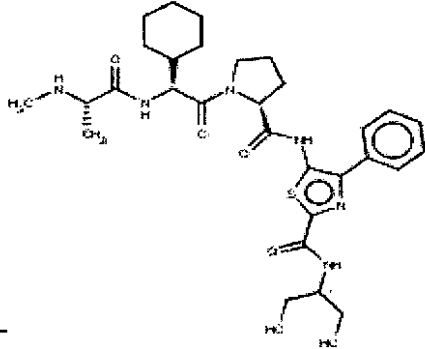
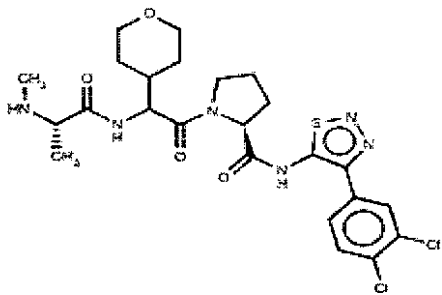
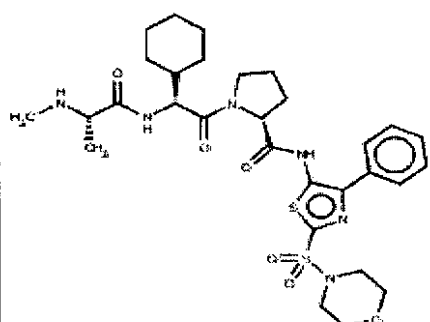
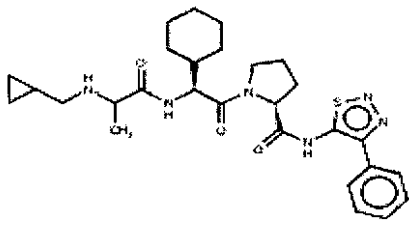
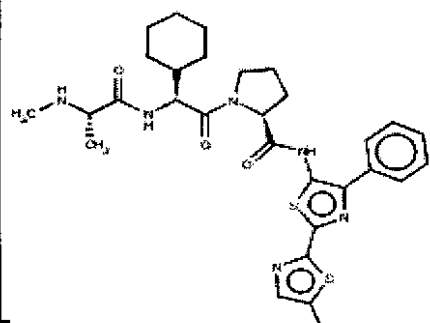
163		0,0879
164		0,0881
165		0,0881
166		0,0888
167		0,0888

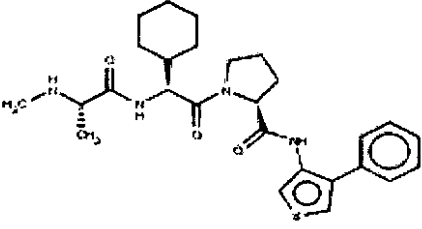
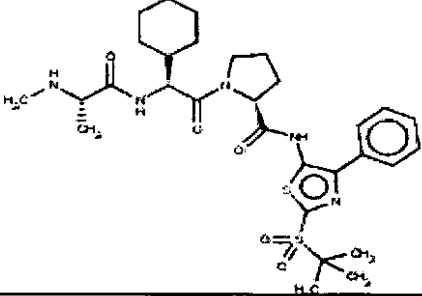
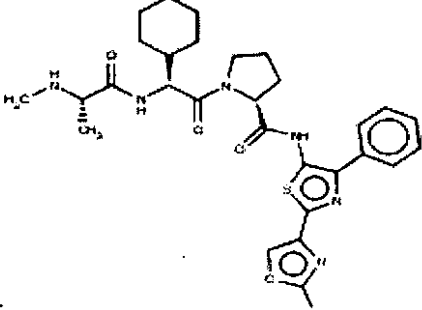
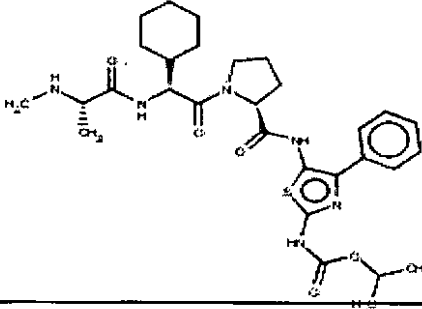
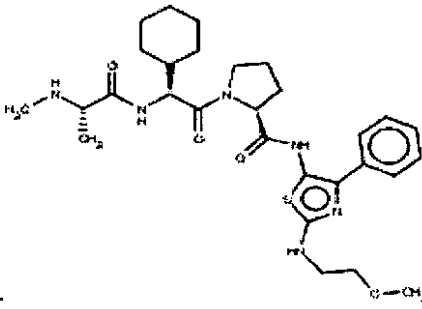
168		0,0889
169		0,0891
170		0,0892
171		0,0898
172		0,0899

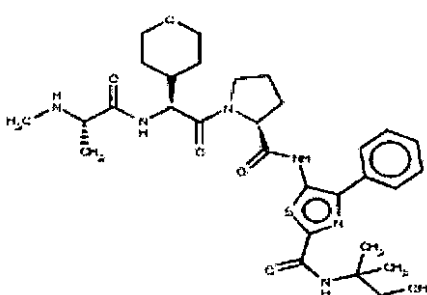
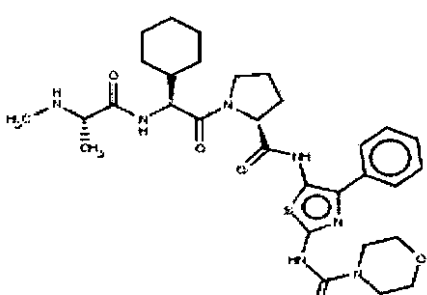
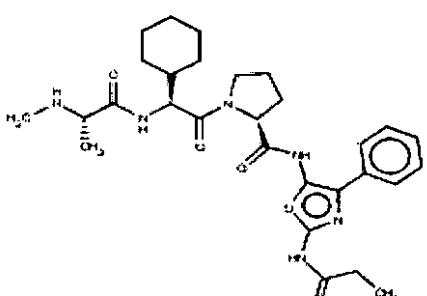
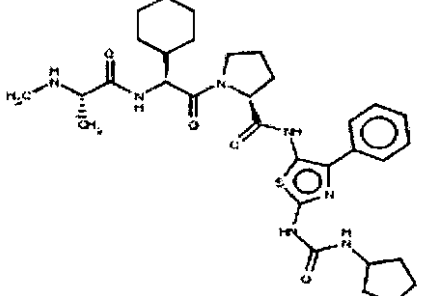
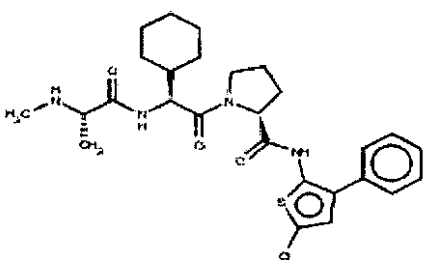
173		0,090
174		0,0904
175		0,0907
176		0,0922
177		0,0936

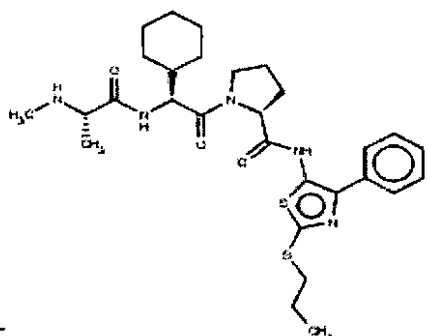
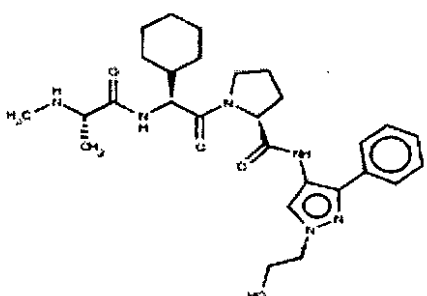
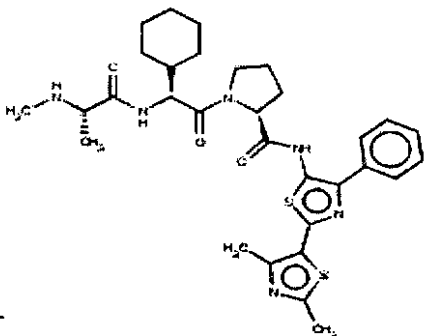
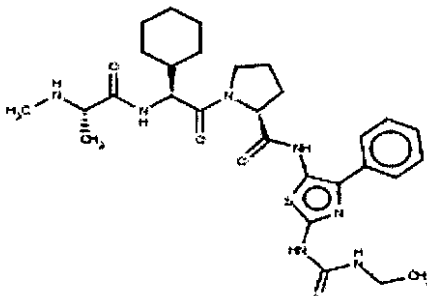
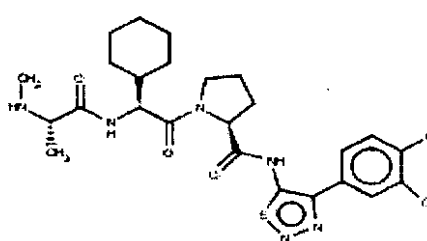
178	 <chem>CC1=CC=C(C=C1)c2nc(C3=CC=CC=C3)c(NC(=O)N4CCCC4C(=O)N[C@@H](C5=CCCCC5)C(=O)N[C@@H](C)C(=O)N)cs2</chem>	0,0939
179	 <chem>CC(C)(C)c1cc(C2=CC=CC=C2)c(NC(=O)N3CCCC3C(=O)N[C@@H](C4=CCCCC4)C(=O)N[C@@H](C)C(=O)N)cs1</chem>	0,0941
180	 <chem>CCCCS1=CC=C(C=C1)c2nc(C3=CC=CC=C3)c(NC(=O)N4CCCC4C(=O)N[C@@H](C5=CCCCC5)C(=O)N[C@@H](C)C(=O)N)cs2</chem>	0,0945
181	 <chem>Nc1ccc(cc1)-c2nc(C3=CC=CC=C3)c(NC(=O)N4CCCC4C(=O)N[C@@H](C5=CCCCC5)C(=O)N[C@@H](C)C(=O)N)cs2</chem>	0,0953
182	 <chem>C1CC1CC2=CC=C(C=C2)c3nc(C4=CC=CC=C4)c(NC(=O)N5CCCC5C(=O)N[C@@H](C6=CCCCC6)C(=O)N[C@@H](C)C(=O)N)cs3</chem>	0,0957

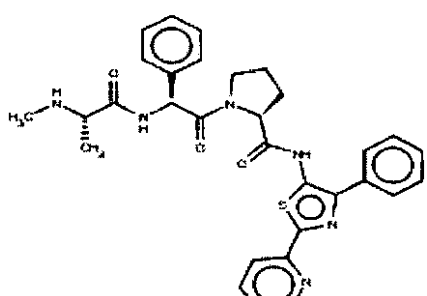
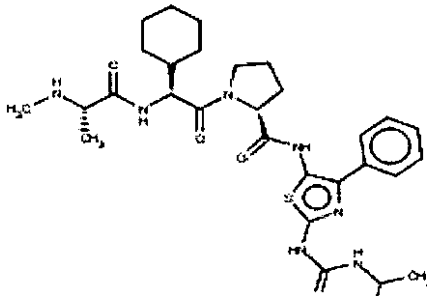
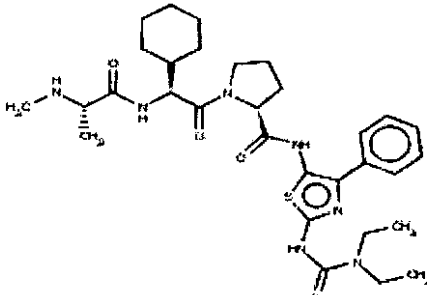
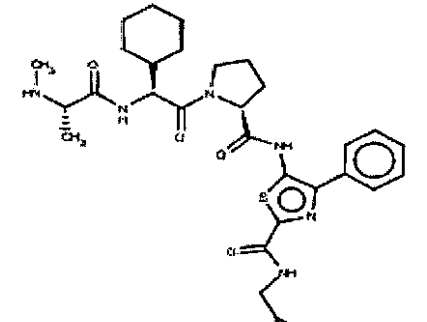
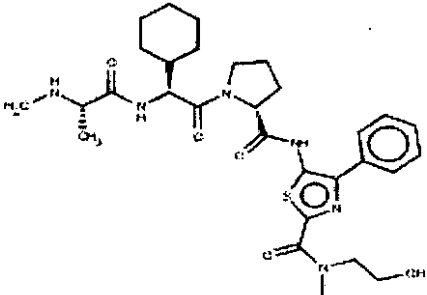
183		0,0958
184		0,096
185		0,0962
186		0,0962
187		0,0965

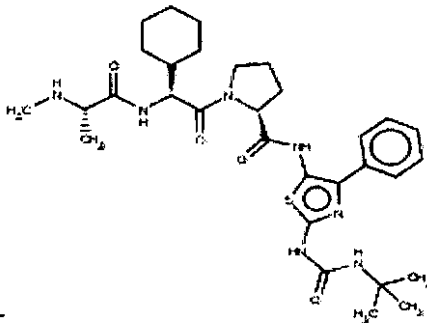
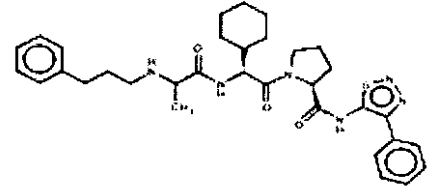
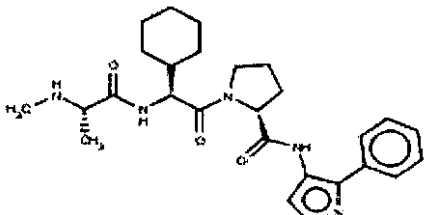
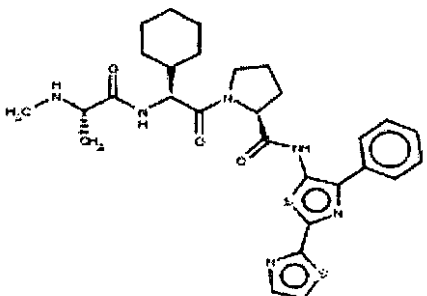
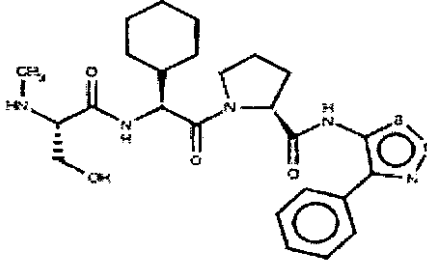
188		0,0966
189		0,0974
190		0,0976
191		0,0978
192		0,0981

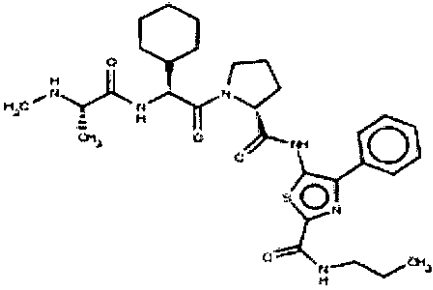
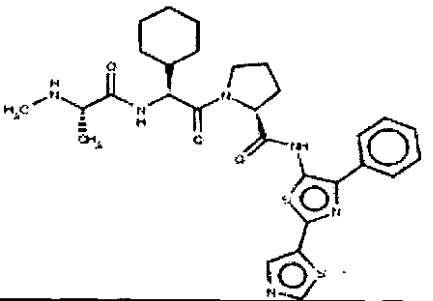
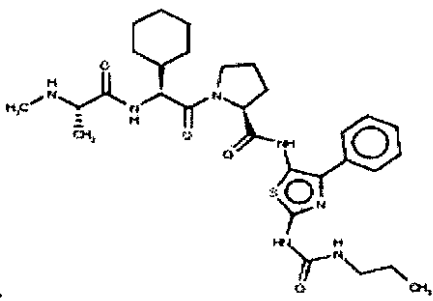
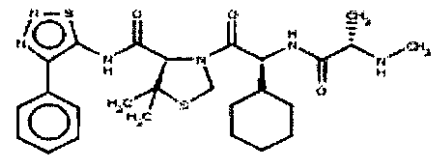
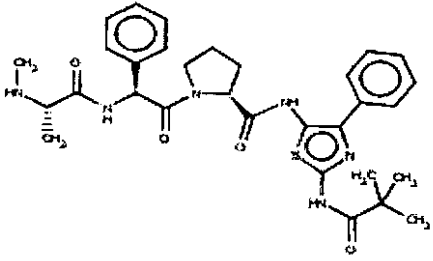
193		0,0981
194		0,0984
195		0,0984
196		0,0993
197		0,10

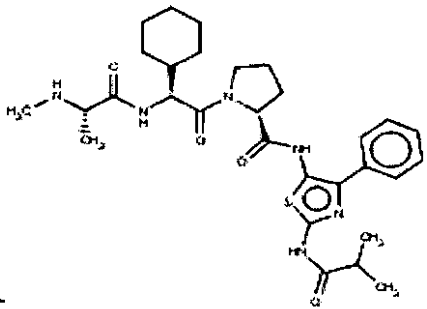
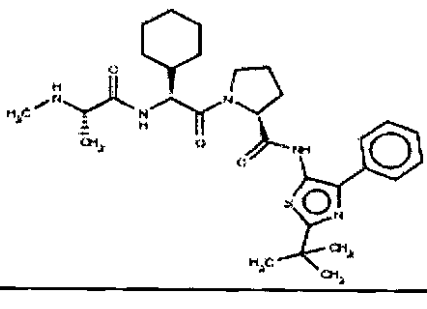
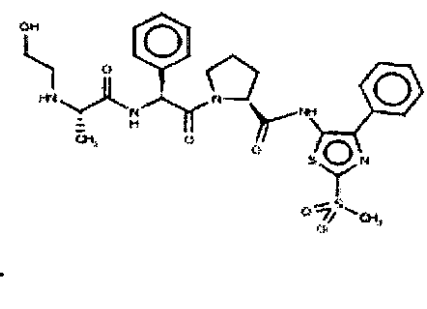
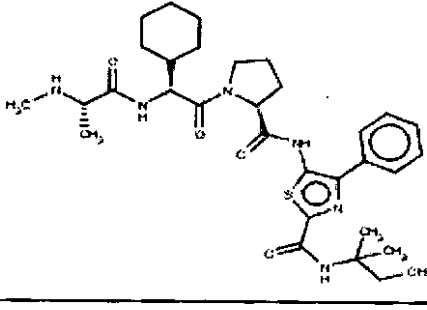
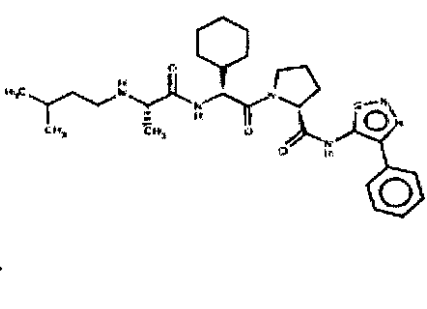
198		0,10
199		0,10
200		0,101
201		0,101
202		0,102

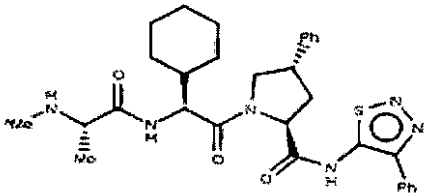
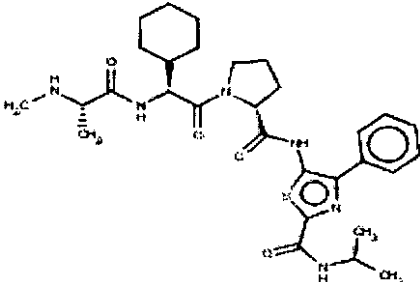
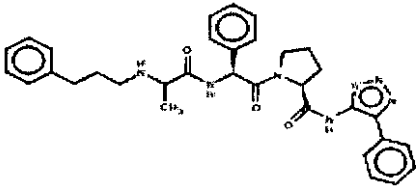
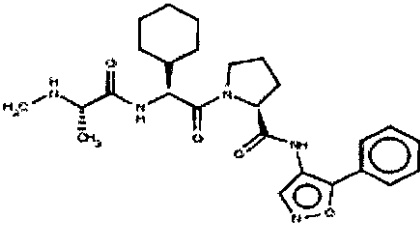
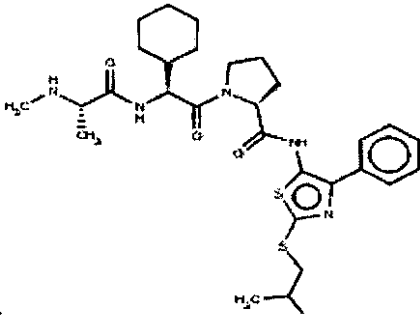
203		0,103
204		0,104
205		0,104
206		0,104
207		0,106

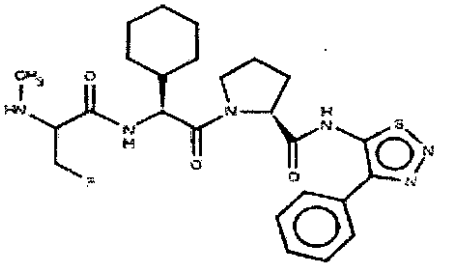
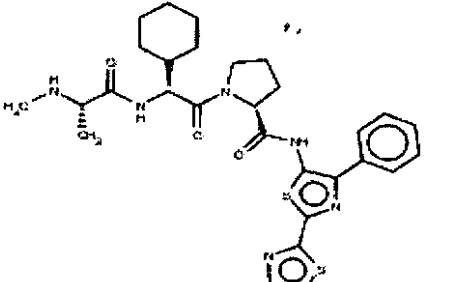
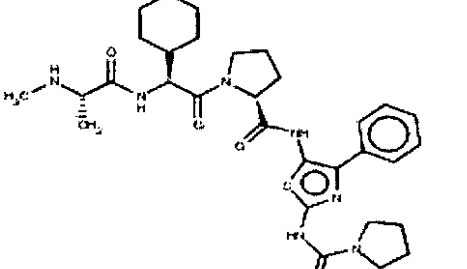
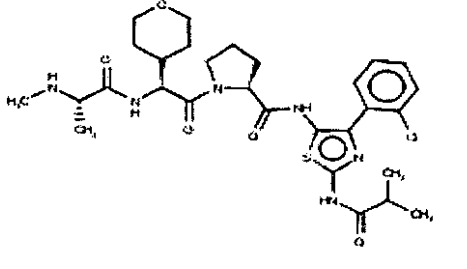
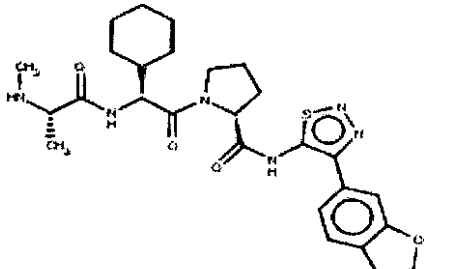
208		0,106
209		0,11
210		0,11
211		0,111
212		0,111

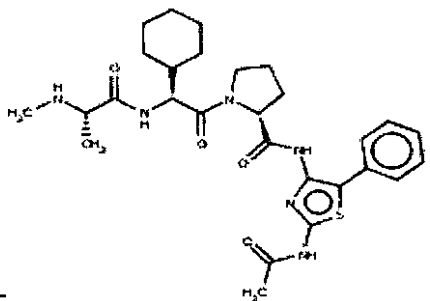
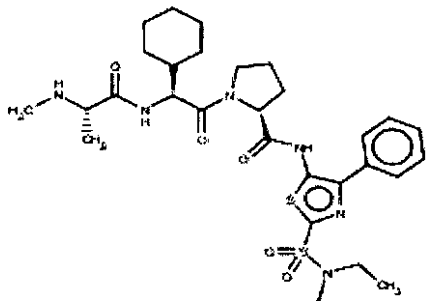
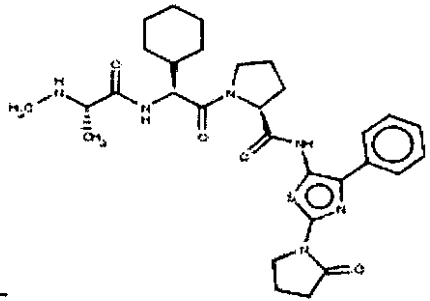
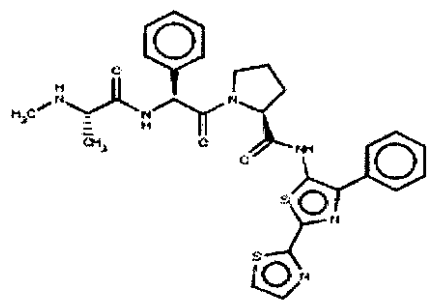
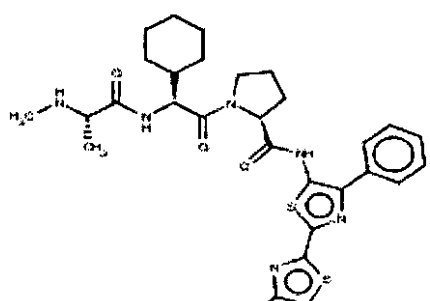
213		0,112
214		0,112
215		0,112
216		0,113
217		0,114

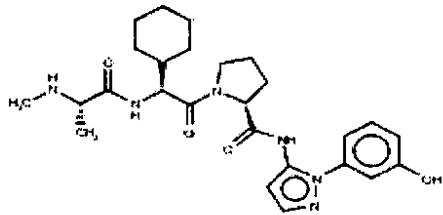
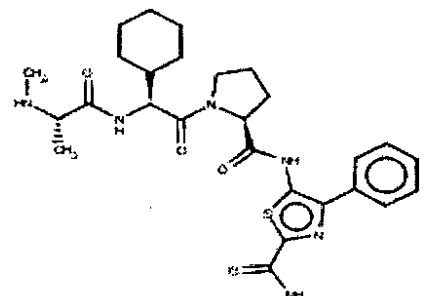
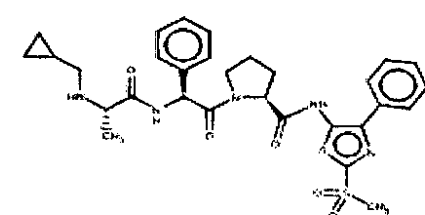
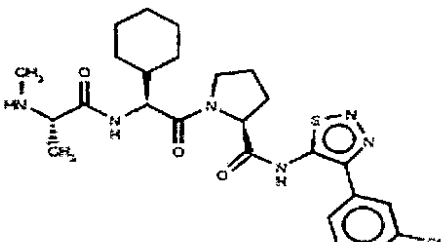
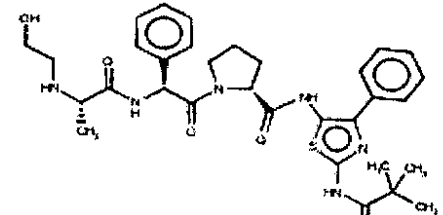
218		0,115
219		0,115
220		0,116
221		0,116
222		0,118

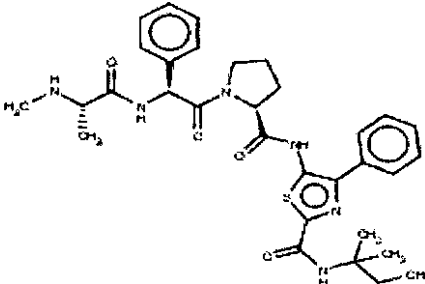
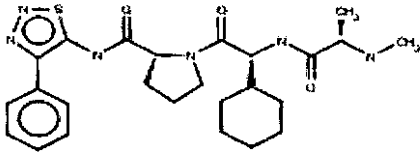
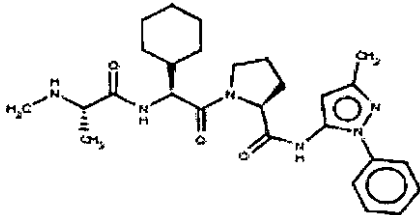
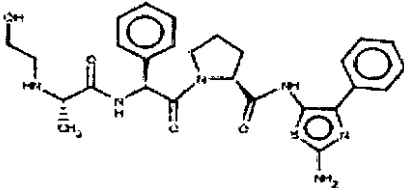
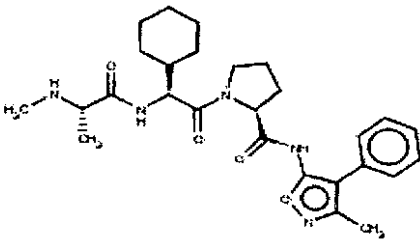
223		0,118
224		0,119
225		0,123
226		0,123
227		0,127

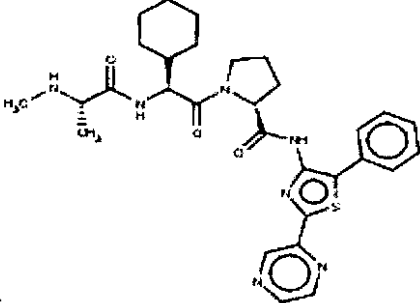
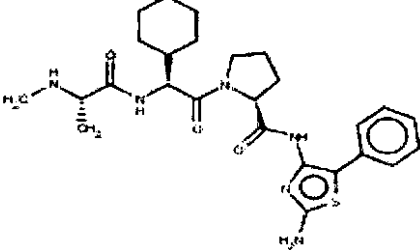
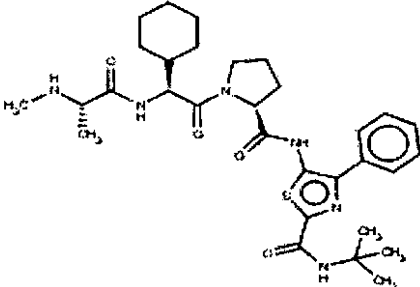
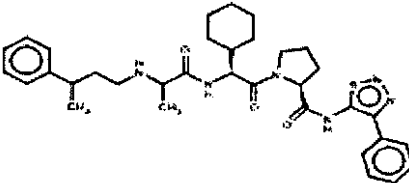
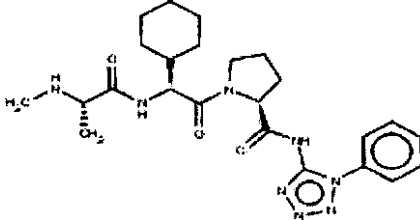
228		0,127
229		0,127
230		0,13
231		0,13
232		0,131

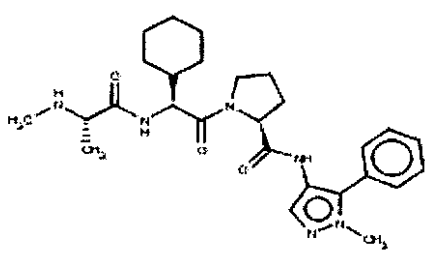
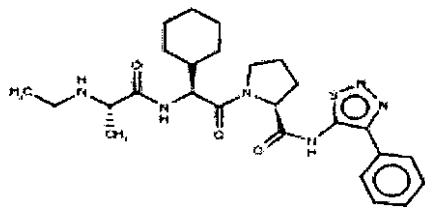
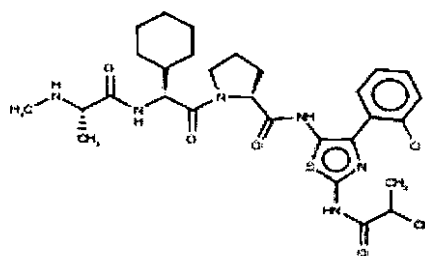
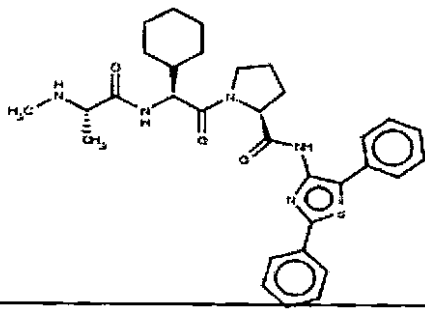
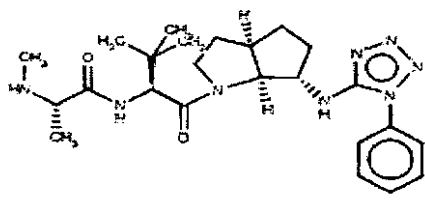
233		0,132
234		0,133
235		0,133
236		0,136
237		0,138

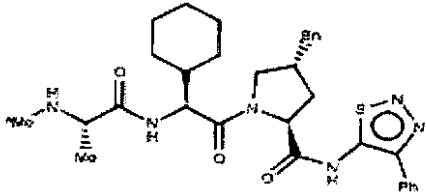
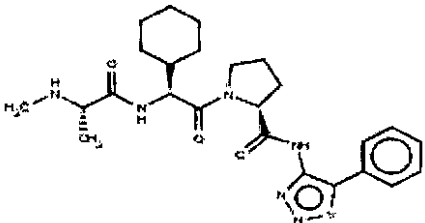
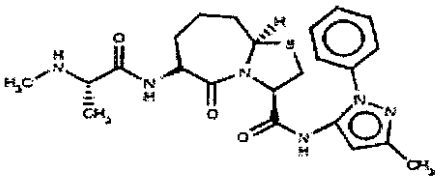
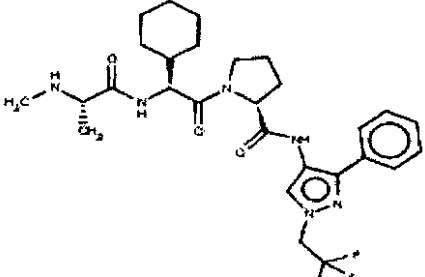
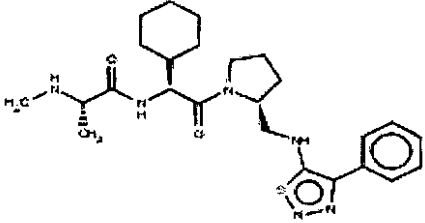
238		0,14
239		0,142
240		0,144
241		0,148
242		0,15

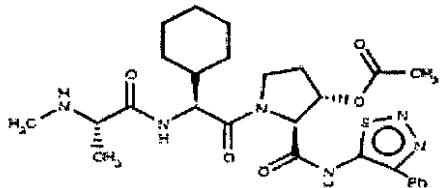
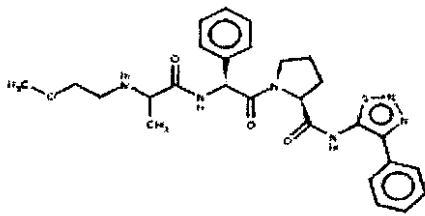
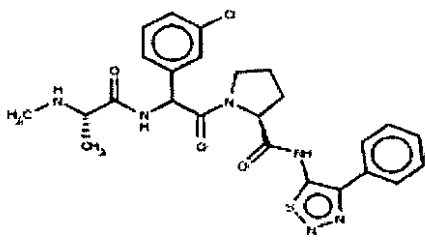
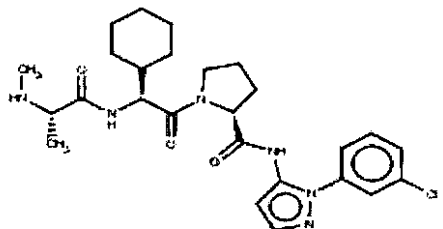
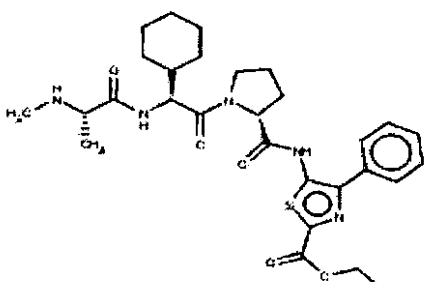
243		0,151
244		0,156
245		0,157
246		0,158
247		0,159

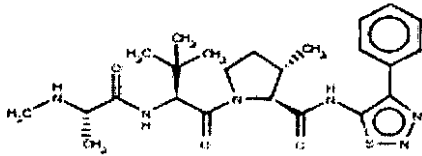
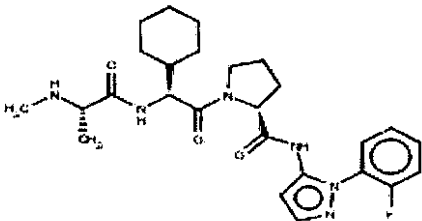
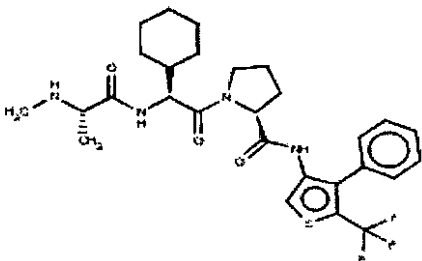
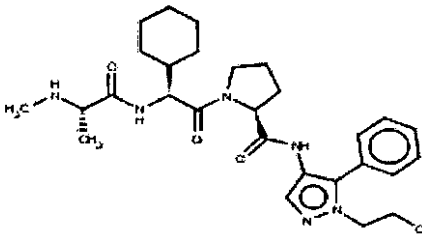
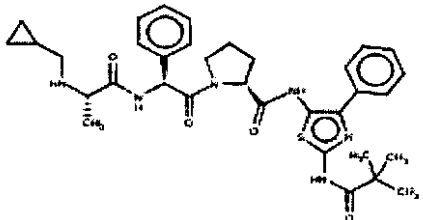
248		0,16
249		0,162
250		0,164
251		0,172
252		0,177

253		0,178
254		0,179
255		0,185
256		0,19
257		0,203

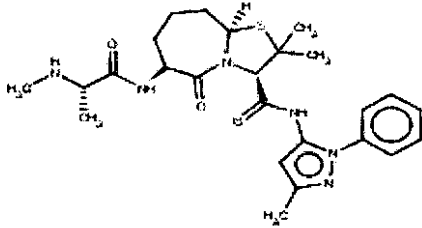
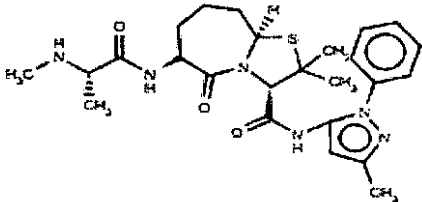
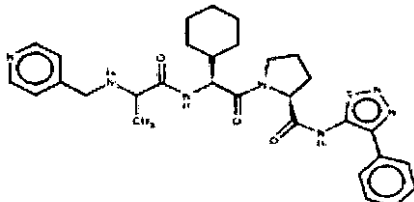
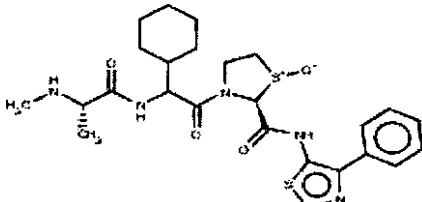
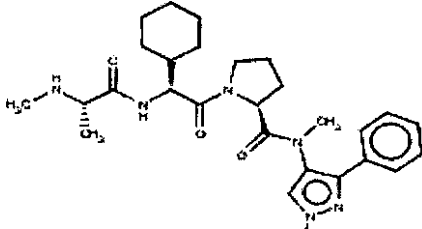
258		0,22
259		0,221
260		0,226
261		0,242
262		0,252

263		0,257
264		0,264
265		0,269
266		0,278
267		0,299

268	 <chem>CCN[C@@H](C(=O)N[C@H](C1CCCCC1)C(=O)N2CCCC2C(=O)Nc3cc[nH]3)C(=O)OC</chem>	0,302
269	 <chem>CCCCN[C@@H](C(=O)N[C@H](c1ccccc1)C(=O)N2CCCC2C(=O)Nc3cc[nH]3)C</chem>	0,308
270	 <chem>CCN[C@@H](C(=O)N[C@H](c1cccc(Cl)c1)C(=O)N2CCCC2C(=O)Nc3cc[nH]3)C</chem>	0,343
271	 <chem>CCN[C@@H](C(=O)N[C@H](C1CCCCC1)C(=O)N2CCCC2C(=O)Nc3cc[nH]3)C(=O)Nc4ccc(Cl)cc4</chem>	0,351
272	 <chem>CCN[C@@H](C(=O)N[C@H](C1CCCCC1)C(=O)N2CCCC2C(=O)Nc3cc[nH]3)C(=O)Nc4c[nH]c5ccccc45</chem>	0,353

273		0,37
274		0,389
275		0,39
276		0,41
277		0,413

278	<chem>C[C@H](Cc1ccccc1)C(=O)N[C@@H](C2CCCCC2)C(=O)N3CC[C@H](C)C3C(=O)Nc4nc5ccccc5nn4</chem>	0,416
279	<chem>C[C@H](Cc1ccccc1)C(=O)N[C@@H](C2CCCCC2)C(=O)N3CC[C@H](C)C3C(=O)Nc4nc5ccccc5nn4</chem>	0,431
280	<chem>C[C@H](Cc1ccccc1)C(=O)N[C@@H](C2CCCCC2)C(=O)N3CC[C@H](C)C3C(=O)Nc4nc5ccccc5nn4</chem>	0,484
281	<chem>C[C@H](Cc1ccccc1)C(=O)N[C@@H](C2CCCCC2)C(=O)N3CC[C@H](C)C3C(=O)Nc4nc5ccccc5nn4</chem>	0,495
282	<chem>C[C@H](Cc1ccccc1)C(=O)N[C@@H](C2CCCCC2)C(=O)N3CC[C@H](C)C3C(=O)Nc4nc5ccccc5nn4</chem>	0,614

283		0,704
284		0,774
285		0,875
286		0,88
287		0,904