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Thiophene derivatives as antiviral agents for flavivirus infection

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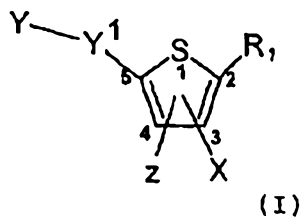
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ABSTRACT

The present invention provides novel compounds represented by
 formula I: or pharmaceutically acceptable salts thereof for
 5 treating flaviviridae viral infection.



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AUSTRALIA

Patents Act 1990

COMPLETE SPECIFICATION

Standard Patent

Applicant(s):

Vertex Pharmaceuticals (Canada) Incorporated

Invention Title:

*THIOPHENE DERIVATIVES AS ANTIVIRAL AGENTS FOR FLAVIVIRUS
INFECTION*

The following statement is a full description of this invention,
including the best method for performing it known to me/us:

COMPOUNDS AND METHODS FOR THE TREATMENT OR
PREVENTION OF FLAVIVIRUS INFECTIONS

FIELD OF THE INVENTION

The present invention relates to novel compounds and a method for the treatment or prevention of *Flavivirus* infections using novel compounds.

BACKGROUND OF THE INVENTION

Hepatitis is a disease occurring throughout the world. It is generally of viral nature, although there are other causes known. Viral hepatitis is by far the most common form of hepatitis.

Nearly 750,000 Americans are affected by hepatitis each year, and out of those, more than 150,000 are infected with the hepatitis C virus ("HCV").

HCV is a positive-stranded RNA virus belonging to the *Flaviviridae* family and has closest relationship to the pestiviruses that include hog cholera virus and bovine viral diarrhea virus (BVDV). HCV is believed to replicate through the production of a complementary negative-strand RNA template. Due to the lack of efficient culture replication system for the virus, HCV particles were isolated from pooled human plasma and shown, by electron microscopy, to have a diameter of about 50-60 nm. The HCV genome is a single-stranded, positive-sense RNA of about 9,600 bp coding for a polyprotein of 3009-3030 amino-acids, which is cleaved co and post-translationally by cellular and two viral proteinases into mature viral proteins (core, E1, E2, p7, NS2, NS3, NS4A, NS4B, NS5A, NS5B). It is believed that the structural proteins, E1 and E2, the major glycoproteins are embedded into a viral lipid envelope and form stable heterodimers. It is also believed that the structural core protein interacts with the viral RNA genome to form the nucleocapsid. The nonstructural proteins designated NS2 to NS5 include proteins with enzymatic functions involved in virus replication and protein processing including a polymerase, protease and helicase.

The main source of contamination with HCV is blood. The magnitude

of the HCV infection as a health problem is illustrated by the prevalence among high-risk groups. For example, 60% to 90% of hemophiliacs and more than 80% of intravenous drug abusers in western countries are chronically infected with HCV. For

5 intravenous drug abusers, the prevalence varies from about 28% to 70% depending on the population studied. The proportion of new HCV infections associated with post-transfusion has been markedly reduced lately due to advances in diagnostic tools used to screen blood donors.

10

The only treatment currently available for HCV infection is interferon- α (IFN- α). However, according to different clinical studies, only 70% of treated patients normalize alanine aminotransferase (ALT) levels in the serum and after

15 discontinuation of IFN, 35% to 45% of these responders relapse. In general, only 20% to 25% of patients have long-term responses to IFN. Clinical studies have shown that combination treatment with IFN and ribavirin (RIBA) results in a superior clinical response than IFN alone. Different genotypes of HCV respond differently to

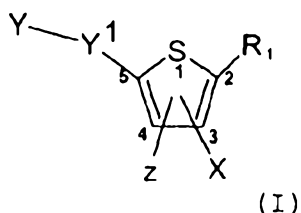
20 IFN therapy, genotype 1b is more resistant to IFN therapy than type 2 and 3.

There is therefore a great need for the development of anti-viral agents.

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SUMMARY OF THE INVENTION

In one aspect, the present invention provides novel compounds represented by formula I:

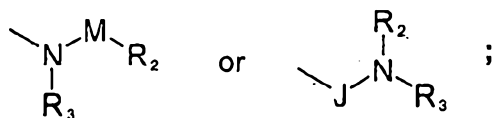


30

or pharmaceutically acceptable salts thereof;

wherein,

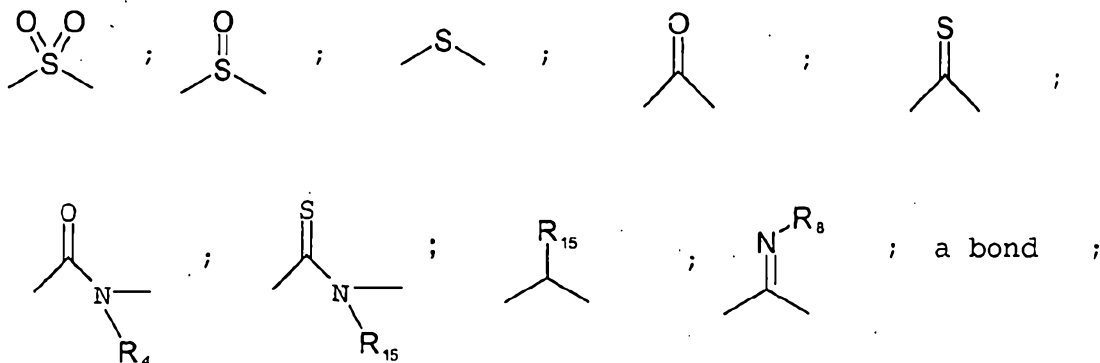
X is chosen from:



wherein,

M is chosen from:

5

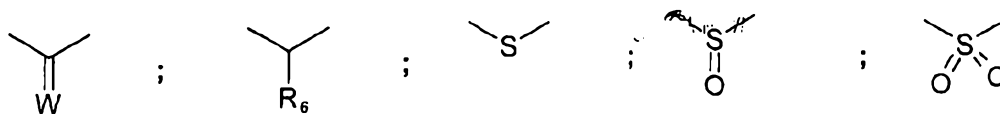


wherein,

R₄ is C₁₋₆ alkyl;

10 R₈ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₂ heteroaralkyl, C₆₋₁₆ aralkyl; and
R₁₅ is chosen from H or C₁₋₆ alkyl;

J is chosen from:



wherein W is chosen from O, S or NR_7 ,

wherein R_7 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl;

and R_6 is chosen from H, C_{1-12} alkyl, C_{6-14} aryl or C_{6-16} aralkyl;

Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{1-6} alkynyl;

Y is chosen from COOR_{16} , COCOOR_5 , $\text{P}(\text{O})\text{OR}_a\text{OR}_b$, $\text{S}(\text{O})\text{OR}_5$, $\text{S}(\text{O})_2\text{OR}_5$, tetrazole, $\text{CON}(\text{R}_9)\text{CH}(\text{R}_5)\text{COOR}_5$, $\text{CONR}_{10}\text{R}_{11}$, $\text{CON}(\text{R}_9)-\text{SO}_2-\text{R}_5$, CONR_9OH or halogen, wherein R_9 , R_5 , R_{10} and R_{11} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;

or R_{10} and R_{11} are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_a and R_b are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

or R_a and R_b are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R_{16} is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

provided that R_{16} is other than methyl or ethyl;

R_1 is chosen from C_{2-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

R_2 is chosen from C_{2-12} alkyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, or C_{6-18} aralkyl;

R_3 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

Z is chosen from H, halogen, C_{1-6} alkyl;

with the proviso that:

i) when X is 4-Chloro-2,6-dimethyl-benzenesulfonamide and, R₁ is phenyl, and R₂ is H, and Y¹ is a bond, then Y is other than CONH₂;
5 compound #580

ii) when X is Toluene-4-sulfonamide and R₁ is 4-chloro-phenyl, and R₂ is H, and Y¹ is a bond, then Y is other than CONH₂;
10 compound #563

iii) when X is Toluene-4-sulfonamide and R₁ is 4-fluoro-phenyl, and R₂ is H, and Y¹ is a bond, then Y is other than CONH₂;
15 compound #564

iv) when X is Toluene-4-sulfonamide and R₁ is 4-methoxy-phenyl, and R₂ is H, and Y¹ is a bond, then Y is other than CONH₂;
20 compound #565

v) when X is Benzamide and R₁ is phenyl Y¹ is a bond and Y is COOH then R₂ is other than hydrogen.

The compounds of the present invention are useful in therapy, particularly as antivirals.

25 In another aspect, there is provided a method of treating viral infections in a subject in need of such treatment comprising administering to the subject a therapeutically effective amount of a compound of formula (I) or composition of the invention.

30 In still another aspect, there is provided a method of treating viral infections in a subject in need of such treatment comprising administering to the subject a combination comprising at least one compound of formula (I) and at least one further therapeutic agent.

35 In another aspect, there is provided a pharmaceutical formulation comprising the compound of the invention in combination with a pharmaceutically acceptable carrier or excipient.

Another aspect of the invention is the use of a compound according to formula (I), for the manufacture of a medicament for the treatment of viral infections.

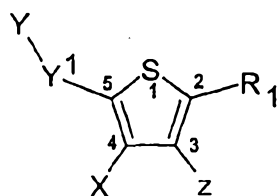
- 5 In another aspect, there is provided a method for inhibiting or reducing the activity of viral polymerase in a host comprising administering a therapeutically effective amount of a compound of formula (I).

10 DETAILED DESCRIPTION OF THE INVENTION

In one embodiment, compounds of the present invention comprise those wherein the following embodiments are present, either independently or in combination.

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In one embodiment, the present invention provides novel compounds of formula (Ia):



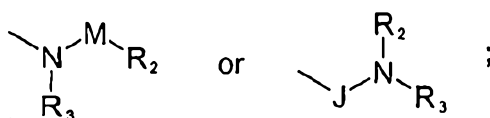
(Ia)

or pharmaceutically acceptable salts thereof;

20

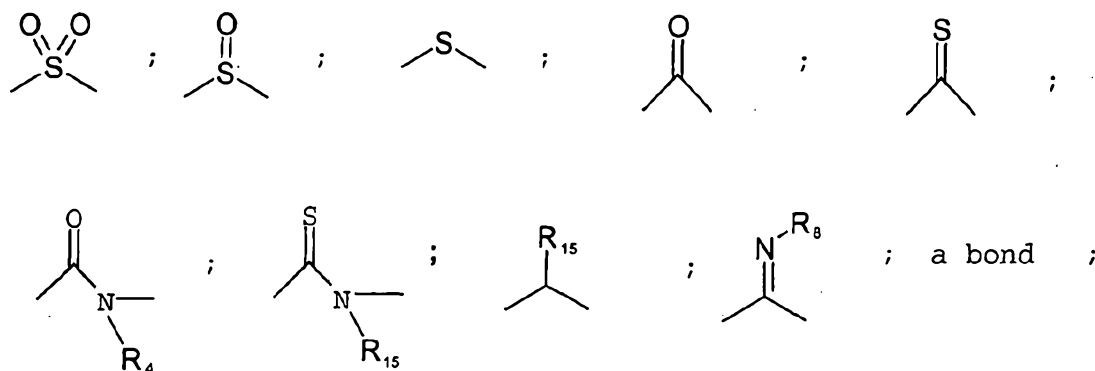
wherein,

X is chosen from:



wherein,

M is chosen from:



5 wherein,

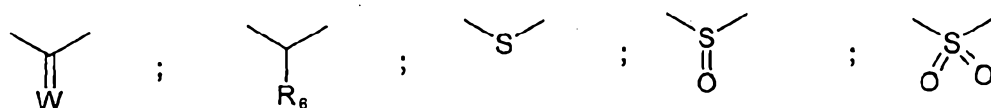
R_4 is C_{1-6} alkyl;

R_8 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl; and

R_{15} is chosen from H or C_{1-6} alkyl;

10

J is chosen from:



wherein W is chosen from O, S or NR_7 ,

15 wherein R_7 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl;

and R_6 is chosen from H, C_{1-12} alkyl, C_{6-14} aryl or C_{6-16} aralkyl;

Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

20

Y is chosen from COOR_{16} , COCOOR_5 , $\text{P}(\text{O})\text{OR}_a\text{OR}_b$, $\text{S}(\text{O})\text{OR}_5$, $\text{S}(\text{O})_2\text{OR}_5$, tetrazole, $\text{CON}(\text{R}_9)\text{CH}(\text{R}_5)\text{COOR}_5$, $\text{CONR}_{10}\text{R}_{11}$, $\text{CON}(\text{R}_9)-\text{SO}_2-\text{R}_5$, CONR_9OH or halogen, wherein R_9 , R_5 , R_{10} and R_{11} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-18} aralkyl;

25

or R_{10} and R_{11} are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_a and R_b are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

or R_a and R_b are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R_{16} is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl; provided that R_{16} is other than methyl or ethyl;

R_1 is chosen from C_{2-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

R_2 is chosen from C_{2-12} alkyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, or C_{6-18} aralkyl;

R_3 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

Z is chosen from H, halogen, C_{1-6} alkyl;

with the proviso that:

i) when X is 4-Chloro-2,6-dimethyl-benzenesulfonamide and, R_1 is phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than $CONH_2$;

compound #580

ii) when X is Toluene-4-sulfonamide and R_1 is 4-chloro-phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than $CONH_2$;

compound #563

iii) when X is Toluene-4-sulfonamide and R_1 is 4-fluoro-phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than $CONH_2$;

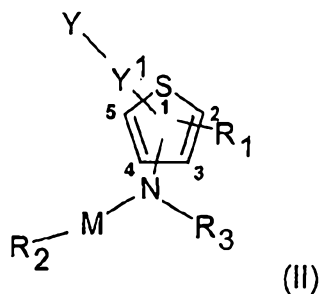
compound #564

iv) when X is Toluene-4-sulfonamide and R_1 is 4-methoxy-phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than $CONH_2$;

compound #565

v) when X is Benzamide and R_1 is phenyl Y^1 is a bond and Y is COOH then R_3 is other than hydrogen.

In a further aspect, the present invention provides novel
5 compounds represented by formula II:

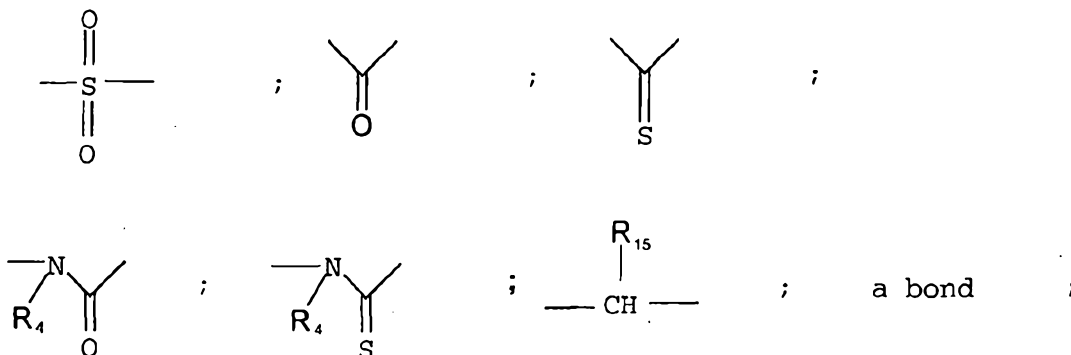


and pharmaceutically acceptable salts thereof,

wherein,

10

M is chosen from:



15 Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

Y is chosen from $COOR_{16}$, $CO-COOR_5$, $PO_3R_4R_5$, SO_3R_5 , tetrazole,
 $CON(R_9)CH(R_5)-COOR_5$, $CON R_{10}R_{11}$ or $CONR_5OH$, wherein

each R_5 , R_9 , R_{10} , R_{11} , R_{16} , R_4 , and R_5 are independently chosen
20 from H or C_{1-6} alkyl,;

R_1 is chosen from C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-12} aryl,
 C_{3-10} heterocycle, C_{3-10} heteroaralkyl, C_{6-12} aralkyl, or a halogen;

R_2 is chosen from C_{6-12} aryl, C_{3-10} heterocycle, C_{6-12} aralkyl or C_{3-10} heteroaralkyl;

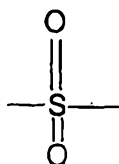
R_3 is chosen from H or C_{1-6} alkyl; C_{6-12} aralkyl or C_{3-10} heteroaralkyl;

R_4 is chosen from H or C_{1-6} alkyl;

R_{15} is chosen from H or C_{1-6} alkyl

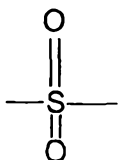
with the proviso that:

i) when M is



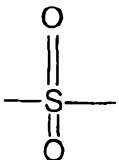
and R_2 is 4-chloro-2,5-dimethyl-phenyl, R_1 is phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than CONH_2 ; compound #580

ii) when M is



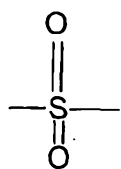
and R_2 is 4-methylphenyl, R_1 is 4-chloro-phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than CONH_2 ; compound #563

iii) when M is



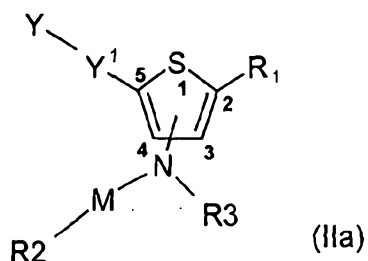
and R_2 is 4-methylphenyl, R_1 is 4-fluoro-phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than CONH_2 ; compound #564

iv) when M is



and R_2 is 4-methylphenyl, R_1 is 4-methoxy-phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than CONH_2 ; compound #565

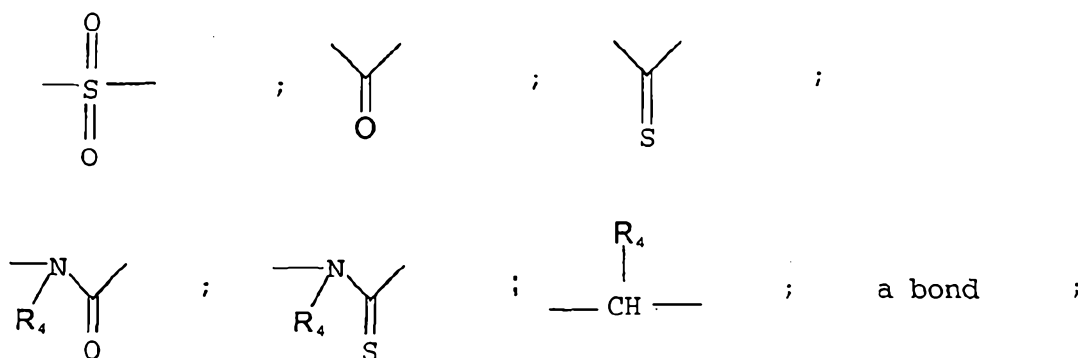
In still a further embodiment, the present invention provides
5 novel compounds of formula (IIa):



wherein,

M is chosen from:

10



Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

- 15 Y is chosen from COOR_5 , CO-COOR_5 , $\text{PO}_3\text{R}_6\text{R}_b$, SO_3R_5 , tetrazole, $\text{CON}(\text{R}_9)\text{CH}(\text{R}_5)\text{-COOR}_5$, $\text{CON} \text{R}_{10}\text{R}_{11}$ or CONR_5OH , wherein each R_5 , R_9 , R_{10} , R_{11} , R_{16} , R_a , and R_b are independently chosen from H or C_{1-6} alkyl,;
- R_1 is chosen from C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-12} aryl,
- 20 C_{3-10} heterocycle, C_{3-10} heteroaralkyl, C_{6-12} aralkyl, or a halogen;

R_2 is chosen from C_{6-12} aryl, C_{3-10} heterocycle, C_{6-12} aralkyl or C_{3-10} heteroaralkyl;

5 R_3 is chosen from H or C_{1-6} alkyl; C_{6-12} aralkyl or C_{3-10} heteroaralkyl;

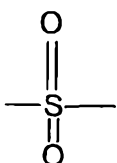
R_4 is chosen from H or C_{1-6} alkyl;

R_{15} is chosen from H or C_{1-6} alkyl;

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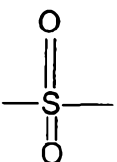
with the proviso that:

i) when M is



15 and R_2 is 4-chloro-2,5-dimethyl-phenyl, R_1 is phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than CONH_2 ; compound #580

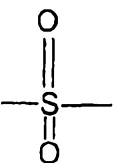
ii) when M is



and R_2 is 4-methylphenyl, R_1 is 4-chloro-phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than CONH_2 ; compound #563

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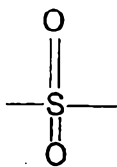
iii) when M is



and R_2 is 4-methylphenyl, R_1 is 4-fluoro-phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than CONH_2 ; compound #564

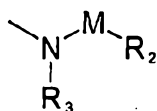
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iv) when M is



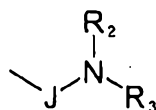
and R_2 is 4-methylphenyl, R_1 is 4-methoxy-phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than CONH_2 ; compound #565.

In one embodiment, X is:



5

In a further embodiment, X is:



In one embodiment, Z is chosen from H, halogen, C_{1-6} alkyl.

In further embodiments,

10 Z is H

Z is halogen

Z is fluoride

Z is C_{1-6} alkyl

Z is chosen from methyl, trifluoromethyl, ethyl, propyl,

15 isopropyl, cyclopropyl, butyl, isobutyl, cyclobutyl, pentyl, neopentyl, cyclopentyl, hexyl or cyclohexyl.

20 In further embodiments;

R_1 is chosen from C_{2-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl.

R_1 is chosen from a C_{2-12} alkyl, C_{6-14} aryl or C_{3-12} heterocycle.

R_1 is a C_{2-12} alkyl.

25 R_1 is a C_{6-14} aryl.

R_1 is a C_{3-12} heterocycle.

R_1 is chosen from t-butyl, isobutyl, allyl, ethynyl, 2-phenylethenyl, isobutenyl, benzyl, phenyl, phenethyl,

benzodioxolyl, thienyl, thiophenyl, pyridinyl, isoxazolyl, thiazolyl, pyrazolyl, tetrazolyl, benzofuranyl, indolyl, furanyl, or benzothiophenyl any of which can be optionally substituted by one or more substituent chosen from halogen, nitro, nitroso, SO_2R_{12} , PO_3RCRd , $\text{CONR}_{13}\text{R}_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-12} aralkyl, C_{6-12} aryl, C_{1-6} alkyloxy, C_{2-6} alkenyloxy, C_{2-6} alkynyloxy, C_{6-12} aryloxy, $\text{C}(\text{O})\text{C}_{1-6}$ alkyl, $\text{C}(\text{O})\text{C}_{2-6}$ alkenyl, $\text{C}(\text{O})\text{C}_{2-6}$ alkynyl, $\text{C}(\text{O})\text{C}_{6-12}$ aryl, $\text{C}(\text{O})\text{C}_{6-12}$ aralkyl, C_{3-10} heterocycle, hydroxyl, $\text{NR}_{13}\text{R}_{14}$, $\text{C}(\text{O})\text{OR}_{12}$, cyano, azido, amidino or guanido;

- wherein R_{12} , Rc , Rd , R_{13} and R_{14} are each independently chosen from H , C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl; or Rc and Rd are taken together with the oxygens to form a 5 to 10 membered heterocycle; or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

R_1 is chosen from thienyl, t-butyl, phenyl or pyridinyl.

R_1 is isoxazolyl substituted by at least one methyl.

R_1 is pyridinyl.

In one embodiment, R_1 is chosen from a C_{1-6} alkyl, C_{6-12} aryl or C_{3-10} heterocycle.

In one embodiment, R_1 is chosen from t-butyl, isobutyl, allyl, ethynyl, 2-phenylethenyl, isobutenyl, benzyl, phenyl, phenethyl, benzodioxolyl, thienyl, thiophenyl, pyridinyl, isoxazolyl, thiazolyl, pyrazolyl, tetrazolyl, benzofuranyl, indolyl, furanyl, or benzothiophenyl, any of which can be substituted by at least one substituent chosen from C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{3-10} heterocycle, halogen, nitro, $\text{CONR}_{13}\text{R}_{14}$, $\text{NR}_{13}\text{R}_{14}$, amidino, guanido, Cyano, $\text{SO}_2\text{-C}_{1-6}$ alkyl, $\text{C}(\text{O})\text{OR}_{12}$, C_{1-6} alkyloxy, C_{2-6} alkenyloxy, C_{2-6} alkynyloxy, or C_{6-12} aryloxy;

wherein R_{12} , R_{13} and R_{14} are each independently chosen from H , C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;

or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

5 In one embodiment, R_1 is chosen from thienyl, t-butyl, phenyl, thiophenyl, pyridinyl, isoxazolyl, any of which can be substituted by at least one substituent chosen from a halogen, C_{1-6} alkyl, C_{1-6} alkyloxy, C_{2-6} alkenyl, C_{2-6} alkynyl, nitro, cyano, SO_2 - C_{1-6} alkyl, NO - C_{1-6} alkyl.

10 In further embodiments;

R_1 is phenyl.

R_1 is phenyl substituted with fluoride.

R_1 is phenyl substituted with at least one fluoride

R_1 is phenyl di-substituted with fluoride.

15 R_1 is phenyl substituted with chloride.

R_1 is phenyl substituted with at least one chloride

R_1 is phenyl di-substituted with chloride.

R_1 is phenyl substituted with fluoride and chloride.

R_1 is phenyl substituted with nitro..

20 R_1 is phenyl substituted with at least one nitro.

R_1 is phenyl substituted with methoxy.

R_1 is phenyl substituted with OCF_3 .

R_1 is phenyl substituted with CF_3 .

R_1 is phenyl substituted with methyl.

25 R_1 is phenyl substituted with at least one methyl.

R_1 is phenyl substituted with CN.

R_1 is phenyl substituted with SO_2 - CH_3 .

R_1 is phenyl substituted with $NH(CO)$ - CH_3 .

30 In further embodiments,

R_1 is thiophenyl.

R_1 is thiophenyl substituted by at least one halogen.

R_1 is thiophenyl substituted by at least one chloride.

R_1 is thiophenyl substituted by at least one methyl.

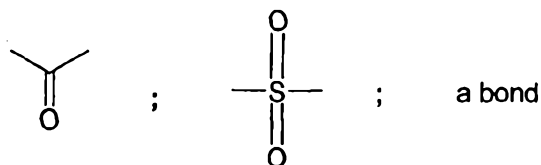
35 R_1 is thiophenyl substituted by at least one methyl and one chloride.

In further embodiments,

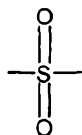
R_1 is thienyl.

R_1 is thienyl substituted by at least one halogen.
 R_1 is thienyl substituted by at least one chloride.
 R_1 is thienyl substituted by at least one methyl.
 R_1 is thienyl substituted by at least one methyl and one
 5 chloride.
 R_1 is isoxazole di-substituted with CH_3 .
 R_1 is pyridine.

In one embodiment, M is chosen from:



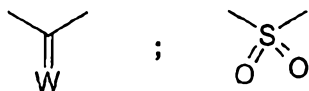
10 In a further embodiment, M is:



In an alternative embodiment, M is:



15 In one embodiment, J is chosen from:



wherein, W is as defined above.

In an alternative embodiment, J is:



20 In a further embodiment, J is:



In one embodiment, Y is chosen from COOR_{16} , COCOOR_5 , $\text{P(O)OR}_2\text{OR}_6$, $\text{S(O)}_2\text{OR}_3$, tetrazole, $\text{CON(R}_5\text{)CH(R}_5\text{)COOR}_5$, $\text{CONR}_{10}\text{R}_{11}$, CONR_3OH .

- 5 In a further embodiment, any of R_5 , R_a , R_b , R_9 , R_{10} , R_{11} and R_{16} are each independently chosen from H or C_{1-6} alkyl; provided that R_{16} is other than methyl or ethyl.

In one embodiment, Y is chosen from COOR_{16} , $\text{CONR}_{10}\text{R}_{11}$ or $\text{CON(R}_5\text{)CH(R}_5\text{)-COOR}_5$.

- 10 In a further embodiment, any of R_5 , R_9 , R_{10} , R_{11} and R_{16} are each independently chosen from H or C_{1-6} alkyl; provided that R_{16} is other than methyl or ethyl.

- In a further embodiment, Y is chosen from COOR_{16} , $\text{CONR}_{10}\text{R}_{11}$ or CON
15 $\text{R}_9\text{CH}_2\text{COOR}_5$.

In a further embodiment, Y is chosen from COOR_5 , CONR_5R_5 or $\text{CON(R}_5\text{)CH(R}_5\text{)COOR}_5$.

In a further embodiment, Y is COOH .

In a further embodiment, Y is CONH_2 .

- 20 In a further embodiment, Y is $\text{CONHCH}_2\text{COOH}$.
In a further embodiment, Y is COOCH_3 .

In a further embodiment, Y^1 is chosen from CH_2 , C=CH , CH-CH_2 or a bond.

25

In further embodiments;

R_3 is chosen from H, C_{1-12} alkyl, C_{6-18} aralkyl, C_{3-12} heterocycle or C_{3-18} heteroaralkyl.

R_3 is chosen from H, C_{1-12} alkyl, C_{6-18} aralkyl or C_{3-12} heterocycle.

- 30 R_3 is C_{1-12} alkyl.

R_3 is C_{6-18} aralkyl.

R_3 is C_{3-12} heterocycle.

R_3 is chosen from H, methyl, ethyl, i-propyl, cyclopropyl, cyclohexyl, allyl, piperidinyl, piperazinyl, pyrrolidinyl, azetidiny, aziridinyl, pyridinyl, piperidinylmethyl, dioxanyl, dioxolanyl, azepanyl or benzyl; any of which can be optionally substituted by one or more substituent chosen from halogen, nitro, nitroso, SO_3R_{12} , PO_3RcRd , $CONR_{13}R_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-12} aralkyl, C_{6-12} aryl, C_{1-6} alkyloxy, C_{2-6} alkenyloxy, C_{2-6} alkynyloxy, C_{6-12} aryloxy, $C(O)C_{1-6}$ alkyl, $C(O)C_{2-6}$ alkenyl, $C(O)C_{2-6}$ alkynyl, $C(O)C_{6-12}$ aryl, $C(O)C_{6-12}$ aralkyl, C_{3-10} heterocycle, hydroxyl, $NR_{13}R_{14}$, $C(O)OR_{12}$, cyano, azido, amidino or guanido; wherein R_{12} , Rc , Rd , R_{13} and R_{14} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl; or Rc and Rd are taken together with the oxygens to form a 5 to 10 membered heterocycle; or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

R_3 is chosen from H or Methyl, isopropyl, piperidinyl, piperidinylmethyl, dioxolanyl or cyclohexyl.

In a further embodiment, R_3 is H or methyl.

In a further embodiment, R_3 is H.

In a further embodiment, R_3 is methyl.

In a further embodiment, R_3 is benzyl, thiophenylmethyl, furanylmethyl.

In additional embodiments;

R_2 is C_{2-12} alkyl, C_{6-14} aryl or C_{3-12} heterocycle;

R_2 is C_{3-6} heterocycle.

R_2 is chosen from thienyl, furanyl, pyridinyl, oxazolyl, thiazolyl, pyrrolyl, benzofuranyl, indolyl, benzoxazolyl, benzothienyl, benzothiazolyl, piperazinyl, pyrrolidinyl or quinolinyl any of which can be optionally substituted by one or more substituent chosen from halogen, nitro, nitroso, SO_3R_{12} ,

PO_3RcRd , $CONR_{13}R_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-12} aralkyl, C_{6-12} aryl, C_{1-6} alkyloxy, C_{2-6} alkenyloxy, C_{2-6} alkynyloxy, C_{6-12} aryloxy,

$C(O)C_{1-6}$ alkyl, $C(O)C_{2-6}$ alkenyl, $C(O)C_{2-6}$ alkynyl, $C(O)C_{6-12}$ aryl, $C(O)C_{6-12}$ aralkyl, C_{3-10} heterocycle, hydroxyl, $NR_{13}R_{14}$, $C(O)OR_{12}$, cyano, azido, amidino or guanido;

wherein R_{12} , R_c , R_d , R_{13} and R_{14} are each independently chosen from

5 H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;

or R_c and R_d are taken together with the oxygens to form a 5 to 10 membered heterocycle;

or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 10 membered heterocycle.

R_2 is a heterocycle chosen from thienyl, furanyl, pyridinyl, pyrrolyl, indolyl, piperazinyl or benzothienyl.

R_2 is C_{2-12} alkyl.

R_2 is chosen from cyclopropyl, cyclobutyl, cyclopentyl, 15 cyclopentenyl cyclohexyl, cycloheptyl, 2-(cyclopentyl)-ethyl, methyl, ethyl, vinyl, propyl, propenyl, isopropyl, butyl, butenyl isobutyl, pentyl, neopentyl or t-butyl any of which can be optionally substituted by one or more substituent chosen from halogen, nitro, nitroso, SO_3R_{12} , $PO_3R_cR_d$, $CONR_{13}R_{14}$, C_{1-6} alkyl, C_{2-6} 20 alkenyl, C_{2-6} alkynyl, C_{6-12} aralkyl, C_{6-12} aryl, C_{1-6} alkyloxy, C_{2-6} alkenyloxy, C_{2-6} alkynyloxy, C_{6-12} aryloxy, $C(O)C_{1-6}$ alkyl, $C(O)C_{2-6}$ alkenyl, $C(O)C_{2-6}$ alkynyl, $C(O)C_{6-12}$ aryl, $C(O)C_{6-12}$ aralkyl, C_{3-10} heterocycle, hydroxyl, $NR_{13}R_{14}$, $C(O)OR_{12}$, cyano, azido, amidino or guanido;

25 wherein R_{12} , R_c , R_d , R_{13} and R_{14} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;

or R_c and R_d are taken together with the oxygens to form a 5 to 10 membered heterocycle;

30 or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

R_2 is C_{6-12} aryl.

R_2 is an aryl chosen from indenyl, naphthyl or biphenyl.

R_2 is phenyl substituted by one or more substituent chosen from halogen, nitro, nitroso, SO_3R_{12} , PO_3RcRd , $CONR_{13}R_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-12} aralkyl, C_{6-12} aryl, C_{1-6} alkyloxy, C_{2-6} alkenyloxy, C_{2-6} alkynyloxy, C_{6-12} aryloxy, $C(O)C_{1-6}$ alkyl, $C(O)C_{2-6}$ alkenyl, $C(O)C_{2-6}$ alkynyl, $C(O)C_{6-12}$ aryl, $C(O)C_{6-12}$ aralkyl, C_{3-10} heterocycle, hydroxyl, $NR_{13}R_{14}$, $C(O)OR_{12}$, cyano, azido, amidino or guanido;

wherein R_{12} , Rc , Rd , R_{13} and R_{14} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;

or Rc and Rd are taken together with the oxygens to form a 5 to 10 membered heterocycle;

or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

R_2 is phenyl substituted by one or two substituents chosen from halogen, nitro, nitroso, SO_3R_{12} , PO_3RcRd , $CONR_{13}R_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-12} aralkyl, C_{6-12} aryl, C_{1-6} alkyloxy, C_{2-6} alkenyloxy, C_{2-6} alkynyloxy, C_{6-12} aryloxy, $C(O)C_{1-6}$ alkyl, $C(O)C_{2-6}$ alkenyl, $C(O)C_{2-6}$ alkynyl, $C(O)C_{6-12}$ aryl, $C(O)C_{6-12}$ aralkyl, C_{3-10}

heterocycle, hydroxyl, $NR_{13}R_{14}$, $C(O)OR_{12}$, cyano, azido, amidino or guanido;

wherein R_{12} , Rc , Rd , R_{13} and R_{14} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;

or Rc and Rd are taken together with the oxygens to form a 5 to 10 membered heterocycle;

or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

R_2 is phenyl substituted by one or more substituent chosen from

halogen, nitro, $CONR_{13}R_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{1-6} alkyloxy,

$C(O)C_{1-6}$ alkyl, C_{6-12} aryl, C_{3-10} heterocycle, hydroxyl, $NR_{13}R_{14}$,

$C(O)OR_{12}$, cyano, azido, wherein R_{12} , R_{13} and R_{14} are each

independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl,

C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;

or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

R_2 is phenyl substituted by one or two substituents chosen from halogen, nitro, $\text{CONR}_{13}\text{R}_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{1-6} alkyloxy,

5 $\text{C}(\text{O})\text{C}_{1-6}$ alkyl, C_{6-12} aryl, C_{3-10} heterocycle, hydroxyl, $\text{NR}_{13}\text{R}_{14}$,

$\text{C}(\text{O})\text{OR}_{12}$, cyano, azido, wherein R_{12} , R_{13} and R_{14} are each

independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl,

C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;

or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to

10 10 membered heterocycle.

R_2 is phenyl substituted by one or two substituents chosen from

halogen, C_{1-6} alkyl, $\text{NR}_{13}\text{R}_{14}$, nitro, $\text{CONR}_{13}\text{R}_{14}$, $\text{C}(\text{O})\text{OC}_{1-6}$ alkyl, COOH or

C_{1-6} alkyloxy $\text{C}(\text{O})\text{OR}_{12}$, cyano, azido, wherein R_{12} , R_{13} and R_{14} are

each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12}

15 alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18}

aralkyl;

or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to

10 membered heterocycle.

In one embodiment, R_2 is chosen from C_{6-14} aryl, C_{3-12} heterocycle,

20 C_{6-12} aralkyl or C_{3-10} heteroaralkyl.

In a further embodiment, R_2 is chosen from a C_{6-12} aryl or C_{3-10} heterocycle.

In a further embodiment, R_2 is a C_6 aryl or a C_{3-6} heterocycle.

In a further embodiment, R_2 is chosen from phenyl, pyridinyl,

25 thiophenyl, benzofuran, thiazole, pyrazole, substituted with at

least one substituent chosen from a halogen, C_{1-6} alkyl, C_{1-6}

alkyloxy, CF_3 , COOH , COOC_{1-6} alkyl, cyano, NH_2 , nitro, $\text{NH}(\text{C}_{1-6}$

alkyl), $\text{N}(\text{C}_{1-6} \text{ alkyl})_2$ or a C_{3-9} heterocycle.

R_2 is chosen from thienyl, furanyl, pyridyl, oxazolyl, thiazolyl,

30 pyrrolyl, benzofuranyl, indolyl, benzoxazolyl, benzothienyl,

benzothiazolyl or quinolinyl any of which can be substituted by

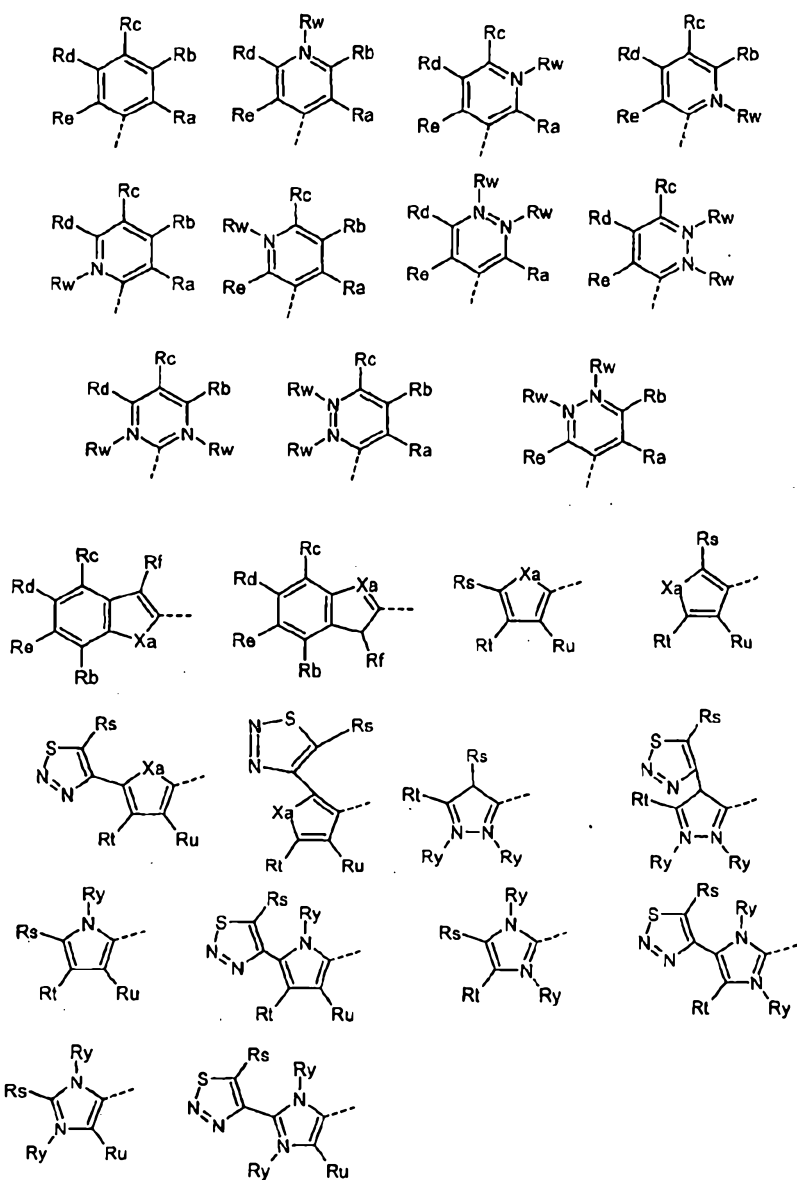
at least one substituent chosen from C_{1-6} alkyl, amino, halogen,

nitro, amido, CN, COOC_{1-6} alkyl, or C_{1-6} alkyloxy.

R_2 is methylphenyl.

35 R_2 is dichlorophenyl.

In a further embodiment, R_2 is chosen from:



wherein:

5

R_w is H, O or methyl;

R_y is H or methyl;

R_w is H;

R_w is methyl;

10 R_y is H;

R_y is methyl;

and wherein, Xa is S, N, O or carbon.

In a further embodiment, each of Ra, Rb, Rc, Rd, Re, and Rf are independently chosen from, H, Cl, Br, I, F, C₁₋₆ alkyl, OC₁₋₆ alkyl, CF₃, COOH, COOC₁₋₆ alkyl, CN, NH₂, NO₂, NH(C₁₋₆ alkyl), N(C₁₋₆ alkyl)₂.

In a further embodiment, each of Ra, Rb, Rc, Rd, Re, and Rf are independently chosen from, H, Cl, Br, I, F, methyl, O-methyl, CF₃, COOH, COOCH₃, CN, NH₂, NO₂, NH(CH₃) or N(CH₃)₂.

In a further embodiment, each of Ra, Rb, Rc, Rd, Re, and Rf are independently chosen from, H, Cl, Br, I, F, methyl, O-methyl, CF₃, COOH, COOCH₃, CN, NH₂, or NO₂.

In a further embodiment, each of Ra, Rb, Rc, Rd, Re, and Rf are independently chosen from, H, Cl, methyl, O-methyl, CF₃, COOH, COOCH₃, CN, NH₂, or NO₂.

In a further embodiment, each of Ra, Rb, Rc, Rd, Re, and Rf are independently chosen from, H, Cl, F, methyl, CF₃ or O-methyl.

In one embodiment, Rf is H or methyl.

In another embodiment, Rf is H.

In another embodiment, Rf is methyl.

In a further embodiment, each of Ra, Rb, Rc, Rd and Re is independently chosen from, H or Cl.

In a further embodiment, each of Ra, Rb, Rc, Rd and Re is H.

In one embodiment:

Ra is chosen from Cl, F, methyl or O-methyl;

Rb is H;

Rc is chosen from Cl, F, methyl or O-methyl;

Rd is H;

Re is chosen from Cl, F, methyl or O-methyl.

In one embodiment:

Ra is methyl;

Rb is H;

5 Rc is Cl;

Rd is H;

Re is methyl.

10 In a further embodiment, each of Rs, Rt, Ru, are independently chosen from, H, Cl, Br, I, F, C₁₋₆ alkyl, OC₁₋₆ alkyl, CF₃, COOH, COOC₁₋₆ alkyl, CN, NH₂, NO₂, NH(C₁₋₆ alkyl), N(C₁₋₆ alkyl)₂.

15 In a further embodiment, each of Rs, Rt, Ru, are independently chosen from, H, Cl, Br, I, F, methyl, O-methyl, CF₃, COOH, COOCH₃, CN, NH₂, NO₂, NH(CH₃) or N(CH₃)₂.

20 In a further embodiment, each of Rs, Rt, Ru, are independently chosen from, H, Cl, Br, I, F, methyl, O-methyl, CF₃, COOH, COOCH₃, CN, NH₂, or NO₂.

In a further embodiment, each of Rs, Rt, Ru, are independently chosen from, H, Cl, Br, I, F, methyl, O-methyl, CF₃, COOH, COOCH₃, CN, NH₂, or NO₂.

25 In a further embodiment, each of Rs, Rt, Ru, are independently chosen from, H, Cl, methyl, O-methyl, CF₃, COOH, COOCH₃, CN, NH₂, or NO₂.

30 In a further embodiment, each of Rs, Rt, Ru, are independently chosen from, H, Cl, F, methyl, CF₃ or O-methyl.

In a further embodiment, each of Rs, Rt, Ru, are independently chosen from, H or Cl.

35 In a further embodiment, each of Rs, Rt, Ru, are H.

In one embodiment:

Rs and Ru are Cl and Rt is H.

Rs is Cl, Rt and Ru are H.

5

In one embodiment, the viral infection is chosen from Flavivirus infections.

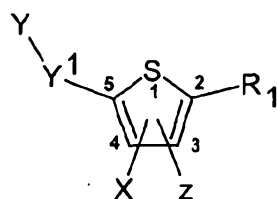
In one embodiment, the Flavivirus infection is chosen from

10 Hepatitis C virus (HCV), bovine viral diarrhea virus (BVDV), hog cholera virus and yellow fever virus.

In another embodiment, the Flavivirus infection is Hepatitis C viral infection.

15

In one embodiment, there is provided a method for treating or preventing a Flaviviridae viral infection in a host comprising administering to the host a therapeutically effective amount of at least one compound of formula (III)



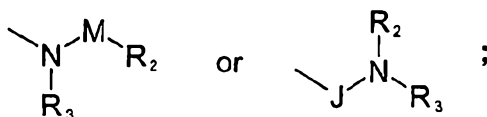
(III)

20

or pharmaceutically acceptable salts thereof;

wherein,

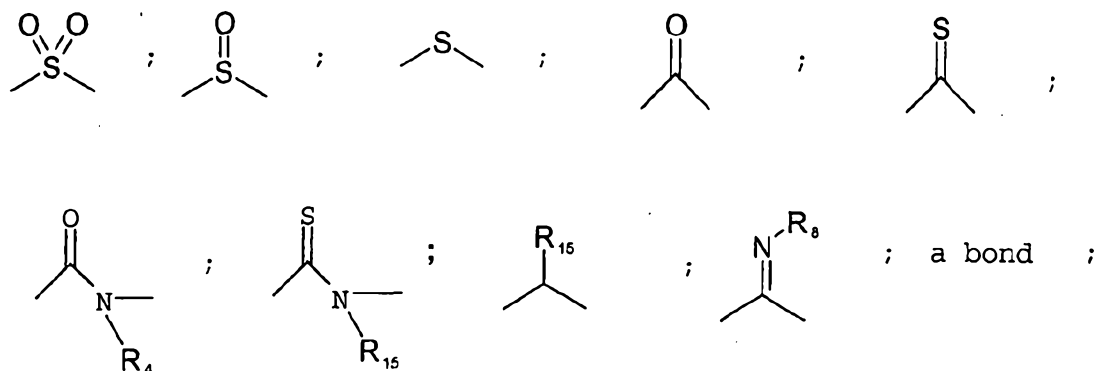
X is chosen from:



25

wherein,

M is chosen from:

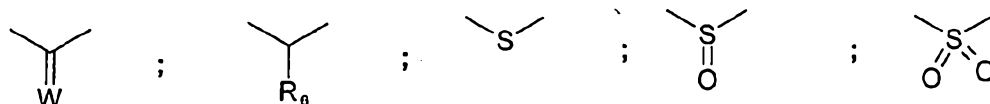


wherein,

R_4 is chosen from H or C_{1-6} alkyl;

- 5 R_8 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl; and R_{15} is chosen from H or C_{1-6} alkyl;

J is chosen from:



10

wherein

W is chosen from O, S or NR_7 ,

wherein R_7 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-12} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl;

15

and R_6 is chosen from H, C_{1-12} alkyl, C_{6-12} aryl or C_{6-16} aralkyl;

Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

- 20 Y is chosen from COOR_{16} , COCOOR_5 , $\text{P(O)OR}_4\text{OR}_5$, S(O)OR_5 , $\text{S(O)}_2\text{OR}_5$, tetrazole, $\text{CON(R}_9\text{)CH(R}_5\text{)COOR}_5$, $\text{CONR}_{10}\text{R}_{11}$, $\text{CON(R}_9\text{)-SO}_2\text{-R}_5$, CONR_9OH or halogen, wherein R_9 , R_5 , R_{10} and R_{11} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;
- 25 or R_{10} and R_{11} are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_a and R_b are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

or R_a and R_b are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R_{16} is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

R_1 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl, or halogen;

R_2 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, or C_{6-18} aralkyl;

R_3 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

Z is chosen from H, halogen, C_{1-6} alkyl.

In one embodiment, there is provided a method for treating or preventing Flaviviridae viral infection in a host comprising administering to the host a therapeutically effective amount of at least one compound of formula (III), further comprising at least one antiviral agent.

In one embodiment, the antiviral agent is chosen from a viral serine protease inhibitor, viral polymerase inhibitor and viral helicase inhibitor.

In a further embodiment, the antiviral agent is chosen from interferon α and ribavirin.

In a further embodiment, said compound of formula (III) and said antiviral agent are administered sequentially.

In a further embodiment, said compound of formula (III) and said antiviral agent are administered simultaneously.

In one embodiment, there is provided a method for treating or preventing Flaviviridae viral infection in a host comprising administering to the host a therapeutically effective amount of
 5 at least one compound of formula (III) and at least one additional agent chosen from immunomodulating agent, antioxidant agent, antibacterial agent or antisense agent.

In another embodiment, the additional agent is chosen from
 10 silybum marianum, interleukine-12, amantadine, ribozyme, thymosin, N-acetyl cysteine or cyclosporin.

In further embodiments;

The compound of formula (III) and the additional agent are administered sequentially.

15 The compound of formula (III) and the additional agent are administered simultaneously.

In one embodiment, the present invention further provides A pharmaceutical composition comprising at least one compound
 20 having the formula III or pharmaceutically acceptable salts thereof; and at least one pharmaceutically acceptable carrier or excipient.

In a further embodiment, the pharmaceutical composition, is
 25 further comprising one or more additional agent chosen from antiviral agent, immunomodulating agent, antioxidant agent, antibacterial agent or antisense agent.

In one embodiment, the antiviral agent is chosen from a viral serine protease inhibitor, viral polymerase inhibitor and viral
 30 helicase inhibitor.

In one embodiment, the antiviral agent is chosen from interferon α and ribavirin.

In one embodiment, the additional agent is chosen from silybum marianum, interleukine-12, amantadine, ribozyme, thymosin, N-
 35 acetyl cysteine or cyclosporin.

In one embodiment, the invention further provides the use of a compound having the formula III for the manufacture of a medicament for treating or preventing a viral Flaviridea infection in a host

In one embodiment, there is provided the use of a compound having the formula III or pharmaceutically acceptable salts thereof in therapy

In one embodiment, the invention provides the use of a compound having the formula III for treating or preventing Flaviviridae viral infection in a host.

In one embodiment, the use of a compound having the compound of formula III for treating or preventing Flaviviridae viral infection in a host is further comprising one or more additional agent chosen from antiviral agent, immunomodulating agent, antioxydant agent, antibacterial agent or antisense agent.

In one embodiment, the antiviral agent is chosen from a viral serine protease inhibitor, viral polymerase inhibitor and viral helicase inhibitor.

In one embodiment, the antiviral agent is chosen from interferon α and ribavirin.

In one embodiment, the additional agent is chosen from silybum marianum, interleukine-12, amantadine, ribozyme, thymosin, N-acetyl cysteine or cyclosporin.

In one embodiment, the compound of formula III and the additional agent are administered sequentially.

In one embodiment, the compound of formula III and the additional agent are administered simultaneously.

In one embodiment, there is provided a method for inhibiting or reducing the activity of viral polymerase in a host comprising administering a therapeutically effective amount of a compound of formula (III).

In one embodiment, the method for inhibiting or reducing the activity of viral polymerase in a host comprising administering a therapeutically effective amount of a compound of formula (III) is further comprising one or more viral polymerase inhibitor.

In further embodiments;

The viral polymerase is a Flaviviridae viral polymerase.

The viral polymerase is a RNA-dependant RNA-polymerase.

The viral polymerase is HCV polymerase.

In one embodiment, the invention provides a method for inhibiting or reducing the activity of viral helicase in a host comprising administering a therapeutically effective amount of a compound having the formula III.

In one embodiment, the invention provides a method for inhibiting or reducing the activity of viral helicase in a host comprising administering a therapeutically effective amount of a compound chosen from:

Compound #14 3-(4-Chloro-2,5-dimethyl-benzenesulfonylamino)-5-(4-chloro-phenyl)-thiophene-2-carboxylic acid

Compound #19 3-(4-Chloro-2,5-dimethyl-benzenesulfonylamino)-5-(4-isobutyl-phenyl)-thiophene-2-carboxylic acid

Compound #223 3-(4-Bromo-2-fluorobenzenesulfo-nylamino)-5-(4-isobutylphenyl)-thiophene-2-carboxylic acid

Compound #224 3-(4-Bromo-2-methylbenzenesulfo-nylamino)-5-(4-isobutylphenyl)-thiophene-2-carboxylic acid

Compound #225 5-(4-Isobutylphenyl 3-(3-methoxy-benzenesulfonylamino)-thiophene-2-carboxylic acid

Compound #581 5-(4-Isobutyl-phenyl)-3-[5-(5-trifluoromethyl-isoxazol-3-yl)-thiophene-2-sulfonylamino]-thiophene-2-carboxylic acid

Compound #227 3-[2,5-Bis-(2,2,2-trifluoroethoxy)-benzenesulfonylamino]-5-(4-isobutyl-phenyl)-thiophene-2-carboxylic acid

Compound #228 3-(2-Chloro-4-cyanobenzenesulfonylamino)-5-(4-isobutylphenyl)-thiophene-2-carboxylic acid

Compound #582 5-(4-Isobutyl-phenyl)-3-(2,3,4-trifluoro-
benzenesulfonylamino)-thiophene-2-carboxylic acid
or pharmaceutically acceptable salts thereof.

In further embodiments;

The viral helicase is a flaviviridea helicase.

The viral helicase is HCV helicase.

5

In a further embodiment, there is provided the use of a compound
having the formula III for inhibiting or reducing the activity
of viral polymerase in a host.

10 In still a further embodiment, there is provided the use of a
compound having the formula III for inhibiting or reducing the
activity of viral polymerase in a host, further comprising one
or more viral polymerase inhibitor.

In further embodiments;

15 The viral polymerase is Flaviviridae viral polymerase.

The viral polymerase is RNA-dependant RNA-polymerase.

The viral polymerase is HCV polymerase.

20 In one embodiment, the invention provides the use of a compound
having the formula III for inhibiting or reducing the activity
of viral helicase in a host.

In one embodiment, the invention provides the use of a compound
chosen from:

Compound #14 3-(4-Chloro-2,5-dimethyl-benzenesulfonylamino)-5-(4-
chloro-phenyl)-thiophene-2-carboxylic acid

Compound #19 3-(4-Chloro-2,5-dimethyl-benzenesulfonylamino)-5-(4-
isobutyl-phenyl)-thiophene-2-carboxylic acid

Compound #223 3-(4-Bromo-2-fluorobenzenesulfo-nylamino)-5-(4-
isobutylphenyl)-thiophene-2-carboxylic acid

Compound #224 3-(4-Bromo-2-methylbenzenesulfo-nylamino)-5-(4-
isobutylphenyl)-thiophene-2-carboxylic acid

Compound #225 5-(4-Isobutylphenyl 3-(3-methoxy-benzenesulfonyl-

amino)-thiophene-2-carboxylic acid

Compound #581 5-(4-Isobutyl-phenyl)-3-[5-(5-trifluoromethyl-
isoxazol-3-yl)-thiophene-2-sulfonylamino]-thiophene-
2-carboxylic acid

Compound #227 3-[2,5-Bis-(2,2,2-trifluoroethoxy)-
benzenesulfonylamino]-5-(4-isobutyl-phenyl)-
thiophene-2-carboxylic acid

Compound #228 3-(2-Chloro-4-cyanobenzenesulfonylamino)-5-(4-
isobutylphenyl)-thiophene-2-carboxylic acid

Compound #582 5-(4-Isobutyl-phenyl)-3-(2,3,4-trifluoro-
benzenesulfonylamino)-thiophene-2-carboxylic acid

or pharmaceutically acceptable salts thereof for inhibiting or
reducing the activity of viral helicase in a host.

In one embodiment, the invention provides the use of a compound
5 having the formula III for inhibiting or reducing the activity
of viral helicase in a host further comprising one or more viral
helicase inhibitor.

In further embodiments;

10 The viral helicase is Flaviviridae viral helicase.

The viral helicase is HCV helicase.

In one embodiment, the present invention provides a combination
comprising a compound having the formula III and one or more
15 additionnal agent chosen from viral serine protease inhibitor,
viral polymerase inhibitor and viral helicase inhibitor,
immunomodulating agent, antioxydant agent, antibacterial agent
or antisense agent.

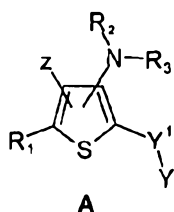
20 In a further embodiment, the additional agent is chosen from
silybum marianum, interleukine-12, amantadine, ribozyme,
thymosin, N-acetyl cysteine, cyclosporin, interferon α and
ribavirin.

In further embodiments;

The compound of formula (III) and the additionnal agent are administered sequentially.

The compound of formula (III) and the additionnal agent are administered simultaneously.

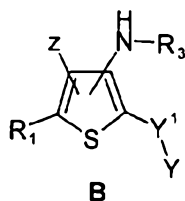
In still a further embodiment, the present invention provides_a process for preparing a compound of formula A:



10 said process comprising the steps of adding:

- an enol ether;
- an hydride donating agent; and
- an organic carboxylic acid;

15 to a compound of formula B:



wherein;

20 Y¹ is chosen from a bond, C₁₋₆ alkyl, C₂₋₆ alkenyl or C₂₋₆ alkynyl;

Y is chosen from COOR₁₆, COCOOR₅, P(O)OR_aOR_b, S(O)OR₅, S(O)₂OR₅, tetrazole, CON(R₉)CH(R₅)COOR₅, CONR₁₀R₁₁, CON(R₉)-SO₂-R₅, CONR₉OH or
 25 halogen, wherein R₉, R₅, R₁₀ and R₁₁ are each independently chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl;

or R₁₀ and R₁₁ are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_a and R_b are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

or R_a and R_b are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R_{1a} is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

R_1 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl or halogen;

R_2 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, or C_{6-18} aralkyl;

R_3 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

Z is chosen from H, halogen, C_{1-6} alkyl.

It will be appreciated by those skilled in the art that the compounds of formula (I) or (Ia) can contain a chiral centre on the general formula (I). The compounds of formula (I) or (Ia) thus exist in the form of two different optical isomers (i.e. (+) or (-) enantiomers). All such enantiomers and mixtures thereof including racemic mixtures are included within the scope of the invention. The single optical isomer or enantiomer can be obtained by method well known in the art, such as chiral HPLC, enzymatic resolution and chiral auxiliary.

In accordance with the present invention, the compounds of formula (I) or (Ia) include:

- | | |
|------------|---|
| Compound 1 | 3-[(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYL)-(3-iodo-benzyl)-amino]-5-phenyl-thiophene-2-carboxylic acid |
| Compound 2 | 3-[(3-benzofuran-2-yl-benzyl)-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYL)-amino]-5-phenyl-thiophene-2-carboxylic acid |
| Compound 3 | 3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-phenyl-thiophene-2-carboxylic acid |
| Compound 4 | 3-[(2,4-DICHLORO-BENZOYL)-[5-(3-TRIFLUOROMETHYL-PHENYL)-FURAN-2-ylmethyl]-amino]-5-phenyl-thiophene-2-carboxylic acid |

	Compound	5	3-[(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	6	5-(4-FLUORO-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	7	3-(2,4-DICHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	8	3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	9	3-[(2,4-DICHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	10	5-TERT-BUTYL-3-(4-CHLORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	11	4-(TOLUENE-4-SULFONYLAMINO)-[2,3']BITHIOPHENYL-5-CARBOXYLIC ACID
	Compound	12	3-[(5-BENZOFURAN-2-YL-THIOPHEN-2-YLMETHYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	13	5-PHENYL-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	14	3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-CHLORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	15	5-PHENYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	16	5-PHENYL-3-(TOLUENE-3-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	17	3-BENZENESULFONYLAMINO-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	18	3-(4-CHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	19	3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	20	5-TERT-BUTYL-3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	21	3-(2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	22	3-(4-METHOXY-2,3,6-TRIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	23	5-PHENYL-3-(THIOPHENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	24	4-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-[2,3']BITHIOPHENYL-5-CARBOXYLIC ACID
60	Compound	25	5-(3,5-BIS-TRIFLUOROMETHYL-PHENYL)-3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID

	Compound	26	8-CHLORO-3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-4H-1,5-DITHIA-CYCLOPENTA[A]NAPHTHALENE-2-CARBOXYLIC ACID
5	Compound	27	3-(2,4-DIFLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	28	3-[3-(2,6-DICHLORO-PYRIDIN-4-YL)-UREIDO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	29	3-(2-CHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	30	3-(2-FLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	31	5-PHENYL-3-(2-TRIFLUOROMETHOXY-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	32	3-(4-TERT-BUTYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	33	3-(4-CHLORO-PHENOXYCARBONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	34	3-(3,4-DICHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	35	5-PHENYL-3-(2-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	36	3-(5-BROMO-6-CHLORO-PYRIDINE-3-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	37	3-(5-CHLORO-THIOPHENE-2-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	38	3-(5-CHLORO-3-METHYL-BENZO[B]THIOPHENE-2-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	39	3-(4-BROMO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	40	3-(3-CHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	41	3-(5-CHLORO-1,3-DIMETHYL-1H-PYRAZOLE-4-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	42	3-(3-BROMO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	43	3-(4-ISOPROPYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	44	3-(2,6-DICHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	45	3-(2-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	46	5-PHENYL-3-(5-[1,2,3]THIADIAZOL-4-YL-THIOPHENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID

	Compound	47	5-PHENYL-3-(PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	48	3-(2,4-DICHLORO-BENZYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	49	3-(3-FLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	50	5-PHENYL-3-(3-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	51	3-(2-CARBOXY-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID METHYL ESTER
15	Compound	52	5-PHENYL-3-(4-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	53	3-(2,5-DIFLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	54	3-(2-CYANO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	55	3-(2,5-DICHLORO-THIOPHENE-3-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	56	4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
30	Compound	57	5'-CHLORO-4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
	Compound	58	5-(2,4-DICHLORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	59	5-(4-NITRO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	60	3-(TOLUENE-2-SULFONYLAMINO)-5-(4-TRIFLUOROMETHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	61	5-QUINOLIN-8-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	62	5-PHENYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	63	5-(3-NITRO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	64	3-(TOLUENE-2-SULFONYLAMINO)-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	65	5-(3-CHLORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	66	5-(4-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	67	5-(3-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID

	Compound	68	5-(4-CHLORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	69	5-(3,5-DIFLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	70	5-(3,4-DIFLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	71	3-(TOLUENE-2-SULFONYLAMINO)-5-VINYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	72	3-(4-CHLORO-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	73	3-[(4-CHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	74	5-PHENYL-3-[(THIOPHENE-2-CARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	75	3-[METHYL-(THIOPHENE-2-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	76	3-(2-BROMO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	77	3-(2,4-DIFLUORO-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	78	3-[(2,4-DIFLUORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	79	3-(TOLUENE-2-SULFONYLAMINO)-5-TRIMETHYLSILANYLETHYNYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	80	5-ETHYNYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	81	3-(TOLUENE-2-SULFONYLAMINO)-5-(3-TRIFLUOROMETHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	82	5-BENZOYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	83	5-(4-CYANO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	84	5-(3-CHLORO-4-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	85	5-(3,4-DICHLORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	86	5-PYRIDIN-4-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	87	5-PYRIDIN-3-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	88	3-(TOLUENE-2-SULFONYLAMINO)-5-(4-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID

	Compound	89	5-(4-METHANESULFONYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	90	5-(3-ACETYLAMINO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	91	5-(3-CHLORO-4-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	92	3-(4-METHYL-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	93	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	94	3-(3,5-DIMETHYL-ISOXAZOLE-4-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	95	3-[(2-CHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	96	3-(2-METHYL-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	97	3-[METHYL-(2-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	98	5-PHENYL-3-(5-TRIFLUOROMETHYL-PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	99	5-PHENYLETHYNYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	100	3-(2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	101	5-(2-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	102	5-(2-CYANO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	103	5-(2-ETHOXYCARBONYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	104	5-(2-METHOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	105	3'-METHYL-4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
50	Compound	106	3-(TOLUENE-2-SULFONYLAMINO)-5-(2-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	107	3-(2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	108	5-STYRYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	109	3-(2,4-DIFLUORO-BENZENESULFONYLAMINO)-5-(4-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	110	3-(2,4-DIFLUORO-BENZENESULFONYLAMINO)-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID

	Compound	111	3-([5-(3-CHLORO-4-FLUORO-PHENYL)-THIOPHEN-2-YLMETHYL]-(2,4-DICHLORO-BENZOYL)-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	112	3-([4-(4-OXO-1-PHENYL-1,3,8-TRIAZA-SPIRO[4.5]DECANE-8-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	113	3-([4-(2-OXO-2,3-DIHYDRO-BENZOIMIDAZOL-1-YL)-PIPERIDINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	114	3-([4-(4-NITRO-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	115	5-(2-CARBOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	116	5-(4-CHLORO-PHENYL)-3-(PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	117	5-(3-CYANO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	118	3-(2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-P-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	119	3-(2,4-DIFLUORO-BENZENESULFONYLAMINO)-5-P-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	120	5-PHENETHYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	121	5-(3-ETHOXYCARBONYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	122	5-(4-METHOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	123	5-(3-METHOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	124	5-(4'-BROMO-BIPHENYL-4-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	125	5-(4-HYDROXYMETHYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	126	5-FURAN-3-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	127	5-BENZOFURAN-2-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	128	5-PYRIDIN-2-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	129	5-(4-NITRO-PHENYL)-3-(PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	130	3-[(BENZOFURAN-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	131	3-[(2,4-DIMETHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	132	3-[[5-(2-CYANO-PHENYL)-THIOPHEN-2-YLMETHYL]-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	133	5-(4-FLUORO-PHENYL)-3-(PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	134	5-[2-(4-CHLORO-PHENYL)-VINYL]-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	135	3-BENZENESULFONYLAMINO-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	136	3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	137	5-PHENYL-3-(2-VINYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	138	3-(4-BROMO-2,5-DIFLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	139	3-(2-ACETYLAMINO-4-METHYL-THIAZOLE-5-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	140	3-(4-ACETYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	141	3-(4-FLUORO-2-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	142	3-(2-METHOXY-4-METHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	143	3-(3,4-DIFLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	144	4-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-5-(4-CHLORO-PHENYL)-2-METHYL-FURAN-3-CARBOXYLIC ACID ETHYL ESTER
40	Compound	145	3-(4-FLUORO-3-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	146	3-(2-AMINO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	147	3-(3-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	148	3-(4-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	149	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	150	5-(3-CYANO-BENZYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	151	5-PHENYL-3-(2,4,6-TRIFLUORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	152	3-(4-METHOXY-2-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	153	5-PHENYL-3-(2,3,4-TRICHLORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	154	5-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-2-METHYL-FURAN-3-CARBOXYLIC ACID METHYL ESTER
	Compound	155	4-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-2-METHYL-1,5-DIPHENYL-1H-PYRROLE-3-CARBOXYLIC ACID ETHYL ESTER
10	Compound	156	5-PHENYL-3-([4-(3-TRIFLUOROMETHYL-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	157	3-([4-(4-FLUORO-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	158	3-([4-(2,6-DIMETHYL-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	159	3-([4-(2-CHLORO-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	160	3-([4-(3-CHLORO-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	161	4,4'-BIS-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5,5'-DICARBOXYLIC ACID
	Compound	162	3-[ALLYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	163	5-(1-DIMETHYLSULFAMOYL-1H-PYRAZOL-4-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	164	5-(3-AMINO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	165	5-(4-AMINO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	166	5-(4-ACETYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	167	4-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-2,5-DIMETHYL-1H-PYRROLE-3-CARBOXYLIC ACID ETHYL ESTER
45	Compound	168	4-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-5-(4-CHLORO-PHENYL)-3-METHYL-1-PHENYL-1H-PYRROLE-2-CARBOXYLIC ACID ETHYL ESTER
50	Compound	169	3-(3,5-DICHLORO-4-HYDROXY-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	170	5-(1H-PYRAZOL-4-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	171	5-(3-HYDROXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	172	3-[METHYL-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	173	3-([2-(4-FLUORO-PHENYL)-ACETYL]-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	174	3-(4-PENTYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	175	3-(METHYL-PHENYLACETYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	176	3-[2,5-BIS-(2,2,2-TRIFLUORO-ETHOXY)-BENZENESULFONYLAMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	177	3-(4-METHYL-2-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	178	5-THIAZOL-2-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	179	5-PHENYL-3-[3-(3-PHENYL-PROPYL)-UREIDO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	180	3-[(3,4-DIHYDRO-1H-ISOQUINOLINE-2-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	181	3-([4-(4-METHOXY-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	182	3-([4-(6-METHYL-PYRIDIN-2-YL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID HYDROCHLORIDE
30	Compound	183	3-([4-(4-CHLORO-BENZYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID HYDROCHLORIDE
35	Compound	184	5-(5-METHYL-PYRIDIN-2-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	185	3-[ETHYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	186	3-[(3-CHLORO-THIOPHENE-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	187	3-[(2-BROMO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	188	3-[(4-BUTYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	189	3-(2-CHLOROMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	190	5-(4-HYDROXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	191	5-(5-CHLORO-PYRIDIN-2-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	192	5-(4-CHLORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	193	5-(4-CYANO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID

	Compound	194	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(4-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	195	5-(4-HYDROXYMETHYL-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	196	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(3-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	197	5-(4-FLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	198	5-(4-METHOXY-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	199	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-P-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	200	5-(4-AMINO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	201	3-[CYCLOPENTYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	202	5-BENZO[1,3]DIOXOL-5-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	203	3-[(2-HYDROXY-ETHYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	204	3-[(2,4-DICHLORO-BENZOYL)-ISOBUTYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	205	3-[(2-METHOXY-4-METHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	206	5-(3-CYANO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	207	5-(2-CHLORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	208	3-[(2,4-DICHLORO-BENZOYL)-PHENYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	209	3-[4-(TRIFLUOROMETHYL-BENZOYL)METHYLAMINE]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	210	3-[(4-CHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	211	3-[ISOPROPYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	212	5-(3,5-DIFLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	213	5-(3-FLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	214	5-(2,4-DIFLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID

	Compound	215	5-(4-HYDROXY-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	216	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(4-TRIFLUOROMETHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	217	5-(2-HYDROXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	218	3-[(2-CHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	219	3-[(3,5-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	220	3-(4-BROMO-2-METHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	221	3-(5-CARBOXY-4-CHLORO-2-FLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	222	5-PHENYL-3-(2,3,4-TRIFLUORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	223	3-(4-BROMO-2-FLUORO-BENZENESULFONYLAMINO)-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	224	3-(4-BROMO-2-METHYL-BENZENESULFONYLAMINO)-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	225	5-(4-ISOBUTYL-PHENYL)-3-(3-METHOXY-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	226	3-[(4-FLUORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	227	3-[2,5-BIS-(2,2,2-TRIFLUORO-ETHOXY)-BENZENESULFONYLAMINO]-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	228	3-(2-CHLORO-4-CYANO-BENZENESULFONYLAMINO)-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	229	5'-ACETYL-4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
45	Compound	230	5-BENZO[B]THIOPHEN-2-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	231	5-(4-BUTYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	232	5-(4-ETHYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	233	3-[BENZYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	234	3-[(4-CHLORO-2-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	235	3-[(2,4-DIMETHYL-BENZENESULFONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	236	5-(4-ACETYL-PHENYL)-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	237	5-(4-ACETYL-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	238	5-(4-ACETYL-PHENYL)-3-(4-CHLORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	239	5-(4-CARBOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID TERT-BUTYL ESTER
	Compound	240	3-[(2,4-DIMETHYL-BENZENESULFONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	241	3-[ACETYL-(4-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	242	3-ETHANESULFONYLAMINO-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	243	3-[ISOPROPYL-(4-TRIFLUOROMETHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	244	3-[(2,4-DICHLORO-BENZOYL)-(3-METHYL-BUT-2-ENYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	245	3-[(2,6-DICHLORO-PYRIDINE-3-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	246	3-[(6-CHLORO-PYRIDINE-3-CARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	247	3-[(4-TERT-BUTYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	248	5-(4-CARBOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	249	5-(4-ETHOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	250	3-[(2,6-DICHLORO-PYRIDINE-3-CARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	251	3-[(BENZO[B]THIOPHENE-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	252	3-[METHYL-(NAPHTHALENE-2-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	253	3-[(3,4-DICHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	254	3-[(3,5-DICHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	255	3-[(4-BROMO-3-METHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	256	3-[(3-CHLORO-BENZO[B]THIOPHENE-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	256	3-[(3-CHLORO-BENZO[B]THIOPHENE-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	257	3-[METHYL-(4-METHYL-3-NITRO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	258	5-(4-CARBAMOYL-PHENYL)-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	259	5-(4-CARBAMOYL-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	260	5-(1H-INDOL-5-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	261	3-[SEC-BUTYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	262	3-[(2,4-DIMETHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	263	5-(4-AZIDO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	264	3-[(2,4-DICHLORO-BENZOYL)-(1-PHENYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	265	5-(4-CARBAMOYL-PHENYL)-3-(4-CHLORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	266	5-(2-FLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	267	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-O-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	268	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	269	5-(3-CHLORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	270	5-(3,4-DIFLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	271	5-(3-AMINO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	272	5-(3-ACETYL-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	273	5-(3-HYDROXY-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	274	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	275	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(4-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	276	3-[(3,4-DIMETHOXY-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	277	3-[METHYL-(2,4,6-TRIFLUORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	278	3-[(2,3-DIFLUORO-4-TRIFLUOROMETHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	279	3-[(3-FLUORO-4-TRIFLUOROMETHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	280	3-[(2,3-DIFLUORO-4-METHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	281	3-[(2-FLUORO-4-TRIFLUOROMETHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	282	5-(4-CARBAMOYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	283	5-(4-FLUORO-PHENYL)-3-[ISOPROPYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	284	3-[(2-BROMO-4-CHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	285	3-(2,6-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	286	3-[METHYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	287	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID METHYL ESTER
30	Compound	288	5-(4-CYANO-PHENYL)-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	289	3-(4-CHLORO-BENZENESULFONYLAMINO)-5-(4-CYANO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	290	5-(4-CYANO-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	291	5'-ACETYL-4-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
40	Compound	292	5'-ACETYL-4-(2,6-DIMETHYL-BENZENESULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
45	Compound	293	3-[METHYL-(4-METHYL-THIOPHENE-2-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	294	5-(3-CHLORO-PHENYL)-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	295	5'-CYANO-4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
	Compound	296	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-PYRIDIN-2-YL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	297	3-[(2,4-DICHLORO-THIOBENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	298	5-PHENYL-3-(2,4,6-TRIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID

	Compound	299	3-[(1-CARBOXY-ETHYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	300	3-[(4-METHYL-BENZOYL)-(3-METHYL-BUT-2-ENYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	301	3-[(2-HYDROXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	302	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-PYRIDIN-3-YL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	303	5'-ACETYL-4-[METHYL-(4-METHYL-BENZOYL)-AMINO]-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
15	Compound	304	3-[ISOPROPYL-(4-METHYL-BENZOYL)-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	305	3-[ISOPROPYL-(4-METHYL-BENZOYL)-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	306	3-[(2-BROMO-4-CHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	307	3-[(4-CHLORO-2-FLUORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	308	3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-4-METHYL-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	309	3-[(2-BROMO-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	310	3-[(4-CHLORO-2-iodo-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	311	3-[(4-CYANO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	312	3-[ALLYL-(4-METHYL-BENZOYL)-AMINO]-5-[4-(2-CARBOXY-VINYL)-PHENYL]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	313	3-[(4-CHLORO-2-HYDROXY-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	314	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-4-METHYL-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	315	5-TERT-BUTYL-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	316	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	317	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	318	5-[4-(2-CARBOXY-ETHYL)-PHENYL]-3-[(4-METHYL-BENZOYL)-PROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	319	5-BENZOFURAN-2-YL-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID

	Compound	320	3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-HYDROXYMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	321	3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-METHANESULFONYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	322	5-[4-(2-CARBOXY-VINYL)-PHENYL]-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	323	3-[ALLYL-(4-METHYL-BENZOYL)-AMINO]-5-[3-(2-CARBOXY-VINYL)-PHENYL]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	324	3-[ISOPROPYL-(2,4,6-TRIMETHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	325	5-[3-(2-CARBOXY-ETHYL)-PHENYL]-3-[(4-METHYL-BENZOYL)-PROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	326	3-[(2-FLUORO-4-TRIFLUOROMETHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	327	3-[TERT-BUTYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	328	3-[(2-AMINO-4-CHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	329	3-[(4-CHLORO-2-NITRO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	330	3-[(4-METHYL-BENZOYL)-(3-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	331	3-[(3-FLUORO-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	332	5-(4-CARBOXY-PHENYL)-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	333	3-[CYCLOPROPYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	334	3-[(3-TERT-BUTYL-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	335	3-[(3-CHLORO-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	336	3-[(2,4-DIFLUORO-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	337	3-[(4-CHLORO-2,5-DIFLUORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	338	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(2-METHYL-ALLYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	339	3-(ALLYL-[2-(4-CHLORO-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	340	3-[BENZYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	341	3-[(4-CHLORO-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	342	3-[(4-METHYL-BENZOYL)-(4-NITRO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	343	3-[(4-METHYL-BENZOYL)-(2-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	344	3-[(3-METHOXY-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	345	3-[(2-CHLORO-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	346	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-ISOBUTYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	347	3-[ALLYL-(2-NAPHTHALEN-2-YL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	348	3-(ALLYL-[2-(2,4-DICHLORO-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	349	3-(ALLYL-[2-(2-CHLORO-4-FLUORO-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	350	3-(ALLYL-[2-(3,4-DICHLORO-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	351	3-(ALLYL-[2-(2,4-DIFLUORO-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	352	3-(ALLYL-[2-(4-TRIFLUOROMETHYL-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	353	3-(ALLYL-[2-(2,6-DICHLORO-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	354	3-[ALLYL-(2-M-TOLYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	355	5-(4-ACETYL-PHENYL)-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	356	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	357	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	358	5'-ACETYL-4-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
55	Compound	359	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	360	4-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5'-METHYL-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
60	Compound	361	3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-METHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID

	Compound	362	3- (CYCLOHEXANECARBONYL-ISOPROPYL-AMINO) -5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	363	3- { (2,4-DICHLORO-BENZOYL) - [1- (2,4-DICHLORO-BENZOYL) -PIPERIDIN-4-YL] -AMINO} -5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	364	4- [(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL) - (4-METHYL-BENZOYL) -AMINO] -PIPERIDINE-1-CARBOXYLIC ACID TERT -BUTYL ESTER
10	Compound	365	4- [(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL) - (2,4-DICHLORO-BENZOYL) -AMINO] -PIPERIDINE-1-CARBOXYLIC ACID TERT -BUTYL ESTER
	Compound	366	3- [(4-METHYL-BENZOYL) -PIPERIDIN-4-YL-AMINO] -5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	367	5'-ACETYL-4- (2,4-DIMETHYL-BENZENESULFONYLAMINO) - [2,3'] BITHIOPHENYL-5-CARBOXYLIC ACID
20	Compound	368	3- [(2,4-DICHLORO-BENZOYL) -PIPERIDIN-4-YL-AMINO] -5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	369	5- (4-METHANESULFONYLAMINO-PHENYL) -3- (TOLUENE-2-SULFONYLAMINO) -THIOPHENE-2-CARBOXYLIC ACID
25	Compound	370	3- (4-FLUORO-2-METHYL-BENZENESULFONYLAMINO) -5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	371	3- [(3-METHYL-CYCLOHEXANECARBONYL) -ISOPROPYL-AMINO] -5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	372	3- (4-CHLORO-2-METHYL-BENZENESULFONYLAMINO) -5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	373	3- [(2,4-DICHLORO-BENZOYL) -ISOPROPYL-AMINO] -5- (4-METHANESULFONYL-PHENYL) -THIOPHENE-2-CARBOXYLIC ACID
35	Compound	374	3- [(2,4-DICHLORO-BENZOYL) -ISOPROPYL-AMINO] -5- (4-METHANESULFINYL-PHENYL) -THIOPHENE-2-CARBOXYLIC ACID
40	Compound	375	5- (4-CARBOXY-PHENYL) -3- [(2,4-DICHLORO-BENZOYL) -ISOPROPYL-AMINO] -THIOPHENE-2-CARBOXYLIC ACID
	Compound	376	5-BENZOFURAN-2-YL-3- [(2,4-DICHLORO-BENZOYL) -ISOPROPYL-AMINO] -THIOPHENE-2-CARBOXYLIC ACID
45	Compound	377	3- [(2-ACETOXY-4-METHYL-BENZOYL) -ISOPROPYL-AMINO] -5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	378	3- [ISOPROPYL- (2-METHYL-CYCLOHEXANECARBONYL) -AMINO] -5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	379	3- [ISOPROPYL- (2-METHYL-CYCLOHEXANECARBONYL) -AMINO] -5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	380	3- (CYCLOHEPTANECARBONYL-ISOPROPYL-AMINO) -5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	381	3- [ISOPROPYL- (4-METHYL-CYCLOHEXANECARBONYL) -AMINO] -5- (3-TRIFLUOROMETHYL-PHENYL) -THIOPHENE-2-CARBOXYLIC ACID
60	Compound	382	3- [(2,4-DICHLORO-BENZOYL) -ISOPROPYL-AMINO] -5-METHYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	383	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-(3-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	384	3-[(3-CYCLOPENTYL-PROPIONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	385	3-(BUTYRYL-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	386	3-(METHYL-PENT-4-ENOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	387	3-[ISOPROPYL-(5-METHYL-3-OXO-3H-ISOINDOL-1-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	388	3-[METHYL-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	389	3-(METHYL-PENTANOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	390	3-[METHYL-(4-METHYL-PENTANOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	391	3-(CYCLOPENTANECARBONYL-ETHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	392	3-[(3-CYCLOPENTYL-PROPIONYL)-ETHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	393	3-(CYCLOBUTANECARBONYL-ETHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	394	3-(BUT-2-ENOYL-ETHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	395	3-[ISOPROPYL-(4-METHYL-2-VINYLBENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	396	3-[ISOPROPYL-(4-METHYL-CYCLOHEX-1-ENECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	397	3-(ALLYL-HEXANOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	398	3-(ALLYL-CYCLOBUTANECARBONYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	399	3-(ALLYL-PENTANOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	400	3-[ALLYL-(4-METHYL-PENTANOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	401	3-[ALLYL-(2-CYCLOPENTYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	402	3-[(2-HYDROXY-4-METHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	403	3-[(2,4-DICHLORO-BENZOYL)-(1-PHENYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	404	3-[(2,4-DICHLORO-BENZOYL)-(1-PHENYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	405	3-[ISOPROPYL-(3-METHYL-CYCLOPENT-3-ENECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	406	3-[(2-BENZYLOXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	407	3-[(2,4-DIMETHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	408	3-[ISOPROPYL-(3-METHYL-CYCLOPENTANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	409	3-[(2-HYDROXY-4-METHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	410	5-PHENYL-3-[PROPIONYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	411	3-[ISOBUTYRYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	412	3-[(3-METHYL-BUTYRYL)-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	413	3-[CYCLOPROPANECARBONYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	414	3-[CYCLOBUTANECARBONYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	415	3-[BUTYRYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	416	3-[(2-CYCLOPENTYL-ACETYL)-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	417	3-[(4-TERT-BUTYL-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	418	3-[(4-NITRO-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	419	3-[(3-METHYL-BUTYRYL)-(4-NITRO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	420	3-[CYCLOPROPANECARBONYL-(4-NITRO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	421	3-[(2-CHLORO-BENZYL)-ISOBUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	422	3-[(2-CHLORO-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	423	3-[(2-CHLORO-BENZYL)-CYCLOPROPANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	424	3-[(ADAMANTANE-1-CARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	425	3-[(2-CHLORO-BENZYL)-CYCLOBUTANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	426	3-[ACETYL-(2-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	427	3-[(2-METHYL-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	428	3-[(2-HYDROXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	429	3-[(1-ACETYL-PIPERIDIN-4-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	430	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-[4-(1 H -TETRAZOL-5-YL)-PHENYL]-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	431	3-[(2-CYANO-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	432	3-[CYCLOBUTANECARBONYL-(2-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	433	3-[BUTYRYL-(2-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	434	3-[ACETYL-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	435	3-[CYCLOBUTANECARBONYL-(4-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	436	3-[CYCLOHEXANECARBONYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	437	3-[(4-TERT-BUTYL-BENZYL)-ISOBUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	438	3-[(4-TERT-BUTYL-BENZYL)-CYCLOPROPANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	439	3-[(4-TERT-BUTYL-BENZYL)-CYCLOBUTANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	440	3-[(4-TERT-BUTYL-BENZYL)-BUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	441	3-[(4-TERT-BUTYL-BENZYL)-CYCLOHEXANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	442	3-[(4-TERT-BUTYL-BENZYL)-(2-CYCLOPENTYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	443	3-[(2-CYCLOPENTYL-ACETYL)-(4-NITRO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	444	3-[(2-CHLORO-BENZYL)-CYCLOHEXANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	445	3-[(2-CYCLOPENTYL-ACETYL)-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	446	3-[BUTYRYL-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	447	3-[BUTYRYL-(2-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	448	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	449	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-THIAZOL-2-YL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	450	3-(ACETYL-BENZYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	451	3-(BENZYL-PROPIONYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	452	3-[BENZYL-(2-METHOXY-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	453	3-[BENZYL-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	454	3-(BENZYL-CYCLOPROPANECARBONYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	455	3-[ACETYL-(4-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	456	3-[(4-CHLORO-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	457	3-[(4-CHLORO-BENZYL)-ISOBUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	458	3-[(4-CHLORO-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	459	3-[(4-CHLORO-BENZYL)-CYCLOPROPANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	460	5-(4-ACETYL-PHENYL)-3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	461	3-[(4-CHLORO-BENZYL)-CYCLOBUTANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	462	3-[BUTYRYL-(4-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	463	3-[(4-CHLORO-BENZYL)-(2-CYCLOPENTYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	464	3-[ACETYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	465	3-[ISOBUTYRYL-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	466	3-[CYCLOPROPANECARBONYL-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	467	3-[(4-METHYL-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	468	3-[ISOBUTYRYL-(4-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	469	3-[CYCLOPROPANECARBONYL-(4-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	470	3-[BUTYRYL-(4-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	471	3-[(3-METHOXY-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	472	3-[(3-METHOXY-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	473	3-[CYCLOBUTANECARBONYL-(3-METHOXY-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	474	3-[(2-CARBAMOYL-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	475	3-[BUTYRYL-(3-METHOXY-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	476	3-[ACETYL-(3-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	477	3-[(3-CHLORO-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	478	3-[(3-CHLORO-BENZYL)-(2-METHOXY-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	479	3-[(3-CHLORO-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	480	3-[(3-CHLORO-BENZYL)-CYCLOPROPANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	481	3-[(3-CHLORO-BENZYL)-CYCLOBUTANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	482	3-[BUTYRYL-(3-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	483	3-[ACETYL-(2,4-DIFLUORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	484	3-[(2,4-DIFLUORO-BENZYL)-(2-METHOXY-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	485	3-[(2,4-DIFLUORO-BENZYL)-ISOBUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	486	3-[(2,4-DIFLUORO-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	487	3-[BENZYL-(2-CYCLOPENTYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	488	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(1H-INDOL-5-YL)-THIOPHENE-2-CARBOXYLIC ACID

	Compound	489	3-(BENZYL-CYCLOBUTANECARBONYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	490	3-[CYCLOHEXANECARBONYL-(2,4-DIFLUORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	491	3-(ALLYL-[2-(4-METHOXY-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	492	3-(ETHYL-HEXANOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	493	3-(BUTYRYL-ETHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	494	3-[ETHYL-(4-METHYL-PENTANOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	495	3-[CYCLOBUTANECARBONYL-(2,4-DIFLUORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	496	3-[BUTYRYL-(2,4-DIFLUORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	497	3-(CYCLOPENTANECARBONYL-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	498	3-(CYCLOHEXANECARBONYL-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	499	3-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-PYRROLIDINE-1-CARBOXYLIC ACID TERT-BUTYL ESTER
	Compound	500	3-[(1,4-DIMETHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	501	5-(4-ETHYL-PHENYL)-3-[(2-HYDROXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	502	3-[(2-HYDROXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	503	3-[(2,4-DICHLORO-BENZOYL)-PYRROLIDIN-3-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	504	4-{5-CARBOXY-4-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHEN-2-YL}-3,6-DIHYDRO-2H-PYRIDINE-1-CARBOXYLIC ACID BENZYL ESTER
	Compound	505	3-[(2-(HYDROXYIMINO-METHYL)-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
50	Compound	506	3-[(1-CARBAMIMIDOYL-PIPERIDIN-4-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	507	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-AZEPANE-1-CARBOXYLIC ACID TERT-BUTYL ESTER
	Compound	508	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-METHYL-PIPERIDINE-1-CARBOXYLIC ACID BENZYL ESTER
60	Compound	509	3-[AZEPAN-4-YL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	510	3-[(4-METHYL-CYCLOHEXANECARBONYL)-PIPERIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID LITHIUM SALT
5	Compound	511	3-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-PIPERIDINE-1-CARBOXYLIC ACID TERT-BUTYL ESTER
	Compound	512	3-[(4-BENZYLOXYCARBONYLAMINO-CYCLOHEXYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	513	3-[ISOPROPYL-(4-METHYL-2-OXO-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	514	3-[(2,4-DICHLORO-BENZOYL)-PIPERIDIN-3-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID; COMPOUND WITH GENERIC INORGANIC NEUTRAL COMPONENT
	Compound	515	3-[(4-BENZYLOXYCARBONYLAMINO-CYCLOHEXYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	516	3-[(2-BENZYLOXY-1-METHYL-ETHYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	517	3-[(2,2-DIMETHYL-[1,3]DIOXAN-5-YL)-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	518	3-[(2,4-DICHLORO-BENZOYL)-(2-HYDROXY-1-HYDROXYMETHYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	519	3-[(2,4-DICHLORO-BENZOYL)-PIPERIDIN-4-YLMETHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	520	3-[(2-CHLORO-BENZOYL)-PIPERIDIN-4-YLMETHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	521	3-[(4,6-DICHLORO-1H-INDOLE-2-CARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	522	3-[(2,4-DICHLORO-BENZOYL)-(2-HYDROXY-1-METHYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	523	4-{1-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-ETHYL}-PIPERIDINE-1-CARBOXYLIC ACID BENZYL ESTER
45	Compound	524	4-{5-CARBOXY-4-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHEN-2-YL}-3,6-DIHYDRO-2 H -PYRIDINE-1-CARBOXYLIC ACID BENZYL ESTER
50	Compound	525	3-[(4-METHYL-CYCLOHEXANECARBONYL)-PYRIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	526	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PIPERIDIN-4-YL-THIOPHENE-2-CARBOXYLIC ACID; COMPOUND WITH TRIFLUORO-ACETIC ACID
55	Compound	527	3-[ISOPROPYL-(4-PROPYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	528	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-CYCLOHEXYL-AMMONIUM; TRIFLUORO-ACETATE

	Compound	529	3-[(2,4-DICHLORO-BENZOYL)-(1-PIPERIDIN-4-YL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID; COMPOUND WITH TRIFLUORO-ACETIC ACID
5	Compound	530	3-[(CYCLOHEX-3-ENECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	531	3-[(4-ETHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	532	3-[(4-CHLORO-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	533	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-3-METHYL-PIPERIDINE-1-CARBOXYLIC ACID BENZYL ESTER
15	Compound	534	3-[(2,4-DICHLORO-BENZOYL)-(2-METHOXY-CYCLOHEXYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	535	3-[(2,4-DICHLORO-BENZOYL)-(2,2-DIMETHYL-[1,3]DIOXAN-5-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	536	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-(1-METHYL-PIPERIDIN-4-YL)-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	537	3-[(2,4-DICHLORO-BENZOYL)-(3-METHYL-PIPERIDIN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID; COMPOUND WITH TRIFLUORO-ACETIC ACID
	Compound	538	3-[(2,4-DICHLORO-BENZOYL)-(2-HYDROXY-CYCLOHEXYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	539	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(4-METHYLCYCLOHEXANE CARBONYL)-AMINO]-METHYL)-PIPERIDINE-1-CARBOXYLIC ACID BENZYL ESTER
35	Compound	540	3-[(1R,2S,4R)-2-HYDROXY-4-METHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	541	3-[ISOPROPYL-[1-(4-METHOXY-2,3,6-TRIMETHYL-BENZENESULFONYL)-5-METHYL-1,2,3,6-TETRAHYDRO-PYRIDINE-2-CARBONYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	542	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-4-FLUORO-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	543	3-[(2,4-DICHLORO-BENZOYL)-(1-METHYL-PIPERIDIN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	544	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(4-METHYLCYCLOHEXANE CARBONYL)-AMINO]-METHYL)-PIPERIDINIUM; TRIFLUORO-ACETATE
50	Compound	545	3-[(2-TERT-BUTOXYCARBONYLAMINO-1-METHYL-ETHYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
55	Compound	546	2-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-PROPYL-AMINE TRIFLUOROACETIC ACID SALT
	Compound	547	3-[(3-CARBOXY-CYCLOPENTYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	548	3-[(3-CARBOXY-CYCLOPENTYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

	Compound	549	2-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-CYCLOHEXYL-AMMONIUM CHLORIDE
5	Compound	550	3-(BENZOYL-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	551	{[5-PHENYL-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBONYL]-AMINO}-ACETIC ACID
10	Compound	552	5-BROMO-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	553	3-[CYCLOHEXYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	554	3-[[1,3]DIOXAN-5-YL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	555	3-[[2-(TERT-BUTYL-DIMETHYL-SILANYLOXY)-1-METHYL-2-PHENYL-ETHYL]-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	556	3-[[2-(TERT-BUTYL-DIMETHYL-SILANYLOXY)-1-METHYL-2-PHENYL-ETHYL]-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	557	3-[(2,4-DICHLORO-BENZOYL)-(2-DIETHYLAMINO-THIAZOL-5-YLMETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	558	(5-({(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO}-METHYL)-THIAZOL-2-YL)-DIETHYL-AMMONIUM; CHLORIDE
35	Compound	559	5-(4-FLUORO-PHENYL)-3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
	Compound	560	3-[(1S,2R,4S)-2-HYDROXY-4-METHYL-CYCLOHEXANECARBONYL]-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	561	3-[(2,4-DICHLORO-BENZOYL)-(2-METHOXY-1-METHYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	562	3-[(4S)-ISOPROPYL-(4-METHYL-CYCLOHEX-1-ENECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
45	Compound	566	3-METHYL-(4-METHYLBENZOYL)-AMINO)-5-PHENYL THIOPHENE-2-CARBOXYLIC ACID (2-HYDROXY-ETHYL) AMIDE
50	Compound	567	5-PHENYL-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID CYCLOBUTYLAMIDE
	Compound	568	3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID AMIDE
55	Compound	569	5-BROMO-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
60	Compound	570	5-(4-CHLORO-PHENYL)-3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID

	Compound	571	5-(4'-CHLORO-BIPHENYL-4-YL)-3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
5	Compound	572	3-[(4-METHYL-CYCLOHEXANECARBONYL)-(TETRAHYDRO-PYRAN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	573	3-[(4-METHYL-CYCLOHEXANECARBONYL)-(1-METHYL-PIPERIDIN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
10	Compound	574	3-[(4-METHYL-CYCLOHEXANECARBONYL)-PIPERIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	575	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-(4-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
15	Compound	576	5-(4-CYANO-PHENYL)-3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
20	Compound	577	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-(4-METHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	578	3-[(2-METHOXY-1-METHYL-ETHYL)-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
25	Compound	579	3-[CYCLOHEXYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
	Compound	581	5-(4-ISOBUTYL-PHENYL)-3-[5-(5-TRIFLUOROMETHYL-ISOXAZOL-3-YL)-THIOPHENE-2-SULFONYLAMINO]-THIOPHENE-2-CARBOXYLIC ACID
30	Compound	582	5-(4-ISOBUTYL-PHENYL)-3-(2,3,4-TRIFLUORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
	Compound	583	3-[(2,4-DICHLORO-PHENYL)-ISOPROPYL-CARBAMOYL]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
35	Compound	584	3-(METHYL-P-TOLYL-CARBAMOYL)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
40	Compound	585	3-[(2,4-DICHLORO-PHENYL)-METHYL-CARBAMOYL]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

or pharmaceutically acceptable salts thereof.

Preferably, the compounds of the present invention are provided in the form of a single enantiomer at least 95%, more preferably at least 97% and most preferably at least 99% free of the corresponding enantiomer.

More preferably the compound of the present invention are in the form of the (+) enantiomer at least 95% free of the corresponding (-) enantiomer.

More preferably the compound of the present invention are in the form of the (+) enantiomer at least 97% free of the corresponding (-) enantiomer.

5 More preferably the compound of the present invention are in the form of the (+) enantiomer at least 99% free of the corresponding (-) enantiomer.

10 In a more preferred embodiment, the compound of the present invention are in the form of the (-) enantiomer at least 95% free of the corresponding (+) enantiomer.

Most preferably the compound of the present invention are in the form of the (-) enantiomer at least 97% free of the corresponding (+) enantiomer.

15 More preferably the compound of the present invention are in the form of the (-) enantiomer at least 99% free of the corresponding (+) enantiomer.

20 There is also provided a pharmaceutically acceptable salts of the present invention. By the term pharmaceutically acceptable salts of compounds of general formula (I) or (Ia) are meant those derived from pharmaceutically acceptable inorganic and organic acids and bases. Examples of suitable acids include

25 hydrochloric, hydrobromic, sulphuric, nitric, perchloric, fumaric, maleic, phosphoric, glycollic, lactic, salicylic, succinic, toleune-p-sulphonic, tartaric, acetic, trifluoroacetic, citric, methanesulphonic, formic, benzoic, malonic, naphthalene-2-sulphonic and benzenesulphonic acids.

30 Other acids such as oxalic, while not in themselves pharmaceutically acceptable, may be useful as intermediates in obtaining the compounds of the invention and their pharmaceutically acceptable acid addition salts.

35 Salts derived from appropriate bases include alkali metal (e.g. sodium), alkaline earth metal (e.g. magnesium), ammonium and NR_4^+ (where R is C_{1-4} alkyl) salts.

References hereinafter to a compound according to the invention includes compounds of the general formula (I) or (Ia) and their pharmaceutically acceptable salts.

5 Unless otherwise defined, all technical and scientific terms used herein have the same meaning as commonly understood by one of ordinary skill in the art to which this invention belongs. All publications, patent applications, patents, and other references mentioned herein are incorporated by reference in
10 their entirety. In case of conflict, the present specification, including definitions, will control. In addition, the materials, methods, and examples are illustrative only and not intended to be limiting.

15 As used in this application, the term "alkyl" represents a straight chain, branched chain or cyclic hydrocarbon moiety which may optionally be substituted by one or more of: halogen, nitro, nitroso, SO₃R₁₂, PO₃R_cR_d, CONR₁₃R₁₄, C₁-6 alkyl, C₂-6 alkenyl, C₂-6 alkynyl, C₆-12 aralkyl, C₆-12 aryl, C₁-6 alkyloxy, C₂-6
20 alkenyloxy, C₂-6 alkynyloxy, C₆-12 aryloxy, C(O)C₁-6 alkyl, C(O)C₂-6 alkenyl, C(O)C₂-6 alkynyl, C(O)C₆-12 aryl, C(O)C₆-12 aralkyl, C₃-10 heterocycle, hydroxyl, NR₁₃R₁₄, C(O)OR₁₂, cyano, azido, amidino or guanido;

25 wherein R₁₂, R_c, R_d, R₁₃ and R₁₄ are each independently chosen from H, C₁-12 alkyl, C₂-12 alkenyl, C₂-12 alkynyl, C₆-14 aryl, C₃-12 heterocycle, C₃-18 heteroaralkyl, C₆-18 aralkyl; or R_c and R_d are taken together with the oxygens to form a 5 to
10 membered heterocycle;

30 or R₁₃ and R₁₄ are taken together with the nitrogen to form a 3 to 10 membered heterocycle. Useful examples of alkyls include isopropyl, ethyl, fluorohexyl or cyclopropyl. The term alkyl is also meant to include alkyls in which one or more hydrogen atoms
35 is replaced by an oxygen, (e.g. a benzoyl) or an halogen, more preferably, the halogen is fluoro (e.g. CF₃- or CF₃CH₂-).

The terms "alkenyl" and "alkynyl" represent an alkyl containing at least one unsaturated group (e.g. allyl, acetylene,
40 ethylene).

The term "aryl" represents a carbocyclic moiety containing at least one benzenoid-type ring which may optionally be substituted by one or more of halogen, nitro, nitroso, S03R12, P03RcRd, CONR13R14, C1-6 alkyl, C2-6 alkenyl, C2-6 alkynyl, C6-12 aralkyl, C6-12 aryl, C1-6 alkyloxy, C2-6 alkenyloxy, C2-6 alkynyloxy, C6-12 aryloxy, C(O)C1-6 alkyl, C(O)C2-6 alkenyl, C(O)C2-6 alkynyl, C(O)C6-12 aryl, C(O)C6-12 aralkyl, C3-10 heterocycle, hydroxyl, NR13R14, C(O)OR12, cyano, azido, amidino or guanido;

wherein R12, Rc, Rd, R13 and R14 are each independently chosen from H, C1-12 alkyl, C2-12 alkenyl, C2-12 alkynyl, C6-14 aryl, C3-12 heterocycle, C3-18 heteroaralkyl, C6-18 aralkyl;

or Rc and Rd are taken together with the oxygens to form a 5 to 10 membered heterocycle;

or R13 and R14 are taken together with the nitrogen to form a 3 to 10 membered heterocycle. Examples of aryl include phenyl and naphthyl.

The term "aralkyl" represents an aryl group attached to the adjacent atom by a C1-6alkyl, C1-6alkenyl, or C1-6alkynyl (e.g., benzyl).

The term "heterocycle" represents a saturated or unsaturated, cyclic moiety wherein said cyclic moiety is interrupted by at least one heteroatom, (e.g. oxygen, sulfur or nitrogen) which may optionally be substituted halogen, nitro, nitroso, S03R12, P03RcRd, CONR13R14, C1-6 alkyl, C2-6 alkenyl, C2-6 alkynyl, C6-12 aralkyl, C6-12 aryl, C1-6 alkyloxy, C2-6 alkenyloxy, C2-6 alkynyloxy, C6-12 aryloxy, C(O)C1-6 alkyl, C(O)C2-6 alkenyl, C(O)C2-6 alkynyl, C(O)C6-12 aryl, C(O)C6-12 aralkyl, C3-10 heterocycle, hydroxyl, NR13R14, C(O)OR12, cyano, azido, amidino or guanido;

wherein R12, Rc, Rd, R13 and R14 are each independently chosen from H, C1-12 alkyl, C2-12 alkenyl, C2-12 alkynyl, C6-14 aryl, C3-12 heterocycle, C3-18 heteroaralkyl, C6-18 aralkyl; or Rc and Rd are taken together with the oxygens to form a 5 to 10 membered heterocycle;

or R13 and R14 are taken together with the nitrogen to form a 3 to 10 membered heterocycle. It is understood that the term heterocyclic ring represents a mono or polycyclic (e.g., bicyclic) ring. Examples of heterocyclic rings include but are not limited to epoxide; furan; benzofuran; isobenzofuran; oxathiolane; dithiolane; dioxolane; pyrrole; pyrrolidine; imidazole; pyridine; pyrimidine; indole; piperidine; morpholine; thiophene and thiomorpholine.

- 10 The term "heteroaralkyl" represents an heterocycle group attached to the adjacent atom by a C₁₋₆ alkyl, C₁₋₆ alkenyl, or C₁₋₆ alkynyl.

When there is a sulfur atom present, the sulfur atom can be at different oxidation levels, ie. S, SO, or SO₂. All such oxidation levels are within the scope of the present invention.

The term "independently" means that a substituent can be the same or different definition for each item.

20

As used in this application, the term "hydride donating agent" means a suitable ionic or covalent inorganic compound of hydrogen with another element (e.g. boron, sodium, lithium or aluminum) allowing the process to occur under the reaction conditions without causing adverse effect on the reagents or product. Useful examples of hydride donating agent include but are not limited to sodium borohydride (NaBH₄), sodium cyanoborohydride (NaCNBH₃), sodium triacetoxyborohydride (Na(OAc)₃BH) and borane-pyridine complexe (BH₃-Py).

- 25
30 Alternatively, resin or polymer supported hydride donating agent on a may be used.

The term "organic carboxylic acid" include but is not limited to aliphatic acid (e.g. acetic, formic, trifluoroacetic), aromatic acid (e.g. benzoic and salicylic), dicarboxylic acid (e.g. oxalic and phthalic). It will be apparent to one of ordinary skill that resin supported organic carboxylic acid may also be used.

The term "enol ether" as used herein have the same meaning as commonly understood by one of ordinary skill in the art to which this invention belongs. Enol ethers may be obtained commercially or prepared by well-known methods. Non-limiting examples of preparation include alkylation or silylation of enolates obtained from carbonyl compounds such as aldehydes, ketones, esters.

It will be appreciated that the amount of a compound of the invention required for use in treatment will vary not only with the particular compound selected but also with the route of administration, the nature of the condition for which treatment is required and the age and condition of the patient and will be ultimately at the discretion of the attendant physician or veterinarian. In general however a suitable dose will be in the range of from about 0.1 to about 750 mg/kg of body weight per day, preferably in the range of 0.5 to 60 mg/kg/day, most preferably in the range of 1 to 20 mg/kg/day.

The desired dose may conveniently be presented in a single dose or as divided dose administered at appropriate intervals, for example as two, three, four or more doses per day.

The compound is conveniently administered in unit dosage form; for example containing 10 to 1500 mg, conveniently 20 to 1000 mg, most conveniently 50 to 700 mg of active ingredient per unit dosage form.

Ideally the active ingredient should be administered to achieve peak plasma concentrations of the active compound of from about 1 to about 75 μ M, preferably about 2 to 50 μ M, most preferably about 3 to about 30 μ M. This may be achieved, for example, by the intravenous injection of a 0.1 to 5% solution of the active ingredient, optionally in saline, or orally administered as a bolus containing about 1 to about 500 mg of the active ingredient. Desirable blood levels may be maintained by a continuous infusion to provide about 0.01 to about 5.0

mg/kg/hour or by intermittent infusions containing about 0.4 to about 15 mg/kg of the active ingredient.

While it is possible that, for use in therapy, a compound of the invention may be administered as the raw chemical it is preferable to present the active ingredient as a pharmaceutical formulation. The invention thus further provides a pharmaceutical formulation comprising compounds of formula (I) or (Ia) or a pharmaceutically acceptable derivative thereof together with one or more pharmaceutically acceptable carriers therefor and, optionally, other therapeutic and/or prophylactic ingredients. The carrier(s) must be "acceptable" in the sense of being compatible with the other ingredients of the formulation and not deleterious to the recipient thereof.

Pharmaceutical formulations include those suitable for oral, rectal, nasal, topical (including buccal and sub-lingual), transdermal, vaginal or parenteral (including intramuscular, sub-cutaneous and intravenous) administration or in a form suitable for administration by inhalation or insufflation. The formulations may, where appropriate, be conveniently presented in discrete dosage units and may be prepared by any of the methods well known in the art of pharmacy. All methods include the step of bringing into association the active compound with liquid carriers or finely divided solid carriers or both and then, if necessary, shaping the product into the desired formulation.

Pharmaceutical formulation suitable for oral administration may conveniently be presented as discrete units such as capsules, cachets or tablets each containing a predetermined amount of the active ingredient; as a powder or granules; as a solution, a suspension or as an emulsion. The active ingredient may also be presented as a bolus, electuary or paste. Tablets and capsules for oral administration may contain conventional excipients such as binding agents, fillers, lubricants, disintegrants, or wetting agents. The tablets may be coated according to methods well known in the art. Oral liquid preparations may be in the form of, for example, aqueous or oily suspensions, solutions, emulsions, syrups or elixirs, or may be presented as a dry

product for constitution with water or other suitable vehicle before use. Such liquid preparations may contain conventional additives such as suspending agents, emulsifying agents, non-aqueous vehicles (which may include edible oils), or
5 .preservatives.

The compounds according to the invention may also be formulated for parenteral administration (e.g. by injection, for example bolus injection or continuous infusion) and may be presented in
10 unit dose form in ampoules, pre-filled syringes, small volume infusion or in multi-dose containers with an added preservative. The compositions may take such forms as suspensions, solutions, or emulsions in oily or aqueous vehicles, and may contain
15 formulatory agents such as suspending, stabilizing an/or dispersing agents. Alternatively, the active ingredient may be in powder form, obtained by aseptic isolation of sterile solid or by lyophilisation from solution, for constitution with a suitable vehicle, e.g. sterile, pyrogen-free water, before use.

20 For topical administration to the epidermis, the compounds according to the invention may be formulated as ointments, creams or lotions, or as a transdermal patch. Such transdermal patches may contain penetration enhancers such as linalool, carvacrol, thymol, citral, menthol and t-anethole. Ointments and
25 creams may, for example, be formulated with an aqueous or oily base with the addition of suitable thickening and/or gelling agents. Lotions may be formulated with an aqueous or oily base and will in general also contain one or more emulsifying agents, stabilizing agents, dispersing agents, suspending agents,
30 thickening agents, or colouring agents.

Formulations suitable for topical administration in the mouth include lozenges comprising active ingredient in a flavoured base, usually sucrose and acacia or tragacanth; pastilles
35 comprising the active ingredient in an inert base such as gelatin and glycerin or sucrose and acacia; and mouthwashes comprising the active ingredient in a suitable liquid carrier.

Pharmaceutical formulations suitable for rectal administration
40 wherein the carrier is a solid are most preferably presented as

unit dose suppositories. Suitable carriers include cocoa butter and other materials commonly used in the art, and the suppositories may be conveniently formed by admixture of the active compound with the softened or melted carrier(s) followed
5 by chilling and shaping in moulds.

Formulations suitable for vaginal administration may be presented as pessaries, tampons, creams, gels, pastes, foams or
10 sprays containing in addition to the active ingredient such carriers as are known in the art to be appropriate.

For intra-nasal administration the compounds of the invention may be used as a liquid spray or dispersible powder or in the form of drops. Drops may be formulated with an aqueous or non-
15 aqueous base also comprising one more dispersing agents, solubilising agents or suspending agents. Liquid sprays are conveniently delivered from pressurized packs.

For administration by inhalation the compounds according to the
20 invention are conveniently delivered from an insufflator, nebulizer or a pressurized pack or other convenient means of delivering an aerosol spray. Pressurized packs may comprise a suitable propellant such as dichlorodifluoromethane, trichlorofluoromethane, dichlorotetrafluoroethane, carbon
25 dioxide or other suitable gas. In the case of a pressurized aerosol the dosage unit may be determined by providing a valve to deliver a metered amount.

Alternatively, for administration by inhalation or insufflation,
30 the compounds according to the invention may take the form of a dry powder composition, for example a powder mix of the compound and a suitable powder base such as lactose or starch. The powder composition may be presented in unit dosage form in, for example, capsules or cartridges or e.g. gelatin or blister packs
35 from which the powder may be administered with the aid of an inhalator or insufflator.

When desired the above described formulations adapted to give sustained release of the active ingredient may be employed.

The compounds of the invention may also be used in combination with other antiviral agents or in combination with any additional agents useful in therapy and may be administered sequentially or simultaneously.

In one aspect of the invention, the compounds of the invention may be employed together with at least one other antiviral agent chosen from protease inhibitors, polymerase inhibitors, and helicase inhibitors.

In another aspect of the invention, the compounds of the invention may be employed together with at least one other antiviral agent chosen from Interferon- α and Ribavirin.

The combinations referred to above may conveniently be presented for use in the form of a pharmaceutical formulation and thus pharmaceutical formulations comprising a combination as defined above together with a pharmaceutically acceptable carrier therefor comprise a further aspect of the invention.

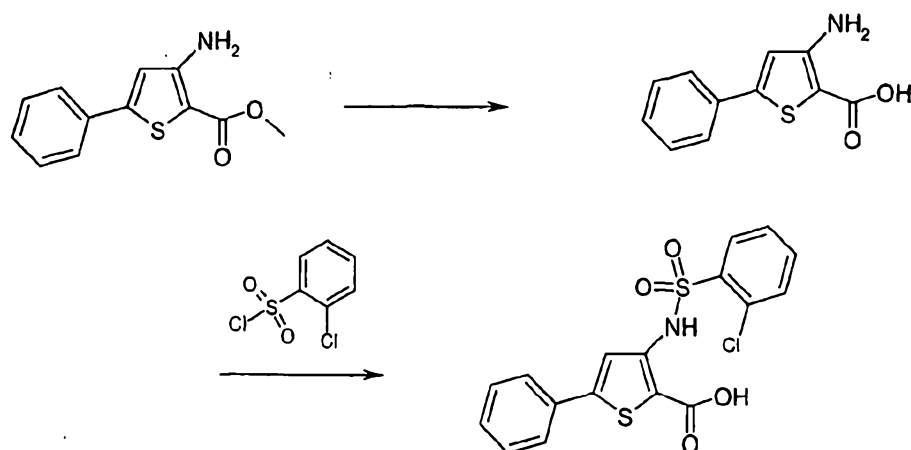
The individual components of such combinations may be administered either sequentially or simultaneously in separate or combined pharmaceutical formulations.

When the compounds of formula (I) or (Ia) or a pharmaceutically acceptable salts thereof is used in combination with a second therapeutic agent active against the same virus the dose of each compound may be either the same as or differ from that when the compound is used alone. Appropriate doses will be readily appreciated by those skilled in the art.

The following general schemes and examples are provided to illustrate various embodiments of the present invention and shall not be considered as limiting in scope.

Example 1

Preparation of 3-(2-Chloro-benzenesulfonylamino)-5-phenyl-thiophene-2-carboxylic acid, compound #29



STEP I

3-Amino-5-phenyl-thiophene-2-carboxylic acid.

- 5 To a suspension of 3-Amino-5-phenyl-thiophene-2-carboxylic acid methyl ester (5 g, 21.459 mmol) in a mixture of THF:MeOH:H₂O (3:2:1, 75 mL), 1N aqueous solution of LiOH·H₂O (64 mL, 64.378 mmol) was added. The reaction mixture was stirred at 85°C (external temperature) for 4h. Solvents were removed under reduced pressure and the residue was partitioned between water and ethyl acetate. The water layer was separated and acidified with 1N HCl solution and then ethyl acetate was added to it. The organic phase was separated, dried (Na₂SO₄) and concentrated to obtain 3-Amino-5-phenyl-thiophene-2-carboxylic acid (4.15 g, 88%) as a yellowish solid. ¹H NMR (DMSO-D₆, 400 MHz): 7.59 (d, 2H), 7.40 (m, 3H), 6.92 (s, 1H).

STEP II

- 3-(2-Chloro-benzenesulfonylamino)-5-phenyl-thiophene-2-carboxylic acid
- 3-Amino-5-phenyl-thiophene-2-carboxylic acid (100mg, 0.457 mmol) was taken in a mixture of dioxane and water (1:1, 25 mL) and then added sodium carbonate (242 mg, 2.285 mmol) and 1-chlorobenzenesulfonyl chloride (289 mg, 1.369 mmol). The reaction mixture was stirred at room temperature for 12 h. Half of the solvent was removed under reduced pressure and then diluted with water and ether in a separatory funnel. The ether layer was separated and the aqueous layer was acidified with 10% KHSO₄ solution. Ethyl acetate was added to the aqueous phase to dissolve the white precipitate. The ethyl acetate layer was

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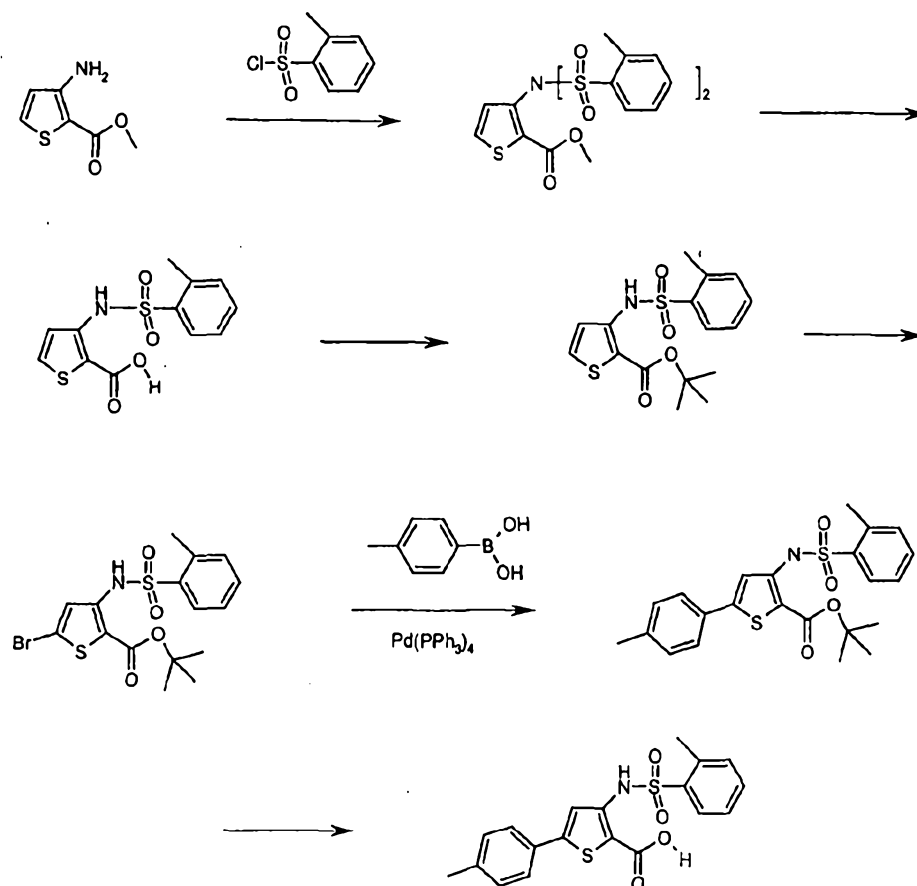
separated, dried (Na_2SO_4) and concentrated to 5 mL. The white solid was filtered and then washed with cold ethyl acetate to obtain 3-(2-Chloro-benzenesulfonylamino)-5-phenyl-thiophene-2-carboxylic acid (125 mg, 69%). ^1H NMR ($\text{DMSO}-d_6$, 400 MHz): 10.51 (bs, 1H), 8.30 (d, 1H), 7.72-7.60 (m, 4H), 7.57 (m, 1H), 7.44 (m, 4H).

The following compounds were prepared in a similar manner as described in general scheme 1:

- 10 Compound #3, Compound #5, Compound #7, Compound #13, Compound #15, Compound #16, Compound #17, Compound #18, Compound #21, Compound #22, Compound #23, Compound #29, Compound #30, Compound #34, Compound #37, Compound #38, Compound #39, Compound #40, Compound #41, Compound #42, Compound #44, Compound #45, Compound
- 15 #46, Compound #49, Compound #50, Compound #52, Compound #53, Compound #54, Compound #55, Compound #76, Compound #94

Example 2

3-(Toluene-2-sulfonylamino)-5-p-tolyl-thiophene-2-carboxylic acid, compound #62



5

STEP I

3-(bis-(Toluene-2-sulfonyl)-amino)-thiophene-2-carboxylic acid methyl ester

- 10 To a cold (0°C) stirred sodium hypochlorite (NaOCl, 10.8% commercial bleach, 124 mL, 180.00 mmol) solution was added o-thiocresol (2.23 g, 2.12 mL, 18.0 mmol). To this vigorous stirred solution was added conc. Sulfuric acid (caution! extremely exothermic, 92 g, 50 mL, 938 mmol) dropwise. The resultant yellow reaction mixture was stirred for 2 h at the same temperature, diluted with water (50 mL) and dichloromethane 50 mL. The organic solution was separated, aqueous solution was extracted with CH₂Cl₂ (2 x 50 mL). The combined organic extracts were washed with water, brine and dried.
- 20 Evaporation of the solvent under reduced pressure furnished the

2-methylsulfonyl chloride (3.13 g, 91.5% yield), which was used in the next step without purification. ¹H NMR (CDCl₃, 300 MHz) 8.07 (td, *J* = 7.3, 1.5 Hz, 1H), 7.61 (tt, *J* = 7.5, 1.1 Hz, 1H), 7.44-7.40 (m, 2H), 2.80 (s, 3H).

To a stirred solution of the methyl 3-amino-thiophene-2-carboxylic acid (1.0 g, 6.36 mmol) and DMAP (776 mg, 6.36 mmol) in CH₂Cl₂, was sequentially added triethyl amine (1.61 g, 15.9 mmol, 2.5 eq) and o-toluenesulfonyl chloride (3.02 g, 15.9 mmol, 2.5 eq), stirred for 24 h. The reaction mixture was diluted with EtOAc (100 mL), washed with 1.2 N HCl (2 x 50 mL), 6 N HCl (40 mL), saturated NaHCO₃ solution, brine and dried. Evaporation of the solvent under reduced pressure yielded 3-(bis-(Toluene-2-sulfonyl)-amino)-thiophene-2-carboxylic acid methyl ester (2.78 g, 93.3%) as a solid. The crude product was used in the next step without purification. ¹H NMR (CDCl₃, 300 MHz) 8.198 (dd, *J* = 8.0, 1.2 Hz, 2H), 7.52 (d, *J* = 5.3 Hz, 1H), 7.5 (dt, *J* = 7.5 Hz, 1.1 Hz, 2H), 7.36 (t, *J* = 7.5 Hz, 3H), 7.28 (d, *J* = 7.6 Hz, 2H), 7.16 (d, *J* = 5.3 Hz, 1H), 3.44 (s, 3H), 2.43 (s, 3H).

STEP II

3-(Toluene-2-sulfonylamino)-thiophene-2-carboxylic acid

To a stirred mixture of 3-(bis-(Toluene-2-sulfonyl)-amino)-thiophene-2-carboxylic acid methyl ester (2.5 g, 5.35 mmol) in 1,4-dioxane/MeOH/water (3:1:1; 62.5 mL) was added aq. 1 N NaOH solution (16.05 mL, 16.05 mmol, 3.0 eq) and heated at 85°C for 3.5 h and it was then cooled to rt. To the reaction mixture was added 1.2 N HCl (16.0 mL), extracted with CHCl₃ (3 x 30 mL), washed with brine and dried. Evaporation of the solvent gave 3-(Toluene-2-sulfonylamino)-thiophene-2-carboxylic acid (1.5 g, 99%) as a white solid. ¹H NMR (DMSO-d₆, 300 MHz) 7.94 (dd, *J* = 7.9 Hz, 1.3 Hz, 1H), 7.76 (d, *J* = 5.5 Hz, 1H), 7.55 (dt, *J* = 7.5 Hz, 1.3 Hz, 1H), 7.42-7.37 (m, 2H), 7.1 (d, *J* = 5.5 Hz, 1H), 2.57 (s, 3H).

STEP III

3-(Toluene-2-sulfonylamino)-thiophene-2-carboxylic acid tert-butyl ester

To a cold (-40°C) mixture of 3-(Toluene-2-sulfonylamino)-thiophene-2-carboxylic acid (1.5 g, 5.05 mmol) in 1,4-dioxane/CHCl₃ (1:2, 12 mL) was bubbled 2-methyl-2-propene gas (15 mL) in a sealed tube. To this was added Conc. H₂SO₄ (0.070 mL, 1.3 mmol) and slowly warmed up to room temperature. The resultant reaction mixture was heated at 70°C for 2.5 days in a sealed tube, cooled to -40°C, stopper was removed. The reaction mixture was slowly brought up to room temperature and stirred until the excess gas is released. The mixture was extracted with EtOAc, washed with aq. NaHCO₃ solution, brine and dried. Evaporation of the solvent and purification of the residue on silica gel using EtOAc/hexane (1:10) as an eluent furnished 3-(Toluene-2-sulfonylamino)-thiophene-2-carboxylic acid tert-butyl ester (1.31 g, 73.5% based on 90% conversion). ¹H NMR (CDCl₃, 300 MHz) 9.89 (s, 1H), 8.01 (d, *J* = 7.9 Hz, 1H), 7.43 (dt, *J* = 7.5 Hz, 1.5 Hz, 1H), 7.3-7.25 (m, 3H), 7.2 (d, *J* = 5.4 Hz, 1H), 2.69 (s, 3H), 1.56 (s, 9H).

20 STEP IV

5-Bromo-3-(toluene-2-sulfonylamino)-thiophene-2-carboxylic acid tert-butyl ester

To a cold (-30°C) stirred solution of diisopropylamine (1.345 g, 1.86 mL, 13.3 mmol, 3.6 eq) in THF (74.0 mL) was added n-BuLi (1.6 M in hexane, 7.63 mL, 12.21 mmol, 3.3 eq) dropwise and stirred for 20 min. To the cold (-78°C) LDA solution was added a solution of 3-(Toluene-2-sulfonylamino)-thiophene-2-carboxylic acid tert-butyl ester (1.31 g, 3.7 mmol, 1.0 eq) in THF (20 mL) dropwise and the solution was stirred for 2h at the same temperature. The resultant red colored solution was then quenched with 1,2-dibromotetrafluoroethane (5.77 g, 2.65 mL, 22.2 mmol, 6.0 eq, passed through K₂CO₃ prior to use) in one portion, stirred for 1 h before being added sat. NH₄Cl solution (15.0 mL). The reaction mixture was warmed up to rt, extracted with EtOAc, washed with brine and dried. Evaporation of the solvent and purification of the residue over silica gel column furnished 5-Bromo-3-(toluene-2-sulfonylamino)-thiophene-2-

carboxylic acid tert-butyl ester (1.2 g, 75% yield). ¹H NMR (CDCl₃, 300 MHz) 9.72 (s, 1H), 8.0 (dd, *J* = 7.8, 1.3 Hz, 1H), 7.47 (dt, *J* = 7.5, 1.2 Hz, 1H), 7.35-7.30 (m, 2H), 7.24 (s, 1H), 2.68 (s, 3H), 1.53 (s, 9H).

5 STEP V

3-(Toluene-2-sulfonylamino)-5-p-tolyl-thiophene-2-carboxylic acid tert-butyl ester

- 10 To the mixture of 4-methylbenzeneboronic acid (38.0 mg, 0.279 mmol) and 5-Bromo-3-(toluene-2-sulfonylamino)-thiophene-2-carboxylic acid tert-butyl ester (40 mg, 0.0925 mmol) in 5:1 mixture of toluene/MeOH (2.0 mL) was added a solution of Pd(PPh₃)₄ (12.0 mg, 0.01 mmol, 10 mol%) in toluene (1.0 mL)
- 15 followed by aqueous 2M Na₂CO₃ solution (0.1 mL, 0.2 mmol). The resultant reaction mixture was heated at 70°C for 16 h, cooled to room temperature, filtered off through MgSO₄ and washed with EtOAc. Evaporation of the solvent and purification of the residue over preparative TLC (1 mm, 60A°) using ethyl
- 20 acetate/hexane (1:10) as an eluent furnished 3-(Toluene-2-sulfonylamino)-5-p-tolyl-thiophene-2-carboxylic acid tert-butyl ester (36.0 mg, 81% yield). ¹H NMR (CDCl₃, 300 MHz) 9.94 (s, 1H), 8.05 (d, *J* = 8.0 Hz, 1H), 7.44-7.25 (m, 6H), 7.18 (d, *J* = 8.1 Hz, 2H), 2.71 (s, 3H), 2.36 (s, 3H), 1.56 (s, 9H).

25 STEP VI

3-(Toluene-2-sulfonylamino)-5-p-tolyl-thiophene-2-carboxylic acid

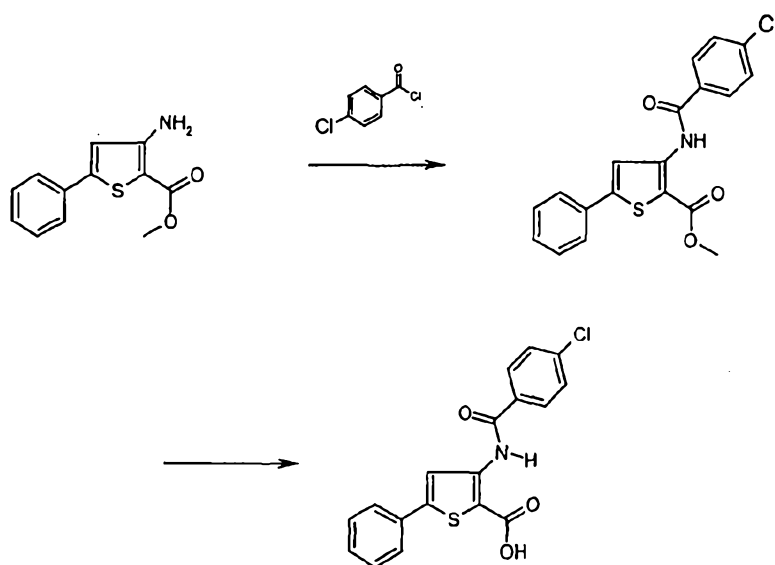
- 30 To a stirred solution of 3-(Toluene-2-sulfonylamino)-5-p-tolyl-thiophene-2-carboxylic acid tert-butyl ester (36.0 mg, 0.081 mmol) in CH₂Cl₂ (1.0 mL) was added TFA (0.5 mL), stirred for 1 h at room temperature and diluted with hexane. Evaporation of the solvent under reduced pressure gave essentially the pure product
- 35 as a solid. The product was purified by triturating with hexane/CH₂Cl₂ furnished 3-(Toluene-2-sulfonylamino)-5-p-tolyl-thiophene-2-carboxylic acid (28.0 mg, 89% yield). ¹H NMR (DMSO-d₆, 300 MHz) 10.21 (br s, 1H), 8.06 (d, *J* = 7.9 Hz, 1H), 7.56-7.36 (m, 6H), 7.24 (d, *J* = 7.9 Hz, 2H), 2.59 (s, 3H), 2.48 (s, 3H).

The following compounds were prepared in a similar manner as described in general scheme 2:

- 5 Compound #6, Compound #8, Compound #11, Compound #14, Compound #24, Compound #56, Compound #57, Compound #58, Compound #59, Compound #60, Compound #62, Compound #63, Compound #64, Compound #65, Compound #66, Compound #67, Compound #68, Compound #69, Compound #70, Compound #71, Compound #552, Compound #79,
- 10 Compound #80, Compound #81, Compound #83, Compound #84, Compound #85, Compound #86, Compound #87, Compound #88, Compound #89, Compound #90, Compound #91

Example 3

- 15 3-(4-Chloro-benzoylamino)-5-phenyl-thiophene-2-carboxylic acid compound #72



STEP I

3-(4-Chloro-benzoylamino)-5-phenyl-thiophene-2-carboxylic acid methyl ester

- 5 To mixture of methyl-3-amino-5-phenylthiophene-2-carboxylate (100 mg, 0.428 mmol) in anhydrous pyridine (4.3 ml) was added p-chlorobenzoyl chloride (71 μ l, 0.556 mmol). The mixture was stirred for 3 hours at room temperature and concentrated. Purification chromatography (silica gel, hexane to hexane: ethyl acetate; 95:5) gave 145 mg (91% yield) of 3-(4-Chloro-
- 10 benzoylamino)-5-phenyl-thiophene-2-carboxylic acid methyl ester. ^1H NMR (CDCl_3 , 400 MHz) 8.54(s, 1H), 7.99-7.96 (m, 2H), 7.73-7.71 (m, 2H), 7.52-7.50 (m, 2H), 7.46-7.39 (m, 3H), 3.95 (s, 3H).

15

STEP II

3-(4-Chlorobenzoylamino)-5-phenyl-thiophene-2-carboxylic acid

- To a mixture of 3-(4-Chloro-benzoylamino)-5-phenyl-thiophene-2-
- 20 carboxylic acid methyl ester (30 mg, 0.081 mmol) in 1 ml of a 3:2:1 solution made with tetrahydrofuran, methanol and water respectively was added lithium hydroxide monohydrated (20 mg, 0.484 mmol). The mixture was stirred 30 minutes at 60°C, cooled to room temperature, diluted with water and washed with ether
- 25 (2x). The collected aqueous layer was then acidified with KHSO_4 20% to pH 3 and extracted with ethyl acetate (3x). The combined ethyl acetate layers were washed with brine, dried (Na_2SO_4) and concentrated. The resulting crude was taken in ethyl acetate and reextracted with NaOH 0.5 N (2x). The combined aqueous layers
- 30 were then back-washed with ethyl acetate and acidified to pH3 with KHSO_4 20% and back-extracted with ethyl acetate (2x). The combined organic layers were washed with brine and dried (Na_2SO_4). ^1H NMR ($\text{DMSO}-d_6$, 400 MHz) 8.35 (s, 1H), 8.02-7.99 (m, 2H), 7.71-7.68 (m, 2H), 7.56-7.53 (m, 2H), 7.43-7.39 (m, 2H),
- 35 7.35-7.31 (m, 1H).

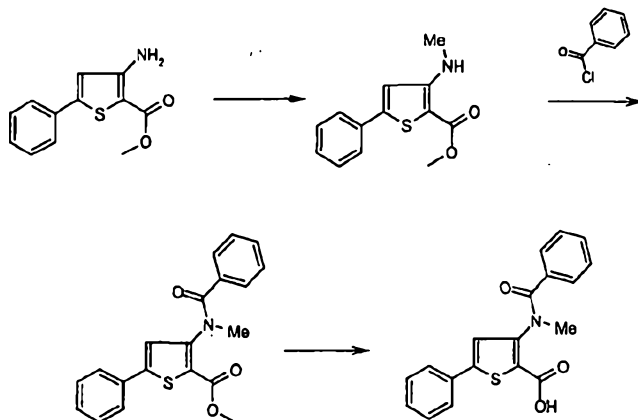
The following compounds were prepared in a similar manner as described in example 3:

Compound #74, Compound #77, Compound #92, Compound #96 ;

Example 4

3-(Benzoyl-methyl-amino)-5-phenyl-thiophene-2-carboxylic acid;

5 compound #550



STEP I

3-Methylamino-5-phenyl-thiophene-2-carboxylic acid methyl ester

10 To a mixture of methyl-3-amino-5-phenylthiophene-2-carboxylate (200 mg, 0.855 mmol) in anhydrous N,N-dimethylformamide (4.6 ml) were added 4.2 ml (8.55 mmol) of 2M iodomethane solution in t-buthylmethylether. The mixture was stirred at 60°C for 18 hours, concentrated and purified using biotage technics (silica gel,
15 hexane to hexane:ethyl acetate;95:5 containing few drops of triethylamine) to give 68 mg (32% yield) of 3-methylamino-5-phenyl-thiophene-2-carboxylic acid methyl ester. ¹H NMR (CDCl₃, 400 MHz) 7.65-7.62 (m, 2H), 7.42-7.36 (m, 3H), 6.86 (broad s, 1H), .
3.83 (s, 3H), 3.04 (d, 3H)

20

STEP II

3-(Benzoyl-methyl-amino)-5-phenyl-thiophene-2-carboxylic acid methyl ester

25 This compound was prepared in a similar manner as for Example 3, Step I; 3-(Benzoyl-methyl-amino)-5-phenyl-thiophene-2-carboxylic acid methyl ester was obtained ¹H NMR (CDCl₃, 400 MHz) 7.60-7.49 (m, 2H), 7.47-7.35 (m, 5H), 7.28-7.20 (m, 3H), 7.11 (broad s, 1H), 3.83 (s, 3H), 3.44 (s, 3H)

30

STEP III

3-(Benzoyl-methyl-amino)-5-phenyl-thiophene-2-carboxylic acid

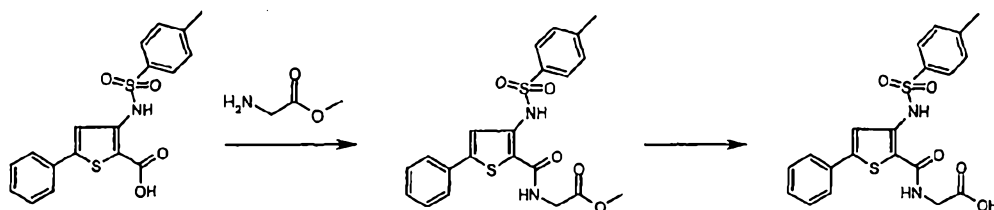
This compound was prepared in a similar manner as in Example 3,
 5 step II; 3-(Benzoyl-methyl-amino)-5-phenyl-thiophene-2-carboxylic acid was obtained; ¹H NMR (CD₃OD, 400 MHz) 7.64-7.62 (m, 2H), 7.47 (s, 1H), 7.44-7.36 (m, 5H), 7.29-7.20 (m, 3H), 3.42 (s, 3H)

10 The following compounds were prepared in a similar manner as described in example 4:

Compound #9 ; Compound #73 Compound #75 ; Compound #75 ;
 Compound #78 ; Compound #93 ; Compound #95.

Example 5

{[5-Phenyl-3-(toluene-4-sulfonylamino)-thiophene-2-carbonyl]-amino}-acetic acid , compound #551



STEP I

{[5-Phenyl-3-(toluene-4-sulfonylamino)-thiophene-2-carbonyl]-amino}-acetic acid methyl ester

To a mixture of 5-phenyl-3-(toluene-4-sulfonylamino)-thiophene-2-carboxylic acid (prepared according to example 2) (50 mg, 0.134 mmol) in anhydrous dimethylformamide (1.4 ml) were added HATU 152 mg, 0.402 mmol), glycine methyl ester hydrochloride (20 mg, 0.161 mmol) followed by collidine (124 μl, 0.938 mmol). The mixture was stirred at room temperature for 1 hour,
 30 concentrated and pre-absorbed on SiO₂. Purification chromatography (hexane to hexane: ethyl acetate; 6:4 to dichloromethane: methanol; 95:5) gave 47 mg of a mixture of {[5-Phenyl-3-(toluene-4-sulfonylamino)-thiophene-2-carbonyl]-amino}-acetic acid methyl ester and collidine. ¹H NMR (CDCl₃, 400 MHz)

7.76-7.73 (m, 2H), 7.61 (s, 1H), 7.57-7.54 (m, 2H), 7.42-7.36 (m, 3H), 7.24-7.22 (m, 2H), 6.19-6.17 (m, 1H), 4.14-4.12 (m, 2H), 3.79 (s, 3H), 2.35 (s, 3H).

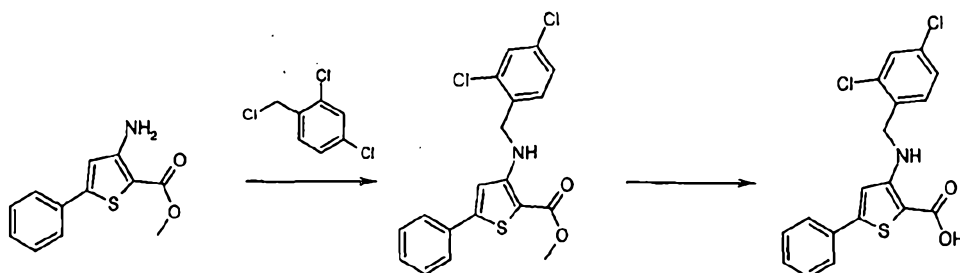
5 STEP II

{[5-Phenyl-3-(toluene-4-sulfonylamino)-thiophene-2-carbonyl]-amino}-acetic acid

Following the procedure described for example 3 (STEP II), 28 mg
 10 (88% yield) of {[5-phenyl-3-(toluene-4-sulfonylamino)-thiophene-2-carbonyl]-amino}-acetic acid were isolated from 33 mg (0.075 mmol) of the {[5-Phenyl-3-(toluene-4-sulfonylamino)-thiophene-2-carbonyl]-amino}-acetic acid methyl ester. ¹H NMR (CD₃OD, 400 MHz): 7.73-7.71 (m, 2H), 7.63-7.61 (m, 2H), 7.54 (s, 1H), 7.45-
 15 7.39 (m, 3H), 7.33-7.31 (m, 2H), 4.88 (s, 2H), 2.36 (s, 3H).

Example 6

3-(2,4-Dichloro-benzylamino)-5-phenyl-thiophene-2-carboxylic acid Compound #48



20

STEP I

3-(2,4-Dichloro-benzylamino)-5-phenyl-thiophene-2-carboxylic acid methyl ester

25 Sodium hydride (60% dispersion in oil, 180 mg, 4.72 mmol) was added to an ice-cold solution of 3-Amino-5-phenyl-thiophene-2-carboxylic acid methyl ester (1000 mg, 4.29 mmol) in 25 ml of dimethylformamide in an atmosphere of N₂. After 5 min, 2,4-dichloro-1-chloromethyl-benzene (755 mg, 3.86 mmol) was added
 30 to the solution and then the reaction mixture was stirred for 30 min at 0°C and 30 min at room temperature. The mixture was partitioned between ether (20 mL) and water (20 mL) and the organic layer was separated. The aqueous phase was washed twice

with ether (2X20 mL) and the combined ether layer was dried (MgSO₄) and concentrated. The residue obtained was then purified by precipitation. The crude product was taken in 25 ml of ethyl acetate, a yellow precipitate came out which was filtered to

- 5 obtain 3-(2,4-Dichloro-benzylamino)-5-phenyl-thiophene-2-carboxylic acid methyl ester, 835 mg (55%). ¹H-NMR (DMSO, 400 MHz): 7,67 ppm (m, 2H, H_{aro}); 7,44-7,35 ppm (m, 6H, H_{aro}); 7,26 ppm (s, 1H, H_{aro}); 4,63 ppm (d, 2H, N-CH₂); 3,75 ppm (s, 3H, O-CH₃)

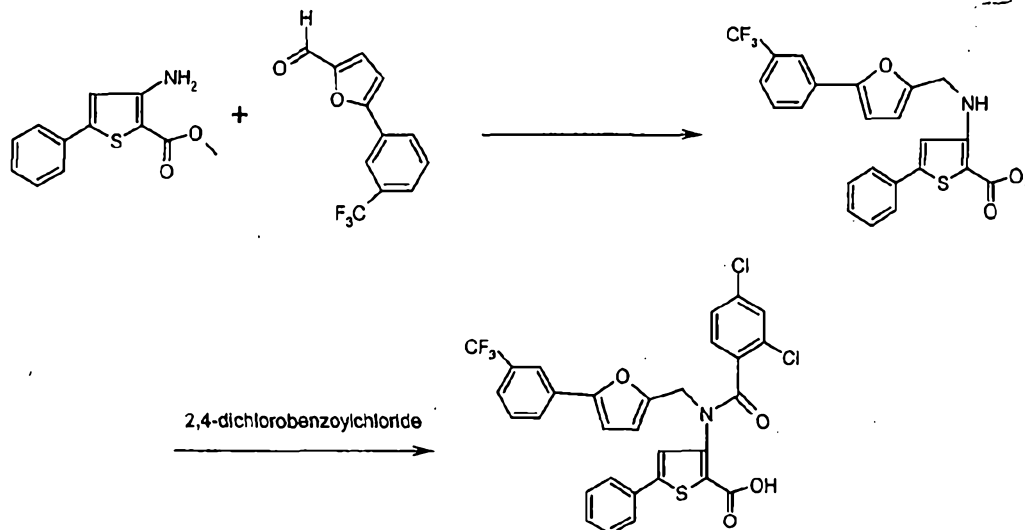
10 STEP II

3-(2,4-Dichloro-benzylamino)-5-phenyl-thiophene-2-carboxylic acid

- 3-(2,4-Dichloro-benzylamino)-5-phenyl-thiophene-2-carboxylic
15 acid methyl ester (70 mg, 0,18 mmol) was dissolved in a mixture of THF-MeOH-H₂O (3:2:1) (20 mL) and then 1080 ul of LiOH 1N was added to it. After 16 h of stirring at temperature of 100°C, solvents were removed and then partitioned between 10 ml of H₂O, 2 ml of KHSO₄ 5% and 10 ml of EtOAc. The organic layer was
20 separated and the aqueous phase was washed twice with ethyl acetate (2 X 10 mL). The combined ethyl acetate layer was dried (MgSO₄) and concentrated to obtain 43 mg (63%) of 3-(2,4-Dichloro-benzylamino)-5-phenyl-thiophene-2-carboxylic acid ¹H-NMR (DMSO, 400 MHz): δ 7,65 ppm (m, 3H, H_{aro}); 7,43-7,32 ppm (m, 5H, H_{aro}); 7,23 ppm (s, 1H, H_{aro}); 4,61 ppm (d, 2H, N-CH₂).
25

Example 7

3-{{(2,4-Dichloro-benzoyl)-[5-(3-trifluoromethyl-phenyl)-furan-2-ylmethyl]-amino}-5-phenyl-thiophene-2-carboxylic acid, Compound #4



5

STEP I

5-Phenyl-3-{{[5-(3-trifluoromethyl-phenyl)-furan-2-ylmethyl]amino}-thiophene-2-carboxylic acid methylester

- 10 To a stirred solution of 3-Amino-5-phenyl-thiophene-2-carboxylic acid methyl ester (100 mg, 0.416mmol) in dichloromethane (15 mL) were added 5-(trifluoromethyl-phenyl)-furan-2-carbaldehyde (100 mg, 0.429 mmol) and molecular sieves. The reaction mixture was stirred at room temperature overnight. The solution was filtered
- 15 over celite and the filtrate was evaporated under reduced pressure. The residue was dissolved in anhydrous methanol (15 mL). and cooled to 0°C in an ice bath. Sodium borohydride (18 mg, 1.1 eq.) was added. The reaction mixture was stirred at this temperature for 2 h. Saturated ammonium chloride (10 mL) was added
- 20 and stirring was continued for an additional 15 min. at room temperature. Methanol was removed and the resulted mixture was extracted with dichloromethane (3 x 30 mL). The organic solution was washed with water, brine and was dried over sodium sulfate. Solvent was evaporated and the crude product was purified on
- 25 silica gel using hexane : ethylacetate 9:1 as eluent to provide the desired product in 34% yield (65 mg).

¹HNMR(CDCl₃, 400MHz): 7.80 (s, 1H), 7.73 (m, 1H), 7.55 (m, 2H), 7.41 (m, 2H), 7.33 (m, 3H), 6.93 (s, 1H), 6.48 (d, 1H), 6.24 (d,

1H), 4.43 (s, 2H), 3.76 (s, 3H).

STEP II

3-((2,4-Dichloro-benzoyl) -[5-(3-trifluoromethyl-phenyl)-furan-2-ylmethyl]-amino)-5-phenyl-thiophene-2-carboxylic acid methyl ester

To a stirred solution of 5-Phenyl-3-[[5-(3-trifluoromethyl-phenyl)-furan-2-ylmethyl]-amino]-thiophene-2-carboxylic acid methylester (65 mg 0.142 mmol) in dichloromethane (3 ml) and saturated NaHCO₃ solution (3 ml) was added a solution of 2,4-dichloro-benzoyl chloride (36 mg, 1.2 eq.) in dichloromethane (0.9 ml). The reaction mixture was stirred vigorously at room temperature for overnight. The organic phase was collected and the aqueous phase was extracted twice with methylene chloride (2 x 15 ml). The organic layers were combined, washed with water, brine and dried over anhydrous Na₂SO₄. Solvent was removed and residue was purified on silica gel using Hexane : EtOAc 9:1 as eluant to give the desired product in 78% yield (70 mg). The proton NMR indicated the presence of rotamers.

¹HNMR(CDCl₃, 400MHz): 7.80 (s, 1H), 7.73 (m, 1H), 7.55 (m, 2H), 7.45 (m, 2H), 7.33 (m, 3H), 7.20 (m, 2H), 7.12 (m, 1H), 6.93 (s, 1H), 6.62 (d, 1H), 6.42 (d, 1H), 5.60 (bd, 1H), 4.70 (bd, 1H), 3.76 (s, 3H).

STEP III

3-((2,4-Dichloro-benzoyl) -[5-(3-trifluoromethyl-phenyl)-furan-2-ylmethyl]-amino)-5-phenyl-thiophene-2-carboxylic acid

3-((2,4-Dichloro-benzoyl) -[5-(3-trifluoromethyl-phenyl)-furan-2-ylmethyl]-amino)-5-phenyl-thiophene-2-carboxylic acid methyl ester (62 mg, 0.098 mmol) was dissolved in THF (5 mL) and water (2 mL). A solution of lithium hydroxide (13 mg, 3eq. in 2 mL of water) was added dropwise. After first few drop, a pink color appeared and disappeared. Mixture was stirred for 5 hrs and acidified with 1N HCl-solution. The product was extracted into ethyl acetate, washed once with water, dried over magnesium sulfate. Solvent was evaporated and the residue was purified on silica gel (Bond-Elute 2 g). The product was elute with a 20 mL gradient of Hexane:EtOAc 9:1. 4:1, 7:3, 3:2, 1:1, 2:3 and EtOAc to give the desired product

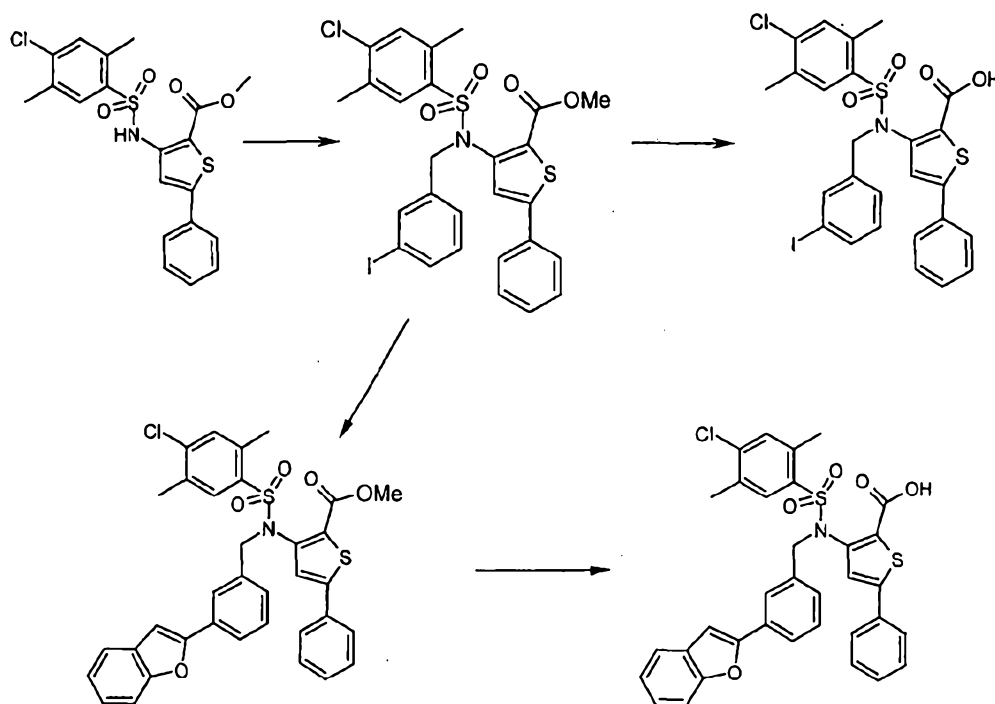
in 76% yield (46 mg).

¹HNMR(CD₃OD, 400MHz): 7.90 (s, 1H), 7.83 (m, 1H), 7.55 (m, 2H),
7.40-7.20 (m, 8H), 7.10 (s, 1H), 6.82 (d, 1H), 6.42 (d, 1H), 5.60
5 (bd, 1H), 4.70 (bd, 1H), 3.86 (s, 3H).

Example 8

Preparation of 3-[(4-Chloro-2,5-dimethyl-benzenesulfonyl)-(3-iodo-benzyl)-amino]-5-phenyl-thiophene-2-carboxylic acid

10 Compound #1 and 3-[(3-Benzofuran-2-yl-benzyl)-(4-chloro-2,5-dimethyl-benzenesulfonyl)-amino]-5-phenyl-thiophene-2-carboxylic acid compound #2.



15

STEP I

To a solution of 3-(4-Chloro-2,5-dimethyl-benzenesulfonylamino)-5-phenyl-thiophene-2-carboxylic acid methyl ester (100 mg, 0.229
20 mmol) in anhydrous DMF (6 mL), 3-iodobenzyl bromide (82 mg, 0.276 mmol) and cesium carbonate (88 mg, 0.276 mmol) were added and the reaction mixture was stirred at room temperature under a N₂ atmosphere for 12 h. The reaction mixture was partitioned between water and ether. The ether layer was separated, dried (Na₂SO₄),

concentrated. The residue was purified by silica gel column chromatography using ethyl acetate and hexane (1:3) as eluent to obtain 3-[(4-Chloro-2,5-dimethyl-benzenesulfonyl)-(3-iodo-benzyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester (130 mg, 87%) as a syrup.

STEP II

3-[(4-Chloro-2,5-dimethyl-benzenesulfonyl)-(3-iodo-benzyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester (25 mg, 0.038 mmol) was taken in a mixture of THF:MeOH:H₂O (3:2:1, 3 mL) and then added 1N aqueous solution of LiOH.H₂O (0.24 mL, 0.228 mmol). The reaction mixture was stirred at room temperature for 12 h. Solvents were removed and the residue was partitioned between water and ethyl acetate. The aqueous layer was acidified using 10 % KHSO₄ solution. The organic layer was separated, dried (Na₂SO₄) and concentrated. The residue was purified by silica gel column chromatography using dichloromethane and methanol (9:1) to obtain 3-[(4-Chloro-2,5-dimethyl-benzene-sulfonyl)-(3-iodo-benzyl)-amino]-5-phenyl-thiophene-2-carboxylic acid (22 mg, 88%) as a white solid. ¹H NMR (CDCl₃, 400 MHz): 7.69(m, 3H), 7.57 (m, 3H), 7.42 (m, 3H), 7.33 (d, 1H), 7.16 (s, 1H), 6.04 (dd, 1H), 4.90 (bs, 2H), 2.36 (s, 6H).

Compound #5 was prepared in a similar manner;

3-[(3-Benzofuran-2-yl-benzyl)-(4-chloro-2,5-dimethyl-benzenesulfonyl)-amino]-5-phenyl-thiophene-2-carboxylic acid compound #2

STEP I

To a degassed solution of 3-[(4-Chloro-2,5-dimethyl-benzenesulfonyl)-(3-iodo-benzyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester (110 mg, 0.169 mmol) and benzofuran-2-boronic acid (55 mg, 0.185 mmol) in a mixture of DME (8 mL) and 2M aqueous Na₂CO₃ (4 mL), Pd(PPh₃)₄ (9 mg) was added and the reaction mixture was stirred at reflux conditions for 2h under a N₂ atmosphere. The reaction mixture was diluted with ethyl acetate and water. The organic layer was separated, dried (Na₂SO₄) and concentrated. 3-[(3-Benzofuran-2-yl-benzyl)-(4-chloro-2,5-dimethyl-benzenesulfonyl)-amino]-5-phenyl-

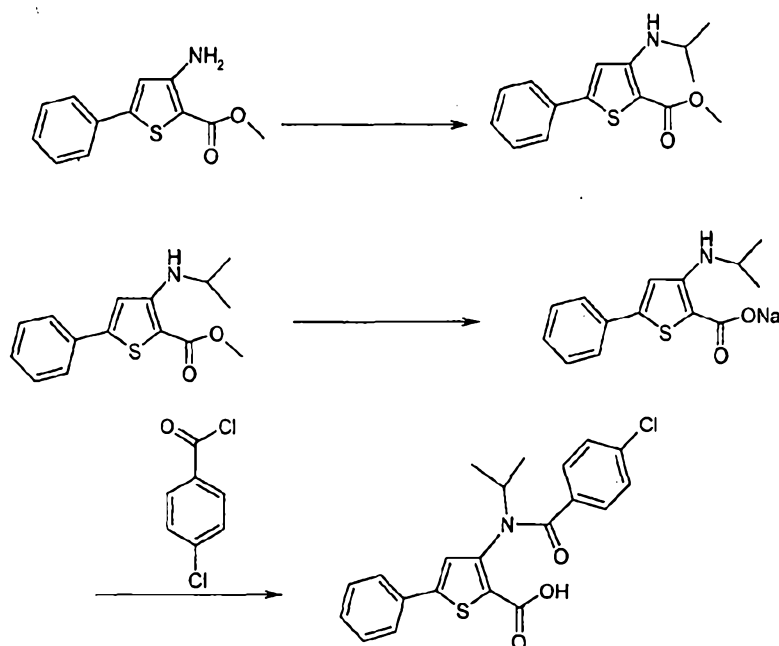
thiophene-2-carboxylic acid methyl ester (107 mg, 100%) was isolated as a thick syrup and used for the next reaction without any further purification.

5 STEP II

3-[(3-Benzofuran-2-yl-benzyl)-(4-chloro-2,5-dimethyl-benzenesulfonyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester (20 mg, 0.031 mmol) was taken in a mixture of THF:MeOH:H₂O (3:2:1, 3 mL) and then added 1N aqueous solution of
10 LiOH.H₂O (0.20 mL, 0.186 mmol). The reaction mixture was stirred at room temperature for 12 h. Solvents were removed and the residue was partitioned between water and ethyl acetate. The aqueous layer was acidified using 10 % KHSO₄ solution. The organic layer was separated, dried (Na₂SO₄) and concentrated. The
15 residue was purified by silica gel column chromatography using dichloromethane and methanol (9:1) to obtain 3-[(3-Benzofuran-2-yl-benzyl)-(4-chloro-2,5-dimethyl-benzene-sulfonyl)-amino]-5-phenyl-thiophene-2-carboxylic acid (14 mg, 70%) as a white
solid. ¹H NMR (DMSO, 400 MHz): δ7.93(s, 1H), 7.84 (s, 1H), 7.74
20 (bd, 1H), 7.65-7.22 (m, 14H), 4.95 (s, 2H), 2.33, 2.23 (2s, 6H).

Example 9

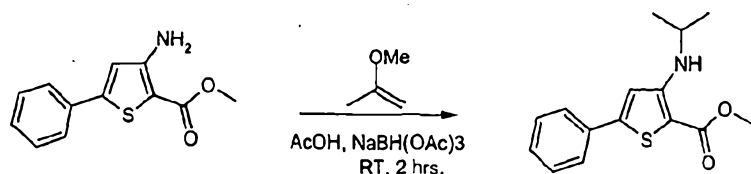
3-[(4-Chloro-benzoyl)-isopropyl-amino]-5-phenyl-thiophene-2-carboxylic acid compound #210 .



STEP I

Method A

A DMF (15 mL) solution of 3-Amino-5-phenyl-thiophene-2-carboxylic acid methyl ester (500 mg, 21.5 mmol) was cooled to 0 °C and then isopropyl iodide (2.57 mL) and NaH (60%, 775 mg, 32.3 mmol) were added under an atmosphere of N₂. The ice bath was removed and the reaction mixture was stirred at room temperature for 1h. The mixture was partitioned between ether and water, the ether layer was separated, dried (Na₂SO₄) and concentrated. The residue was purified by silica gel column chromatography using ethyl acetate and hexane (5:95) as eluent to obtain 3-isopropylamino-5-phenyl-thiophene-2-carboxylic acid methyl ester (189 mg, 32%) as a solid. ¹H NMR (CDCl₃, 400 MHz): 7.62 (d, 2H), 7.40 (m, 3H), 6.91 (s, 1H), 3.84 (s, 3H), 1.35 (d, 6H).

Method B

To a stirred solution of 3-Amino-5-phenyl-thiophene-2-carboxylic acid methyl ester (1.82 g, 7.8 mmol) in 1,2-dichloroethane (40 mL) was added sequentially 2-methoxypropene (3.0 mL, 31.2 mmol), AcOH (1.8 mL, 31.2 mmol) and $\text{NaBH}(\text{OAc})_3$ (3.31 g, 15.6 mmol) and
5 stirred for 2 hrs. It was then diluted with EtOAc and H_2O . The aqueous solution was adjusted to pH = 7 by adding NaHCO_3 . The aqueous phase was extracted with EtOAc, the combined extract was washed with brine and dried on MgSO_4 and filtered. Purification on bond elute with hexane to 5% EtOAc-hexane furnished 3-Amino-
10 5-phenyl-thiophene-2-carboxylic acid methyl ester (2.07 g, 96% yield).

The intermediate compounds 3-Cyclohexylamino-5-phenyl-thiophene-2-carboxylic acid methyl ester, 3-(1-Methyl-piperidin-4-ylamino)-5-phenyl-thiophene-2-carboxylic acid methyl ester and
15 3-(1-Methyl-piperidin-4-ylamino)-5-phenyl-thiophene-2-carboxylic acid methyl ester were prepared in a similar manner as described and used as intermediates in the synthesis of compound #543, compound #553 and compound #573

20 STEP II

To a suspension of 3-isopropylamino-5-phenyl-thiophene-2-carboxylic acid methyl ester (1.2 g, 4.364 mmol) in a mixture of H_2O (22 mL) and dioxane (35 mL), 1N aqueous solution of NaOH (13
25 mL, 13.00 mmol) was added. The reaction mixture was stirred at 100°C for 3h. The reaction mixture was used for the next reaction without any further purification.

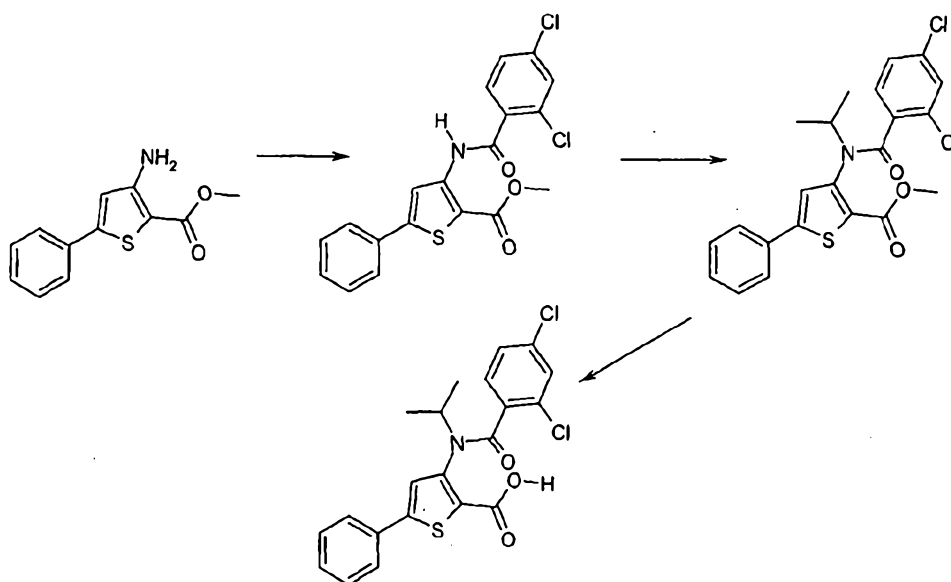
To this reaction mixture of 3-Amino-5-phenyl-thiophene-2-carboxylic acid sodium salt (23 mL, 1.41 mmol), 4-chlorobenzoyl
30 chloride (0.269 mL, 2.11 mmol) was added at 0°C . The pH of the solution was maintained at 9 by adding 1N NaOH solution and then stirred at room temperature for 5h. The reaction mixture was diluted with ethyl acetate and water. The water layer was acidified by adding 1N HCl solution. The organic layer was
35 separated, dried (Na_2SO_4) and concentrated. The crude product was

purified by recrystallization from ethyl acetate to obtain the pure 3-[(4-Chloro-benzoyl)-isopropyl-amino]-5-phenyl-thiophene-2-carboxylic acid (45 mg) as a white solid. ^1H NMR (DMSO- D_6 , 400 MHz): 7.58 (d, 2H), 7.38-7.26 (m, 6H), 7.13 (d, 1H), 4.77 (m, 1H), 1.25 (d, 3H), 1.02 (d, 3H). ESI $^-$ (M-H): 398.

Similarly, the following compounds were made: Compound #218 , Compound #219 , Compound #226 , Compound #234 , Compound #243 , Compound #246 , Compound #250 , Compound #262 , Compound #324 , Compound 326 , Compound #331 .

Example 10

3-[(2,4-Dichloro-benzoyl)-isopropyl-amino]-5-phenyl-thiophene-2-carboxylic acid compound #149



Step I

3-(2,4-Dichloro-benzoylamino)-5-phenyl-thiophene-2-carboxylic acid methyl ester.

To a ice-cold solution of 3-Amino-5-phenyl-thiophene-2-carboxylic acid methyl ester 1 (5 g, 21.5 mmol) and triethylamine (4.56 g, 45.0 mmol) in dichloromethane (100 ml)

was added 2,4-dichlorobenzoyl chloride (3.90 g, 19.4 mmol). The reaction mixture was stirred for 30 min at 0°C and 16 h at room temperature. Then, the reaction mixture was partitioned between 25 ml of H₂O, 50 ml sat. NaHCO₃ and 50 ml of CH₂Cl₂. The organic layer was separated and the aqueous phase was washed twice with CH₂Cl₂ (2 X 50 mL). The combined dichloromethane layer was dried (MgSO₄), concentrated and the residue was purified by recrystallization in CH₂Cl₂ to obtain 5.832 g (74%) as a white solid of 3-(2,4-Dichloro-benzoylamino)-5-phenyl-thiophene-2-carboxylic acid methyl ester. NMR ¹H (CDCl₃, 400 MHz): 8,30 ppm (s, 1H, H_{aro}); 7,74-7,66 ppm (m, 3H, H_{aro}); 7,51 ppm (d, 1H, H_{aro}); 7,46-7,34 ppm (m, 4H, H_{aro}); 3,91 ppm (s, 3H).

Step II

3-[(2,4-Dichloro-benzoyl)-isopropyl-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester.

Sodium Hydride (60% dispersion in oil, 190 mg, 5,2 mmol) was added to an ice-cold solution of 3-(2,4-Dichloro-benzoylamino)-5-phenyl-thiophene-2-carboxylic acid methyl ester (2) (1.5 g, 3,69 mmol) in 350 ml of *N,N*-dimethylformamide in an atmosphere of N₂. After 5 min, 2-Iodo-propane (941 mg, 5.54 mmol) was added to the solution and then the reaction mixture was stirred for 30 min at 0°C and 64 h at room temperature. The mixture was

partitioned between ether (200 mL) and water (350 mL) and the organic layer was separated. The aqueous phase was washed twice with ether (2 X 70 mL) and the combined ether layer was dried (MgSO₄), concentrated and the residue was purified by flash chromatography (10% EtOAc/Hexane) to obtain 908 mg (55%) of 3-[(2,4-Dichloro-benzoyl)-isopropyl-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester. NMR ¹H (CDCl₃, 400 MHz): Rotamere 95/05 : 7,54 ppm (dd, 2H, H_{aro}); 7,49-7,35 ppm (m, 3H, H_{aro}); 7,29-7,25 ppm (m, 2H, H_{aro}); 7,15 ppm (d, 1H, H_{aro}); 7,05 ppm (d, 1H, H_{aro}); 5,09 ppm (hex, 1H, N-CH(CH₃), major rotamere); 3,99 ppm (hex, N-CH(CH₃), minor rotamere); 3,89 ppm (s, 3H); 1,40 ppm (d,

3H, N-CH(CH₃), major rotamere); 1,28 ppm (d, N-CH(CH₃), minor rotamere); 1,09 ppm (d, 3H, N-CH(CH₃), major rotamere); 1,01 ppm (d, N-CH(CH₃), minor rotamere).

5 Step III

3-[(2,4-Dichloro-benzoyl)-isopropyl-amino]-5-phenyl-thiophene-2-carboxylic acid.

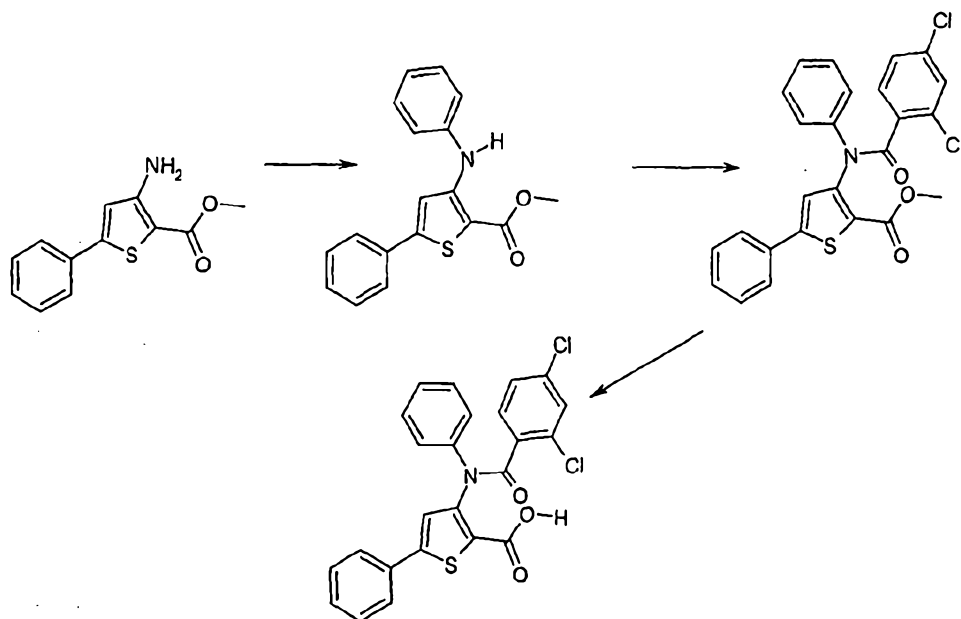
3-[(2,4-Dichloro-benzoyl)-isopropyl-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester (3) (345 mg, 0.77 mmol) was dissolved in a mixture of THF-MeOH-H₂O (3:2:1) (30 mL) and then 4,6 ml of LiOH 1N was added to it. After 120 min of stirring at room temperature, solvent was removed and then partitioned between 25 ml of H₂O, 4 ml of KHSO₄ 5% and 25 ml of EtOAc. The organic layer was separated and the aqueous phase was washed twice with ethyl acetate (2 X 10 mL). The combined ethyl acetate layer was dried (MgSO₄), concentrated and the residue was purified by preparative chromatography (10% MeOH/CH₂Cl₂) to obtain 175 mg (53%) as a white solid of 3-[(2,4-Dichloro-benzoyl)-isopropyl-amino]-5-phenyl-thiophene-2-carboxylic acid. NMR ¹H (DMSO, 400 MHz): Rotamer 95/05 : 7,82 ppm (m, H_{aro}, minor rotamer); 7,69 ppm (d, 2H, H_{aro}); 7,61 ppm (d, 1H, H_{aro}); 7,51-7,37 ppm (m, 4H, H_{aro}); 7,35-7,28 ppm (m, 2H, H_{aro}); 4,89 ppm (hex, 1H, N-CH(CH₃), major rotamer); 3,84 ppm (hex, N-CH(CH₃), minor rotamer); 1,36 ppm (d, 3H, N-CH(CH₃), major rotamer); 1,25 ppm (d, N-CH(CH₃), minor rotamer); 1,03 ppm (d, 3H, N-CH(CH₃), major rotamer); 0,93 ppm (d, N-CH(CH₃), minor rotamere).

The following compounds were prepared in a similar manner:

30 Compound #201 , Compound #204 , Compound #233 , Compound #244 , Compound #261 , Compound #264 , Compound #299 .

Example 11

3-[(2,4-Dichloro-benzoyl)-phenyl-amino]-5-phenyl-thiophene-2-carboxylic acid. Compound #208.



Step I

5-Phenyl-3-phenylamino-thiophene-2-carboxylic acid methyl ester.

5

To a solution of 3-Amino-5-phenyl-thiophene-2-carboxylic acid methyl ester (1 g, 4.29 mmol) in dichloromethane (50 ml) was added phenyl boronic acid (1.05 g, 8.6 mmol), pyridine (680 mg, 8.6 mmol) and copper(II) acetate (1.18 g, 6.5 mmol). The reaction mixture was stirred for 16 h at room temperature. Then, the reaction mixture was filtered through celite, concentrated and the residue was purified by flash chromatography (9:1 Hexane/EtOAc) to obtain 435 mg (33%) of 5-Phenyl-3-phenylamino-thiophene-2-carboxylic acid methyl ester. NMR ^1H (CDCl_3 , 400 MHz): 7.38 ppm (dd, 2H, H_{aro}); 7.35-7.26 ppm (m, 5H, H_{aro}); 7.19 ppm (s, 1H, H_{aro}); 7.15 ppm (dd, 2H, H_{aro}); 7.02 ppm (ddt, 1H, H_{aro}); 3.82 ppm (s, 3H).

15

Step II

3-[(2,4-Dichloro-benzoyl)-phenyl-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester.

20

Sodium Hydride (60% dispersion in oil, 80 mg, 1,5 mmol) was added to an ice-cold solution 5-Phenyl-3-phenylamino-thiophene-2-carboxylic acid methyl ester (2) (230 mg, 0,74 mmol) in 20 ml of *N,N*-dimethylformamide in an atmosphere of N_2 . After 5 min, 2,4-Dichloro-benzoyl chloride (310 mg, 1.48 mmol) was added to the solution and then the reaction mixture was stirred for 30 min at 0°C and 16 h at room temperature. The mixture was partitioned between ether (20 mL) and water (20 mL) and the organic layer was separated. The aqueous phase was washed twice with ether (2 X 10 mL) and the combined ether layer was dried ($MgSO_4$), concentrated and the residue was purified by preparative chromatography (30% EtOAc/Hexane) to obtain 58 mg (16%) of 3-[(2,4-Dichloro-benzoyl)-phenyl-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester. NMR 1H ($CDCl_3$, 400 MHz): 7,65-7,10 ppm (m, 14H, H_{aro}); 3,77 ppm (s, 3H).

Step III

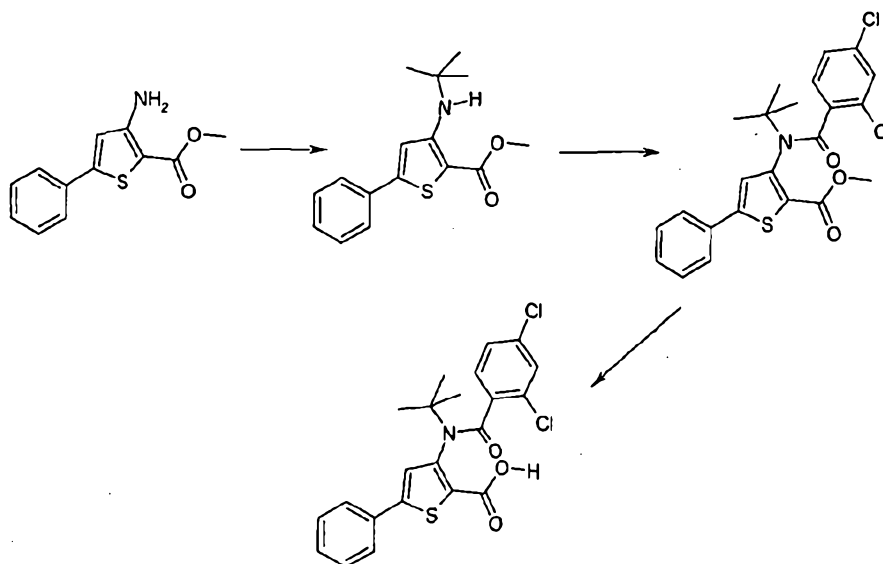
3-[(2,4-Dichloro-benzoyl)-phenyl-amino]-5-phenyl-thiophene-2-carboxylic acid.

3-[(2,4-Dichloro-benzoyl)-phenyl-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester (55 mg, 0.11 mmol) was dissolved in a mixture of THF-MeOH- H_2O (3:2:1) (15 mL) and then 0.66 ml of LiOH 1N was added to it. After 60 min of stirring at room temperature, solvents were removed and then partitioned between 15 ml of H_2O , 4 ml of $KHSO_4$ 5% and 15 ml of EtOAc. The organic layer was separated and the aqueous phase was washed twice with ethyl acetate (2 X 10 mL). The combined ethyl acetate layer was dried ($MgSO_4$), concentrated and the residue was purified by preparative chromatography (10% MeOH/ CH_2Cl_2) to obtain 32 mg (60%) of 3-[(2,4-Dichloro-benzoyl)-phenyl-amino]-5-phenyl-thiophene-2-carboxylic acid. NMR 1H (DMSO, 400 MHz): Rotamer : 7,75 ppm (d, 1H, H_{aro}); 7,68 ppm (2H, H_{aro}); 7,53 ppm (d, H_{aro} , minor rotamer); 7,51-7.23 ppm (m, 11H, H_{aro} , minor rotamer); 7,17 ppm (H_{aro} , minor rotamer).

Compound #525 was prepared in a similar manner.

Example 12

- 5 3-[tert-Butyl-(2,4-dichloro-benzoyl)-amino]-5-phenyl-thiophene-2-carboxylic acid compound #327



Step I

- 10 3-tert-Butylamino-5-phenyl-thiophene-2-carboxylic acid methyl ester.

Concentrated sulfuric acid (10 drop) was added to a solution of 3-Amino-5-phenyl-thiophene-2-carboxylic acid methyl ester (500
15 mg, 2,15 mmol) in 20 ml of dioxane/ chloroforme (2 :3) in a sealed tube. After cooling the solution at -78 °C, put 20 ml of isobutene gaz. The sealed tube was closed and then the reaction mixture was stirred for 6 days at 60 °C. The solvent was removed and then partitioned between 15 ml of sat. Na₂CO₃ solution and 15
20 ml of EtOAc. The organic layer was separated, the aqueous phase was washed twice with ethyl acetate and the combined ethyl acetate layer was dried (MgSO₄), concentrated and the residue was purified by flash chromatography (5% EtOAc/Hexane) to obtain 385 mg (62%) of 3-tert-Butylamino-5-phenyl-thiophene-2-carboxylic

acid methyl ester. NMR ^1H (CDCl_3 , 400 MHz): 7,65 ppm (d, 2H, H_{aro}); 7,44-7,38 ppm (m, 3H, H_{aro}); 7,07 ppm (s, 1H, H_{aro}); 3,86 ppm (s, 3H); 1,48 ppm (s, 9H).

5 Step II

3-[tert-Butyl-(2,4-dichloro-benzoyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester.

To a solution of 3-tert-Butylamino-5-phenyl-thiophene-2-carboxylic acid methyl ester (100 mg, 0.35 mmol) in
10 dichloroethane (10 ml) in an atmosphere of N_2 was added 2,4-dichloro-benzoyl chloride (79 mg, 0.38 mmol). The reaction mixture was stirred for 16 h at reflux. Then, the solvents were removed and the residue was purified by flash chromatography
15 (9:1 Hexane/EtOAc) to obtain 112 mg (69%) of 3-[tert-Butyl-(2,4-dichloro-benzoyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester. NMR ^1H (CDCl_3 , 400 MHz): 7,50 ppm (m, 2H, H_{aro}); 7,44-7,34 ppm (m, 3H, H_{aro}); 7,27 ppm (s, 1H, H_{aro}); 7,18 ppm (dl, 1H, H_{aro}); 7,14 ppm (d, 1H, H_{aro}); 7,00 ppm (dd, 1H, H_{aro});
20 H_{aro}); 3,93 ppm (s, 3H); 1,56 ppm (s, 9H).

Step III

3-[tert-Butyl-(2,4-dichloro-benzoyl)-amino]-5-phenyl-thiophene-2-carboxylic acid.

25

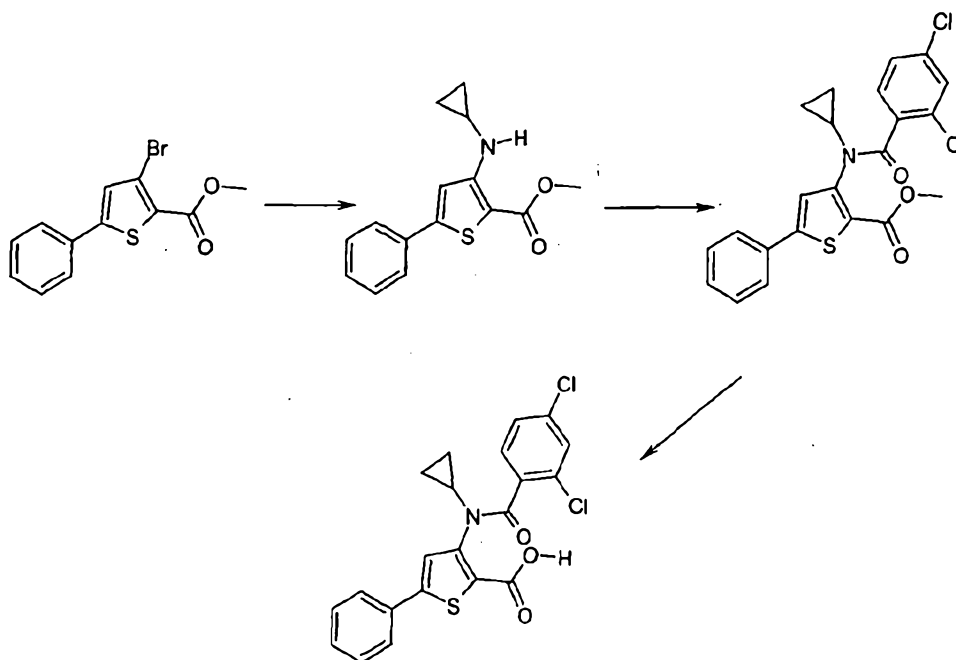
3-[tert-Butyl-(2,4-dichloro-benzoyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester (112 mg, 0.24 mmol) was dissolved in a mixture of THF-MeOH- H_2O (3:2:1) (15 mL) and then 1.5 ml of LiOH 1N was added to it. After 3 h of stirring at room
30 temperature, solvent was removed and then partitioned between 15 ml of H_2O , 4 ml of KHSO_4 5% and 15 ml of EtOAc. The organic layer was separated and the aqueous phase was washed twice with ethyl acetate (2 X 10 mL). The combined ethyl acetate layer was dried (MgSO_4), concentrated and the residue was purified by preparative
35 chromatography (10% MeOH/ CH_2Cl_2) to obtain 32 mg (29 %) of 3-

[tert-Butyl-(2,4-dichloro-benzoyl)-amino]-5-phenyl-thiophene-2-carboxylic acid. NMR ^1H (DMSO, 400 MHz): 7,62 ppm (d, 2H, H_{aro}); 7,44-7,34 ppm (m, 4H, H_{aro}); 7,32-7,12 ppm (m, 3H, H_{aro}); 2,48 ppm (s, 9H).

5

Example 13

3-[Cyclopropyl-(2,4-dichloro-benzoyl)-amino]-5-phenyl-thiophene-2-carboxylic acid. Compound #333



10 Step I

3-Cyclopropylamino-5-phenyl-thiophene-2-carboxylic acid methyl ester.

To a solution of 3-Bromo-5-phenyl-thiophene-2-carboxylic acid methyl ester (250 mg, 0.89 mmol) in toluene (25 ml) was added cyclopropylamine (57 mg, 1.0 mmol), cesium carbonate (382 mg, 1.2 mmol), BINAP (50 mg, 0.08 mmol) and tris (dibenzylidenacetone)dipaladium (0) (38 mg, 0.04 mmol). The reaction mixture was stirred for 16 h at 110 °C in a sealed tube. The mixture was partitioned between toluene (20 mL) and water (20 mL) and the organic layer was separated. The aqueous phase was

washed twice with toluene (2 X 10 mL) and the combined toluene layer was dried (MgSO₄), concentrated and the residue was purified by preparative chromatography (10% EtOAc/Hexane) to obtain 52 mg (22 %) of 3-Cyclopropylamino-5-phenyl-thiophene-2-carboxylic acid methyl ester. NMR ¹H (CDCl₃, 400 MHz): 7,67-7,62 ppm (m, 2H, H_{aro}); 7,43-7,32 ppm (m, 3H, H_{aro}); 7,16 ppm (s, 1H, H_{aro}); 3,82 ppm (s, 3H); 2,65 ppm (m, 1H); 0,62 ppm (m, 2H); 0,35 ppm (m, 2H).

10 Step II

3-[Cyclopropyl-(2,4-dichloro-benzoyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester.

To a solution of 3-Cyclopropylamino-5-phenyl-thiophene-2-carboxylic acid methyl ester (52 mg, 0.19 mmol) in dichloroethane (10 ml) in an atmosphere of N₂ was added 2,4-dichlorobenzoyl chloride (45 mg, 0.21 mmol). The reaction mixture was stirred for 16 h at reflux. Then, the solvent was removed and the residue was purified by flash chromatography (8:2 Hexane/EtOAc) to obtain 85 mg (99%) of 3-[Cyclopropyl-(2,4-dichloro-benzoyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester. NMR ¹H (CDCl₃, 400 MHz): 7,64 ppm (d, 2H, H_{aro}); 7,47 ppm (m, 2H, H_{aro}); 7,44-7,33 ppm (m, 3H, H_{aro}); 7,21-7,12 ppm (m, 2H, H_{aro}); 3,89 ppm (s, 3H); 3,33 ppm (m, minor rotamer); 3,13 ppm (m, 1H, major rotamer) 1,01-0,49 ppm (m, 4H).

25 Step III

3-[Cyclopropyl-(2,4-dichloro-benzoyl)-amino]-5-phenyl-thiophene-2-carboxylic acid.

30 3-[Cyclopropyl-(2,4-dichloro-benzoyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester (85mg, 0.19mmol) was dissolved in a mixture of THF-MeOH-H₂O (3:2:1) (10 mL) and then 1.2 ml of LiOH 1N was added to it. After 60 min of stirring at room temperature, solvent was removed and then partitioned between 15 ml of H₂O, 4 ml of KHSO₄ 5% and 15 ml of EtOAc. The organic layer

was separated and the aqueous phase was washed twice with ethyl acetate (2 X 10 mL). The combined ethyl acetate layer was dried (MgSO₄), concentrated and the residue was purified by preparative chromatography (10% MeOH/CH₂Cl₂) to obtain 22 mg (27 %) of 3-

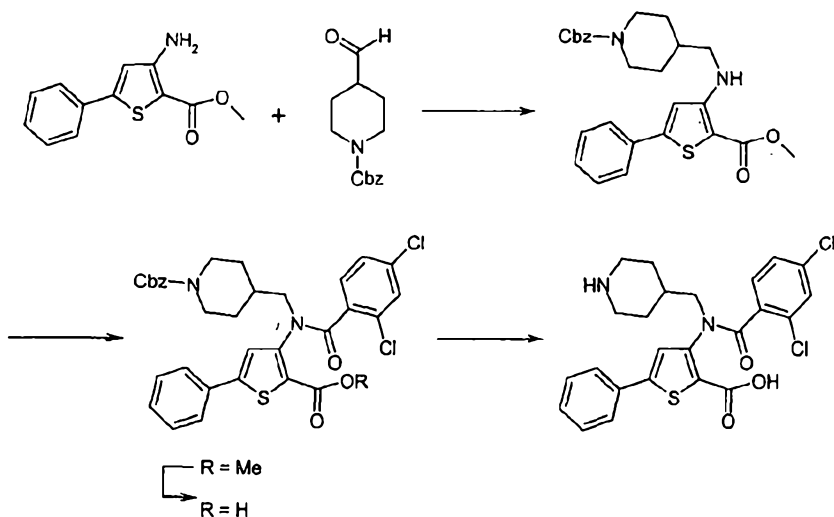
5 [Cyclopropyl-(2,4-dichloro-benzoyl)-amino]-5-phenyl-thiophene-2-carboxylic acid. NMR ¹H (DMSO, 400 MHz): rotamer : 7,75 ppm (m, 2H, H_{aro}); 7,68 ppm (m, H_{aro}, minor rotamer) ; 7,62-7,55 ppm (m, 2H, H_{aro}); 7,52 ppm (m, H_{aro}, minor rotamer) ; 7,48-7,27 ppm (m, 5H, H_{aro}); 3,14 ppm (m, minor rotamer); 3,04 ppm (m, 1H, major rotamer); 0,87-0,42 ppm (m, 4H,).

The following compounds were prepared in a similar manner:

Compound #403 , Compound #404

Example 14

15 3-[(2,4-dichloro-benzoyl)-piperidin-4-ylmethylamino]-5-phenyl-thiophene-2-carboxylic acid Compound #519 .



STEP I

20 A suspension of 3-amino-5-phenyl-thiophene-2-carboxylic acid methyl ester (0.70 g, 3 mmol) and 4-formyl N-Cbz-piperidine (0.74 g, 3 mmol) in THF (1.2 mL) was treated with dibutyltin dichloride (46 mg, 0.15 mol) followed by phenylsilane (0.41 mL, 3.3 mmol). The mixture was stirred for 2 days at room temperature. The solvent was then evaporated and the residue was purified by silica gel column chromatography using

CH₂Cl₂:hexanes:EtOAc as eluent to provide 4-[(2-Methoxycarbonyl-5-phenyl-thiophen-3-ylamino)-methyl]-piperidine-1-carboxylic acid benzyl ester (0.6906 g, 50% yield).

5 STEP II

4-[(2-Methoxycarbonyl-5-phenyl-thiophen-3-ylamino)-methyl]-piperidine-1-carboxylic acid benzyl ester (133 mg, 0.28 mmol) was dissolved in 1,2-dichloroethane (2.8 mL) and was treated with 2,4-dichlorobenzoyl chloride (60 µL, 0.43 mmol). The solution was
10 heated at reflux for 1 day. The solvent was then evaporated and the residue purified by silica gel column chromatography using hexanes:EtOAc as eluent to provide 4-[[[(2,4-Dichloro-benzoyl)-(2-methoxycarbonyl-5-phenyl-thiophen-3-yl)-amino]-methyl]-piperidine-1-carboxylic acid benzyl ester (0.156 g, 85% yield).

15 STEP III

4-[[[(2,4-Dichloro-benzoyl)-(2-methoxycarbonyl-5-phenyl-thiophen-3-yl)-amino]-methyl]-piperidine-1-carboxylic acid benzyl ester (150 mg, 0.24 mmol) was dissolved in a mixture of THF:MeOH:H₂O (3:2:1, 2.4 mL) and treated with LiOH.H₂O (29.6 mg, 0.7 mmol). The
20 solution was heated at 55 °C for 2 h. The solvents were removed and the residue was acidified using HCl. The product was extracted with EtOAc and the organic layers were washed with brine and dried. The residue was purified by silica gel column chromatography using EtOAc:MeOH:AcOH as eluent to provide 4-[[[(2-
25 Carboxy-5-phenyl-thiophen-3-yl)-(2,4-dichloro-benzoyl)-amino]-methyl]-piperidine-1-carboxylic acid benzyl ester (124 mg, 85% yield).

STEP IV

30 4-[[[(2-Carboxy-5-phenyl-thiophen-3-yl)-(2,4-dichloro-benzoyl)-amino]-methyl]-piperidine-1-carboxylic acid benzyl ester (124 mg, 0.2 mmol) was dissolved in MeOH (2 mL) and treated with 10% Pd/C (200 mg) under H₂ balloon. The reaction was stirred at room temperature for 18 h and the mixture was filtered on celite.
35 The solution was evaporated to a residue that was purified by

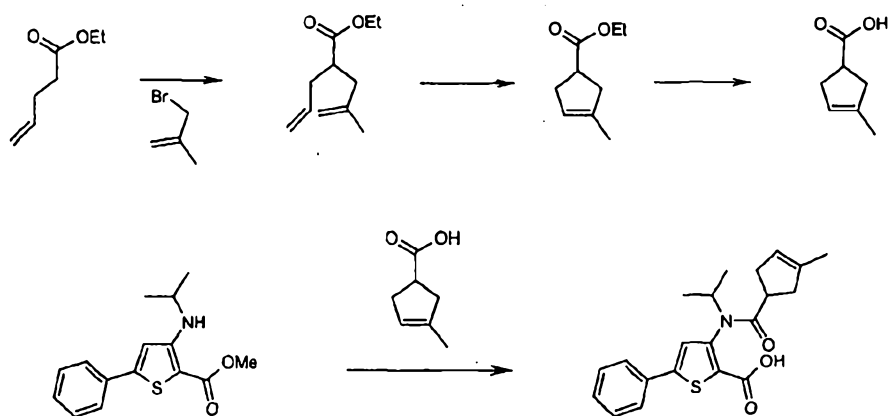
reverse-phase HPLC to provide 3-[(2,4-Dichloro-benzoyl)-piperidin-4-ylmethyl-amino]-5-phenyl-thiophene-2-carboxylic acid (17.3 mg, 18% yield). ^1H NMR (CD_3OD , 300 MHz): 7.55 (d, 1 H), 7.50 (m, 2 H), 7.27-7.39 (m, 4 H), 7.25 (s, 1 H), 7.18 (dd, 1 H), 4.12 (m, 1 H), 3.75 (m, 1 H), 3.43 (m, 2 H), 2.96 (q, 2 H), 2.65 (d, 2 H), 2.05 (m, 1 H), 1.62 (m, 2 H).

The following compounds were prepared in a similar manner:

Compound #503 , Compound #509 , Compound #519 , Compound #529 ,
 10 Compound #537 , Compound #538 , Compound #516 , Compound #522 ,
 Compound #535 .

Example 15

3-[Isopropyl-(3-methyl-cyclopent-3-enecarbonyl)-amino]-5 phenyl-
 15 thiophene-2-carboxylic acid Compound #405



Step I:

To a cold (-78°C) stirred solution of LDA (generated from DIPA (1.42 mL, 10.14 mmol), BuLi (5.85 mL, 9.36 mmol) in THF at -78°C for 20 min) in THF (31 mL) was added a solution of Pent-4-enoic acid ethyl ester (1.0 g, 7.8 mmol, 1.2 eq.) in THF (9.0 mL). After stirred for 1 h, neat 3-Bromo-2-methyl-propene (2.03 g, 15.0 mmol, 1.51 mL) was added and slowly warmed up to room
 25 temperature for overnight. The reaction mixture was then quenched with saturated NH_4Cl solution, extracted with ether, washed with brine and dried. Evaporation of the solution furnished the 2-Allyl-4-methyl-pent-4-enoic acid ethyl ester

(1.45 g, 100%) as an oil which was used in the next step without purification. ^1H NMR (400 MHz, CDCl_3), 5.78-5.71 (m, 1H), 5.05 (d, $J = 18.6$ Hz, 1H), 5.02 (d, $J = 9.4$ Hz, 1H), 4.76 (brs, 1H), 4.70 (s, 1H), 4.11 (dq, $J = 7.2, 1.0$ Hz, 2H), 2.66-2.13 (m, 5H), 1.72 (s, 3H), 1.23 (dt, $J = 7.2, 1.3$ Hz, 3H).

Step II:

To a refluxing stirred solution of the 2-Allyl-4-methyl-pent-4-enoic acid ethyl ester (364 mg, 2.0 mmol) in CH_2Cl_2 (100 mL, 0.02 M solution) was added drop wise a solution of the tricyclohexylphosphine (1,3-Bis(2,4,6-trimethylphenyl)-4,5-dihydroimidazol-2-ylidene)(benzylidene)ruthenium (IV)dichloride (85 mg, 0.1 mmol) in CH_2Cl_2 (3.0 mL). After 50 min, the reaction mixture was cooled to room temperature, concentrated and purified on silica gel bond elute using EtOAc/hexane (1:20) as an eluent furnished the 3-Methyl-cyclopent-3-enecarboxylic acid ethyl ester (286 mg, 93% yield) as an oil. ^1H NMR (CDCl_3 , 400 MHz), 5.25 (brs, 1H), 4.17 (q, $J = 7.1$ Hz, 2H), 3.2-3.1 (m, 1H), 2.65-2.46 (m, 4H), 1.74 (s, 3H), 1.28 (t, $J = 7.1$ Hz, 3H).

Step III:

A solution of the 3-Methyl-cyclopent-3-enecarboxylic acid ethyl ester (255 mg, 1.65 mmol) in MeOH (4.0 mL) and 10% aq. NaOH (3.3 mL, 8.25 mmol) was heated at 50°C for 16 h, reaction mixture was cooled to room temperature, solvent was evaporated, diluted with water. The aqueous solution was washed with ether, and acidified with aq. 1 N HCl, extracted with ether. The ethereal solution was washed with brine and dried. Evaporation of the solvent furnished the 3-Methyl-cyclopent-3-enecarboxylic acid (200 mg, 97% yield). ^1H NMR (CDCl_3 , 400 MHz) 5.27 (brs, 1H), 3.26-3.17 (m, 1H), 2.7-2.55 (m, 4H), 1.74 (s, 3H).

Step IV:

The coupling of the 3-Isopropylamino-5-phenyl-thiophene-2-carboxylic acid methyl ester (82 mg, 0.3 mmol) and the 3-Methyl-

cyclopent-3-enecarboxylic acid (45 mg, 0.357 mmol) using PPh_3 (95.4 mg, 0.363 mmol) and NCS (48.5 mg, 0.363 mmol) furnished the 3-[Isopropyl-(3-methyl-cyclopent-3-enecarbonyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester (70 mg, 61%

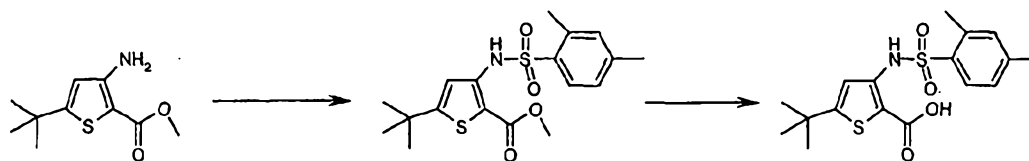
5 yield) ^1H NMR (CDCl_3 , 400 MHz 1:1 mixture of rotamers), 7.68-7.64 (m, 4H), 7.5-7.4 (m, 6H), 7.1 (s, 1H), 7.09 (s, 1H), 5.2 (s, 1H), 5.1 (s, 1H), 5.06-4.98 (m, 2H), 3.88 (s, 3H), 3.87 (s, 3H), 3.08-3.0 (m, 2H), 2.85-2.76 (m, 2H), 2.5-2.42 (m, 2H), 2.3-2.1 (m, 4H), 1.69 (s, 3H), 1.64 (s, 3H), 1.24 (d, $J = 6.7$ Hz, 3H), 1.23 (d, $J = 6.7$ Hz, 3H), 1.01 (d, $J = 6.9$ Hz, 3H), 1.007 (d, $J = 6.8$ Hz, 3H).

Saponification of 3-[Isopropyl-(3-methyl-cyclopent-3-enecarbonyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester (50 mg, 0.13 mmol) using $\text{LiOH}\cdot\text{H}_2\text{O}$ (22 mg) as previously described furnished the 3-[Isopropyl-(3-methyl-cyclopent-3-enecarbonyl)-amino]-5-phenyl-thiophene-2-carboxylic acid (30 mg, 62.5% yield) as a solid.

^1H NMR (CD_3OD , 400 MHz 1:1 mixture of rotamers) 7.73-7.70 (m, 4H), 7.47-7.35 (m, 6H), 7.29 (s, 1H), 7.27 (s, 1H), 5.16 (s, 1H), 5.08 (s, 1H), 4.9-4.8 (m, 2H), 3.15-3.05 (m, 2H), 2.76-2.65 (m, 2H), 2.42-2.12 (m, 6H), 1.65 (s, 3H), 1.61 (s, 3H), 1.25 (d, $J = 6.6$ Hz, 3H), 1.24 (d, $J = 6.6$ Hz, 3H), 1.03 (d, $J = 6.9$ Hz, 6H).

Example 16

5-tert-Butyl-3-(2,4-dimethyl-benzenesulfonylamino)-thiophene-2-carboxylic acid Compound #315 :



STEP I

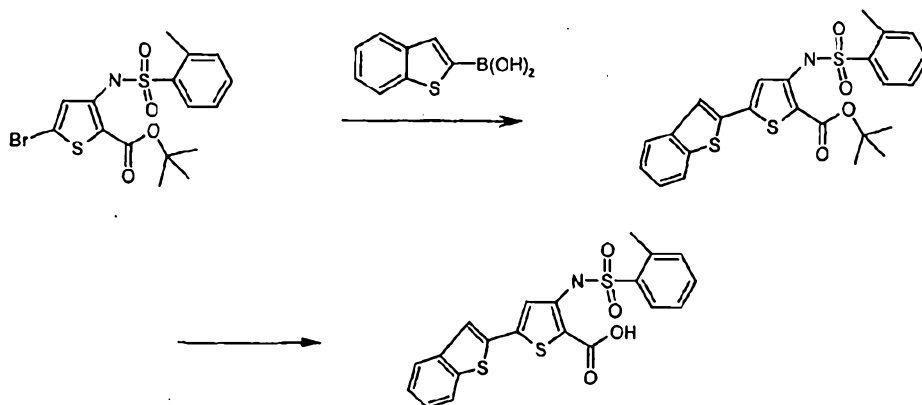
A mixture of 3-Amino-5-tert-butyl-thiophene-2-carboxylic acid methyl ester (106.5 mg, 0.5 mmol) and 2,4-dimethylsulfonyl chloride (156 mg, 0.75 mmol) in pyridine (1.5 mL) was heated at 72 °C for 16 h. The reaction mixture was diluted with EtOAc, washed with aq. 1N HCl, brine and dried. Evaporation of the solvent and purification of the residue on silica gel bond elute using EtOAc (1:20 to 1:10) as an eluent furnished the 5-tert-Butyl-3-(2,4-dimethyl-benzenesulfonylamino)-thiophene-2-carboxylic acid methyl ester (188 mg, 99% yield). ¹H NMR (CDCl₃, 400 MHz) 9.73 (s, 1H), 7.89 (d, *J* = 8.6 Hz, 1H), 7.04-7.08 (m, 2H), 7.03 (s, 1H), 3.82 (s, 3H), 2.62 (s, 3H), 2.33 (s, 3H), 1.28 (s, 9H).

STEP II

Hydrolysis of the 5-tert-Butyl-3-(2,4-dimethyl-benzenesulfonylamino)-thiophene-2-carboxylic acid methyl ester (55 mg, 0.14 mmol) using LiOH.H₂O (22 mg) as previously described provided the 5-tert-Butyl-3-(2,4-dimethyl-benzenesulfonylamino)-thiophene-2-carboxylic acid (36 mg, 70% yield) as a solid. ¹H NMR (CD₃OD, 400 MHz) 7.85 (d, *J* = 8.6 Hz, 1H), 7.14-7.10 (m, 2H), 7.0 (s, 1H), 2.56 (s, 3H), 2.31 (s, 3H), 1.27 (s, 9H).

Example 17

5-Benzo[b]thiophen-2-yl-3-(toluene-2-sulfonylamino)-thiophene-2-carboxylic acid Compound #230



STEP I

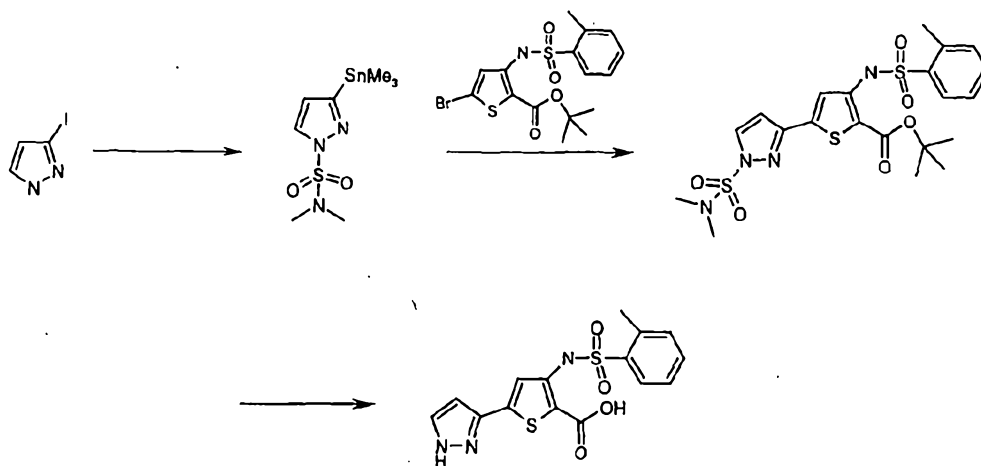
Suzuki coupling of 5-Bromo-3-(toluene-2-sulfonylamino)-thiophene-2-carboxylic acid tert-butyl ester (43 mg, 0.1 mmol) and benzo[b]thiophene-2-boronic acid (53.4 mg, 0.3 mmol) was carried out using $\text{Pd}(\text{PPh}_3)_4$ and Na_2CO_3 (as described in example 2) resulted in 5-Benzo[b]thiophen-2-yl-3-(toluene-2-sulfonylamino)-thiophene-2-carboxylic acid tert-butyl ester (27 mg, 55% yield). ^1H NMR (CDCl_3 , 400 MHz) 9.92 (s, 1H), 8.07 (d, $J = 7.8$ Hz, 1H), 7.79-7.71 (m, 2H), 7.45-7.24 (m, 7H), 2.7 (s, 3H), 1.56 (s, 9H).

STEP II

5-Benzo[b]thiophen-2-yl-3-(toluene-2-sulfonylamino)-thiophene-2-carboxylic acid tert-butyl ester was hydrolyzed to the acid using TFA as described for example 2 providing 5-Benzo[b]thiophen-2-yl-3-(toluene-2-sulfonylamino)-thiophene-2-carboxylic acid (24 mg, 99% yield). ^1H NMR ($\text{DMSO}-d_6$, 400 MHz) 10.19 (s, 1H), 8.0 (d, $J = 7.7$ Hz, 1H), 7.79-7.74 (m, 1H), 7.86 (s, 1H), 7.84-7.81 (m, 1H), 7.54 (t, $J = 7.7$ Hz, 1H), 7.53-7.36 (m, 4H), 7.32 (s, 1H), 2.58 (s, 3H).

Example 18

5-(1H-Pyrazol-3-yl)-3-(toluene-2-sulfonylamino)-thiophene-2-carboxylic acid Compound #170



Step I

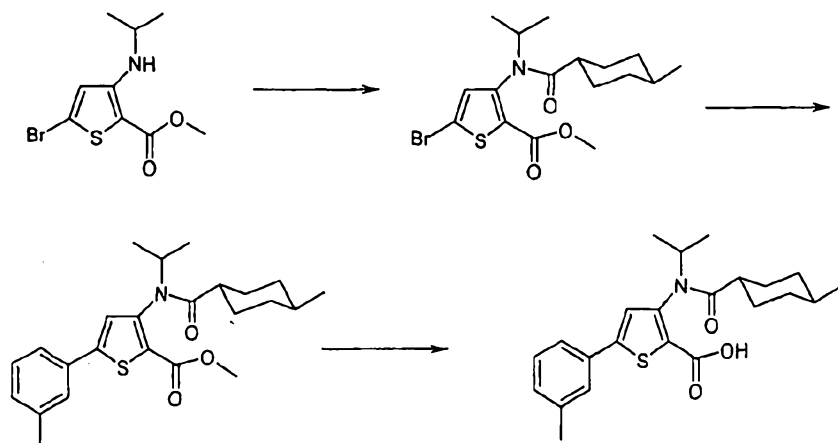
To a stirred solution of 5-Bromo-3-(toluene-2-sulfonylamino)-thiophene-2-carboxylic acid tert-butyl ester (43mg, 0.1mmol) in toluene (3.0 mL) was sequentially added a solution of Pd(PPh₃)₄ (12 mg, 0.01 mmol) in toluene (1.0 mL) and 3-Trimethylstannanyl-pyrazole-1-sulfonic acid dimethylamide (prepared according to *J. Med. Chem.* (1998), **41**, p-2019) (75 mg, 0.2 mmol, 2.0 eq), and heated the resulting overnight at 80°C. It was then cooled to room temperature, the solvent was evaporated and the crude was purified on preparative TLC using EtOAc/hexane (1:5). 5-(1-Dimethylsulfamoyl-1H-pyrazol-3-yl)-3-(toluene-2-sulfonylamino)-thiophene-2-carboxylic acid tert-butyl ester (35 mg, 66.5% yield) was isolated. ¹H NMR (CDCl₃, 400 MHz) 9.93 (s, 1H), 8.11 (d, *J* = 0.7 Hz, 1H), 8.02 (dd, *J* = 6.7, 1.32 Hz, 1H), 7.84 (d, *J* = 0.7 Hz, 1H), 7.45 (dt, *J* = 7.5, 1.3 Hz, 1H), 7.31 (t, *J* = 8.2 Hz, 2H), 7.26 (d, *J* = 1.0 Hz, 1H), 2.98 (s, 6H), 2.7 (s, 3H), 1.55 (s, 9H).

Step II

A reaction mixture of 5-(1-Dimethylsulfamoyl-1H-pyrazol-3-yl)-3-(toluene-2-sulfonylamino)-thiophene-2-carboxylic acid tert-butyl ester (10 mg, 0.019 mmol) and 4N HCl (0.3 mL) solution in dioxane in MeOH (0.3 mL) was stirred at room temperature 26 h. Reaction mixture was then diluted with water and extracted with EtOAc, concentrated and purified on preparative TLC using MeOH/CH₂Cl₂/AcOH (5:95:1) furnished the 5-(1H-Pyrazol-3-yl)-3-(toluene-2-sulfonylamino)-thiophene-2-carboxylic acid (4.5 mg, 65.2% yield). ¹H NMR (CD₃OD, 400 MHz) 7.99 (d, *J* = 7.9 Hz, 1H), 7.81 (s, 1H), 7.43 (t, *J* = 7.5, 1.3 Hz, 1H), 7.42-7.26 (m, 2H), 7.19 (s, 1H), 2.69 (s, 3H).

Example 19

3-Isopropyl-[(4-methyl-cyclohexanecarbonyl)-amino]-5-m-tolyl-thiophene-2-carboxylic acid Compound #448



STEP I

Trans-4-methyl-cyclohexanecarbonyl chloride was prepared by heating to reflux trans-4-methyl-cyclohexanecarboxylic acid (5g, 0.035 mmol) in thionylchloride (5.0 ml) for 2h followed by purification of the corresponding acyl chloride under reduced pressure in a Kugel-Rhorr apparatus collecting the fraction distilling at 95 °C yielding 5.1 g of the desired material which was used in the next step without further purification. This acyl chloride (1.5 ml, aprox. 10 mmol) was dissolved along with 5-Bromo-3-isopropylamino-thiophene-2-carboxylic acid methyl ester (2 g, 7.12 mmol) in anhydrous dichloroethane (2 mL) and heated at 80 °C (closed vial) for 12h. The solvents were evaporated, the resulting crude material was dissolved in methanol and left 30 min. at room temperature, concentrated and purified via flash chromatography on silica gel using a 5% EtOAc 95% hexanes mixture of eluents, in this manner 600 mg (21%) of 5-Bromo-3-[isopropyl-(4-methyl-cyclohexanecarbonyl)-amino]-thiophene-2-carboxylic acid methyl ester the was isolated. ¹H NMR(CDCl₃, 300 MHz): 6.78 (s, 1H), 4.93 (m, 1H), 3.69 (s, 3H), 2.00-1.20 (m, 8H), 1.14 (d, 3H), 0.93 (d, 3H), 0.81 (d, 3H), 0.72-0.70 (m, 2H).

STEP II

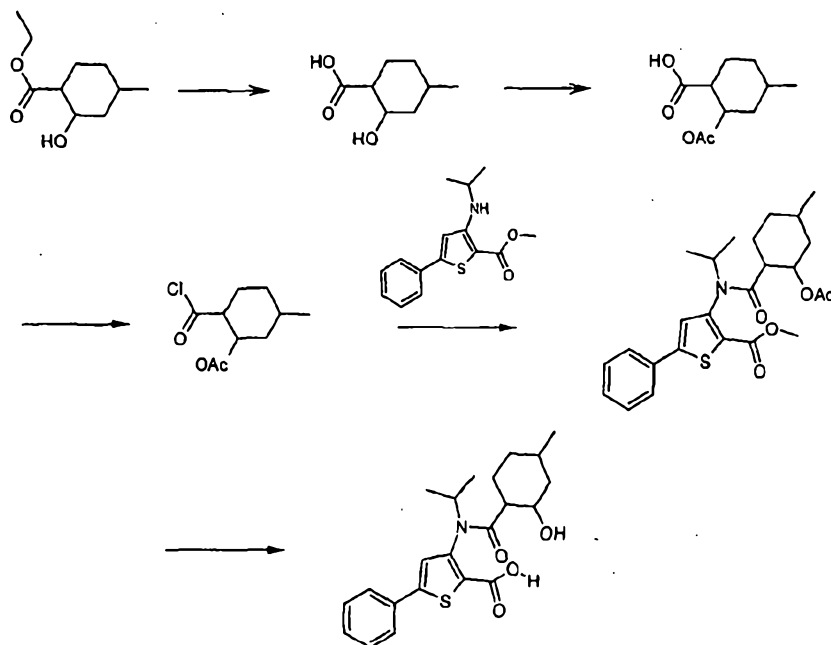
To a degassed solution of 5-Bromo-3-[isopropyl-(4-methyl-cyclohexanecarbonyl)-amino]-thiophene-2-carboxylic acid methyl ester (100 mg, 0.249 mmol) and 3-methyl boronic acid (38 mg, 0.279 mmol) in a mixture of DME (6 mL) and 2M aqueous Na₂CO₃ (3 mL), Pd(PPh₃)₄ (12 mg) was added and the reaction mixture was stirred at reflux conditions for 12h under a N₂ atmosphere. The reaction mixture was diluted with ethyl acetate and water. The organic layer was separated, dried (Na₂SO₄), concentrated. The residue was purified by column chromatography using ethyl acetate and hexane (1:3) as eluent. 35 mg (34%) of 3-Isopropyl-[(4-methyl-cyclohexanecarbonyl)-amino]-5-m-tolyl-thiophene-2-carboxylic acid methyl ester was isolated. ¹H NMR (CDCl₃, 400 MHz): 7.45 (bs, 2H), 7.36 (t, 1H), 7.23 (m, 1H), 7.01 (s, 1H), 4.99 (m, 1H), 3.83 (s, 3H), 2.41 (s, 3H), 2.01-0.61 (m, 20H).

Step III

3-Isopropyl-[(4-methyl-cyclohexanecarbonyl)-amino]-5-m-tolyl-thiophene-2-carboxylic acid methyl ester (30 mg, 0.073 mmol) was taken in a mixture of THF:MeOH:H₂O (3:2:1, 3 mL) and then added 1N aqueous solution of LiOH.H₂O (0.44 mL, 0.438 mmol). The reaction mixture was stirred at room temperature for 12 h. Solvents were removed and the residue was partitioned between water and ethyl acetate. The aqueous layer was acidified using 10 % KHSO₄ solution. The organic layer was separated, dried (Na₂SO₄) and concentrated. The residue was purified by preparative TLC using chloroform:methanol:acetic acid (9:1:0.1) to obtain 3-Isopropyl-[(4-methyl-cyclohexanecarbonyl)-amino]-5-m-tolyl-thiophene-2-carboxylic acid (15 mg, 52%) as a white solid. ¹H NMR (CDCl₃, 400 MHz): (s, 2H), 7.38 (t, 1H), 7.24 (m, 1H), 7.08 (s, 1H), 5.01 (s, 1H), 2.42 (s, 3H), 2.10-0.62 (m, 20H). ESI⁻ (M-H): 398.

Example 20

(1R,2S,4R)-3-[Isopropyl-(2-hydroxy-4-methyl-cyclohexanecarbonyl)-amino]-5-phenyl-thiophene-2-carboxylic acid
Compound #402



5

(1R,2S,4R)-2-Hydroxy-4-methyl-cyclohexanecarboxylic acid methyl ester was prepared as described in J. Org. Chem, (1993), 58, pp.6255-6265. NMR ^1H (CDCl_3 , 400 MHz): 4,26 ppm (s, 1H); 4,19-4,13 ppm (m, 2H); 3,16 ppm (s, 1H); 2,35-2,29 ppm (m, 1H); 1,92-1,74 ppm (m, 5H); 1,31-1,24 ppm (m, 3H); 1,08-1,01 ppm (m, 1H); 0,96-0,92 ppm (m, 1H); 0,88 ppm (d, 3H).

STEP I

To a solution of (1R,2S,4R)-2-Hydroxy-4-methyl-

cyclohexanecarboxylic acid methyl ester (450 mg, 2.42 mmol) in methanol (12 ml) was added a 2.5 M solution of sodium hydroxide (9.7 ml, 24.2 mmol). The reaction mixture was stirred for 4 h at 50 °C. Then, the solvents were removed and the residue was partitioned between 20 ml of H_2O acidified to pH 4 and 20 ml of EtOAc. The organic layer was separated and the aqueous phase was washed with ethyl acetate (2 X 20 ml). The combined ethyl acetate layers were dried (Na_2SO_4) and concentrated to obtain 313 mg (82 %) of (1R,2S,4R)-2-Hydroxy-4-methyl-cyclohexanecarboxylic

acid. NMR ^1H (CDCl_3 , 400 MHz): 4,34 ppm (s, 1H); 2,43-2,39 ppm (m, 1H); 1,96-1,76 ppm (m, 5H); 1,14-1,08 ppm (m, 1H); 1,02-0,93 ppm (m, 1H); 0,90 ppm (d, 3H).

5 Step II

To a solution of (1R,2S,4R)-2-Hydroxy-4-methyl-cyclohexanecarboxylic acid (162 mg, 1.02 mmol) in dichloromethane (5 ml) was added pyridine (495 μl , 6.12 mmol) followed by acetic anhydride (385 μl , 4.08 mmol). The reaction
10 mixture was stirred for 20 h at room temperature. Then, the solvents were removed and 10 ml of 3N HCl solution was added. This mixture was stirred for 30 minutes and then a saturated solution of NaHCO_3 was slowly added until pH = 9-10. This solution was then extracted with ethyl acetate (2 X 5 ml). The
15 aqueous phase was then acidified with a 10% HCl solution and extracted with ethyl acetate (3X5 ml). The following ethyl acetate layers were combined, dried (Na_2SO_4) and concentrated to obtain 109 mg (53 %) of (1R,2S,4R)-2-Acetoxy-4-methyl-cyclohexanecarboxylic acid. NMR ^1H (CDCl_3 , 400 MHz): 5,45 ppm
20 (s, 1H); 2,46-2,42 ppm (m, 1H); 2,02 ppm (s, 3H); 2,02-1,96 ppm (m, 1H); 1,91-1,76 ppm (m, 3H); 1,70-1,61 ppm (m, 1H); 1,16-1,08 ppm (m, 1H); 0,99-0,88 ppm (m, 1H); 0,87 ppm (d, 3H).

Step III

25 To a solution of (1R,2S,4R)-2-Acetoxy-4-methyl-cyclohexanecarboxylic acid (109 mg, 0.54 mmol) in dichloromethane (2.7 ml) was added oxalyl chloride (545 μl , 1.09 mmol) followed by 1 drop of dimethylformamide. The reaction mixture was stirred for 4 h at room temperature. The solvents
30 were then removed to obtain 119 mg (99%) of (1R,2S,4R)-2-Acetoxy-4-methyl-cyclohexanecarboxylic acid chloride.

Step IV

To a solution of 3-Isopropylamino-5-phenyl-thiophene-2-carboxylic
35 acid methyl ester (136 mg, 0.50 mmol) in 1,2-dichloroethane (1.0

ml) was added (1R,2S,4R)-2-Acetoxy-4-methyl-cyclohexanecarboxylic acid chloride (119 mg, 0.54 mmol) dissolved in 1,2-dichloroethane (0.6 ml) followed by PPh₃ (136 mg, 0.52 mmol). The resulting solution was stirred for 20 h at 90 °C and then cooled to room temperature. It was then diluted with ethyl acetate (10 ml) and a solution of saturated NaHCO₃ (10 ml). The aqueous phase was separated and washed with ethyl acetate (2x10 ml) and the combined organic layers were dried (Na₂SO₄), filtered and concentrated. The residue was purified by flash chromatography (0% to 25% EtOAc/Hexane) to obtain 110 mg (45%) of (1R,2S,4R)-3-[Isopropyl-(2-Acetoxy-4-methyl-cyclohexanecarbonyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester. NMR ¹H (CDCl₃, 400 MHz): 1.5:1.0 mixture of rotamers 7.73-7.70 ppm (m, 2H, H_{aro}); 7.69-7.63 ppm (m, 1H, H_{aro}); 7.51-7.41 ppm (m, 4H, H_{aro}); 7.13 ppm (s, 0.6H, H_{aro}, major rotamer); 5.79 ppm (s, 0.4H, minor rotamer); 5.21 ppm (s, 0.6H, major rotamer); 4.95-4.88 ppm (m, 1H); 3.88 ppm (s, 1.8H, major rotamer); 3.87 ppm (s, 1.2H, minor rotamer); 2.40-2.36 ppm (m, 0.6H, major rotamer); 2.11 ppm (s, 3H); 1.78-0.77 ppm (m, 16H).

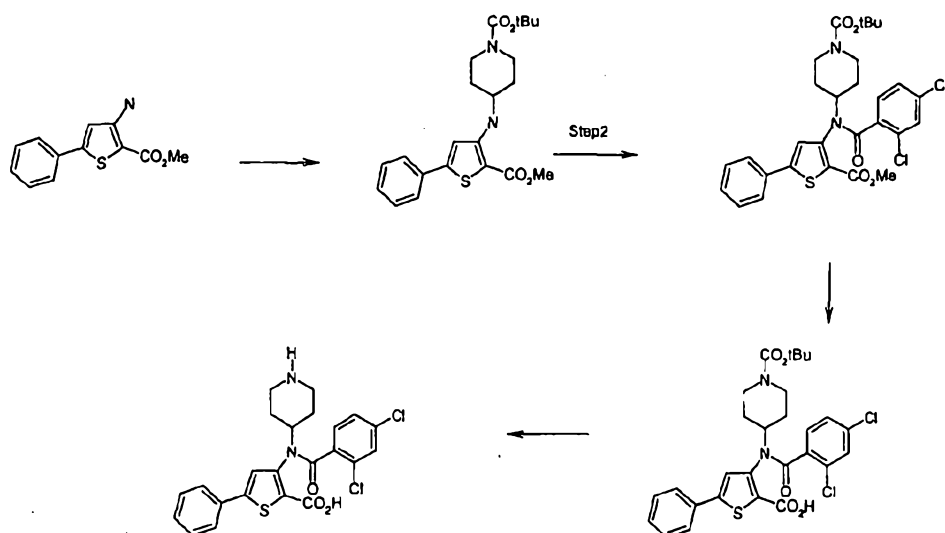
Step V

(1R,2S,4R)-3-[Isopropyl-(2-Acetoxy-4-methyl-cyclohexanecarbonyl)-amino]-5-phenyl-thiophene-2-carboxylic acid methyl ester (36 mg, 0.17 mmol) was dissolved in a mixture of dioxane:H₂O (4:1) (700 µl) and then 470 µl of LiOH 1N was added to it. After 3 h at 50 °C the reaction mixture was cooled to room temperature and the solvents were removed. The residue was then partitioned between 10 ml of H₂O acidified to pH 4 and 10 ml of EtOAc. The organic layer was separated and the aqueous phase was washed with ethyl acetate (2 X 10 ml). The combined ethyl acetate layers were dried (Na₂SO₄), concentrated and the residue was purified by preparative chromatography to obtain 9 mg (29 %) of (1R,2S,4R)-3-[Isopropyl-(2-hydroxy-4-methyl-cyclohexanecarbonyl)-amino]-5-phenyl-thiophene-2-carboxylic acid. NMR ¹H (CDCl₃, 400 MHz): 3:2 mixture of rotamers 7.76-

7,73 ppm (m, 2H, H_{aro}); 7,50-7,38 ppm (m, 3H, H_{aro}); 7,36 ppm (s, 1H, H_{aro}); 4,93-4,87 ppm (m, 1H); 4,25 ppm (s, 0.70H, major rotamer); 3,97 ppm (s, 0.3H, minor rotamer); 2,35-2,28 ppm (m, 1H); 1,99-1,53 ppm (m, 5H); 1,28 ppm (d, 0,6H, minor rotamer);
 5 1,25 ppm (d, 1,4H, major rotamer); 1,06-1,03 ppm (m, 3H), 0,96-0,72 ppm (m, 1H); 0,79 ppm (d, 3H); 0,67-0,56 ppm (m, 1H).

Example 21

3-[(2,4-Dichloro-benzoyl)-piperidin-4-yl-amino]-5-phenyl-
 10 thiophene-2-carboxylic acid hydrochloride salt compound #368



Step I

15 A suspension of 3-amino-5-phenyl-thiophene-2-carboxylic acid methyl ester (745 mg, 3.2 mmol) in dry THF (1.3 ml), at 21 °C, under nitrogen, was treated with tert-butyl 4-oxo-1-piperidine carboxylate (673 mg, 3.2 mmol), followed by dibutyltin dichloride (19 mg, 0.064 mmol, 0.02 eq.). After 5 min the
 20 reaction was treated with phenyl silane (435 µL, 380 mg, 3.52 mmol, 1.1 eq). The mixture was left to stir for 74h when a clear solution resulted. The reaction was stripped off solvent to leave a thick bright yellow gum (1.59 g). The crude material was purified by column chromatography using (CH₂Cl₂ :Hexane :EtOAc)
 25 = 15 : 5 :1 as eluent to provide 4-(2-Methoxycarbonyl-5-phenyl-

thiophen-3-ylamino)-piperidine-1-carboxylic acid tert-butyl ester as a yellow foam (713 mg, 54%). ^1H NMR (CDCl_3 , 400 MHz) 7.63-7.60 (m, 2H), 7.74-7.36 (m, 3H), 6.90-6.84 (bs, 1H), 6.84 (s, 1H), 3.97-4.01 (m, 2H), 3.80 (s, 3H), 3.48 (bs, 1H), 3.06-2.99 (m, 2H), 2.03-1.99 (m, 2H), 1.51-1.48 (m, 2H), 1.47 (bs, 9H)

Step II

4-(2-Methoxycarbonyl-5-phenyl-thiophen-3-ylamino)-piperidine-1-carboxylic acid tert-butyl ester (200 mg, 0.48 mmol) was treated with 2,4 dichlorobenzoylchloride (202 μL , 302 mg, 1.44 mmol, 3 eq) under previously described conditions (e.g. Example 14) to provide, after column chromatography using (CH_2Cl_2 : Hexane : EtOAc = 15 : 5 : 1) as eluent, 4-[(2,4-Dichloro-benzoyl)-(2-methoxycarbonyl-5-phenyl-thiophen-3-yl)-amino]-piperidine-1-carboxylic acid tert-butyl ester as a pale yellow foam (165 mg, 58%), ^1H NMR (CDCl_3 , 400 MHz) 7.54-7.51 (m, 2H), 7.45-7.39 (m, 3H), 7.27-7.25 (m, 2H), 7.17 (d, $J = 1.96\text{Hz}$, 1H), 7.06 (dd, $J = 1.92\text{Hz}$, $J = 8.34\text{Hz}$, 1H), 4.86-4.92 (m, 1H), 4.11-4.21 (m, 2H), 3.89 (s, 3H), 2.82-2.89 (m, 2H), 2.17-2.20 (m, 1H), 1.89-1.92 (m, 1H), 1.49-1.61 (m, 1H), 1.40 (bs, 9H), 1.19-1.25 (m, 1H)

Step III

A suspension of 4-[(2,4-Dichloro-benzoyl)-(2-methoxycarbonyl-5-phenyl-thiophen-3-yl)-amino]-piperidine-1-carboxylic acid tert-butyl ester (160 mg, 0.27 mmol) above in dioxane: water (4 : 1, 3 ml) was treated with lithium hydroxide (2M aqueous solution, 41 μL , 341 mg, 0.814 mmol, 3 eq) and the reaction allowed to stir overnight for 18h. The reaction was stripped-off solvent and the residue partitioned between EtOAc : water (4 : 1). The aqueous phase was separated and extracted several times, with EtOAc, following acidification to pH 5.5 with 0.1N HCl. The combined organic extract was evaporated to a solid. The solid was taken into EtOAc and the above acid wash repeated to give, after drying and evaporation, 4-[(2-Carboxy-5-phenyl-thiophen-3-yl)-(2,4-

dichloro-benzoyl)-amino]-piperidine-1-carboxylic acid tert-butyl ester as a colourless solid (128 mg, 91%), ¹H NMR (Acetone, 400 MHz) 7.75-7.70 (m, 1H), 7.64 (s, 1H), 7.52-7.40 (m, 3H), 7.52 (d, J = 1.98 Hz, 1H), 7.21 (dd, J = 1.96 Hz, J = 8.19 Hz, 1H), 4.80-4.71 (m, 1H), 4.26-4.01 (m, 2H), 2.71- 2.30 (bs, 3H), 2.25-2.17 (m, 1H), 1.82-1.69 (m, 1H), 1.40 (bs, 9H), 1.33-1.24 (m, 1H).

Step IV

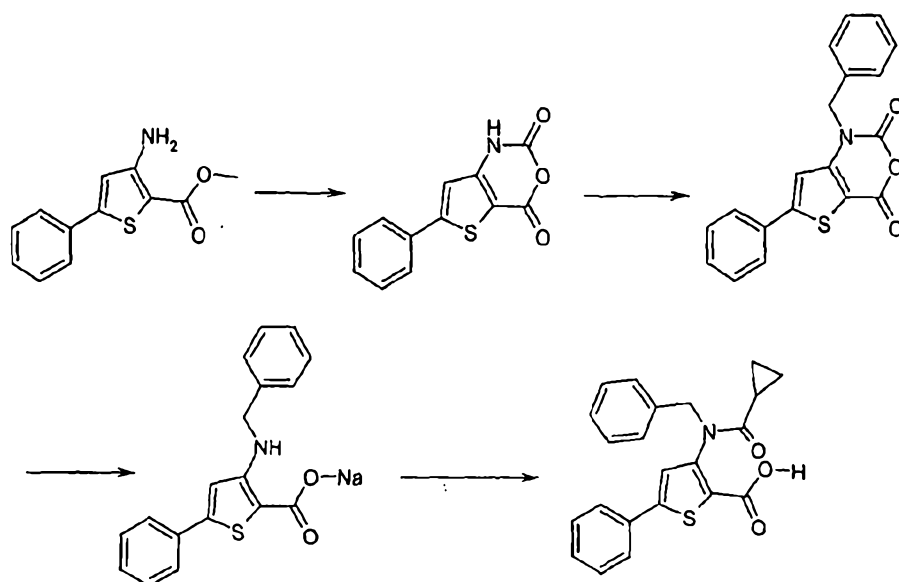
A solution of 4-[(2-Carboxy-5-phenyl-thiophen-3-yl)-(2,4-dichloro-benzoyl)-amino]-piperidine-1-carboxylic acid tert-butyl ester (240 mg, 0.42 mmol) in dioxane (4 ml) at 21 °C under nitrogen was treated with anhydrous 4M HCl (3 ml, 12.6 mmol, 30 eq). After 4h the reaction was stripped off solvent and the residue triturated with ether to give 3-[(2,4-Dichloro-benzoyl)-piperidin-4-yl-

amino]-5-phenyl-thiophene-2-carboxylic acid as a pale yellow powder (214 mg, 100 %) ¹H NMR (Acetone, 400 MHz) 7.76-7.73 (m, 2H), 7.64 (s, 1H), 7.45-7.38 (m, 3H), 7.30 (bs, 1H), 7.28-7.24 (m, 1H), 4.93-4.84 (m, 1H), 3.56-3.49 (m, 2H), 3.25-3.14 (m, 2H), 3.05-2.55 (bs, 1H), 2.50-2.37 (m, 2H), 2.13-1.83 (m, 1H).

Similarly prepared were Compound #366 , Compound #553 , Compound #543

Example 22

3-(Benzyl-cyclopropanecarbonyl-amino)-5-phenyl-thiophene-2-carboxylic acid. Compound #454



Step I

A solvent mixture of THF/MeOH/H₂O (3:2:1) was added to 3.04 g of methyl (3-amino-5-phenyl)thiophene-2-carboxylate (13 mmol) and 1.64 g of lithium hydroxide monohydrate (39 mmol). The mixture was refluxed for 8 hours and concentrated *in vacuo*. The crude material was taken in 100 ml of water, washed with ethyl acetate (2 x 100 ml) and transferred into a multineck flask. A 20% phosgene solution in toluene (11 ml, 39 mmol) was added dropwise at 0°C. A precipitate was then collected by filtration and sequentially washed by trituration with a saturated solution of bicarbonate, water, acetone and diethyl ether. 2.52 g (79%) of 6-phenyl-1H-thieno[3,2-d][1,3]oxazine-2,4-dione were isolated as a white solid. NMR ¹H (DMSO D₆, 400 MHz): 7.79-7.76 ppm (m, 2H, H_{aro}); 7.52-7.47 ppm (m, 3H, H_{aro}); 7.25 ppm (s, 1H, H_{azole}); 0.4 ppm (s, 1H, NH).

Step II

A solution of 6-phenyl-1H-thieno[3,2-d][1,3]oxazine-2,4-dione (1 g, 4.1 mmol) and anhydrous sodium carbonate (477 mg, 4.5 mmol) diluted in 15 ml of anhydrous dimethylacetamide was stirred for one hour under nitrogen before adding benzyl bromide (785 mg, 4.5 mmol). The mixture was stirred overnight at room temperature. 912 mg (66.3%) of 1-benzyl-6-phenyl-1H-thieno[3,2-

d][1,3]oxazine-2,4-dione were obtained as a pale yellow solid after filtration and washing the precipitate with acetone and pentane. NMR ^1H (DMSO D_6 , 400 MHz): 7.8-7.76 ppm (m, 3H, H_{aro}); 7.51-7.45 ppm (m, 3H, H_{aro}); 7.43-7.41 ppm (m, 2H, H_{aro}); 7.35-7.3 ppm (m, 2H, H_{aro}); 7.28-7.24 ppm (m, 1H, H_{aro}); 5.22 ppm (s, 1H, NCH_2).

Step III

To a solution of 1-benzyl-6-phenyl-1H-thieno[3,2-d][1,3]oxazine-2,4-dione (880 mg, 2.62 mmol) were successively added 32 ml of dioxane and 7.87 ml of NaOH 1N aqueous solution. The mixture was vigorously stirred for 2 h and then the solvents were concentrated *in vacuo*. Dichloromethane was added to the crude material and sodium 3-benzylamino-5-phenyl-thiophene-2-carboxylate (1.07 g, 100%) precipitated as a pale yellow solid. NMR ^1H (DMSO D_6 , 400 MHz): 7.76 ppm (t, 1H, $J = 6.4$ Hz, NH); 7.53-7.51 ppm (m, 2H, H_{aro}); 7.33-7.26 ppm (m, 6H, H_{aro}); 7.23-7.16 ppm (m, 2H, H_{aro}); 7.07 ppm (s, 1H, H_{azole}); 4.36 ppm (d, 2H, $J = 6.4$ Hz, NHCH_2).

Step IV

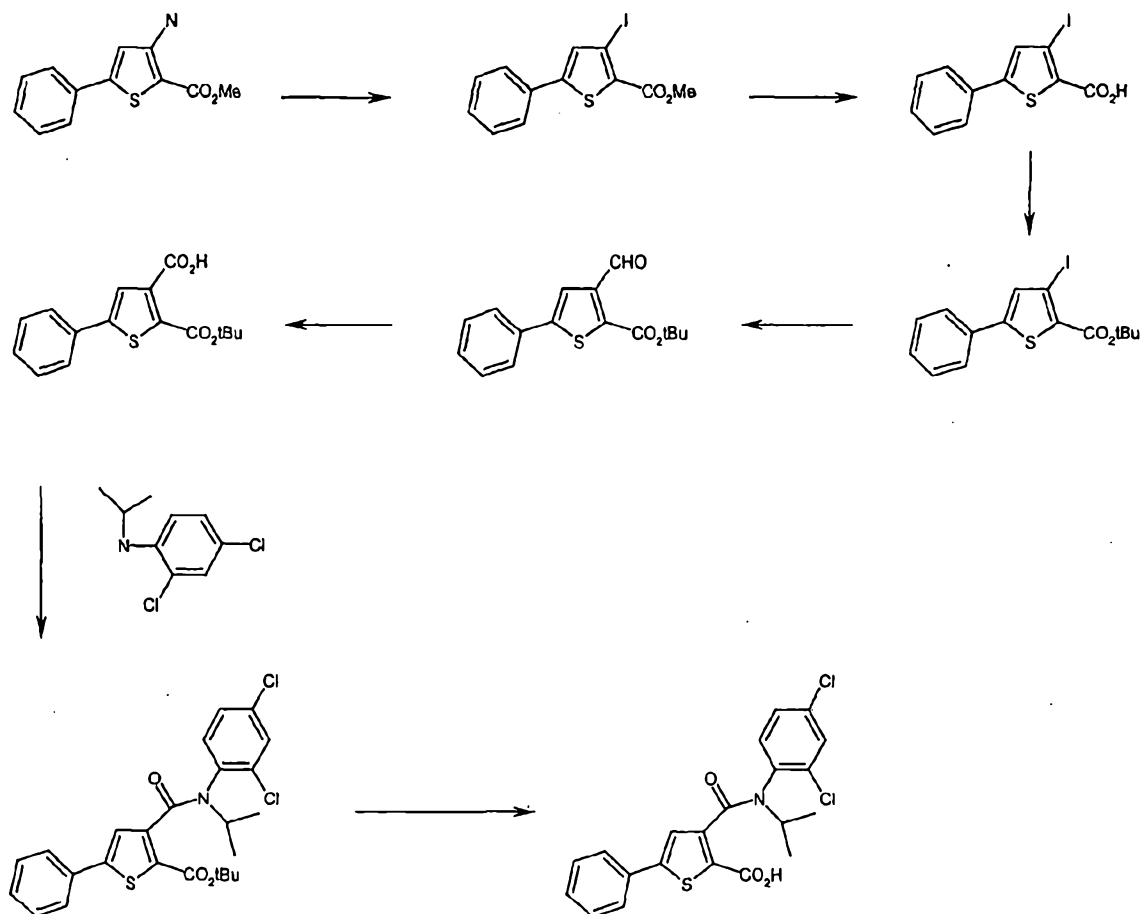
To a solution of sodium 3-benzylamino-5-phenyl-thiophene-2-carboxylate (41.1 mg, 0.1 mmol) was added 32 mg (0.3 mmol) of cyclopropanecarbonyl chloride, 1.5 ml of dioxane and 0.5 ml of water. The mixture was stirred overnight at room temperature and concentrated *in vacuo*. A 4N hydrogen chloride solution in dioxane (1 ml) was added and the mixture was stirred for one hour at room temperature. The mixture was again concentrated and the crude material was purified by reverse phase HPLC giving access to 11.9 mg (31.5%) of 3-(benzyl-cyclopropanecarbonylamino)-5-phenyl-thiophene-2-carboxylic acid as a pale yellow solid. NMR ^1H (DMSO D_6 , 400 MHz): 7.56-7.54 ppm (m, 2H, H_{aro}); 7.39-7.13 ppm (m, 10H, H_{aro} , H_{azole} and COOH); 5.27 ppm (d, 1H, $J = 15.2$ Hz); 4.48 ppm (d, 1H, $J = 15.2$ Hz); 1.49 ppm (m, 1H); 0.77 ppm (m, 2H); 0.61 ppm (m, 2H).

The following compounds were prepared in a similar manner: '

Compound #172 , Compound #173 , Compound #175 , Compound #186 ,
Compound #187 , Compound #188 , Compound #241 , Compound #247 ,
5 Compound #251 , Compound #252 , Compound #253 , Compound #254 ,
Compound #255 , Compound #256 , Compound #257 , Compound #276 ,
Compound #277 , Compound #278 , Compound #279 , Compound #280 ,
Compound #281 , Compound #330 , Compound #334 , Compound #335 ,
Compound #336 , Compound #339 , Compound #340 , Compound #341 ,
10 Compound #342 , Compound #343 , Compound #344 , Compound #345 ,
Compound #347 , Compound #349 , Compound #350 , Compound #351 ,
Compound #352 , Compound #353 BCH-23932, Compound #354 ,
Compound #384 , Compound #385 , Compound #386 , Compound #388 ,
Compound #389 , Compound #390 , Compound #391 , Compound #392 ,
15 Compound #393 , Compound #394 , Compound #397 , Compound #398 ,
Compound #399 , Compound #400 , Compound #401 .

Example 23

3-[(2,4-Dichloro-phenyl)-isopropyl-carbamoyl]-5-phenyl-
20 thiophene-2-carboxylic acid



Step I

5 3-Iodo-5-phenyl-thiophene-2-carboxylic acid methyl ester

A suspension of 3-Amino-5-phenyl-thiophene-2-carboxylic acid methyl ester (10 g, 43 mmol) in anhydrous benzene (200 ml), at 21 °C, under N₂, was treated with t-butyl nitrite (21.8 g, 86 mmol) and the dark mixture cooled to 0 °C and treated dropwise, over 15 min, with iodine (21.8 ml, 184 mmol) . After 30 min at 0 °C, the solution was allowed to warm-up to ambient temperature and stirred for 2h. The reaction mixture was then poured into water (300 ml) and stirred vigorously for 15 min. The organic phase was separated and washed several times with 20% sodium thiosulfate (4x100 ml) . The resulting emulsion was filtered through celite. The celite pad was washed with EtOAc and the combined filtrate and washings were washed with more sodium

thiosulfate (100 ml) to give an orange solution which was washed with brine and dried. Evaporation of the solvent afforded an oil (7.4 g). The crude oil was purified by biotage flash chromatography using Hexane/CH₂Cl₂/EtOAc (20/2/1) as eluent to give 4.42g (29%) of 3-Iodo-5-phenyl-thiophene-2-carboxylic acid methyl ester as a pale yellow oil. NMR ¹H (CDCl₃, 400 MHz,) : 7.62-7.57 (m, 2H); 7.58 (s, 1H); 7.50-7.36 (m, 3H); 3.91 (s, 3H)

Step II

10 3-Iodo-5-phenyl-thiophene-2-carboxylic acid

A solution of 3-Iodo-5-phenyl-thiophene-2-carboxylic acid methyl ester (4.4 g, 12.78 mmol) in dioxane/water 4/1 (50 ml), at 21 °C, under N₂, was treated with lithium hydroxyde (2N, 19.3 ml, 38 mmol) and the solution left to stir for 21.5 h. The reaction mixture was evaporated to dryness and the residue partitioned between EtOAc (75 ml) and water (25 ml) and acidified with 2N HCl to pH 5.5. The aqueous phase was separated and extracted with EtOAc (3x50 ml). The combined organic extract were washed with brine, dried and evaporated to give 4.12 g (97%) of 3-Iodo-5-phenyl-thiophene-2-carboxylic acid as a pale yellow solid. NMR ¹H (CD₃OD, 400 MHz) : 7.69-7.67 (m, 2H); 7.55 (s, 1H); 7.46-7.39 (m, 3H).

25 Step III

3-Iodo-5-phenyl-thiophene-2-carboxylic acid tert-butyl ester

A suspension of magnesium sulfate (4.61 g, 38.32 mmol) in dichloromethane (37 ml) at 21 °C, under N₂, was treated with conc H₂SO₄ (510 µl, 9.58 mmol). After 15 min solid 3-Iodo-5-phenyl-thiophene-2-carboxylic acid (3.7 g, 9.58 mmol) was added followed by t-butanol (4.55 ml, 47.9 mmol) and the flask was stoppered and left over-night for 19.5 h. The reaction mixture was treated with saturated bicarbonate aqueous solution, and filtered. The solid was washed with CH₂Cl₂ and the filtrate dried

and concentrated to an oil. The crude material was purified by flash chromatography using Hexane/CH₂Cl₂ (3:1) as eluent to give 1.63 g (44%) of 3-Iodo-5-phenyl-thiophene-2-carboxylic acid tert-butyl ester as a colorless solid. NMR ¹H (CDCl₃, 400 MHz) 7.61-7.59 (m, 2H); 7.43-7.35 (m, 3H), 7.25 (s, 1H), 1.60 (bs, 9H).

Step IV

3-Formyl-5-phenyl-thiophene-2-carboxylic acid tert-butyl ester

10

A solution of 3-Iodo-5-phenyl-thiophene-2-carboxylic acid tert-butyl ester (1.41 g, 3.65 mmol) in dry THF (37 ml) at -78°C, under nitrogen, was treated dropwise, over 5 min with n-butyl lithium (4.8 ml, 7.66 mmol). The reaction gradually darkened to a red-brown color. After 15 min at -78 °C dimethylformamide (1.7 ml, 21.9 mmol) was added dropwise over 7 min. The dark solution was allowed to stir for 2 h then quenched with saturated NH₄Cl solution (10 ml) and allowed to reach 21°C. The aqueous phase was separated and extracted with EtOAc (3x50 ml). The combined organic extracts were evaporated and the residue taken into EtOAc and washed with water, brine, dried and concentrated to give 1.14 g of a brownn oil. The crude material was purified by flash chromatography using Hexane/CH₂Cl₂ (1/1) as eluent to provide 303 mg (28%) of 3-Formyl-5-phenyl-thiophene-2-carboxylic acid tert-butyl ester as a colorless solid. NMR ¹H : (CDCl₃, 400 MHz) : 10.62 (s, 1H); 7.78 (s, 1H); 7.64-7.62 (m, 2H); 7.48-7.38 (m, 3H); 1.62 (bs, 9H).

Step V

30 5-Phenyl-thiophene-2,3-dicarboxylic acid 2-tert-butyl ester

A solution of 3-Formyl-5-phenyl-thiophene-2-carboxylic acid tert-butyl ester (300 mg, 1.04 mmol) in dry THF (20 ml), at 0 °C, under nitrogen, was treated with methyl sulfide (10% w/w in THF, 3.8 ml, 5.2 mmol) followed by sodium dihydrogenphosphate (30%

aqueous solution, 9.56 ml, 2.05 mmol). After 0.5 h, the solution was treated with sodium chlorite (30% w/w aqueous solution, 1.9 ml, 2.08 mmol) added over 1 min via a syringe. The pale yellow solution was stirred for 1.5 h at 0 °C, then diluted
5 with water (20 ml) and extracted with EtOAc (4x 40 ml). The aqueous phase was separated, extracted with more EtOAc (40 ml) and the combined extracts were washed with brine dried and concentrated to give 316 mg (100 %) of 5-Phenyl-thiophene-2,3-dicarboxylic acid 2-tert-butyl ester as a pale brown solid. NMR
10 ¹H (CD₃CO; 400 MHz): 7.87 (s, 1H); 7.83-7.81 (m, 2H); 7.17-7.53 (m, 3H); 1.65 (bs, 9H).

Step VI

3-[(2,4-Dichloro-phenyl)-isopropyl-carbamoyl]-5-phenyl-
15 thiophene-2-carboxylic acid tert-butyl ester

A solution of 5-Phenyl-thiophene-2,3-dicarboxylic acid 2-tert-butyl ester (40 mg, 0.13 mmol) in CH₂Cl₂ (1.3 ml), under nitrogen, at 0°C, was treated with diisopropylethylamine (27 μ L, 0.16 mmol) followed by dimethylformamide (10 μ L, 0.13 mmol) and
20 oxalyl chloride (170 μ L, 0.34 mmol). Slight effervescence was observed. The reaction was kept at 0 °C for 30 min before being treated with (2,4-Dichloro-phenyl)-isopropyl-amine (described previously) (79 mg, 0.39 mmol). The reaction was allowed to
25 reach 21 °C and then placed in a bath at 90 °C for 15 h. Solvent was removed to leave a pale brown gum (144 mg). The crude material was purified on bond-elute using Hexane/CH₂Cl₂/EtOAc (12.5/2/1) as eluent to give 39 mg, (62%) of 3-[(2,4-Dichloro-phenyl)-isopropyl-carbamoyl]-5-phenyl-thiophene-2-carboxylic
30 acid tert-butyl ester as a pale brown solid. NMR ¹H (CDCl₃; 400 MHz) 7.50-7.48 (m, 2H); 7.38-7.25 (m, 6H); 7.10-7.03 (m, 1H); 5.05 (quint, J = 6.88 Hz, 1H); 1.57 (bs, 9H); 1.40 (d, J = 6.88 Hz, 3H); 1.12 (d, J = 6.88 Hz, 3H)

35 Step VII

3-[(2,4-Dichloro-phenyl)-isopropyl-carbamoyl]-5-phenyl-thiophene-2-carboxylic acid

A solution of 3-[(2,4-Dichloro-phenyl)-isopropyl-carbamoyl]-5-phenyl-thiophene-2-carboxylic acid tert-butyl ester (37 mg, 0.08 mmol) in CH₂Cl₂ (0.2 ml) at room temperature, under nitrogen was treated with trifluoroacetic acid (0.8 ml). After 1 h the reaction was concentrated the residue was taken into EtOAc and washed sequentially with 2N HCl (2x15 ml), water, brine dried and evaporated to a foam (33 mg). The foam was redissolved in EtOAc and above acidic wash was repeated to yield 27 mg (84 %) of 3-[(2,4-Dichloro-phenyl)-isopropyl-carbamoyl]-5-phenyl-thiophene-2-carboxylic acid compound as pale brown foam. NMR ¹H : (CD₃OD; 400 MHz) 7.57-7.55 (m, 2H); 7.49-7.36 (m, 6H), 7.30-7.27 (m, 1H); 4.89 (quint, J = 6.73 Hz, 1H); 1.42 (d, J = 6.73 Hz, 3H); 1.12 (d, J = 6.73 Hz, 3H).

Example 24 The following compound was obtained from Discovery Technology:

3-(4-Chloro-2,5-dimethyl-benzenesulfonylamino)-5-phenyl-thiophene-2-carboxylic acid amide Compound #580

Example 25 The following compounds were obtained from Maybridge:

5-(4-Chloro-phenyl)-3-(toluene-4-sulfonylamino)-thiophene-2-carboxylic acid amide, Compound #563

5-(4-Fluoro-phenyl)-3-(toluene-4-sulfonylamino)-thiophene-2-carboxylic acid amide, Compound #564, GK 01137

5-(4-Methoxy-phenyl)-3-(toluene-4-sulfonylamino)-thiophene-2-carboxylic acid amide, Compound #565, GK 01175

Example 26 Evaluation of compounds in The HCV RNA-Dependent RNA Polymerase Assay

The following references are all incorporated by reference:

1. Behrens, S., Tomei, L., De Francesco, R. (1996) *EMBO* 15, 12-22

2. Harlow, E, and Lane, D. (1988) *Antibodies: A Laboratory Manual*. Cold Spring Harbord Laboratory. Cold Spring Harbord. NY.
3. Lohmann, V., Körner, F., Herian, U., and Bartenschlager, R. (1997) *J. Virol.* 71, 8416-8428
4. Tomei, L., Failla, C., Santolini, E., De Francesco, R., and La Monica, N. (1993) *J Virol* 67, 4017-4026

Compounds were evaluated using an *in vitro* polymerase assay containing purified recombinant HCV RNA-dependent RNA polymerase (NS5B protein). HCV NS5B was expressed in insect cells using a recombinant baculovirus as vector. The experimental procedures used for the cloning, expression and purification of the HCV NS5B protein are described below. Follows, are details of the RNA-dependent RNA polymerase assays used to test the compounds.

Expression of the HCV NS5B protein in insect cells:

The cDNA encoding the entire NS5B protein of HCV-Bk strain, genotype 1b, was amplified by PCR using the primers NS5Nhe5' (5'-GCTAGCGCTAGCTCAATGTCCTACACATGG-3') and XhoNS53' (5'-CTCGAGCTCGAGCGTCCATCGGTTGGGGAG-3') and the plasmid pCD 3.8-9.4 as template (Tomei et al, 1993). NS5Nhe5' and XhoNS53' contain two *NheI* and *XhoI* sites (underlined sequences), respectively, at their 5' end. The amplified DNA fragment was cloned in the bacterial expression plasmid pET-21b (Novagen) between the restriction sites *NheI* and *XhoI*, to generate the plasmid pET/NS5B. This plasmid was later used as template to PCR-amplify the NS5B coding region, using the primers NS5B-H9 (5'-ATACATATGGCTAGCATGTCAATGTCCTACACATGG-3') and NS5B-R4 (5'-GGATCCGGATCCCGTTCATCGGTTGGGGAG-3'). NS5B-H9 spans a region of 15 nucleotides in the plasmid pET-21b followed by the translation initiation codon (ATG) and 8 nucleotides corresponding to the 5' end of the NS5B coding region (nt. 7590-7607 in the HCV sequence with the accession number M58335). NS5B-R4 contains two *BamHI* sites (underlined) followed by 18 nucleotides corresponding to the region around the stop codon in the HCV genome (nt. 9365-9347). The amplified sequence, of 1.8 kb, was digested with *NheI* and *BamHI* and ligated to a predigested pBlueBacII plasmid (Invitrogen). The resulting recombinant plasmid was designated pBac/NS5B. Sf9 cells were

co-transfected with 3 µg of pBac/NS5B, together with 1 µg of linearized baculovirus DNA (Invitrogen), as described in the manufacturer's protocol. Following two rounds of plaque purification, an NS5B-recombinant baculovirus, BacNS5B, was isolated. The presence of the recombinant NS5B protein was determined by western blot analysis (Harlow and Lane, 1988) of BacNS5B-infected Sf9 cells, using a rabbit polyclonal antiserum (anti-NS5B) raised against a His-tagged version of the NS5B protein expressed in *E. coli*. Infections of Sf9 cells with this plaque purified virus were performed in one-liter spinner flasks at a cell density of 1.2×10^6 cells/ml and a multiplicity of infection of 5.

Preparation of a soluble recombinant NS5B protein

Sf9 cells were infected as described above. Sixty hours post-infection, cells were harvested then washed twice with phosphate buffer saline (PBS). Total proteins were solubilized as described in Lohmann et al. (1997) with some modifications. In brief, proteins were extracted in three steps, S1, S2, S3, using lysis buffers (LB) I, LB II and LB III (Lohmann et al, 1997). The composition of LBII was modified to contain 0.1 % triton X-100 and 150 mM NaCl to reduce the amount of solubilized NS5B protein at this step. In addition, sonication of cell extracts was avoided throughout the protocol to preserve the integrity of the protein structure.

Purification of recombinant NS5B using fast protein liquid chromatography (FPLC):

Soluble NS5B protein in the S3 fraction was diluted to lower the NaCl concentration to 300 mM, then it incubated batchwise with DEAE sepharose beads (Amersham-Pharmacia) for 2 hrs at 4°C, as described by Behrens et al. (1996). Unbound material was cleared by centrifugation for 15 min at 4°C, at 25 000 rpm using a SW41 rotor (Beckman). The supernatant was further diluted to lower the NaCl concentration to 200 mM and subsequently loaded, with a flow rate of 1 ml/min, on a 5 ml HiTrap® heparin column (Amersham-Pharmacia) connected to an FPLC® system (Amersham-Pharmacia). Bound proteins were eluted in 1 ml fractions, using a continuous NaCl gradient of 0.2 to 1 M, over a 25 ml volume.

NS5B-containing fractions were identified by sodium dodecyl sulfate polyacrylamide gel electrophoresis (SDS-PAGE), followed by western blotting using the anti-NS5B antiserum at a dilution of 1:2000. Positive fractions were pooled and the elution buffer was exchanged against a 50 mM NaPO₄ pH 7.0, 20 % glycerol, 0.5 % triton X-100 and 10 mM DTT, using a PD-10 column (Amersham-Pharmacia). The sample was then loaded onto a 1 ml HiTrap[®] SP column (Amersham-Pharmacia), with a flow rate of 0.1 ml/min. Bound proteins were eluted using a continuous 0 to 1 M NaCl gradient over a 15 ml volume. Eluted fractions were analyzed by SDS-PAGE and western blotting. Alternatively, proteins were visualized, following SDS-PAGE, by silver staining using the Silver Stain Plus kit (BioRad) as described by the manufacturer. Positive fractions were tested for RdRp activity (see below) and the most active ones were pooled, and stored as a 40 % glycerol solution at -70°C.

In vitro HCV RdRp Flashplate scintillation proximity assay (STREP-FLASH ASSAY) used to evaluate analogues:

This assay consists on measuring the incorporation of [³H] radiolabelled UTP in a polyrA/ biotinylated-oligo dT template-primer, captured on the surface of streptavidin-coated scintillant-embeded microtiter Flashplates[™] (NEN Life Science Products inc, MA, USA, SMP 103A). In brief, a 400 ng/μl polyrA solution (Amersham Pharmacia Biotech) was mixed volume-to-volume with 5' biotin-oligo dT₁₅ at 20 pmol/μl. The template and primers were denatured at 95 C for 5 minutes then incubated at 37 C for 10 minutes. Annealed template-primers were subsequently diluted in a Tris-HCl containing buffer and allowed to bind to streptavidin-coated flashplates overnight. Unbound material was discarded, compounds were added in a 10 μl solution followed by a 10 μl of a solution containing 50 mM MgCl₂, 100 mM Tris-HCl pH 7.5, 250 mM NaCl and 5 mM DTT. The enzymatic reaction was initiated upon addition of a 30 μl solution containing the enzyme and substrate to obtain the following concentrations: 25 μM UTP, 1 μCi [³H] UTP and 100 nM recombinant HCV NS5B. RdRp reactions were allowed to proceed for 2 hrs at room temperature after which wells were washed three times with a 250μL of 0.15 M NaCl solution, air dried at 37 C, and counted

using a liquid scintillation counter (Wallac Microbeta Trilex, Perkin-Elmer, MA, USA). Results are shown in Table 1.

In vitro HCV RdRp filtration assay used to evaluate analogues
5 RdRp assays were conducted using the homopolymeric
template/primer polyA/oligo dT. All RdRp reactions were
performed in a total volume of 50 μ l, and in a basic buffer
consisting of 20 mM Tris-HCl pH 7.5, 1mM DTT, 50 mM NaCl, 5 mM
MgCl₂, 0.5 μ Ci [γ^{32} P]-UTP (3000 Ci/mmol), 15 μ M cold UTP and 20 U
10 RNasin (Promega). Standard HCV RdRp reactions contained 200 ng
of purified NS5B protein. PolyA RNAs (Amersham-Pharmacia) was
resuspended at 400 ng/ μ l. The primer oligodT₁₅ (Canadian life
technologies) was diluted to a concentration of 20 pmol/ μ l (7.6
ng/ml). Templates and primers were mixed volume to volume,
15 denatured at 95°C for 5 min and annealed at 37°C for 10 min.
Following a two hour incubation at 22°C, reactions were stopped
by the addition of 100 μ g of sonicated salmon sperm DNA (Life
Technologies) and 1 ml of 10 % trichloroacetic acid-0.5 %
tetrasodium pyrophosphate (TCA-PPI). Nucleic acids were
20 precipitated at 4°C for 30 min after which samples were filtered
on GF/C glass microfiber filters (Millipore). Membranes were
subsequently washed with 25 ml of a 1% TCA-0.1 % PPI solution,
then air dried. Incorporated radioactivity was quantified using
a liquid scintillation counter (1450-Microbeta, Wallac). Results
25 are shown in Table 1.

Example 27 Evaluation of Analogues for measurement of ATPase
activity of HCV NS3 helicase

30 Malachite Green Assay:

The measurement of ATPase activity was performed by measuring
the amount of free inorganic phosphate released during the
conversion of ATP to ADP by the HCV NS3 ATPase activity. The
35 assay is as follows: In a 96-well microtiter-plate, compounds
were dissolved at various concentrations in a final volume of 25
 μ L of ATPase buffer containing 400 μ M ATP. The enzymatic

reaction was initiated by the addition of 25 μ l of ATPase buffer containing 6 nM of HCV NS3 enzyme without ATP to the wells followed by an incubation of 30 min. at 37 C. Essentially, the final concentration of the ATPase buffer components are as

5 follows: 44 mM MOPS pH 7.0, 8.8 mM NaCl, 2.2 mM MgCl₂, 125 μ g/ml poly A, 1% DMSO, 200 μ M ATP, and 3 nM HCV NS3 enzyme. The

reaction was stopped by the addition of 100 μ l of Biomol Green™ reagent (BIOMOL® Research Laboratories Inc., Plymouth Meeting, PA). In order to allow the development of the green color, the

10 plate was incubated for 15 min. at room temperature. Then the plate was read on a micro-plate reader at 620 nm. The 50% inhibitory concentration (IC₅₀) for anti-ATPase activity was

defined as the concentration of compound that resulted in a 50 % reduction of the signal compared to the signal observed in

15 control sample without compound. The signal recorded was also

corrected from the background signal obtained with control

samples with compound only. The IC₅₀ was determined from dose-response curves using six to eight concentrations per compound.

Curves were fitted to data points using a non-linear regression

20 analysis, and IC₅₀s were interpolated from the resulting curves

using GraphPad Prism software, version 2.0 (GraphPad Software Inc, San Diego, CA).

HPLC Assay:

25 The measurement of HCV NS3 ATPase activity was performed by

measuring the amount of ADP produced during the conversion of

ATP to ADP by the HCV NS3 enzyme using paired-ion HPLC on a

reverse phase column. The assay is as follows: The same

30 protocol as mentioned above was used except that the final

concentration of HCV NS3 enzyme was reduced to 1 nM in a 50 μ l

reaction mixture and that the ATPase reaction was stopped by the

addition of 12.5 μ l of 0.5 M EDTA. A modular liquid

chromatography system (TSP Spectrasystem®, ThermoQuest

35 Corporation, San Diego, USA) using a ChromQuest™ software

(ThermoQuest Corporation, San Diego, USA) controlled the

autosampling of 25 μ l from each reaction. The mobile phase was

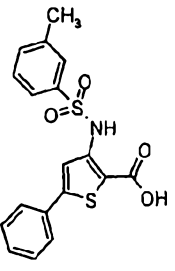
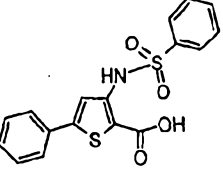
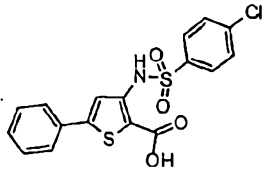
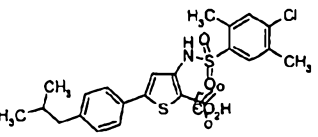
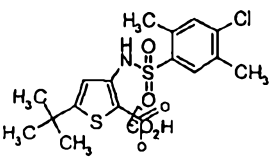
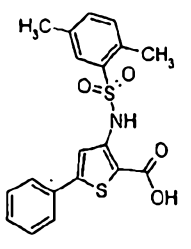
an isocratic solution of 0.15 M triethylamine, 6% methanol, and phosphoric acid to pH 5.5. ADP and ATP peaks were resolved using the Aqua 5 μ , C18, 125 Å, (150 X 4.6 mm) reverse phase column. The extent of ATP conversion to ADP was evaluated by measuring the area under the ADP peak produced which was detected at 259 nm. The amount of ADP was corrected for the presence of ADP contaminant in the original ATP solution. The 50% inhibitory concentration (IC_{50}) for anti-ATPase activity was defined as the concentration of compound that resulted in a 50 % reduction of the ADP peak area compared to the ADP peak area observed in control sample without compound. The IC_{50} was determined from dose-response curves using six to eight concentrations per compound. Curves were fitted to data points using a non-linear regression analysis, and IC_{50} s were interpolated from the resulting curves using GraphPad Prism software, version 2.0 (GraphPad Software Inc, San Diego, CA).

EXAMPLE 27 List of compounds and related polymerase activity *

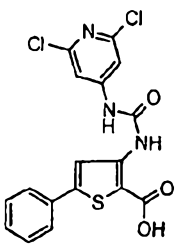
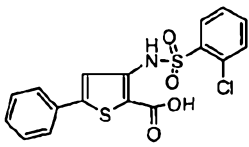
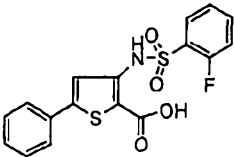
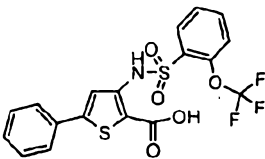
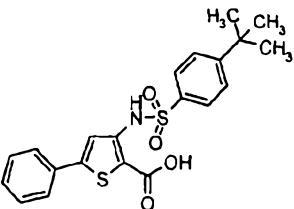
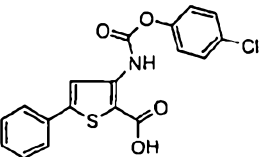
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1		3-[(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYL)-(3-iodo-benzyl)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
2		3-[(3-BENZOFURAN-2-YL-BENZYL)-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
3		3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
4		3-[(2,4-DICHLORO-BENZOYL)-[5-(3-TRIFLUOROMETHYL-PHENYL)-FURAN-2-YLMETHYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

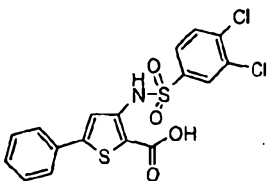
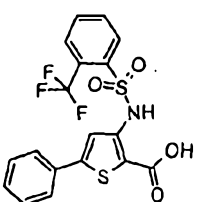
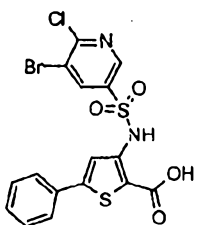
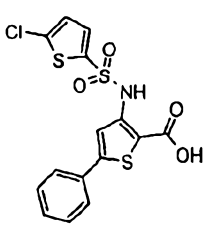
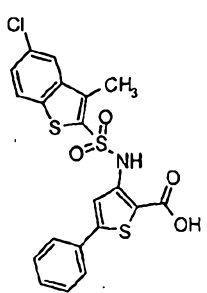
	MOLSTRUCTURE	COMPOUND NAME	IC50
5		3-[(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
6		5-(4-FLUORO-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
7		3-(2,4-DICHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
8		3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
9		3-[(2,4-DICHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
10		5-#TERT!BUTYL-3-(4-CHLORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++

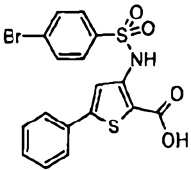
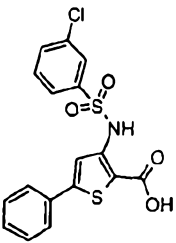
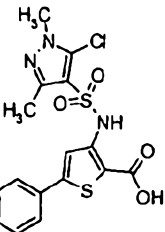
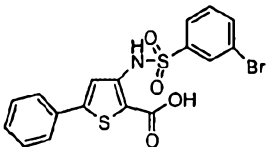
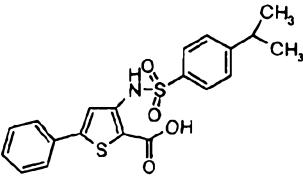
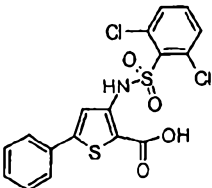
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12		3-[(5-BENZOFURAN-2-YL-THIOPHEN-2- YLMETHYL)-(2,4-DICHLORO-BENZOYL)- AMINO]-5-PHENYL-THIOPHENE-2- CARBOXYLIC ACID	++
13		5-PHENYL-3-(TOLUENE-4- SULFONYLAMINO)-THIOPHENE-2- CARBOXYLIC ACID	+++
14		3-(4-CHLORO-2,5-DIMETHYL- BENZENESULFONYLAMINO)-5-(4- CHLORO-PHENYL)-THIOPHENE-2- CARBOXYLIC ACID	+++
15		5-PHENYL-3-(TOLUENE-2- SULFONYLAMINO)-THIOPHENE-2- CARBOXYLIC ACID	+

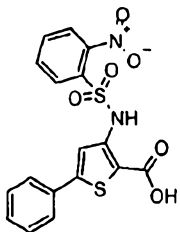
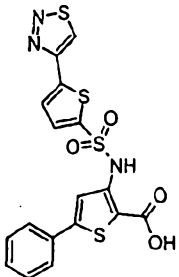
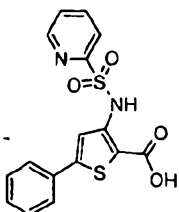
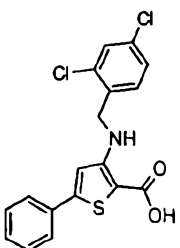
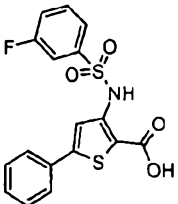
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17		3-BENZENESULFONYLAMINO-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
18		3-(4-CHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
19		3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	++
20		5-TERT-BUTYL-3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
21		3-(2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

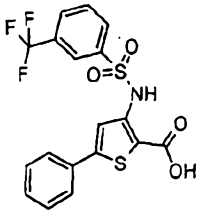
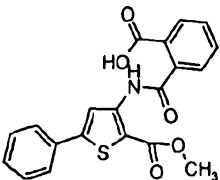
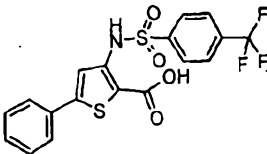
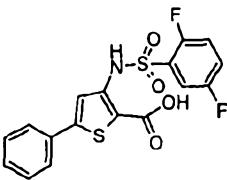
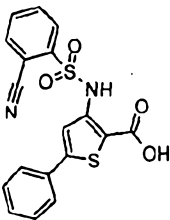
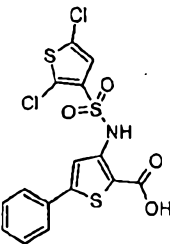
	MOLSTRUCTURE	COMPOUND NAME	IC50
22		3-(4-METHOXY-2,3,6-TRIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
23		5-PHENYL-3-(THIOPHENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
24		4-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-[2,3']BITHIOPHENYL-5-CARBOXYLIC ACID	+++
25		5-(3,5-BIS-TRIFLUOROMETHYL-PHENYL)-3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+
26		8-CHLORO-3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-4H-1,5-DITHIA-CYCLOPENTA[A]NAPHTHALENE-2-CARBOXYLIC ACID	++
27		3-(2,4-DIFLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

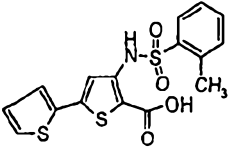
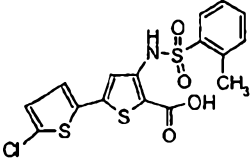
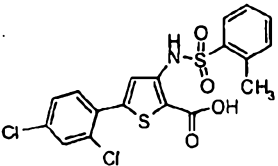
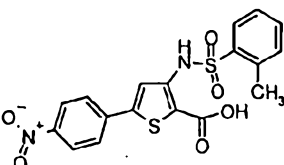
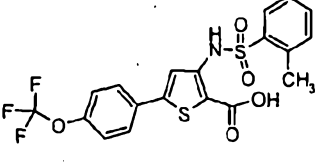
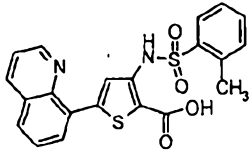
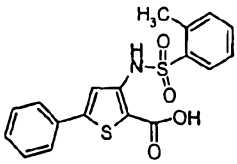
	MOLSTRUCTURE	COMPOUND NAME	IC50
28		3-[3-(2,6-DICHLORO-PYRIDIN-4-YL)-UREIDO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
29		3-(2-CHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
30		3-(2-FLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
31		5-PHENYL-3-(2-TRIFLUOROMETHOXY-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+
32		3-(4-#TERT!-BUTYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
33		3-(4-CHLORO-PHENOXYCARBONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

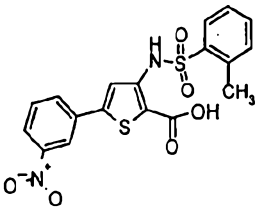
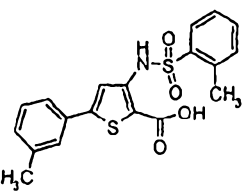
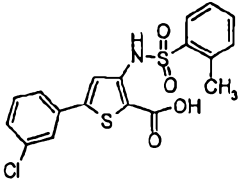
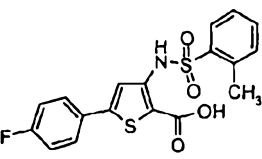
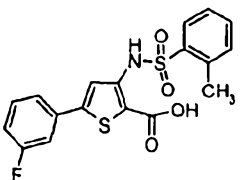
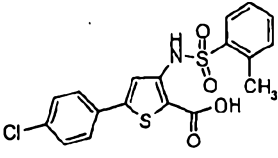
	MOLSTRUCTURE	COMPOUND NAME	IC50
34		3-(3,4-DICHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
35		5-PHENYL-3-(2-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+
36		3-(5-BROMO-6-CHLORO-PYRIDINE-3-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
37		3-(5-CHLORO-THIOPHENE-2-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
38		3-(5-CHLORO-3-METHYL-BENZO[1,2-b:4,5-b']THIOPHENE-2-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

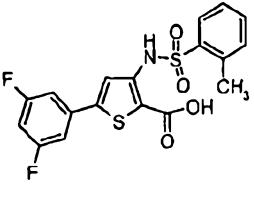
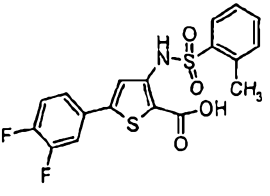
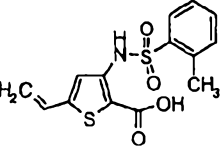
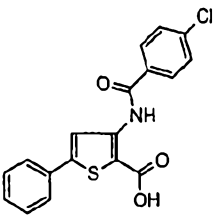
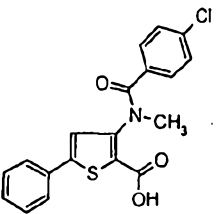
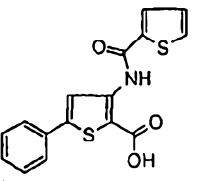
	MOLSTRUCTURE	COMPOUND NAME	IC50
39		3-(4-BROMO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
40		3-(3-CHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
41		3-(5-CHLORO-1,3-DIMETHYL-1H-PYRAZOLE-4-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
42		3-(3-BROMO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
43		3-(4-ISOPROPYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
44		3-(2,6-DICHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
45		3-(2-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
46		5-PHENYL-3-(5-[1,2,3]THIADIAZOL-4-YL-THIOPHENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
47		5-PHENYL-3-(PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+
48		3-(2,4-DICHLORO-BENZYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
49		3-(3-FLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
50		5-PHENYL-3-(3-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
51		3-(2-CARBOXY-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID METHYL ESTER	++
52		5-PHENYL-3-(4-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
53		3-(2,5-DIFLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
54		3-(2-CYANO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
55		3-(2,5-DICHLORO-THIOPHENE-3-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
56		4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID	+++
57		5'-CHLORO-4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID	+++
58		5-(2,4-DICHLORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
59		5-(4-NITRO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
60		3-(TOLUENE-2-SULFONYLAMINO)-5-(4-TRIFLUOROMETHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
61		5-QUINOLIN-8-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+
62		5-PHENYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++

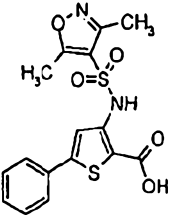
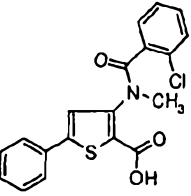
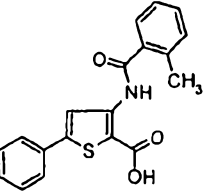
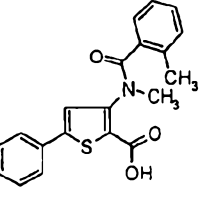
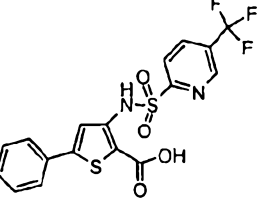
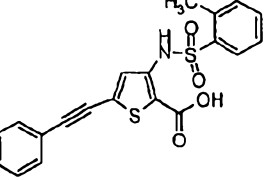
	MOLSTRUCTURE	COMPOUND NAME	IC50
63		5-(3-NITRO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
64		3-(TOLUENE-2-SULFONYLAMINO)-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID	+++
65		5-(3-CHLORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
66		5-(4-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
67		5-(3-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
68		5-(4-CHLORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++

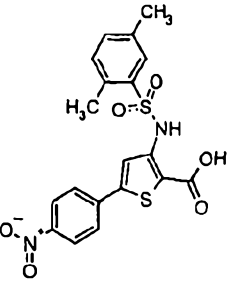
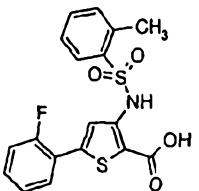
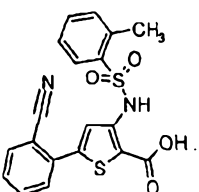
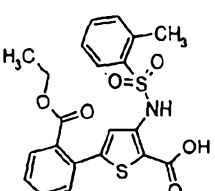
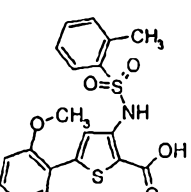
	MOLSTRUCTURE	COMPOUND NAME	IC50
69		5-(3,5-DIFLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
70		5-(3,4-DIFLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
71		3-(TOLUENE-2-SULFONYLAMINO)-5-VINYL-THIOPHENE-2-CARBOXYLIC ACID	++
72		3-(4-CHLORO-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
73		3-[(4-CHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
74		5-PHENYL-3-[(THIOPHENE-2-CARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
75		3-[METHYL-(THIOPHENE-2-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
76		3-(2-BROMO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
77		3-(2,4-DIFLUORO-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
78		3-[(2,4-DIFLUORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
79		3-(TOLUENE-2-SULFONYLAMINO)-5-TRIMETHYLSILANYLETHYNYL-THIOPHENE-2-CARBOXYLIC ACID	+++
80		5-ETHYNYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
81		3-(TOLUENE-2-SULFONYLAMINO)-5-(3-TRIFLUOROMETHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	++
82		5-BENZOYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
83		5-(4-CYANO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
84		5-(3-CHLORO-4-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
85		5-(3,4-DICHLORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
86		5-PYRIDIN-4-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
87		5-PYRIDIN-3-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
88		3-(TOLUENE-2-SULFONYLAMINO)-5-(4-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
89		5-(4-METHANESULFONYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
90		5-(3-ACETYLAMINO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
91		5-(3-CHLORO-4-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
92		3-(4-METHYL-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
93		3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
94		3-(3,5-DIMETHYL-ISOXAZOLE-4-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
95		3-[(2-CHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
96		3-(2-METHYL-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
97		3-[METHYL-(2-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
98		5-PHENYL-3-(5-TRIFLUOROMETHYL-PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
99		5-PHENYLETHYNYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
100		3-(2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
101		5-(2-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
102		5-(2-CYANO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
103		5-(2-ETHOXYCARBONYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
104		5-(2-METHOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++

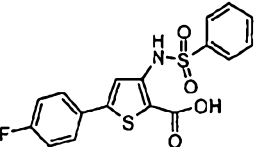
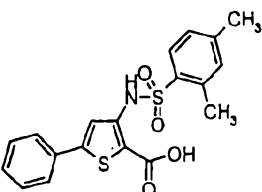
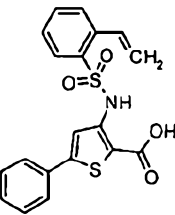
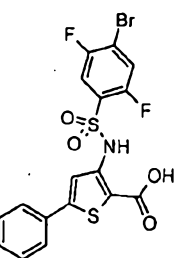
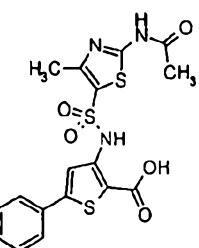
	MOLSTRUCTURE	COMPOUND NAME	IC50
105		3'-METHYL-4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID	++
106		3-(TOLUENE-2-SULFONYLAMINO)-5-(2-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	++
107		3-(2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
108		5-STYRYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
109		3-(2,4-DIFLUORO-BENZENESULFONYLAMINO)-5-(4-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
110		3-(2,4-DIFLUORO-BENZENESULFONYLAMINO)-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++

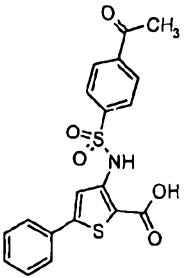
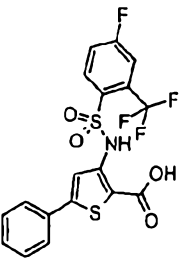
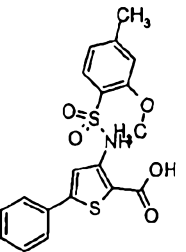
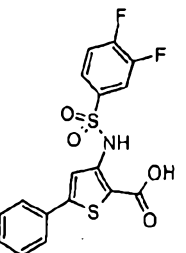
	MOLSTRUCTURE	COMPOUND NAME	IC50
111		3-[[5-(3-CHLORO-4-FLUORO-PHENYL)-THIOPHEN-2-YLMETHYL]-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
112		3-[(4-OXO-1-PHENYL-1,3,8-TRIAZA-SPIRO[4.5]DECANE-8-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
113		3-[[4-(2-OXO-2,3-DIHYDRO-BENZOIMIDAZOL-1-YL)-PIPERIDINE-1-CARBONYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
114		3-[[4-(4-NITRO-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
115		5-(2-CARBOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
116		5-(4-CHLORO-PHENYL)-3-(PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
117		5-(3-CYANO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
118		3-(2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-P-TOLYL-THIOPHENE-2-CARBOXYLIC ACID	+++
119		3-(2,4-DIFLUORO-BENZENESULFONYLAMINO)-5-P-TOLYL-THIOPHENE-2-CARBOXYLIC ACID	+++
120		5-PHENETHYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
121		5-(3-ETHOXYCARBONYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++

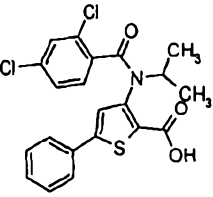
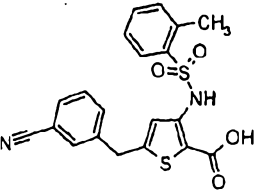
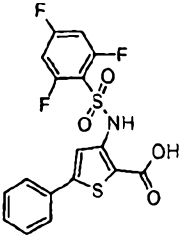
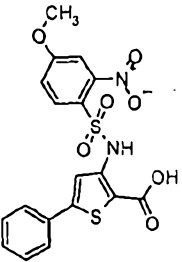
	MOLSTRUCTURE	COMPOUND NAME	IC50
122		5-(4-METHOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
123		5-(3-METHOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
124		5-(4'-BROMO-BIPHENYL-4-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
125		5-(4-HYDROXYMETHYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
126		5-FURAN-3-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
127		5-BENZOFURAN-2-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
128		5-PYRIDIN-2-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++

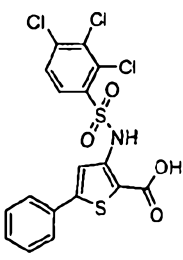
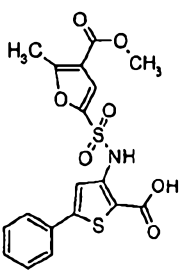
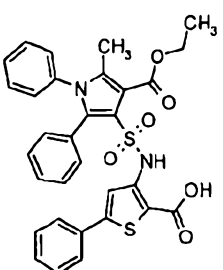
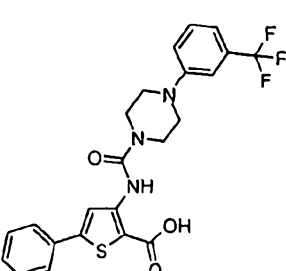
	MOLSTRUCTURE	COMPOUND NAME	IC50
129		5-(4-NITRO-PHENYL)-3-(PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
130		3-[(BENZOFURAN-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
131		3-[(2,4-DIMETHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
132		3-[[5-(2-CYANO-PHENYL)-THIOPHEN-2-YLMETHYL]- (2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
133		5-(4-FLUORO-PHENYL)-3-(PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
134		5-[2-(4-CHLORO-PHENYL)-VINYL]-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++

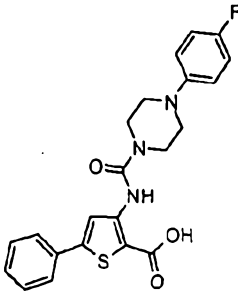
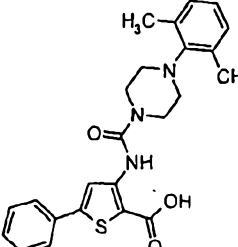
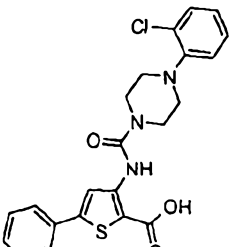
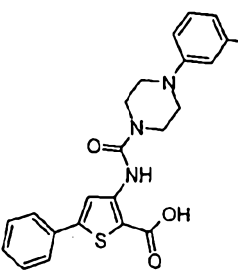
	MOLSTRUCTURE	COMPOUND NAME	IC50
135		3-BENZENESULFONYLAMINO-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
136		3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
137		5-PHENYL-3-(2-VINYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
138		3-(4-BROMO-2,5-DIFLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
139		3-(2-ACETYLAMINO-4-METHYL-THIAZOLE-5-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
140		3-(4-ACETYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
141		3-(4-FLUORO-2-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
142		3-(2-METHOXY-4-METHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
143		3-(3,4-DIFLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
144		4-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-5-(4-CHLORO-PHENYL)-2-METHYL-FURAN-3-CARBOXYLIC ACID ETHYL ESTER	++
145		3-(4-FLUORO-3-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
146		3-(2-AMINO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
147		3-(3-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
148		3-(4-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

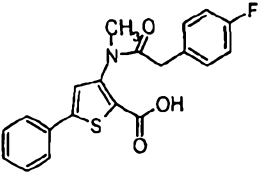
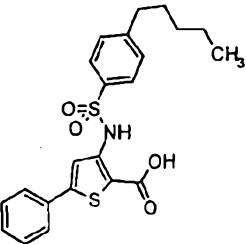
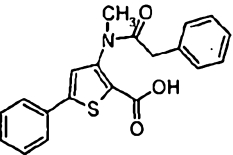
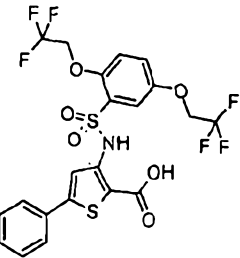
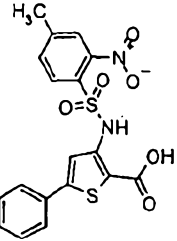
	MOLSTRUCTURE	COMPOUND NAME	IC50
149		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
150		5-(3-CYANO-BENZYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+
151		5-PHENYL-3-(2,4,6-TRIFLUORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
152		3-(4-METHOXY-2-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

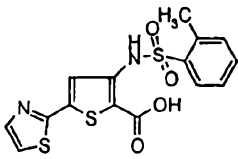
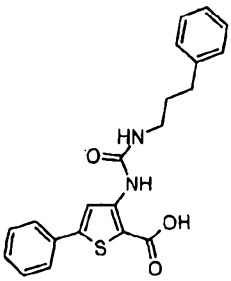
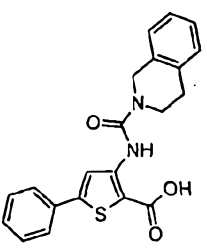
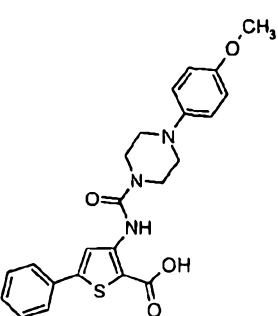
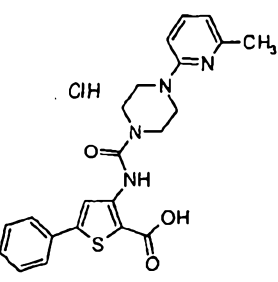
	MOLSTRUCTURE	COMPOUND NAME	IC50
153		5-PHENYL-3-(2,3,4-TRICHLORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
154		5-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-2-METHYL-FURAN-3-CARBOXYLIC ACID METHYL ESTER	+++
155		4-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-2-METHYL-1,5-DIPHENYL-1H-PYRROLE-3-CARBOXYLIC ACID ETHYL ESTER	+++
156		5-PHENYL-3-([4-(3-TRIFLUOROMETHYL-PHENYL)-PIPERAZINE-1-CARBONYL]-AMIN)-THIOPHENE-2-CARBOXYLIC ACID	++

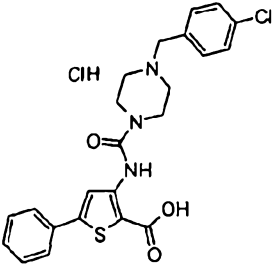
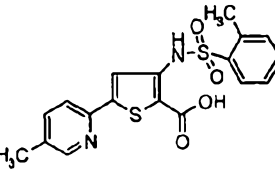
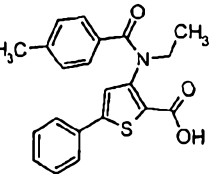
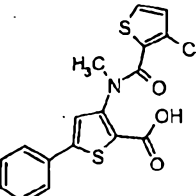
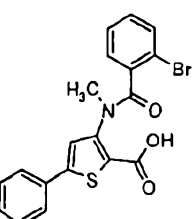
	MOLSTRUCTURE	COMPOUND NAME	IC50
157		3-([4-(4-FLUORO-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
158		3-([4-(2,6-DIMETHYL-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
159		3-([4-(2-CHLORO-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
160		3-([4-(3-CHLORO-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

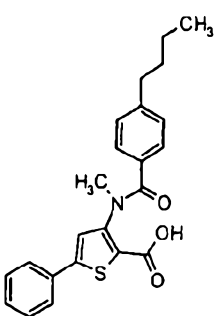
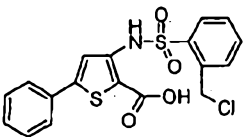
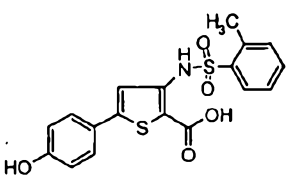
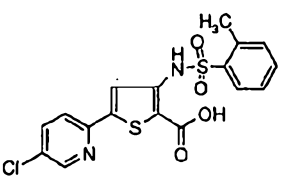
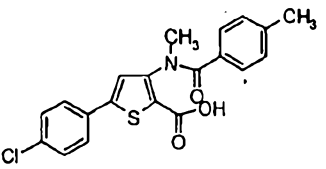
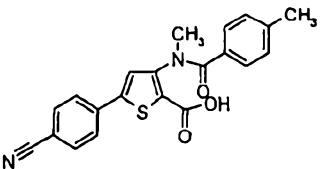
	MOLSTRUCTURE	COMPOUND NAME	IC50
161		4,4'-BIS-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5,5'-DICARBOXYLIC ACID	+++
162		3-[ALLYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
163		5-(1-DIMETHYLSULFAMOYL-1#H-PYRAZOL-4-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
164		5-(3-AMINO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
165		5-(4-AMINO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
166		5-(4-ACETYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
167		4-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-2,5-DIMETHYL-1H-PYRROLE-3-CARBOXYLIC ACID ETHYL ESTER	++
168		4-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-5-(4-CHLORO-PHENYL)-3-METHYL-1-PHENYL-1H-PYRROLE-2-CARBOXYLIC ACID ETHYL ESTER	++
169		3-(3,5-DICHLORO-4-HYDROXY-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
170		5-(1#H-1-PYRAZOL-4-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
171		5-(3-HYDROXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
172		3-[METHYL-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

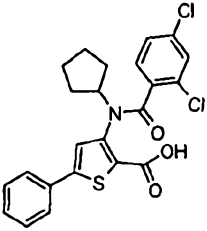
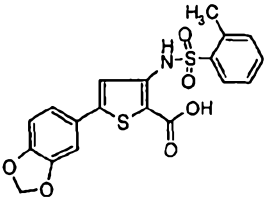
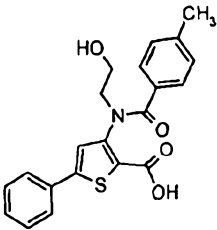
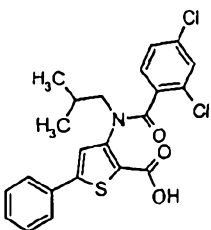
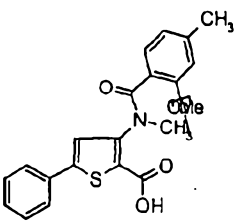
	MOLSTRUCTURE	COMPOUND NAME	IC50
173		3-[[2-(4-FLUORO-PHENYL)-ACETYL]-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
174		3-(4-PENTYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
175		3-(METHYL-PHENYLACETYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
176		3-[2,5-BIS-(2,2,2-TRIFLUORO-ETHOXY)-BENZENESULFONYLAMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
177		3-(4-METHYL-2-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

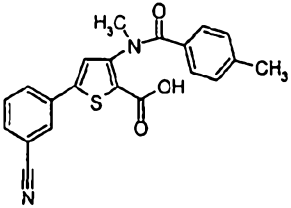
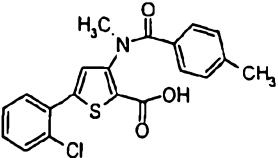
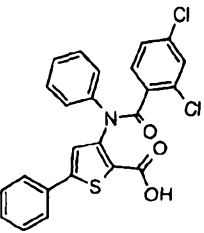
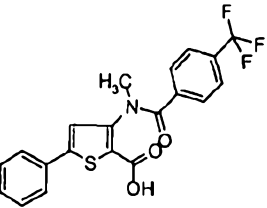
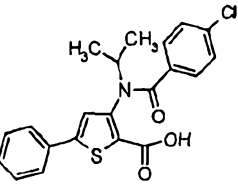
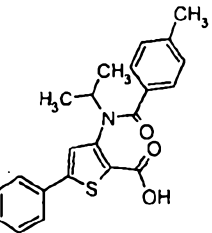
	MOLSTRUCTURE	COMPOUND NAME	IC50
178		5-THIAZOL-2-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
179		5-PHENYL-3-[3-(3-PHENYL-PROPYL)-UREIDO]-THIOPHENE-2-CARBOXYLIC ACID	++
180		3-[(3,4-DIHYDRO-1H-ISOQUINOLINE-2-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
181		3-[[4-(4-METHOXY-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
182		3-[[4-(6-METHYL-PYRIDIN-2-YL)-PIPERAZINE-1-CARBONYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID HYDROCHLORIDE	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
183		3-([4-(4-CHLORO-BENZYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID HYDROCHLORIDE	++
184		5-(5-METHYL-PYRIDIN-2-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
185		3-[ETHYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
186		3-[(3-CHLORO-THIOPHENE-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
187		3-[(2-BROMO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

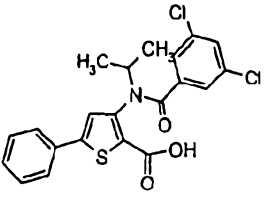
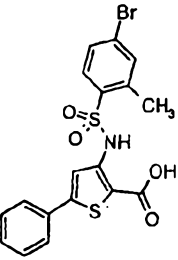
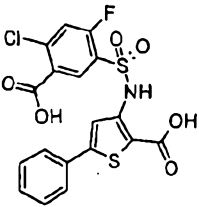
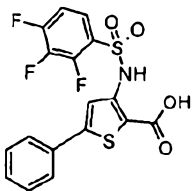
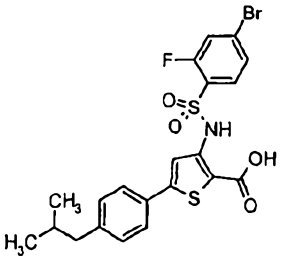
	MOLSTRUCTURE	COMPOUND NAME	IC50
188		3-[(4-BUTYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
189		3-(2-CHLOROMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
190		5-(4-HYDROXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
191		5-(5-CHLORO-PYRIDIN-2-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
192		5-(4-CHLORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	++
193		5-(4-CYANO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	++

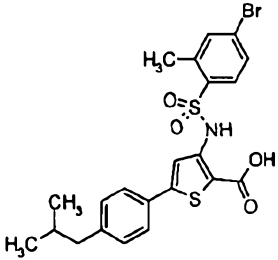
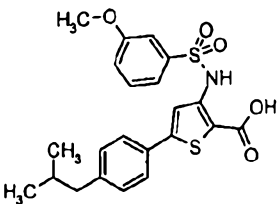
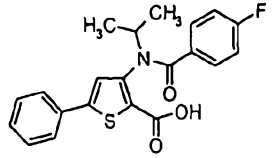
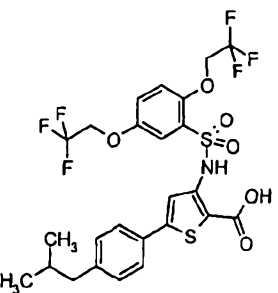
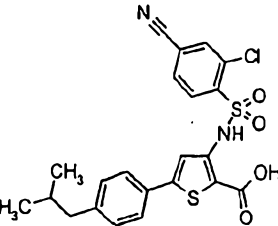
	MOLSTRUCTURE	COMPOUND NAME	IC50
194		3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(4-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	++
195		5-(4-HYDROXYMETHYL-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	++
196		3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(3-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
197		5-(4-FLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
198		5-(4-METHOXY-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	++
199		3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-#P1-TOLYL-THIOPHENE-2-CARBOXYLIC ACID	++
200		5-(4-AMINO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
201		3-[(CYCLOPENTYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
202		5-BENZO[1,3]DIOXOL-5-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
203		3-[(2-HYDROXY-ETHYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
204		3-[(2,4-DICHLORO-BENZOYL)-ISOBUTYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
205		3-[(2-METHOXY-4-METHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
206		5-(3-CYANO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
207		5-(2-CHLORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	++
208		3-[(2,4-DICHLORO-BENZOYL)-PHENYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
209		3-[4-(TRIFLUOROMETHYL-BENZOYL)METHYLAMINE]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
210		3-[(4-CHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
211		3-[ISOPROPYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

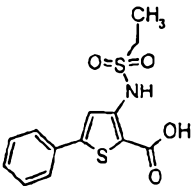
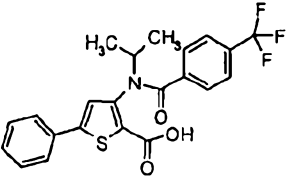
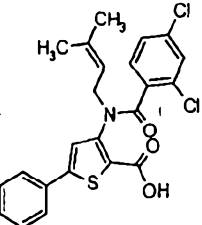
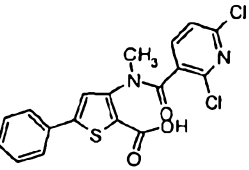
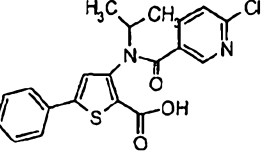
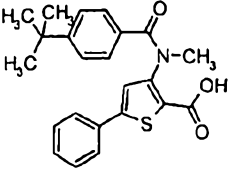
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212		5-(3,5-DIFLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
213		5-(3-FLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
214		5-(2,4-DIFLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	++
215		5-(4-HYDROXY-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
216		3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(4-TRIFLUOROMETHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	++
217		5-(2-HYDROXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
218		3-[(2-CHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

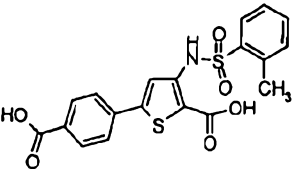
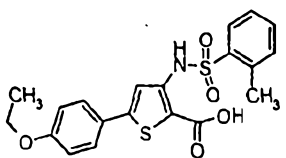
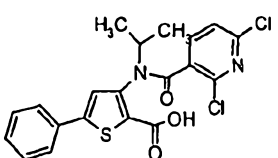
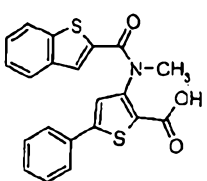
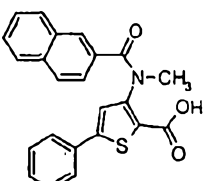
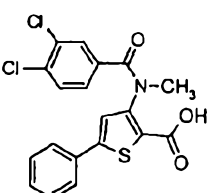
	MOLSTRUCTURE	COMPOUND NAME	IC50
219		3-[(3,5-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
220		3-(4-BROMO-2-METHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
221		3-(5-CARBOXY-4-CHLORO-2-FLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
222		5-PHENYL-3-(2,3,4-TRIFLUORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
223		3-(4-BROMO-2-FLUORO-BENZENESULFONYLAMINO)-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
224		3-(4-BROMO-2-METHYL-BENZENESULFONYLAMINO)-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+
225		5-(4-ISOBUTYL-PHENYL)-3-(3-METHOXY-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+
226		3-[(4-FLUORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
227		3-[2,5-BIS-(2,2,2-TRIFLUORO-ETHOXY)-BENZENESULFONYLAMINO]-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+
228		3-(2-CHLORO-4-CYANO-BENZENESULFONYLAMINO)-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+

	MOLSTRUCTURE	COMPOUND NAME	IC50
229		5'-ACETYL-4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID	+++
230		5-BENZO[B]THIOPHEN-2-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
231		5-(4-BUTYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
232		5-(4-ETHYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
233		3-[BENZYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
234		3-[(4-CHLORO-2-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
235		3-[(2,4-DIMETHYL-BENZENESULFONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
236		5-(4-ACETYL-PHENYL)-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
237		5-(4-ACETYL-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
238		5-(4-ACETYL-PHENYL)-3-(4-CHLORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
239		5-(4-CARBOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID #TERTI-BUTYL ESTER	++
240		3-[(2,4-DIMETHYL-BENZENESULFONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
241		3-[ACETYL-(4-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
242		3-ETHANESULFONYLAMINO-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
243		3-[ISOPROPYL-(4-TRIFLUOROMETHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
244		3-[(2,4-DICHLORO-BENZOYL)-(3-METHYL-BUT-2-ENYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
245		3-[(2,6-DICHLORO-PYRIDINE-3-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
246		3-[(6-CHLORO-PYRIDINE-3-CARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
247		3-[(4-TERT-BUTYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
248		5-(4-CARBOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
249		5-(4-ETHOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
250		3-[(2,6-DICHLORO-PYRIDINE-3-CARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
251		3-[(BENZO[B]THIOPHENE-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
252		3-[METHYL-(NAPHTHALENE-2-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
253		3-[(3,4-DICHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
254		3-[(3,5-DICHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
255		3-[(4-BROMO-3-METHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
256		3-[(3-CHLORO-BENZO[B]THIOPHENE-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
257		3-[METHYL-(4-METHYL-3-NITRO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
258		5-(4-CARBAMOYL-PHENYL)-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
259		5-(4-CARBAMOYL-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++

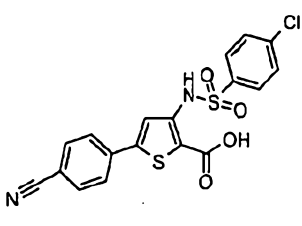
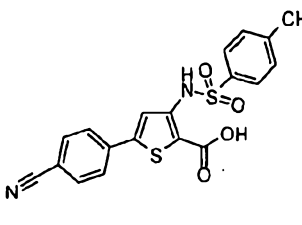
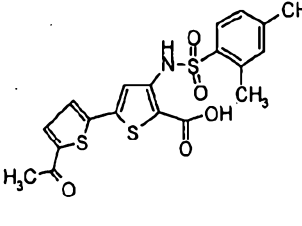
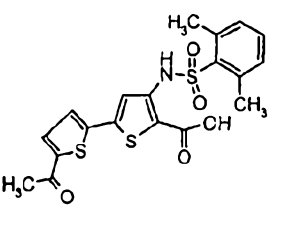
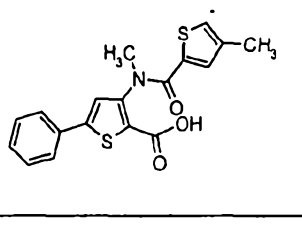
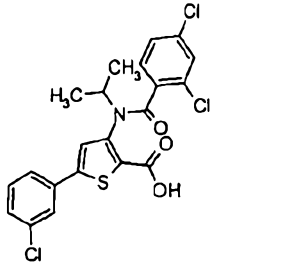
	MOLSTRUCTURE	COMPOUND NAME	IC50
260		5-(1H-INDOL-5-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
261		3-[(S)-1-(2,4-DICHLORO-BENZOYL)-2-PHENYL-ETHYL]-AMINO-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
262		3-[(2S)-2-(4-DIMETHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
263		5-(4-AZIDO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
264		3-[(2S)-2-(2,4-DICHLORO-BENZOYL)-(1-PHENYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
265		5-(4-CARBAMOYL-PHENYL)-3-(4-CHLORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
266		5-(2-FLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
267		3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID	+++
268		3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID	+++
269		5-(3-CHLORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
270		5-(3,4-DIFLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
271		5-(3-AMINO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
272		5-(3-ACETYL-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	++
273		5-(3-HYDROXY-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
274		3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
275		3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(4-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
276		3-[(3,4-DIMETHOXY-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
277		3-[METHYL-(2,4,6-TRIFLUORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
278		3-[(2,3-DIFLUORO-4-TRIFLUOROMETHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
279		3-[(3-FLUORO-4-TRIFLUOROMETHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
280		3-[(2,3-DIFLUORO-4-METHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
281		3-[(2-FLUORO-4-TRIFLUOROMETHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
282		5-(4-CARBAMOYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
283		5-(4-FLUORO-PHENYL)-3-[ISOPROPYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
284		3-[(2-BROMO-4-CHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
285		3-(2,6-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
286		3-[METHYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
287		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID METHYL ESTER	++
288		5-(4-CYANO-PHENYL)-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
289		3-(4-CHLORO-BENZENESULFONYLAMINO)-5-(4-CYANO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
290		5-(4-CYANO-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
291		5'-ACETYL-4-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID	+++
292		5'-ACETYL-4-(2,6-DIMETHYL-BENZENESULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID	+++
293		3-[METHYL-(4-METHYL-THIOPHENE-2-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
294		5-(3-CHLORO-PHENYL)-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++

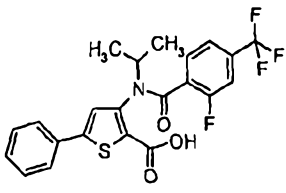
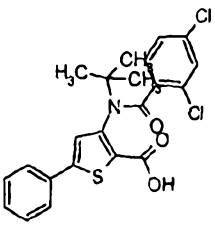
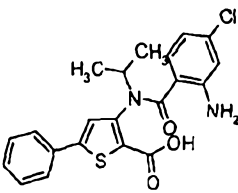
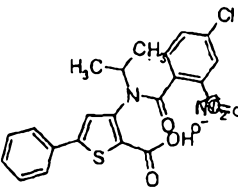
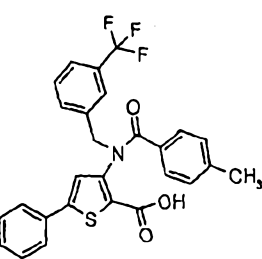
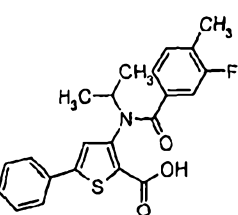
	MOLSTRUCTURE	COMPOUND NAME	IC50
295		5'-CYANO-4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID	+++
296		3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-PYRIDIN-2-YL-THIOPHENE-2-CARBOXYLIC ACID	++
297		3-[(2,4-DICHLORO-THIOBENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
298		5-PHENYL-3-(2,4,6-TRIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
299		3-[(1-CARBOXY-ETHYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
300		3-[(4-METHYL-BENZOYL)-(3-METHYL-BUT-2-ENYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
301		3-[(2-HYDROXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
302		3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-PYRIDIN-3-YL-THIOPHENE-2-CARBOXYLIC ACID	++
303		5'-ACETYL-4-[METHYL-(4-METHYL-BENZOYL)-AMINO]-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID	+++
304		3-[ISOPROPYL-(4-METHYL-BENZOYL)-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
305		3-[ISOPROPYL-(4-METHYL-BENZOYL)-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID	+++
306		3-[(2-BROMO-4-CHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
307		3-[(4-CHLORO-2-FLUORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
308		3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-4-METHYL-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
309		3-[(2-BROMO-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
310		3-[(4-CHLORO-2-iodo-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
311		3-[(4-CYANO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
312		3-[ALLYL-(4-METHYL-BENZOYL)-AMINO]-5-[4-(2-CARBOXY-VINYL)-PHENYL]-THIOPHENE-2-CARBOXYLIC ACID.	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
313		3-[(4-CHLORO-2-HYDROXY-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
314		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-4-METHYL-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
315		5-#TERT!BUTYL-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
316		3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
317		3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
318		5-[4-(2-CARBOXY-ETHYL)-PHENYL]-3-[(4-METHYL-BENZOYL)-PROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
319		5-BENZOFURAN-2-YL-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
320		3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-HYDROXYMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
321		3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-METHANESULFONYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
322		5-[4-(2-CARBOXY-VINYL)-PHENYL]-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
323		3-ALLYL-(4-METHYL-BENZOYL)-AMINO]-5-[3-(2-CARBOXY-VINYL)-PHENYL]-THIOPHENE-2-CARBOXYLIC ACID	++
324		3-ISOPROPYL-(2,4,6-TRIMETHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
325		5-[3-(2-CARBOXY-ETHYL)-PHENYL]-3-[(4-METHYL-BENZOYL)-PROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID.	++

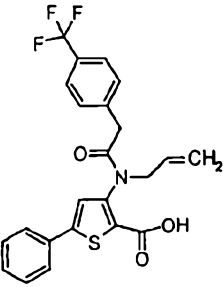
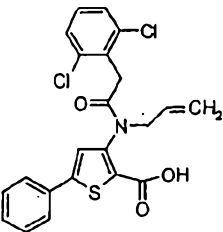
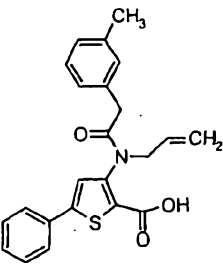
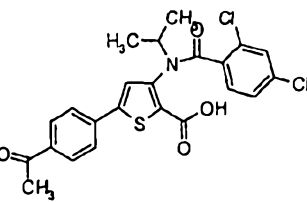
	MOLSTRUCTURE	COMPOUND NAME	IC50
326		3-[(2-FLUORO-4-TRIFLUOROMETHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
327		3-[(TERT-BUTYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
328		3-[(2-AMINO-4-CHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
329		3-[(4-CHLORO-2-NITRO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
330		3-[(4-METHYL-BENZOYL)-(3-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
331		3-[(3-FLUORO-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
332		5-(4-CARBOXY-PHENYL)-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	+++
333		3-[CYCLOPROPYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
334		3-[(3-TERT-BUTYL-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
335		3-[(3-CHLORO-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
336		3-[(2,4-DIFLUORO-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

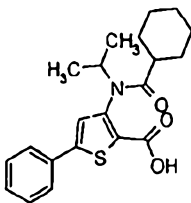
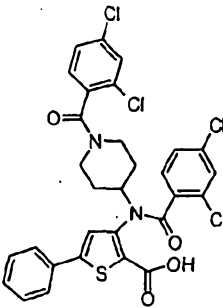
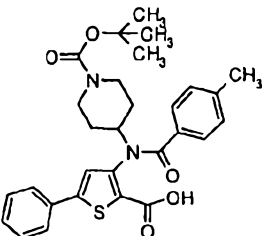
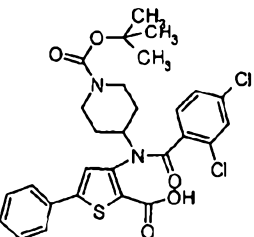
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337		3-[(4-CHLORO-2,5-DIFLUORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
338		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(2-METHYL-ALLYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
339		3-{ALLYL-[2-(4-CHLORO-PHENYL)-ACETYL]-AMINO}-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
340		3-[BENZYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
341		3-[(4-CHLORO-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

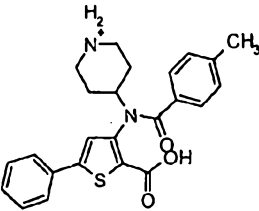
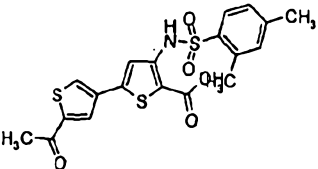
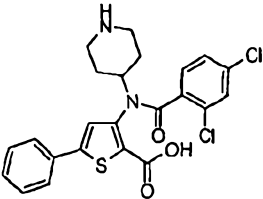
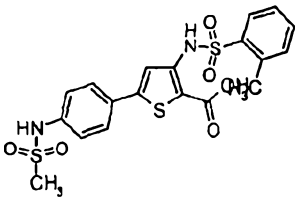
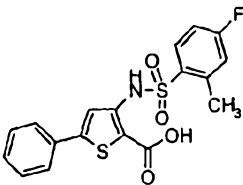
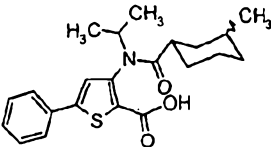
	MOLSTRUCTURE	COMPOUND NAME	IC50
342		3-[(4-METHYL-BENZOYL)-(4-NITRO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
343		3-[(4-METHYL-BENZOYL)-(2-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
344		3-[(3-METHOXY-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
345		3-[(2-CHLORO-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
346		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-ISOBUTYL-THIOPHENE-2-CARB OXYLIC ACID	+++

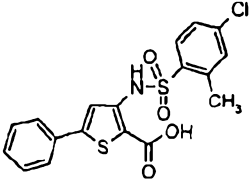
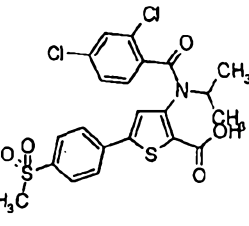
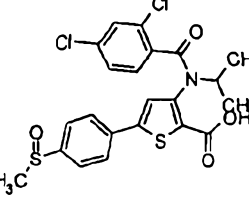
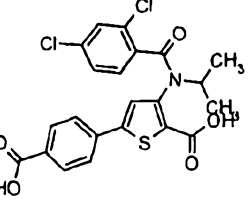
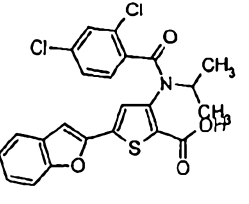
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347		3-[ALLYL-(2-NAPHTHALEN-2-YL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
348		3-(ALLYL-[2-(2,4-DICHLORO-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
349		3-(ALLYL-[2-(2-CHLORO-4-FLUORO-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
350		3-(ALLYL-[2-(3,4-DICHLORO-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
351		3-(ALLYL-[2-(2,4-DIFLUORO-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
352		3-{ALLYL-[2-(4-TRIFLUOROMETHYL-PHENYL)-ACETYL]-AMINO}-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
353		3-{ALLYL-[2-(2,6-DICHLORO-PHENYL)-ACETYL]-AMINO}-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
354		3-[ALLYL-(2-M-TOLYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
355		5-(4-ACETYL-PHENYL)-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++

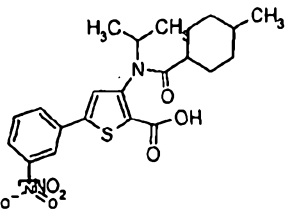
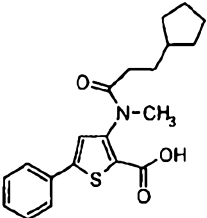
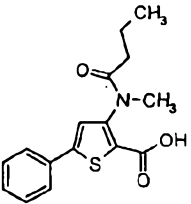
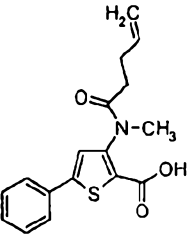
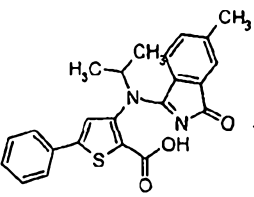
	MOLSTRUCTURE	COMPOUND NAME	IC50
356		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
357		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID	+++
358		5'-ACETYL-4-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID	+++
359		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
360		4-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5'-METHYL-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID	+++
361		3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-METHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++

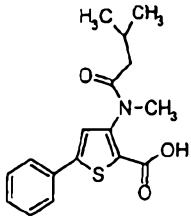
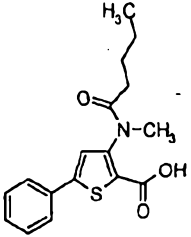
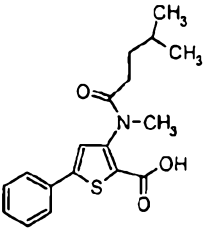
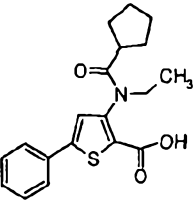
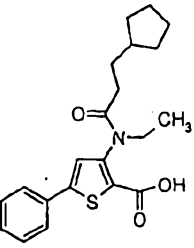
	MOLSTRUCTURE	COMPOUND NAME	IC50
362		3-(CYCLOHEXANECARBONYL-ISOPROPYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
363		3-((2,4-DICHLORO-BENZOYL)-[1-(2,4-DICHLORO-BENZOYL)-PIPERIDIN-4-YL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
364		4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(4-METHYL-BENZOYL)-AMINO]-PIPERIDINE-1-CARBOXYLIC ACID #TERTI-BUTYL ESTER	+++
365		4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-PIPERIDINE-1-CARBOXYLIC ACID #TERTI-BUTYL ESTER	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
366		3-[(4-METHYL-BENZOYL)-PIPERIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
367		5'-ACETYL-4-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-[2,3']BITHIOPHENYL-5-CARBOXYLIC ACID	+++
368		3-[(2,4-DICHLORO-BENZOYL)-PIPERIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
369		5-(4-METHANESULFONYLAMINO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++
370		3-(4-FLUORO-2-METHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
371		3-[(3-METHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+

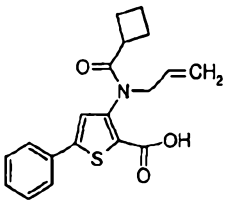
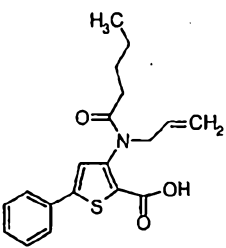
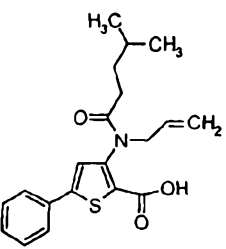
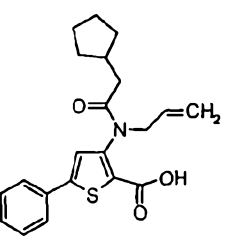
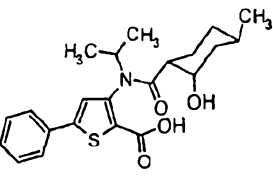
	MOLSTRUCTURE	COMPOUND NAME	IC50
372		3-(4-CHLORO-2-METHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
373		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(4-METHANESULFONYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
374		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(4-METHANESULFINYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
375		5-(4-CARBOXY-PHENYL)-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
376		5-BENZOFURAN-2-YL-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
377		3-[(2-ACETOXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
378		3-[ISOPROPYL-(2-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
379		3-[ISOPROPYL-(2-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
380		3-(CYCLOHEPTANECARBONYL-ISOPROPYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
381		3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
382		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-METHYL-THIOPHENE-2-CARBOXYLIC ACID	+++

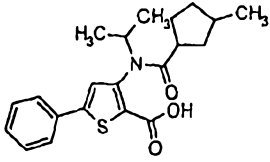
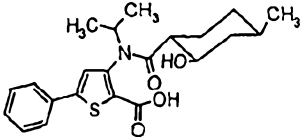
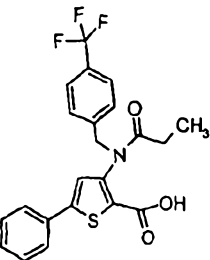
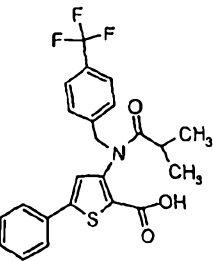
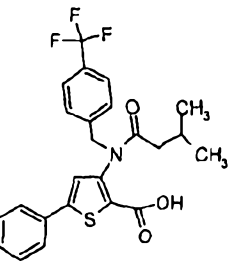
	MOLSTRUCTURE	COMPOUND NAME	IC50
383		3-[(ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-(3-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
384		3-[(3-CYCLOPENTYL-PROPIONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
385		3-(BUTYRYL-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
386		3-(METHYL-PENT-4-ENOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
387		3-[(ISOPROPYL-(5-METHYL-3-OXO-3H-INDOL-1-YL)-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

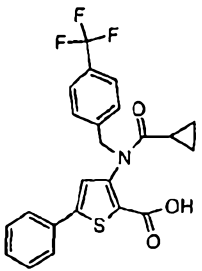
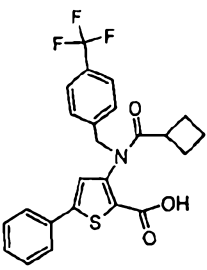
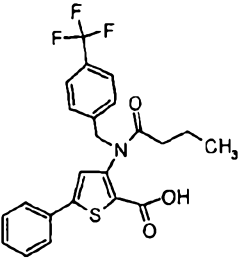
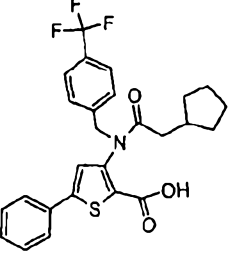
	MOLSTRUCTURE	COMPOUND NAME	IC50
388		3-[METHYL-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
389		3-(METHYL-PENTANOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
390		3-[METHYL-(4-METHYL-PENTANOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
391		3-(CYCLOPENTANECARBONYL-ETHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
392		3-[(3-CYCLOPENTYL-PROPIONYL)-ETHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

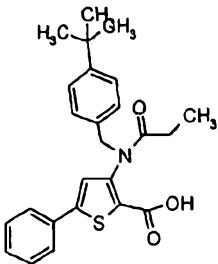
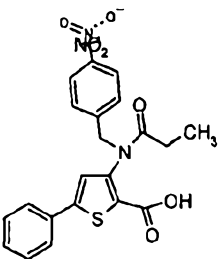
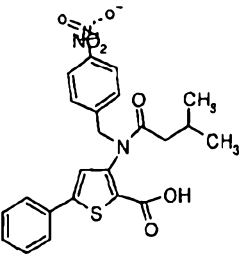
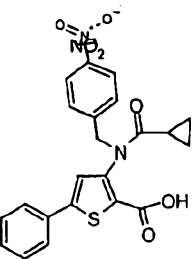
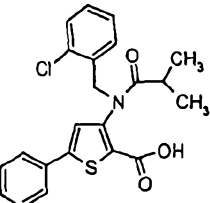
	MOLSTRUCTURE	COMPOUND NAME	IC50
393		3-(CYCLOBUTANECARBONYL-ETHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
394		3-(BUT-2-ENOYL-ETHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
395		3-[ISOPROPYL-(4-METHYL-2-VINYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
396		3-[ISOPROPYL-(4-METHYL-CYCLOHEX-1-ENECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
397		3-(ALLYL-HEXANOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

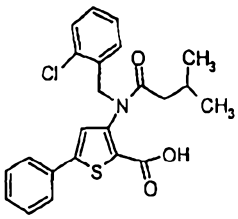
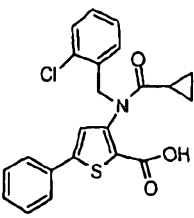
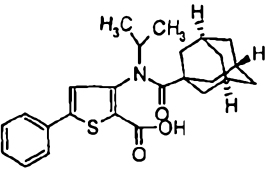
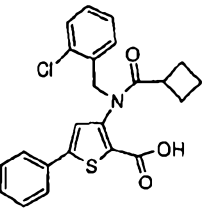
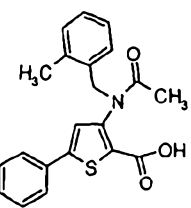
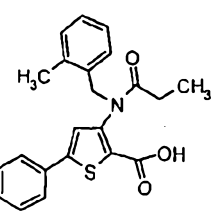
	MOLSTRUCTURE	COMPOUND NAME	IC50
398		3-(ALLYL-CYCLOBUTANECARBONYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
399		3-(ALLYL-PENTANOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
400		3-[ALLYL-(4-METHYL-PENTANOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
401		3-[ALLYL-(2-CYCLOPENTYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
402		3-[(2-HYDROXY-4-METHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

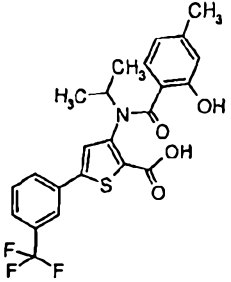
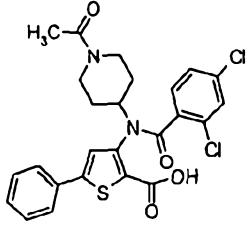
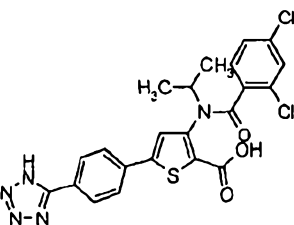
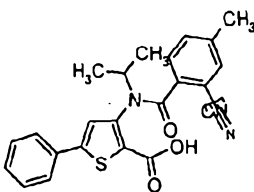
	MOLSTRUCTURE	COMPOUND NAME	IC50
403		3-[(2,4-DICHLORO-BENZOYL)-(1-PHENYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
404		3-[(2,4-DICHLORO-BENZOYL)-(1-PHENYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
405		3-[ISOPROPYL-(3-METHYL-CYCLOPENT-3-ENECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
406		3-[(2-BENZYLOXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
407		3-[(2,4-DIMETHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

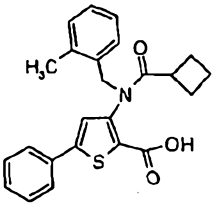
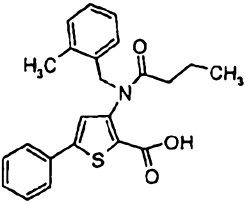
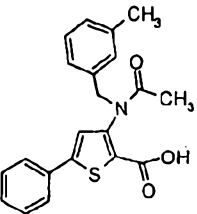
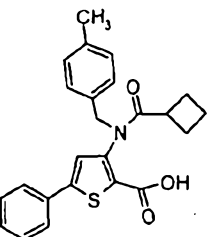
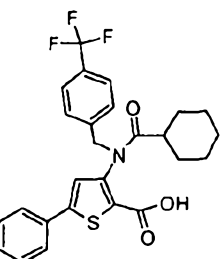
	MOLSTRUCTURE	COMPOUND NAME	IC50
408		3-[[ISOPROPYL-(3-METHYL-CYCLOPENTANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
409		3-[[2-HYDROXY-4-METHYL-CYCLOHEXANECARBONYL]-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
410		5-PHENYL-3-[[PROPIONYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	++
411		3-[[ISOBUTYRYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
412		3-[[3-METHYL-BUTYRYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

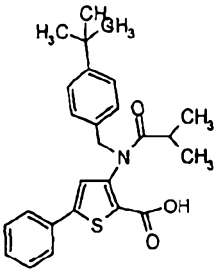
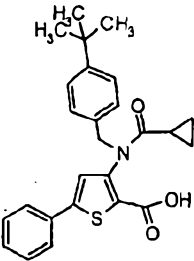
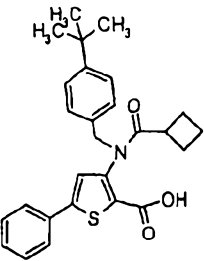
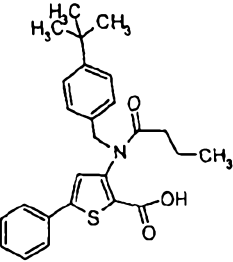
	MOLSTRUCTURE	COMPOUND NAME	IC50
413		3-[(CYCLOPROPANECARBONYL)-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
414		3-[(CYCLOBUTANECARBONYL)-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
415		3-[(BUTYRYL)-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
416		3-[(2-CYCLOPENTYL-ACETYL)-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

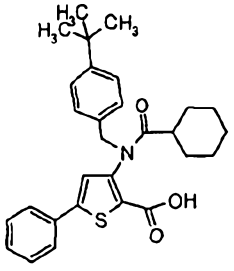
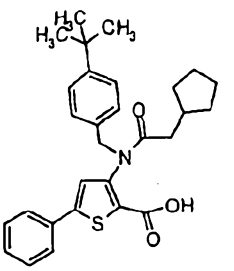
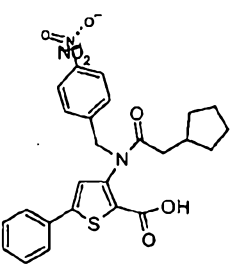
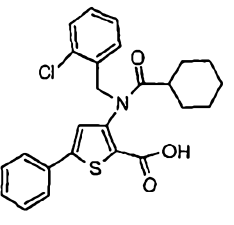
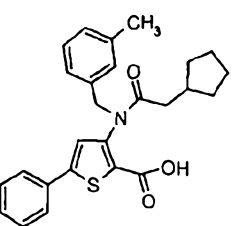
	MOLSTRUCTURE	COMPOUND NAME	IC50
417		3-[(4-TERT-BUTYL-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
418		3-[(4-NITRO-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
419		3-[(3-METHYL-BUTYRYL)-(4-NITRO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
420		3-[CYCLOPROPANECARBONYL-(4-NITRO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
421		3-[(2-CHLORO-BENZYL)-ISOBUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

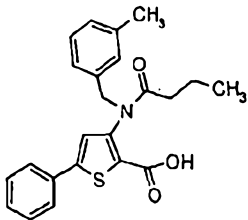
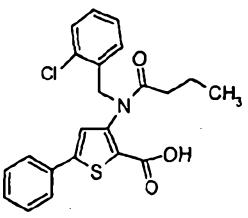
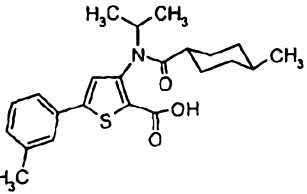
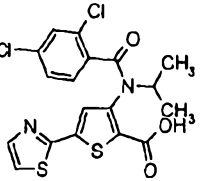
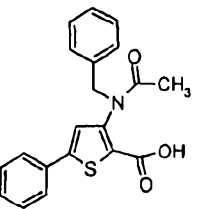
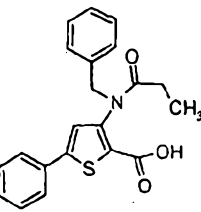
	MOLSTRUCTURE	COMPOUND NAME	IC50
422		3-[(2-CHLORO-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
423		3-[(2-CHLORO-BENZYL)-CYCLOPROPANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
424		3-[(ADAMANTANE-1-CARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
425		3-[(2-CHLORO-BENZYL)-CYCLOBUTANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
426		3-[ACETYL-(2-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
427		3-[(2-METHYL-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

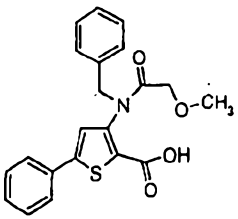
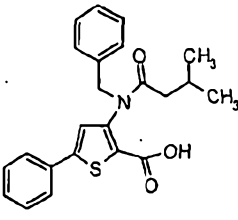
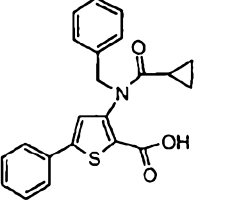
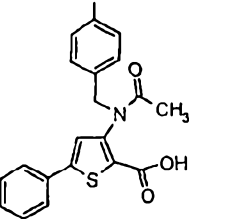
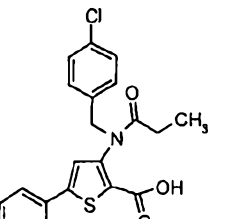
	MOLSTRUCTURE	COMPOUND NAME	IC50
428		3-[(2-HYDROXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	++
429		3-[(1-ACETYL-PIPERIDIN-4-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
430		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-[4-(1H-TETRAZOL-5-YL)-PHENYL]-THIOPHENE-2-CARBOXYLIC ACID	+++
431		3-[(2-CYANO-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

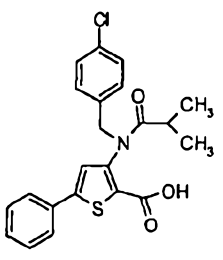
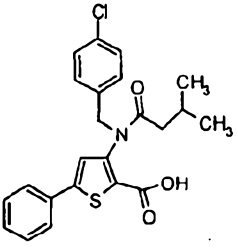
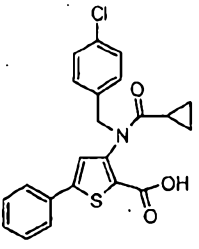
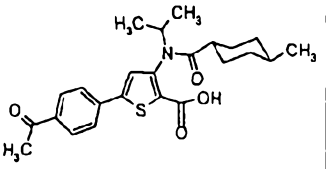
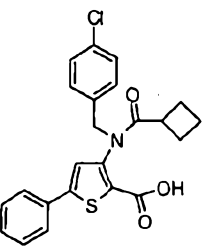
	MOLSTRUCTURE	COMPOUND NAME	IC50
432		3-[CYCLOBUTANECARBONYL-(2-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
433		3-[BUTYRYL-(2-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
434		3-[ACETYL-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
435		3-[CYCLOBUTANECARBONYL-(4-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
436		3-[CYCLOHEXANECARBONYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

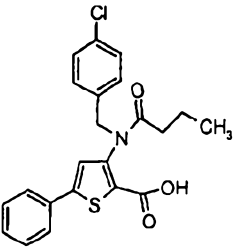
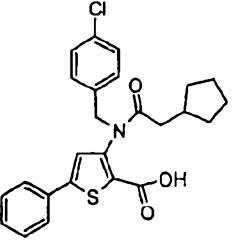
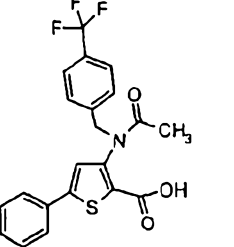
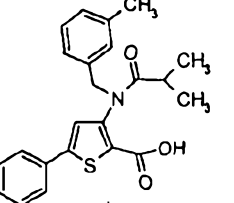
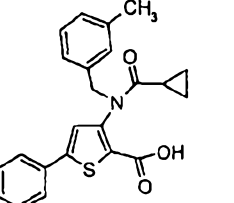
	MOLSTRUCTURE	COMPOUND NAME	IC50
437		3-[(4-TERT-BUTYL-BENZYL)- ISOBUTYRYL-AMINO]-5-PHENYL- THIOPHENE-2-CARBOXYLIC ACID	++
438		3-[(4-TERT-BUTYL-BENZYL)- CYCLOPROPANECARBONYL-AMINO]-5- PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
439		3-[(4-TERT-BUTYL-BENZYL)- CYCLOBUTANECARBONYL-AMINO]-5- PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
440		3-[(4-TERT-BUTYL-BENZYL)-BUTYRYL- AMINO]-5-PHENYL-THIOPHENE-2- CARBOXYLIC ACID	++

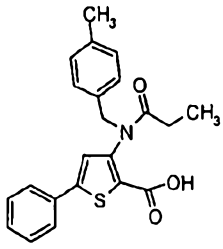
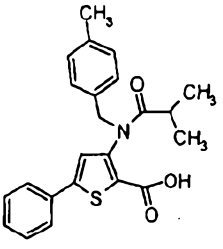
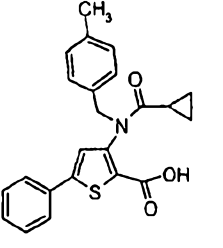
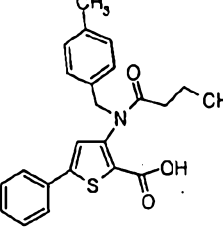
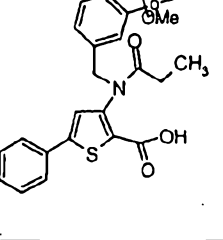
	MOLSTRUCTURE	COMPOUND NAME	IC50
441		3-[(4-TERT-BUTYL-BENZYL)-CYCLOHEXANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
442		3-[(4-TERT-BUTYL-BENZYL)-(2-CYCLOPENTYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
443		3-[(2-CYCLOPENTYL-ACETYL)-(4-NITRO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
444		3-[(2-CHLORO-BENZYL)-CYCLOHEXANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
445		3-[(2-CYCLOPENTYL-ACETYL)-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
446		3-[BUTYRYL-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
447		3-[BUTYRYL-(2-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
448		3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID	+++
449		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-THIAZOL-2-YL-THIOPHENE-2-CARBOXYLIC ACID	+++
450		3-(ACETYL-BENZYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
451		3-(BENZYL-PROPIONYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

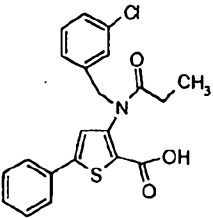
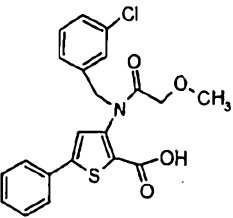
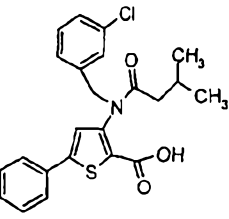
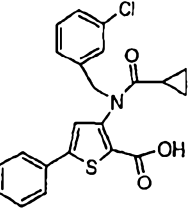
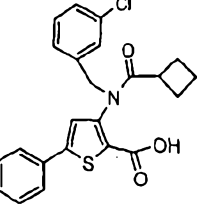
	MOLSTRUCTURE	COMPOUND NAME	IC50
452		3-[BENZYL-(2-METHOXY-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
453		3-[BENZYL-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
454		3-(BENZYL-CYCLOPROPANECARBONYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
455		3-[ACETYL-(4-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
456		3-[(4-CHLORO-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

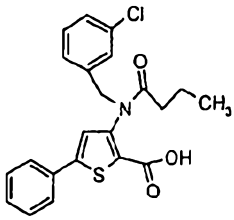
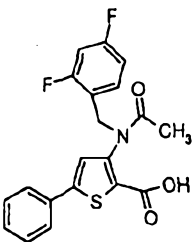
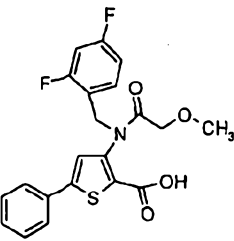
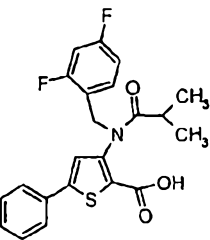
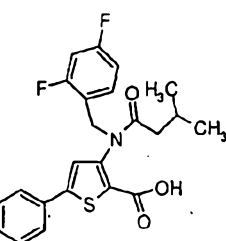
	MOLSTRUCTURE	COMPOUND NAME	IC50
457		3-[(4-CHLORO-BENZYL)-ISOBUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
458		3-[(4-CHLORO-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
459		3-[(4-CHLORO-BENZYL)-CYCLOPROPANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
460		5-(4-ACETYL-PHENYL)-3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
461		3-[(4-CHLORO-BENZYL)-CYCLOBUTANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

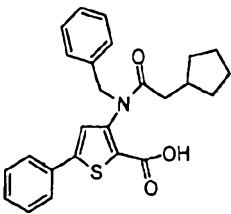
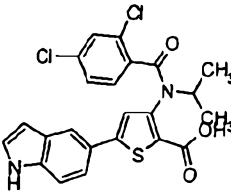
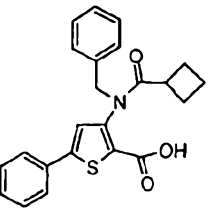
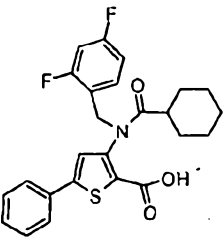
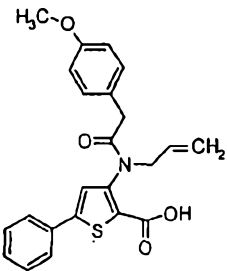
	MOLSTRUCTURE	COMPOUND NAME	IC50
462		3-[BUTYRYL-(4-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
463		3-[(4-CHLORO-BENZYL)-(2-CYCLOPENTYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
464		3-[ACETYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
465		3-[ISOBUTYRYL-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
466		3-[CYCLOPROPANECARBONYL-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
467		3-[(4-METHYL-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
468		3-[ISOBUTYRYL-(4-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
469		3-[CYCLOPROPANECARBONYL-(4-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
470		3-[BUTYRYL-(4-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
471		3-[(3-METHOXY-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+

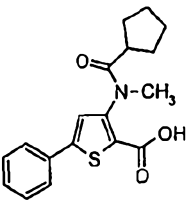
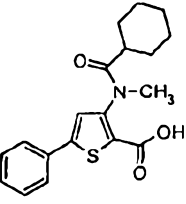
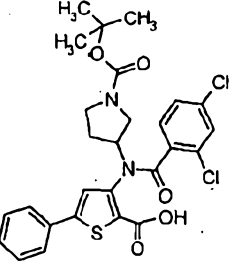
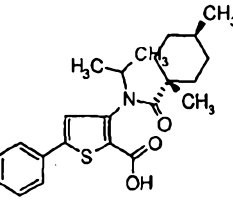
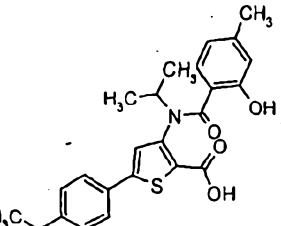
	MOLSTRUCTURE	COMPOUND NAME	IC50
472		3-[(3-METHOXY-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
473		3-[CYCLOBUTANECARBONYL-(3-METHOXY-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
474		3-[(2-CARBAMOYL-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
475		3-[BUTYRYL-(3-METHOXY-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
476		3-[ACETYL-(3-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+

	MOLSTRUCTURE	COMPOUND NAME	IC50
477		3-[(3-CHLORO-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
478		3-[(3-CHLORO-BENZYL)-(2-METHOXY-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
479		3-[(3-CHLORO-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
480		3-[(3-CHLORO-BENZYL)-CYCLOPROPANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
481		3-[(3-CHLORO-BENZYL)-CYCLOBUTANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
482		3-[BUTYRYL-(3-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
483		3-[ACETYL-(2,4-DIFLUORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
484		3-[(2,4-DIFLUORO-BENZYL)-(2-METHOXY-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
485		3-[(2,4-DIFLUORO-BENZYL)-ISOBUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
486		3-[(2,4-DIFLUORO-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
487		3-[BENZYL-(2-CYCLOPENTYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
488		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(1H-INDOL-5-YL)-THIOPHENE-2-CARBOXYLIC ACID	+++
489		3-(BENZYL-CYCLOBUTANECARBONYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
490		3-[CYCLOHEXANECARBONYL-(2,4-DIFLUORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
491		3-(ALLYL-[2-(4-METHOXY-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+

	MOLSTRUCTURE	COMPOUND NAME	IC50
492		3-(ETHYL-HEXANOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
493		3-(BUTYRYL-ETHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
494		3-[ETHYL-(4-METHYL-PENTANOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
495		3-[CYCLOBUTANECARBONYL-(2,4-DIFLUORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
496		3-[BUTYRYL-(2,4-DIFLUORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
497		3-(CYCLOPENTANECARBONYL-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
498		3-(CYCLOHEXANECARBONYL-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
499		3-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-PYRROLIDINE-1-CARBOXYLIC ACID #TERTI-BUTYL ESTER	+++
500		3-[(1,4-DIMETHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
501		5-(4-ETHYL-PHENYL)-3-[(2-HYDROXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
502		3-[(2-HYDROXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-(4-METHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
503		3-[(2,4-DICHLORO-BENZOYL)-PYRROLIDIN-3-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
504		4-[5-CARBOXY-4-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHEN-2-YL]-3,6-DIHYDRO-2H-PYRIDINE-1-CARBOXYLIC ACID BENZYL ESTER	+++
505		3-[[2-(HYDROXYIMINO-METHYL)-4-METHYL-BENZOYL]-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
506		3-[(1-CARBAMIMIDOYL-PIPERIDIN-4-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
507		4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-AZEPANE-1-CARBOXYLIC ACID TERTI-BUTYL ESTER	+++
508		4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-METHYL-PIPERIDINE-1-CARBOXYLIC ACID BENZYL ESTER	+++
509		3-[AZEPAN-4-YL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
510		3-[(4-METHYL-CYCLOHEXANECARBONYL)-PIPERIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID LITHIUM SALT	++
511		3-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-PIPERIDINE-1-CARBOXYLIC ACID #TERTI-BUTYL ESTER	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
512		3-[(4-BENZYLOXYCARBONYLAMINO-CYCLOHEXYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
513		3-[ISOPROPYL-(4-METHYL-2-OXO-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
514		3-[(2,4-DICHLORO-BENZOYL)-PIPERIDIN-3-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID; COMPOUND WITH GENERIC INORGANIC NEUTRAL COMPONENT	+++
515		3-[(4-BENZYLOXYCARBONYLAMINO-CYCLOHEXYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
516		3-[(2-BENZYLOXY-1-METHYL-ETHYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
517		3-[(2,2-DIMETHYL-[1,3]DIOXAN-5-YL)-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
518		3-[(2,4-DICHLORO-BENZOYL)-(2-HYDROXY-1-HYDROXYMETHYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
519		3-[(2,4-DICHLORO-BENZOYL)-PIPERIDIN-4-YLMETHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
520		3-[(2-CHLORO-BENZOYL)-PIPERIDIN-4-YLMETHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
521		3-[(4,6-DICHLORO-1#HI-INDOLE-2-CARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
522		3-[(2,4-DICHLORO-BENZOYL)-(2-HYDROXY-1-METHYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
523		4-{1-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-ETHYL}-PIPERIDINE-1-CARBOXYLIC ACID BENZYL ESTER	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
524		4-{5-CARBOXY-4-[(ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHEN-2-YL}-3,6-DIHYDRO-2#H]-PYRIDINE-1-CARBOXYLIC ACID BENZYL ESTER	+++
525		3-[(4-METHYL-CYCLOHEXANECARBONYL)-PYRIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
526		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PIPERIDIN-4-YL-THIOPHENE-2-CARBOXYLIC ACID; COMPOUND WITH TRIFLUORO-ACETIC ACID	+
527		3-[ISOPROPYL-(4-PROPYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
528		4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-CYCLOHEXYL-AMMONIUM; TRIFLUORO-ACETATE	+++
529		3-[(2,4-DICHLORO-BENZOYL)-(1-PIPERIDIN-4-YL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID; COMPOUND WITH TRIFLUORO-ACETIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
530		3-[(CYCLOHEX-3-ENECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
531		3-[(4-ETHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
532		3-[(4-CHLORO-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
533		4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-3-METHYL-PIPERIDINE-1-CARBOXYLIC ACID BENZYL ESTER	+++
534		3-[(2,4-DICHLORO-BENZOYL)-(2-METHOXY-CYCLOHEXYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
535		3-[(2,4-DICHLORO-BENZOYL)-(2,2-DIMETHYL-[1,3]DIOXAN-5-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
536		3-[[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-(1-METHYL-PIPERIDIN-4-YL)-THIOPHENE-2-CARBOXYLIC ACID	+++
537		3-[(2,4-DICHLORO-BENZOYL)-(3-METHYL-PIPERIDIN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID; COMPOUND WITH TRIFLUORO-ACETIC ACID	+++
538		3-[(2,4-DICHLORO-BENZOYL)-(2-HYDROXY-CYCLOHEXYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
539		4-[[[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-METHYL]-PIPERIDINE-1-CARBOXYLIC ACID BENZYL ESTER	+++
540		3-[[[(1R,2S,4R)-2-HYDROXY-4-METHYL-CYCLOHEXANECARBONYL]-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
541		3-[[ISOPROPYL-[1-(4-METHOXY-2,3,6-TRIMETHYL-BENZENESULFONYL)-5-METHYL-1,2,3,6-TETRAHYDRO-PYRIDINE-2-CARBONYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
542		3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-4-FLUORO-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
543		3-[(2,4-DICHLORO-BENZOYL)-(1-METHYL-PIPERIDIN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
544		4-[[[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-METHYL]-PIPERIDIUM; TRIFLUORO-ACETATE	+++
545		3-[(2-TERT-BUTOXYCARBONYLAMINO-1-METHYL-ETHYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
546		2-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-PROPYL-AMINE TRIFLUOROACETIC ACID SALT	+++

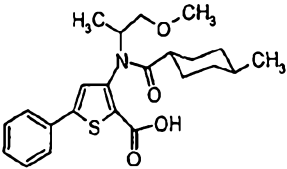
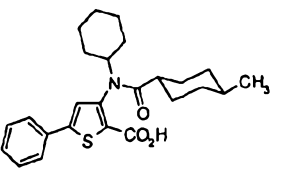
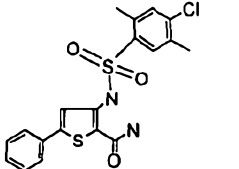
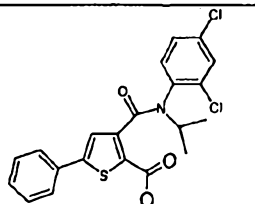
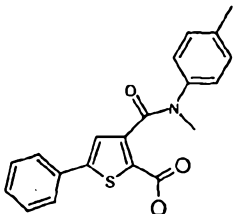
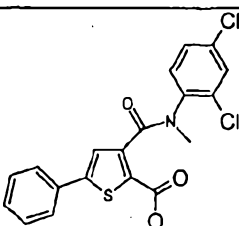
	MOLSTRUCTURE	COMPOUND NAME	IC50
547		3-[(3-CARBOXY-CYCLOPENTYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
548		3-[(3-CARBOXY-CYCLOPENTYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
549		2-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-CYCLOHEXYL-AMMONIUM; CHLORIDE	+++
550		3-(BENZOYL-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
551		[(5-PHENYL-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBONYL)-AMINO]-ACETIC ACID	++
552		5-BROMO-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
553		3-[CYCLOHEXYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
554		3-[[1,3]DIOXAN-5-YL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
555		3-[[2-(TERT-BUTYL-DIMETHYL-SILANYLOXY)-1-METHYL-2-PHENYL-ETHYL]-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
556		3-[[2-(TERT-BUTYL-DIMETHYL-SILANYLOXY)-1-METHYL-2-PHENYL-ETHYL]-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
557		3-[(2,4-DICHLORO-BENZOYL)-(2-DIETHYLAMINO-THIAZOL-5-YLMETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
558		(5-[[[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-METHYL]-THIAZOL-2-YL]-DIETHYL-AMMONIUM; CHLORIDE	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
559		5-(4-FLUORO-PHENYL)-3-[(ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
560		3-[[[(1S,2R,4S)-2-HYDROXY-4-METHYL-CYCLOHEXANECARBONYL]-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
561		3-[[[2,4-DICHLORO-BENZOYL]-(2-METHOXY-1-METHYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
562		3-[[[4S]-ISOPROPYL-(4-METHYL-CYCLOHEX-1-ENECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
563		5-(4-CHLORO-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID AMIDE	++
564		5-(4-FLUORO-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID AMIDE	++

	MOLSTRUCTURE	COMPOUND NAME	IC50
565		5-(4-METHOXY-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID AMIDE	++
566		3-METHYL-(4-METHYLBENZOYL)-AMINO)5-PHENYL THIOPHENE-2-CARBOXYLIC ACID (2-HYDROXY-ETHYL)AMIDE	++
567		5-PHENYL-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID CYCLOBUTYLAMIDE	++
568		3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID AMIDE	++
569		5-BROMO-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++
570		5-(4-CHLORO-PHENYL)-3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID	+++

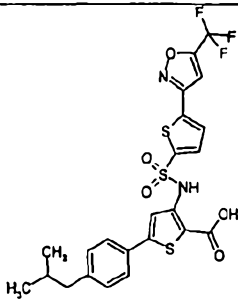
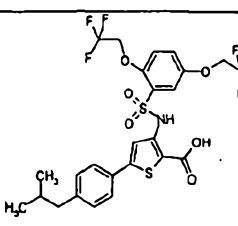
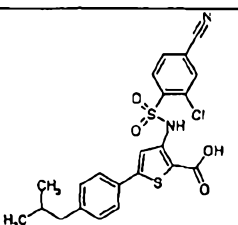
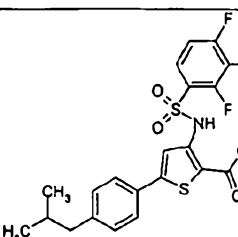
	MOLSTRUCTURE	COMPOUND NAME	IC50
571		5-(4'-CHLORO-BIPHENYL-4-YL)-3-[(ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO)]-THIOPHENE-2-CARBOXYLIC ACID	+++
572		3-[(4-METHYL-CYCLOHEXANECARBONYL)-(TETRAHYDRO-PYRAN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
573		3-[(4-METHYL-CYCLOHEXANECARBONYL)-(1-METHYL-PIPERIDIN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
574		3-[(4-METHYL-CYCLOHEXANECARBONYL)-PIPERIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+
575		3-[(ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO)]-5-(4-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++
576		5-(4-CYANO-PHENYL)-3-[(ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO)]-THIOPHENE-2-CARBOXYLIC ACID	+++
577		3-[(ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO)]-5-(4-METHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID	+++

	MOLSTRUCTURE	COMPOUND NAME	IC50
578		3-[(2-METHOXY-1-METHYL-ETHYL)-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
579		3-[CYCLOHEXYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
580		3-(4-CHLORO-2,5 DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID AMIDE	+++
583		3-[(2,4-DICHLORO-PHENYL)-ISOPROPYL-CARBAMOYL]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	+++
584		3-(METHYL-P-TOLYL-CARBAMOYL)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++
585		3-[(2,4-DICHLORO-PHENYL)-METHYL-CARBAMOYL]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID	++

* : +++ IC₅₀ <5μM
 ++ IC₅₀ 5μM-20μM
 + IC₅₀ >20μM

EXAMPLE 28 List of compounds having anti-helicase activity *

Compound #	Compound name	Structure	Anti-ATPase (Malachite Green assay) EC ₅₀	Anti-ATPase (HPLC method) EC ₅₀
Compound #14	3-(4-Chloro-2,5-dimethyl-benzenesulfonylamin o)-5-(4-chloro-phenyl)-thiophene-2-carboxylic acid		+	ND
Compound #19	3-(4-Chloro-2,5-dimethyl-benzenesulfonylamin o)-5-(4-isobutyl-phenyl)-thiophene-2-carboxylic acid		+++	++
Compound #223	3-(4-Bromo-2-fluorobenzenesulfo-nylamino)-5-(4-isobutylphenyl)-thiophene-2-carboxylic acid		ND	+++
Compound #224	3-(4-Bromo-2-methylbenzenesulfo-nylamino)-5-(4-isobutylphenyl)-thiophene-2-carboxylic acid		ND	++
Compound #225	5-(4-Isobutylphenyl) 3-(3-methoxy-benzenesulfonyl-amino)-thiophene-2-carboxylic acid		ND	+

Compound #581	5-(4-Isobutyl-phenyl)-3-[5-(5-trifluoromethyl-isoxazol-3-yl)-thiophene-2-sulfonylamino]-thiophene-2-carboxylic acid		ND	++
Compound #227	3-[2,5-Bis-(2,2,2-trifluoroethoxy)-benzenesulfonylamino]-5-(4-isobutyl-phenyl)-thiophene-2-carboxylic acid		ND	+
Compound #228	3-(2-Chloro-4-cyanobenzenesulfonylamino)-5-(4-isobutylphenyl)-thiophene-2-carboxylic acid		ND	+
Compound #582	5-(4-Isobutyl-phenyl)-3-(2,3,4-trifluoro-benzenesulfonylamino)-thiophene-2-carboxylic acid		ND	+

*: +++ IC₅₀ <5μM
 ++ IC₅₀ 5μM-20μM
 + IC₅₀ >20μM

It is to be understood that, if any prior art publication is referred to herein, such reference does not constitute an admission that the publication forms a part of the common general knowledge in the art, in Australia or any other country.

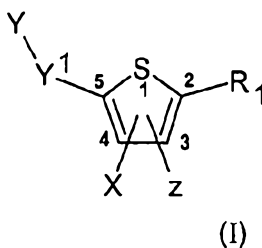
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In the claims which follow and in the preceding description of the invention, except where the context requires otherwise due to express language or necessary implication, the word

10 "comprise" or variations such as "comprises" or "comprising" is used in an inclusive sense, i.e. to specify the presence of the stated features but not to preclude the presence or addition of further features in various embodiments of the invention.

We claim:

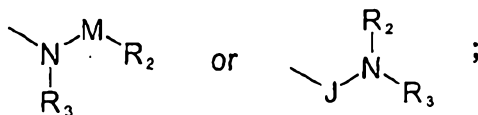
1. A compound having the formula I:



or pharmaceutically acceptable salts thereof;

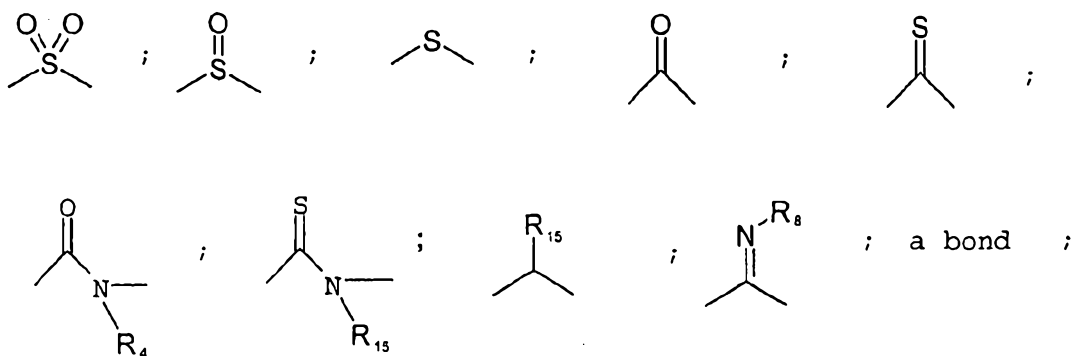
wherein,

X is chosen from:



wherein,

M is chosen from:



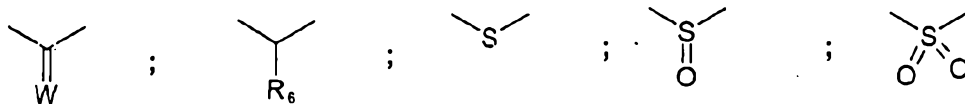
wherein,

R₄ is C₁₋₆ alkyl;

R₈ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₂ heteroaralkyl, C₆₋₁₆ aralkyl; and

R₁₅ is chosen from H or C₁₋₆ alkyl;

J is chosen from:



wherein, W is chosen from O, S or NR₇,

wherein R₇ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₂ heteroaralkyl, C₆₋₁₆ aralkyl;

and R₆ is chosen from H, C₁₋₁₂ alkyl, C₆₋₁₄ aryl or C₆₋₁₆ aralkyl;

Y¹ is chosen from a bond, C₁₋₆ alkyl, C₂₋₆ alkenyl or C₂₋₆ alkynyl;

Y is chosen from COOR₁₆, COCOOR₅, P(O)OR₄OR₆, S(O)OR₅, S(O)₂OR₅, tetrazole, CON(R₉)CH(R₅)COOR₅, CONR₁₀R₁₁, CON(R₉)-SO₂-R₅, CONR₉OH or halogen, wherein R₉, R₅, R₁₀ and R₁₁ are each independently chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl;

or R₁₀ and R₁₁ are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R₄ and R₆ are each independently chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl and C₆₋₁₈ aralkyl;

or R₄ and R₆ are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R₁₆ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl and C₆₋₁₈ aralkyl; provided that R₁₆ is other than methyl or ethyl;

R₁ is chosen from C₂₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl or C₆₋₁₈ aralkyl;

R₂ is chosen from C₂₋₁₂ alkyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, or C₆₋₁₈ aralkyl;

R_3 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-10} aralkyl;

Z is chosen from H, halogen, C_{1-6} alkyl;

with the proviso that:

i) when X is 4-Chloro-2,6-dimethyl-benzenesulfonamide and, R_1 is phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than $CONH_2$; compound #580

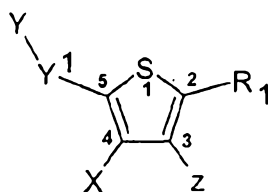
ii) when X is Toluene-4-sulfonamide and R_1 is 4-chloro-phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than $CONH_2$; compound #563

iii) when X is Toluene-4-sulfonamide and R_1 is 4-fluoro-phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than $CONH_2$; compound #564

iv) when X is Toluene-4-sulfonamide and R_1 is 4-methoxy-phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than $CONH_2$; compound #565

v) when X is Benzamide and R_1 is phenyl Y^1 is a bond and Y is $COOH$ then R_3 is other than hydrogen.

2. A compound having the formula Ia:

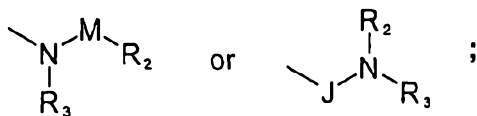


(Ia)

or pharmaceutically acceptable salts thereof;

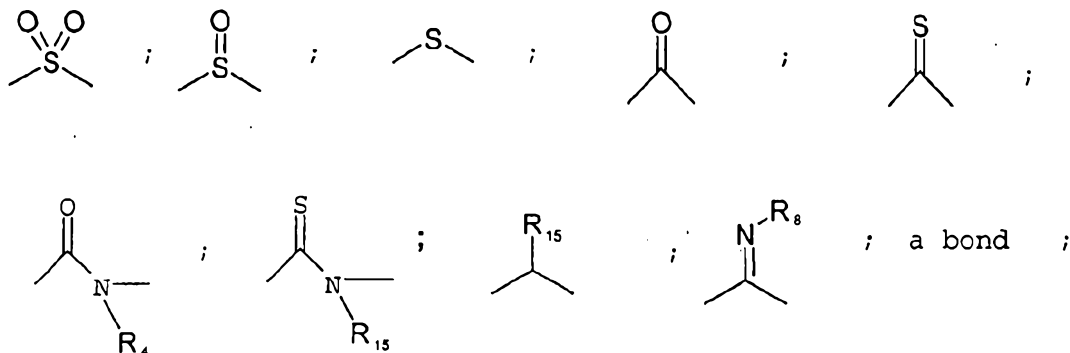
wherein,

X is chosen from:



wherein,

M is chosen from:



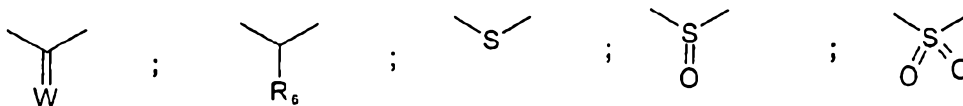
wherein,

R_4 is C_{1-6} alkyl;

R_8 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl; and

R_{15} is chosen from H or C_{1-6} alkyl;

J is chosen from:



wherein W is chosen from O, S or NR_7 ,

wherein R_7 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl;

and R_6 is chosen from H, C_{1-12} alkyl, C_{6-14} aryl or C_{6-16} aralkyl;

Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

Y is chosen from $COOR_{16}$, $COCOOR_5$, $P(O)OR_aOR_b$, $S(O)OR_5$, $S(O)_2OR_5$, tetrazole, $CON(R_9)CH(R_3)COOR_5$, $CONR_{10}R_{11}$, $CON(R_9)-SO_2-R_5$, $CONR_3OH$ or halogen, wherein R_9 , R_5 , R_{10} and R_{11} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;

or R_{10} and R_{11} are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_a and R_b are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

or R_a and R_b are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R_{16} is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl; provided that R_{16} is other than methyl or ethyl;

R_1 is chosen from C_{2-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

R_2 is chosen from C_{2-12} alkyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, or C_{6-18} aralkyl;

R_3 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

Z is chosen from H, halogen, C_{1-6} alkyl;

with the proviso that:

i) when X is 4-Chloro-2,6-dimethyl-benzenesulfonamide and, R_1 is phenyl, and R_3 is H, and Y^1 is a bond, then Y is other than $CONH_2$; compound #580

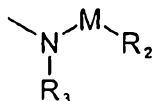
ii) when X is Toluene-4-sulfonamide and R_1 is 4-chlorophenyl, and R_2 is H, and Y^1 is a bond, then Y is other than CONH_2 ; compound #563

iii) when X is Toluene-4-sulfonamide and R_1 is 4-fluorophenyl, and R_2 is H, and Y^1 is a bond, then Y is other than CONH_2 ; compound #564

iv) when X is Toluene-4-sulfonamide and R_1 is 4-methoxyphenyl, and R_2 is H, and Y^1 is a bond, then Y is other than CONH_2 ; compound #565

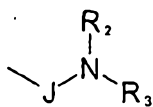
v) when X is Benzamide and R_1 is phenyl Y^1 is a bond and Y is COOH then R_3 is other than hydrogen.

3. A compound as defined in claims 1 or 2, wherein X is:



4. A compound as defined in claims 1 or 2, wherein

X is:

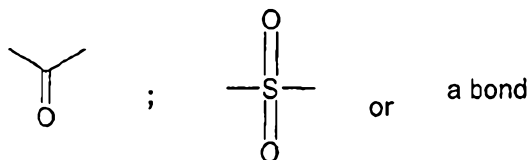


5. The compound as defined in claims 1 or 2, wherein Z is H.
6. The compound as defined in claims 1 or 2, wherein Y^1 is a bond.
7. The compound as defined in anyone of claims 1 or 2, wherein R_1 is chosen from C_{2-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl.

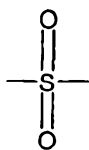
8. The compound as defined in anyone of claims 1 or 2, wherein R_1 is chosen from a C_{2-12} alkyl, C_{6-14} aryl or C_{3-12} heterocycle.
9. The compound as defined in anyone of claims 1 or 2, wherein R_1 is a C_{2-12} alkyl.
10. The compound as defined in anyone of claims 1 or 2, wherein R_1 is a C_{6-14} aryl.
11. The compound as defined in anyone of claims 1 or 2, wherein R_1 is a C_{3-12} heterocycle.
12. The compound as defined in anyone of claims 1 or 2, wherein R_1 is chosen from t-butyl, isobutyl, allyl, ethynyl, 2-phenylethenyl, isobutenyl, benzyl, phenyl, phenethyl, benzodioxolyl, thienyl, thiophenyl, pyridinyl, isoxazolyl, thiazolyl, pyrazolyl, tetrazolyl, benzofuranyl, indolyl, furanyl, or benzothiophenyl any of which can be optionally substituted by one or more substituent chosen from halogen, nitro, nitroso, SO_3R_{12} , PO_3RcRd , $CONR_{13}R_{14}$, $COOH$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-12} aralkyl, C_{6-12} aryl, C_{1-5} alkyloxy, C_{2-6} alkenyloxy, C_{2-6} alkynyloxy, C_{6-12} aryloxy, $C(O)C_{1-6}$ alkyl, $C(O)C_{2-6}$ alkenyl, $C(O)C_{2-6}$ alkynyl, $C(O)C_{6-12}$ aryl, $C(O)C_{6-12}$ aralkyl, C_{3-10} heterocycle, hydroxyl, $NR_{13}R_{14}$, $C(O)OR_{12}$, cyano, azido, amidino or guanido;
 wherein R_{12} , Rc , Rd , R_{13} and R_{14} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;
 or Rc and Rd are taken together with the oxygens to form a 5 to 10 membered heterocycle;
 or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

13. The compound as defined in claim 12, wherein R_1 is chosen from thienyl, t-butyl, phenyl or pyridinyl.
14. The compound as defined in claim 12, wherein R_1 is phenyl.
15. The compound as defined in anyone of claims 1 or 2, wherein R_1 is phenyl substituted by at least one fluoride.
16. The compound as defined in anyone of claims 1 or 2, wherein R_1 is phenyl substituted by at least one chloride.
17. The compound as defined in anyone of claims 1 or 2, wherein R_1 is phenyl substituted by at least one nitro.
18. The compound as defined in anyone of claims 1 or 2, wherein R_1 is phenyl substituted by at least one methyl.
19. The compound as defined in anyone of claims 1 or 2, wherein R_1 is phenyl substituted by at least one methoxy.
20. The compound as defined in anyone of claims 1 or 2, wherein R_1 is thienyl.
21. The compound as defined in anyone of claims 1 or 2, wherein R_1 is thienyl substituted by at least one halogen.
22. The compound as defined in anyone of claims 1 or 2, wherein R_1 is thienyl substituted by at least one chloride.
23. The compound as defined in anyone of claims 1 or 2, wherein R_1 is thienyl substituted by at least one methyl.

24. The compound as defined in anyone of claims 1 or 2, wherein R_1 is thienyl substituted by at least one methyl and one chloride.
25. The compound as defined in anyone of claims 1 or 2, wherein R_1 is isoxazolyl substituted by at least one methyl.
26. The compound as defined in anyone of claims 1 or 2, wherein R_1 is pyridinyl.
27. The compound as defined in anyone of claims 1 or 2, wherein M is chosen from:



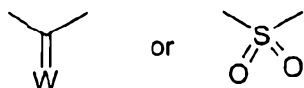
28. The compound as defined in anyone of claims 1 or 2, wherein M is:



29. The compound as defined anyone of claims 1 or 2, wherein M is:



30. The compound as defined in anyone of claims 1 or 2, wherein J is chosen from:



wherein W is as defined in claim 1.

31. The compound as defined in anyone of claims 1 or 2, wherein J is:



32. The compound as defined in anyone of claims 1 or 2, wherein J is:



33. The compound as defined in anyone of claims 1 or 2, wherein Y is chosen from COOR_{16} , COCOOR_5 , $\text{P(O)OR}_a\text{OR}_b$, $\text{S(O)}_2\text{OR}_5$, tetrazole, $\text{CON(R}_5\text{)CH(R}_5\text{)COOR}_5$, $\text{CONR}_{10}\text{R}_{11}$, CONR_5OH .
34. The compound as defined in claim 33, wherein any of R_5 , R_a , R_b , R_9 , R_{10} , R_{11} and R_{16} are each independently chosen from H or C_{1-6} alkyl; provided that R_{16} is other than methyl or ethyl.
35. The compound as defined in anyone of claims 1 or 2, wherein Y is chosen from COOR_{16} , $\text{CONR}_{10}\text{R}_{11}$ or $\text{CON(R}_5\text{)CH(R}_5\text{)-COOR}_5$.
36. The compound as defined in claim 33, wherein any of R_5 , R_9 , R_{10} , R_{11} and R_{16} are each independently chosen from H or C_{1-6} alkyl; provided that R_{16} is other than methyl or ethyl.
37. The compound as defined in anyone of claims 1 or 2, wherein Y is COOH .

38. The compound as defined in anyone of claims 1 or 2, wherein Y is $\text{CONHCH}_2\text{COOH}$.
39. The compound as defined in anyone of claims 1 or 2, wherein Y is COOCH_3 .
40. The compound as defined in anyone of claims 1 or 2, wherein Y is COONH_2 .
41. The compound as defined in anyone of claims 1 or 2, wherein R_3 is chosen from H, C_{1-12} alkyl, C_{6-18} aralkyl, C_{3-12} heterocycle or C_{3-18} heteroaralkyl.
42. The compound as defined in anyone of claims 1 or 2, wherein R_3 is chosen from H, C_{1-12} alkyl, C_{6-18} aralkyl or C_{3-12} heterocycle.
43. The compound as defined in anyone of claims 1 or 2, wherein R_3 is C_{1-12} alkyl.
44. The compound as defined in anyone of claims 1 or 2, wherein R_3 is C_{6-18} aralkyl.
45. The compound as defined in anyone of claims 1 or 2, wherein R_3 is C_{3-12} heterocycle.
46. The compound as defined in anyone of claims 1 or 2, wherein R_3 is chosen from H, methyl, ethyl, i-propyl, cyclopropyl, cyclohexyl, allyl, piperidinyl, piperazinyl, pyrrolidinyl, azetidiny, aziridinyl, pyridinyl, piperidinylmethyl, dioxanyl, azepanyl or benzyl; any of which can be optionally substituted by one or more substituent chosen from halogen, nitro, nitroso, SO_3R_{12} , PO_3RCRd , $\text{CONR}_{13}\text{R}_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-12} aralkyl, C_{6-12} aryl, C_{1-6} alkyloxy, C_{2-6} alkenyloxy,

C_{2-6} alkynyloxy, C_{6-12} aryloxy, $C(O)C_{1-6}$ alkyl, $C(O)C_{2-6}$ alkenyl, $C(O)C_{2-6}$ alkynyl, $C(O)C_{6-12}$ aryl, $C(O)C_{6-12}$ aralkyl, C_{3-10} heterocycle, hydroxyl, $NR_{13}R_{14}$, $C(O)OR_{12}$, cyano, azido, amidino or guanido; wherein R_{12} , R_c , R_d , R_{13} , and R_{14} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl; or R_c and R_d are taken together with the oxygens to form a 5 to 10 membered heterocycle; or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

47. The compound as defined in claim 43, wherein R_1 is chosen from H or Methyl, isopropyl, piperidinyl, piperidinylmethyl or cyclohexyl.
48. The compound as defined in anyone of claims 1 or 2, wherein R_1 is H.
49. The compound as defined in anyone of claims 1 or 2, wherein R_1 is Methyl.
50. The compound as defined in anyone of claims 1 or 2, wherein R_1 is C_{2-12} alkyl, C_{6-14} aryl or C_{3-12} heterocycle.
51. The compound as defined in anyone of claims 1 or 2, wherein R_1 is C_{3-6} heterocycle.
52. The compound as defined in anyone of claims 1 or 2, wherein R_1 is chosen from thienyl, furanyl, pyridinyl, oxazolyl, thiazolyl, pyrrolyl, benzofuranyl, indolyl, benzoxazolyl, benzothienyl, benzothiazolyl, piperazinyl, pyrrolidinyl or quinolinyl any of which can be optionally substituted by one or more substituent chosen from halogen, nitro, nitroso, SO_3R_{12} , $PO_3R_cR_d$, $CONR_{13}R_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-12}

- aralkyl, C₆₋₁₂ aryl, C₁₋₆ alkyloxy, C₂₋₆ alkenyloxy, C₂₋₆ alkynyloxy, C₆₋₁₂ aryloxy, C(O)C₁₋₆ alkyl, C(O)C₂₋₆ alkenyl, C(O)C₂₋₆ alkynyl, C(O)C₆₋₁₂ aryl, C(O)C₆₋₁₂ aralkyl, C₃₋₁₀ heterocycle, hydroxyl, NR₁₃R₁₄, C(O)OR₁₂, cyano, azido, amidino or guanido; wherein R₁₂, Rc, Rd, R₁₃ and R₁₄ are each independently chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl; or Rc and Rd are taken together with the oxygens to form a 5 to 10 membered heterocycle; or R₁₃ and R₁₄ are taken together with the nitrogen to form a 3 to 10 membered heterocycle.
53. The compound as defined in claim 49, wherein R₂ is a heterocycle chosen from thienyl, furanyl, pyridinyl, pyrrolyl, indolyl, piperazinyl or benzothienyl.
54. The compound as defined in anyone of claims 1 or 2, wherein R₂ is C₂₋₁₂ alkyl.
55. The compound as defined in anyone of claims 2 to 4, wherein R₂ is chosen from cyclopropyl, cyclobutyl, cyclopentyl, cyclopentenyl cyclohexyl, cycloheptyl, 2-(cyclopentyl)-ethyl, methyl, ethyl, vinyl, propyl, propenyl, isopropyl, butyl, butenyl isobutyl, pentyl, neopentyl or t-butyl any of which can be optionally substituted by one or more substituent chosen from halogen, nitro, nitroso, SO₃R₁₂, PO₃RcRd, CONR₁₃R₁₄, C₁₋₆ alkyl, C₂₋₆ alkenyl, C₂₋₆ alkynyl, C₆₋₁₂ aralkyl, C₆₋₁₂ aryl, C₁₋₆ alkyloxy, C₂₋₆ alkenyloxy, C₂₋₆ alkynyloxy, C₆₋₁₂ aryloxy, C(O)C₁₋₆ alkyl, C(O)C₂₋₆ alkenyl, C(O)C₂₋₆ alkynyl, C(O)C₆₋₁₂ aryl, C(O)C₆₋₁₂ aralkyl, C₃₋₁₀ heterocycle, hydroxyl, NR₁₃R₁₄, C(O)OR₁₂, cyano, azido, amidino or guanido;

wherein R_{12} , R_c , R_d , R_{13} and R_{14} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;

or R_c and R_d are taken together with the oxygens to form a 5 to 10 membered heterocycle;

or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

56. The compound as defined in anyone of claims 1 or 2, wherein R_2 is C_{6-12} aryl.
57. The compound as defined in anyone of claims 1 or 2, wherein R_2 is an aryl chosen from indenyl, naphthyl or biphenyl.
58. The compound as defined in anyone of claims 2 to 4, wherein R_2 is phenyl substituted by one or more substituent chosen from halogen, nitro, nitroso, SO_3R_{12} , $PO_3R_cR_d$, $CONR_{13}R_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-12} aralkyl, C_{6-12} aryl, C_{1-6} alkyloxy, C_{2-6} alkenyloxy, C_{2-6} alkynyloxy, C_{6-12} aryloxy, $C(O)C_{1-6}$ alkyl, $C(O)C_{2-6}$ alkenyl, $C(O)C_{2-6}$ alkynyl, $C(O)C_{6-12}$ aryl, $C(O)C_{6-12}$ aralkyl, C_{3-10} heterocycle, hydroxyl, $NR_{13}R_{14}$, $C(O)OR_{12}$, cyano, azido, amidino or guanido;
 wherein R_{12} , R_c , R_d , R_{13} and R_{14} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;
 or R_c and R_d are taken together with the oxygens to form a 5 to 10 membered heterocycle;
 or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.
59. The compound as defined in anyone of claims 1 or 2, wherein R_2 is phenyl substituted by one or two substituents chosen from

halogen, nitro, nitroso, SO_3R_{12} , PO_3RcRd , $\text{CONR}_{13}\text{R}_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-12} aralkyl, C_{6-12} aryl, C_{1-6} alkyloxy, C_{2-6} alkenyloxy, C_{2-6} alkynyloxy, C_{6-12} aryloxy, $\text{C}(\text{O})\text{C}_{1-6}$ alkyl, $\text{C}(\text{O})\text{C}_{2-6}$ alkenyl, $\text{C}(\text{O})\text{C}_{2-6}$ alkynyl, $\text{C}(\text{O})\text{C}_{6-12}$ aryl, $\text{C}(\text{O})\text{C}_{6-12}$ aralkyl, C_{3-10} heterocycle, hydroxyl, $\text{NR}_{13}\text{R}_{14}$, $\text{C}(\text{O})\text{OR}_{12}$, cyano, azido, amidino or guanido;

wherein R_{12} , Rc, Rd, R_{13} and R_{14} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;

or Rc and Rd are taken together with the oxygens to form a 5 to 10 membered heterocycle;

or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

60. The compound as defined in anyone of claims 1 or 2, wherein R_2 is phenyl substituted by one or more substituent chosen from halogen, nitro, $\text{CONR}_{13}\text{R}_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{1-6} alkyloxy, $\text{C}(\text{O})\text{C}_{1-6}$ alkyl, C_{6-12} aryl, C_{3-10} heterocycle, hydroxyl, $\text{NR}_{13}\text{R}_{14}$, $\text{C}(\text{O})\text{OR}_{12}$, cyano, azido, wherein R_{12} , R_{13} and R_{14} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl;
- or R_{13} and R_{14} are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

61. The compound as defined in anyone of claims 1 or 2, wherein R_2 is phenyl substituted by one or two substituents chosen from halogen, nitro, $\text{CONR}_{13}\text{R}_{14}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{1-6} alkyloxy, $\text{C}(\text{O})\text{C}_{1-6}$ alkyl, C_{6-12} aryl, C_{3-10} heterocycle, hydroxyl, $\text{NR}_{13}\text{R}_{14}$, $\text{C}(\text{O})\text{OR}_{12}$, cyano, azido, wherein R_{12} , R_{13} and R_{14} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12}

alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl;

or R₁₃ and R₁₄ are taken together with the nitrogen to form a 3 to 10 membered heterocycle.

62. The compound as defined in anyone of claims 1 or 2, wherein R₂ is phenyl substituted by one or two substituents chosen from halogen, C₁₋₆ alkyl, NR₁₃R₁₄, nitro, CONR₁₃R₁₄, C(O)OC₁₋₆ alkyl, COOH or C₁₋₆ alkyloxy C(O)OR₁₂, cyano, azido, wherein R₁₂, R₁₃ and R₁₄ are each independently chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl;
- or R₁₃ and R₁₄ are taken together with the nitrogen to form a 3 to 10 membered heterocycle.
63. The compound as defined in anyone of claims 1 or 2, wherein R₂ is methylphenyl.
64. The compound as defined in anyone of claims 1 or 2, wherein R₂ is dichlorophenyl.
65. The compound as defined in anyone of claims 1 or 2, wherein R₂ is chlorophenyl.

66. A compound chosen from:

Compound 1	3-[(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYL)-(3-iodo-benzyl)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 2	3-[(3-BENZOFURAN-2-YL-BENZYL)-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 3	3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 4	3-[(2,4-DICHLORO-BENZOYL)-[5-(3-TRIFLUOROMETHYL-PHENYL)-FURAN-2-YLMETHYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 5	3-[(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 6	5-(4-FLUORO-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 7	3-(2,4-DICHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 8	3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 9	3-[(2,4-DICHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 10	5-TERT-BUTYL-3-(4-CHLORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 11	4-(TOLUENE-4-SULFONYLAMINO)-[2,3']BITHIOPHENYL-5-CARBOXYLIC ACID
Compound 12	3-[(5-BENZOFURAN-2-YL-THIOPHEN-2-YLMETHYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 13	5-PHENYL-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 14	3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-CHLORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 15	5-PHENYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 16	5-PHENYL-3-(TOLUENE-3-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 17	3-BENZENESULFONYLAMINO-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 18	3-(4-CHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 19	3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 20	5-TERT-BUTYL-3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 21	3-(2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 22	3-(4-METHOXY-2,3,6-TRIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 23	5-PHENYL-3-(THIOPHENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 24	4-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-[2,3']BITHIOPHENYL-5-CARBOXYLIC ACID

Compound 25	5-(3,5-BIS-TRIFLUOROMETHYL-PHENYL)-3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 26	8-CHLORO-3-(4-CHLORO-2,5-DIMETHYL-BENZENESULFONYLAMINO)-4H-1,5-DITHIA-CYCLOPENTA[A]NAPHTHALENE-2-CARBOXYLIC ACID
Compound 27	3-(2,4-DIFLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 28	3-[3-(2,6-DICHLORO-PYRIDIN-4-YL)-UREIDO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 29	3-(2-CHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 30	3-(2-FLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 31	5-PHENYL-3-(2-TRIFLUOROMETHOXY-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 32	3-(4- TERT -BUTYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 33	3-(4-CHLORO-PHENOXYCARBONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 34	3-(3,4-DICHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 35	5-PHENYL-3-(2-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 36	3-(5-BROMO-6-CHLORO-PYRIDINE-3-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 37	3-(5-CHLORO-THIOPHENE-2-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 38	3-(5-CHLORO-3-METHYL-BENZO[B]THIOPHENE-2-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 39	3-(4-BROMO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 40	3-(3-CHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 41	3-(5-CHLORO-1,3-DIMETHYL-1H-PYRAZOLE-4-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 42	3-(3-BROMO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 43	3-(4-ISOPROPYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 44	3-(2,6-DICHLORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 45	3-(2-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 46	5-PHENYL-3-(5-[1,2,3]THIADIAZOL-4-YL-THIOPHENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 47	5-PHENYL-3-(PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 48	3-(2,4-DICHLORO-BENZYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 49	3-(3-FLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 50	5-PHENYL-3-(3-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 51	3-(2-CARBOXY-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID METHYL ESTER
Compound 52	5-PHENYL-3-(4-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 53	3-(2,5-DIFLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 54	3-(2-CYANO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 55	3-(2,5-DICHLORO-THIOPHENE-3-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 56	4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
Compound 57	5'-CHLORO-4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
Compound 58	5-(2,4-DICHLORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 59	5-(4-NITRO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 60	3-(TOLUENE-2-SULFONYLAMINO)-5-(4-TRIFLUOROMETHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 61	5-QUINOLIN-8-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 62	5-PHENYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 63	5-(3-NITRO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID

Compound 64	3-(TOLUENE-2-SULFONYLAMINO)-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 65	5-(3-CHLORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 66	5-(4-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 67	5-(3-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 68	5-(4-CHLORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 69	5-(3,5-DIFLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 70	5-(3,4-DIFLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 71	3-(TOLUENE-2-SULFONYLAMINO)-5-VINYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 72	3-(4-CHLORO-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 73	3-[(4-CHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 74	5-PHENYL-3-[(THIOPHENE-2-CARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 75	3-[METHYL-(THIOPHENE-2-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 76	3-(2-BROMO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 77	3-(2,4-DIFLUORO-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 78	3-[(2,4-DIFLUORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 79	3-(TOLUENE-2-SULFONYLAMINO)-5-TRIMETHYLSILANYLETHYNYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 80	5-ETHYNYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 81	3-(TOLUENE-2-SULFONYLAMINO)-5-(3-TRIFLUOROMETHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 82	5-BENZOYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID

Compound 83	5-(4-CYANO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 84	5-(3-CHLORO-4-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 85	5-(3,4-DICHLORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 86	5-PYRIDIN-4-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 87	5-PYRIDIN-3-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 88	3-(TOLUENE-2-SULFONYLAMINO)-5-(4-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 89	5-(4-METHANESULFONYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 90	5-(3-ACETYLAMINO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 91	5-(3-CHLORO-4-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 92	3-(4-METHYL-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 93	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 94	3-(3,5-DIMETHYL-ISOXAZOLE-4-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 95	3-[(2-CHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 96	3-(2-METHYL-BENZOYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 97	3-[METHYL-(2-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 98	5-PHENYL-3-(5-TRIFLUOROMETHYL-PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 99	5-PHENYLETHINYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 100	3-(2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 101	5-(2-FLUORO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 102	5-(2-CYANO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID

Compound 103	5-(2-ETHOXYCARBONYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 104	5-(2-METHOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 105	3'-METHYL-4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
Compound 106	3-(TOLUENE-2-SULFONYLAMINO)-5-(2-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 107	3-(2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 108	5-STYRYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 109	3-(2,4-DIFLUORO-BENZENESULFONYLAMINO)-5-(4-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 110	3-(2,4-DIFLUORO-BENZENESULFONYLAMINO)-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 111	3-[[5-(3-CHLORO-4-FLUORO-PHENYL)-THIOPHEN-2-YLMETHYL]-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 112	3-[(4-OXO-1-PHENYL-1,3,8-TRIAZA-SPIRO[4.5]DECANE-8-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 113	3-[[4-(2-OXO-2,3-DIHYDRO-BENZOIMIDAZOL-1-YL)-PIPERIDINE-1-CARBONYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 114	3-[[4-(4-NITRO-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 115	5-(2-CARBOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 116	5-(4-CHLORO-PHENYL)-3-(PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 117	5-(3-CYANO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 118	3-(2,5-DIMETHYL-BENZENESULFONYLAMINO)-5-P-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 119	3-(2,4-DIFLUORO-BENZENESULFONYLAMINO)-5-P-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 120	5-PHENETHYL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 121	5-(3-ETHOXYCARBONYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID

Compound 122	5-(4-METHOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 123	5-(3-METHOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 124	5-(4'-BROMO-BIPHENYL-4-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 125	5-(4-HYDROXYMETHYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 126	5-FURAN-3-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 127	5-BENZOFURAN-2-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 128	5-PYRIDIN-2-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 129	5-(4-NITRO-PHENYL)-3-(PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 130	3-[(BENZOFURAN-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 131	3-[(2,4-DIMETHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 132	3-[[5-(2-CYANO-PHENYL)-THIOPHEN-2-YLMETHYL]-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 133	5-(4-FLUORO-PHENYL)-3-(PYRIDINE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 134	5-[2-(4-CHLORO-PHENYL)-VINYL]-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 135	3-BENZENESULFONYLAMINO-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 136	3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 137	5-PHENYL-3-(2-VINYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 138	3-(4-BROMO-2,5-DIFLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 139	3-(2-ACETYLAMINO-4-METHYL-THIAZOLE-5-SULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 140	3-(4-ACETYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 141	3-(4-FLUORO-2-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 142	3-(2-METHOXY-4-METHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 143	3-(3,4-DIFLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 144	4-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-5-(4-CHLORO-PHENYL)-2-METHYL-FURAN-3-CARBOXYLIC ACID ETHYL ESTER
Compound 145	3-(4-FLUORO-3-TRIFLUOROMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 146	3-(2-AMINO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 147	3-(3-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 148	3-(4-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 149	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 150	5-(3-CYANO-BENZYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 151	5-PHENYL-3-(2,4,6-TRIFLUORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 152	3-(4-METHOXY-2-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 153	5-PHENYL-3-(2,3,4-TRICHLORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 154	5-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-2-METHYL-FURAN-3-CARBOXYLIC ACID METHYL ESTER
Compound 155	4-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-2-METHYL-1,5-DIPHENYL-1H-PYRROLE-3-CARBOXYLIC ACID ETHYL ESTER
Compound 156	5-PHENYL-3-([4-(3-TRIFLUOROMETHYL-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 157	3-([4-(4-FLUORO-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 158	3-([4-(2,6-DIMETHYL-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 159	3-([4-(2-CHLORO-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 160	3-([4-(3-CHLORO-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 161	4,4'-BIS-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5,5'-DICARBOXYLIC ACID
Compound 162	3-[ALLYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 163	5-(1-DIMETHYLSULFAMOYL-1H-PYRAZOL-4-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 164	5-(3-AMINO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 165	5-(4-AMINO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 166	5-(4-ACETYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 167	4-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-2,5-DIMETHYL-1H-PYRROLE-3-CARBOXYLIC ACID ETHYL ESTER
Compound 168	4-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YLSULFAMOYL)-5-(4-CHLORO-PHENYL)-3-METHYL-1-PHENYL-1H-PYRROLE-2-CARBOXYLIC ACID ETHYL ESTER
Compound 169	3-(3,5-DICHLORO-4-HYDROXY-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 170	5-(1H-PYRAZOL-4-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 171	5-(3-HYDROXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 172	3-[METHYL-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 173	3-([2-(4-FLUORO-PHENYL)-ACETYL]-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 174	3-(4-PENTYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 175	3-(METHYL-PHENYLACETYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 176	3-[2,5-BIS-(2,2,2-TRIFLUORO-ETHOXY)-BENZENESULFONYLAMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 177	3-(4-METHYL-2-NITRO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 178	5-THIAZOL-2-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 179	5-PHENYL-3-[3-(3-PHENYL-PROPYL)-UREIDO]-THIOPHENE-2-CARBOXYLIC ACID

Compound 180	3-[(3,4-DIHYDRO-1H-ISOQUINOLINE-2-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 181	3-[[4-(4-METHOXY-PHENYL)-PIPERAZINE-1-CARBONYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 182	3-[[4-(6-METHYL-PYRIDIN-2-YL)-PIPERAZINE-1-CARBONYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID HYDROCHLORIDE
Compound 183	3-[[4-(4-CHLORO-BENZYL)-PIPERAZINE-1-CARBONYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID HYDROCHLORIDE
Compound 184	5-(5-METHYL-PYRIDIN-2-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 185	3-[ETHYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 186	3-[(3-CHLORO-THIOPHENE-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 187	3-[(2-BROMO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 188	3-[(4-BUTYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 189	3-(2-CHLOROMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 190	5-(4-HYDROXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 191	5-(5-CHLORO-PYRIDIN-2-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 192	5-(4-CHLORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 193	5-(4-CYANO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 194	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(4-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 195	5-(4-HYDROXYMETHYL-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 196	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(3-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 197	5-(4-FLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 198	5-(4-METHOXY-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 199	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-P-TOLYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 200	5-(4-AMINO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 201	3-[CYCLOPENTYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 202	5-BENZO[1,3]DIOXOL-5-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 203	3-[(2-HYDROXY-ETHYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 204	3-[(2,4-DICHLORO-BENZOYL)-ISOBUTYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 205	3-[(2-METHOXY-4-METHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 206	5-(3-CYANO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 207	5-(2-CHLORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 208	3-[(2,4-DICHLORO-BENZOYL)-PHENYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 209	3-[4-(TRIFLUOROMETHYL-BENZOYL)METHYLAMINE]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 210	3-[(4-CHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 211	3-[ISOPROPYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 212	5-(3,5-DIFLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 213	5-(3-FLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 214	5-(2,4-DIFLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 215	5-(4-HYDROXY-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 216	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(4-TRIFLUOROMETHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 217	5-(2-HYDROXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 218	3-[(2-CHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 219	3-[(3,5-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 220	3-(4-BROMO-2-METHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 221	3-(5-CARBOXY-4-CHLORO-2-FLUORO-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 222	5-PHENYL-3-(2,3,4-TRIFLUORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 223	3-(4-BROMO-2-FLUORO-BENZENESULFONYLAMINO)-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 224	3-(4-BROMO-2-METHYL-BENZENESULFONYLAMINO)-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 225	5-(4-ISOBUTYL-PHENYL)-3-(3-METHOXY-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 226	3-[(4-FLUORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 227	3-[2,5-BIS-(2,2,2-TRIFLUORO-ETHOXY)-BENZENESULFONYLAMINO]-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 228	3-(2-CHLORO-4-CYANO-BENZENESULFONYLAMINO)-5-(4-ISOBUTYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 229	5'-ACETYL-4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
Compound 230	5-BENZO[B]THIOPHEN-2-YL-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 231	5-(4-BUTYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 232	5-(4-ETHYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 233	3-[BENZYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 234	3-[(4-CHLORO-2-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 235	3-[(2,4-DIMETHYL-BENZENESULFONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 236	5-(4-ACETYL-PHENYL)-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 237	5-(4-ACETYL-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 238	5-(4-ACETYL-PHENYL)-3-(4-CHLORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID

Compound 239	5-(4-CARBOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID TERT-BUTYL ESTER
Compound 240	3-[(2,4-DIMETHYL-BENZENESULFONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 241	3-[ACETYL-(4-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 242	3-ETHANESULFONYLAMINO-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 243	3-[ISOPROPYL-(4-TRIFLUOROMETHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 244	3-[(2,4-DICHLORO-BENZOYL)-(3-METHYL-BUT-2-ENYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 245	3-[(2,6-DICHLORO-PYRIDINE-3-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 246	3-[(6-CHLORO-PYRIDINE-3-CARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 247	3-[(4-TERT-BUTYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 248	5-(4-CARBOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 249	5-(4-ETHOXY-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 250	3-[(2,6-DICHLORO-PYRIDINE-3-CARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 251	3-[(BENZO[B]THIOPHENE-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 252	3-[METHYL-(NAPHTHALENE-2-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 253	3-[(3,4-DICHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 254	3-[(3,5-DICHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 255	3-[(4-BROMO-3-METHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 256	3-[(3-CHLORO-BENZO[B]THIOPHENE-2-CARBONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 257	3-[METHYL-(4-METHYL-3-NITRO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 258	5-(4-CARBAMOYL-PHENYL)-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID

Compound 259	5-(4-CARBAMOYL-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 260	5-(1H-INDOL-5-YL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 261	3-[SEC-BUTYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 262	3-[(2,4-DIMETHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 263	5-(4-AZIDO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 264	3-[(2,4-DICHLORO-BENZOYL)-(1-PHENYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 265	5-(4-CARBAMOYL-PHENYL)-3-(4-CHLORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 266	5-(2-FLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 267	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-O-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 268	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 269	5-(3-CHLORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 270	5-(3,4-DIFLUORO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 271	5-(3-AMINO-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 272	5-(3-ACETYL-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 273	5-(3-HYDROXY-PHENYL)-3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 274	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 275	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-(4-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 276	3-[(3,4-DIMETHOXY-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 277	3-[METHYL-(2,4,6-TRIFLUORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 278	3-[(2,3-DIFLUORO-4-TRIFLUOROMETHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 279	3-[(3-FLUORO-4-TRIFLUOROMETHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 280	3-[(2,3-DIFLUORO-4-METHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 281	3-[(2-FLUORO-4-TRIFLUOROMETHYL-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 282	5-(4-CARBAMOYL-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 283	5-(4-FLUORO-PHENYL)-3-[ISOPROPYL-(4-METHYL-BENZOYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 284	3-[(2-BROMO-4-CHLORO-BENZOYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 285	3-(2,6-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 286	3-[METHYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 287	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID METHYL ESTER
Compound 288	5-(4-CYANO-PHENYL)-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 289	3-(4-CHLORO-BENZENESULFONYLAMINO)-5-(4-CYANO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 290	5-(4-CYANO-PHENYL)-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 291	5'-ACETYL-4-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
Compound 292	5'-ACETYL-4-(2,6-DIMETHYL-BENZENESULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
Compound 293	3-[METHYL-(4-METHYL-THIOPHENE-2-CARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 294	5-(3-CHLORO-PHENYL)-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 295	5'-CYANO-4-(TOLUENE-2-SULFONYLAMINO)-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
Compound 296	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-PYRIDIN-2-YL-THIOPHENE-2-CARBOXYLIC ACID
Compound 297	3-[(2,4-DICHLORO-THIOBENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 298	5-PHENYL-3-(2,4,6-TRIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 299	3-[(1-CARBOXY-ETHYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 300	3-[(4-METHYL-BENZOYL)-(3-METHYL-BUT-2-ENYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 301	3-[(2-HYDROXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 302	3-[METHYL-(4-METHYL-BENZOYL)-AMINO]-5-PYRIDIN-3-YL-THIOPHENE-2-CARBOXYLIC ACID
Compound 303	5'-ACETYL-4-[METHYL-(4-METHYL-BENZOYL)-AMINO]-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
Compound 304	3-[ISOPROPYL-(4-METHYL-BENZOYL)-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 305	3-[ISOPROPYL-(4-METHYL-BENZOYL)-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 306	3-[(2-BROMO-4-CHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 307	3-[(4-CHLORO-2-FLUORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 308	3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-4-METHYL-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 309	3-[(2-BROMO-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 310	3-[(4-CHLORO-2-iodo-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 311	3-[(4-CYANO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 312	3-[ALLYL-(4-METHYL-BENZOYL)-AMINO]-5-[4-(2-CARBOXY-VINYL)-PHENYL]-THIOPHENE-2-CARBOXYLIC ACID
Compound 313	3-[(4-CHLORO-2-HYDROXY-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 314	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-4-METHYL-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 315	5-TERT-BUTYL-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 316	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 317	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 318	5-[4-(2-CARBOXY-ETHYL)-PHENYL]-3-[(4-METHYL-BENZOYL)-PROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 319	5-BENZOFURAN-2-YL-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 320	3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-HYDROXYMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 321	3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-METHANESULFONYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 322	5-[4-(2-CARBOXY-VINYL)-PHENYL]-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 323	3-[ALLYL-(4-METHYL-BENZOYL)-AMINO]-5-[3-(2-CARBOXY-VINYL)-PHENYL]-THIOPHENE-2-CARBOXYLIC ACID
Compound 324	3-[ISOPROPYL-(2,4,6-TRIMETHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 325	5-[3-(2-CARBOXY-ETHYL)-PHENYL]-3-[(4-METHYL-BENZOYL)-PROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 326	3-[(2-FLUORO-4-TRIFLUOROMETHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 327	3-[TERT-BUTYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 328	3-[(2-AMINO-4-CHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 329	3-[(4-CHLORO-2-NITRO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 330	3-[(4-METHYL-BENZOYL)-(3-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 331	3-[(3-FLUORO-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 332	5-(4-CARBOXY-PHENYL)-3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 333	3-[CYCLOPROPYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 334	3-[(3-TERT-BUTYL-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 335	3-[(3-CHLORO-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 336	3-[(2,4-DIFLUORO-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 337	3-[(4-CHLORO-2,5-DIFLUORO-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 338	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(2-METHYL-ALLYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 339	3-{ALLYL-[2-(4-CHLORO-PHENYL)-ACETYL]-AMINO}-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 340	3-[BENZYL-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 341	3-[(4-CHLORO-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 342	3-[(4-METHYL-BENZOYL)-(4-NITRO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 343	3-[(4-METHYL-BENZOYL)-(2-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 344	3-[(3-METHOXY-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 345	3-[(2-CHLORO-BENZYL)-(4-METHYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 346	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-ISOBUTYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 347	3-[ALLYL-(2-NAPHTHALEN-2-YL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 348	3-{ALLYL-[2-(2,4-DICHLORO-PHENYL)-ACETYL]-AMINO}-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 349	3-{ALLYL-[2-(2-CHLORO-4-FLUORO-PHENYL)-ACETYL]-AMINO}-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 350	3-{ALLYL-[2-(3,4-DICHLORO-PHENYL)-ACETYL]-AMINO}-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 351	3-{ALLYL-[2-(2,4-DIFLUORO-PHENYL)-ACETYL]-AMINO}-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 352	3-{ALLYL-[2-(4-TRIFLUOROMETHYL-PHENYL)-ACETYL]-AMINO}-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 353	3-{ALLYL-[2-(2,6-DICHLORO-PHENYL)-ACETYL]-AMINO}-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 354	3-[ALLYL-(2-M-TOLYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 355	5-(4-ACETYL-PHENYL)-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID

Compound 356	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(4-FLUORO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 357	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 358	5'-ACETYL-4-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
Compound 359	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 360	4-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5'-METHYL-[2,2']BITHIOPHENYL-5-CARBOXYLIC ACID
Compound 361	3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-(4-METHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 362	3-(CYCLOHEXANECARBONYL-ISOPROPYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 363	3-[(2,4-DICHLORO-BENZOYL)-(1-(2,4-DICHLORO-BENZOYL)-PIPERIDIN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 364	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(4-METHYL-BENZOYL)-AMINO]-PIPERIDINE-1-CARBOXYLIC ACID TERT -BUTYL ESTER
Compound 365	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-PIPERIDINE-1-CARBOXYLIC ACID TERT -BUTYL ESTER
Compound 366	3-[(4-METHYL-BENZOYL)-PIPERIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 367	5'-ACETYL-4-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-[2,3']BITHIOPHENYL-5-CARBOXYLIC ACID
Compound 368	3-[(2,4-DICHLORO-BENZOYL)-PIPERIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 369	5-(4-METHANESULFONYLAMINO-PHENYL)-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 370	3-(4-FLUORO-2-METHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 371	3-[(3-METHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 372	3-(4-CHLORO-2-METHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 373	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(4-METHANESULFONYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 374	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(4-METHANESULFINYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID

Compound 375	5-(4-CARBOXY-PHENYL)-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 376	5-BENZOFURAN-2-YL-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 377	3-[(2-ACETOXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 378	3-[ISOPROPYL-(2-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 379	3-[ISOPROPYL-(2-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 380	3-(CYCLOHEPTANECARBONYL-ISOPROPYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 381	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 382	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-METHYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 383	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-(3-NITRO-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 384	3-[(3-CYCLOPENTYL-PROPIONYL)-METHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 385	3-(BUTYRYL-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 386	3-(METHYL-PENT-4-ENOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 387	3-[ISOPROPYL-(5-METHYL-3-OXO-3H-ISOINDOL-1-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 388	3-[METHYL-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 389	3-(METHYL-PENTANOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 390	3-[METHYL-(4-METHYL-PENTANOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 391	3-(CYCLOPENTANECARBONYL-ETHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 392	3-[(3-CYCLOPENTYL-PROPIONYL)-ETHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 393	3-(CYCLOBUTANECARBONYL-ETHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 394	3-(BUT-2-ENOYL-ETHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 395	3-[ISOPROPYL-(4-METHYL-2-VINYL-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 396	3-[ISOPROPYL-(4-METHYL-CYCLOHEX-1-ENECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 397	3-(ALLYL-HEXANOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 398	3-(ALLYL-CYCLOBUTANECARBONYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 399	3-(ALLYL-PENTANOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 400	3-(ALLYL-(4-METHYL-PENTANOYL)-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 401	3-[ALLYL-(2-CYCLOPENTYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 402	3-[(2-HYDROXY-4-METHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 403	3-[(2,4-DICHLORO-BENZOYL)-(1-PHENYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 404	3-[(2,4-DICHLORO-BENZOYL)-(1-PHENYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 405	3-[ISOPROPYL-(3-METHYL-CYCLOPENT-3-ENECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 406	3-[(2-BENZYLOXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 407	3-[(2,4-DIMETHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 408	3-[ISOPROPYL-(3-METHYL-CYCLOPENTANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 409	3-[(2-HYDROXY-4-METHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 410	5-PHENYL-3-[PROPIONYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 411	3-[ISOBUTYRYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 412	3-[(3-METHYL-BUTYRYL)-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 413	3-[CYCLOPROPANECARBONYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 414	3-[CYCLOBUTANECARBONYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 415	3-[BUTYRYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 416	3-[(2-CYCLOPENTYL-ACETYL)-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 417	3-[(4-TERT-BUTYL-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 418	3-[(4-NITRO-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 419	3-[(3-METHYL-BUTYRYL)-(4-NITRO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 420	3-[CYCLOPROPANECARBONYL-(4-NITRO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 421	3-[(2-CHLORO-BENZYL)-ISOBUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 422	3-[(2-CHLORO-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 423	3-[(2-CHLORO-BENZYL)-CYCLOPROPANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 424	3-[(ADAMANTANE-1-CARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 425	3-[(2-CHLORO-BENZYL)-CYCLOBUTANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 426	3-[ACETYL-(2-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 427	3-[(2-METHYL-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 428	3-[(2-HYDROXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-(3-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 429	3-[(1-ACETYL-PIPERIDIN-4-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 430	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-[4-(1 H -TETRAZOL-5-YL)-PHENYL]-THIOPHENE-2-CARBOXYLIC ACID
Compound 431	3-[(2-CYANO-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 432	3-[CYCLOBUTANECARBONYL-(2-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 433	3-[BUTYRYL-(2-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 434	3-[ACETYL-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 435	3-[CYCLOBUTANECARBONYL-(4-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 436	3-[CYCLOHEXANECARBONYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 437	3-[(4-TERT-BUTYL-BENZYL)-ISOBUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 438	3-[(4-TERT-BUTYL-BENZYL)-CYCLOPROPANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 439	3-[(4-TERT-BUTYL-BENZYL)-CYCLOBUTANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 440	3-[(4-TERT-BUTYL-BENZYL)-BUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 441	3-[(4-TERT-BUTYL-BENZYL)-CYCLOHEXANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 442	3-[(4-TERT-BUTYL-BENZYL)-(2-CYCLOPENTYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 443	3-[(2-CYCLOPENTYL-ACETYL)-(4-NITRO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 444	3-[(2-CHLORO-BENZYL)-CYCLOHEXANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 445	3-[(2-CYCLOPENTYL-ACETYL)-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 446	3-[BUTYRYL-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 447	3-[BUTYRYL-(2-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 448	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 449	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-THIAZOL-2-YL-THIOPHENE-2-CARBOXYLIC ACID
Compound 450	3-(ACETYL-BENZYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 451	3-(BENZYL-PROPIONYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 452	3-[BENZYL-(2-METHOXY-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 453	3-[BENZYL-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 454	3-(BENZYL-CYCLOPROPANECARBONYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 455	3-[ACETYL-(4-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 456	3-[(4-CHLORO-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 457	3-[(4-CHLORO-BENZYL)-ISOBUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 458	3-[(4-CHLORO-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 459	3-[(4-CHLORO-BENZYL)-CYCLOPROPANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 460	5-(4-ACETYL-PHENYL)-3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 461	3-[(4-CHLORO-BENZYL)-CYCLOBUTANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 462	3-[BUTYRYL-(4-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 463	3-[(4-CHLORO-BENZYL)-(2-CYCLOPENTYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 464	3-[ACETYL-(4-TRIFLUOROMETHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 465	3-[ISOBUTYRYL-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 466	3-[CYCLOPROPANECARBONYL-(3-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 467	3-[(4-METHYL-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 468	3-[ISOBUTYRYL-(4-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 469	3-[CYCLOPROPANECARBONYL-(4-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 470	3-[BUTYRYL-(4-METHYL-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 471	3-[(3-METHOXY-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 472	3-[(3-METHOXY-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 473	3-[CYCLOBUTANECARBONYL-(3-METHOXY-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 474	3-[(2-CARBAMOYL-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 475	3-[BUTYRYL-(3-METHOXY-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 476	3-[ACETYL-(3-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 477	3-[(3-CHLORO-BENZYL)-PROPIONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 478	3-[(3-CHLORO-BENZYL)-(2-METHOXY-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 479	3-[(3-CHLORO-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 480	3-[(3-CHLORO-BENZYL)-CYCLOPROPANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 481	3-[(3-CHLORO-BENZYL)-CYCLOBUTANECARBONYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 482	3-[BUTYRYL-(3-CHLORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 483	3-[ACETYL-(2,4-DIFLUORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 484	3-[(2,4-DIFLUORO-BENZYL)-(2-METHOXY-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 485	3-[(2,4-DIFLUORO-BENZYL)-ISOBUTYRYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 486	3-[(2,4-DIFLUORO-BENZYL)-(3-METHYL-BUTYRYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 487	3-[BENZYL-(2-CYCLOPENTYL-ACETYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 488	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-(1H-INDOL-5-YL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 489	3-(BENZYL-CYCLOBUTANECARBONYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 490	3-[CYCLOHEXANECARBONYL-(2,4-DIFLUORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 491	3-(ALLYL-[2-(4-METHOXY-PHENYL)-ACETYL]-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 492	3-(ETHYL-HEXANOYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 493	3-(BUTYRYL-ETHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 494	3-[ETHYL-(4-METHYL-PENTANOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 495	3-[CYCLOBUTANECARBONYL-(2,4-DIFLUORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 496	3-[BUTYRYL-(2,4-DIFLUORO-BENZYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 497	3-(CYCLOPENTANECARBONYL-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 498	3-(CYCLOHEXANECARBONYL-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 499	3-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-PYRROLIDINE-1-CARBOXYLIC ACID TERT-BUTYL ESTER
Compound 500	3-[(1,4-DIMETHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 501	5-(4-ETHYL-PHENYL)-3-[(2-HYDROXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 502	3-[(2-HYDROXY-4-METHYL-BENZOYL)-ISOPROPYL-AMINO]-5-M-TOLYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 503	3-[(2,4-DICHLORO-BENZOYL)-PYRROLIDIN-3-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 504	4-{5-CARBOXY-4-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHEN-2-YL}-3,6-DIHYDRO-2H-PYRIDINE-1-CARBOXYLIC ACID BENZYL ESTER
Compound 505	3-[[2-(HYDROXYIMINO-METHYL)-4-METHYL-BENZOYL]-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 506	3-[(1-CARBAMIMIDOYL-PIPERIDIN-4-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 507	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-AZEPANE-1-CARBOXYLIC ACID TERT-BUTYL ESTER
Compound 508	4-[[2-(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-METHYL]-PIPERIDINE-1-CARBOXYLIC ACID BENZYL ESTER
Compound 509	3-[AZEPAN-4-YL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 510	3-[(4-METHYL-CYCLOHEXANECARBONYL)-PIPERIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID LITHIUM SALT

Compound 511	3-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-PIPERIDINE-1-CARBOXYLIC ACID TERT -BUTYL ESTER
Compound 512	3-[(4-BENZYLOXYCARBONYLAMINO-CYCLOHEXYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 513	3-[ISOPROPYL-(4-METHYL-2-OXO-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 514	3-[(2,4-DICHLORO-BENZOYL)-PIPERIDIN-3-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID; COMPOUND WITH GENERIC INORGANIC NEUTRAL COMPONENT
Compound 515	3-[(4-BENZYLOXYCARBONYLAMINO-CYCLOHEXYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 516	3-[(2-BENZYLOXY-1-METHYL-ETHYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 517	3-[(2,2-DIMETHYL-{1,3}DIOXAN-5-YL)-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 518	3-[(2,4-DICHLORO-BENZOYL)-(2-HYDROXY-1-HYDROXYMETHYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 519	3-[(2,4-DICHLORO-BENZOYL)-PIPERIDIN-4-YLMETHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 520	3-[(2-CHLORO-BENZOYL)-PIPERIDIN-4-YLMETHYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 521	3-[(4,6-DICHLORO-1H-INDOLE-2-CARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 522	3-[(2,4-DICHLORO-BENZOYL)-(2-HYDROXY-1-METHYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 523	4-{1-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-ETHYL}-PIPERIDINE-1-CARBOXYLIC ACID BENZYL ESTER
Compound 524	4-{5-CARBOXY-4-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHEN-2-YL}-3,6-DIHYDRO-2 H -PYRIDINE-1-CARBOXYLIC ACID BENZYL ESTER
Compound 525	3-[(4-METHYL-CYCLOHEXANECARBONYL)-PYRIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 526	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-5-PIPERIDIN-4-YL-THIOPHENE-2-CARBOXYLIC ACID; COMPOUND WITH TRIFLUORO-ACETIC ACID
Compound 527	3-[ISOPROPYL-(4-PROPYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 528	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-CYCLOHEXYL-AMMONIUM; TRIFLUORO-ACETATE

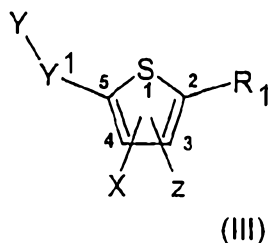
Compound 529	3-[(2,4-DICHLORO-BENZOYL)-(1-PIPERIDIN-4-YL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID; COMPOUND WITH TRIFLUORO-ACETIC ACID
Compound 530	3-[(CYCLOHEX-3-ENECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 531	3-[(4-ETHYL-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 532	3-[(4-CHLORO-CYCLOHEXANECARBONYL)-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 533	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-3-METHYL-PIPERIDINE-1-CARBOXYLIC ACID BENZYL ESTER
Compound 534	3-[(2,4-DICHLORO-BENZOYL)-(2-METHOXY-CYCLOHEXYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 535	3-[(2,4-DICHLORO-BENZOYL)-(2,2-DIMETHYL-[1,3]DIOXAN-5-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 536	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-(1-METHYL-PIPERIDIN-4-YL)-THIOPHENE-2-CARBOXYLIC ACID
Compound 537	3-[(2,4-DICHLORO-BENZOYL)-(3-METHYL-PIPERIDIN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID; COMPOUND WITH TRIFLUORO-ACETIC ACID
Compound 538	3-[(2,4-DICHLORO-BENZOYL)-(2-HYDROXY-CYCLOHEXYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 539	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(4-METHYLCYCLOHEXANE CARBONYL)-AMINO]-METHYL-PIPERIDINE-1-CARBOXYLIC ACID BENZYL ESTER
Compound 540	3-[(1R,2S,4R)-2-HYDROXY-4-METHYL-CYCLOHEXANECARBONYL]-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 541	3-[ISOPROPYL-[1-(4-METHOXY-2,3,6-TRIMETHYL-BENZENESULFONYL)-5-METHYL-1,2,3,6-TETRAHYDRO-PYRIDINE-2-CARBONYL]-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 542	3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-4-FLUORO-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 543	3-[(2,4-DICHLORO-BENZOYL)-(1-METHYL-PIPERIDIN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 544	4-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(4-METHYLCYCLOHEXANE CARBONYL)-AMINO]-METHYL-PIPERIDINIUM; TRIFLUORO-ACETATE
Compound 545	3-[(2-TERT-BUTOXYCARBONYLAMINO-1-METHYL-ETHYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

Compound 546	2-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-PROPYL-AMINE TRIFLUOROACETIC ACID SALT
Compound 547	3-[(3-CARBOXY-CYCLOPENTYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 548	3-[(3-CARBOXY-CYCLOPENTYL)-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 549	2-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-CYCLOHEXYL-AMMONIUM CHLORIDE
Compound 550	3-(BENZOYL-METHYL-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 551	((5-PHENYL-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBONYL)-AMINO)-ACETIC ACID
Compound 552	5-BROMO-3-(TOLUENE-2-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound 553	3-[CYCLOHEXYL-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 554	3-[[1,3]DIOXAN-5-YL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 555	3-[[2-(TERT-BUTYL-DIMETHYL-SILANYLOXY)-1-METHYL-2-PHENYL-ETHYL]-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 556	3-[[2-(TERT-BUTYL-DIMETHYL-SILANYLOXY)-1-METHYL-2-PHENYL-ETHYL]-(2,4-DICHLORO-BENZOYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 557	3-[(2,4-DICHLORO-BENZOYL)-(2-DIETHYLAMINO-THIAZOL-5-YLMETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 558	(5-[(2-CARBOXY-5-PHENYL-THIOPHEN-3-YL)-(2,4-DICHLORO-BENZOYL)-AMINO]-METHYL)-THIAZOL-2-YL)-DIETHYL-AMMONIUM; CHLORIDE
Compound 559	5-(4-FLUORO-PHENYL)-3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound 560	3-[(1S,2R,4S)-2-HYDROXY-4-METHYL-CYCLOHEXANECARBONYL]-ISOPROPYL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 561	3-[(2,4-DICHLORO-BENZOYL)-(2-METHOXY-1-METHYL-ETHYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 562	3-[(4S)-ISOPROPYL-(4-METHYL-CYCLOHEX-1-ENECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound 566	3-METHYL-(4-METHYLBENZOYL)-AMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID (2-HYDROXY-ETHYL)AMIDE
Compound 567	5-PHENYL-3-(TOLUENE-4-SULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID CYCLOBUTYLAMIDE

Compound	568	3-(2,4-DIMETHYL-BENZENESULFONYLAMINO)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID AMIDE
Compound	569	5-BROMO-3-[(2,4-DICHLORO-BENZOYL)-ISOPROPYL-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound	570	5-(4-CHLORO-PHENYL)-3-[ISOPROPYL-(4-METHYL-CYCLOHEXANE-CARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound	571	5-(4'-CHLORO-BIPHENYL-4-YL)-3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound	572	3-[(4-METHYL-CYCLOHEXANECARBONYL)-(TETRAHYDRO-PYRAN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound	573	3-[(4-METHYL-CYCLOHEXANECARBONYL)-(1-METHYL-PIPERIDIN-4-YL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound	574	3-[(4-METHYL-CYCLOHEXANECARBONYL)-PIPERIDIN-4-YL-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound	575	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-(4-TRIFLUOROMETHYL-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound	576	5-(4-CYANO-PHENYL)-3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound	577	3-[ISOPROPYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-(4-METHOXY-PHENYL)-THIOPHENE-2-CARBOXYLIC ACID
Compound	578	3-[(2-METHOXY-1-METHYL-ETHYL)-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound	579	3-[CYCLOHEXYL-(4-METHYL-CYCLOHEXANECARBONYL)-AMINO]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound	581	5-(4-ISOBUTYL-PHENYL)-3-[5-(5-TRIFLUOROMETHYL-ISOXAZOL-3-YL)-THIOPHENE-2-SULFONYLAMINO]-THIOPHENE-2-CARBOXYLIC ACID
Compound	582	5-(4-ISOBUTYL-PHENYL)-3-(2,3,4-TRIFLUORO-BENZENESULFONYLAMINO)-THIOPHENE-2-CARBOXYLIC ACID
Compound	583	3-[(2,4-DICHLORO-PHENYL)-ISOPROPYL-CARBAMOYL]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound	584	3-(METHYL-P-TOLYL-CARBAMOYL)-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID
Compound	585	3-[(2,4-DICHLORO-PHENYL)-METHYL-CARBAMOYL]-5-PHENYL-THIOPHENE-2-CARBOXYLIC ACID

or pharmaceutically acceptable salts thereof.

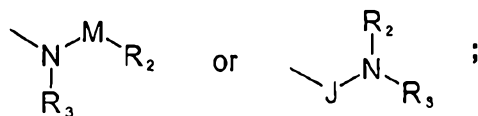
67. A method for treating or preventing a Flaviviridae viral infection in a host comprising administering to the host a therapeutically effective amount of at least one compound having the formula III:



or pharmaceutically acceptable salts thereof;

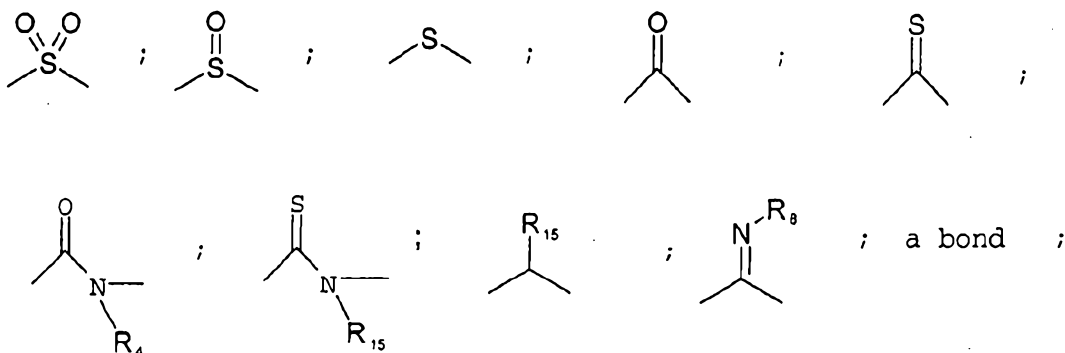
wherein,

X is chosen from:



wherein,

M is chosen from:



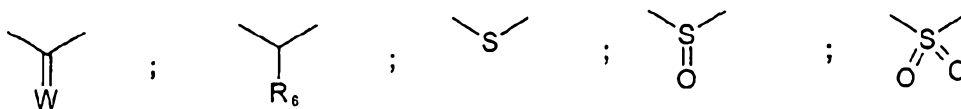
wherein,

R_4 is chosen from H or C_{1-6} alkyl;

R_8 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl; and

R_{15} is chosen from H or C_{1-6} alkyl;

J is chosen from:



wherein,

W is chosen from O, S or NR₇,

wherein R₇ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₂ aryl, C₃₋₁₂ heterocycle, C₃₋₁₂ heteroaralkyl, C₆₋₁₆ aralkyl;

and R₆ is chosen from H, C₁₋₁₂ alkyl, C₆₋₁₂ aryl or C₆₋₁₆ aralkyl;

Y¹ is chosen from a bond, C₁₋₆ alkyl, C₂₋₆ alkenyl or C₂₋₆ alkynyl;

Y is chosen from COOR₁₆, COCOOR₅, P(O)OR₄OR₃, S(O)OR₅, S(O)₂OR₅, tetrazole, CON(R₉)CH(R₅)COOR₅, CONR₁₀R₁₁, CON(R₉)-SO₂-R₅, CONR₉OH or halogen, wherein R₉, R₅, R₁₀ and R₁₁ are each independently chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl; or R₁₀ and R₁₁ are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R₄ and R₅ are each independently chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl and C₆₋₁₈ aralkyl;

or R₄ and R₅ are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R₁₆ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl and C₆₋₁₈ aralkyl;

R₁ is chosen from C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl, or halogen;

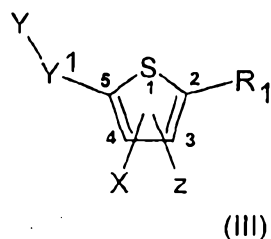
R_2 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, or C_{6-18} aralkyl;

R_3 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

Z is chosen from H, halogen, C_{1-6} alkyl.

68. The method of claim 65, further comprising at least one antiviral agent.
69. The method according to claim 66, wherein the antiviral agent is chosen from a viral serine protease inhibitor, viral polymerase inhibitor and viral helicase inhibitor.
70. The method according to claim 66, wherein the antiviral agent is chosen from interferon α and ribavirin.
71. The method according to anyone of claims 66 to 68, wherein said compound and said antiviral agent are administered sequentially.
72. The method according to anyone of claims 66 to 68, wherein said compound and said antiviral agent are administered simultaneously.
73. The method of claim 65, further comprising at least one additional agent chosen from immunomodulating agent, antioxidant agent, antibacterial agent or antisense agent.
74. The method of claim 71, wherein said additional agent is chosen from silybum marianum, interleukine-12, amantadine, ribozyme, thymosin, N-acetyl cysteine or cyclosporin.

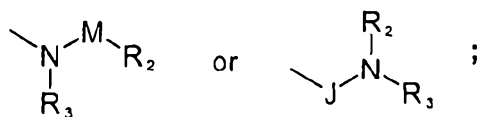
75. The method according to anyone of claims 71 or 72, wherein said compound and said additionnal agent are administered sequentially.
76. A method according to anyone of claims 71 or 72, wherein said compound and said additionnal agent are administered simultaneously.
77. The method as defined in anyone of claims 65 to 74, wherein said Flaviviridea viral infection is hepatitis C viral infection (HCV).
78. A pharmaceutical composition comprising at least one compound having the formula III:



or pharmaceutically acceptable salts thereof;

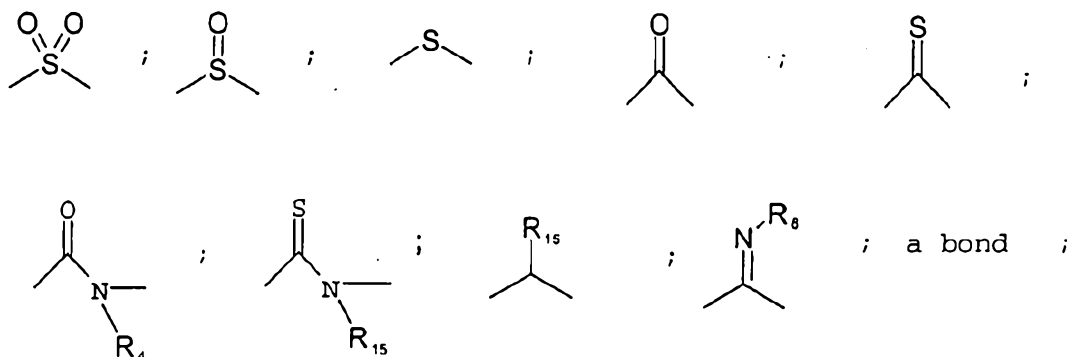
wherein,

X is chosen from:



wherein,

M is chosen from:



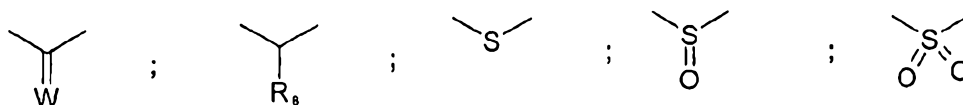
wherein,

R_4 is chosen from H or C_{1-6} alkyl;

R_8 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl; and

R_{15} is chosen from H or C_{1-6} alkyl;

J is chosen from:



wherein W is chosen from O, S or NR_8 ,

wherein R_8 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-12} aryl, C_{3-12} heterocycle, C_{3-12}

heteroaralkyl, C_{6-16} aralkyl;

and R_6 is chosen from H, C_{1-12} alkyl, C_{6-12} aryl or C_{6-16} aralkyl;

Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

Y is chosen from COOR_{16} , COCOOR_5 , $\text{P(O)OR}_4\text{OR}_5$, S(O)OR_5 , $\text{S(O)}_2\text{OR}_5$, tetrazole, $\text{CON(R}_3\text{)CH(R}_5\text{)COOR}_5$, $\text{CONR}_{10}\text{R}_{11}$, $\text{CON(R}_3\text{)-SO}_2\text{-R}_5$, CONR_3OH or halogen, wherein R_3 , R_5 , R_{10} and R_{11} are each independently

chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl; or R₁₀ and R₁₁ are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_a and R_b are each independently chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl and C₆₋₁₈ aralkyl;

or R_a and R_b are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R₁₆ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl and C₆₋₁₈ aralkyl;

R₁ is chosen from C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl or halogen;

R₂ is chosen from C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, or C₆₋₁₈ aralkyl;

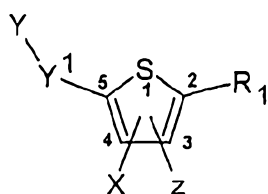
R₃ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl or C₆₋₁₈ aralkyl;

Z is chosen from H, halogen, C₁₋₆ alkyl; and

at least one pharmaceutically acceptable carrier or excipient.

79. A pharmaceutical composition as defined in claim 76, further comprising one or more additional agent is chosen from antiviral agent, immunomodulating agent, antioxydant agent, antibacterial agent or antisense agent.
80. The pharmaceutical composition as defined in claim 77, wherein the antiviral agent is chosen from a viral serine protease inhibitor, viral polymerase inhibitor and viral helicase inhibitor.

81. The pharmaceutical composition as defined in claim 77, wherein the antiviral agent is chosen from interferon α and ribavirin.
82. The pharmaceutical composition as defined in claim 77, wherein said additional agent is chosen from silybum marianum, interleukine-12, amantadine, ribozyme, thymosin, N-acetyl cysteine or cyclosporin.
83. The composition as defined in anyone of claims 76-80 wherein said Flaviviridae viral infection is hepatitis C viral infection (HCV).
84. The use of a compound having the formula III:

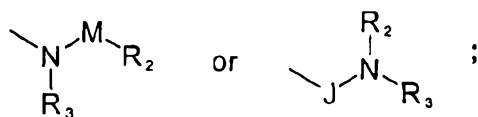


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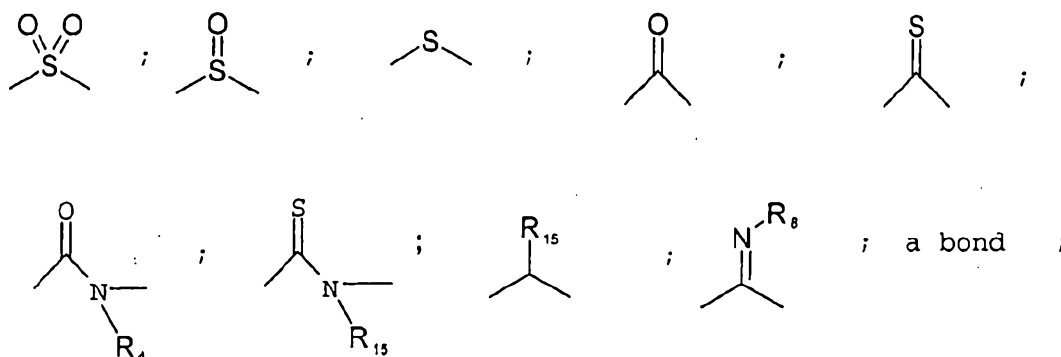
or pharmaceutically acceptable salts thereof;

wherein,

X is chosen from:

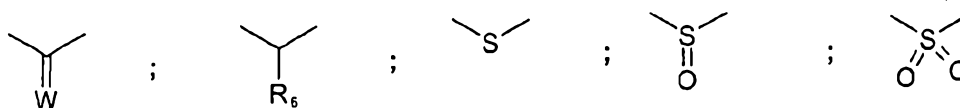


wherein,
M is chosen from:



wherein,
 R_4 is chosen from H or C_{1-6} alkyl;
 R_8 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl; and
 R_{15} is chosen from H or C_{1-6} alkyl;

J is chosen from:



wherein W is chosen from O, S or NR_7 ,
 wherein R_7 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-12} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl;
 and R_6 is chosen from H, C_{1-12} alkyl, C_{6-12} aryl or C_{6-16} aralkyl;

Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

Y is chosen from COOR_{16} , COCOOR_5 , $\text{P}(\text{O})\text{OR}_a\text{OR}_b$, $\text{S}(\text{O})\text{OR}_5$, $\text{S}(\text{O})_2\text{OR}_5$, tetrazole, $\text{CON}(\text{R}_9)\text{CH}(\text{R}_5)\text{COOR}_5$, $\text{CONR}_{10}\text{R}_{11}$, $\text{CON}(\text{R}_9)-\text{SO}_2-\text{R}_5$, CONR_9OH or halogen, wherein R_9 , R_5 , R_{10} and R_{11} are each independently

chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl; or R₁₀ and R₁₁ are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_a and R_b are each independently chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl and C₆₋₁₈ aralkyl;

or R_a and R_b are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R₁₆ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl and C₆₋₁₈ aralkyl;

R₁ is chosen from C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl or halogen;

R₂ is chosen from C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, or C₆₋₁₈ aralkyl;

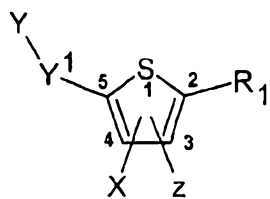
R₃ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl or C₆₋₁₈ aralkyl;

Z is chosen from H, halogen, C₁₋₆ alkyl;

for the manufacture of a medicament for treating or preventing a viral Flaviridea infection in a host.

85. The use as defined in claim 84, wherein said Flaviviridae viral infection is hepatitis C viral infection (HCV).

86. The use of a compound having the formula III:

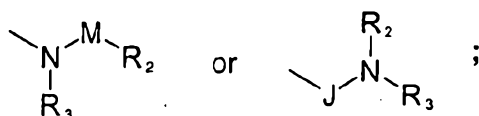


(III)

or pharmaceutically acceptable salts thereof in therapy;

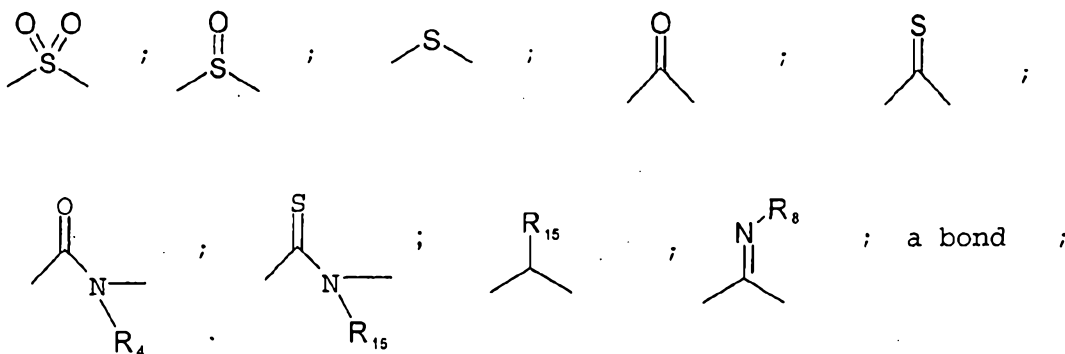
wherein,

X is chosen from:



wherein,

M is chosen from:



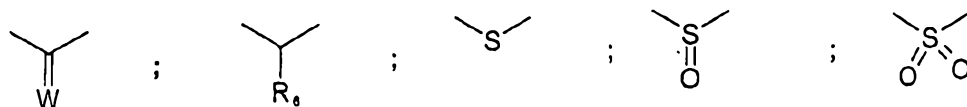
wherein,

R_1 is chosen from H or C_{1-6} alkyl;

R_2 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl; and

R_{15} is chosen from H or C_{1-6} alkyl;

J is chosen from:



wherein W is chosen from O, S or NR_7 ,

wherein R_7 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-12} aryl, C_{3-12} heterocycle, C_{3-12}

heteroaralkyl, C_{6-16} aralkyl;

and R_6 is chosen from H, C_{1-12} alkyl, C_{6-12} aryl or C_{6-16} aralkyl;

Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

Y is chosen from COOR_{16} , COCOOR_5 , $\text{P}(\text{O})\text{OR}_6\text{OR}_b$, $\text{S}(\text{O})\text{OR}_5$, $\text{S}(\text{O})_2\text{OR}_5$, tetrazole, $\text{CON}(\text{R}_9)\text{CH}(\text{R}_5)\text{COOR}_5$, $\text{CONR}_{10}\text{R}_{11}$, $\text{CON}(\text{R}_9)-\text{SO}_2-\text{R}_5$, CONR_9OH or halogen, wherein R_9 , R_5 , R_{10} and R_{11} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl; or R_{10} and R_{11} are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_a and R_b are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

or R_a and R_b are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R_{16} is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

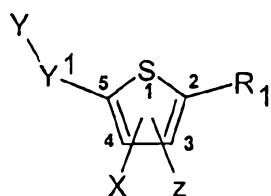
R_1 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl or halogen;

R_2 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, or C_{6-18} aralkyl;

R_3 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

Z is chosen from H, halogen, C_{1-6} alkyl.

87. The use of a compound having the formula III:

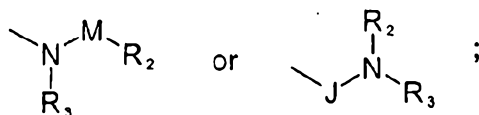


(III)

or pharmaceutically acceptable salts thereof;

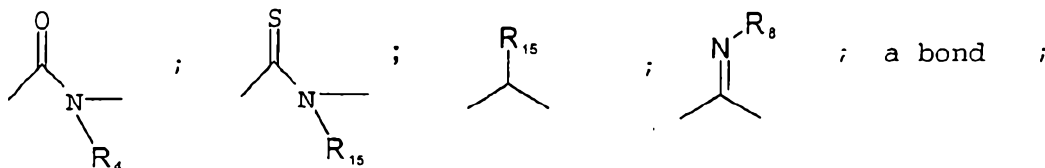
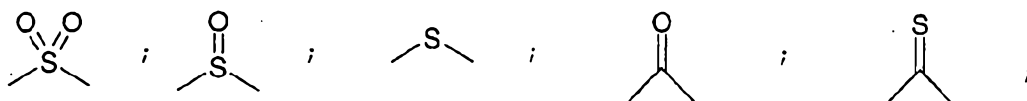
wherein,

X is chosen from:



wherein,

M is chosen from:



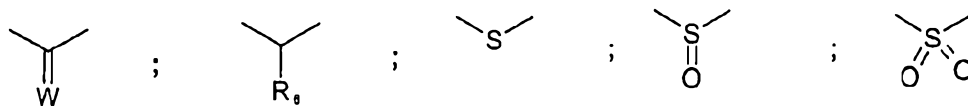
wherein,

R_4 is chosen from H or C_{1-6} alkyl;

R_8 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl; and

R_{15} is chosen from H or C_{1-6} alkyl;

J is chosen from:



wherein W is chosen from O, S or NR_7 ,

wherein R_7 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-12} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl;

and R_8 is chosen from H, C_{1-12} alkyl, C_{6-12} aryl or C_{6-16} aralkyl;

Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

Y is chosen from $COOR_{16}$, $COCOOR_5$, $P(O)OR_4OR_6$, $S(O)OR_5$, $S(O)_2OR_5$, tetrazole, $CON(R_9)CH(R_5)COOR_5$, $CONR_{10}R_{11}$, $CON(R_9)-SO_2-R_5$, $CONR_9OH$ or halogen, wherein R_9 , R_5 , R_{10} and R_{11} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl; or R_{10} and R_{11} are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_4 and R_6 are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

or R_4 and R_6 are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R_{16} is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

R_1 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl, or halogen;

R₂ is chosen from C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, or C₆₋₁₈ aralkyl;

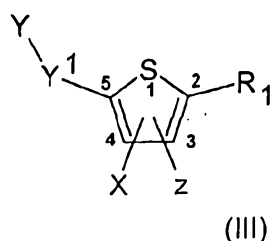
R₃ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl or C₆₋₁₈ aralkyl;

Z is chosen from H, halogen, C₁₋₆ alkyl;

for treating or preventing Flaviviridae viral infection in a host.

88. The use of a compound as defined in claim 85, further comprising one or more additional agent chosen from antiviral agent, immunomodulating agent, antioxydant agent, antibacterial agent or antisense agent.
89. The use as defined in claim 86, wherein said antiviral agent is chosen from a viral serine protease inhibitor, viral polymerase inhibitor and viral helicase inhibitor.
90. The use as defined in claim 86, wherein said antiviral agent is chosen from interferon α and ribavirin.
91. The use of as defined in claim 86 wherein said additional agent is chosen from silybum marianum, interleukine-12, amantadine, ribozyme, thymosin, N-acetyl cysteine or cyclosporin.
92. The use as defined in anyone of claims 86 to 89, wherein said compound and said additionnal agent are administered sequentially.

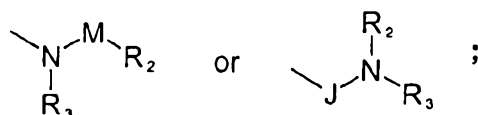
93. The use as defined in anyone of claims 86 to 89, wherein said compound and said additionnal agent are administered simultaneously.
94. The use as defined in anyone of claims 85 to 91, wherein said Flaviviridea viral infection is hepatitis C viral infection (HCV).
95. A method for inhibiting or reducing the activity of viral polymerase in a host comprising administering a therapeutically effective amount of a compound having the formula III:



or pharmaceutically acceptable salts thereof;

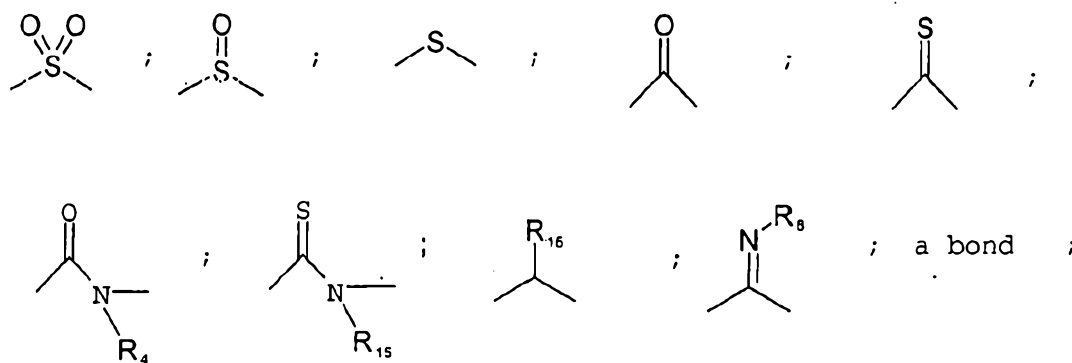
wherein,

X is chosen from:



wherein,

M is chosen from:



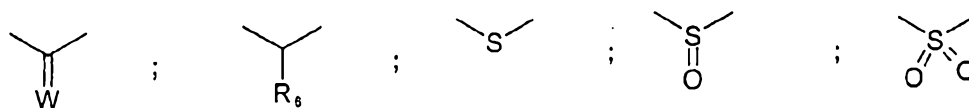
wherein,

R_4 is chosen from H or C_{1-6} alkyl;

R_8 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl; and

R_{15} is chosen from H or C_{1-6} alkyl;

J is chosen from:



wherein W is chosen from O, S or NR_7 ,

wherein R_7 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-12} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl;

and R_6 is chosen from H, C_{1-12} alkyl, C_{6-12} aryl or C_{6-16} aralkyl;

Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

Y is chosen from COOR_{16} , COCOOR_5 , $\text{P(O)OR}_6\text{OR}_7$, S(O)OR_5 , $\text{S(O)}_2\text{OR}_5$, tetrazole, $\text{CON(R}_3\text{)CH(R}_5\text{)COOR}_5$, $\text{CONR}_{10}\text{R}_{11}$, $\text{CON(R}_3\text{)-SO}_2\text{-R}_5$, CONR_3OH or halogen, wherein R_3 , R_5 , R_{10} and R_{11} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl;

or R_{10} and R_{11} are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_4 and R_5 are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

or R_4 and R_5 are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R_6 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

R_1 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl, or halogen;

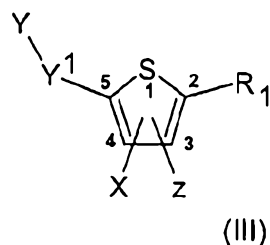
R_2 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, or C_{6-18} aralkyl;

R_3 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

Z is chosen from H, halogen, C_{1-6} alkyl.

96. The method as defined in claim 93, further comprising one or more viral polymerase inhibitor.
97. The method as defined in anyone of claims 93 or 94, wherein said viral polymerase is a Flaviviridae viral polymerase.
98. The method as defined in anyone of claims 93 or 94, wherein said viral polymerase is a RNA-dependant RNA-polymerase.
99. The method as defined in anyone of claims 93 or 94, wherein said viral polymerase is HCV polymerase.

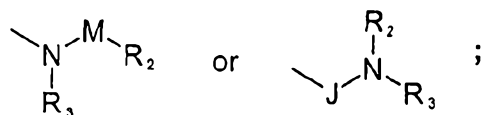
100. A method for inhibiting or reducing the activity of viral helicase in a host comprising administering a therapeutically effective amount of a compound having the formula III:



or pharmaceutically acceptable salts thereof;

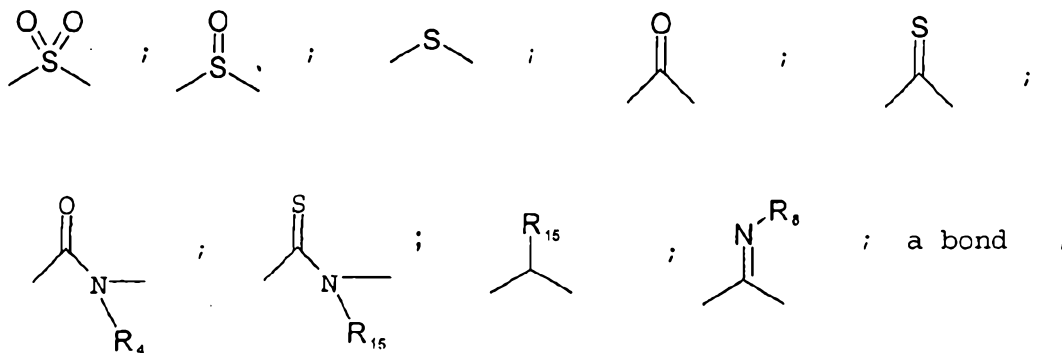
wherein,

X is chosen from:



wherein,

M is chosen from:



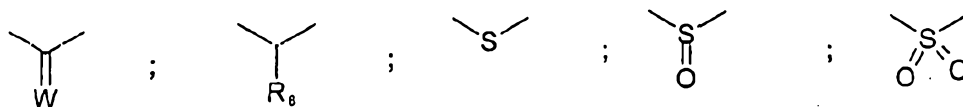
wherein,

R₄ is chosen from H or C₁₋₆ alkyl;

R₈ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₂ heteroaralkyl, C₆₋₁₆ aralkyl; and

R₁₅ is chosen from H or C₁₋₆ alkyl;

J is chosen from:



wherein W is chosen from O, S or NR_7 ,

wherein R_7 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-12} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl;

and R_6 is chosen from H, C_{1-12} alkyl, C_{6-12} aryl or C_{6-16} aralkyl;

Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

Y is chosen from COOR_{16} , COCOOR_5 , $\text{P}(\text{O})\text{OR}_4\text{OR}_5$, $\text{S}(\text{O})\text{OR}_5$, $\text{S}(\text{O})_2\text{OR}_5$, tetrazole, $\text{CON}(\text{R}_9)\text{CH}(\text{R}_5)\text{COOR}_5$, $\text{CONR}_{10}\text{R}_{11}$, $\text{CON}(\text{R}_9)-\text{SO}_2-\text{R}_5$, CONR_9OH or halogen, wherein R_9 , R_5 , R_{10} and R_{11} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl; or R_{10} and R_{11} are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_6 and R_5 are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

or R_6 and R_5 are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R_{16} is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

R_1 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl or halogen;

R_2 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, or C_{6-18} aralkyl;

R₃ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl or C₆₋₁₈ aralkyl;

Z is chosen from H, halogen, C₁₋₆ alkyl.

101. The method as defined in claim 98, wherein said compound is chosen from:

Compound #14 3-(4-Chloro-2,5-dimethyl-benzenesulfonylamino)-5-(4-chloro-phenyl)-thiophene-2-carboxylic acid

Compound #19 3-(4-Chloro-2,5-dimethyl-benzenesulfonylamino)-5-(4-isobutyl-phenyl)-thiophene-2-carboxylic acid

Compound #223 3-(4-Bromo-2-fluorobenzenesulfo-nylamino)-5-(4-isobutylphenyl)-thiophene-2-carboxylic acid

Compound #224 3-(4-Bromo-2-methylbenzenesulfo-nylamino)-5-(4-isobutylphenyl)-thiophene-2-carboxylic acid

Compound #225 5-(4-Isobutylphenyl 3-(3-methoxy-benzenesulfonylamino)-thiophene-2-carboxylic acid

Compound #581 5-(4-Isobutyl-phenyl)-3-[5-(5-trifluoromethyl-isoxazol-3-yl)-thiophene-2-sulfonylamino]-thiophene-2-carboxylic acid

Compound #227 3-[2,5-Bis-(2,2,2-trifluoroethoxy)-benzenesulfonylamino]-5-(4-isobutyl-phenyl)-thiophene-2-carboxylic acid

Compound #228 3-(2-Chloro-4-cyanobenzenesulfonylamino)-5-(4-isobutylphenyl)-thiophene-2-carboxylic acid

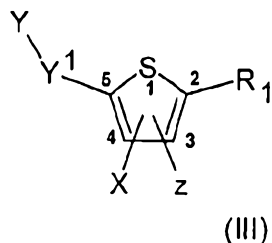
Compound #582 5-(4-Isobutyl-phenyl)-3-(2,3,4-trifluoro-benzenesulfonylamino)-thiophene-2-carboxylic acid

or pharmaceutically acceptable salts thereof.

102. The method as defined in anyone of claims 98 or 99, wherein said viral helicase is a flaviviridea helicase

103. The method as defined in anyone of claims 98 or 99, wherein said viral helicase is HCV helicase.

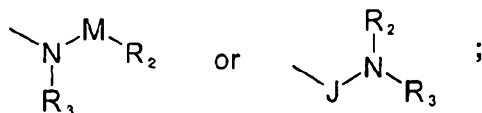
104. The use of a compound having the formula III:



or pharmaceutically acceptable salts thereof;

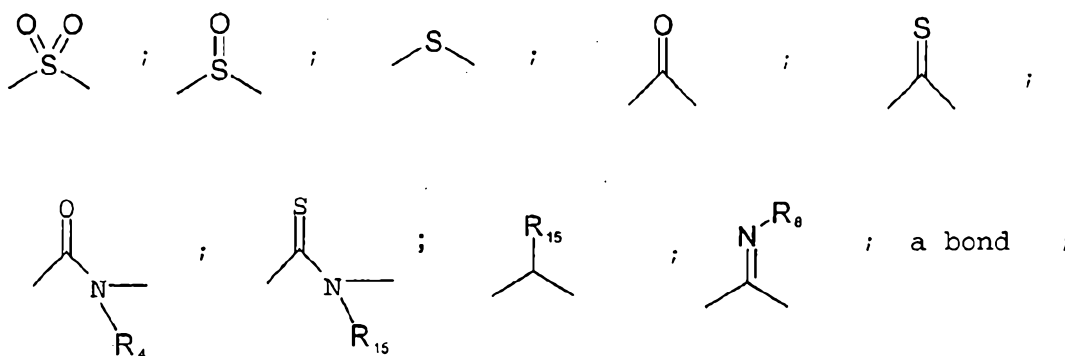
wherein,

X is chosen from:



wherein,

M is chosen from:



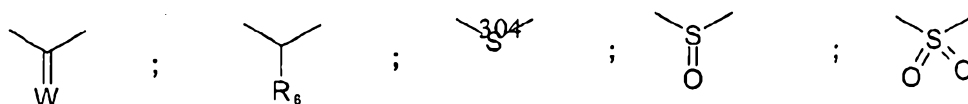
wherein,

R₄ is chosen from H or C₁₋₆ alkyl;

R₉ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₂ heteroaralkyl, C₅₋₁₆ aralkyl; and

R₁₅ is chosen from H or C₁₋₆ alkyl;

J is chosen from:



wherein W is chosen from O, S or NR₇,

wherein R₇ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₂ aryl, C₃₋₁₂ heterocycle, C₃₋₁₂ heteroaralkyl, C₆₋₁₆ aralkyl;

and R₈ is chosen from H, C₁₋₁₂ alkyl, C₆₋₁₂ aryl or C₆₋₁₆ aralkyl;

Y¹ is chosen from a bond, C₁₋₆ alkyl, C₂₋₆ alkenyl or C₂₋₆ alkynyl;

Y is chosen from COOR₁₆, COCOOR₅, P(O)OR₈OR₉, S(O)OR₅, S(O)₂OR₅, tetrazole, CON(R₉)CH(R₅)COOR₅, CONR₁₀R₁₁, CON(R₉)-SO₂-R₅, CONR₉OH or halogen, wherein R₉, R₅, R₁₀ and R₁₁ are each independently chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl;

or R₁₀ and R₁₁ are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R₈ and R₉ are each independently chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl and C₆₋₁₈ aralkyl;

or R₈ and R₉ are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R₁₆ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl and C₆₋₁₈ aralkyl;

R₁ is chosen from C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl or halogen;

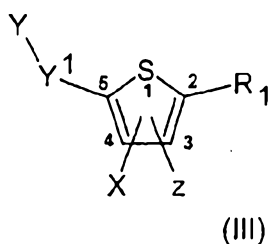
R₂ is chosen from C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, or C₆₋₁₈ aralkyl;

R₃ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl or C₆₋₁₈ aralkyl;

Z is chosen from H, halogen, C₁₋₆ alkyl;

for inhibiting or reducing the activity of viral polymerase in a host.

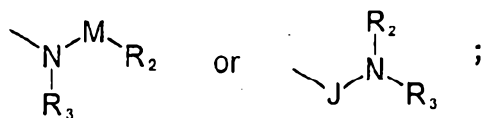
105. The use as defined in claim 102 further comprising one or more viral polymerase inhibitor.
106. The use as defined in anyone of claims 102 or 103, wherein said viral polymerase is Flaviviridae viral polymerase.
107. The use as defined in anyone of claims 102 or 103 wherein said viral polymerase is RNA-dependant RNA-polymerase.
108. The use as defined in anyone of claims 102 or 103, wherein said viral polymerase is HCV polymerase.
109. The use of a compound having the formula III:



or pharmaceutically acceptable salts thereof;

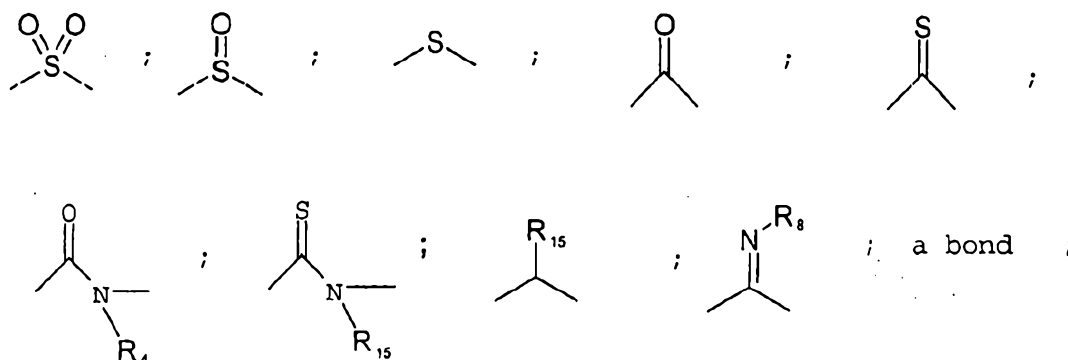
wherein,

X is chosen from:



wherein,

M is chosen from:



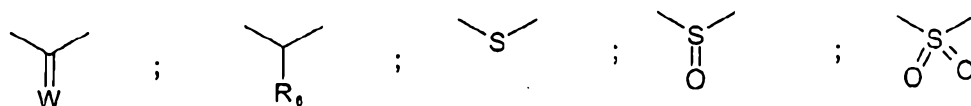
wherein,

R_4 is chosen from H or C_{1-6} alkyl;

R_8 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl; and

R_{15} is chosen from H or C_{1-6} alkyl;

J is chosen from:



wherein W is chosen from O, S or NR_1 ,

wherein R_1 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-12} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl;

and R_6 is chosen from H, C_{1-12} alkyl, C_{6-12} aryl or C_{6-16} aralkyl;

Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

Y is chosen from COOR_{16} , COCOOR_5 , $\text{P(O)OR}_5\text{OR}_5$, S(O)OR_5 , $\text{S(O)}_2\text{OR}_5$, tetrazole, $\text{CON(R}_3\text{)CH(R}_5\text{)COOR}_5$, $\text{CONR}_{10}\text{R}_{11}$, $\text{CON(R}_3\text{)-SO}_2\text{-R}_5$, CONR_3OH or halogen, wherein R_3 , R_5 , R_{10} and R_{11} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl;

or R_{10} and R_{11} are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_a and R_b are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

or R_a and R_b are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R_{16} is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

R_1 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl or halogen;

R_2 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, or C_{6-18} aralkyl;

R_3 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

Z is chosen from H, halogen, C_{1-6} alkyl;

for inhibiting or reducing the activity of viral helicase in a host.

110. The use as defined in claim 109, wherein said compound is chosen from:

Compound #14 3-(4-Chloro-2,5-dimethyl-benzenesulfonylamino)-5-(4-chloro-phenyl)-thiophene-2-carboxylic acid

Compound #19 3-(4-Chloro-2,5-dimethyl-benzenesulfonylamino)-5-(4-isobutyl-phenyl)-thiophene-2-carboxylic acid

Compound #223 3-(4-Bromo-2-fluorobenzenesulfo-nylamino)-5-(4-isobutylphenyl)-thiophene-2-carboxylic acid

Compound #224 3-(4-Bromo-2-methylbenzenesulfo-nylamino)-5-(4-isobutylphenyl)-thiophene-2-carboxylic acid

Compound #225 5-(4-Isobutylphenyl 3-(3-methoxy-benzenesulfonyl-

amino)-thiophene-2-carboxylic acid

Compound #581 5-(4-Isobutyl-phenyl)-3-[5-(5-trifluoromethyl-isoxazol-3-yl)-thiophene-2-sulfonylamino]-thiophene-2-carboxylic acid

Compound #227 3-[2,5-Bis-(2,2,2-trifluoroethoxy)-benzenesulfonylamino]-5-(4-isobutyl-phenyl)-thiophene-2-carboxylic acid

Compound #228 3-(2-Chloro-4-cyanobenzenesulfonylamino)-5-(4-isobutylphenyl)-thiophene-2-carboxylic acid

Compound #582 5-(4-Isobutyl-phenyl)-3-(2,3,4-trifluorobenzenesulfonylamino)-thiophene-2-carboxylic acid

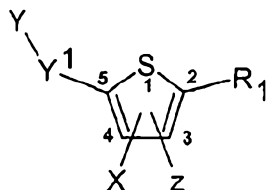
or pharmaceutically acceptable salts thereof.

111. The use as defined in anyone of claims 109 and 110 further comprising one or more viral helicase inhibitor.

112. The use as defined in anyone of claims 109 or 111, wherein said viral helicase is Flaviviridae viral helicase.

113. The use as defined in anyone of claims 109 or 111, wherein said viral helicase is HCV helicase.

114. A combination comprising a compound having the formula III:

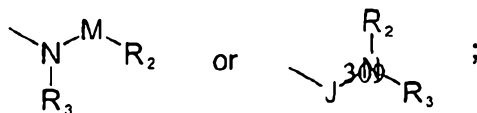


(III)

or pharmaceutically acceptable salts thereof;

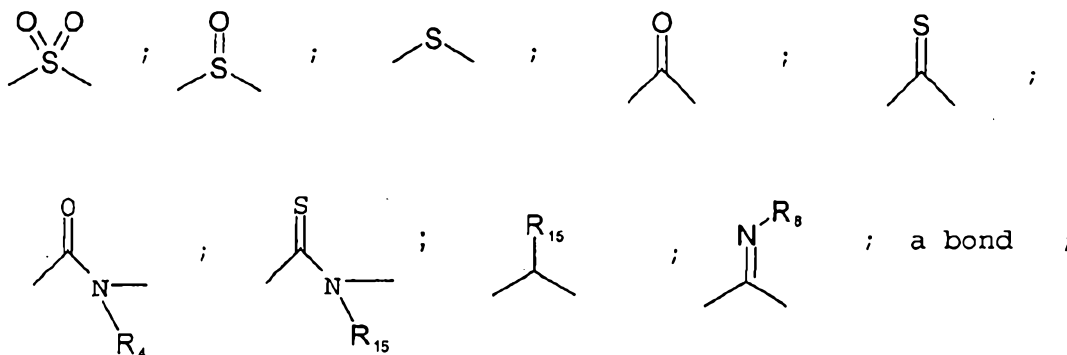
wherein,

X is chosen from:



wherein,

M is chosen from:



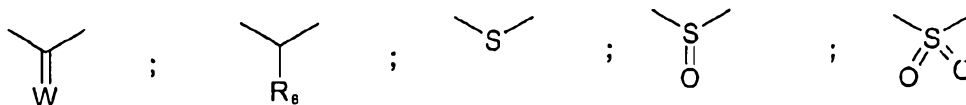
wherein,

R_4 is chosen from H or C_{1-6} alkyl;

R_8 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl; and

R_{15} is chosen from H or C_{1-6} alkyl;

J is chosen from:



wherein W is chosen from O, S or NR_7 ,

wherein R_7 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-12} aryl, C_{3-12} heterocycle, C_{3-12} heteroaralkyl, C_{6-16} aralkyl;

and R_6 is chosen from H, C_{1-12} alkyl, C_{6-12} aryl or C_{6-16} aralkyl;

Y^1 is chosen from a bond, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl;

Y is chosen from COOR_{16} , COCOOR_5 , $\text{P}(\text{O})\text{OR}_6\text{OR}_6$, $\text{S}(\text{O})\text{OR}_5$, $\text{S}(\text{O})_2\text{OR}_5$, tetrazole, $\text{CON}(\text{R}_9)\text{CH}(\text{R}_5)\text{COOR}_5$, $\text{CONR}_{10}\text{R}_{11}$, $\text{CON}(\text{R}_9)-\text{SO}_2-\text{R}_5$, CONR_9OH or halogen, wherein R_9 , R_5 , R_{10} and R_{11} are each independently

chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl; or R₁₀ and R₁₁ are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_a and R_b are each independently chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl and C₆₋₁₈ aralkyl;

or R_a and R_b are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R₁₆ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl and C₆₋₁₈ aralkyl;

R₁ is chosen from C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, C₆₋₁₈ aralkyl, or halogen;

R₂ is chosen from C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl, or C₆₋₁₈ aralkyl;

R₃ is chosen from H, C₁₋₁₂ alkyl, C₂₋₁₂ alkenyl, C₂₋₁₂ alkynyl, C₆₋₁₄ aryl, C₃₋₁₂ heterocycle, C₃₋₁₈ heteroaralkyl or C₆₋₁₈ aralkyl;

Z is chosen from H, halogen, C₁₋₆ alkyl;

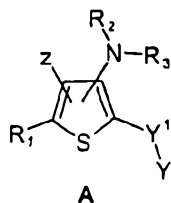
and one or more additional agent chosen from viral serine protease inhibitor, viral polymerase inhibitor and viral helicase inhibitor, immunomodulating agent, antioxydant agent, antibacterial agent or antisense agent.

115. The combination as defined in claim 114, wherein said additional agent is chosen from silybum marianum, interleukine-12, amantadine, ribozyme, thymosin, N-acetyl cysteine, cyclosporin, interferon α and ribavirin.

116. The combination as defined in anyone of claims 114 or 115, wherein said compound and said additionnal agent are administered sequentially.

117. The combination as defined in anyone of claims 114 or 115, wherein said compound and said additionnal agent are administered simultaneously.

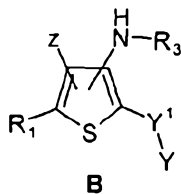
118. A process for preparing a compound of formula A:



said process comprising the steps of adding:

- an enol ether;
- an hydride donating agent; and
- an organic carboxylic acid;

to a compound of formula B:



wherein,

Y¹ is chosen from a bond, C₁₋₆ alkyl, C₃₋₆ alkenyl or C₂₋₆ alkynyl;

Y is chosen from COOR₆, COCOOR₅, P(O)OR₆OR₆, S(O)OR₅, S(O)₂OR₅, tetrazole, CON(R₉)CH(R₅)COOR₅, CONR₁₀R₁₁, CON(R₉)-SO₂-R₅, CONR₅OH

or halogen, wherein R_3 , R_5 , R_{10} and R_{11} are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl; or R_{10} and R_{11} are taken together with the nitrogen to form a 3 to 10 membered heterocycle;

R_4 and R_6 are each independently chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

or R_4 and R_6 are taken together with the oxygens to form a 5 to 10 membered heterocycle;

R_{16} is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl and C_{6-18} aralkyl;

R_1 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, C_{6-18} aralkyl or halogen;

R_2 is chosen from C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl, or C_{6-18} aralkyl;

R_3 is chosen from H, C_{1-12} alkyl, C_{2-12} alkenyl, C_{2-12} alkynyl, C_{6-14} aryl, C_{3-12} heterocycle, C_{3-18} heteroaralkyl or C_{6-18} aralkyl;

Z is chosen from H, halogen, C_{1-6} alkyl.