



(12) **CORRECTED EUROPEAN PATENT SPECIFICATION**

Note: Bibliography reflects the latest situation

- (15) Correction information:
Corrected version no 1 (W1 B1)
Corrections, see page(s) 4-7, 27, 28, 34-36, 42-44, 51
- (48) Corrigendum issued on:
28.12.2005 Bulletin 2005/52
- (45) Date of publication and mention of the grant of the patent:
04.05.2005 Bulletin 2005/18
- (21) Application number: **99955931.3**
- (22) Date of filing: **27.10.1999**
- (51) Int Cl.7: **C07D 277/46**, C07D 417/12, A61K 31/426, A61K 31/427, A61P 35/00
- (86) International application number:
PCT/EP1999/008306
- (87) International publication number:
WO 2000/026202 (11.05.2000 Gazette 2000/19)

(54) **2-AMINO-THIAZOLE DERIVATIVES, PROCESS FOR THEIR PREPARATION, AND THEIR USE AS ANTITUMOR AGENTS**

2-AMINOTHIAZOLEDERIVATE, VERFAHREN ZU IHRER HERSTELLUNG UND IHRE VERWENDUNG ALS ANTITUMORMITTEL

DERIVES 2-AMINO-THIAZOLE, LEUR PROCEDE DE PREPARATION ET LEUR UTILISATION EN TANT QU'AGENTS ANTITUMORAUX

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| <p>(84) Designated Contracting States:
AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE
Designated Extension States:
AL LT LV MK RO SI</p> <p>(30) Priority: 30.10.1998 GB 9823871</p> <p>(43) Date of publication of application:
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January 1956 (1956-01-10) Columbus, Ohio, US;
abstract no. 964e, S.R.M.BUSHBY ET AL: "The
antitrichomonal activity of amidonitrothiazoles"
page 964; XP002130674 & J.PHARM. AND
PHARMACOL., vol. 7, 1955, pages 112-117, | EP-A- 0 261 503 | EP-A- 0 412 404 | WO-A-98/04536 | WO-A-99/65884 | DE-A- 1 642 352 | DE-A- 2 128 941 | FR-A- 1 488 625 | FR-A- 1 499 557 | US-A- 3 374 082 | US-A- 3 427 318 | US-A- 3 591 600 | US-A- 4 027 031 |
| EP-A- 0 261 503 | EP-A- 0 412 404 | | | | | | | | | | | | |
| WO-A-98/04536 | WO-A-99/65884 | | | | | | | | | | | | |
| DE-A- 1 642 352 | DE-A- 2 128 941 | | | | | | | | | | | | |
| FR-A- 1 488 625 | FR-A- 1 499 557 | | | | | | | | | | | | |
| US-A- 3 374 082 | US-A- 3 427 318 | | | | | | | | | | | | |
| US-A- 3 591 600 | US-A- 4 027 031 | | | | | | | | | | | | |

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- CHEMICAL ABSTRACTS, vol. 61, no. 3, 3 August 1964 (1964-08-03) Columbus, Ohio, US; abstract no. 3087, MAX ROBBA ET AL: "Synthesis of thiazoles and isothiazoles. Their action on Trichomonas vaginalis and Candida albicans" XP002130675 & ANN. PHARM. FRANCO., vol. 22, no. 3, 1964, pages 201-210,
- PETER J. ISLIP ET AL: "Schistosomicidal 5-nitro-4-thiazolines" JOURNAL OF MEDICINAL CHEMISTRY., vol. 15, no. 9, 1972, pages 951-954, XP002130668 AMERICAN CHEMICAL SOCIETY. WASHINGTON., US ISSN: 0022-2623
- ROGER D. WESTLAND ET AL: "Novel schistosomicides. S-2-[[2-(2-thiazolylcarbonyl)ethyl]amino]ethyl hydrogen thiosulfate and related compounds" JOURNAL OF MEDICINAL CHEMISTRY., vol. 14, no. 10, 1971, pages 916-920, XP002130669 AMERICAN CHEMICAL SOCIETY. WASHINGTON., US ISSN: 0022-2623
- LEIF GREHN: "A method for nitration of thiazoles" JOURNAL OF HETEROCYCLIC CHEMISTRY., vol. 14, no. 5, August 1977 (1977-08), pages 917-919, XP002130670 HETEROCORPORATION. PROVO., US ISSN: 0022-152X
- H.ERLENMEYER ET AL: "Zur Kenntnis der Thiazol-4-sulfonsäure und der Thiazol-5-sulfonsäure" HELVETICA CHIMICA ACTA., vol. 28, 1945, pages 985-991, XP002130671 VERLAG HELVETICA CHIMICA ACTA. BASEL., CH ISSN: 0018-019X
- CHARLES D. HURD ET AL: "The 2-aminothiazoles" JOURNAL OF THE AMERICAN CHEMICAL SOCIETY., vol. 71, December 1949 (1949-12), pages 4007-4010, XP002130672 AMERICAN CHEMICAL SOCIETY, WASHINGTON, DC., US ISSN: 0002-7863
- TIMOTHY N. BIRKINSHAW ET AL: "Tautomerism in 2-trichloro- and 2-trifluoro-acetamidothiazoles" JOURNAL OF THE CHEMICAL SOCIETY, PERKIN TRANSACTIONS 1., 1982, pages 939-943, XP002130673 CHEMICAL SOCIETY. LETCHWORTH., GB ISSN: 0300-922X
- CHEMICAL ABSTRACTS, vol. 81, no. 5, 5 August 1974 (1974-08-05) Columbus, Ohio, US; abstract no. 22258q, page 156; XP002130676 & JP 48 027467 B (SANKYO CO LTD) 22 August 1973 (1973-08-22) cited in the application

Description

- 5 **[0001]** The present invention relates to 2-amino-thiazole derivatives, to a process for their preparation, to pharmaceutical compositions containing them and to their use as therapeutic agents, particularly in the treatment of cancer and cell proliferative disorders.
- [0002]** Several cytotoxic drugs such as, e.g. fluorouracil (5-FU), doxorubicin and camptothecins result to damage DNA or to affect cellular metabolic pathways and thus cause, in many cases, an indirect block of the cell cycle.
- [0003]** Therefore, by producing an irreversible damage to both normal and tumor cells, these agents result in a significant toxicity and side-effects.
- 10 **[0004]** In this respect, compounds capable of being highly specific antitumor agents by selectively leading to tumor cell arrest and apoptosis, with comparable efficacy but reduced toxicity than the currently available drugs, are desirable.
- [0005]** It is well known in the art that progression through the cell cycle is governed by a series of checkpoint controls, otherwise referred to as restriction points, which are regulated by a family of enzymes known as the cyclin-dependent kinases (cdk).
- 15 **[0006]** In their turn, the cdks themselves are regulated at many levels such as, for instance, binding to cyclins.
- [0007]** A normal progression through the cell cycle is controlled by the coordinated activation and inactivation of different cyclin/cdk complexes. In G1, both cyclin D/cdk4 and cyclin
- [0008]** E/cdk2 are thought to mediate the onset of S-phase.
- [0009]** Progression through S-phase requires the activity of cyclin A/cdk2 whereas the activation of cyclin A/cdc2 (cdk1) and cyclin B/cdc2 are required for the onset of metaphases.
- 20 **[0010]** For a general reference to cyclins and cyclin-dependent kinases see, for instance, Kevin R. Webster et al. in Exp.
- [0011]** Opin. Invest. Drugs, 1998, Vol. 7(6), 865-887.
- [0012]** Checkpoint controls are defective in tumor cells due, in part, to dysregulation of cdk activity. For example, altered expression of cyclin E and cdk's has been observed in tumor cells, and deletion of the cdk inhibitor p27 KIP gene in mice has been shown to result in a higher incidence of cancer.
- 25 **[0013]** Increasing evidence supports the idea that the cdks are rate-limiting enzymes in cell cycle progression and, as such, represent molecular targets for therapeutic intervention. In particular, the direct inhibition of cdk/cyclin kinase activity should be helpful in restricting the unregulated proliferation of a tumor cell.
- [0014]** It has now been found that the 2-amino-1,3-thiazoles of the invention are endowed with cdk/cyclin kinase inhibitory activity and are thus useful in therapy as antitumor agents whilst lacking, in terms of both toxicity and side effects, the aforementioned drawbacks known for currently available antitumor drugs.
- 30 **[0015]** More specifically, the compounds of this invention are useful in the treatment of a variety of cancers including, but not limited to: carcinoma such as bladder, breast, colon, kidney, liver, lung, including small cell lung cancer, esophagus, gall-bladder, ovary, pancreas, stomach, cervix, thyroid, prostate, and skin, including squamous cell carcinoma; hematopoietic tumors of lymphoid lineage, including leukemia, acute lymphocytic leukemia, acute lymphoblastic leukemia, B-cell lymphoma, T-cell-lymphoma, Hodgkin's lymphoma, non-Hodgkin's lymphoma, hairy cell lymphoma and Burkett's lymphoma; hematopoietic tumors of myeloid lineage, including acute and chronic myelogenous leukemias, myelodysplastic syndrome and promyelocytic leukemia; tumors of mesenchymal origin, including fibrosarcoma and rhabdomyosarcoma; tumors of the central and peripheral nervous system, including astrocytoma, neuroblastoma, glioma and schwannomas; other tumors, including melanoma, seminoma, teratocarcinoma, osteosarcoma, xeroderma pigmentosum, keratocanthoma, thyroid follicular cancer and Kaposi's sarcoma.
- 35 **[0016]** Due to the key role of cdks in the regulation of cellular proliferation, these 2-amino-1,3-thiazole derivatives are also useful in the treatment of a variety of cell proliferative disorders such as, for instance, benign prostate hyperplasia, familial adenomatosis, polyposis, neuro-fibromatosis, psoriasis, vascular smooth cell proliferation associated with atherosclerosis, pulmonary fibrosis, arthritis, glomerulonephritis and post-surgical stenosis and restenosis.
- [0017]** The compounds of the invention can be useful in the treatment of Alzheimer's disease, as suggested by the fact that cdk5 is involved in the phosphorylation of tau protein (*J.Biochem.*, 117, 741-749, 1995).
- 40 **[0018]** The compounds of this invention, as modulators of apoptosis, could be useful in the treatment of cancer, viral infections, prevention of AIDS development in HIV-infected individuals, autoimmune diseases and neurodegenerative disorder.
- [0019]** The compounds of this invention could be useful in inhibiting tumor angiogenesis and metastasis.
- [0020]** The compounds of this invention may also act as inhibitors of other protein kinases, e.g. protein kinase C, her2, raf1, MEK1, MAP kinase, EGF receptor, PDGF receptor, IGF receptor, PI3 kinase, weel kinase, Src, Abl and thus
- 45 **[0021]** be effective in the treatment of diseases associated with other protein kinases.
- 50 **[0021]** Several 2-amino-1,3-thiazole derivatives are known in the art. Just few examples among them are 2-acetamido-, 2-propionamido- or 2-butyramido-1,3-thiazole derivatives further substituted by halogen atoms in position 5 of the thiazole ring, which are reported as herbicides in JP 73027467 (Sankyo Co. Ltd.) or US 3,374,082 (The Upjohn

Co.); 5-nitro-2-benzamido-1,3-thiazole is reported as pesticide in Ann. Rech. Vet., 22(4), 359-63, 1991; 5-phenyl-2-acetamido-1,3-thiazoles further substituted onto phenyl ring are reported as synthetic intermediates (Chemical Abstracts, 1980, 92:128793); and 5-dimethylaminomethyl- or 5-diethylaminomethyl-2-acetamido-1,3-thiazole, both reported as herbicides in JP 71018564 (Japan Gas Chem Co.).

[0022] Several other 2-amino-1,3-thiazole derivatives have been reported in the art as useful therapeutic agents.

[0023] In particular, 5-methyl-1,3-thiazoles further substituted in position 2 of the thiazole ring by a benzothiazinyl-carbonylamino moiety, or derivatives thereof, have been described as cyclooxygenase inhibitors; see, for instance, C. A. 126(1997):301540.

[0024] 2-Benzamido-1,3-thiazoles are disclosed in EP-A-261503 (Valeas S.p.A.) as anti-allergic agents; 5-Alkyl-2-phenylalkylcarbonylamino-1,3-thiazoles further substituted onto the phenyl ring with an alkenylcarbonyl or alkynyl-carbonyl moieties are disclosed in WO 98/04536 (Otsuka Pharmaceutical Co.) as protein kinase C inhibitors. 5-Arylthio-2-acylamino-1,3-thiazole derivatives are disclosed in EP-A-412404 (Fujisawa Pharm. Co.) as antitumor agents.

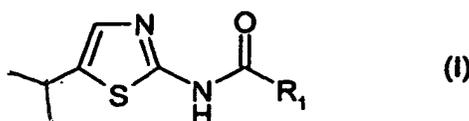
[0025] In addition, among the compounds reported in the art as therapeutic agents, DE 2128941 (Melle-Bezons) discloses 2-aminomethylcarbonylamino-5-chloro-1,3-thiazoles as anti-inflammatory, sedative and analgesic agents; the compound 2-diethylaminomethylcarbonylamino-5-chloro-1,3-thiazole being specifically exemplified therein.

[0026] Moreover, 5-halogeno-thiazoles with anti-inflammatory activity are disclosed in US 4,027,031. Thiazoles being substituted in position 5 by halogen atoms or nitro groups are also disclosed in C.A. 61(1964):3087; C.A. 50(1956):964e; C.A. 81(1974):22258q; Helvetica Chimica Acta 28, 1945, 985-91; J. Am. Chem. Soc., 71, 1949, 4007-10; J.

[0027] Med. Chem., 15(9), 1972, 951-954; J. Med. Chem., 14(10), 1971, 916-20; US 3,427,318; FR 1499557; US 3,591,600; FR 1488625; and DE 1642352.

[0028] Finally, 5-methyl-, ethyl- or benzyl-thiazoles are also disclosed in J. Heterocycl. Chem., 14(5), 1977, 917-19; and J. Chem. Soc. Perkin Trans., (1982), 939-43.

[0029] Accordingly, the present invention provides the use of a compound which is a 2-amino-1,3-thiazole derivative of formula (I)



wherein

R₁ is an optionally substituted group selected from:

- i) aryl;
- ii) arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

or a pharmaceutically acceptable salt thereof; in the manufacture of a medicament for treating cell proliferative disorders associated with an altered cell dependent kinase activity.

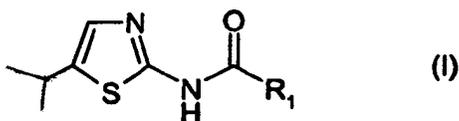
[0030] According to a preferred embodiment of the invention, the said cell proliferative disorder is selected from the group consisting of cancer, Alzheimer's disease, viral infections, auto-immune diseases or neurodegenerative disorders.

[0031] Preferably, the cancer is selected from the group consisting of carcinoma, squamous cell carcinoma, hematopoietic tumors of myeloid or lymphoid lineage, tumors of mesenchymal origin, tumors of the central and peripheral nervous system, melanoma, seminoma, teratocarcinoma, osteosarcoma, xenoderoma pigmentosum, keratocanthoma, thyroid follicular cancer and Kaposi's sarcoma.

[0032] According to another preferred embodiment of the invention, the cell proliferative disorder is selected from the group consisting of benign prostate hyperplasia, familial adenomatous polyposis, neuro-fibromatosis, psoriasis, vascular smooth cell proliferation associated with atherosclerosis, pulmonary fibrosis, arthritis glomerulonephritis and post-surgical stenosis and restenosis.

[0033] In addition, being useful in the treatment of cell proliferative disorders associated with an altered cell dependent kinase activity, hence cell cycle inhibition or cdk/cyclin dependent inhibition, the compounds of formula (I) of the invention also enable tumor angiogenesis and metastasis inhibition.

[0034] Therefore, it is a further object of the present invention a compound which is a 2-amino-1,3-thiazole derivative of formula (I)



wherein

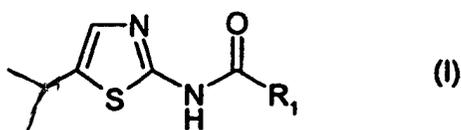
10 R_1 is an optionally substituted group selected from:

- i) aryl;
- ii) arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

15 or a pharmaceutically acceptable salt thereof; for use as a medicament.

[0035] The compounds of formula (I) above reported result to be novel.

[0036] Therefore, the present invention further provides a compound which is a 2-amino-1,3-thiazole derivative of formula (I)



25 wherein

30 R_1 is an optionally substituted group selected from:

- i) aryl;
- ii) arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

or a pharmaceutically acceptable salt thereof.

35 **[0037]** The compounds of formula (I) may have asymmetric carbon atoms and may therefore exist either as racemic admixtures or as individual optical isomers.

[0038] Accordingly, all the possible isomers and their admixtures and of both the metabolites and the pharmaceutically acceptable bio-precursors (otherwise referred to as pro-drugs) of the compounds of formula (I), as well as the uses thereof, are also within the scope of the present invention.

40 **[0039]** According to the above indicated substituent meanings, any of the above R_1 groups may be optionally substituted in any of the free positions by one or more groups, for instance 1 to 6 groups, selected from: halogen, nitro, oxo groups (=O), carboxy, cyano, alkyl, perfluorinated alkyl, alkenyl, alkynyl, cycloalkyl, aryl, heterocycyl; amino groups and derivatives thereof such as, for instance, alkylamino, alkoxy carbonylalkylamino, dialkylamino, arylamino, diarylamino or arylureido; carbonylamino groups and derivatives thereof such as, for instance, hydrogenocarbonylamino (HCONH-), alkylcarbonylamino, alkenylcarbonylamino, arylcarbonylamino, alkoxy carbonylamino; oxygen-substituted oximes such as, for instance, alkoxy carbonylalkoxyimino or alkoxyimino; hydroxy groups and derivatives thereof such as, for instance, alkoxy, aryloxy, alkylcarbonyloxy, arylcarbonyloxy, cycloalkenyloxy; carbonyl groups and derivatives thereof such as, for instance, alkylcarbonyl, arylcarbonyl, alkoxy carbonyl, aryloxy carbonyl, cycloalkyloxy carbonyl, aminocarbonyl, alkylaminocarbonyl, dialkylaminocarbonyl; sulfurated derivatives such as, for instance, alkylthio, arylthio, alkylsulphonyl, arylsulphonyl, alkylsulphinyl, arylsulphinyl, arylsulphonyloxy, aminosulphonyl, alkylaminosulphonyl or dialkylaminosulphonyl. In their turn, whenever appropriate, each of the above possible substituents on R_1 may be further substituted by one or more of the aforementioned groups.

[0040] In the present description, unless otherwise specified, with the term halogen atom we intend a chlorine, bromine, fluorine or iodine atom.

55 **[0041]** With the term optionally substituted amino group we intend an amino group wherein one or both hydrogen atoms are optionally replaced by other substituents which are the same or different, as set forth below.

[0042] With the term straight or branched C_1 - C_8 alkyl we intend a group such as, for instance, methyl, ethyl, n.propyl, isopropyl, n.butyl, isobutyl, sec-butyl, tert-butyl, n.pentyl, n.hexyl, n.heptyl, n.octyl and the like.

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[0043] With the term straight or branched C₂-C₆ alkenyl or alkynyl we intend a group such as, for instance, vinyl, allyl, isopropenyl, 1-, 2- or 3-butenyl, isobutylenyl, pentenyl, hexenyl, ethynyl, 1- or 2-propynyl, butynyl, pentynyl, hexynyl and the like.

[0044] With the term C₃-C₆ cycloalkyl we intend a cyclopropyl, cyclobutyl, cyclopentyl or cyclohexyl group.

[0045] With the term aryl, either as such or as arylalkyl, arylalkenyl, arylcarbonyl and the like, we intend a mono-, bi- or poly- either carbocyclic as well as heterocyclic hydrocarbon with from 1 to 4 ring moieties, either fused or linked to each other by single bonds, wherein at least one of the carbocyclic or heterocyclic rings is aromatic.

[0046] Examples of aryl groups are phenyl, indanyl, biphenyl, α - or β -naphthyl, fluorenyl, 9,10-dihydroanthracenyl, pyridyl, pyrazinyl, pyrimidinyl, pyridazinyl, indolyl, imidazolyl, 1,2-methylenedioxyphenyl, thiazolyl, isothiazolyl, pyrrolyl, pyrrolyl-phenyl, furyl, phenyl-furyl, benzo-tetrahydrofuran, oxazolyl, isoxazolyl, pyrazolyl, chromenyl, thienyl, benzothienyl, isoindolinyl, benzoimidazolyl, tetrazolyl, tetrazolylphenyl, pyrrolidinyl-tetrazolyl, isoindolinyl-phenyl, quinolinyl, isoquinolinyl, 2,6-diphenyl-pyridyl, quinoxalyl, pyrazinyl, phenyl-quinolyl, benzofurazanyl, 1,2,3-triazole, 1-phenyl-1,2,3-triazole, and the like.

[0047] With the term 3 to 6 membered carbocycle, hence encompassing but not limited to C₃-C₆ cycloalkyl groups, we also intend an unsaturated carbocyclic hydrocarbon such as, for instance, cyclopentylene or cyclohexylene.

[0048] With the term 5 to 7 membered heterocycle, hence encompassing aromatic heterocycles also referred to as aryl groups, we further intend a saturated or partially unsaturated 5 to 7 membered carbocycle wherein one or more carbon atoms are replaced by heteroatoms such as nitrogen, oxygen and sulphur.

[0049] Examples of 5 to 7 membered heterocycles, optionally benzocondensed or further substituted, are 1,3-dioxolane, pyran, pyrrolidine, pyrroline, imidazolidine, pyrazolidine, pyrazoline, piperidine, piperazine, N-alkyl-piperazine, morpholine, tetrahydrofuran and the like.

[0050] Examples of compounds of formula (I) wherein R₁ groups are substituted by one or more of the aforementioned substituents which, in their turn, are optionally further substituted as set forth above, are given below.

[0051] Pharmaceutically acceptable salts of the compounds of formula (I) are the acid addition salts with inorganic or organic, e.g. nitric, hydrochloric, hydrobromic, sulphuric, perchloric, phosphoric, acetic, trifluoroacetic, propionic, glycolic, lactic, oxalic, malonic, malic, maleic, tartaric, citric, benzoic, cinnamic, mandelic, methanesulphonic, isethionic and salicylic acid, as well as the salts with inorganic or organic bases, e.g. alkali or alkaline-earth metals, especially sodium, potassium, calcium or magnesium hydroxides, carbonates or bicarbonates, acyclic or cyclic amines, preferably methylamine, ethylamine, diethylamine, triethylamine or piperidine.

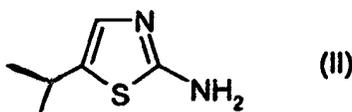
[0052] The compounds of formula (I) may have asymmetric carbon atoms and may therefore exist either as racemic admixtures or as individual optical isomers.

[0053] Accordingly, the use as an antitumor agent of all the possible isomers and their admixtures and of both the metabolites and the pharmaceutically acceptable bio-precursors (otherwise referred to as pro-drugs) of the compounds of formula (I) are also within the scope of the present invention.

[0054] Preferred compounds of formula (I), according to the present invention, are 2-amino-1,3-thiazole derivatives wherein ~~R₁~~ is an optionally substituted group selected from aryl or arylalkyl with from 1 to 4 carbon atoms within the alkyl chain.

[0055] Examples of preferred compounds of the invention, whenever appropriate in the form of pharmaceutically acceptable salts, e.g. hydrobromide or hydrochloride salt, are reported in the experimental section and claims.

[0056] The compounds of formula (I) object of the present invention and the salts thereof can be obtained, for instance, by a process comprising reacting a compound of formula (II)



with a compound of formula (III)



wherein R₁ are as defined above and X is hydroxy or a suitable leaving group;

and, if desired, converting a 2-amino-1,3-thiazole derivative of formula (I) into another such derivative of formula (I), and/or into a salt thereof.

[0057] Examples of specific compounds of formula (III) wherein X is a suitable leaving group are those wherein X

represents a halogen atom, preferably chlorine or bromine.

[0058] It is clear to the man skilled in the art that if the compound of formula (I), prepared according to the above process is obtained as an admixture of isomers, its separation into the single isomers according to conventional techniques is still within the scope of the present invention.

[0059] Likewise, the conversion into the free compound (I) of a corresponding salt thereof, according to well-known procedures in the art, is still within the scope of the invention.

[0060] The above process is an analogy process which can be carried out according to well known methods.

[0061] The reaction between a compound of formula (II) and a carboxylic acid of formula (III) wherein X is a hydroxy group, can be carried out in the presence of a coupling agent or a polymer supported coupling agent such as, for instance, carbodiimide, i.e. 1,3-dicyclohexylcarbodiimide, 1,3-diisopropylcarbodiimide, 1-(3-dimethylaminopropyl)-3-ethylcarbodiimide, or N-Cyclohexylcarbodiimide N'-methylpolystyrene in a suitable solvent such as, for instance, dichloromethane, chloroform, tetrahydrofuran, diethyl ether, 1,4-dioxane, acetonitrile, toluene, or N,N-dimethylformamide at a temperature ranging from about -10°C to reflux for a suitable time, i.e. from about 30 min. to about 96 hours.

[0062] The reaction between a compound of formula (II) and a compound of formula (III) can be also carried out, for example, by a mixed anhydride method, using an alkyl chloroformate, such as ethyl, iso-butyl, or iso-propyl chloroformate, in the presence of a tertiary base, such as triethylamine, N,N-diisopropylethylamine or pyridine, in a suitable solvent such as, for instance, toluene, dichloromethane, chloroform, tetrahydrofuran, acetonitrile, diethyl ether, 1,4-dioxane, or N,N-dimethylformamide, at a temperature ranging from about -30°C to room temperature.

[0063] The reaction between a compound of formula (II) and a carboxylic acid derivative of formula (III) wherein X is a suitable leaving group can be carried out in the presence of a tertiary base, such as triethylamine, N,N-diisopropylethylamine or pyridine, in a suitable solvent, such as toluene, dichloromethane, chloroform, diethyl ether, tetrahydrofuran, acetonitrile, or N,N-dimethylformamide, at a temperature ranging from about -10°C to reflux.

[0064] Also the optional conversion of a compound of formula (I) into another compound of formula (I) can be carried out according to known methods.

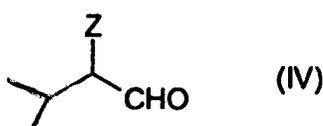
[0065] As an example, the nitro group of a compound of formula (I) may be converted into an amino group by treatment, for example, with stannous chloride in concentrated hydrochloric acid and by using, if necessary, an organic solvent such as acetic acid, 1,4-dioxane and tetrahydrofuran, at a temperature varying between room temperature and about 100°C.

[0066] Likewise, an alkylthio or an arylthio group may be converted into the corresponding alkylsulfonyl and arylsulfonyl group by reaction, for example, with m-chloroperbenzoic acid in a suitable solvent such as dichloromethane or chloroform, at a temperature varying between about -5°C and room temperature.

[0067] The optional salification of a compound of formula (I) or the conversion of a salt into the free compound as well as the separation of a mixture of isomers into the single isomers may be carried out by conventional methods.

[0068] The compounds of formula (II) and (III) according to the process object of the present invention are known compounds or can be obtained according to known methods.

[0069] For example, a compound of formula (II) can be obtained by reacting a compound of formula (IV)



wherein Z is a bromine or chlorine atom, with thiourea in a suitable solvent such as methanol, ethanol, tetrahydrofuran, 1,4-dioxane or toluene, at a temperature varying between room temperature and reflux, for a suitable time ranging from about 1 hour to about 24 hours.

[0070] A compound of formula (III) wherein X is a leaving group as defined above can be obtained according to conventional techniques from the corresponding carboxylic acids of formula (III) wherein X is hydroxy.

[0071] When preparing the compounds of formula (I) according to the process object of the present invention, optional functional groups within both the starting materials or the intermediates thereof, which could give rise to unwanted side reactions, need to be properly protected according to conventional techniques.

[0072] Likewise, the conversion of these latter into the free deprotected compounds may be carried out according to known procedures.

Pharmacology

[0073] The compounds of formula (I) are active as cdk/cyclin inhibitors as they gave positive results when tested according to the following procedure.

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[0074] The inhibiting activity of putative cdk/cyclin inhibitors and the potency of selected compounds was determined through a method of assay based on the use of the MultiScreen-PH 96 well plate (Millipore), in which a phosphocellulose filter paper was placed at each well bottom allowing binding of positive charged substrate after a washing/filtration step.

[0075] When a radioactivity labelled phosphate moiety was transferred by the ser/threo kinase to the filter-bound histone, light emitted was measured in a scintillation counter.

[0076] The inhibition assay of cdk2/Cyclin A activity was performed according to the following protocol:

Kinase reaction: 1.5 μ M histone H1 substrate, 25 μ M ATP (0.5 μ Ci $P^{33}\gamma$ -ATP), 30 ng Cyclin A/cdk2 complex, 10 μ M inhibitor in a final volume of 100 μ l buffer (TRIS HCl 10 mM pH 7.5, $MgCl_2$ 10 mM, 7.5 mM DTT) were added to each well of a 96 U bottom well plate. After 10 min at 37 °C incubation, reaction was stopped by 20 μ l EDTA 120 mM.

Capture: 100 μ l were transferred from each well to MultiScreen plate, to allow substrate binding to phosphocellulose filter. Plates were then washed 3 times with 150 μ l/well PBS Ca^{++}/Mg^{++} free and filtered by MultiScreen filtration system.

Detection: filters were allowed to dry at 37°C, then 100 μ l/well scintillant were added and ^{33}P labelled histone H1 was detected by radioactivity counting in the Top-Count instrument.

Results: data were analysed and expressed as % inhibition referred to total activity of enzyme (=100%).

[0077] All compounds showing inhibition ≥ 50 % were further analysed in order to study and define potency (IC50) as well as the kinetic-profile of inhibitor through K_i calculation.

[0078] IC50 determination: the protocol used was the same described above, where inhibitors were tested at concentrations ranging from 0.0045 to 10 μ M. Experimental data were analyzed by the computer program GraphPad Prism.

[0079] K_i calculation: either the concentration of ATP and histone H1 substrate were varied: 4, 8, 12, 24, 48 μ M for ATP (containing proportionally diluted $P^{33}\gamma$ -ATP) and 0.4, 0.8, 1.2, 2.4, 4.8 μ M for histone were used in absence and presence of two different, properly chosen inhibitor concentrations.

[0080] Experimental data were analysed by the computer program SigmaPlot for K_i determination, using a random bireactant system equation:

$$v = \frac{V_{max} \frac{(A)(B)}{aK_A K_B}}{1 + \frac{(A)}{K_A} + \frac{(B)}{K_B} + \frac{(A)(B)}{aK_A K_B}}$$

where A=ATP and B=histone H1.

[0081] Following the method above described, a representative compound of formula (I) of the invention, which is 2-[4-(dimethylamino)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide, showed an inhibiting activity towards the cdk2/cyclin A complex corresponding to 0.1 μ M (K_i).

[0082] In addition, the inhibiting activity of putative cdk/cyclin inhibitors and the potency of selected compounds was determined through a method of assay based on the use of a SPA (Scintillation Proximity Assay) 96 well plate assay. The assay is based on the ability of streptavidin coated SPA beads to capture a biotinylated peptide derived from a phosphorylation site of histone.

[0083] When a radioactivity labelled phosphate moiety was transferred by the ser/threo kinase to the biotinylated histone peptide, light emitted was measured in a scintillation counter.

[0084] The inhibition assay of cdk5/p25 activity was performed according to the following protocol:

Kinase reaction: 1.0 μ M biotinylated histone peptide substrate, 0.25 μ Ci $P^{33}\gamma$ -ATP, 4 nM cdk2/p25 complex, 0-100 μ M inhibitor in a final volume of 100 μ l buffer (Hepes 20 mM pH 7.5, $MgCl_2$ 15 mM, 1 mM DTT) were added to each well of a 96 U bottom well plate. After 20 min at 37 °C incubation, the reaction was stopped by the addition of 500 μ g SPA beads in phosphate-buffered saline containing 0.1% Triton X-100, 50 μ M ATP and 5 mM EDTA. The beads were allowed to settle, and the radioactivity incorporated in the ^{33}P -labelled peptide was detected in a Top Count scintillation counter.

Results: Data were analyzed and expressed as % Inhibition using the formula:

$$100X(1 - (\text{Unknown} - \text{Bkgd})/(\text{Enz. Control} - \text{Bkgd}))$$

IC50 values were calculated using a variation of the four parameter logistics equation:

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$$Y = 100/[1 + 10^{((\text{LogEC50} - X) \cdot \text{Slope})}]$$

Where X =log(uM) and Y = % Inhibition.

5

[0085] The compounds of formula (I) are therefore useful to restrict the unregulated proliferation of tumor cells, hence in therapy in the treatment of various tumors such as, for instance, carcinomas, e.g. mammary carcinoma, lung carcinoma, bladder carcinoma, colon carcinoma, ovary and endometrial tumors, sarcomas, e.g. soft tissue and bone sarcomas, and the hematological malignancies such as, e.g., leukemias.

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[0086] In addition, the compounds of formula (I) are also useful in the treatment of other cell proliferative disorders such as psoriasis, vascular smooth cell proliferation associated with atherosclerosis and post-surgical stenosis and restenosis and in the treatment of Alzheimer's disease.

15

[0087] The compounds of the present invention can be administered either as single agents or, alternatively, in combination with known anticancer treatments such as radiation therapy or chemotherapy regimen in combination with cytostatic or cytotoxic agents.

[0088] As an example, the above compounds can be administered in combination with one or more chemotherapeutic agents such as, for instance, taxane, taxane derivatives, CPT-11, camptothecin derivatives, anthracycline glycosides, e.g. doxorubicin or epirubicin, etoposide, navelbine, vinblastine, carboplatin, cisplatin and the like, optionally within liposomal formulations thereof.

20

[0089] The compounds of formula (I) of the present invention, suitable for administration to a mammal, e.g. to humans, can be administered by the usual routes and the dosage level depends upon the age, weight, conditions of the patient and the administration route.

[0090] For example, a suitable dosage adopted for oral administration of a compound of formula (I) may range from about 10 to about 500 mg pro dose, from 1 to 5 times daily.

25

[0091] The compounds of the invention can be administered in a variety of dosage forms, e.g. orally, in the form of tablets, capsules, sugar or film coated tablets, liquid solutions or suspensions; rectally in the form of suppositories; parenterally, e.g. intramuscularly, or by intravenous and/or intrathecal and/or intraspinal injection or infusion.

[0092] The present invention also includes pharmaceutical compositions comprising a compound of formula (I), or a pharmaceutically acceptable salt thereof, in association with a pharmaceutically acceptable excipient (which can be a carrier or a diluent).

30

[0093] The pharmaceutical compositions containing the compounds of the invention are usually prepared following conventional methods and are administered in a pharmaceutically suitable form.

[0094] For example, the solid oral forms may contain, together with the active compound, diluents, e.g. lactose, dextrose, saccharose, sucrose, cellulose, corn starch or potato starch; lubricants, e.g. silica, talc, stearic acid, magnesium or calcium stearate, and/or polyethylene glycols; binding agents, e.g. starches, arabic gum, gelatine, methylcellulose, carboxymethylcellulose or polyvinyl pyrrolidone; disaggregating agents, e.g. a starch, alginic acid, alginates or sodium starch glycolate; effervescent mixtures; dyestuffs; sweeteners; wetting agents such as lecithin, polysorbates, laurylsulphates; and, in general, non-toxic and pharmacologically inactive substances used in pharmaceutical formulations. Said pharmaceutical preparations may be manufactured in known manner, for example, by means of mixing, granulating, tableting, sugar-coating, or film-coating processes.

35

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[0095] The liquid dispersions for oral administration may be e.g. syrups, emulsions and suspensions.

[0096] The syrups may contain as carrier, for example, saccharose or saccharose with glycerine and/or mannitol and/or sorbitol.

[0097] The suspensions and the emulsions may contain as carrier, for example, a natural gum, agar, sodium alginate, pectin, methylcellulose, carboxymethylcellulose, or polyvinyl alcohol.

45

[0098] The suspension or solutions for intramuscular injections may contain, together with the active compound, a pharmaceutically acceptable carrier, e.g. sterile water, olive oil, ethyl oleate, glycols, e.g. propylene glycol, and, if desired, a suitable amount of lidocaine hydrochloride. The solutions for intravenous injections or infusions may contain as carrier, for example, sterile water or preferably they may be in the form of sterile, aqueous, isotonic saline solutions or they may contain as a carrier propylene glycol.

50

[0099] The suppositories may contain together with the active compound a pharmaceutically acceptable carrier, e.g. cocoa butter, polyethylene glycol, a polyoxyethylene sorbitan fatty acid ester surfactant or lecithin.

[0100] The following examples illustrate but do not limit the present invention.

55

Example 1

Preparation of N-(5-Bromo-thiazol-2-yl)-2-ethoxy-acetamide

5 **[0101]** EDCI (0.53 g, 2.78 mmol) was added to a solution of ethoxyacetic acid (0.29 g, 2.78 mmol) in CH₂Cl₂ (5 ml) under ice-cooling.

[0102] After stirring for 1 h, a solution of 2-amino-5-bromothiazole hydrobromide (0.60 g, 2.31 mmol) and diisopropylethylamine (0.40 ml, 2.34 mmol) in CH₂Cl₂ (5 ml) was added dropwise, and the entire mixture was kept at 0°C for 1 h, then at room temperature overnight.

10 **[0103]** The solution was evaporated and the residue partitioned between ethyl acetate and water. The ethyl acetate layer was further washed with water, 5% citric acid, water, saturated sodium bicarbonate, and water.

[0104] Drying over sodium sulfate and evaporation gave a solid which was triturated with isopropyl ether to give the title compound as a beige solid (0.43 g; 70%)
m.p. 100-102°C

15 ¹H-NMR (CDCl₃) δ ppm: 9.64 (bs, 1H, CONH); 7.38 (s, 1H, thiazole CH); 4.16 (s, 2H, COCH₂O); 3.65 (q, J = 6.8 Hz, 2H, OCH₂CH₃); 1.29 (t, J = 6.8 Hz, 3H, OCH₂CH₃).

[0105] Analogously, the following products can be prepared:

tert-butyl 4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}phenylcarbamate

20 tert-butyl 4-[[[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]phenylcarbamate

tert-butyl 4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}-1,3-thiazol-2-ylcarbamate;

N-(5-isopropyl-1,3-thiazol-2-yl)-2-bromoacetamide;

2-N-[2-(3-pyridyl)-acetyl-amino]-5-isopropyl-thiazole

m.p. 178-180°C (dec.)

25 ¹H-NMR (DMSO-d₆) δ ppm: 12.20 (bs, 1H, CONH); 8.45, 7.7, 7.35 (m, 4H, Py); 7.17 (s, 1H, thiazole CH); 3.78 (s, 2H, COCH₂); 3.14 (m, 1H, CH(Me)₂); 1.22 (d, 6H, CHMe₂);

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide

m.p. 206-208°C.

30 ¹H-NMR (DMSO-d₆) δ ppm: 12.1 (bs, 1H, CONH); 9.34 (s, 1H, OH); 7.14 (s, 1H, thiazole CH); 7.1 (t, 1H, H₅ Ph); 6.6-6.7

(m, 3H, H₂, H₄, H₆ Ph); 3.6 (s, 2H, COCH₂); 3.08 (ept, 1H, CHMe₂); 1.22 (d, 6H, CHMe₂);

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide

m.p. 97-98°C.

35 ¹H-NMR (DMSO-d₆) δ ppm: 12.12 (s, 1H, CONH); 7.21 (dd, 1H, H₅ Ph); 7.14 (d, 1H, thiazole CH); 6.87 (m, 2H, H₂, H₆ Ph); 6.81 (ddd, 1H, H₄ Ph); 3.72 (s, 3H, OMe); 3.67 (s, 2H, COCH₂); 3.07 (m, 1H, CHMe₂); 1.22 (d, 6H, CHMe₂);

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide m.p. 116-118°C.

¹H-NMR (CDCl₃) δ ppm: 11.8 (bs, 1H, CONH); 7.32 (s, 1H, H₂ Ph); 7.24 (m, 3H, H₄, H₅, H₆ Ph); 7.04 (s, 1H, thiazole CH); 3.76 (s, 2H, COCH₂); 3.13 (m, 1H, CHMe₂); 1.31 (d, 6H, CHMe₂);

40 N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamide

¹H-NMR (DMSO-d₆) δ ppm: 12.07 (bs, 1H, CONH); 9.33 (sb, 1H, OH); 7.17-6.7 (m, 5H, Ar+ CHthiazole); 3.60 (s, 2H, COCH₂); 3.1 (m, 1H, CHMe₂); 1.23 (d, 6H, CHMe₂);

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dihydroxyphenyl)acetamide

m.p. 168-169°C.

45 ¹H-NMR (DMSO-d₆) δ ppm: 12.01 (bs, 1H, CONH); 8.79 (sb, 2H, 2 OH); 7.12 (s, 1H, thiazole CH); 6.69 (d, 1H, H₂ Ph); 6.63 (d, 1H, H₅ Ph); 6.52 (dd, 1H, H₆ Ph); 3.48 (s, 2H, COCH₂); 3.06 (m, 1H, CHMe₂); 1.22 (d, 6H, CHMe₂);

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxy-3-methoxyphenyl)acetamide

m.p. 115-116°C.

50 ¹H-NMR (DMSO-d₆) δ ppm: 12.0 (bs, 1H, CONH); 8.80 (s, 1H, OH); 7.12 (d, 1H, thiazole CH); 6.88 (s, 1H, H₂ Ph); 6.68 (m, 2H, H₅, H₆ Ph); 3.73 (s, 3H, OMe); 3.56 (s, 2H, COCH₂); 3.07 (m, 1H, CHMe₂); 1.22 (d, 6H, CHMe₂);

N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide

m.p. 129-130°C.

¹H-NMR (DMSO-d₆) δ ppm: 12.08 (s, 1H, CONH); 7.21 (dd, 2H, H₂, H₆ Ph); 7.13 (d, 1H, thiazole CH); 6.87 (dd, 2H, H₃, H₅ Ph); 3.70 (s, 3H, OMe); 3.62 (s, 2H, COCH₂); 3.06 (m, 1H, CHMe₂); 1.22 (d, 6H, CHMe₂);

55 N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-acetamide

m.p. 135-137°C

¹H-NMR (DMSO-d₆) δ ppm: 12.20 (bs, 1H, CONH); 7.29 (m, 5H, Ph); 7.13 (s, 1H, thiazole CH); 3.70 (s, 2H, COCH₂); 3.07 (m, 1H, CHMe₂); 1.22 (d, 6H, CHMe₂);

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2-[3-(3-chloropropoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide m.p. 91-92°C

¹H-NMR (DMSO-d₆) δ ppm: 12.08 (bs, 1H, CONH); 7.21 (t, 1H, H5 Ph); 7.13 (s, 1H, thiazole CH); 6.8-6.9 (m, 3H, H2, H4, H6 Ph); 4.05 (t, 2H, OCH₂CH₂CH₂Cl); 3.77 (t, 2H, OCH₂CH₂CH₂Cl); 3.67 (s, 2H, COCH₂); 3.07 (ept, 1H, CHMe₂);

2.14 (quint, 2H, OCH₂CH₂CH₂Cl); 1.22 (d, 6H, CHMe₂); and 2-[3-(2-chloroethoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

m.p. 134-135°C

¹H-NMR (DMSO-d₆) δ ppm: 12.09 (bs, 1H, CONH); 7.22 (t, 1H, H5 Ph); 7.13 (s, 1H, thiazole CH); 6.8-6.9 (m, 3H, H2, H4, H6 Ph); 4.2 (t, 2H, OCH₂CH₂Cl); 3.91 (t, 2H, OCH₂CH₂Cl); 3.67 (s, 2H, COCH₂); 3.07 (ept, 1H, CHMe₂);

1.22 (d, 6H, CHMe₂).

Example 2

Preparation of N-(5-bromo-thiazol-2-yl)-4-sulfamoyl-benzamide.

[0106] To a mixture of 4-sulfamoylbenzoic acid (1.0 g, 4.97 mmol), Et₃N (1.5 ml, 10.78 mmol), DMF (5 ml) and THF (5 ml) isobutyl chloroformate (0.70 ml, 5.36 mmol) was added dropwise at -10°C.

[0107] After stirring for 1 h, a solution of 2-amino-5-bromothiazole hydrobromide (1.55 g, 5.96 mmol) and Et₃N (0.83 ml, 5.96 mmol) in DMF (6 ml) and THF (4 ml) was added dropwise to the mixture at the same temperature.

[0108] The resulting mixture was gradually warmed to room temperature over a period of 3 h and then concentrated by evaporation of the solvent in vacuo. To the resultant residue AcOEt and 5% aqueous NaHCO₃ were added. The separated organic phase was washed with water, dried over anhydrous Na₂SO₄, and concentrated under reduced pressure. The residual solid was purified by flash chromatography (dichloromethane/methanol/30% aqueous ammonia=95:5:0.5) to afford the title compound as a yellow solid (0.77 g, 43%)

m.p. 268-270°C

¹H-NMR (DMSO-d₆) δ ppm: 7.54 (s, 2H, SO₂NH₂); 7.67 (s, 1H, thiazole CH); 7.94 (d, J=8.8 Hz, 2H, H3 and H5 Ph); 8.21 (d, J=8.8 Hz, 2H, H2 and H6 Ph); 13.10 (bs, 1H, CONH).

[0109] Analogously, the following product can be prepared:

N-(5-isopropyl-thiazol-2-yl)-4-sulfamoyl-benzamide

m.p. 222-224°C.

¹H-NMR (DMSO-d₆) δ ppm: 12.65 (bs, 1H, CONH); 8.18 (dd, 2H, H2, H6 Ph); 7.92 (dd, 2H, H3, H5 Ph); 7.51 (s, 2H, SO₂NH₂); 7.25 (s, 1H, thiazole CH); 3.13 (m, 1H, CHMe₂); 1.28 (d, 6H, CHMe₂).

Example 3

Preparation of 3-amino-N-(5-bromo-1,3-thiazol-2-yl)propionamide hydrochloride

[0110] A solution 3.6 N HCl in isopropanol (14 ml) was added to tert-butyl 3-[(5-bromo-1,3-thiazol-2-yl)amino]-4-oxopropylcarbamate (0.90 g, 2.57 mmol) and the mixture was stirred at room temperature overnight. The solvent was evaporated and the residual solid was triturated in ether, filtered and dried in vacuo to afford the title compound as a white solid (0.73 g, quantitative yield)

m.p. 255°C ca.(dec.)

¹H-NMR (DMSO-d₆) δ ppm: 2.83 (t, J=6.8 Hz, 2H, COCH₂CH₂NH₂); 3.07 (q, J=6.4 Hz, 2H, COCH₂CH₂NH₂); 7.55 (s, 1H, thiazole CH); 7.96 (bs, 3H, NH₃⁺); 12.58 (s, 1H, CONH).

[0111] Analogously, the following compounds can be prepared:

2-(4-aminophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

¹H-NMR (DMSO-d₆) δ ppm: 1.22 (d, 6H, CHMe₂); 3.07 (m, 1H, CHMe₂); 3.47 (s, 2H, COCH₂); 4.94 (s, 2H, NH₂); 6.48 (m, 2H, H3, H5 Ph); 6.93 (m, 2H, H2, H6 Ph); 7.12 (d, 1H, CH thiazole); 12.00 (s, 1H, CONH).

4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

¹H-NMR (DMSO-d₆) δ ppm: 1.29 (d, 6H, CHMe₂); 3.12 (m, 1H, CHMe₂); 6.58 (m, 2H, H3, H5 Ph); 7.18 (d, 1H, CH thiazole); 7.82 (m, 2H, H2, H6 Ph); 12.80 (bs, 1H, CONH).

2-(2-amino-1,3-thiazol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

m.p. 204-206°C ca.(dec.)

¹H-NMR (DMSO-d₆) δ ppm: 1.24 (d, 6H, CHMe₂); 3.10 (m, 1H, CHMe₂); 3.54 (s, 2H, COCH₂); 6.30 (s, 1H, H5 thiazole); 6.88 (s, 2H, NH₂); 7.13 (s, 1H, H4 thiazole); 11.90 (s, 1H, CONH).

Example 4**2-amino-5-isopropyl-1,3-thiazole**

5 [0112] 2 ml (18.6 mmol) of 3-methylbutyraldehyde were dissolved in 15 ml of dioxane. 40.4 ml (18.6 mmol) of a solution 2 % v/v of bromine in dioxane was dropped therein at 0°C.

[0113] The mixture was maintained at room temperature under stirring for 2 hours, then 2.83 g (37.2 mmol) of thiourea and 5 ml of ethanol were added.

10 [0114] After 6 hours at room temperature the solution was evaporated to dryness, the residue was dissolved in methylene chloride and the product extracted with 1M hydrochloric acid; the aqueous layer was made basic by using 30% ammonium hydrate and extracted again with methylene chloride. The organic phase was dried over sodium sulfate and evaporated under vacuum. The residue was chromatographed on a silica gel column, eluting with cyclohexane-ethylacetate to give 1.1 g (42% yield) of the title compound.

15 ¹H-NMR (DMSO-d₆) δ ppm: 6.6 (s, 2H, NH₂); 6.58 (s, 1H, thiazole CH); 2.9 (m, 1H, CHMe₂); 1.18 (s, 3H, MeCHMe); 1.17 (s, 3H, MeCHMe).

Example 5**4-ethoxy-1-butanol**

20 [0115] 85 mg (0.004 mmol) of sodium were dissolved in 50 ml of methanol and 8.7 g (0.23 mol) of sodium borohydride were added. A solution of 4.6 g (0.032 mol) of methyl 4-ethoxy-butanoate in 20 ml of methanol was dropped to the mixture under stirring. The reaction is maintained at reflux for 6 hours, then 300 ml of brine were added and the product was extracted with methylene chloride. The organic layer was dried over anhydrous sodium sulfate and evaporated to dryness to give 2.25 g (61% yield) of the title compound.

25 [0116] Analogously the following products can be prepared starting from the suitable ester:

30 2-cyclopropyl-1-ethanol;
3-(3-indolyl)-1-propanol; and
5-dimethylaminoimino-1-hexanol.

Example 6**Methyl 3-(3-indolyl)-propanoate**

35 [0117] 2 g (10.57 mmol) of 3-indolepropionic acid were dissolved in 50 ml of methanol. The solution was cooled to 0°C and 5 ml of sulfuric acid 96% were dropped under stirring. The solution was maintained at room temperature overnight and then poured onto ice-water, basified with 30 % ammonium hydrate and finally extracted with methylene chloride. The organic layer was dried over anhydrous sodium sulfate and evaporated to dryness to give 2.3 g of an oily product (93% yield).

40 [0118] Analogously the following products can be prepared starting from the suitable carboxylic acid:

45 Methyl 4-ethoxy butanoate;
Methyl cyclopropylacetate; and
5-methoxycarbonylethyl-2-amino-1,3-thiazole.

Example 7**4-methyl-pentanal**

50 [0119] 1.24 ml (14.18 mmol) of oxalyl chloride were dissolved in 10 ml of methylene chloride and after cooling to -60°C, 2.31 ml of DMSO (35 mmoles) were dropped.

55 [0120] After 5 minutes at the same temperature, a solution of 1 ml (11.9 mmol) of 4-methyl-1-pentanol in 10 ml of methylene chloride was slowly dropped. The mixture was maintained under stirring for 30 minutes at the same temperature, then 8.3 ml (59.5 mmol) of triethylamine were added. After 2 hours at 0°C water was added. The mixture was diluted with methylene chloride and washed successively with 1M hydrochloric acid, water, saturated sodium bicarbonate and finally with brine. The organic layer was dried over anhydrous sodium sulfate and evaporated to dryness to give 0.7 g (25% yield) of the title compound.

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[0121] Analogously the following products can be prepared starting from the suitable alcohol:

2-cyclopropyl-1-ethanal;
4-methylthio-1-butanal;
4-ethoxy-1-butanal;
5-methoxy-1-pentanal; and
5-dimethylaminoimino-1-hexanal.

Example 8

5-benzyloxy-1-methoxy-pentane

[0122] 1.6 g (0.039 mol) of 55% sodium hydride in oil were added to 50 ml of dimethylformamide under stirring at room temperature. 5 ml (0.026 mol) of 5-benzyloxy-1-pentanol and 2.43 ml (0.039 mol) of methyl iodide were then added successively. After a night the excess of sodium hydride was decomposed with water and the solvent evaporated under vacuum. The residue was redissolved with methylene chloride and washed with water. The organic layer was finally dried over anhydrous sodium sulfate and evaporated to give 3.5 g (70% yield) of the title compound.

[0123] Analogously, by using ethyl iodide, the following compound can be prepared:

4-ethoxy-butanoic acid.

Example 9

5-methoxy-1-pentanol

[0124] 3.5 g (0.018 mol) of 5-benzyloxy-1-methoxy-pentane were dissolved in 50 ml of ethanol and 400 mg of 10% palladium on activated charcoal were added. The mixture was hydrogenated at 40 psi at room temperature for 5 hours, then filtered on celite and evaporated under vacuum to give 1.77 g (84% yield) of the title compound.

Example 10

Ethyl 5-dimethylaminoimino-hexanoate

[0125] 15.8 g (100 mmol) of ethyl 4-acetyl-butanoate and 6 g (100 mmol) of anhydrous N,N-dimethyl hydrazine in 50 ml of toluene containing 0.1 ml of trifluoroacetic acid were heated at 70 °C for 5 hours. The mixture was then washed with water, dried over anhydrous sodium sulfate and evaporated to give 12.3 g (79% yield) of the title compound.

Example 11

Preparation of methyl 2-[3-(3-chloropropoxy)phenyl]acetate

[0126] A mixture of methyl (m-hydroxyphenyl)acetate ((5 g, 0.03 moles), 1-bromo-3-chloropropane (3.26 ml, 0.03 moles) and anhydrous potassium carbonate (6.4 g) in anhydrous acetone (60 ml) was refluxed for 40 hours. After cooling, the precipitate was filtered off and the solution was evaporated to dryness to give the product as an oil, which was purified by flash chromatography with hexane:AcOEt (97:3) as eluent (6.2 g, 85% yield).

[0127] Analogously, the following product can be prepared:

methyl 2-[3-(2-chloroethoxy)phenyl]acetate.

Example 12

Preparation of 2-[3-(3-chloropropoxy)phenyl]acetic acid

[0128] A mixture of methyl 2-[3-(3-chloropropoxy)phenyl]acetate (4.95 g, 0.02 moles) and a solution of 1N sodium hydroxide (0.02 moles) was stirred at room temperature for 24 hours. After acidification the acid separated as white powder (4.53 g, 97% yield)
m.p. 83-84°C

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[0129] Analogously, the following product can be prepared:

2-[3-(2-chloroethoxy)phenyl]acetic acid

m.p. 100-101°C.

5 Example 13

Preparation of N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[3-(4-morpholinyl)propoxy]phenyl}acetamide

[0130] A mixture of 2-[3-(3-chloropropoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide (1.00 g, 2.8 mmoles), morpholine (1.24 ml, 14.2 mmoles), potassium iodide (0.24 g, 1.4 mmoles) in anhydrous dimethylformamide (3.5 ml) was heated at 100°C for 6 hours. The solution was acidified and extracted with ether to eliminate unreacted products; then the solution was basified and extracted with ether. The solvent was evaporated to dryness to give the product as an oily semisolid which was purified by flash chromatography with dichloromethane:methanol (97:3) as eluent (1.0 g, 87% yield)

¹H-NMR (DMSO-d₆) δ ppm: 12.09 (bs, 1H, CONH); 7.21 (t, 1H, H₅ Ph); 7.13 (s, 1H, thiazole CH); 6.8-6.9 (m, 3H, H₂, H₄, H₆ Ph); 3.97 (t, 2H, OCH₂CH₂CH₂N); 3.66 (s, 2H, COCH₂); 3.54 (t, 2H, OCH₂CH₂N); 3.07 (ept, 1H, CHMe₂); 2.39 (t, 2H, OCH₂CH₂CH₂N); 2.33 (t, 2H, OCH₂CH₂N); 2.14 (quint, 2H, OCH₂CH₂CH₂N); 1.22 (d, 6H, CHMe₂).

[0131] Analogously, the following product can be prepared:

N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[2-(4-morpholinyl)ethoxy]phenyl}acetamide

¹H-NMR (DMSO-d₆) δ ppm: 12.11 (bs, 1H, CONH); 7.20 (t, 1H, H₅ Ph); 7.13 (d, 1H, thiazole CH); 6.7-6.9 (m, 3H, H₂, H₄, H₆ Ph); 4.04 (t, 2H, OCH₂CH₂N); 3.66 (s, 2H, COCH₂); 3.55 (m, 4H, OCH₂CH₂N morpholine); 3.08 (m, 1H, CHMe₂); 2.66 (t, 2H,

OCH₂CH₂N); 2.44 (m, 4H, OCH₂CH₂N morpholine); 1.22 (d, 6H, CHMe₂);

N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[3-(1-pyrrolidiny)propoxy]phenyl}acetamide

¹H-NMR (DMSO-d₆) δ ppm: 12.1 (bs, 1H, CONH); 7.19 (t, 1H, H₅ Ph); 7.13 (d, 1H, thiazole CH); 6.7-6.9 (m, 3H, H₂, H₄, H₆ Ph); 3.97 (t, 2H, OCH₂CH₂CH₂N); 3.66 (s, 2H, COCH₂); 3.08 (m, 1H, CHMe₂); 2.50 (m, 2H, OCH₂CH₂CH₂N); 2.41 (m, 4H, CH₂N pyrrolidine); 1.85 (m, 2H, OCH₂CH₂CH₂N); 1.65 (m, 4H, CH₂CH₂N pyrrolidine); 1.23 (d, 6H, CHMe₂);

N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[3-(4-methyl-1-piperazinyl)propoxy]phenyl}acetamide

¹H-NMR (DMSO-d₆) δ ppm: 12.1 (bs, 1H, CONH); 7.19 (t, 1H, H₅ Ph); 7.13 (d, 1H, thiazole CH); 6.7-6.9 (m, 3H, H₂, H₄, H₆ Ph); 3.95 (t, 2H, OCH₂CH₂CH₂N); 3.66 (s, 2H, COCH₂); 3.08 (m, 1H, CHMe₂); 2.15-2.45 (m, 10H, OCH₂CH₂CH₂N+piperazine); 2.11 (s, 3H, NMe); 1.82 (m, 2H, OCH₂CH₂CH₂N); 1.22 (d, 6H, CHMe₂).

2-{3-[2-(dimethylamino)ethoxy]phenyl}-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

¹H-NMR (DMSO-d₆) δ ppm: 12.08 (bs, 1H, CONH); 7.2-6.90 (m, 5H, Ph+thiazole CH); 4.00 (t, 2H, OCH₂CH₂N); 3.66 (s, 2H, COCH₂); 3.07 (m, 1H, CHMe₂); 2.59 (t, 2H, OCH₂CH₂N); 2.11 (s, 3H, NMe); 2.19 (s, 6H, Me₂N); 1.22 (d, 6H, CHMe₂);

2-{3-[3-(dimethylamino)propoxy]phenyl}-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide

¹H-NMR (DMSO-d₆) δ ppm: 12.05 (bs, 1H, CONH); 7.19-6.79 (m, 5H, Ph+thiazole CH); 3.95 (t, 2H, OCH₂CH₂CH₂N); 3.66 (s, 2H, COCH₂); 3.08 (m, 1H, CHMe₂); 2.32 (t, 2H, OCH₂CH₂CH₂N); 2.11 (s, 3H, NMe₂); 1.81 (m, 2H, OCH₂CH₂CH₂N); 1.22 (d, 6H, CHMe₂).

Example 14

Preparation of 2-[N-[2'-N'-(ethoxycarbonyl-methyl)-amino]-acetyl]-amino-5-bromo-thiazole

[0132] A solution of N-(5-bromo-1,3-thiazol-2-yl)-2-bromoacetamide (0.35 g, 1.17 mmol) in DMF (5 ml) was added dropwise to a solution of glycine ethyl ester hydrochloride (0.33 g, 2.33 mmol) and triethylamine (0.49 ml, 3.5 mmol) in DMF (10 ml). After 3 hours at room temperature, the reaction mixture was heated at 40°C for about 5 hours and then diluted with water and extracted with methylene chloride. The combined organic layers were washed with brine, dried, concentrated and chromatographed on silica gel using cyclohexane: ethyl acetate 7:3 as eluent. The title compound was obtained as a colourless solid (0.15 g, 43%)

m.p. 115-116°C

¹H-NMR (DMSO-d₆) δ ppm: 7.46 (s, 1H, H₄thiaz), 4.05 (q, 2H, OCH₂CH₃), 3.49 (s, 2H, NHCOCH₂), 3.4 (s, 2H, NHCH₂), 1.18 (t, 3H, OCH₂CH₃).

[0133] Analogously, the following compound can be prepared:

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2-anilino-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide: m.p. 143-145°C

¹H-NMR (DMSO-d₆) δ ppm: 11.92 (s, 1H, NHCO), 7.13 (s, 1H, H4thiaz), 7.06-6.6 (m, 5H, Ph), 6.0 (t, 1H, NHCH₂), 3.95 (d, 2H, NHCH₂), 3.08 (m, 1H, CHMe₂), 1.23 (d, 6H, CHMe₂).

5 Example 15

Preparation of N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-bromophenyl)acetamide

[0134] To a suspension of resin N-Cyclohexylcarbodiimide N'-methylpolystyrene (0.251 g, 2.39 mmol g⁻¹, 0.6 mmol), previously washed with DCM (3X5 ml), in DCM (4 ml) at room temperature, 2-bromophenylacetic acid (0.086 g, 0.4 mmol) was added. After 10 min., a solution of 2-amino-5-isopropyl-1,3-thiazole (0.0284 g, 0.2 mmol) in DCM (4 ml) was added. The mixture was shaken for 24 hours at room temperature, the resin filtered and washed with DCM (3X10 ml). The filtrate were combined, washed with water, 5% HCl, water, saturated sodium bicarbonate and water, dried over sodium sulfate and evaporated.

¹H-NMR (DMSO-d₆) δ ppm: 10.05 (s broad, 1H, NHCOCH₂), 7.6-7.2 (m, 4H, Ar), 7.08 (s, 1H, H4thiaz), 3.98 (s, 2H, NHCOCH₂),

3.11 (m, 1H, CHMe₂), 1.31 (d, 6H, CHMe₂)

[0135] All the compounds were characterised by Mass Spectroscopy (MS). LC-MS confirmed that in each case the principle component had a molecular ion corresponding to the expected product.

[0136] Chromatography: Reverse phase HPLC with UV detection were run.

Mobile A: water (0.1% TFA)

Mobile B: acetonitrile:water 95:5 (0.1% TFA)

Flow rate: 1ml/min

Gradient: 10-100% B in 12 minutes, hold 100% B 3 min, return 10% B in 5 min

Detection: UV monitor 215, 254 and 300 nm

[0137] Sample were prepared as dilute solutions in acetonitrile (1-1.5 mM).

[0138] The compounds showed an HPLC area % ranging from 40 to 100%.

[0139] Starting from the suitable carboxylic acid, the following compounds can be prepared:

1. 2-N-[2-(3-pyridyl)-acetyl-amino]-5-isopropyl-thiazole;
2. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide;
3. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide;
4. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide;
5. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamide;
6. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dihydroxyphenyl)acetamide;
7. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxy-3-methoxyphenyl)acetamide;
8. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide;
9. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide;
10. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-acetamide;
11. N-(5-isopropyl-thiazol-2-yl)-4-sulfamoyl-benzamide;
12. 2-[3-(3-chloropropoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;
13. 2-[3-(2-chloroethoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;
14. 2-(4-aminophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;
15. 4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;
16. 2-(2-amino-1,3-thiazol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;
17. N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[3-(4-morpholinyl)propoxy]phenyl}acetamide;
18. N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[2-(4-morpholinyl)ethoxy]phenyl}acetamide;
19. N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[3-(1-pyrrolidinyl)propoxy]phenyl}acetamide;
20. N-(5-isopropyl-1,3-thiazol-2-yl)-2-{3-[3-(4-methyl-1-piperazinyl)propoxy]phenyl}acetamide;
21. 2-{3-[2-(dimethylamino)ethoxy]phenyl}-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;
22. 2-{3-[3-(dimethylamino)propoxy]phenyl}-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
23. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(2-methoxyethoxy)-phenyl]acetamide
24. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methyl-1-piperazinyl)benzamide

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- 25.2-anilino-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
26. (R)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide
27. (S)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide
28. N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
5 29. 2,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
30. 3,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
31. 3,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
32. 2,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
33. 2,3-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
10 34. 3-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
35. 2-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
36. 4-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
37. 3-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
38. 4-chloro-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
15 39. 5-bromo-2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
40. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
41. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
42. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
43. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
20 44. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
45. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
46. 2,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
47. 3,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
48. 2,3,4,5,6-pentafluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
25 49. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-3-nitrobenzamide
50. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-nitrobenzamide
51. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-nitrobenzamide
52. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-nitrobenzamide
53. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-2-nitrobenzamide
30 54. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-2-nitrobenzamide
55. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-3-nitrobenzamide
56. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-nitrobenzamide
57. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dinitrobenzamide
58. 5-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl-2-nitrophenyl octanoate
35 59. N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide
60. N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide
61. N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide
62. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)-3-nitrobenzamide
63. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide
40 64. 6-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide
65. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide
66. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide
67. 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide
68. 2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-5-nitrobenzamide
45 69. 4-fluoro-N-(5-isopropyl-2,3-thiazol-2-yl)-3-nitrobenzamide
70. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide
71. N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitro-4-(trifluoromethyl)benzamide
72. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-bis(trifluoromethyl)benzamide
73. N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-bis(trifluoromethyl)benzamide
50 74. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)benzamide
75. N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide
76. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide
77. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide
78. 5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide
55 79. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide
80. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide
81. methyl 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylbenzoate
82. methyl 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonylbenzoate

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83. 4-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
84. 3-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
85. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methylbenzamide
86. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methylbenzamide
5 87. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide
88. N-(5-isopropyl-1,3-thiazol-2-yl)-4-vinylbenzamide
89. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(2-phenylethynyl)benzamide
90. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-methylbenzamide
91. 2-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
10 92. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenethylbenzamide
93. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylbenzamide
94. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbenzamide
95. 4-(tert-butyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
96. N-(5-isopropyl-1,3-thiazol-2-yl)-4-isopropylbenzamide
15 97. N-(5-isopropyl-1,3-thiazol-2-yl)-4-pentylbenzamide
98. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide
99. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethylbenzamide
100. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethylbenzamide
101. 4-acetyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
20 102. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)benzamide
103. 5-(aminosulfonyl)-2,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
104. 5-(aminosulfonyl)-4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
105. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide
106. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide
25 107. 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide
108. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide
109. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxybenzamide
110. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide
111. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethoxybenzamide
30 112. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethoxybenzamide
113. N-(5-isopropyl-1,3-thiazol-2-yl)-2,4-dimethoxybenzamide
114. N-(5-isopropyl-1,3-thiazol-2-yl)-2,3-dimethoxybenzamide
115. N-(5-isopropyl-1,3-thiazol-2-yl)-3-phenoxybenzamide
116. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxybenzamide
35 117. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenoxybenzamide
118. 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
119. 4-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
120. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4,5-trimethoxybenzamide
121. 3,4-diethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
40 122. 3,4,5-triethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
123. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-(methoxymethoxy)benzamide
124. 4-butoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
125. N-(5-isopropyl-1,3-thiazol-2-yl)-4-propoxybenzamide
126. 4-isopropoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
45 127. N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-benzodioxole-5-carboxamide
128. 4-(benzyloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
129. 4-(2-cyclohexen-1-yloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
130. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethoxy)benzamide
131. 4-(difluoromethoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
50 132. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)benzamide
133. 2-[(4-chlorophenyl)sulfinyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
134. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[(4-nitrophenyl)sulfinyl]benzamide
135. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(4-methylphenyl)sulfonyl]-3-nitrobenzamide
136. N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(trifluoromethyl)sulfonyl]benzamide
55 137. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-4-(methylsulfonyl)benzamide
138. 2-[(2-cyanophenyl)sulfonyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
139. N~1~,N~1~-diethyl-3,6-difluoro-N~2~- (5-isopropyl-1,3-thiazol-2-yl)phthalamide
140. 4-formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide

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141. 2-formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
142. 4-[[[(2,5-dimethoxyanilino)carbonyl]amino]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
143. 4-(hydroxymethyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
144. 4-[[[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]-2-nitrobenzyl acetate
5 145. 4-[[[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]-2-nitrobenzyl 4-(acetylamino)-3-iodobenzoate
146. 4-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
147. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-phenylacetyl)amino]benzamide
148. 4-(acetylamino)-3-iodo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
149. 4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
10 150. 4-(dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
151. 3-(dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
152. 2-(methylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
153. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluoromethyl)anilino]benzamide
154. 3-[[[(5-bromo-1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)methyl]amino]-N-(5-isopropyl-1,3-thiazol-2-yl)benza-
15 mide
155. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(1H-pyrrol-1-yl)benzamide
156. 2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamide
157. 2-(4-bromophenyl)-6-(4-iodophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamide
158. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluoromethyl)anilino]nicotinamide
20 159. 2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide
160. 5,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide
161. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-6-methylnicotinamide
162. 2,6-dichloro-5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide
163. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxy nicotinamide
25 164. N-(5-isopropyl-1,3-thiazol-2-yl)-6-(2,2,2-trifluoroethoxy)nicotinamide
165. N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-dimethoxynicotinamide
166. N-(5-isopropyl-1,3-thiazol-2-yl)-2-quinoxalinecarboxamide
167. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-pyrazinecarboxamide
168. N-(5-isopropyl-1,3-thiazol-2-yl)-8-quinolinecarboxamide
30 169. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-4-quinolinecarboxamide
170. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1-phenyl-1H-pyrazole-4-carboxamide
171. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1H-pyrazole-3-carboxamide
172. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-pyrazole-4-carboxamide
173. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-phenyl-2H-1,2,3-triazole-4-carboxamide
35 174. 2-[[[(2,1,3-benzoxadiazol-5-yloxy)methyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxam-
ide
175. N-(5-isopropyl-1,3-thiazol-2-yl)-9H-fluorene-1-carboxamide
176. N-(5-isopropyl-1,3-thiazol-2-yl)-7-methoxy-1-benzofuran-2-carboxamide
177. N-(5-isopropyl-1,3-thiazol-2-yl)-1-[(4-methylphenyl)sulfonyl]-1H-pyrrole-3-carboxamide
40 178. 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide
179. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide
180. N-(5-isopropyl-1,3-thiazol-2-yl)-2-naphthamide
181. N-(5-isopropyl-1,3-thiazol-2-yl)-9,10-dioxo-9,10-dihydro-2-anthracenecarboxamide
182. N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-4-carboxamide
45 183. N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-1-carboxamide
184. N-(5-isopropyl-1,3-thiazol-2-yl)-8-oxo-5,6,7,8-tetrahydro-2-naphthalenecarboxamide
185. N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-dioxo-1,3-dihydro-2-benzofuran-5-carboxamide
186. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide
187. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-4-carboxamide
50 188. N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-2-phenyl-1H-indole-5-carboxamide
189. 2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-5-carboxamide
190. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-6-carboxamide
191. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methoxy-1H-indole-2-carboxamide
192. 1-allyl-2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide
55 193. N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-2-carboxamide
194. 1-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-1H-indole-5-carboxamide
195. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-1,2,3-benzotriazole-5-carboxamide
196. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-isoxazolecarboxamide

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197. N-(5-isopropyl-1,3-thiazol-2-yl)-3-thiophenecarboxamide
 198. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-thiophenecarboxamide
 199. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-thiophenecarboxamide
 200. 5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-thiophenecarboxamide
 201. N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(2,3,3-trichloroacryloyl)amino]-2-thiophenecarboxamide
 202. 5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide
 203. N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide
 204. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-nitrophenyl)-2-furamide
 205. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(2-nitrophenyl)-2-furamide
 206. 5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide
 207. N-(5-isopropyl-1,3-thiazol-2-yl)-5-[3-(trifluoromethyl)phenyl]-2-furamide
 208. 5-(4-chloro-2-nitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide
 209. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-methyl-2-nitrophenyl)-2-furamide
 210. 5-[2-chloro-5-(trifluoromethyl)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide
 211. tert-butyl (1R)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethylcarbamate
 212. (1R)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate
 213. (1S)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate
 214. (R,S)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
 215. (R)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
 216. (S)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
 217. 2-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
 218. (R,S)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
 219. (R)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
 220. (S)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
 221. 3,3,3-trifluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-2-phenylpropanamide
 222. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(1-naphthyl)acetamide
 223. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-naphthyl)acetamide
 224. 2-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 225. 2-(1,3-benzodioxol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 226. 2-(2,4-dinitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 227. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methyl-1H-indol-3-yl)acetamide
 228. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methyl-1H-indol-3-yl)acetamide
 229. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-methoxy-1H-indol-3-yl)acetamide
 230. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-benzyloxy-1H-indol-3-yl)acetamide
 231. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxy-2-methyl-1H-indol-3-yl)acetamide
 232. 2-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamide
 233. 2-(5-bromo-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 234. 2-(5-fluoro-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 235. 2-[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 236. 3-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide
 237. 4-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide
 238. N-(5-isopropyl-1,3-thiazol-2-yl)-3-(2-thienyl)propanamide
 239. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-thienyl)acetamide
 240. N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-(2-thienyl)acetamide
 241. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-thienyl)acetamide
 242. 2-(5-chloro-1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 243. 2-(1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 244. 2-[2-(formylamino)-1,3-thiazol-4-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide
 245. 2-{2-[(2-chloroacetyl)amino]-1,3-thiazol-4-yl}-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide
 246. 2-chloro-N-(4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}-1,3-thiazol-2-yl)acetamide
 247. ethyl 2-{{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(1H-pyrazol-3-yl)ethylidene}amino}oxy}acetate
 248. 2-(2-furyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamide
 249. 2-(5-bromo-3-pyridinyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 250. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(7-methoxy-2-oxo-2H-chromen-4-yl)acetamide
 251. N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxo-4-(4-methyl-phenyl)butanamide
 252. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-nitrophenyl)butanamide
 253. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbutanamide
 254. benzyl 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-oxobutylcarbamate

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255. methyl 5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate
 256. 4-(1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide
 257. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methoxy-1-naphthyl)-4-oxobutanamide
 5 258. 3-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide
 259. 3-(4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide
 260. 3-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide
 261. N-(5-isopropyl-1,3-thiazol-2-yl)-5-oxo-5-phenylpentanamide
 262. 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate
 263. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[4-(1-oxo-1,3-dihydro-2H-isoindol-2-yl)phenyl]propanamide
 10 264. 1-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide
 265. 1-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide
 266. 2-(3-bromo-4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 267. 2-(2-nitro-4-trifluoromethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 268. 5-cyclohexyl 1-(4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}benzyl) (2S)-2-[(tert-butoxycarbonyl)amino]pentanedioate
 15 269. 2-(5,6-dimethyl-1H-benzimidazol-1-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 270. 2-[5-(4-chlorophenyl)-2H-1,2,3,4-tetraazol-2-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 271. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[5-(1-pyrrolidinyl)-2H-1,2,3,4-tetraazol-2-yl]acetamide
 272. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methyl-1-benzothiophen-2-yl)acetamide
 20 273. N-(4-bromo-6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexyl)benzamide
 274. benzyl 6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexylcarbamate
 275. 4-(2,4-dimethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide
 276. 4-(4-benzyloxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide
 277. 4-(thiophen-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide
 25 278. benzyl 2-[(benzyloxy)carbonylamino]-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate
 279. 4-(1H-indol-3-yl)-N-{3-[(5-isopropyl-1,3-thiazol-2-yl)amino]-3-oxopropyl}butanamide
 280. 4-[[5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]phenyl 4-chlorobenzenesulfonate
 281. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-methoxyanilino)carbonylamino]benzamide
 282. 4-[[2-(isopropylsulfonyl)acetyl]amino]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
 30 283. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[[2-(phenylsulfonyl)acetyl]amino]benzamide
 284. 4-[(diethylamino)sulfonyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
 285. 2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
 286. 3,5-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
 287. 3-[[2-fluoroanilino)carbonylamino]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
 35 288. N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenyl-5-propyl-1H-pyrazole-4-carboxamide
 289. 3-chloro-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfonyl)-2-thiophenecarboxamide
 290. 3-iodo-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfonyl)-2-thiophenecarboxamide
 291. 2-[[4-chlorophenyl)sulfonyl]methyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxamide
 292. 5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)-3-furamide
 40 293. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,3,4,5,6-pentafluorophenyl)acetamide
 294. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-fluorophenyl)acetamide
 295. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-bromophenyl)acetamide
 296. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chlorophenyl)acetamide
 297. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenyl)acetamide
 45 298. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-trifluoromethylphenyl)acetamide
 299. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methoxyphenyl)acetamide
 300. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-dimethoxyphenyl)acetamide
 301. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-difluorophenyl)acetamide
 302. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4,5-trimethoxyphenyl)acetamide
 50 303. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,6-dichlorophenyl)acetamide
 304. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chloro-6-fluorophenyl)acetamide
 305. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-dimethoxyphenyl)acetamide
 306. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-difluorophenyl)acetamide
 307. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-bis-trifluoromethylphenyl)acetamide
 55 308. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylthiophenyl)acetamide
 309. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide
 310. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-bromophenyl)acetamide
 311. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide

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312. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-fluorophenyl)acetamide
313. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)acetamide
314. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-trifluoromethylphenyl)acetamide
315. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylphenyl)acetamide
5 316. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-dimethylaminophenyl)acetamide
317. 2-[1,1'-biphenyl]-4-yl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
318. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-trifluoromethylphenyl)acetamide
319. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-bromophenyl)acetamide
10 320. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide
321. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-nitrophenyl)acetamide
322. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide
323. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dinitrophenyl)acetamide
324. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dichlorophenyl)acetamide
325. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-difluorophenyl)acetamide
15 326. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-benzyloxy-3-methoxyphenyl) acetamide 327. N-(5-isopropyl-1,3-thiazol-
2-yl)-2-(3,4-dichlorophenyl)acetamide
328. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-difluorophenyl)acetamide
329. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dimethoxyphenyl)acetamide
330. 2-(2,3-dihydro-1H-inden-5-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
20 331. N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenylcyclopropanecarboxamide
332. 2-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
333. 2-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
334. N-(5-isopropyl-1,3-thiazol-2-yl)- 2,2-diphenylacetamide
335. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)propanamide
25 336. N-(5-isopropyl-2,3-thiazol-2-yl)-2-phenyl)propanamide
337. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-isobutylphenyl)propanamide
338. N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-phenylacetamide
339. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-phenylpentanamide
340. N-(5-isopropyl-1,3-thiazol-2-yl)bicyclo[4.2.0]octa-1,3,5-triene-7-carboxamide
30 341. N-(5-isopropyl-2,3-thiazol-2-yl)-3-oxo-1-indanecarboxamide
342. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl)butanamide
343. tert-butyl (1S)-1-(1H-indol-3-ylmethyl)-2-[(5-isopropyl-2,3-thiazol-2-yl)amino]-2-oxoethylcarbamate
344. tert-butyl (1S)-1-(4-benzyloxybenzyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate.

35 **[0140]** Following the same procedure as reported in Example 1, the compounds described in the table (I) below can
be prepared:

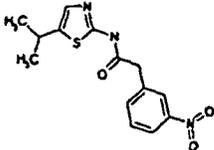
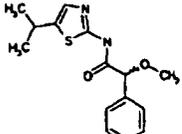
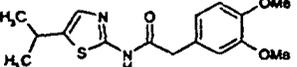
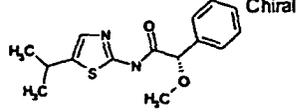
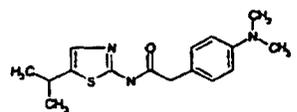
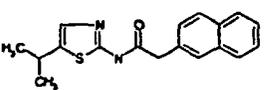
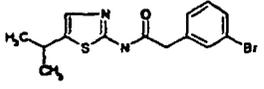
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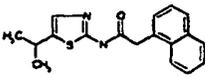
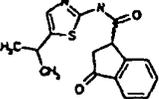
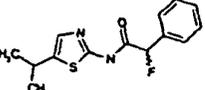
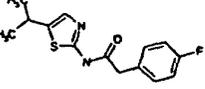
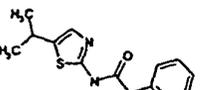
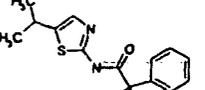
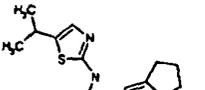
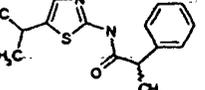
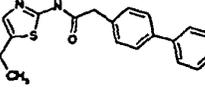
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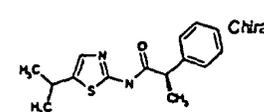
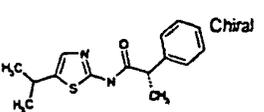
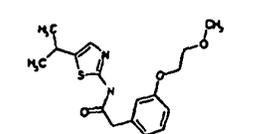
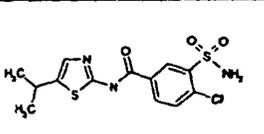
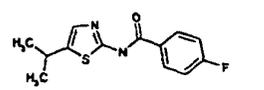
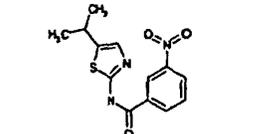
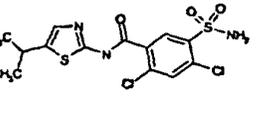
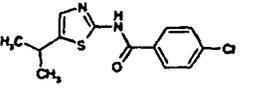
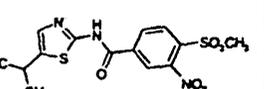
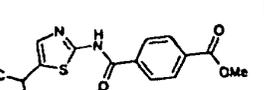
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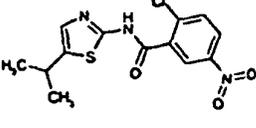
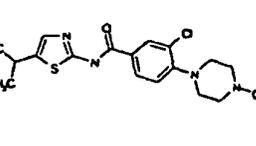
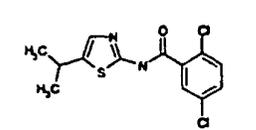
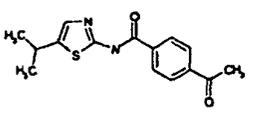
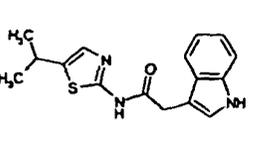
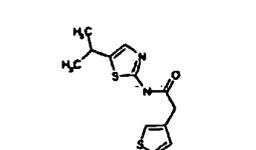
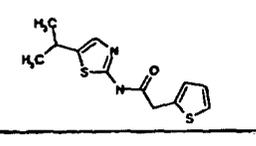
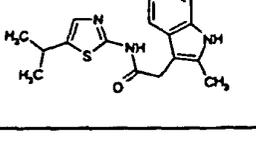
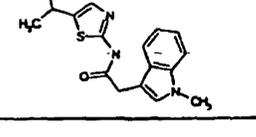
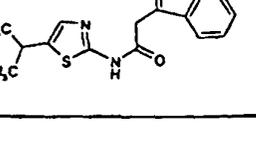
Table I

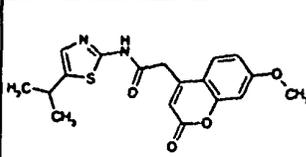
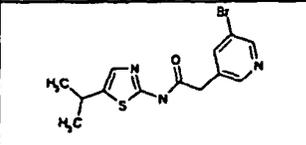
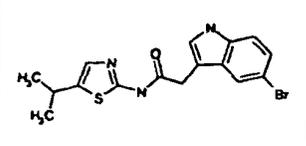
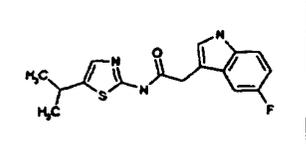
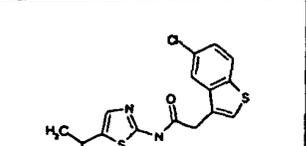
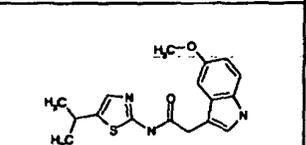
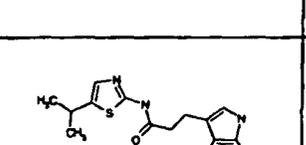
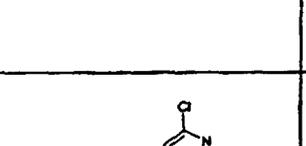
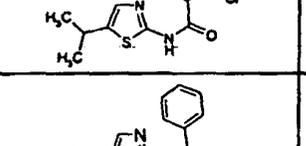
MOLSTRUCTURE	m p °C	¹ H-NMR	Sol vent
		12.23 (s broad, 1H, <u>NHCOCH₂</u>), 8.22-7.62 (m, 4H, Ar), 7.15 (s, 1H, H4thiaz), 3.91 (s, 2H, <u>NHCOCH₂</u>), 3.08 (m, 1H, <u>CHMe₂</u>), 1.22 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
		9.81 (s broad, 1H, <u>NHCOCH₂</u>), 7.5-7.3 (m, 4H, Ar), 7.11 (s, 1H, H4thiaz), 4.83 (s, 1H, <u>NHCOCH</u>), 3.44 (s, 3H, Ome) 3.11 (m, 1H, <u>CHMe₂</u>), 1.3 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
	124 - 125	12.06 (s broad, 1H, <u>NHCO</u>), 7.13 (s, 1H, H4thiaz) 6.92-6.81 (m, 3H, Ar), 3.72 (s, 3H, OMe), 3.70 (s, 3H, OMe), 3.61 (s, 2H, <u>NHCOCH₂</u>), 3.07 (m, 1H, <u>CHMe₂</u>), 1.22 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
 Chiral	77- 78	12.05 (s broad, 1H, <u>NHCO</u>), 7.38-7.29 (m, 5H, Ar), 7.12 (s, 1H, H4thiaz), 4.95 (s, 1H, <u>CHOMe</u>), 3.23 (s, 2H, <u>CHOMe</u>), 3.05 (m, 1H, <u>CHMe₂</u>), 1.20 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
	136 - 137	12.08 (s broad, 1H, <u>NHCOCH₂</u>), 7.28 (d, 2H, Ar), 7.13 (s, 1H, H4thiaz), 7.1 (d, 2H, Ar), 3.65 (s, 2H, <u>NHCOCH₂</u>), 3.06 (m, 1H, <u>CHMe₂</u>), 2.98 (s, 6H, <u>NMe₂</u>), 1.22 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
	130 - 131	12.22 (s, 1H, <u>NHCO</u>), 7.85-7.48 (m, 7H, Ar), 7.14 (s, 1H, H4thiaz), 3.89 (s, 2H, <u>CH₂CO</u>), 3.07 (m, 1H, <u>CHMe₂</u>), 1.22 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
	130 - 131	12.16 (s, 1H, <u>NHCO</u>), 7.52-7.29 (m, 4H, Ar), 7.14 (s, 1H, H4thiaz), 3.73 (s, 2H, <u>CH₂CO</u>), 3.08 (m, 1H, <u>CHMe₂</u>), 1.22 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶

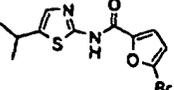
5		177 - 178	8.07-7.48 (m, 7H, Ar), 7.15 (s, 1H, H4thiaz), 4.22 (s, 2H, CH ₂ CO), 3.06 (m, 1H, CHMe ₂), 1.20 (d, 6H, CHMe ₂)	DMSO-d ⁶
10		223 - 224	12.61 (s, 1H, NHCO), 7.69-7.51 (m, 4H, Ar), 7.19 (s, 1H, H4thiaz), 4.55 (dd, 1H, CHCO), 3.08 (m, 1H, CHMe ₂), 2.89 (m, 2H, COCH ₂ CH), 1.22 (d, 6H, CHMe ₂)	DMSO-d ⁶
15		105 - 106	12.50 (s, 1H, NHCO), 7.53-7.51 (m, 5H, Ar), 7.18 (s, 1H, H4thiaz), 6.12 (d, 1H, J _{H-F} = 46.8, CHF), 3.09 (m, 1H, CHMe ₂), 1.22 (d, 6H, CHMe ₂)	DMSO-d ⁶
20		150 - 152	11.20 (s broad, 1H, NHCO), 7.28-7.07 (m, 5H, Ar+H4thiaz), 3.80 (s, 2H, CH ₂ CO), 3.13 (m, 1H, CHMe ₂), 1.32 (d, 6H, CHMe ₂)	DMSO-d ⁶
25		164 - 166	11.45 (s broad, 1H, NHCO), 7.37-7.14 (m, 5H, Ar+ H4thiaz), 3.88 (s, 2H, NHCOCH ₂), 3.12 (m, 1H, CHMe ₂), 1.32 (d, 6H, CHMe ₂)	DMSO-d ⁶
30		98- 100	8.35 (s broad, 1H, NHCO), 7.40 (m, 5H, Ar), 6.99 (s, 1H, H4thiaz), 3.10 (m, 1H, CHMe ₂), 1.78 (m, 2H, CH ₂), 1.29 (m, 2H, CH ₂), 1.25 (d, 6H, CHMe ₂)	CDCl ₃
35		130 - 132	12.06 (s broad, 1H, NHCOCH ₂), 7.13 (s, 1H, H4thiaz), 6.86-6.75 (m, 3H, Ar), 5.96 (s, 2H, OCH ₂ O), 3.60 (s, 2H, NHCOCH ₂), 3.05 (m, 1H, CHMe ₂), 1.22 (d, 6H, CHMe ₂)	DMSO-d ⁶
40		100 - 102	12.1 (s broad, 1H, NHCOCH ₂), 7.2-7 (m, 4H, Ar+ H4thiaz), 3.64 (s, 2H, NHCOCH ₂), 3.07 (m, 1H, CHMe ₂), 2.8-1.97 (m, 6H, -CH ₂ CH ₂ CH ₂ -), 1.22 (d, 6H, CHMe ₂)	DMSO-d ⁶
45		98- 100	12.06 (s broad, 1H, NHCO), 7.3 (m, 5H, Ar), 7.03 (s, 1H, H4thiaz), 3.79 (q, 1H, CHMe), 3.10 (m, 1H, CHMe ₂), 1.59 (d, 3H, CHMe), 1.30 (d, 6H, CHMe ₂)	DMSO-d ⁶
50		167 - 169	10 (s broad, 1H, NHCOCH ₂), 7.6-7.4 (m, 9H, Ar), 7.04 (s, 1H, H4thiaz), 3.84 (s, 2H, NHCOCH ₂), 3.11 (m, 1H, CHMe ₂), 1.31 (d, 6H, CHMe ₂)	DMSO-d ⁶
55				

5		115 - 116	12.06 (s broad, 1H, <u>NHCO</u>), 7.26 (m, 5H, Ar), 6.99 (s, 1H, H4thiaz), 3.79 (q, 1H, <u>CHMe</u>), 3.10 (m, 1H, <u>CHMe₂</u>), 1.59 (d, 3H, <u>CHMe</u>), 1.30 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
10		112 - 114	12.06 (s broad, 1H, <u>NHCO</u>), 7.33 (m, 5H, Ar), 7.11 (s, 1H, H4thiaz), 3.93 (q, 1H, <u>CHMe</u>), 3.07 (m, 1H, <u>CHMe₂</u>), 1.40 (d, 3H, <u>CHMe</u>), 1.22 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
15		88- 90	12.08 (s broad, 1H, <u>NHCO</u>), 7.20-6.81 (m, 5H, Ar+H4thiaz), 4.01 (dd, 2H, <u>OCH₂CH₂OMe</u>), 3.68 (s, 2H, <u>NHCOCH₂</u>), 3.61 (dd, 2H <u>OCH₂CH₂OMe</u>), 3.3 (s, 3H, <u>OCH₂CH₂OMe</u>), 3.05 (m, 1H, <u>CHMe₂</u>), 1.22 (s, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
20		230 - 231	12.81 (s broad, 1H, <u>NHCO</u>), 8.63-7.79 (m, 3H, Ar), 7.71 (s, 2H, <u>NH₂</u>), 7.24 (s, 1H, H4thiaz), 3.12 (m, 1H, <u>CHMe₂</u>), 1.27 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
25		181 - 182	12.47 (s broad, 1H, <u>NHCO</u>), 8.13-7.37 (m, 4H, Ar), 7.23 (s, 1H, H4thiaz), 3.13 (m, 1H, <u>CHMe₂</u>), 1.27 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
30			12.0 (s broad, 1H, <u>NHCO</u>), 8.89-7.82 (m, 4H, Ar), 7.27 (s, 1H, H4thiaz), 3.13 (m, 1H, <u>CHMe₂</u>), 1.28 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
35		263 - 264	12.74 (s broad, 1H, <u>NHCO</u>), 8.11-8.0 (2s, 2H, Ar), 7.82 (s, 2H, <u>NH₂</u>), 7.24 (s, 1H, H4thiaz), 3.15 (m, 1H, <u>CHMe₂</u>), 1.27 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
40		204 - 206	12.6 (s broad, 1H, <u>NHCO</u>), 8.06-7.60 (m, 3H, Ar), 7.23 (s, 1H, H4thiaz), 3.12 (m, 1H, <u>CHMe₂</u>), 1.27 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
45		148 - 150	8.54-8.31 (m, 3H, Ar), 6.98 (s, 1H, H4thiaz), 3.43 (s, 3H, <u>SO₂Me</u>), 3.14 (m, 1H, <u>CHMe₂</u>), 1.35 (d, 6H, <u>CHMe₂</u>)	CDCl ₃
50		173 - 175	8.16-8.06 (2d, 4H, Ar), 7.25 (s, 1H, H4thiaz), 3.88 (s, 3H, <u>COOMe</u>), 3.14 (m, 1H, <u>CHMe₂</u>), 1.28 (d, 6H, <u>CHMe₂</u>)	DMSO-d ⁶
55				

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	<p>164 - 166</p>	<p>8.50-7.86 (m, 3H, Ar), 7.24 (s, 1H, H4thiaz), 3.15 (m, 1H, CHMe₂), 1.28 (d, 6H, CHMe₂)</p>	<p>DMSO-d⁶</p>
	<p>178 - 179</p>	<p>12.4 (s broad, 1H, NHCO), 8.12-7.21 (m, 3H, Ar), 7.22 (s, 1H, H4thiaz), 3.2-2.48 (m, 5H, CHMe₂ + piperazine), 2.22 (s, 3H, NMe), 1.27 (d, 6H, CHMe₂)</p>	<p>DMSO-d⁶</p>
		<p>12.6 (s broad, 1H, NHCO), 7.73-7.57 (m, 3H, Ar), 7.22 (s, 1H, H4thiaz), 3.15 (m, 1H, CHMe₂), 1.27 (d, 6H, CHMe₂)</p>	<p>DMSO-d⁶</p>
		<p>12.6 (s broad, 1H, NHCO), 8.16-8.05 (m, 4H, Ar), 7.24 (s, 1H, H4thiaz), 3.13 (m, 1H, CHMe₂), 2.62 (s, 3H, COMe), 1.28 (d, 6H, CHMe₂)</p>	<p>DMSO-d⁶</p>
	<p>207 - 209</p>	<p>9.4 (s broad, 1H, NHCO), 8.3 (s, 1H, NH), 7.55-6.98 (m, 6H, indole+H4thiaz), 3.96 (s, 2H, COCH₂), 3.10 (m, 1H, CHMe₂), 1.30 (d, 6H, CHMe₂)</p>	<p>CDCl₃</p>
	<p>116 - 118</p>	<p>9.80 (s broad, 1H, NHCO), 7.37-7.05 (m, 3H, Ar), 7.04 (d, 1H, H4thiaz), 3.84 (s, 2H, COCH₂), 3.11 (m, 1H, CHMe₂), 1.32 (d, 6H, CHMe₂)</p>	<p>CDCl₃</p>
	<p>148 - 150</p>	<p>10.20 (s broad, 1H, NHCO), 7.28-7.01 (m, 4H, Ar+H4thiaz), 4.02 (s, 2H, COCH₂), 3.13 (m, 1H, CHMe₂), 1.32 (d, 6H, CHMe₂)</p>	<p>CDCl₃</p>
	<p>170 - 172</p>	<p>12.05 (s broad, 1H, NHCO), 10.82 (s, 1H, NH), 7.48-6.90 (m, 5H, indole+H4thiaz), 3.74 (s, 2H, COCH₂), 3.06 (m, 1H, CHMe₂), 2.36 (s, 3H, Me), 1.21 (d, 6H, CHMe₂)</p>	<p>DMSO-d⁶</p>
	<p>163 - 165</p>	<p>12.07 (s broad, 1H, NHCO), 7.57-7.01 (m, 6H, indole+H4thiaz), 3.79 (s, 2H, COCH₂), 3.74 (s, 3H, NMe), 3.05 (m, 1H, CHMe₂), 1.21 (d, 6H, CHMe₂)</p>	<p>DMSO-d⁶</p>
	<p>155 - 157</p>	<p>10.20 (s broad, 1H, NHCO), 7.88-7.40 (m, 5H, Ar), 6.95 (s, 1H, H4thiaz), 4.04 (s, 2H, COCH₂), 3.07 (m, 1H, CHMe₂), 1.27 (d, 6H, CHMe₂)</p>	<p>DMSO-d⁶</p>

5		234 - 236	11.3 (s broad, 1H, NHCO), 7.52-6.28 (m, 5H, Ar+H4thiaz), 3.93 (s, 2H, COCH ₂), 3.87 (s, 3H, OMe), 3.10 (m, 1H, CHMe ₂), 1.27 (d, 6H, CHMe ₂)	DMSO-d ⁶
10		164 - 167	8.63-7.9 (m, 5H, Ar), 7.11 (s, 1H, H4thiaz), 3.85 (s, 2H, COCH ₂), 3.15 (m, 1H, CHMe ₂), 1.29 (d, 6H, CHMe ₂)	CDCl ₃
15		217 - 220	12.09 (s broad, 1H, NHCO), 11.5 (s, 1H, NH), 7.78-7.16 (m, 4H, indole), 7.13 (s, 1H, H4thiaz), 3.78 (s, 2H, COCH ₂), 3.07 (m, 1H, CHMe ₂), 1.21 (d, 6H, CHMe ₂)	DMSO-d ⁶
20		222 - 225 dec	12.07 (s, 1H, NHCO), 11.03 (s, 1H, NH), 7.3-6.80 (m, 5H, indole+H4thiaz), 3.77 (s, 2H, COCH ₂), 3.06 (m, 1H, CHMe ₂), 1.22 (d, 6H, CHMe ₂)	DMSO-d ⁶
25		172 - 173	12.25 (s, 1H, NHCO), 8.02-7.4 (m, 4H, Ar), 7.15 (s, 1H, H4thiaz), 4.0 (s, 2H, COCH ₂), 3.07 (m, 1H, CHMe ₂), 1.22 (d, 6H, CHMe ₂)	DMSO-d ⁶
30		203 - 204	12.05 (s, 1H, NHCO), 10.77 (s, 1H, NH), 7.22-6.70 (m, 5H, indole+H4thiaz), 3.75 (s, 2H, COCH ₂), 3.72 (s, 3H, OMe), 3.07 (m, 1H, CHMe ₂), 1.22 (d, 6H, CHMe ₂)	DMSO-d ⁶
35		163 - 164	12.89 (s, 1H, NHCO), 10.75 (s, 1H, NH), 7.12-6.97 (m, 5H, indole+H4thiaz), 3.10 (m, 1H, CHMe ₂), 3.01 (t, 2H, CH ₂ CH ₂ CO), 2.78 (t, 2H, CH ₂ CH ₂ CO), 1.25 (d, 6H, CHMe ₂)	DMSO-d ⁶
40		186 - 187	12.7 (s broad, 1H, NHCO), 8.18 (d, 1H, J=7.8, Ar), 7.71 (d, 1H, J=7.8, Ar), 7.24 (s, 1H, H4thiaz), 3.15 (m, 1H, CHMe ₂), 1.27 (d, 6H, CHMe ₂)	DMSO-d ⁶
45			12.4 (s broad, 1H, NHCO), 8.05-7.51 (m, 5H, Ph), 7.23 (s, 1H, H4thiaz), 3.13 (m, 1H, CHMe ₂), 1.28 (d, 6H, CHMe ₂)	DMSO-d ⁶
50				
55				

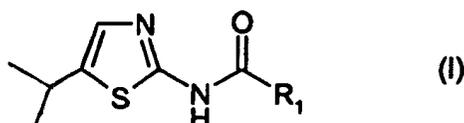
	<p>12.7 (s broad, 1H, <u>NHCO</u>), 7.54-6.82 (m, 3H, H4thiaz+furane), 3.10 (m, 1H, <u>CHMe₂</u>), 1.26 (d, 6H, <u>CHMe₂</u>),</p>	DMSO-d ⁶
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5

10 **Claims**

1. The use of a compound which is a 2-amino-1,3-thiazole derivative of formula (I)

15



20

wherein

R₁ is an optionally substituted group selected from:

25

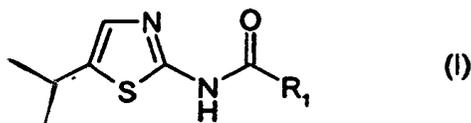
- i) aryl;
- ii) arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

or a pharmaceutically acceptable salt thereof; in the manufacture of a medicament for treating cell proliferative disorders associated with an altered cell cycle dependent kinase activity.

30

2. Use according to claim 1 wherein the said cell proliferative disorder is selected from the group consisting of cancer, Alzheimer's disease, viral infections, auto-immune diseases or neurodegenerative disorders.
3. Use according to claim 2 wherein the cancer is selected from the group consisting of carcinoma, squamous cell carcinoma, hematopoietic tumors of myeloid or lymphoid lineage, tumors of mesenchymal origin, tumors of the central and peripheral nervous system, melanoma, seminoma, teratocarcinoma, osteosarcoma, xenoderoma pigmentosum, keratoanthoma, thyroid follicular cancer and Kaposi's sarcoma.
4. Use according to claim 1 wherein the cell proliferative disorder is selected from the group consisting of benign prostate hyperplasia, familial adenomatosis polyposis, neuro-fibromatosis, psoriasis, vascular smooth cell proliferation associated with atherosclerosis, pulmonary fibrosis, arthritis glomerulonephritis and post-surgical stenosis and restenosis.
5. Use according to any one of the preceding claims wherein the medicament enables tumor angiogenesis and metastasis inhibition.
6. A compound which is a 2-amino-1,3-thiazole derivative of formula (I)

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55

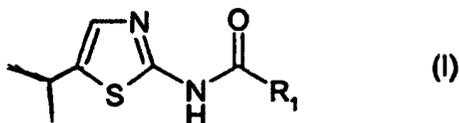
wherein

R₁ is an optionally substituted group selected from:

- i) aryl;
- ii) arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

or a pharmaceutically acceptable salt thereof; for use as a medicament.

7. A compound which is a 2-amino-1,3-thiazole derivative of formula (I)



wherein

R₁ is an optionally substituted group selected from:

- i) aryl;
- ii) arylalkyl with from 1 to 8 carbon atoms within the straight or branched alkyl chain;

or a pharmaceutically acceptable salt thereof.

8. A compound of formula (I), according to claim 7, wherein R is isopropyl and R₁ is an optionally substituted group selected from aryl or arylalkyl with from 1 to 4 carbon atoms within the alkyl chain.
9. A compound of formula (I) according to any one of the preceding claims, whenever appropriate in the form of a pharmaceutically acceptable salt, selected from the group consisting of:

1. 2-N-[2-(3-pyridyl)-acetyl-amino]-5-isopropyl-thiazole;
2. N-(5-isopropyl-2,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamide;
3. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide;
4. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide;
5. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamide;
6. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dihydroxyphenyl)acetamide;
7. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxy-3-methoxyphenyl)acetamide;
8. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide;
9. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide;
10. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide;
11. N-(5-isopropyl-thiazol-2-yl)-4-sulfamoylbenzamide;
12. 2-[3-(3-chloropropoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;
13. 2-[3-(2-chloroethoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;
14. 2-(4-aminophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;
15. 4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;
16. 2-(2-amino-1,3-thiazol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;
17. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-[3-(4-morpholinyl)propoxy]phenyl]acetamide;
18. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-[2-(4-morpholinyl)ethoxy]phenyl]acetamide;
19. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-[3-(1-pyrrolidinyl)propoxy]phenyl]acetamide;
20. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-[3-(4-methyl-1-piperazinyl)propoxy]phenyl]acetamide;
21. 2-[3-[2-(dimethylamino)ethoxy]phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;
22. 2-[3-[3-(dimethylamino)propoxy]phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;
23. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(2-methoxyethoxy)-phenyl]acetamide;
24. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methyl-1-piperazinyl)benzamide;
25. 2-anilino-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide;
26. (R)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide;
27. (S)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide;
28. N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;
29. 2,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide;

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- 30.3,5-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
31.3,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
32.2,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
33.2,3-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
5 34.3-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
35.2-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide.
36.4-iodio-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
37.3-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
38.4-chloro-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
10 39.5-bromo-2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
40.3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
41.2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
42.4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
43.3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
15 44.2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
45.4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
46.2,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
47.3,4-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
48.2,3,4,5,6-pentafluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
20 49.N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-3-nitrobenzamide
50.N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-nitrobenzamide
51.N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-nitrobenzamide
52.N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-nitrobenzamide
53.N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-2-nitrobenzamide
25 54.N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-2-nitrobenzamide
55.N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxy-3-nitrobenzamide
56.N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-nitrobenzamide
57.N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dinitrobenzamide
58.5-[[[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]-2-nitrophenyl octanoate
30 59.N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide
60.N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide
61.N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide
62. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)-3-nitrobenzamide
63. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide
35 64. 6-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide
65. 4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide
66. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamide
67. 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide
68. 2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-5-nitrobenzamide
40 69. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamide
70. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamide
71. N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitro-4-(trifluoromethyl)benzamide
72. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-bis(trifluoromethyl)benzamide
73. N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-bis(trifluoromethyl)benzamide
45 74. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)benzamide
75. N-(5-isopropyl-2,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide
76. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide
77. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide
78. 5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide
50 79. 2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethyl)benzamide
80. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluoromethyl)benzamide
81. methyl 4-[[[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]benzoate
82. methyl 2-[[[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]benzoate
83. 4-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
55 84. 3-cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
85. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methylbenzamide
86. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methylbenzamide
87. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide

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88. N-(5-isopropyl-1,3-thiazol-2-yl)-4-vinylbenzamide
89. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(2-phenylethynyl)benzamide
90. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-methylbenzamide
5 91. 2-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
92. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenethylbenzamide
93. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylbenzamide
94. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbenzamide
95. 4-(tert-butyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
10 96. N-(5-isopropyl-1,3-thiazol-2-yl)-4-isopropylbenzamide
97. N-(5-isopropyl-1,3-thiazol-2-yl)-4-pentylbenzamide
98. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamide
99. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethylbenzamide
100. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethylbenzamide
15 101. 4-acetyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
102. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)benzamide
103. 5-(aminosulfonyl)-2,4-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
104. 5-(aminosulfonyl)-4-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
105. 3-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide
20 106. 3-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide
107. 5-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide
108. N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamide
109. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxybenzamide
110. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamide
25 111. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4-dimethoxybenzamide
112. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethoxybenzamide
113. N-(5-isopropyl-1,3-thiazol-2-yl)-2,4-dimethoxybenzamide
114. N-(5-isopropyl-1,3-thiazol-2-yl)-2,3-dimethoxybenzamide
115. N-(5-isopropyl-1,3-thiazol-2-yl)-3-phenoxybenzamide
30 116. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxybenzamide
117. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenoxybenzamide
118. 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
119. 4-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
120. N-(5-isopropyl-1,3-thiazol-2-yl)-3,4,5-trimethoxybenzamide
121. 3,4-diethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
35 122. 3,4,5-triethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
123. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-(methoxymethoxy)benzamide
124. 4-butoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
125. N-(5-isopropyl-1,3-thiazol-2-yl)-4-propoxybenzamide
126. 4-isopropoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
40 127. N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-benzodioxole-5-carboxamide
128. 4-(benzyloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
129. 4-(2-cyclohexen-1-yloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
130. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluoromethoxy)benzamide
131. 4-(difluoromethoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
45 132. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)benzamide
133. 2-[(4-chlorophenyl)sulfonyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
134. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[(4-nitrophenyl)sulfonyl]benzamide
135. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(4-methylphenyl)sulfonyl]-3-nitrobenzamide
136. N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(trifluoromethyl)sulfonyl]benzamide
50 137. N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-4-(methylsulfonyl)benzamide
138. 2-[(2-cyanophenyl)sulfonyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
139. N-1~,N~1~-diethyl-3,6-difluoro-N~2~--(5-isopropyl-1,3-thiazol-2-yl)phthalamide
140. 4-formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
141. 2-formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
55 142. 4-[[2,5-dimethoxyanilino]carbonyl]amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
143. 4-(hydroxymethyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
144. 4-[[5-isopropyl-1,3-thiazol-2-yl]amino]carbonyl-2-nitrobenzyl acetate
145. 4-[[5-isopropyl-1,3-thiazol-2-yl]amino]carbonyl-2-nitrobenzyl 4-(acetylamino)-3-iodobenzoate

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146. 4-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
147. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-phenylacetyl)amino]benzamide
148. 4-(acetylamino)-3-iodo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
149. 4-amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
5 150. 4-(dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
151. 3-(dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
152. 2-(methylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
153. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluoromethyl)anilino]benzamide
154. 3-[[[5-bromo-1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)methyl]amino]-N-(5-isopropyl-1,3-thiazol-2-yl)ben-
10 zamide
155. N-(S-isopropyl-1,3-thiazol-2-yl)-4-(1H-pyrrol-1-yl)benzamide
156. 2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamide
157. 2-(4-bromophenyl)-6-(4-iodophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamide
158. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluoromethyl)anilino]nicotinamide
15 159. 2,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide
160. 5,6-dichloro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide
161. 2-chloro-N-(5-isopropyl-1,3-thiazol-2-yl)-6-methylnicotinamide
162. 2,6-dichloro-5-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamide
163. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenoxy nicotinamide
20 164. N-(5-isopropyl-1,3-thiazol-2-yl)-6-(2,2,2-trifluoroethoxy)nicotinamide
165. N-(5-isopropyl-1,3-thiazol-2-yl)-2,6-dimethoxynicotinamide
166. N-(5-isopropyl-1,3-thiazol-2-yl)-2-quinoxalinecarboxamide
167. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-pyrazinecarboxamide
168. N-(5-isopropyl-1,3-thiazol-2-yl)-8-quinolinecarboxamide
25 169. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-4-quinolinecarboxamide
170. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1-phenyl-1H-pyrazole-4-carboxamide
171. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-1H-pyrazole-3-carboxamide
172. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-pyrazole-4-carboxamide
173. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-phenyl-2H-1,2,3-triazole-4-carboxamide
30 174. 2-[(2,1,3-benzoxadiazol-5-yloxy)methyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-car-
boxamide
175. N-(5-isopropyl-1,3-thiazol-2-yl)-9H-fluorene-1-carboxamide
176. N-(5-isopropyl-1,3-thiazol-2-yl)-7-methoxy-1-benzofuran-2-carboxamide
177. N-(5-isopropyl-1,3-thiazol-2-yl)-1-[(4-methylphenyl)sulfonyl]-1H-pyrrole-3-carboxamide
35 178. 2-ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide
179. 4-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamide
180. N-(5-isopropyl-1,3-thiazol-2-yl)-2-naphthamide
181. N-(5-isopropyl-1,3-thiazol-2-yl)-9,10-dioxo-9,10-dihydro-2-anthracenecarboxamide
182. N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-4-carboxamide
40 183. N-(5-isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluorene-1-carboxamide
184. N-(5-isopropyl-1,3-thiazol-2-yl)-8-oxo-5,6,7,8-tetrahydro-2-naphthalenecarboxamide
185. N-(5-isopropyl-1,3-thiazol-2-yl)-1,3-dioxo-1,3-dihydro-2-benzofuran-5-carboxamide
186. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide
187. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-4-carboxamide
45 188. N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-2-phenyl-1H-indole-5-carboxamide
189. 2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-5-carboxamide
190. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-6-carboxamide
191. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methoxy-1H-indole-2-carboxamide
192. 1-allyl-2-butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1H-indole-5-carboxamide
50 193. N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indole-2-carboxamide
194. 1-benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-1H-indole-5-carboxamide
195. N-(5-isopropyl-1,3-thiazol-2-yl)-1H-1,2,3-benzotriazole-5-carboxamide
196. N-(5-isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-isoxazolecarboxamide
197. N-(5-isopropyl-1,3-thiazol-2-yl)-3-thiophenecarboxamide
55 198. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-thiophenecarboxamide
199. N-(5-isopropyl-1,3-thiazol-2-yl)-5-methyl-2-thiophenecarboxamide
200. 5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-thiophenecarboxamide
201. N-(5-isopropyl-1,3-thiazol-2-yl)-3-[(2,3,3-trichloroacryloyl)amino]-2-thiophenecarboxamide

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202. 5-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide
203. N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide
204. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-nitrophenyl)-2-furamide 205. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(2-nitrophenyl)-2-furamide
5 206. 5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide
207. N-(5-isopropyl-1,3-thiazol-2-yl)-5-[3-(trifluoromethyl)phenyl]-2-furamide
208. 5-(4-chloro-2-nitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide
209. N-(5-isopropyl-1,3-thiazol-2-yl)-5-(4-methyl-2-nitrophenyl)-2-furamide
210. 5-[2-chloro-5-(trifluoromethyl)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamide
10 211. tert-butyl (1R)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethylcarbamate
212. (1R)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate
213. (1S)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate
214. (R,S)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
215. (R)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
15 216. (S)-2-fluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
217. 2-(acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
218. (R,S)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
219. (R)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
220. (S)-2-(methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
20 221. 3,3,3-trifluoro-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-2-phenylpropanamide
222. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(1-naphthyl)acetamide
223. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-naphthyl)acetamide
224. 2-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
225. 2-(1,3-benzodioxol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
25 226. 2-(2,4-dinitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
227. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methyl-1H-indol-3-yl)acetamide
228. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(1-methyl-1H-indol-3-yl)acetamide
229. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-methoxy-1H-indol-3-yl)acetamide
230. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(5-benzyloxy-1H-indol-3-yl)acetamide
30 231. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxy-2-methyl-1H-indol-3-yl)acetamide
232. 2-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamide
233. 2-(5-bromo-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
234. 2-(5-fluoro-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
235. 2-[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
35 236. 3-(1H-indol-3-yl)-N-(5-isopropyl-2,3-thiazol-2-yl)propanamide
237. 4-(1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide
238. N-(5-isopropyl-1,3-thiazol-2-yl)-3-(2-thienyl)propanamide
239. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-thienyl)acetamide
240. N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-(2-thienyl)acetamide
40 241. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-thienyl)acetamide
242. 2-(5-chloro-1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
243. 2-(1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
244. 2-[2-(formylamino)-1,3-thiazol-4-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide
245. 2-[2-[(2-chloroacetyl)amino]-1,3-thiazol-4-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamide
45 246. 2-chloro-N-(4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}-1,3-thiazol-2-yl)acetamide
247. ethyl 2-({[2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(1H-pyrazol-3-yl)ethylidene]amino}oxy)acetate
248. 2-(2-furyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamide
249. 2-(5-bromo-3-pyridinyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
50 250. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(7-methoxy-2-oxo-2H-chromen-4-yl)acetamide
251. N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxo-4-(4-methyl-phenyl)butanamide
252. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-nitrophenyl)butanamide
253. N-(5-isopropyl-1,3-thiazol-2-yl)-4-phenylbutanamide
254. benzyl 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-oxobutylcarbamate
55 255. methyl 5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate
256. 4-(1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamide
257. N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methoxy-1-naphthyl)-4-oxobutanamide
258. 3-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide

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259. 3-(4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide
 260. 3-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamide
 261. N-(5-isopropyl-1,3-thiazol-2-yl)-5-oxo-5-phenylpentanamide
 262. 2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethyl acetate
 5 263. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[4-(1-oxo-1,3-dihydro-2H-isoindol-2-yl)phenyl]propanamide
 264. 1-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide
 265. 1-phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)cyclopentanecarboxamide
 266. 2-(3-bromo-4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 267. 2-(2-nitro-4-trifluoromethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 10 268. 5-cyclohexyl 1-(4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}benzyl) (2S)-2-[(tert-butoxycarbonyl)amino]pentanedioate
 269. 2-(5,6-dimethyl-1H-benzimidazol-1-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 270. 2-[5-(4-chlorophenyl)-2H-1,2,3,4-tetraazol-2-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 271. N-(5-isopropyl-1,3-thiazol-2-yl)-2-[5-(1-pyrrolidinyl)-2H-1,2,3,4-tetraazol-2-yl]acetamide
 15 272. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methyl-1-benzothiophen-2-yl)acetamide
 273. N-{4-bromo-6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexyl}benzamide
 274. benzyl 6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexylcarbamate
 275. 4-(2,4-dimethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide
 276. 4-(2-benzyloxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide 277. 4-(thiophen-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamide
 20 278. benzyl 2-[(benzyloxy)carbonyl]amino-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate
 279. 4-(1H-indol-3-yl)-N-{3-[(5-isopropyl-1,3-thiazol-2-yl)amino]-3-oxopropyl}butanamide
 280. 4-[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl}phenyl 4-chlorobenzenesulfonate
 281. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[(2-methoxyanilino)carbonyl]amino}benzamide
 25 282. 4-[[2-(isopropylsulfonyl)acetyl]amino]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
 283. N-(5-isopropyl-1,3-thiazol-2-yl)-4-[[2-(phenylsulfonyl)acetyl]amino]benzamide
 284. 4-[(diethylamino)sulfonyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
 285. 2-bromo-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
 286. 3,5-difluoro-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
 30 287. 3-[(2-fluoroanilino)carbonyl]amino]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamide
 288. N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenyl-5-propyl-1H-pyrazole-4-carboxamide
 289. 3-chloro-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfonyl)-2-thiophenecarboxamide
 290. 3-iodo-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfonyl)-2-thiophenecarboxamide
 35 291. 2-[[4-(4-chlorophenyl)sulfonyl]methyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazole-5-carboxamide
 292. 5-(4-chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluoromethyl)-3-furamide
 293. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,3,4,5,6-pentafluorophenyl)acetamide
 294. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-fluorophenyl)acetamide
 40 295. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-bromophenyl)acetamide
 296. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chlorophenyl)acetamide
 297. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenyl)acetamide
 298. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-trifluoromethylphenyl)acetamide
 299. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-methoxyphenyl)acetamide
 45 300. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-dimethoxyphenyl)acetamide
 301. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-difluorophenyl)acetamide
 302. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4,5-trimethoxyphenyl)acetamide
 303. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,6-dichlorophenyl)acetamide
 304. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2-chloro-6-fluorophenyl)acetamide
 50 305. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-dimethoxyphenyl)acetamide
 306. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,5-difluorophenyl)acetamide
 307. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,5-bis-trifluoromethylphenyl)acetamide
 308. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylthiophenyl)acetamide
 309. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamide
 55 310. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-bromophenyl)acetamide
 311. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-chlorophenyl)acetamide
 312. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-fluorophenyl)acetamide
 313. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)acetamide

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314. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-trifluoromethylphenyl)acetamide
 315. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-methylphenyl)acetamide
 316. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-dimethylaminophenyl)acetamide
 5 317. 2-[1,1'-biphenyl]-4-yl-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 318. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-trifluoromethylphenyl)acetamide
 319. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-bromophenyl)acetamide
 320. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-chlorophenyl)acetamide
 321. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-nitrophenyl)acetamide
 10 322. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamide
 323. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dinitrophenyl)acetamide
 324. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-dichlorophenyl)acetamide
 325. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(2,4-difluorophenyl)acetamide
 326. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-benzyloxy-3-methoxyphenyl)acetamide
 327. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dichlorophenyl)acetamide
 15 328. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-difluorophenyl)acetamide
 329. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(3,4-dimethoxyphenyl)acetamide
 330. 2-(2,3-dihydro-1H-inden-5-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamide
 331. N-(5-isopropyl-1,3-thiazol-2-yl)-1-phenylcyclopropanecarboxamide
 332. 2-cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
 20 333. 2-cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamide
 334. N-(5-isopropyl-1,3-thiazol-2-yl)-2,2-diphenylacetamide
 335. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)propanamide
 336. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamide
 337. N-(5-isopropyl-1,3-thiazol-2-yl)-2-(4-isobutylphenyl)propanamide
 25 338. N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxo-2-phenylacetamide
 339. N-(5-isopropyl-1,3-thiazol-2-yl)-3-methyl-2-phenylpentanamide
 340. N-(5-isopropyl-1,3-thiazol-2-yl)bicyclo[4.2.0]octa-1,3,5-triene-7-carboxamide
 341. N-(5-isopropyl-1,3-thiazol-2-yl)-3-oxo-1-indanecarboxamide
 342. N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylbutanamide
 30 343. tert-butyl (1S)-1-(1H-indol-3-ylmethyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate
 344. tert-butyl (1S)-1-(4-benzyloxybenzyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate

and the pharmaceutically acceptable salts thereof.

- 35 **10.** A process for producing a compound of formula (I), as defined in claim 7, which process comprises reacting a compound of formula (II)



45 with a compound of formula (III)



50 wherein R_1 are as defined in claim 7 and X is hydroxy or a suitable leaving group;
 and, if desired, converting a 2-amino-1,3-thiazole derivative of formula (I) into another such derivative of formula (I), and/or into a salt thereof.

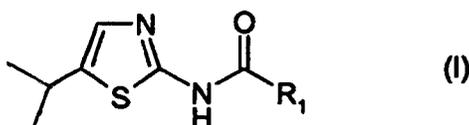
- 11.** A process according to claim 10 wherein X is hydroxy, bromine or chlorine.
- 55 **12.** A pharmaceutical composition comprising one or more pharmaceutically acceptable carriers and/or diluents and, as the active principle, an effective amount of a compound of formula (I) as defined in claim 7.

13. A compound of formula (I), according to claim 7, wherein R₁ is optionally substituted by one or more groups selected from: halogen, nitro, oxo groups (=O), carboxy, cyano, alkyl, perfluorinated alkyl, alkenyl, alkynyl, cycloalkyl, aryl, heterocycyl; amino groups and derivatives thereof such as alkylamino, alkoxy-carbonylalkylamino, dialkylamino, arylamino, diarylamino or arylureido; carbonylamino groups and derivatives thereof such as hydrogenocarbonylamino (HCONH-), alkylcarbonylamino, arylcarbonylamino, alkoxy-carbonylamino; oxygen-substituted oximes such as alkoxy-carbonylalkoxyimino or alkoxyimino; hydroxy groups and derivatives thereof such as alkoxy, aryloxy, alkylcarbonyloxy, arylcarbonyloxy, cycloalkenyloxy; carbonyl groups and derivatives thereof such as alkylcarbonyl, arylcarbonyl, alkoxy-carbonyl, aryloxy-carbonyl, cycloalkyloxy-carbonyl, aminocarbonyl, alkylaminocarbonyl, dialkylaminocarbonyl; sulfurated derivatives such as alkylthio, arylthio, alkylsulphonyl, arylsulphonyl, alkylsulphinyl, arylsulphinyl, arylsulphonyloxy, aminosulfonyl, alkylaminosulphonyl or dialkylaminosulphonyl.

14. A compound of formula (I), according to claim 13, wherein R₁ is optionally substituted by one or more groups selected from: halogen, nitro, oxo groups (=O), carboxy, alkyl, perfluorinated alkyl, cycloalkyl, aryl, heterocycyl; amino groups; carbonylamino groups such as alkylcarbonylamino, arylcarbonylamino or alkoxy-carbonylamino; hydroxy groups and derivatives thereof such as alkoxy, aryloxy or alkylcarbonyloxy; carbonyl groups and derivatives thereof such as alkylcarbonyl or arylcarbonyl.

Patentansprüche

1. Die Verwendung einer Verbindung, welche ein 2-Amino-1,3-thiazol-Derivat ist mit der Formel (I)



worin

R₁ eine wahlweise substituierte Gruppe ist gewählt aus:

- i) Aryl;
- ii) Arylalkyl mit 1 bis 8 Kohlenstoffatomen innerhalb der geradkettigen oder verzweigten Alkylkette;

oder ein pharmazeutisch verträgliches Salz davon; in der Herstellung eines Medikamentes für die Behandlung von Zellproliferationskrankheiten, die in Verbindung stehen mit einer veränderten Zell-Zyklus-abhängigen Kinaseaktivität.

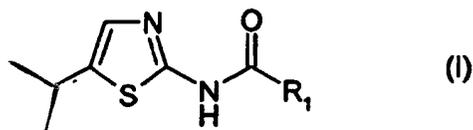
2. Verwendung gemäß Anspruch 1, worin die Zellproliferationskrankheit gewählt ist aus der Gruppe bestehend aus Krebs, Alzheimer-Krankheit, Virusinfektionen, Autoimmunkrankheiten oder neurodegenerativen Krankheiten.

3. Verwendung gemäß Anspruch 2, worin der Krebs gewählt ist aus der Gruppe bestehend aus Karzinom, Plattenepithelkarzinom, hämatopoietische Tumoren myeloischer oder lymphoider Abstammung, Tumoren mesenchymalen Ursprungs, Tumoren des zentralen und peripheren Nervensystems, Melanom, Seminom, Teratokarzinom, Osteosarkom, Xeroderma Pigmentosum, Keratoakanthom, follikulärer Schilddrüsenkrebs und Karposi-Sarkom.

4. Verwendung gemäß Anspruch 1, worin die Zellproliferationskrankheit gewählt ist aus der Gruppe bestehend aus benigner Prostatahyperplasie, familiärer adenomatöser Polypose, Neurofibromatosis, Schuppenflechte, vaskulärer glatter Muskelzellproliferation verbunden mit Atherosklerose, Lungenfibrose, Arthritis-Glomerulonephritis und postoperativer Stenose und Restenose.

5. Verwendung gemäß irgendeinem der vorhergehenden Ansprüche, worin das Medikament Tumorangiogenese- und Metastasehemmung ermöglicht.

6. Eine Verbindung, welche ein 2-Amino-1,3-thiazolderivat ist mit der Formel (I)



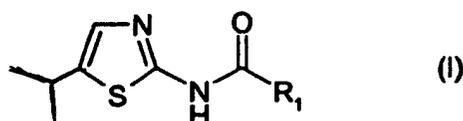
worin

10 R_1 eine wahlweise substituierte Gruppe ist gewählt aus:

- i) Aryl;
- ii) Arylalkyl mit 1 bis 8 Kohlenstoffatomen innerhalb der geraden oder verzweigten Alkylkette;

15 oder ein pharmazeutisch verträgliches Salz davon; für die Verwendung als ein Medikament.

7. Eine Verbindung, welche ein 2-Amino-1,3-thiazolderivat ist mit der Formel (I)



25 worin

R_1 eine wahlweise substituierte Gruppe ist gewählt aus:

- i) Aryl;
- ii) Arylalkyl mit 1 bis 8 Kohlenstoffatomen innerhalb der geraden oder verzweigten Alkylkette;

oder ein pharmazeutisch verträgliches Salz davon.

8. Eine Verbindung von Formel (I) gemäß Anspruch 7, worin (R Isopropyl ist und) R_1 eine wahlweise substituierte Gruppe ist gewählt aus Aryl oder Arylalkyl mit 1 bis 4 Kohlenstoffatomen innerhalb der Alkylkette.

9. Eine Verbindung von Formel (I) gemäß einem beliebigen der vorhergehenden Ansprüche, wann immer geeignet, in der Form eines pharmazeutisch verträglichen Salzes, gewählt aus der Gruppe bestehend aus:

- 40
1. 2-N-[2-(3-Pyridyl)-acetylamino]-5-isopropylthiazol;
 2. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3-hydroxyphenyl)acetamid;
 3. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl)acetamid;
 4. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3-chlorphenyl)acetamid;
 5. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxyphenyl)acetamid;
 - 45 6. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3,4-dihydroxyphenyl) acetamid;
 7. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-hydroxy-3-methoxyphenyl) acetamid;
 8. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl)acetamid;
 9. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-chlorphenyl)acetamid;
 10. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-phenylacetamid;
 - 50 11. N-(5-Isopropylthiazol-2-yl)-4-sulfamoylbenzamid;
 12. 2-[3-(3-Chlorpropoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
 13. 2-[3-(2-Chlorethoxy)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl) acetamid ;
 14. 2-(4-Aminophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
 15. 4-Amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
 - 55 16. 2-(2-Amino-1,3-thiazol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
 17. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-[3-[3-(4-morpholinyl) propoxy]phenyl]acetamid;
 18. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-[3-[2-(4-morpholinyl) ethoxy]phenyl]acetamid;
 19. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-[3-[3-(1-pirrolidinyl) propoxy]phenyl]acetamid;

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20. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-[3-[3-(4-methyl-1-piperazinyl)propoxy]phenyl]acetamid;
21. 2-[3-[2-(Dimethylamino)ethoxy]phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
22. 2-[3-[3-(Dimethylamino)propoxy]phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
5 23. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-[3-(2-methoxyethoxy)-phenyl]acetamid;
24. 3-Chlor-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(4-methyl-1-piperazinyl)benzamid;
25. 2-Anilino-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
26. (R)-N-(5-Isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamid;
27. (S)-N-(5-Isopropyl-1,3-thiazol-2-yl)-2-phenylpropanamid;
28. N-(5-Isopropyl-1,3-thiazol-2-yl)benzamid;
10 29. 2,5-Dichlor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
30. 3,5-Dichlor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
31. 3,4-Dichlor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
32. 2,4-Dichlor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
33. 2,3-Dichlor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
15 34. 3-Iod-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
35. 2-Iod-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
36. 4-Iod-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
37. 3-Brom-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
38. 4-Chlor-2-fluor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
20 39. 5-Brom-2-chlor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
40. 3-Chlor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
41. 2-Chlor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
42. 4-Chlor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
43. 3-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
25 44. 2-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
45. 4-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
46. 2,4-Difluor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
47. 3,4-Difluor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
48. 2,3,4,5,6-Pentafluor-N-(5-isopropyl-1,3-thiazol-2-yl) benzamid;
30 49. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-methyl-3-nitrobenzamid;
50. N-(5-Isopropyl-1,3-thiazol-2-yl)-5-methyl-2-nitrobenzamid;
51. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-methyl-2-nitrobenzamid;
52. N-(5-Isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-nitrobenzamid;
53. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-methoxy-2-nitrobenzamid;
35 54. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-methoxy-2-nitrobenzamid;
55. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-methoxy-3-nitrobenzamid;
56. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-nitrobenzamid;
57. N-(5-Isopropyl-1,3-thiazol-2-yl)-3,5-dinitrobenzamid;
58. 5-[[5-Isopropyl-1,3-thiazol-2-yl]amino]carbonyl]-2-nitrophenyloctanoat;
40 59. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamid;
60. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamid;
61. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamid;
62. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl)-3-nitrobenzamid;
63. 4-Chlor-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamid;
45 64. 6-Chlor-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamid;
65. 4-Chlor-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamid;
66. 2-Chlor-N-(5-isopropyl-1,3-thiazol-2-yl)-4-nitrobenzamid;
67. 5-Chlor-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamid;
68. 2-Brom-N-(5-isopropyl-1,3-thiazol-2-yl)-5-nitrobenzamid;
50 69. 4-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-3-nitrobenzamid;
70. 4-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-2-nitrobenzamid;
71. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-nitro-4-(trifluormethyl) benzamid;
72. N-(5-Isopropyl-1,3-thiazol-2-yl)-3,5-bis(trifluormethyl) benzamid;
73. N-(5-Isopropyl-1,3-thiazol-2-yl)-2,6-bis(trifluormethyl) benzamid;
55 74. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(trifluormethyl)benzamid;
75. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-(trifluormethyl)benzamid;
76. 3-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluormethyl) benzamid;
77. 2-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluormethyl) benzamid;

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78. 5-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluormethyl) benzamid;
79. 2-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-4-(trifluormethyl) benzamid;
80. 4-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-3-(trifluormethyl) benzamid;
81. Methyl-4-[[5-isopropyl-1,3-thiazol-2-yl]amino]carbonyl} benzoat;
5 82. Methyl-2-[[5-isopropyl-1,3-thiazol-2-yl]amino]carbonyl} benzoat;
83. 4-Cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
84. 3-Cyano-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
85. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-methylbenzamid;
86. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-methylbenzamid;
10 87. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-methylbenzamid;
88. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-vinylbenzamid;
89. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-(2-phenylethynyl) benzamid;
90. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-methylbenzamid;
91. 2-Benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
15 92. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-phenethylbenzamid;
93. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-phenylbenzamid;
94. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-phenylbenzamid;
95. 4-(tert-Butyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
96. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-isopropylbenzamid;
20 97. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-pentylbenzamid;
98. 3-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methylbenzamid;
99. N-(5-Isopropyl-1,3-thiazol-2-yl)-3,4-dimethylbenzamid;
100. N-(5-Isopropyl-1,3-thiazol-2-yl)-3,5-dimethylbenzamid;
101. 4-Acetyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
25 102. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl) benzamid;
103. 5-(Aminosulfonyl)-2,4-dichlor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
104. 5-(Aminosulfonyl)-4-chlor-N-(5-isopropyl-1,3-thiazol-2-yl) benzamid;
105. 3-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamid;
106. 3-Chlor-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamid;
30 107. 5-Chlor-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamid;
108. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-methoxybenzamid;
109. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-methoxybenzamid;
110. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-methoxybenzamid;
111. N-(5-Isopropyl-1,3-thiazol-2-yl)-3,4-dimethoxybenzamid;
35 112. N-(5-Isopropyl-1,3-thiazol-2-yl)-3,5-dimethoxybenzamid;
113. N-(5-Isopropyl-1,3-thiazol-2-yl)-2,4-dimethoxybenzamid;
114. N-(5-Isopropyl-1,3-thiazol-2-yl)-2,3-dimethoxybenzamid;
115. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-phenoxybenzamid;
116. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-phenoxybenzamid;
40 117. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-phenoxybenzamid;
118. 2-Ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
119. 4-Ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
120. N-(5-Isopropyl-1,3-thiazol-2-yl)-3,4,5-trimethoxybenzamid;
121. 3,4-Diethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
45 122. 3,4,5-Triethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
123. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-methoxy-4-(methoxymethoxy)benzamid;
124. 4-Butoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
125. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-propoxybenzamid;
126. 4-Isopropoxy-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
50 127. N-(5-Isopropyl-1,3-thiazol-2-yl)-1,3-benzodioxol-5-carboxamid;
128. 4-(Benzyloxy)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
129. 4-(2-Cyclohexen-1-yloxy)-N-(5-isopropyl-1,3-thiazol-2-yl) benzamid;
130. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-(trifluormethoxy) benzamid;
131. 4-(Difluormethoxy)-N-(5-isopropyl-1,3-thiazol-2-yl) benzamid;
55 132. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-(methylsulfonyl) benzamid;
133. 2-[(4-Chlorphenyl)sulfinyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
134. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-[(4-nitrophenyl) sulfinyl]benzamid;
135. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-[(4-methylphenyl) sulfonyl]-3-nitrobenzamid;

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136. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-[(trifluormethyl) sulfanyl]benzamid;
 137. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-methoxy-4-(methylsulfanyl)benzamid;
 138. 2-[(2-Cyanophenyl)sulfanyl]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
 139. N~1~,N~1~-Diethyl-3,6-difluor-N~2~--(5-isopropyl-1,3-thiazol-2-yl)phthalamid;
 140. 4-Formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
 141. 2-Formyl-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
 142. 4-[(2,5-Dimethoxyanilino)carbonyl]amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
 143. 4-(Hydroxymethyl)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
 144. 4-[(5-Isopropyl-1,3-thiazol-2-yl)amino]carbonyl-2-nitrobenzylacetat;
 145. 4-[(5-Isopropyl-1,3-thiazol-2-yl)amino]carbonyl-2-nitrobenzyl-4-(acetylamino)-3-iodobenzoat;
 146. 4-(Acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
 147. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-[(2-phenylacetyl)amino] benzamid;
 148. 4-(Acetylamino)-3-iod-N-(5-isopropyl-1,3-thiazol-2-yl) benzamid;
 149. 4-Amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
 150. 4-(Dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
 151. 3-(Dimethylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
 152. 2-(Methylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
 153. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluormethyl) anilino]benzamid;
 154. 3-[(5-Brom-1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)methyl] amino-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
 155. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-(1H-pyrrol-1-yl) benzamid;
 156. 2,6-Dichlor-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamid;
 157. 2-(4-Bromphenyl)-6-(4-iodphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)isonicotinamid;
 158. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-[3-(trifluormethyl) anilino]nicotinamid;
 159. 2,6-Dichlor-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamid;
 160. 5,6-Dichlor-N-(5-isopropyl-1,3-thiazol-2-yl)nicotinamid;
 161. 2-Chlor-N-(5-isopropyl-1,3-thiazol-2-yl)-6-methylnicotinamid;
 162. 2,6-Dichlor-5-fluor-N-(5-isopropyl-1,3-thiazol-2-yl) nicotinamid;
 163. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-phenoxy nicotinamid;
 164. N-(5-Isopropyl-1,3-thiazol-2-yl)-6-(2,2,2-trifluorethoxy) nicotinamid;
 165. N-(5-Isopropyl-1,3-thiazol-2-yl)-2,6-dimethoxynicotinamid;
 166. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-chinoxalincarboxamid;
 167. N-(5-Isopropyl-1,3-thiazol-2-yl)-5-methyl-2-pyrazincarboxamid;
 168. N-(5-Isopropyl-1,3-thiazol-2-yl)-8-chinolincarboxamid;
 169. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-phenyl-4-chinolincarboxamid;
 170. N-(5-Isopropyl-1,3-thiazol-2-yl)-5-methyl-1-phenyl-1Hpyrazol-4-carboxamid;
 171. N-(5-Isopropyl-1,3-thiazol-2-yl)-5-methyl-1H-pyrazol-3-carboxamid;
 172. N-(5-Isopropyl-1,3-thiazol-2-yl)-1H-pyrazol-4-carboxamid;
 173. N-(5-Isopropyl-1,3-thiazol-2-yl)-5-methyl-2-phenyl-2H-1,2,3-triazol-4-carboxamid;
 174. 2-[(2,1,3-Benzoxadiazol-5-yloxy)methyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazol-5-carboxamid;
 175. N-(5-Isopropyl-1,3-thiazol-2-yl)-9H-fluoren-1-carboxamid;
 176. N-(5-Isopropyl-1,3-thiazol-2-yl)-7-methoxy-1-benzofuran-2-carboxamid;
 177. N-(5-Isopropyl-1,3-thiazol-2-yl)-1-[(4-methylphenyl) sulfonyl]-1H-pyrrol-3-carboxamid;
 178. 2-Ethoxy-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamid;
 179. 4-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-1-naphthamid;
 180. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-naphthamid;
 181. N-(5-Isopropyl-1,3-thiazol-2-yl)-9,10-dioxo-9,10-dihydro-2-anthracencarboxamid;
 182. N-(5-Isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluoren-4-carboxamid;
 183. N-(5-Isopropyl-1,3-thiazol-2-yl)-9-oxo-9H-fluoren-1-carboxamid;
 184. N-(5-Isopropyl-1,3-thiazol-2-yl)-8-oxo-5,6,7,8-tetrahydro-2-naphthalencarboxamid;
 185. N-(5-Isopropyl-1,3-thiazol-2-yl)-1,3-dioxo-1,3-dihydro-2-benzofuran-5-carboxamid;
 186. N-(5-Isopropyl-1,3-thiazol-2-yl)-1H-indol-5-carboxamid;
 187. N-(5-Isopropyl-1,3-thiazol-2-yl)-1H-indol-4-carboxamid;
 188. N-(5-Isopropyl-1,3-thiazol-2-yl)-1-methyl-2-phenyl-1Hindol-5-carboxamid;
 189. 2-Butyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indol-5-carboxamid;
 190. N-(5-Isopropyl-1,3-thiazol-2-yl)-1H-indol-6-carboxamid;
 191. N-(5-Isopropyl-1,3-thiazol-2-yl)-5-methoxy-1H-indol-2-carboxamid;

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192. 1-Allyl-2-butyl-N-(5-isopropyl-2,3-thiazol-2-yl)-1H-indol-5-carboxamid;
 193. N-(5-Isopropyl-1,3-thiazol-2-yl)-1-methyl-1H-indol-2-carboxamid;
 194. 1-Benzyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenyl-1H-indol-5-carboxamid;
 195. N-(5-Isopropyl-1,3-thiazol-2-yl)-1H-1,2,3-benzotriazol-5-carboxamid;
 196. N-(5-Isopropyl-1,3-thiazol-2-yl)-3,5-dimethyl-4-isoxazolcarboxamid;
 197. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-thiophencarboxamid;
 198. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-methyl-2-thiophencarboxamid;
 199. N-(5-Isopropyl-1,3-thiazol-2-yl)-5-methyl-2-thiophencarboxamid;
 200. 5-Brom-N-(5-isopropyl-1,3-thiazol-2-yl)-2-thiophencarboxamid;
 201. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-[(2,3,3-trichloracryloyl)amino]-2-thiophencarboxamid;
 202. 5-Brom-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamid;
 203. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-furamid;
 204. N-(5-Isopropyl-1,3-thiazol-2-yl)-5-(4-nitrophenyl)-2-furamid;
 205. N-(5-Isopropyl-1,3-thiazol-2-yl)-5-(2-nitrophenyl)-2-furamid;
 206. 5-(4-Chlorphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamid;
 207. N-(5-Isopropyl-1,3-thiazol-2-yl)-5-[3-(trifluormethyl) phenyl]-2-furamid;
 208. 5-(4-Chlor-2-nitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamid;
 209. N-(5-Isopropyl-1,3-thiazol-2-yl)-5-(4-methyl-2-nitrophenyl)-2-furamid;
 210. 5-[2-Chlor-5-(trifluormethyl)phenyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-furamid;
 211. tert-Butyl-(1R)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethylcarbamat;
 212. (1R)-2-[(5-Isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethylacetat;
 213. (1S)-2-[(5-Isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethylacetat;
 214. (R,S)-2-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamid;
 215. (R)-2-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamid;
 216. (S)-2-Fluor-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamid;
 217. 2-(Acetylamino)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamid;
 218. (R,S)-2-(Methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamid;
 219. (R)-2-(Methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamid;
 220. (S)-2-(Methoxy)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamid;
 221. 3,3,3-Trifluor-N-(5-isopropyl-1,3-thiazol-2-yl)-2-methoxy-2-phenylpropanamid;
 222. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(1-naphthyl)acetamid;
 223. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2-naphthyl)acetamid;
 224. 2-(1H-Indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
 225. 2-(1,3-Benzodioxol-4-yl)-N-(5-isopropyl-1,3-thiazol-2-yl) acetamid;
 226. 2-(2,4-Dinitrophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl) acetamid;
 227. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2-methyl-1H-indol-3-yl) acetamid;
 228. N-(5-Isopropyl-2,3-thiazol-2-yl)-2-(1-methyl-1H-indol-3-yl) acetamid;
 229. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(5-methoxy-1H-indol-3-yl)acetamid;
 230. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(5-benzyloxy-1H-indol-3-yl)acetamid;
 231. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3-methoxy-2-methyl-1H-indol-3-yl)acetamid;
 232. 2-(1H-Indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamid;
 233. 2-(5-Brom-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl) acetamid;
 234. 2-(5-Fluor-1H-indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl) acetamid;
 235. 2-[1-(4-Chlorbenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
 236. 3-(1H-Indol-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl) propanamid;
 237. 4-(1H-Indol-3-yl)-N-(5-isopropyl-2,3-thiazol-2-yl) butanamid;
 238. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-(2-thienyl)propanamid;
 239. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2-thienyl)acetamid;
 240. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-oxo-2-(2-thienyl) acetamid;
 241. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3-thienyl)acetamid;
 242. 2-(5-Chlor-1-benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
 243. 2-(1-Benzothiophen-3-yl)-N-(5-isopropyl-1,3-thiazol-2-yl) acetamid;
 244. 2-[2-(Formylamino)-1,3-thiazol-4-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamid;
 245. 2-{2-[(2-Chloracetyl)amino]-1,3-thiazol-4-yl}-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(methoxyimino)acetamid;
 246. 2-Chlor-N-(4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}-1,3-thiazol-2-yl)acetamid;
 247. Ethyl-2-({2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-(1H-pyrazol-3-yl)ethyliden}amino)oxy)acetat;
 248. 2-(2-Furyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-oxoacetamid;
 249. 2-(5-Brom-3-pyridinyl)-N-(5-isopropyl-1,3-thiazol-2-yl) acetamid;

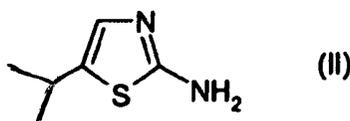
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250. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(7-methoxy-2-oxo-2Hchromen-4-yl)acetamid;
251. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-oxo-4-(4-methylphenyl) butanamid;
252. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-(4-nitrophenyl) butanamid;
253. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-phenylbutanamid;
5 254. Benzyl-4-[(5-isopropyl-1,3-thiazol-2-yl)amino]-4-oxobutylcarbamate;
255. Methyl-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate;
256. 4-(1,3-Dioxo-1,3-dihydro-2H-isoindol-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)butanamid;
257. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-(4-methoxy-1-naphthyl)-4-oxobutanamid;
258. 3-(4-Chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl) propanamid;
10 259. 3-(4-Methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl) propanamid;
260. 3-Phenyl-N-(5-isopropyl-1,3-thiazol-2-yl)propanamid;
261. N-(5-Isopropyl-1,3-thiazol-2-yl)-5-oxo-5-phenylpentanamid;
262. 2-[(5-Isopropyl-1,3-thiazol-2-yl)amino]-2-oxo-1-phenylethylacetate;
263. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-[4-(1-oxo-1,3-dihydro-2H-isoindol-2-yl)phenyl]propanamid;
15 264. 1-(4-Chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl) cyclopentancarboxamid;
265. 1-Phenyl-N-(5-isopropyl-1,3-thiazol-2-yl) cyclopentancarboxamid;
266. 2-(3-Brom-4-methoxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl) acetamid;
267. 2-(2-Nitro-4-trifluormethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
268. 5-Cyclohexyl-1-(4-{2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethyl}benzyl)-(2S)-2-[(tert-butoxycarbo-
20 nyl)amino] pentandioate;
269. 2-(5,6-Dimethyl-1H-benzimidazol-1-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
270. 2-[5-(4-Chlorophenyl)-2H-1,2,3,4-tetraazol-2-yl]-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
271. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-[5-(1-pyrrolidinyl)-2H-1,2,3,4-tetraazol-2-yl]acetamid;
272. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3-methyl-1-benzothiophen-2-yl)acetamid;
25 273. N-{4-Brom-6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexyl}benzamid;
274. Benzyl-6-[(5-isopropyl-1,3-thiazol-2-yl)amino]-6-oxohexylcarbamate;
275. 4-(2,4-Dimethylphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamid;
276. 4-(4-Benzyloxyphenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamid;
277. 4-(Thiophen-2-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)-4-oxobutanamid;
30 278. Benzyl-2-[(benzyloxy)carbonylamino]-5-[(5-isopropyl-1,3-thiazol-2-yl)amino]-5-oxopentanoate;
279. 4-(1H-Indol-3-yl)-N-{3-[(5-isopropyl-1,3-thiazol-2-yl) amino]-3-oxopropyl}butanamid;
280. 4-[(5-Isopropyl-1,3-thiazol-2-yl)amino]carbonyl}phenyl-4-chlorbenzensulfonate;
281. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-[(2-methoxyanilino) carbonylamino]benzamid;
282. 4-[[2-(Isopropylsulfonyl)acetyl]amino]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
35 283. N-(5-Isopropyl-1,3-thiazol-2-yl)-4-[[2-(phenylsulfonyl) acetyl]amino]benzamid;
284. 4-[(Diethylamino)sulfonyl]-N-(5-isopropyl-1,3-thiazol-2-yl) benzamid;
285. 2-Brom-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
286. 3,5-Difluor-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
287. 3-[(2-Fluoranilino)carbonylamino]-N-(5-isopropyl-1,3-thiazol-2-yl)benzamid;
40 288. N-(5-Isopropyl-1,3-thiazol-2-yl)-1-phenyl-5-propyl-1Hpyrazol-4-carboxamid;
289. 3-Chlor-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfonyl)-2-thiophencarboxamid;
290. 3-Iod-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-5-(methylsulfonyl)-2-thiophencarboxamid;
291. 2-[[4-Chlorophenyl)sulfonyl]methyl]-N-(5-isopropyl-1,3-thiazol-2-yl)-4-methyl-1,3-thiazol-5-carboxamid;
292. 5-(4-Chlorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-2-(trifluormethyl)-3-furamid;
45 293. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2,3,4,5,6-pentafluorophenyl)acetamid;
294. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2-fluorophenyl)acetamid;
295. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2-bromophenyl)acetamid;
296. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2-chlorophenyl)acetamid;
297. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2-nitrophenyl)acetamid;
50 298. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2-trifluormethylphenyl) acetamid;
299. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2-methoxyphenyl) acetamid;
300. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2,5-dimethoxyphenyl) acetamid;
301. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2,5-difluorophenyl) acetamid;
302. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3,4,5-trimethoxyphenyl) acetamid;
55 303. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2,6-dichlorophenyl) acetamid;
304. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2-chlor-6-fluorophenyl) acetamid;
305. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3,5-dimethoxyphenyl) acetamid;
306. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3,5-difluorophenyl) acetamid;

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307. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2,5-bis(trifluoromethyl)phenyl)acetamid;
308. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-methylthiophenyl) acetamid;
309. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-methoxyphenyl) acetamid;
5 310. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-bromphenyl)acetamid;
311. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-chlorphenyl)acetamid;
312. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-fluorphenyl)acetamid;
313. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl)acetamid;
314. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-trifluormethylphenyl) acetamid;
315. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-methylphenyl) acetamid;
10 316. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-dimethylaminophenyl) acetamid;
317. 2-[1,1'-Biphenyl]-4-yl-N-(5-isopropyl-1,3-thiazol-2-yl) acetamid;
318. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3-trifluormethylphenyl) acetamid;
319. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3-bromphenyl)acetamid;
320. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3-chlorphenyl)acetamid;
15 321. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3-nitrophenyl)acetamid;
322. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3-methoxyphenyl) acetamid;
323. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2,4-dinitrophenyl) acetamid;
324. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2,4-dichlorphenyl) acetamid;
325. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(2, 4-difluorphenyl) acetamid;
20 326. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-benzyloxy-3-methoxyphenyl)acetamid;
327. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3,4-dichlorphenyl) acetamid;
328. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3,4-difluorphenyl) acetamid;
329. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(3,4-dimethoxyphenyl) acetamid;
330. 2-(2,3-Dihydro-1H-inden-5-yl)-N-(5-isopropyl-1,3-thiazol-2-yl)acetamid;
25 331. N-(5-Isopropyl-1,3-thiazol-2-yl)-1-phenylcyclopropancarboxamid;
332. 2-Cyclopentyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamid;
333. 2-Cyclohexyl-N-(5-isopropyl-1,3-thiazol-2-yl)-2-phenylacetamid;
334. N-(5-Isopropyl-1,3-thiazol-2-yl)-2,2-diphenylacetamid;
335. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-nitrophenyl) propanamid;
30 336. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-phenyl)propanamid;
337. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-(4-isobutylphenyl) propanamid;
338. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-oxo-2-phenylacetamid;
339. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-methyl-2-phenylpentanamid;
340. N-(5-Isopropyl-1,3-thiazol-2-yl)bicyclo[4.2.0]octa-1,3,5-trien-7-carboxamid;
35 341. N-(5-Isopropyl-1,3-thiazol-2-yl)-3-oxo-1-indancarboxamid;
342. N-(5-Isopropyl-1,3-thiazol-2-yl)-2-phenyl)butanamid;
343. tert-Butyl (1S)-1-(1H-indol-3-ylmethyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate;
344. tert-Butyl (1S)-1-(4-benzyloxybenzyl)-2-[(5-isopropyl-1,3-thiazol-2-yl)amino]-2-oxoethylcarbamate und
die pharmazeutisch verträglichen Salze davon.

- 40 10. Ein Verfahren zur Herstellung einer Verbindung von Formel (I), wie in Anspruch 7 definiert, wobei das Verfahren das Reagieren einer Verbindung von Formel (II)



50 mit einer Verbindung von Formel (III)

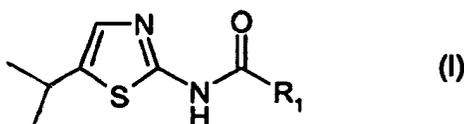


55 umfasst, worin R_1 wie in Anspruch 7 definiert ist und X Hydroxy oder eine geeignete Austrittsgruppe ist; und, falls erwünscht, Umwandeln eines 2-Amino-1,3-thiazolderivats von Formel (I) in ein anderes derartiges Derivat von Formel (I) und/oder in ein Salz davon.

11. Ein Verfahren gemäß Anspruch 10, worin X Hydroxy, Brom oder Chlor ist.
12. Eine pharmazeutische Zusammensetzung, die einen oder mehrere pharmazeutisch verträgliche Träger und/oder Verdünnungsmittel und, als den aktiven Stoff, eine wirksame Menge einer Verbindung von Formel (I), wie in Anspruch 7 definiert, umfasst.
13. Eine Verbindung von Formel (I), gemäß Anspruch 7, worin R₁ wahlweise substituiert ist durch eine oder mehrere Gruppen gewählt aus: Halogen, Nitro, Oxogruppen (=O), Carboxy, Cyano, Alkyl, perfluoriertes Alkyl, Alkenyl, Alkyl, Cycloalkyl, Aryl, Heterocycl; Aminogruppen und Derivate davon wie Alkylamino, Alkoxy-carbonylalkylamino, Dialkylamino, Arylamino, Diarylamino oder Arylureido; Carbonylaminogruppen und Derivate davon wie Hydrogenocarbonylamino (HCONH-), Alkylcarbonylamino, Arylcarbonylamino, Alkoxy-carbonylamino; Sauerstoff-substituierte Oxime wie Alkoxy-carbonylalkoxyimino oder Alkoxyimino; Hydroxygruppen und Derivate davon wie Alkoxy, Aryloxy, Alkylcarbonyloxy, Arylcarbonyloxy, Cycloalkenyloxy; Carbonylgruppen und Derivate davon wie Alkylcarbonyl, Arylcarbonyl, Alkoxy-carbonyl, Aryloxy-carbonyl, Cycloalkyloxy-carbonyl, Aminocarbonyl, Alkylaminocarbonyl, Dialkylaminocarbonyl; sulfurierte Derivate wie Alkylthio, Arylthio, Alkylsulphonyl, Arylsulphonyl, Alkylsulphinyl, Arylsulphinyl, Arylsulphonyloxy, Aminosulfonyl, Alkylaminosulphonyl oder Dialkylaminosulphonyl.
14. Eine Verbindung von Formel (I), gemäß Anspruch 13, worin R₁ wahlweise substituiert ist durch eine oder mehrere Gruppen gewählt aus: Halogen, Nitro, Oxogruppen (=O), Carboxy, Alkyl, perfluoriertes Alkyl, Cycloalkyl, Aryl, Heterocycl; Aminogruppen; Carbonylaminogruppen wie Alkylcarbonylamino, Arylcarbonylamino oder Alkoxy-carbonylamino; Hydroxygruppen und Derivate davon wie Alkoxy, Aryloxy oder Alkylcarbonyloxy; Carbonylgruppen und Derivate davon wie Alkylcarbonyl oder Arylcarbonyl. **[Streichung(en)]**

Revendications

1. Utilisation d'un composé qui est un dérivé de 2-amino-1,3-thiazole de formule (I)



dans laquelle

R¹ représente un groupe, facultativement substitué, choisi entre :

- i) un groupe aryle ;
- ii) un groupe arylalkyle ayant 1 à 8 atomes de carbone

dans la chaîne alkyle droite ou ramifiée ; ou un de ses sels pharmaceutiquement acceptables ; dans la production d'un médicament destiné au traitement de troubles de la prolifération cellulaire associés à une activité de kinase, dépendant du cycle cellulaire, modifiée.

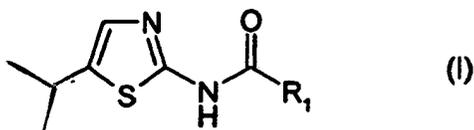
2. Utilisation suivant la revendication 1, dans laquelle ledit trouble de la prolifération cellulaire est choisi dans le groupe consistant en le cancer, la maladie d'Alzheimer, des infections virales, des maladies auto-immunes et des troubles neurodégénératifs.
3. Utilisation suivant la revendication 2, dans laquelle le cancer est choisi dans le groupe consistant en un carcinome, un carcinome à cellules squameuses, des tumeurs hématopoïétiques de la lignée myéloïde ou lymphoïde, des tumeurs d'origine mésenchymateuse, des tumeurs du système nerveux central ou périphérique, un mélanome, un séminome, un tératocarcinome, un ostéosarcome, le xeroderma pigmentosum, un kératoacanthome, un cancer folliculaire thyroïdien et le sarcome de Kaposi.
4. Utilisation suivant la revendication 1, dans laquelle le trouble de la prolifération cellulaire est choisi dans le groupe consistant en l'hyperplasie prostatique bénigne, la polypose adénomateuse familiale, la neurofibromatose, le psoriasis, une prolifération de cellules du tissu lisse vasculaire associée à l'athérosclérose, la fibrose pulmonaire,

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l'arthrose, la glomérulonéphrite et une sténose ou resténose post-chirurgicale.

5. Utilisation suivant l'une quelconque des revendications précédentes, dans laquelle le médicament permet une inhibition de l'angiogenèse et de la métastase de tumeurs.

6. Composé qui est un dérivé de 2-amino-1,3-thiazole de formule (I)



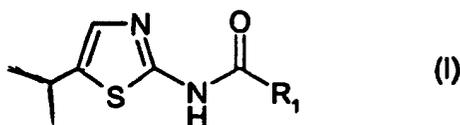
dans laquelle

R¹ représente un groupe, facultativement substitué, choisi entre :

- i) un groupe aryle ;
- ii) un groupe arylalkyle ayant 1 à 8 atomes de carbone

dans la chaîne alkyle droite ou ramifiée ; ou un de ses sels pharmaceutiquement acceptables ; destiné à être utilisé comme médicament.

7. Composé qui est un dérivé de 2-amino-1,3-thiazole de formule (I)



dans laquelle

R¹ représente un groupe, facultativement substitué, choisi entre :

- i) un groupe aryle ;
- ii) un groupe arylalkyle ayant 1 à 8 atomes de carbone

dans la chaîne alkyle droite ou ramifiée ; ou un de ses sels pharmaceutiquement acceptables.

8. Composé de formule (I), suivant la revendication 7, dans lequel R représente un groupe isopropyle et R₁ représente un groupe, facultativement substitué, choisi entre des groupes aryle ou arylalkyle ayant 1 à 4 atomes de carbone dans la chaîne alkyle.

9. Composé de formule (I) suivant l'une quelconque des revendications précédentes, chaque fois sous sa forme appropriée sous forme d'un sel pharmaceutiquement acceptable, choisi dans le groupe consistant en :

- 1. 2-N-[2-(3-pyridyl)-acétylamino]-5-isopropylthiazole ;
- 2. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3-hydroxyphényl)-acétamide ;
- 3. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3-méthoxyphényl)-acétamide ;
- 4. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3-chlorophényl)-acétamide ;
- 5. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-hydroxyphényl)-acétamide ;
- 6. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3,4-dihydroxy-phényl)acétamide ;
- 7. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-hydroxy-3-méthoxyphényl)acétamide ;
- 8. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-méthoxyphényl)-acétamide ;
- 9. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-chlorophényl)-acétamide ;

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10. N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénylacétamide ;
11. N-(5-isopropylthiazole-2-yl)-4-sulfamoylbenzamide ;
12. 2-[3-(3-chloropropoxy)phényl]-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
13. 2-[3-(2-chloroéthoxy)phényl]-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
- 5 14. 2-(4-aminophényl)-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
15. 4-amino-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
16. 2-(2-amino-1,3-thiazole-4-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
17. N-(5-isopropyl-1,3-thiazole-2-yl)-2-{3-[3-(4-morpholinyl)-propoxy]phényl}acétamide ;
18. N-(5-isopropyl-1,3-thiazole-2-yl)-2-{3-[2-(4-morpholinyl)-éthoxy]phényl}acétamide ;
- 10 19. N-(5-isopropyl-1,3-thiazole-2-yl)-2-{3-[3-(1-pyrrolidinyl)-propoxy]phényl}acétamide ;
20. N-(5-isopropyl-1,3-thiazole-2-yl)-2-{3-[3-(4-méthyl-1-pipérazinyl)propoxy]phényl}acétamide ;
21. 2-{3-[2-(diméthylamino)éthoxy]phényl}-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
22. 2-{3-[3-(diméthylamino)propoxy]phényl}-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
23. N-(5-isopropyl-1,3-thiazole-2-yl)-2-[3-(2-méthoxyéthoxy)-phényl]acétamide ;
- 15 24. 3-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)-4-(4-méthyl-1-pipérazinyl)benzamide ;
25. 2-anilino-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
26. (R)-N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénylpropanamide ;
27. (S)-N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénylpropanamide ;
28. N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
- 20 29. 2,5-dichloro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
30. 3,5-dichloro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
31. 3,4-dichloro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
32. 2,4-dichloro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
33. 2,3-dichloro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
- 25 34. 3-iodo-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
35. 2-iodo-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
36. 4-iodo-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
37. 3-bromo-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
38. 4-chloro-2-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-benzamide ;
- 30 39. 5-bromo-2-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)-benzamide ;
40. 3-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
41. 2-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
42. 4-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
43. 3-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
- 35 44. 2-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
45. 4-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
46. 2,4-difluoro-N-(5-isopropyl-2,3-thiazole-2-yl)benzamide ;
47. 3,4-difluoro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
48. 2,3,4,5,6-pentafluoro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
- 40 49. N-(5-isopropyl-1,3-thiazole-2-yl)-4-méthyl-3-nitrobenzamide ;
50. N-(5-isopropyl-2,3-thiazole-2-yl)-5-méthyl-2-nitrobenzamide ;
51. N-(5-isopropyl-1,3-thiazole-2-yl)-3-méthyl-2-nitrobenzamide ;
52. N-(5-isopropyl-1,3-thiazole-2-yl)-3,5-diméthyl-4-nitrobenzamide ;
53. N-(5-isopropyl-1,3-thiazole-2-yl)-4-méthoxy-2-nitrobenzamide ;
- 45 54. N-(5-isopropyl-1,3-thiazole-2-yl)-3-méthoxy-2-nitrobenzamide ;
55. N-(5-isopropyl-1,3-thiazole-2-yl)-4-méthoxy-3-nitrobenzamide ;
56. N-(5-isopropyl-1,3-thiazole-2-yl)-3-méthoxy-4-nitrobenzamide ;
57. N-(5-isopropyl-1,3-thiazole-2-yl)-3,5-dinitrobenzamide ;
58. octanoate de 5-[(5-isopropyl-1,3-thiazole-2-yl) amino]-carbonyl]-2-nitrophényle ;
- 50 59. N-(5-isopropyl-1,3-thiazole-2-yl)-3-nitrobenzamide ;
60. N-(5-isopropyl-1,3-thiazole-2-yl)-2-nitrobenzamide ;
61. N-(5-isopropyl-1,3-thiazole-2-yl)-4-nitrobenzamide ;
62. N-(5-isopropyl-1,3-thiazole-2-yl)-4-(méthylsulfonyl)-3-nitrobenzamide ;
63. 4-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)-3-nitrobenzamide ;
- 55 64. 6-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)-3-nitrobenzamide ;
65. 4-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)-2-nitrobenzamide ;
66. 2-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)-4-nitrobenzamide ;
67. 5-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)-2-nitrobenzamide ;

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68. 2-bromo-N-(5-isopropyl-1,3-thiazole-2-yl)-5-nitro-benzamide ;
69. 4-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-3-nitro-benzamide ;
70. 4-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-2-nitro-benzamide ;
5 71. N-(5-isopropyl-1,3-thiazole-2-yl)-2-nitro-4-(trifluorométhyl) benzamide ;
72. N-(5-isopropyl-1,3-thiazole-2-yl)-3,5-bis(trifluorométhyl)-benzamide ;
73. N-(5-isopropyl-1,3-thiazole-2-yl)-2,6-bis(trifluorométhyl)-benzamide ;
74. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(trifluorométhyl)-benzamide ;
75. N-(5-isopropyl-1,3-thiazole-2-yl)-3-(trifluorométhyl)-benzamide ;
10 76. 3-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-4-(trifluoro-méthyl)benzamide ;
77. 2-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-3-(trifluoro-méthyl)benzamide ;
78. 5-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-3-(trifluoro-méthyl)benzamide ;
79. 2-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-4-(trifluoro-méthyl)benzamide ;
80. 4-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-3-(trifluoro-méthyl)benzamide ;
15 81. 4-[[5-isopropyl-1,3-thiazole-2-yl]amino]carbonyl]benzoate de méthyle ;
82. 2-[[5-isopropyl-1,3-thiazole-2-yl]amino]carbonyl]benzoate de méthyle ;
83. 4-cyano-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
84. 3-cyano-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
85. N-(5-isopropyl-1,3-thiazole-2-yl)-3-méthylbenzamide ;
86. N-(5-isopropyl-1,3-thiazole-2-yl)-2-méthylbenzamide ;
20 87. N-(5-isopropyl-1,3-thiazole-2-yl)-4-méthylbenzamide ;
88. N-(5-isopropyl-1,3-thiazole-2-yl)-4-vinylbenzamide ;
89. N-(5-isopropyl-1,3-thiazole-2-yl)-4-(2-phényléthy-nyl) benzamide ;
90. N-(5-isopropyl-1,3-thiazole-2-yl)-3-méthoxy-4-méthylbenzamide ;
91. 2-benzyl-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
25 92. N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénéthylbenzamide ;
93. N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénylbenzamide ;
94. N-(5-isopropyl-1,3-thiazole-2-yl)-4-phénylbenzamide ;
95. 4-(tertiobutyl)-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
96. N-(5-isopropyl-1,3-thiazole-2-yl)-4-isopropylbenzamide ;
30 97. N-(5-isopropyl-1,3-thiazole-2-yl)-4-pentylbenzamide ;
98. 3-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-4-méthylbenzamide ;
99. N-(5-isopropyl-1,3-thiazole-2-yl)-3,4-diméthylbenzamide ;
100. N-(5-isopropyl-1,3-thiazole-2-yl)-3,5-diméthylbenzamide ;
101. 4-acétyl-N-(5-isopropyl-1,3-thiazole-2-yl)-benzamide ;
35 102. N-(5-isopropyl-1,3-thiazole-2-yl)-4-(méthylsulfonyl)-benzamide ;
103. 5-(aminosulfonyl)-2,4-dichloro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
104. 5-(aminosulfonyl)-4-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
105. 3-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-4-méthoxy-benzamide ;
106. 3-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)-4-méthoxy-benzamide ;
40 107. 5-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)-2-méthoxy-benzamide ;
108. N-(5-isopropyl-1,3-thiazole-2-yl)-4-méthoxybenzamide ;
109. N-(5-isopropyl-1,3-thiazole-2-yl)-3-méthoxybenzamide ;
110. N-(5-isopropyl-1,3-thiazole-2-yl)-2-méthoxybenzamide ;
111. N-(5-isopropyl-1,3-thiazole-2-yl)-3,4-diméthoxybenzamide ;
45 112. N-(5-isopropyl-1,3-thiazole-2-yl)-3,5-diméthoxybenzamide ;
113. N-(5-isopropyl-1,3-thiazole-2-yl)-2,4-diméthoxybenzamide ;
114. N-(5-isopropyl-1,3-thiazole-2-yl)-2,3-diméthoxy-benzamide ;
115. N-(5-isopropyl-1,3-thiazole-2-yl)-3-phénoxybenzamide ;
116. N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénoxybenzamide ;
50 117. N-(5-isopropyl-1,3-thiazole-2-yl)-4-phénoxybenzamide ;
118. 2-éthoxy-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
119. 4-éthoxy-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
120. N-(5-isopropyl-1,3-thiazole-2-yl)-3,4,5-triméthoxybenzamide ;
121. 3,4-diéthoxy-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
55 122. 3,4,5-triéthoxy-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
123. N-(5-isopropyl-1,3-thiazole-2-yl)-3-méthoxy-4-(méthoxy méthoxy)benzamide ;
124. 4-butoxy-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
125. N-(5-isopropyl-1,3-thiazole-2-yl)-4-propoxybenzamide ;

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126. 4-isopropoxy-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 127. N-(5-isopropyl-1,3-thiazole-2-yl)-1,3-benzodioxole-5-carboxamide ;
 128. 4-(benzyloxy)-N-(5-isopropyl-1,3-thiazole-2-yl)propoxy-benzamide ;
 129. 4-(2-cyclohexène-1-yloxy)-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 130. N-(5-isopropyl-1,3-thiazole-2-yl)-4-(trifluorométhoxy)-benzamide ;
 131. 4-(difluorométhoxy)-N-(5-isopropyl-1,3-thiazole-2-yl)-benzamide ;
 132. N-(5-isopropyl-1,3-thiazole-2-yl)-4-(méthylsulfanyl)-benzamide ;
 133. 2-[(4-chlorophényl)sulfinyl]-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 134. N-(5-isopropyl-1,3-thiazole-2-yl)-2-[(4-nitrophényl)-sulfinyl]benzamide ;
 135. N-(5-isopropyl-1,3-thiazole-2-yl)-4-[(4-méthylphényl)-sulfonyl]-3-nitrobenzamide ;
 136. N-(5-isopropyl-1,3-thiazole-2-yl)-3-[(trifluorométhyl)-sulfanyl]benzamide ;
 137. N-(5-isopropyl-1,3-thiazole-2-yl)-2-méthoxy-4-(méthyl-sulfanyl)benzamide ;
 138. 2-[(2-cyanophényl)sulfanyl]-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 139. N~1~,N~1~-diéthyl-3,6-difluoro-N~2~- (5-isopropyl-1,3-thiazole-2-yl)phthalamide ;
 140. 4-formyl-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 141. 2-formyl-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 142. 4-[(2,5-diméthoxyanilino)carbonyl]amino-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 143. 4-(hydroxyméthyl)-N-(5-isopropyl-1,3-thiazole-2-yl)-benzamide ;
 144. Acétate de 4-[[5-isopropyl-1,3-thiazole-2-yl]amino]-carbonyl]-2-nitrobenzyle ;
 145. 4-(acétylamino)-3-iodobenzoate de 4-[[5-isopropyl-1,3-thiazole-2-yl]amino]carbonyl]-2-nitrobenzyle ;
 146. 4-(acétylamino)-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 147. N-(5-isopropyl-1,3-thiazole-2-yl)-4-[(2-phénylacétyl)-amino]benzamide ;
 148. 4-(acétylamino)-3-iodo-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 149. 4-amino-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 150. 4-(diméthylamino)-N-(5-isopropyl-1,3-thiazole-2-yl)-benzamide ;
 151. 3-(diméthylamino)-N-(5-isopropyl-1,3-thiazole-2-yl)-benzamide ;
 152. 2-(méthylamino)-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 153. N-(5-isopropyl-1,3-thiazole-2-yl)-2-[3-(trifluorométhyl)-anilino]benzamide ;
 154. 3-[[5-bromo-1,3-dioxo-1,3-dihydro-2H-isoindole-2-yl)-méthyl]amino]-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 155. N-(5-isopropyl-1,3-thiazole-2-yl)-4-(1H-pyrrol-1-yl)-benzamide ;
 156. 2,6-dichloro-N-(5-isopropyl-1,3-thiazole-2-yl)-isonicotinamide ;
 157. 2-(4-bromophényl)-6-(4-iodophényl)-N-(5-isopropyl-1,3-thiazole-2-yl)isonicotinamide ;
 158. N-(5-isopropyl-1,3-thiazole-2-yl)-2-[3-(trifluorométhyl)anilino]nicotinamide ;
 159. 2,6-dichloro-N-(5-isopropyl-1,3-thiazole-2-yl)-nicotinamide ;
 160. 5,6-dichloro-N-(5-isopropyl-1,3-thiazole-2-yl)-nicotinamide ;
 161. 2-chloro-N-(5-isopropyl-1,3-thiazole-2-yl)-6-méthyl-nicotinamide ;
 162. 2,6-dichloro-5-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl) nicotinamide ;
 163. N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénoxy nicotinamide ;
 164. N-(5-isopropyl-1,3-thiazole-2-yl)-6-(2,2,2-trifluoréthoxy)-nicotinamide ;
 165. N-(5-isopropyl-1,3-thiazole-2-yl)-2,6-diméthoxy-nicotinamide ;
 166. N-(5-isopropyl-1,3-thiazole-2-yl)-2-quinoxalinecarboxamide ;
 167. N-(5-isopropyl-1,3-thiazole-2-yl)-5-méthyl-2-pyrazinecarboxamide ;
 168. N-(5-isopropyl-1,3-thiazole-2-yl)-8-quinoléinecarboxamide ;
 169. N-(5-isopropyl-1,3-thiazole-2-yl)-2-phényl-4-quinoléine-carboxamide ;
 170. N-(5-isopropyl-1,3-thiazole-2-yl)-5-méthyl-1-phényl-1H-pyrazole-4-carboxamide ;
 271. N-(5-isopropyl-1,3-thiazole-2-yl)-5-méthyl-1H-pyrazole-3-carboxamide ;
 172. N-(5-isopropyl-1,3-thiazole-2-yl)-1H-pyrazole-4-carboxamide ;
 173. N-(5-isopropyl-1,3-thiazole-2-yl)-5-méthyl-2-phényl-2H-1,2,3-triazole-4-carboxamide ;
 174. 2-[(2,1,3-benzoxadiazol-5-yloxy)méthyl]-N-(5-isopropyl-1,3-thiazole-2-yl)-4-méthyl-1,3-thiazole-5-carboxamide ;
 175. N-(5-isopropyl-1,3-thiazole-2-yl)-9H-fluorène-1-carboxamide ;
 176. N-(5-isopropyl-1,3-thiazole-2-yl)-7-méthoxy-1-benzofuranne-2-carboxamide ;
 177. N-(5-isopropyl-1,3-thiazole-2-yl)-1-[(4-méthylphényl)-sulfonyl]-1H-pyrrole-3-carboxamide ;
 178. 2-éthoxy-N-(5-isopropyl-1,3-thiazole-2-yl)-1-naphtamide ;
 179. 4-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-1-naphtamide ;
 180. N-(5-isopropyl-1,3-thiazole-2-yl)-2-naphtamide ;
 181. N-(5-isopropyl-1,3-thiazole-2-yl)-9,10-dioxo-9,10-dihydro-2-anthracène-carboxamide ;

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182. N-(5-isopropyl-1,3-thiazole-2-yl)-9-oxo-9H-fluorène-4-carboxamide ;
 183. N-(5-isopropyl-1,3-thiazole-2-yl)-9-oxo-9H-fluorène-1-carboxamide ;
 184. N-(5-isopropyl-1,3-thiazole-2-yl)-8-oxo-5, 6, 7, 8-tétrahydro-2-naphtalèncarboxamide ;
 185. N-(5-isopropyl-1,3-thiazole-2-yl)-1,3-dioxo-1,3-dihydro-2-benzofuranne-5-carboxamide ;
 5 186. N-(5-isopropyl-1,3-thiazole-2-yl)-1H-indole-5-carboxamide ;
 187. N-(5-isopropyl-1,3-thiazole-2-yl)-1H-indole-4-carboxamide ;
 188. N-(5-isopropyl-1,3-thiazole-2-yl)-1-méthyl-2-phényl-1H-indole-5-carboxamide ;
 189. 2-butyl-N-(5-isopropyl-1,3-thiazole-2-yl)-1-méthyl-1H-indole-5-carboxamide ;
 190. N-(5-isopropyl-1,3-thiazole-2-yl)-1H-indole-6-carboxamide ;
 10 191. N-(5-isopropyl-1,3-thiazole-2-yl)-5-méthoxy-1H-indole-2-carboxamide ;
 192. 1-allyl-2-butyl-N-(5-isopropyl-1,3-thiazole-2-yl)-1H-indole-5-carboxamide ;
 193. N-(5-isopropyl-1,3-thiazole-2-yl)-1-méthyl-1H-indole-2-carboxamide ;
 194. 1-benzyl-N-(5-isopropyl-1,3-thiazole-2-yl)-2-phényl-1H-indole-5-carboxamide ;
 195. N-(5-isopropyl-1,3-thiazole-2-yl)-1H-1,2,3-benzotriazole-5-carboxamide ;
 15 196. N-(5-isopropyl-1,3-thiazole-2-yl)-3,5-diméthyl-4-isoxazolecarboxamide ;
 197. N-(5-isopropyl-1,3-thiazole-2-yl)-3-thiophèncarboxamide ;
 198. N-(5-isopropyl-1,3-thiazole-2-yl)-3-méthyl-2-thiophène-carboxamide ;
 199. N-(5-isopropyl-1,3-thiazole-2-yl)-5-méthyl-2-thiophène-carboxamide ;
 200. 5-bromo-N-(5-isopropyl-1,3-thiazole-2-yl)-2-thiophène-carboxamide ;
 20 201. N-(5-isopropyl-1,3-thiazole-2-yl)-3-[(2,3,3-trichloro-acryloyl)amino]-2-thiophèncarboxamide ;
 202. 5-bromo-N-(5-isopropyl-1,3-thiazole-2-yl)-2-furamide ;
 203. N-(5-isopropyl-1,3-thiazole-2-yl)-2-furamide ;
 204. N-(5-isopropyl-1,3-thiazole-2-yl)-5-(4-nitrophényl)-2-furamide ;
 205. N-(5-isopropyl-1,3-thiazole-2-yl)-5-(2-nitrophényl)-2-furamide ;
 25 206. 5-(4-chlorophényl)-N-(5-isopropyl-1,3-thiazole-2-yl)-2-furamide ;
 207. N-(5-isopropyl-1,3-thiazole-2-yl)-5-[3-(trifluorométhyl)-phényl]-2-furamide ;
 208. 5-(4-chloro-2-nitrophényl)-N-(5-isopropyl-1,3-thiazole-2-yl)-2-furamide ;
 209. N-(5-isopropyl-1,3-thiazole-2-yl)-5-(4-méthyl-2-nitro-phényl)-2-furamide ;
 210. 5-[2-chloro-5-(trifluorométhyl)phényl]-N-(5-isopropyl-1,3-thiazole-2-yl)-2-furamide ;
 30 211. (1R)-2-[(5-isopropyl-1,3-thiazole-2-yl)amino]2-oxo-1-phényléthylcarbamate de tertibutyle ;
 212. Acétate de (1R)-2-[(5-isopropyl-1,3-thiazole-2-yl)-amino]-2-oxo-1-phényléthyle ;
 213. Acétate de (1S)-2-[(5-isopropyl-1,3-thiazole-2-yl)-amino]-2-oxo-1-phényléthyle ;
 214. (R,S)-2-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-2-phényl-acétamide ;
 215. (R)-2-fluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-2-phényl-acétamide ;
 35 216. (S)-2-fluoro-(5-isopropyl-1,3-thiazole-2-yl)-2-phényl-acétamide ;
 217. 2-(acétylamino)-N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénylacétamide ;
 218. (R,S)-2-(méthoxy)-N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénylacétamide ;
 219. (R)-2-(méthoxy)-N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénylacétamide ;
 220. (S)-2-(méthoxy)-N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénylacétamide ;
 40 221. 3,3,3-trifluoro-N-(5-isopropyl-1,3-thiazole-2-yl)-2-méthoxy-2-phénylpropanamide ;
 222. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(1-naphtyl)acétamide ;
 223. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2-naphtyl)acétamide ;
 224. 2-(1H-indole-3-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)-acétamide ;
 225. 2-(1,3-benzodioxol-4-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
 45 226. 2-(2,4-dinitrophényl)-N-(5-isopropyl-1,3-thiazole-2-yl) acétamide ;
 227. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2-méthyl-1H-indole-3-yl)acétamide ;
 228. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(1-méthyl-1H-indole-3-yl)acétamide ;
 229. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(5-méthoxyl-1H-indole-3-yl)acétamide ;
 230. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(5-benzyloxy-1H-indole-3-yl)acétamide ;
 50 231. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3-méthoxy-2-méthyl-1H-indole-3-yl)acétamide ;
 232. 2-(1H-indole-3-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)-2-oxoacétamide ;
 233. 2-(5-bromo-1H-indole-3-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
 234. 2-(5-fluoro-1H-indole-3-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
 235. 2-[1-(4-chlorobenzoyl)-5-méthoxy-2-méthyl-1H-indole-3-yl]-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
 55 236. 3-(1H-indole-3-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)-propanamide ;
 237. 4-(1H-indole-3-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)-butanamide ;
 238. N-(5-isopropyl-1,3-thiazole-2-yl)-3-(2-thiényl)-propanamide ;
 239. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2-thiényl)-acétamide ;

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240. N-(5-isopropyl-1,3-thiazole-2-yl)-2-oxo-2-(2-thiényl)-acétamide ;
 241. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3-thiényl)-acétamide ;
 242.2-(5-chloro-1-benzothiophène-3-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
 243.2-(1-benzothiophène-3-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
 5 244.2-[2-(formylamino)-1,3-thiazole-4-yl]-N-(5-isopropyl-1,3-thiazole-2-yl)-2-(méthoxyimino)acétamide ;
 245.2-[2-[(2-chloroacétyl)amino]-1,3-thiazole-4-yl]-N-(5-isopropyl-1,3-thiazole-2-yl)-2-(méthoxyimino)
 acétamide ;
 246.2-chloro-N-(4-{2-[(5-isopropyl-2,3-thiazole-2-yl)-amino]-2-oxoéthyl}-1,3-thiazole-2-yl)acétamide ;
 247.2-[[2-[(5-isopropyl-1,3-thiazole-2-yl)amino]-2-oxo-1-(1H-pyrazole-3-yl)éthylidène]amino]oxy]acétate
 10 d'éthyle ;
 248.2-(2-furyl)-N-(5-isopropyl-1,3-thiazole-2-yl)-2-oxoacétamide ;
 249.2-(5-bromo-3-pyridinyl)-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
 250.N-(5-isopropyl-1,3-thiazole-2-yl)-2-(7-méthoxy-2-oxo-2H-chromène-4-yl)acétamide ;
 251.N-(5-isopropyl-1,3-thiazole-2-yl)-4-oxo-4-(4-méthyl-phényl)acétamide ;
 15 252.N-(5-isopropyl-1,3-thiazole-2-yl)-4-(4-nitrophényl)-butanamide ;
 253.N-(5-isopropyl-1,3-thiazole-2-yl)-4-phénylbutanamide ;
 254.4-[(5-isopropyl-1,3-thiazole-2-yl)amino]-4-oxobutyl-carbamate de benzyle ;
 255.5-[(5-isopropyl-1,3-thiazole-2-yl)amino]-5-oxo-pentanoate de méthyle ;
 256.4-(1,3-dioxo-1,3-dihydro-2H-isoindole-2-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)butanamide ;
 20 257.N-(5-isopropyl-1,3-thiazole-2-yl)-4-(4-méthoxy-1-naphtyl)-4-oxobutanamide ;
 258.3-(4-chlorophényl)-N-(5-isopropyl-1,3-thiazole-2-yl)-propanamide ;
 259.3-(4-méthoxyphényl)-N-(5-isopropyl-1,3-thiazole-2-yl)-propanamide ;
 260.3-phényl-N-(5-isopropyl-1,3-thiazole-2-yl)propanamide ;
 261.N-(5-isopropyl-1,3-thiazole-2-yl)-5-oxo-5-phénylpentanamide ;
 25 262.Acétate de 2-[(5-isopropyl-1,3-thiazole-2-yl)amino]-2-oxo-1-phényléthyle ;
 263.N-(5-isopropyl-1,3-thiazole-2-yl)-2-[4-(1-oxo-1,3-dihydro-2H-isoindole-2-yl)phényl]propanamide ;
 264.1-(4-chlorophényl)-N-(5-isopropyl-1,3-thiazole-2-yl)cyclo-pentanecarboxamide ;
 265.1-phényl-N-(5-isopropyl-1,3-thiazole-2-yl)cyclopentane-carboxamide ;
 266.2-(3-bromo-4-méthoxyphényl)-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
 30 267.2-(2-nitro-4-trifluorométhylphényl)-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
 268.(2S)-2-[(tertobutoxycarbonyl)amino]pentanedioate de 5-cyclohexyle 1-(4-{2-[(5-isopropyl-1,3-thiazole-
 2-yl) amino]-2-oxoéthyl}benzyl) ;
 269.2-(5,6-diméthyl-1H-benzimidazole-1-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
 270.2-[5-(4-chlorophényl)-2H-1,2,3,4-tétrazole-2-yl]-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
 35 271.N-(5-isopropyl-1,3-thiazole-2-yl)-2-[5-(1-pyrrolidinyl)-2H-1,2,3,4-tétrazole-2-yl]acétamide ;
 272.N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3-méthyl-1-benzo-thiophène-2-yl)acétamide ;
 273.N-{4-bromo-6-[(5-isopropyl-1,3-thiazole-2-yl)amino]-6-oxohexyl}benzamide ;
 274.6-[(5-isopropyl-1,3-thiazole-2-yl)amino]-6-oxohexyl-carbamate de benzyle ;
 275.4-(2,4-diméthylphényl)-N-(5-isopropyl-1,3-thiazole-2-yl)-4-oxobutanamide ;
 40 276.4-(4-benzyloxyphényl)-N-(5-isopropyl-1,3-thiazole-2-yl)-4-oxobutanamide ;
 277.4-(thiophène-2-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)-4-oxobutanamide ;
 278.2-[[benzyloxy]carbonyl]amino]-5-[(5-isopropyl-1,3-thiazole-2-yl)amino]-5-oxopentanoate de benzyle ;
 279.4-(1H-indole-3-yl)-N-{3-[(5-isopropyl-1,3-thiazole-2-yl)amino]-3-oxopropyl}butanamide ;
 280.4-chlorobenzènesulfonate de 4-[[5-isopropyl-1,3-thiazole-2-yl)amino]carbonyl]phényle ;
 45 281.N-(5-isopropyl-1,3-thiazole-2-yl)-4-[[2-méthoxyanilino]-carbonyl]amino}benzamide ;
 282.4-[[2-(isopropylsulfonyl)acétyl]amino]-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 283.N-(5-isopropyl-1,3-thiazole-2-yl)-4-[[2-(phénylsulfanyl)-acétyl]amino]benzamide ;
 284.4-[(diéthylamino)sulfonyl]-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 285.2-bromo-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 50 286.3,5-difluoro-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 287.3-[[2-(fluoranilino)carbonyl]amino]-N-(5-isopropyl-1,3-thiazole-2-yl)benzamide ;
 288.N-(5-isopropyl-1,3-thiazole-2-yl)-1-phényl-5-propyl-1H-pyrazole-4-carboxamide ;
 289.3-chloro-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazole-2-yl)-5-(méthylsulfanyl)-2-thiophèncarboxa-
 xamide ;
 55 290.3-iodo-4-(isopropylsulfonyl)-N-(5-isopropyl-1,3-thiazole-2-yl)-5-(méthylsulfanyl)-2-thiophèncarboxa-
 mide ;
 291.2-[[4-chlorophényl)sulfonyl]méthyl]-N-(5-isopropyl-1,3-thiazole-2-yl)-4-méthyl-1,3-thiazole-
 5-carboxamide ;

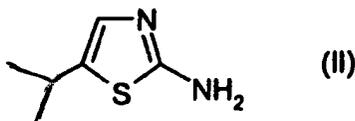
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- 292.5-(4-chlorophényl)-N-(5-isopropyl-1,3-thiazole-2-yl)-2-(trifluorométhyl)-3-furamide ;
293. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2,3,4,5,6-pentafluorophényl)acétamide ;
294. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2-fluorophényl)-acétamide ;
5 295. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2-bromophényl)-acétamide ;
296. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2-chlorophényl)-acétamide ;
297. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2-nitrophényl)-acétamide ;
298. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2-trifluoro-méthylphényl)acétamide ;
299. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2-méthoxyphényl)-acétamide ;
10 300. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2,5-diméthoxyphényl)-acétamide ;
301. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2,5-difluorophényl)-acétamide ;
302. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3,4,5-triméthoxy-phényl)acétamide ;
303. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2,6-dichlorophényl)-acétamide ;
304. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2-chloro-6-fluoro-phényl)acétamide ;
305. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3,5-diméthoxyphényl)-acétamide ;
15 306. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3,5-difluorophényl)-acétamide ;
307. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2,5-bis-trifluoro-méthylphényl)acétamide ;
308. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-méthylthiophényl)-acétamide ;
309. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-méthoxyphényl)-acétamide ;
310. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-bromophényl)-acétamide ;
20 311. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-chlorophényl)-acétamide ;
312. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-fluorophényl)-acétamide ;
313. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-nitrophényl)-acétamide ;
314. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-trifluoro-méthylphényl)acétamide ;
315. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-méthylphényl)-acétamide ;
25 316. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-diméthylamino-phényl)acétamide ;
317. 2-[1,1'-biphényl]-4-yl-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
318. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3-trifluorométhyl-phényl)acétamide ;
319. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3-bromophényl)-acétamide ;
320. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3-chlorophényl)-acétamide ;
30 321. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3-nitrophényl)-acétamide ;
322. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3-méthoxyphényl)-acétamide ;
323. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2,4-dinitrophényl)-acétamide ;
324. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2,4-dichlorophényl)acétamide ;
325. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(2,4-difluorophényl)acétamide ;
35 326. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-benzyloxy-3-méthoxyphényl)acétamide ;
327. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3,4-dichlorophényl)-acétamide ;
328. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3,4-difluorophényl)-acétamide ;
329. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(3,4-diméthoxyphényl)-acétamide ;
330. 2-(2,3-dihydro-1H-indène-5-yl)-N-(5-isopropyl-1,3-thiazole-2-yl)acétamide ;
40 331. N-(5-isopropyl-1,3-thiazole-2-yl)-1-phénylcyclopropane-carboxamide ;
332. 2-cyclopentyl-N-(5-isopropyl-1,3-thiazole-2-yl)-2-phényl-acétamide ;
333. 2-cyclohexyl-N-(5-isopropyl-1,3-thiazole-2-yl)-2-phényl-acétamide ;
334. N-(5-isopropyl-1,3-thiazole-2-yl)-2,2-diphénylacétamide ;
335. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-nitrophényl)-propanamide ;
45 336. N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénylpropanamide ;
337. N-(5-isopropyl-1,3-thiazole-2-yl)-2-(4-isobutylphényl)-propanamide ;
338. N-(5-isopropyl-1,3-thiazole-2-yl)-2-oxo-2-phényl-acétamide ;
339. N-(5-isopropyl-1,3-thiazole-2-yl)-3-méthyl-2-phényl-pentanamide ;
340. N-(5-isopropyl-1,3-thiazole-2-yl)bicyclo[4.2.0]octa-1,3,5-triène-7-carboxamide ;
50 341. N-(5-isopropyl-1,3-thiazole-2-yl)-3-oxo-1-indane-carboxamide ;
342. N-(5-isopropyl-1,3-thiazole-2-yl)-2-phénylbutanamide ;
343. (1S)-1-(1H-indole-3-ylméthyl)-2-[(5-isopropyl-1,3-thiazole-2-yl)amino]-2-oxoéthylcarbamate de tertio-
butyle ;
55 344. (1S)-1-(4-benzyloxybenzyl)-2-[(5-isopropyl-1,3-thiazole-2-yl)amino]-2-oxoéthylcarbamate de tertio-
butyle ;

et leurs sels pharmaceutiquement acceptables.

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10. Procédé pour la production d'un composé de formule (I), répondant à la définition figurant dans la revendication 7, procédé qui comprend la réaction d'un composé de formule (II)



10 avec un composé de formule (III)



15 dans laquelle R_1 est tel que défini dans la revendication 7 et X représente un groupe hydroxy ou un groupe partant convenable ;

et, si cela est désiré, la conversion d'un dérivé de 2-amino-1,3-thiazole de formule (I) en un autre tel dérivé de formule (I), et/ou en un de ses sels.

- 20 11. Procédé suivant la revendication 10, dans lequel X représente un groupe hydroxy, un atome de brome ou un atome de chlore.

- 25 12. Composition pharmaceutique comprenant un ou plusieurs supports et/ou diluants pharmaceutiquement acceptables et, comme principe actif, une quantité efficace d'un composé de formule (I) répondant à la définition figurant dans la revendication 7.

- 30 13. Composé de formule (I), suivant la revendication 7, dans lequel R_1 est facultativement substitué avec un ou plusieurs groupes choisis entre des groupes : halogéno, nitro, oxo, (=O), carboxy, cyano, alkyle, perfluoralkyle, alcényle, alcynyle, cycloalkyle, aryle, hétérocyclyle ; des groupes amino et leurs dérivés tels que des groupes alkylamino, alkoxy-carbonylalkylamino, dialkylamino, arylamino, diarylamino ou aryluréido ; des groupes carbonylamino et leurs dérivés tels que des groupes hydrogénocarbonylamino (HCONH-), alkylcarbonylamino, arylcarbonylamino, alkoxy-carbonylamino ; des oximes substituées au niveau de l'oxygène, telles que des groupes alkoxy-carbonylalkoxyimino ou alkoxyimino ; des groupes hydroxy et leurs dérivés tels que des groupes alkoxy, aryloxy, alkyl-carbonyloxy, arylcarbonyloxy, cycloalcényloxy ; des groupes carbonyle et leurs dérivés tels que des groupes alkyl-carbonyle, arylcarbonyle, alkoxy-carbonyle, aryloxy-carbonyle, cycloalkyloxy-carbonyle, aminocarbonyle, alkylaminocarbonyle, dialkylaminocarbonyle ; des dérivés sulfurés tels que des groupes alkylthio, arylthio, alkylsulfonyle, arylsulfonyle, alkylsulfinyle, arylsulfinyle, arylsulfonyloxy, aminosulfonyle, alkylaminosulfonyle ou dialkylaminosulfonyle.

- 40 14. Composé de formule (I), suivant la revendication 13, dans lequel R_1 est facultativement substitué avec un ou plusieurs groupes choisis entre des groupes : halogéno, nitro, oxo (=O), carboxy, alkyle, perfluoralkyle, cycloalkyle, aryle, hétérocyclyle ; des groupes amino ; des groupes carbonylamino tels que des groupes alkylcarbonylamino, arylcarbonylamino ou alkoxy-carbonylamino ; des groupes hydroxy et leurs dérivés tels que des groupes alkoxy, aryloxy ou alkylcarbonyloxy ; des groupes carbonyle et leurs dérivés tels que des groupes alkylcarbonyle ou aryl-carbonyle.
- 45