PATENT REQUEST :  STANDARD PATENT

I/We being the person(s) identified below as the Applicant(s), request the grant of a patent to the person(s) identified below as the Nominated Person(s), for an invention described in the accompanying standard complete specification.

Full application details follow:

[71/70] Applicant(s)/Nominated Person(s):
Fujisawa Pharmaceutical Co., Ltd.
of
4-7, Doshomachi 3-chome, Chuo-ku, Osaka-shi, Osaka, 541, Japan

[54] Invention Title:
IMIDAZO [1,2-A] PYRIDINE DERIVATIVES AS BRADYKININ ANTAGONISTS, PHARMACEUTICALS AND PROCESSES FOR THEIR PREPARATION

[72] Name(s) of actual inventor(s):
Teruo OKU
Hiroshi KAYAKIRI
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[74] Address for service in Australia:
DAVIES COLLISON CAVE, Patent Attorneys, 1 Little Collins Street, Melbourne, Victoria, Australia. Attorney Code: DM

Basic Convention Application(s) Details:

<table>
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<th>Application Number</th>
<th>Country</th>
<th>Code</th>
<th>Date of Application</th>
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<tr>
<td>9222947.5</td>
<td>United Kingdom</td>
<td>GB</td>
<td>2 November 1992</td>
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<td>9304249.7</td>
<td>United Kingdom</td>
<td>GB</td>
<td>3 March 1993</td>
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DATED this TWENTY SIXTH day of OCTOBER 1993

[Signature]

a member of the firm of DAVIES COLLISON CAVE for and on behalf of the applicant(s)
AUSTRALIA
PATENTS ACT 1990
NOTICE OF ENTITLEMENT

We, Fujisawa Pharmaceutical Co., Ltd., the applicant/Nominated Person named in the accompanying Patent Request state the following:-

The Nominated Person is entitled to the grant of the patent because the Nominated Person derives the invention from the inventors by assignment.

The Nominated Person is entitled to claim priority from the basic applications listed on the patent request because the Nominated Person made the basic applications, and because those applications were the first applications made in a Convention country in respect of the invention.

DATED this TWENTY SIXTH day of OCTOBER 1993

Keith...[signature]
a member of the firm of DAVIES COLLISON CAVE for and on behalf of the applicant(s)

(DCC ref: 1626439)
IMIDAZO (1,2-A) PYRIDINE DERIVATIVES AS BRADYKININ ANTAGONISTS, PHARMACEUTICALS AND PROCESSES FOR THEIR PREPARATION

1. The compound of the formula:

\[
\begin{align*}
\text{R}^3 & \quad \text{N} & \quad \text{R}^1 \\
\text{Q} & \quad \text{A} & \quad \text{R}^2
\end{align*}
\]

wherein \( \text{R}^1 \) is halogen.

\( \text{R}^2 \) and \( \text{R}^3 \) are each hydrogen, lower alkyl, halo (lower) alkyl or acyl.

\( \text{R}^4 \) is aryl substituted with substituent(s) selected from the group consisting of halogen; halo (lower) alkyl; acyl; aryl substituted with halogen or cyano; ar (lower) alkyl substituted with hydroxy; lower alkoxy; nitro; amino; amino substituted with substituent(s) selected from the group consisting of lower alkyl, acyl, ar (lower) alkyl, heterocyclic (lower) alkyl, carboxy (lower) alkyl, lower alkylaminomethylene and N-methylpyrrolidinyldiene; a heterocyclic group and a heterocyclic group substituted with oxo; or a heterocyclic group substituted with substituent(s) selected from the group consisting of halogen; lower alkyl; halo (lower) alkyl; acyl; aryl; aryl substituted with halogen or
cyano; ar (lower) alkyl substituted with hydroxy; lower alkoxy; oxo; nitro; amino; amino substituted with substituent(s) selected from the group consisting of lower alkyl, acyl, ar (lower) alkyl, heterocyclic (lower) alkyl, carboxy (lower) alkyl, lower alkylaminomethylene and N-methylpyrrolidinylidene; a heterocyclic group and a heterocyclic group substituted with oxo.

Q is O or N-R^{11},
in which R^{11} is hydrogen or acyl, and
A is lower alkylene,
and pharmaceutically acceptable salts thereof.
NAME OF APPLICANT(S):
Fujisawa Pharmaceutical Co., Ltd.

ADDRESS FOR SERVICE:

DAVIES COLLISON CAVE
Patent Attorneys
1 Little Collins Street, Melbourne, 3000.

INVENTION TITLE:
Imidazo [1,2-a] pyridine derivatives as bradykinin antagonists, pharmaceuticals and processes for their preparation

The following statement is a full description of this invention, including the best method of performing it known to me/us:-
This invention relates to new heterocyclic compounds and pharmaceutically acceptable salts thereof.

More particularly, it relates to new heterocyclic compounds and pharmaceutically acceptable salts thereof which have activities as bradykinin antagonists, to processes for preparation thereof, to a pharmaceutical composition comprising the same, and to methods of using the same therapeutically in the prevention and/or the treatment of bradykinin or its analogues mediated diseases such as allergy, inflammation, autoimmune disease, shock, pain, or the like, in human being or animals.

One object of this invention is to provide new and useful heterocyclic compounds and pharmaceutically acceptable salts thereof which possess activities as bradykinin antagonists.

Another object of this invention is to provide processes for the preparation of said compounds and salts thereof.

A further object of this invention is to provide a
pharmaceutical composition comprising, as an active ingredient, said heterocyclic compounds and pharmaceutically acceptable salts thereof.

Still further object of this invention is to provide a therapeutical method for the prevention and/or the treatment of bradykinin or its analogues mediated diseases such as allergy, inflammation, autoimmune disease, shock, pain, or the like, using said heterocyclic compounds and pharmaceutically acceptable salts thereof.

Some heterocyclic compounds have been known as described, for example, in J. Med. Chem., 28, 876-892 (1985). However, it is not known that said compounds have activities as bradykinin antagonists.

The object heterocyclic compounds of this invention are new and can be represented by the following general formula [I]:

\[ Q-A-R^4 \]

wherein \( R^1 \) is halogen,
\( R^2 \) and \( R^3 \) are each hydrogen, lower alkyl, halo(lower)alkyl or acyl,
\( R^4 \) is aryl substituted with substituent(s) selected from the group consisting of halogen; halo (lower) alkyl; acyl; aryl; aryl substituted with halogen or cyano; ar (lower) alkyl substituted with hydroxy; lower alkoxy; nitro; amino; amino substituted with substituent(s) selected from the group consisting of lower alkyl, acyl, ar (lower) alkyl, heterocyclic (lower) alkyl, carboxy (lower) alkyl, lower alkylaminomethylene and \( N \)-methylpyrrolidinylidene; a heterocyclic
group and a heterocyclic group substituted with oxo; or a heterocyclic group substituted with substituent(s) selected from the group consisting of halogen; lower alkyl; halo (lower) alkyl; acyl; aryl; aryl substituted with halogen or cyano; ar (lower) alkyl substituted with hydroxy; lower alkoxy; oxo; nitro; amino; amino substituted with substituent(s) selected from the group consisting of lower alkyl, acyl, ar (lower) alkyl, heterocyclic (lower) alkyl, carboxy (lower) alkyl, lower alkylaminomethylene and N-methylpyrrolidinylidene; a heterocyclic group and a heterocyclic group substituted with oxo,

Q is O or N-R\textsuperscript{11},

in which R\textsuperscript{11} is hydrogen or acyl, and

A. is lower alkylene.
The object compound [I] or its salt can be prepared by processes as illustrated in the following reaction schemes.

Process 1

\[ \text{R}^3 \text{Q-A-R}^4 \xrightarrow{\text{Halo}} \text{R}^3 \text{Q-A-R}^4 \]

or its salt

Process 2

\[ \text{X-A-R}^4 [IV] \xrightarrow{\text{OH}} \text{R}^3 \text{Q-A-R}^4 \]

or its salt

or its salt
Process 3

\[
\text{[Ia] or its salt}
\]

Process 4

\[
\text{[Ic] or its salt}
\]
Process 5

5

or its salt

10

Alkylation

[If]

or its reactive derivative

at the carboxy group

or a salt thereof

Process 6

20

or its reactive derivative at the amino group

or a salt thereof

25
[Ig]
or its salt

wherein \( R^5 \) is hydrogen or lower alkyl,
\( R^5, R^7, R^8 \) and \( R^9 \) are each hydrogen or halogen,
\( R^{10} \) is acyl,
\( R^{10}_a \) is acyl having amino,
\( R^{10}_b \) is acyl having acylamino,
\( R^{10}_c \) is acyl having lower alkylamino or acyl having
acyl(lower)alkylamino,
\( R^{12} \) is hydrogen, lower alkyl, lower
alkoxy(lower)alkyl, lower
alkylamino(lower)alkyl,
heterocyclic(lower)alkyl, a heterocyclic
group, protected or unprotected
hydroxy(lower)alkyl or aryl optionally
substituted with lower alkylamino, and
\( R^{13} \) is hydrogen, lower alkyl, lower
alkoxy(lower)alkyl or protected or
unprotected hydroxy(lower)alkyl, or
\( R^{12} \) and \( R^{13} \) are taken together with the attached
nitrogen atom to form a heterocyclic group
optionally having suitable substituent(s);
\((AA)\) is amino acid residue,
\( X \) is a leaving group,
Y is NH or lower alkenylene,
Z is CH or N, and
R¹, R², R³, R⁴, Q and A are each as defined above.

In the above and subsequent description of the present specification, suitable examples of the various definitions to be included within the scope of the invention are explained in detail in the following.

The term "lower" is intended to mean a group having 1 to 6 carbon atom(s), unless otherwise provided.

In this respect, the term "lower" in lower alkenyl moiety, lower alkynyl moiety and ar(lower)alkenyl moiety in the various definitions is intended to mean a group having 2 to 6 carbon atoms.

Further, the term "lower" in lower alkenoyl moiety, lower alkynoyl moiety, cyclo(lower)alkyl moiety, cyclo(lower)alkenyl moiety, ar(lower)alkenoyl moiety, ar(lower)alkynoyl moiety and heterocyclic(lower)alkenoyl moiety in the various definitions is intended to mean a group having 3 to 6 carbon atoms.

Suitable "halogen" may be fluorine, chlorine, bromine and iodine.

Suitable "aryl" may be phenyl, naphthyl, phenyl substituted with lower alkyl [e.g. tolyl, xylyl, mesityl, cumenyl, di(tert-butyl)phenyl, etc.] and the like, in which preferable one is phenyl and tolyl.

Suitable "lower alkyl" and lower alkyl moiety in the terms "heterocyclic(lower)alkyl", and "lower alkylamino" may be straight or branched one such as methyl, ethyl, propyl, isopropyl, butyl, isobutyl, tert-butyl, penty1, hexyl or the like, in which preferable one is C₁-C₄ lower alkyl such as methyl, ethyl, propyl, isobutyl or tert-butyl.

Suitable "lower alkenylene" may be a straight or branched one such as methylene, ethylene, trimethylene,
methylmethylene, tetramethylene, ethylethylene, propylene, pentamethylene, hexamethylene or the like, in which the most preferable one is methylene.

Suitable "halo(lower)alkyl" may be fluoromethyl, difluoromethyl, trifluoromethyl, chloromethyl, dichloromethyl, bromomethyl, fluoroethyl, difluoroethyl, chloroethyl, dichloroethyl, or the like.

Suitable "lower alkenylene" may be a straight or branched C₂-C₆ alkenylene such as vinylene, methylvinylene, propylene, 1,3-butadienylene or the like, in which the most preferable one is vinylene.

Suitable "acyl" may be substituted or unsubstituted alkanoyl such as alkanoyl [e.g. formyl, acetyl, propionyl, butyryl, isobutyryl, valeryl, isovaleryl, pivaloyl, hexanoyl, heptanoyl, 3,3-dimethylbutyryl, etc.], halo(lower)alkanoyl [e.g. chloroacetyl, trifluoroacetyl, bromoacetyl, bromobutyryl, heptafluorobutyryl, etc.], hydroxy(lower)alkanoyl [e.g. glycoloyl, lactoyl, 3-hydroxypropionyl, glyceroyl, etc.], lower alkylsulfonyleoxy(lower)alkanoyl [e.g. mesyloxyacetyl, ethylsulfonyleoxyacetyl, mesyloxypropionyl, etc.], lower alkoxy(lower)alkanoyl [e.g. methoxyacetyl, ethoxyacetyl, methoxypropionyl, ethoxypropionyl, propoxypropionyl, methoxybutyryl, etc.], lower alkylthiio(lower)alkanoyl [e.g. methylthioacetyl, ethylthioacetyl, methylthiopropionyl, ethylthiopropionyl, propylthiopropionyl, methylthiobutyryl, etc.], lower alkanoyloxy(lower)alkanoyl [e.g. acetyloxyacetyl, acetyloxypropionyl, propionyloxyacetyl, etc.], aryloxy(lower)alkanoyl [e.g. phenyloxyacetyl, phenyloxypopionyl, tolyloxyacetyl, naphthoxyacetyl, etc.], aroyl(lower)alkanoyl [e.g. phenoxalyl, benzoylacetyl, benzoylpropionyl, etc.], carboxy(lower)alkanoyl [e.g. oxalo, carboxyacetyl, 3-carboxypropionyl, 3-carboxybutyryl, 4-carboxybutyryl,
4-carboxyvaleryl, etc.), esterified carboxy(lower)alkanoyl, for example, lower alkoxy carbonyl(lower)alkanoyl [e.g. methoxycarbonylacetyl, ethoxycarbonylacetyl, methoxycarbonylpropionyl, ethoxycarbonylpropionyl, etc.], carbamoyl(lower)alkanoyl [e.g. carbamoylacetyl, carbamoylpropionyl, etc.], lower alkyl carbamoyl(lower)alkanoyl [e.g. methyl carbamoylaceteyl, methyl carbamoylpropionyl, ethyl carbamoylpropionyl, dimethyl carbamoylpropionyl, (N-methyl-N-ethyl carbamoyl)-propionyl, etc.], ar(lower)alkanoyl [e.g. phenylacetyl, tolylacetyl, naphthylacetyl, 2-phenylpropionyl, 3-phenylpropionyl, 4-phenylbutyryl, tritylcarbonyl, etc.], optionally substituted heterocyclic(lower)alkanoyl [e.g. morpholinacetyl, thiomorpholinacetyl, morpholinopropionyl, thiomorpholinopropionyl, piperidinopropionyl, piperazinyllpropionyl, pyridylacetyl, pyrrolidinylpropionyl, imidazolidinylpropionyl, piperidinoacetyl, pyrrolidinylacetyl, hexamethyleneiminoacetyl, hexamethyleneiminopropionyl, imidazolylaceteyl, furylacetyl, thienylacetyl, methylpiperazinyacetyl, pyridylpiperazinyacetyl, etc.], heterocyclicthio(lower)alkanoyl [e.g. pyridylthioacetyl, pyrimidinylthioacetyl, imidazolylthioacetyl, etc.], etc., lower alkenoyl [e.g. acryloyl, crotonoyl, isocrotonoyl, 3-butenoyl, 3-pentenoyl, 4-pentenoyl, methacryloyl, etc.], lower alkynoyl [e.g. propioloyl, 2-butylnoyl, 3-butylnoyl, etc.], cyclo(lower)alkylcarbonyl [e.g. cyclopropylcarbonyl, cyclobutylcarbonyl, cyclopentylcarbonyl, cyclohexylcarbonyl, etc.], cyclo(lower)alkenylcarbonyl [e.g. cyclopentenylcarbonyl, cyclohexenylcarbonyl, etc.], carboxy, esterified carboxy such as lower alkoxy carbonyl [e.g. methoxycarbonyl, ethoxycarbonyl, propoxycarbonyl, isopropoxycarbonyl, butyoxycarbonyl, isobutoxycarbonyl, tert-butoxycarbonyl, pentyloxy carbonyl, hexyloxy carbonyl, etc.],
aryloxycarbonyl [e.g. phenoxycarbonyl, etc.], etc., substituted or unsubstituted aroyl such as aroyl [e.g. benzoyl, toluoyl, xyloyl, naphthoyl, etc.], lower alkoxyaroyl [e.g. methoxybenzoyl, etc.], haloaroyl [e.g. chlorobenzoyl, fluorobenzoyl, etc.], acylaroyl, for example, lower alkoxy carbonylaroyl [e.g. methoxycarbonylbenzoyl, etc.], etc., substituted or unsubstituted ar(lower)alkenoyl such as ar(lower)alkenoyl [e.g. cinnamoyl, allocinnamoyl, α-methylcinnamoyl, 4-methylcinnamoyl, etc.], lower alkoxy-ar(lower)alkenoyl [e.g. methoxycinnamoyl, ethoxycinnamoyl, dimethoxycinnamoyl, etc.], lower alkylenedioxy-ar(lower)alkenoyl [e.g. methylenedioxy cinnamoyl, ethylenedioxy cinnamoyl, etc.], nitro-ar(lower)alkenoyl [e.g. nitrocinnamoyl, etc.], halo-ar(lower)alkenoyl [e.g. chlorocinnamoyl, fluorocinnamoyl, etc.], hydroxy-ar(lower)alkenoyl [e.g. hydroxycinnamoyl, etc.], hydroxy(lower)alkoxy-ar(lower)alkenoyl [e.g. hydroxymethoxy cinnamoyl, hydroxyethoxy cinnamoyl, etc.], amino(lower)alkoxy-ar(lower)alkenoyl [e.g. aminothoxy cinnamoyl, etc.], lower alky lamino(lower)alkoxy-ar(lower)alkenoyl [e.g. methylaminomethoxy cinnamoyl, dimethylaminoethoxy cinnamoyl, etc.], heterocyclic(lower)alkoxy-ar(lower)alkenoyl [e.g. pyridylmethoxycinnamoyl, etc.], optionally substituted heterocyclic-ar(lower)alkenoyl [e.g. morpholinocinnamoyl, methylpiperazinylcinnamoyl, pyrrolidinylcinnamoyl, oxopyrrolidinylcinnamoyl, oxopiperidinocinnamoyl, dioxopyrrolidinylcinnamoyl, oxooxazolidinylcinnamoyl, pyrrolylcinnamoyl, etc.], amino-ar(lower)alkenoyl [e.g. aminocinnamoyl, etc.], lower alky lamino-ar(lower)alkenoyl [e.g. methylaminocinnamoyl, dimethylaminocinnamoyl, etc.], acylamino-ar(lower)alkenoyl, for example, lower alkanoylamino-ar(lower)alkenoyl [e.g.
acetylaminocinnamoyl, propionylaminocinnamoyl, etc.), hydroxy(lower)alkanoylamino-ar(lower)alkenoyl [e.g. hydroxyacetylaminocinnamoyl, hydroxypropionylaminocinnamoyl, etc.], lower alkoxy(lower)alkanoylamino-ar(lower)alkenoyl [e.g. methoxyacetylaminocinnamoyl, methoxypropionylaminocinnamoyl, etc.], halo(lower)alkanoylamino-ar(lower)alkenoyl [e.g. chloroacetylaminocinnamoyl, bromobutrylaminocinnamoyl, etc.], amino(lower)alkanoylamino-ar(lower)alkenoyl [e.g. aminoacetylaminocinnamoyl, aminopropionylaminocinnamoyl, etc.], lower alkylamino(lower)alkanoylamino-ar(lower)-alkenoyl [e.g. methylaminoacetylaminocinnamoyl, dimethylaminoacetylaminocinnamoyl, etc.], lower alkanoylamino(lower)alkanoylamino-ar(lower)alkenoyl [e.g. acetylaminoacetylaminocinnamoyl, acetylaminopropionylaminocinnamoyl, etc.], carboxy(lower)alkanoylamino-ar(lower)alkenoyl [e.g. carboxyacetylaminocinnamoyl, carboxypropionylaminocinnamoyl, etc.], lower alkoxy carbonyl(lower)alkanoylamino-ar(lower)alkenoyl [e.g. ethoxy carbonylacetylaminocinnamoyl, ethoxy carbonylpropionylaminocinnamoyl, etc.], lower alkoxy carbonyl(lower)alkanoylamino-ar(lower)alkenoyl [e.g. ethoxy carbonylacryloylaminocinnamoyl, etc.], halo(lower)alkoxy carbonylamo-ar(lower)alkenoyl [e.g. chloroethoxy carbonylaminocinnamoyl, etc.], heterocyclic(lower)alkanoylamino-ar(lower)alkenoyl [e.g. pyridylacetylaminocinnamoyl, etc.], aroylamino-ar(lower)alkenoyl [e.g. benzoylaminocinnamoyl, etc.], heterocycliccarbonylamino-ar(lower)alkenoyl [e.g. nicotinylaminocinnamoyl, isonicotinylaminocinnamoyl, morpholinocarbonylaminocinnamoyl, etc.], lower alkylsulfonylamino-ar(lower)alkenoyl [e.g. mesylaminocinnamoyl, ethylsulfonylaminocinnamoyl, etc.],
etc., N-(lower alkanoyl)-N-(lower alkyl)amino-ar(lower)alkenooyl [e.g. N-acetyl-N-methylaminocinnamoyl, N-acetyl-N-ethylaminocinnamoyl, N-propionyl-N-methylaminocinnamoyl, etc.], N-[lower alkoxy(lower)alkanoyl]-N-(lower alkyl)amino-ar(lower)alkenooyl [e.g. N-methoxyacetyl-N-methylaminocinnamoyl, N-methoxypropionyl-N-methylaminocinnamoyl, etc.], N-(lower alkanoyl)-N-[heterocyclic(lower)alkyl]amino-ar(lower)alkenooyl [e.g. N-acetyl-N-pyridylmethylaminocinnamoyl, etc.], ureido-ar(lower)alkenooyl [e.g. ureidocinnamoyl, etc.], lower alkyureido-ar(lower)alkenooyl [e.g. methylureidocinnamoyl, ethylureidocinnamoyl, dimethylureidocinnamoyl, etc.], heterocyclicureido-ar(lower)alkenooyl [e.g. pyridylureidocinnamoyl, pyrimidinylureidocinnamoyl, thienylureidocinnamoyl, etc.], acyl-ar(lower)alkenooyl, for example, lower alkanoyl-ar(lower)alkenooyl [e.g. formylcinnamoyl, acetylcinnamoyl, propionylcinnamoyl, etc.], carboxy-ar(lower)alkenooyl [e.g. carboxycinnamoyl, etc.], lower alkoxyacyl-ar(lower)alkenooyl [e.g. methoxycarbonylcinnamoyl, ethoxycarbonylcinnamoyl, etc.], carbamoyl-ar(lower)alkenooyl [e.g. carbamoylcinnamoyl, etc.], lower alkylcarbamoyl-ar(lower)alkenooyl [e.g. methylcarbamoylcinnamoyl, ethylcarbamoylcinnamoyl, dimethylcarbamoylcinnamoyl, propylcarbamoylcinnamoyl, isopropylcarbamoylcinnamoyl, diethylcarbamoylcinnamoyl, N-methyl-N-ethylcarbamoylcinnamoyl, etc.], hydroxy(lower)alkylcarbamoyl-ar(lower)alkenooyl [e.g. hydroxyethylcarbamoylcinnamoyl, bis(hydroxyethyl)carbamoylcinnamoyl, etc.], N-[hydroxy(lower)alkyl]-N-(lower alkyl)carbamoyl-ar(lower)alkenooyl [e.g. N-hydroxyethyl-N-methylcarbamoylcinnamoyl, etc.],
lower alkoxy(lower)alkylcarbamoyl-ar(lower)alkenoyl [e.g. methoxymethylcarbamoylcinnamoyl, methoxyethylcarbamoylcinnamoyl, bis(methoxyethyl)carbamoylcinnamoyl, ethoxyethylcarbamoylcinnamoyl, methoxypropylcarbamoylcinnamoyl, bis(ethoxyethyl)carbamoylcinnamoyl, etc.], N-[lower alkoxy(lower)alkyl]-N-(lower alkyl)carbamoyl-ar(lower)alkenoyl [e.g. N-methoxyethyl-N-methylcarbamoylcinnamoyl, etc.], heterocycliccarbamoyl-ar(lower)alkenoyl [e.g. morpholinylcarbamoylcinnamoyl, thiienylcarbamoylcinnamoyl, pyridylcarbamoylcinnamoyl, pyrimidinylcarbamoylcinnamoyl, etc.], heterocycliccarbonyl-ar(lower)alkenoyl [e.g. morpholinocarbonylcinnamoyl, pyrrolidinocarbonylcinnamoyl, piperidinocarbonylcinnamoyl, etc.], ar(lower)alkenoyl [e.g. phenylpropioloyl, et c.], substituted or unsubstituted heterocyclic(lower)alkenoyl such as heterocyclic(lower)alkenoyl [e.g. morpholinylacryloyl, pyridylacryloyl, thiienylacryloyl, etc.], amino-heterocyclic(lower)alkenoyl [e.g. aminopyridylacryloyl, etc.], lower alkylamino-heterocyclic(lower)alkenoyl [e.g. methylaminopyridylacryloyl, dimethylaminopyridylacryloyl, etc.], acylamino-heterocyclic(lower)alkenoyl, for example, lower alkanoylamino-heterocyclic(lower)alkenoyl [e.g. acetylaminopyridylacryloyl, propionylaminopyridylacryloyl, etc.], etc., lower alkylureido-heterocyclic(lower)alkenoyl [e.g. methylureidopyridylacryloyl, etc.], acyl-heterocyclic(lower)alkenoyl, for example, carboxy-heterocyclic(lower)alkenoyl [e.g. carboxy pyridylacryloyl, etc.], lower alkoxy carbonyl-heterocyclic(lower)alkenoyl [e.g. ethoxycarbonylpyridylacryloyl, etc.], lower
alkylcarbamoyl-heterocyclic(lower)alkenoyl [e.g. methylcarbamoylpyridylacryloyl, ethylcarbamoylpyridylacryloyl, dimethylcarbamoylpyridylacryloyl, diethylcarbamoylpyridylacryloyl, isopropylcarbamoylpyridylacryloyl, etc.], lower alkoxy(lower)alkylcarbamoyl-heterocyclic(lower)alkenoyl [e.g. methoxymethylcarbamoylpyridylacryloyl, methoxyethylcarbamoylpyridylacryloyl, methoxypropylcarbamoylpyridylacryloyl, ethoxyethylcarbamoylpyridylacryloyl, bis(methoxyethyl)carbamoylpyridylacryloyl, etc.], hydroxy(lower)alkylcarbamoyl-heterocyclic(lower)alkenoyl [e.g. hydroxymethylcarbamoylpyridylacryloyl, hydroxyethylcarbamoylpyridylacryloyl, bis(hydroxyethyl)carbamoylpyridylacryloyl, etc.], etc., heterocycliccarbonyl which may be substituted with substituent [e.g. furoyl, thenoyl, nicotinoyl, isonicotinoyl, morpholinocarbonyl, piperidinocarbonyl, 4-methyl-1-piperazinylcarbonyl, 4-ethyl-1-piperazinylcarbonyl, dimethylaminopiperidinocarbonyl, 4-methylcarbamoyl-1-piperazinylcarbonyl, 1,2,3,6-tetrahydropyridylcarbonyl, pyrrolidinylcarbonyl, indolylcarbonyl, etc.], aryloxy carbonyl which may be substituted with nitro [e.g. phenyloxycarbonyl, nitrophenyloxycarbonyl, etc.], ar(lower)alkoxy carbonyl which may be substituted with nitro [e.g. benzyloxycarbonyl, nitrobenzyloxycarbonyl, etc.], substituted or unsubstituted carbamoyl or thiocarbamoyl such as carbamoyl, lower alkylcarbamoyl [e.g. methylcarbamoyl, ethylcarbamoyl, propylcarbamoyl, isopropylcarbamoyl, butylcarbamoyl, isobutylcarbamoyl, tert-butylcarbamoyl, penty1carbamoyl, dimethylcarbamoyl, diethylcarbamoyl, N-ethyl-N-methylcarbamoyl, etc.],
carboxy(lower)alkylcarbamoyl [e.g. carboxymethylcarbamoyl, carboxyethylcarbamoyl, etc.], esterified
carboxy(lower)alkylcarbamoyl, for example, lower
alkoxy carbonyl(lower)alkylcarbamoyl [e.g.
  methoxycarbonylmethylcarbamoyl,
ethoxycarbonylmethylcarbamoyl,
ethoxycarbonylethylcarbamoyl, etc.], lower
alkenylcarbamoyl [e.g. vinylcarbamoyl, allylcarbamoyl,
etc.], cyclo(lower)alkylcarbamoyl [e.g.
cyclopropylcarbamoyl, cyclobutylcarbamoyl,
cyclopentylcarbamoyl, cyclohexylcarbamoyl, etc.],
halo(cyclo)lower alkanoylcarbamoyl [e.g.
  trichloroacetylcarbamoyl, etc.],
substituted or unsubstituted arylcarbamoyl, for example,
arylcarbamoyl [e.g. phenylcarbamoyl, tolylcarbamoyl,
  xylylcarbamoyl, naphthylcarbamoyl, ethylphenylcarbamoyl,
etc.], arylthiocarbamoyl [e.g. phenylthiocarbamoyl, etc.],
lower alkoxy-arylcarbamoyl [e.g. methoxyphenylcarbamoyl,
etc.], halo-arylcarbamoyl [e.g. fluorophenylcarbamoyl,
  chlorophenylcarbamoyl, etc.], halo(cyclo)lower alkanoyl-
arylc arbamoyl [e.g. trifluoromethylphenylcarbamoyl, etc.],
nitro-arylcarbamoyl [e.g. nitrophenylcarbamoyl, etc.],
cyano-arylcarbamoyl [e.g. cyanophenylcarbamoyl, etc.],
hydroxy(lower)alkyl-arylcarbamoyl [e.g. hydroxymethyl-
phenylcarbamoyl, hydroxyethylphenylcarbamoyl, etc.],
amino-arylcarbamoyl [e.g. aminophenylcarbamoyl, etc.],
lower alkanoylamino-arylcarbamoyl [e.g.
methylaninophenylcarbamoyl, ethylaninophenylcarbamoyl,
dimethylaninophenylcarbamoyl, etc.], lower
alkanoylamino-arylcarbamoyl [e.g.
  acetylaninophenylcarbamoyl, propionylaninophenylcarbamoyl,
etc.], N-(lower alkanoyl)-N-(lower
alkyl)amino-arylcarbamoyl [e.g.
N-acetyl-N-methylaninophenylcarbamoyl,
N-propionyl-N-methylaninophenylcarbamoyl, etc.],

lower alkoxy(lower)alkanoylamino-arylcarbamoyl [e.g. methoxyacetylaminophenylcarbamoyl, methoxypropionylaminophenylcarbamoyl, etc.], lower alkoxy carbonyl(lower)alkanoylamino-arylcarbamoyl [e.g. ethoxycarbonylacetylaminophenylcarbamoyl, methoxycarbonylpropionylaminophenylcarbamoyl, etc.], carboxyamino-arylcarbamoyl [e.g. carboxyaminophenylcarbamoyl, etc.], lower alkoxy carbonylamino-arylcarbamoyl [e.g. ethoxycarbonylaminophenylcarbamoyl, etc.], ureido-arylcarbamoyl [e.g. ureidophenylcarbamoyl, etc.], lower alkylureido-arylcarbamoyl [e.g. methylureido phenylcarbamoyl, ethylureido phenylcarbamoyl, etc.], hydroxyimino(lower)alkyl-arylcarbamoyl [e.g. hydroxyiminoethylphenylcarbamoyl, etc.], lower alkoxyimino(lower)alkyl-arylcarbamoyl [e.g. methoxyiminoethylphenylcarbamoyl, etc.], lower alkylhydrazono(lower)alkyl-arylcarbamoyl [e.g. methylhydrazonoethylphenylcarbamoyl, dimethylhydrazonoethylphenylcarbamoyl, etc.], optionally substituted heterocyclic-arylcarbamoyl [e.g. oxopyrroldinylphenylcarbamoyl, oxopiperidinophenylcarbamoyl, dioxopyrroldinylphenylcarbamoyl, oxooxazolidinylphenylcarbamoyl, pyrrolylphenylcarbamoyl, etc.], acyl-arylcarbamoyl, for example, carboxy-arylcarbamoyl [e.g. carboxyphenylcarbamoyl, etc.], lower alkoxy carbonyl-arylcarbamoyl [e.g. ethoxycarbonylphenylcarbamoyl, etc.], heterocyclic carbonyl-arylcarbamoyl [e.g. morpholinocarbonylphenylcarbamoyl, pyrroldinycarbonylphenylcarbamoyl, piperidinocarbonylphenylcarbamoyl, 1,2,3,6-tetrahydropyridylcarbonylphenylcarbamoyl, piperazinylcarbonylphenylcarbamoyl,
thiomorpholinocarbonylphenylcarbamoyl, etc.],
heterocycliccarbonyl-arylcarbamoyl substituted with lower alkyl [e.g. methylpiperazinylcarbonylphenylcarbamoyl, ethylpiperazinylcarbonylphenylcarbamoyl, etc.],
heterocycliccarbonyl-arylcarbamoyl substituted with aryl [e.g. phenylpiperazinylcarbonylphenylcarbamoyl, etc.],
heterocycliccarbonyl-arylcarbamoyl substituted with a heterocyclic group [e.g. pyridylpiperazinylcarbonylphenylcarbamoyl, etc.],
heterocycliccarbonyl-arylcarbamoyl substituted with lower alkanoyl [e.g. acetylpiperazinylcarbonylphenylcarbamoyl, etc.],
heterocycliccarbonyl-arylcarbamoyl substituted with lower alkoxyacyl [e.g. ethoxycarbonylpiperazinylcarbonylphenylcarbamoyl, etc.],
heterocycliccarbonyl-arylcarbamoyl substituted with lower alkylamino [e.g. methylaminopiperazinylcarbonylphenylcarbamoyl, dimethylaminopiperidinocarbonylphenylcarbamoyl, etc.],
heterocycliccarbonyl-arylcarbamoyl substituted with lower alkylnocarbamoyl [e.g. carbamoylphenylcarbamoyl, etc.],
carbamoyl-arylcarbamoyl [e.g. carbamoylphenylcarbamoyl, etc.], lower alkylcarbamoyl-arylcarbamoyl [e.g. methylcarbamoylphenylcarbamoyl, ethylcarbamoylphenylcarbamoyl, dimethylcarbamoylphenylcarbamoyl, diethylcarbamoylphenylcarbamoyl, N-ethyl-N-methylcarbamoylphenylcarbamoyl, N-isopropyl-N-methylcarbamoylphenylcarbamoyl, etc.],
hydroxy(lower)alkylcarbamoyl-arylcarbamoyl [e.g. hydroxymethylcarbamoylphenylcarbamoyl, hydroxyethylcarbamoylphenylcarbamoyl, bis(hydroxyethyl)carbamoylphenylcarbamoyl, etc.],
N-[hydroxy(lower)alkyl]-N-(lower alkyl)carbamoyl-arylcarbamoyl [e.g. N-(hydroxyethyl)-N-methylcarbamoylphenylcarbamoyl, etc.],
lower alkoxy(lower)alkylcarbamoyl-arylcarbamoyl [e.g. methoxymethylcarbamoylphenylcarbamoyl,
methoxyethylcarbamoylphenylcarbamoyl, bis(methoxyethyl)carbamoylphenylcarbamoyl,
10 bis(ethoxyethyl)carbamoylphenylcarbamoyl, etc.),
N-[lower alkoxy(lower)alkyl]-N-(lower alkyl)carbamoyl-arylcarbamoyl [e.g.
15 N-(methoxyethyl)-N-methylcarbamoylphenylcarbamoyl,
N-(methoxypropyl)-N-methylcarbamoylphenylcarbamoyl, etc.],
lower alkylamino(lower)alkylcarbamoyl-arylcarbamoyl [e.g.
methylaminoethylcarbamoylphenylcarbamoyl,
dimethylaminoethylcarbamoylphenylcarbamoyl, etc.],
15 N-[lower alkylamino(lower)alkyl]-N-(lower alkyl)carbamoyl-arylcarbamoyl [e.g. N-(dimethylaminoethyl)-N-methylcarbamoylphenylcarbamoyl,
N-(dimethylaminopropyl)-N-methylcarbamoylphenylcarbamoyl, etc.], heterocycliccarbamoyl-arylcarbamoyl [e.g. morpholinylcarbamoylphenylcarbamoyl,
thienylcarbamoylphenylcarbamoyl,
10 pyridylcarbamoylphenylcarbamoyl,
pyrimidinylcarbamoylphenylcarbamoyl, etc.],
N-(heterocyclic)-N-(lower alkyl)carbamoyl-arylcarbamoyl
20 [e.g. N-pyridyl-N-methylcarbamoylphenylcarbamoyl, etc.],
heterocyclic(lower)alkylcarbamoyl-arylcarbamoyl [e.g.
pyridylmethylcarbamoylphenylcarbamoyl,
pyrimidinylcarbamoylphenylcarbamoyl,
25 thienylmethylcarbamoylphenylcarbamoyl, etc.],
N-[heterocyclic(lower)alkyl]-N-(lower alkyl)carbamoyl-arylcarbamoyl [e.g.
N-pyridylmethyl-N-methylcarbamoylphenylcarbamoyl, etc.],
N-[heterocyclic(lower)alkyl]-N-[lower alkoxy(lower)alkyl]-arylcarbamoyl-arylcarbamoyl [e.g.
20 N-pyridylmethyl-N-methoxyethylcarbamoylphenylcarbamoyl, etc.] arylcarbamoyl-arylcarbamoyl [e.g.
phenylcarbamoylphenylcarbamoyl, etc.], lower
30 alkylamino-arylcarbamoyl-arylcarbamoyl [e.g.
dimethylaminophenylcarbamoylphenylcarbamoyl, etc.],
lower alkanoyl-arylcarnbamoyl [e.g. acetylphenylcarbamoyl, propionylphenylcarbamoyl, etc.], etc., etc., ar(lower)alkylcarbamoyl [e.g. benzylcarbamoyl, phenethylcarbamoyl, etc.], heterocycliccarbamoyl [e.g. furylcarbamoyl, thiencarbarbamoyl, pyridylcarbamoyl, quinolylcarbamoyl, isoquinolylcarbamoyl, pyrimidinylcarbamoyl, pyrazolylcarbamoyl, etc.], heterocyclic(lower)alkylcarbamoyl [e.g. pyridylmethy3carbamoyl, pyridylethylcarbamoyl, thienylmethylcarbamoyl, thienylethylcarbamoyl, etc.], arylaminocarbamoyl [e.g. phenylaminocarbamoyl, etc.], arylcarbamoyl [e.g. benzoylcarbamoyl, etc.], lower alkylsulfonyl [e.g. mesyl, ethylsulfonyl, propylsulfonyl, isopropylsulfonyl, tert-butylsulfonyl, pentylsulfonyl, etc.], arylsulfonyl [e.g. tosyl, phenylsulfonyl, etc.], ar(lower)alkylsulfonyl [e.g. benzylsulfonyl, phenethylsulfonyl, etc.], ar(lower)alkenylsulfonyl [e.g. styrylsulfonyl, cinnamylsulfonyl, etc.], phthaloyl, substituted or unsubstituted amino acid residue mentioned below, or the like.

Suitable "amino acid residue" may include natural or artificial ones, and such amino acid may be glycine, sarcosine, alanine, β-alanine, valine, norvaline, leucine, isoleucine, norleucine, serine, threonine, cysteine, methionine, phenylalanine, phenylglycine, tryptophan, tyrosine, proline, hydroxyproline, glutamic acid, aspartic acid, glutamine, asparagine, lysine, arginine, histidine, ornithine, or the like, in which more preferable one is glycine, sarcosine, alanine, β-alanine and proline, and the most preferable one is glycine. And said amino acid residue may be substituted with suitable substituent(s) such as the above-mentioned lower alkyl, the above-mentioned aryl, the above-mentioned acyl, ar(lower)alkyl [e.g. benzyl, phenethyl, trityl, etc.], cycloalkyl [e.g. cyclopropyl, cyclobutyl, cyclopentyl,
cyclohexyl, cycloheptyl, cyclooctyl, adamantyl, etc.], a heterocyclic group mentioned below, heterocyclic(lower)alkyl [e.g. pyridylmethyl, pyridylethyl, imidazolylmethyl, furylmethyl, thienylmethyl, morpholinomethyl, piperidinomethyl, etc.], substituted or unsubstituted amidino [e.g. amidino, methylamidino, N-ethyl-N'-cyanoamidino, etc.], or the like.

Preferred example of said amino acid residue substituted with suitable substituent(s) may be amino acid residue substituted with lower alkyl [e.g. ethylglycyl, isopropylglycyl, dimethylglycyl, diethylglycyl, ethylsarcosyl, isopropylsarcosyl, methylalanyl, methyl-β-alanyl, dimethyl-β-alanyl, etc.], amino acid residue substituted with aryl [e.g. N-phenylglycyl, N-tolylglycyl, N-phenylalanyl, N-phenylsarcosyl, etc.], amino acid residue substituted with ar(lower)alkyl [e.g. benzylglycyl, tritylglycyl, phenethylglycyl, benzylsarcosyl, benzylalanyl, etc.], amino acid residue substituted with a heterocyclic group [e.g. morpholinoglycyl, piperidinoglycyl, pyridylglycyl, etc.], amino acid residue substituted with heterocyclic(lower)alkyl [e.g. pyridylmethylglycyl, imidazolylmethylglycyl, furylmethylglycyl, thienylmethylglycyl, etc.], amino acid residue substituted with cycloalkyl [e.g. cyclopropylglycyl, cyclobutylglycyl, cyclopentylglycyl, cyclohexylglycyl, cycloheptylglycyl, cyclooctylglycyl, adamantylglycyl, cyclohexylsarcosyl, cycloheptylsarcosyl, cyclohexylalanyl, etc.], amino acid residue substituted with optionally substituted amidino [e.g. amidinoglycyl, methylamidinoglycyl, N-ethyl-N'-cyanoamidinoglycyl, etc.], amino acid residue substituted with acyl such as amino acid residue substituted with alkanoyl [e.g. formylglycyl, acetylglycyl, acetylsarcosyl, acetylalanyl,
acetyl-β-alanyl, propionylglycyl, butyrylglycyl, isobutyrylglycyl, valerylglcyyl, isovalerylglcyyl, pivaloylglycyl, hexanoylglycyl, heptanoylglycyl, etc.), amino acid residue substituted with halo(lower)alkanoyl [e.g. trifluoroacetetylglcyyl, trifluoroacetylalsarcosyl, trifluoroacetylalanyl, bromoacetylglcyyl, heptafluorobutyrylglycyl, etc.], amino acid residue substituted with hydroxy(lower)alkanoyl [e.g. glycoloylglycyl, glycoloylsarcosyl, lactoylglycyl, lactoylalanyl, etc.], amino acid residue substituted with lower alkylsulfonyloxy(lower)alkanoyl [e.g. mesyloxyacetylglcyyl, ethylsulfonyloxyacetetylglcyyl, mesyloxyacetylalsarcosyl, etc.], amino acid residue substituted with lower alkoxy(lower)alkanoyl [e.g. methoxyacetetylglcyyl, ethoxyacetetylglcyyl, methoxyacetylalsarcosyl, methoxypropionylalanyl, etc.], amino acid residue substituted with aryloxy(lower)alkanoyl [e.g. phenyloxyacetetylglcyyl, phenyloxypropionylglcyyl, phenyloxyacetylalsarcosyl, etc.], amino acid residue substituted with lower alkylthio(lower)alkanoyl [e.g. methylthioacetylglcyyl, methylthiopropionylglcyyl, etc.], amino acid residue substituted with lower alkylcarbamoyl(lower)alkanoyl [e.g. methylcarbamoylpropionylglcyyl, methylcarbamoylpropionylalanyl, etc.], amino acid residue substituted with lower alkanoyloxy(lower)alkanoyl [e.g. acetyloxyacetetylglcyyl, acetyloxyacetylalsarcosyl, propionyloxyacetylglcyyl, acetyloxypropionylalanyl, etc.], amino acid residue substituted with carboxy(lower)alkanoyl [e.g. carboxyacetetylglcyyl, carboxypropionylglcyyl, carboxypropionylsarcosyl, carboxyacetylalanyl, etc.], amino acid residue substituted with lower alkoxy carbonyl(lower)alkanoyl [e.g. methoxycarbonylacetylglcyyl, ethoxycarbonylpropionylglcyyl, methoxycarbonylacetylalsarcosyl, etc.], amino acid residue substituted with ar(lower)alkanoyl [e.g.
phenylacetylglycyl, phenylacetylsarcosyl, phenylpropionylalanyl, phenylpropionylglycyl, naphthylacetylglycyl, phenylbutyrylglycyl, etc.), amino acid residue substituted with optionally substituted heterocyclic(lower)alkanoyl [e.g. morpholinoacetylglglycyl, thiomorpholinoacetylglglycyl, its oxide or dioxide, pyridylacetylglycyl, morpholinopropionylalanyl, imidazolylacetylglycyl, piperidinoacetylglglycyl, pyrrolidinylacetylglycyl, hexamethyleneiminoacetylglglycyl, methylpiperazinylacetylglycyl, pyridylpiperazinylacetylglycyl, etc.], amino acid residue substituted with lower alkenoyl [e.g. acryloylglycyl, crotonoylglycyl, 3-pentenoylglycyl, 3-butenoylglycyl, 4-pentenoylglycyl, 3-butenoylsarcosyl, etc.], amino acid residue substituted with ar(lower)alkenoyl [e.g. cinnamoylglycyl, allocinnamoylglycyl, a-methylcinnamoylglycyl, 4-methylcinnamoylglycyl, cinnamoylsarcosyl, etc.], amino acid residue substituted with lower alkoxy-ar(lower)alkenoyl [e.g. methoxycinnamoylglycyl, ethoxycinnamoylglycyl, dimethoxycinnamoylglycyl, etc.], amino acid residue substituted with lower alkylendioxy-ar(lower)alkenoyl [e.g. methylenedioxyacetylglglycyl, ethylenedioxyacetylglglycyl, etc.], amino acid residue substituted with nitro-ar(lower)alkenoyl [e.g. nitrocinnamoylglycyl, etc.], amino acid residue substituted with halo-ar(lower)alkenoyl [e.g. chlorocinnamoylglycyl, fluorocinnamoylglycyl, etc.], amino acid residue substituted with hydroxy-ar(lower)alkenoyl [e.g. hydroxycinnamoylglycyl, etc.], amino acid residue substituted with hydroxy(lower)alkoxy-ar(lower)alkenoyl [e.g. hydroxyethoxycinnamoylglycyl, etc.], amino acid residue substituted with amino(lower)alkoxy-ar(lower)alkenoyl [e.g. aminoethoxycinnamoylglycyl, etc.],
amino acid residue substituted with lower alkylamino-(lower)alkoxy-ar-(lower)alkenoyl [e.g. methylaminomethoxycinnamoylglycyl, dimethylaminoethoxycinnamoylglycyl, etc.],
amino acid residue substituted with heterocyclic-(lower)alkoxy-ar-(lower)alkenoyl [e.g. pyridylmethoxycinnamoylglycyl, etc.],
amino acid residue substituted with optionally substituted heterocyclic-ar-(lower)alkenoyl [e.g. morpholinocinnamoylglycyl, methylpiperazinylcinnamoylglycyl, pyrrolidinyl-cinnamoylglycyl, oxopyrrolidinylcinnamoylglycyl, oxopiperidinocinnamoylglycyl, dioxopyrrolidinyl-cinnamoylglycyl, oxooxazolidinylcinnamoylglycyl, pyrrolylcinnamoylglycyl, etc.],
amino acid residue substituted with amino-ar-(lower)alkenoyl [e.g. aminocinnamoylglycyl, etc.],
amino acid residue substituted with lower alkylamino-ar-(lower)alkenoyl [e.g. methylaminocinnamoylglycyl, dimethylaminocinnamoylglycyl, etc.],
amino acid residue substituted with acylamino-ar-(lower)alkenoyl, for example,
amino acid residue substituted with lower alkanoylamino-ar-(lower)alkenoyl [e.g. acetylaminocinnamoylglycyl, propionylaminocinnamoylglycyl, etc.],
amino acid residue substituted with hydroxy-(lower)alkanoylamino-ar-(lower)alkenoyl [e.g. hydroxyacetylaminocinnamoylglycyl, hydroxypropionylaminocinnamoylglycyl, etc.],
amino acid residue substituted with lower alkoxy-(lower)alkanoylamino-ar-(lower)alkenoyl [e.g. methoxyacetylaminocinnamoylglycyl, methoxypropionylaminocinnamoylglycyl, etc.],
amino acid residue substituted with halo-(lower)alkanoylamino-ar-(lower)alkenoyl [e.g. chloroacetylaminocinnamoylglycyl,
bromobutyrylaminocinnamoyl glycyl, etc.),
amino acid residue substituted with
amino(lower) alkanoylamino-ar(lower) alkenoyl [e.g. aminoacetylaminocinnamoyl glycyl,
aminopropionylaminocinnamoylglycyl, etc.],
amino acid residue substituted with lower
alkylamino(lower) alkanoylamino-ar(lower) alkenoyl [e.g. methylaminoacetylaminocinnamoylglycyl,
aminopropionylaminocinnamoylglycyl, etc.],
amino acid residue substituted with lower
alkanoylamino(lower) alkanoylamino-ar(lower) alkenoyl [e.g. acetylaminoacetylaminocinnamoylglycyl,
aminopropionylaminocinnamoylglycyl, etc.],
amino acid residue substituted with
carboxy(lower) alkanoylamino-ar(lower) alkenoyl [e.g. carboxyacetylaminocinnamoyl glycyl,
carboxypropionylaminocinnamoylglycyl, etc.],
amino acid residue substituted with lower
alkoxycarbonyl(lower) alkanoylamino-ar(lower) alkenoyl [e.g. ethoxycarbonylacetylaminocinnamoylglycyl,
ethoxycarbonylpropionylaminocinnamoylglycyl, etc.],
amino acid residue substituted with lower
alkoxycarbonyl(lower) alkenoylamino-ar(lower) alkenoyl [e.g. ethoxycarbonylacryloylaminocinnamoylglycyl, etc.],
amino acid residue substituted with
halo(lower) alkoxy carbonylamino-ar(lower) alkenoyl [e.g. chloroethoxycarbonylaminocinnamoylglycyl, etc.],
amino acid residue substituted with
heterocyclic(lower) alkanoylamino-ar(lower) alkenoyl [e.g. pyridylacetylaminocinnamoylglycyl, etc.],
amino acid residue substituted with
aro lamino-ar(lower) alkenoyl [e.g. benzoylaminoacetylaminocinnamoylglycyl, etc.],
amino acid residue substituted with
heterocyclic carbonylamino-ar(lower) alkenoyl [e.g. pyridylacetylaminocinnamoylglycyl, etc.],
amino acid residue substituted with
nicotinoylaminocinnamoylglycyl,
isonicotinoylaminocinnamoylglycyl,
morpholinocarbonylaminocinnamoylglycyl, etc.),
amino acid residue substituted with lower
alkylsulfonylamino-ar(lower)alkenoyl [e.g.
mesylaminocinnamoylglycyl,
ethylsulfonylaminocinnamoylglycyl, etc.], etc.,
amino acid residue substituted with N-(lower
alkanoyl)-N-(lower alkyl)amino-ar(lower)alkenoyl [e.g.
N-acetyl-N-methylaminocinnamoylglycyl,
N-acetyl-N-ethylaminocinnamoylglycyl,
N-propionyl-N-methylaminocinnamoylglycyl], etc.),
amino acid residue substituted with N-[lower
alkoxy(lower)alkanoyl]-N-(lower alkyl)amino-
ar(lower)alkenoyl [e.g. N-methoxyacetyl-N-
methylnaminocinnamoylglycyl, N-methoxypropionyl-N-
 methylnaminocinnamoylglycyl, etc.],
amino acid residue substituted with N-(lower
[ heterocyclic(lower)alkyl]amino-ar(lower)alkenoyl [e.g.
N-acetyl-N-pyridylmethylaminocinnamoylglycyl, etc.],
amino acid residue substituted with ureido-
ar(lower)alkenoyl [e.g. ureidocinnamoylglycyl, etc.],
amino acid residue substituted with lower
alkylureido-ar(lower)alkenoyl [e.g.
methylureidocinnamoylglycyl, ethylureidocinnamoylglycyl,
dimethylureidocinnamoylglycyl], etc.),
amino acid residue substituted with heterocyclicureido-ar(lower)alkenoyl [e.g.
pyridylureidocinnamoylglycyl,
pyrimidinylureidocinnamoylglycyl,
thienylureidocinnamoylglycyl, etc.],
amino acid residue substituted with acyl-ar(lower)-
alkenoyl, for example, amino acid residue substituted with
lower alkanoyl-ar(lower)alkenoyl [e.g.
formylcinnamoylglycyl, acetylcinnamoylglycyl,
propionylcinnamoylglycyl, etc.,
amino acid residue substituted with carboxy-ar(lower)-
alkenoyl [e.g. carboxycinnamoylglycyl, etc.],
amino acid residue substituted with lower alkoxy carbonyl-ar(lower) alkenoyl [e.g.
methoxycarbonylcinnamoylglycyl,
ethoxycarbonylcinnamoylglycyl, etc.],
amino acid residue substituted with carbamoyl-ar(lower)-
alkenoyl [e.g. carbamoylcinnamoylglycyl, etc.],
amino acid residue substituted with lower alkyl carbamoyl-ar(lower) alkenoyl [e.g.
methylcarbamoylcinnamoylglycyl,
ethylcarbamoylcinnamoylglycyl,
dimethylcarbamoylcinnamoylglycyl,
propylcarbamoylcinnamoylglycyl,
isopropylcarbamoylcinnamoylglycyl,
diethylcarbamoylcinnamoylglycyl,
N-methyl-N-ethylcarbamoylcinnamoylglycyl, etc.],
amino acid residue substituted with hydroxy(lower) alkyl carbamoyl-ar(lower) alkenoyl [e.g.
hydroxyethylcarbamoylcinnamoylglycyl,
bis(hydroxyethyl) carbamoylcinnamoylglycyl, etc.],
amino acid residue substituted with N-[hydroxy(lower) alkyl]-N-(lower alkyl) carbamoyl-ar(lower) alkenoyl [e.g.
N-hydroxyethyl-N-methylcarbamoylcinnamoylglycyl, etc.],
amino acid residue substituted with lower alkoxy(lower) alkyl carbamoyl-ar(lower) alkenoyl [e.g.
methoxyethylcarbamoylcinnamoylglycyl, bis(methoxyethyl)-carbamoylcinnamoylglycyl,
ethoxyethylcarbamoylcinnamoylglycyl,
methoxypropylcarbamoylcinnamoylglycyl,
bis(ethoxyethyl) carbamoylcinnamoylglycyl, etc.],
amino acid residue substituted with N-[lower
alkoxy(lower)alkyl]-N-(lower alkyl)carbamoyl-ar(lower)alkenoyl [e.g. N-methoxyethyl-N-methylcarbamoylcinnamoylglycyl, etc.], amino acid residue substituted with heterocycliccarbamoyl-ar(lower)alkenoyl [e.g. morpholinylcarbamoylcinnamoylglycyl, thienylcarbamoylcinnamoylglycyl, pyridylcarbamoylcinnamoylglycyl, pyrimidinylcarbamoylcinnamoylglycyl, etc.], amino acid residue substituted with heterocycliccarbonyl-ar(lower)alkenoyl [e.g. morpholinocarbonylcinnamoylglycyl, pyrrolidinocarbonylcinnamoylglycyl, piperidinocarbonylcinnamoylglycyl, etc.], etc., amino acid residue substituted with ar(lower)alkynoyl [e.g. phenylpropioloylglycyl, etc.], amino acid residue substituted with heterocyclic(lower)alkenoyl [e.g. morpholinylacryloylglycyl, pyridylacryloylglycyl, thienylacryloylglycyl, etc.], amino acid residue substituted with amino-heterocyclic(lower)alkenoyl [e.g. aminopyridylacryloylglycyl, etc.], amino acid residue substituted with lower alkylamino-heterocyclic(lower)alkenoyl [e.g. methylvaminopyridylacryloylglycyl, dimethylaminopyridylacryloylglycyl, etc.], amino acid residue substituted with acylamino-heterocyclic(lower)alkenoyl, for example, amino acid residue substituted with lower alkanoylamino-heterocyclic(lower)alkenoyl [e.g. acetylaminopyridylacryloylglycyl, propionylaminopyridylacryloylglycyl, etc.], etc., amino acid residue substituted with lower alkyureido-heterocyclic(lower)alkenoyl [e.g. methylureidopyridylacryloylglycyl, etc.], amino acid residue substituted with acyl-heterocyclic(lower)alkenoyl, for example, amino acid residue substituted with carboxy-heterocyclic(lower)alkenoyl [e.g. carboxypyrrolidylacryloylglycyl, etc.], amino acid residue substituted with lower
alkoxycarbonyl-heterocyclic(lower)alkenoyl [e.g. ethoxycarbonylpyridylacryloylglycyl, etc.],
amino acid residue substituted with lower alkylcarbamoyl-heterocyclic(lower)alkenoyl [e.g. methylcarbamoylpyridylacryloylglycyl, ethylcarbamoylpyridylacryloylglycyl, dimethylcarbamoylpyridylacryloylglycyl, diethylcarbamoylpyridylacryloylglycyl, isopropylcarbamoylpyridylacryloylglycyl, etc.], amino acid residue substituted with lower alkoxy(lower)alkylcarbamoyl-heterocyclic(lower)alkenoyl [e.g. methoxymethylcarbamoylpyridylacryloylglycyl, methoxyethylcarbamoylpyridylacryloylglycyl, methoxypropylcarbamoylpyridylacryloylglycyl, ethoxyethylcarbamoylpyridylacryloylglycyl, bis(methoxyethyl)carbamoylpyridylacryloylglycyl, etc.], amino acid residue substituted with hydroxy(lower)alkylcarbamoyl-heterocyclic(lower)alkenoyl [e.g. hydroxymethylcarbamoylpyridylacryloylglycyl, hydroxyethylcarbamoylpyridylacryloylglycyl, bis(hydroxyethyl)carbamoylpyridylacryloylglycyl, etc.], etc., amino acid residue substituted with heterocyclicthio(lower)alkanoyl [e.g. pyridylthioacetylglucyl, pyrimidylinylthioacetylglucyl, imidazolylthiopropionylglucyl, etc.], amino acid residue substituted with optionally substituted heterocycliccarbonyl [e.g. morpholinocarbonylglucyl, indolylcarbonylglucyl, 4-methyl-1-piperazinyl-carbonylglucyl, etc.], amino acid residue substituted with cyclo(lower)alkylcarbonyl [e.g. cyclopropylcarbonylglucyl, cyclopentylcarbonylglucyl, cyclohexylcarbonylglucyl, cyclohexylcarbonylsarcosyl, etc.], amino acid residue substituted with lower alkoxy carbonyl [e.g. methoxycarbonylglucyl, tert-butoxycarbonylglucyl, tert-butoxycarbonylsarcosyl, tert-butoxycarbaonyllalanyl, etc.], amino acid residue substituted with aryloxycarbonyl [e.g. phenoxy carbonylglucyl, etc.], amino
acid residue substituted with aroyl(lower)alkanoyl [e.g. phenyloxalylglycyl, benzoylpropionylglycyl, etc.], amino acid residue substituted with aroyl [e.g. benzoylglycyl, naphtooylglycyl, benzoylsarcosyl, benzoylalanyl, etc.], amino acid residue substituted with nitro-aryloxycarbonyl [e.g. nitrophenyloxycarbonylglycyl, etc.], amino acid residue substituted with carbamoyl [e.g. carbamoylglycyl, carbamoylalanyl, carbamoylsarcosyl, carbamoyl-β-alanyl, etc.], amino acid residue substituted with lower alkylcarbamoyl [e.g. methylcarbamoylglycyl, ethylcarbamoylglycyl, propylcarbamoylglycyl, isopropylcarbamoylglycyl, methylcarbamoylsarcosyl, ethylcarbamoylalanyl, isopropylcarbamoyl-β-alanyl, pentaoylcarbamoylglycyl, etc.], amino acid residue substituted with lower alkoxy carbonyl(lower)alkylcarbamoyl [e.g. methoxycarbonylmethylcarbamoylglycyl, ethoxycarbonylmethylcarbamoylglycyl, etc.], amino acid residue substituted with lower alkenylcarbamoyl [e.g. vinylcarbamoylglycyl, allylcarbamoylglycyl, allylcarbamoylsarcosyl, etc.], amino acid residue substituted with cyclo(lower)alkylcarbamoyl [e.g. cyclopropylcarbamoylglycyl, cyclohexylcarbamoylglycyl, cyclohexylcarbamoylsarcosyl, etc.], amino acid residue substituted with arylcarbamoyl [e.g. phenylcarbamoylglycyl, naphthylcarbamoylglycyl, tolylcarbamoylglycyl, ethylphenylcarbamoylglycyl, phenylcarbamoylalanyl, phenylcarbamoylsarcosyl, etc.], amino acid residue substituted with lower alkoxy-arylcarbamoyl [e.g. methoxyphenylcarbamoylglycyl, ethoxyphenylcarbamoylglycyl, methoxyphenylcarbamoylalanyl, etc.], amino acid residue substituted with halo(lower)alkyl-arylcarbamoyl [e.g. trifluoromethylphenylcarbamoylglycyl, trifluoromethylphenylcarbamoylalanyl, trifluoromethylphenylcarbamoylsarcosyl, etc.], amino acid residue substituted with halo-arylcarbamoyl [e.g.
chlorophenylcarbamoylglycyl, fluorophenylcarbamoylglycyl, fluorophenylcarbamoylalanyl, etc.), amino acid residue substituted with hydroxy(lower)alkyl-arylcarbamoyl [e.g. hydroxymethylphenylcarbamoylglycyl, hydroxyethylphenylcarbamoylglycyl, hydroxyethylphenylcarbamoylalanyl, etc.], amino acid residue substituted with nitro-arylcarbamoyl [e.g. nitrophenylcarbamoylglycyl, etc.], amino acid residue substituted with cyano-arylcarbamoyl [e.g. cyanophenylcarbamoylglycyl, etc.], amino acid residue substituted with amino-arylcarbamoyl [e.g. aminophenylcarbamoylglycyl, etc.], amino acid residue substituted with lower alkylamino-arylcarbamoyl [e.g. methylaminophenylcarbamoylglycyl, ethylaminophenylcarbamoylglycyl, dimethyaminophenylcarbamoylglycyl, etc.], amino acid residue substituted with lower alkanoylamino-arylcarbamoyl [e.g. acetylamino-arylcarbamoylglycyl, propionylamino-arylcarbamoylglycyl, etc.], amino acid residue substituted with N-(lower alkanoyl)-N-(lower alkyl)amino-arylcarbamoyl [e.g. N-acetyl-N-methylnaphenylcarbamoylglycyl, N-propionyl-N-methylnaphenylcarbamoylglycyl, etc.], amino acid residue substituted with lower alkoxy(lower)alkanoylamino-arylcarbamoyl [e.g. methoxyacetylamino-arylcarbamoylglycyl, methoxypropionylaminophenylcarbamoylglycyl, etc.], amino acid residue substituted with lower alkoxy carbonyl(lower)alkanoylamino-arylcarbamoyl [e.g. ethoxycarbonylacetylamino-arylcarbamoylglycyl, methoxycarbonylpropionylaminophenylcarbamoylglycyl, etc.], amino acid residue substituted with carboxyamino-arylcarbamoyl [e.g. carboxyaminophenylcarbamoylglycyl, etc.], amino acid residue substituted with lower alkoxy carbonyl-amino-arylcarbamoyl [e.g. ethoxycarbonylaminophenyl-
carbamoylglycyl, etc.], amino acid residue substituted with ureido-arylcarbamoyl [e.g. ureidophenylcarbamoylglycyl, etc.], amino acid residue substituted with lower alkylureido-arylcarbamoyl [e.g. methylureidophenylcarbamoylglycyl, ethylureidophenylcarbamoylglycyl, etc.], amino acid residue substituted with hydroxyimino(lower)alkyl-arylcarbamoyl [e.g. hydroxyiminoethylphenylcarbamoylglycyl, etc.], amino acid residue substituted with lower alkoxyimino(lower)alkyl-arylcarbamoyl [e.g. methoxyiminoethylphenylcarbamoylglycyl, dimethoxyiminoethoxyphenylcarbamoylglycyl, etc.], amino acid residue substituted with lower alkylhydrazono(lower)alkyl-arylcarbamoyl [e.g. methylhydrazonoethylphenylcarbamoylglycyl, dimethylhydrazonoethoxyphenylcarbamoylglycyl, etc.], amino acid residue substituted with optionally substituted heterocyclic-arylcarbamoyl [e.g. oxopyrrolidinylphenylcarbamoylglycyl, oxopiperidinophenylcarbamoylglycyl, dioxopyrrolidinylphenylcarbamoylglycyl, oxooxazolidinylphenylcarbamoylglycyl, pyrrolylphenylcarbamoylglycyl, etc.], amino acid residue substituted with acyl-arylcarbamoyl, for example, amino acid residue substituted with lower alkanoyl-arylcarbamoyl [e.g. acetylphenylcarbamoylglycyl, propionylphenylcarbamoylglycyl, etc.], amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl [e.g. morpholinocarbonylphenylcarbamoylglycyl, piperidinocarbonylphenylcarbamoylglycyl, piperazinylcarbonylphenylcarbamoylglycyl, thiomorpholinocarbonylphenylcarbamoylalanyl, pyrrolidinycarbonylphenylcarbamoylglycyl, 1,2,3,6-tetrahydropyridylcarbonylphenylcarbamoylglycyl, etc.], amino acid residue substituted with carboxy-arylcarbamoyl [e.g. carboxyphenylcarbamoylglycyl, etc.], amino acid residue substituted with lower
alkoxycarbonyl-arylcarbamoyl [e.g.
methoxycarbonylphenylcarbamoylglycyl, ethoxycarbonyl-
phenylcarbamoylglycyl, etc.], amino acid residue
substituted with lower alkylcarbamoyl-arylcarbamoyl [e.g.
methylcarbamoylphenylcarbamoylglycyl,
dimethylcarbamoylphenylcarbamoylglycyl,
diethylcarbamoylphenylcarbamoylglycyl,
N-ethyl-N-methylcarbamoylphenylcarbamoylglycyl,
N-isopropyl-N-methylcarbamoylphenylcarbamoylglycyl, etc.],
5 amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having lower alkyl
[e.g. methylpiperazinylcarbonylphenylcarbamoylglycyl,
ethylpiperazinylcarbonylphenylcarbamoylglycyl, etc.],
amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having aryl [e.g.
phenylpiperazinylcarbonylphenylcarbamoylglycyl, etc.],
amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having a heterocyclic
group [e.g. pyridylpiperazinylcarbonylphenylcarbamoylglycyl, etc.], amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having lower alkanoyl
[e.g. acetylpiperazinylcarbonylphenylcarbamoylglycyl,
etc.], amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having lower
alkoxycarbonyl [e.g. ethoxycarbonylpiperazinylcarbonyl-
phenylcarbamoylglycyl, etc.], amino acid residue
substituted with heterocycliccarbonyl-arylcarbamoyl having lower alkylamino [e.g. methylaminopiperazinylcarbonyl-
phenylcarbamoylglycyl, dimethylaminopiperidinocarbonyl-
phenylcarbamoylglycyl, etc.],
amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having lower alkylicarbamoyl [e.g. methylcarbamoylpiperazinylcarbonylphenylcarbamoylglycyl, etc.],
amino acid residue substituted with hydroxy(lower)alkylcarbamoyl-arylcarbamoyl [e.g.
hydroxymethylcarbamoylphenylcarbamoylglycyl,
hydroxyethylcarbamoylphenylcarbamoylglycyl,
bis(hydroxyethyl)carbamoylphenylcarbamoylglycyl, etc.],
amino acid residue substituted with
5 N-[hydroxy(lower)alkyl]-N-(lower alkyl)carbamoyl-
arylcarmamoyl [e.g. N-(hydroxyethyl)-N-methylcarbamoyl-
phenylcarbamoylglycyl, etc.], amino acid residue
substituted with lower alkoxy(lower)alkylcarbamoyl-aryl-
carbamoyl [e.g. methoxymethylcarbamoylphenylcarbamoyl-
glycyl, methoxyethylcarbamoylphenylcarbamoylglycyl,
bis(methoxyethyl)carbamoylphenylcarbamoylglycyl,
bis(ethoxyethyl)carbamoylphenylcarbamoylglycyl, etc.],
amino acid residue substituted with N-[lower alkoxy-
(lower)alkyl]-N-(lower alkyl)carbamoyl-arylcarmamoyl [e.g.
N-(methoxyethyl)-N-methylcarbamoylphenylcarbamoylglycyl,
N-(methoxypropyl)-N-methylcarbamoylphenylcarbamoylglycyl,
etc.], amino acid residue substituted with lower alkylamino(lower)alkylcarbamoyl-arylcarmamoyl [e.g.
methylaminoethylcarbamoylphenylcarbamoylglycyl,
dimethylaminoethylcarbamoylphenylcarbamoylglycyl, etc.],
amino acid residue substituted with N-[lower alkylamino-
(lower)alkyl]-N-(lower alkyl)carbamoyl-arylcarmamoyl [e.g. N-(dimethylaminoethyl)-N-
methylcarbamoylphenylcarbamoylglycyl,
N-(dimethylaminopropyl)-N-methylcarbamoylphenylcarbamoyl-
glycyl, etc.], amino acid residue substituted with
heterocycliccarbamoyl-arylcarmamoyl [e.g. morpholinylnlcarbamoylphenylcarbamoylglycyl,
thienylcarbamoylphenylcarbamoylglycyl,
pyridylcarbamoylphenylcarbamoylglycyl,
pyrimidinylcarbamoylphenylcarbamoylglycyl, etc.],
amino acid residue substituted with N-(heterocyclic)-
N-(lower alkyl)carbamoyl-arylcarmamoyl [e.g.
N-pyridyl-N-methylcarbamoylphenylcarbamoylglycyl, etc.],
amino acid residue substituted with
heterocyclic(lower)alkylcarbamoyl-arylcarbamoyl [e.g. pyridylmethylcarbamoylphenylcarbamoylglycyl, pyridylethylcarbamoylphenylcarbamoylglycyl, thienylmethylcarbamoylphenylcarbamoylglycyl, etc.],

5 amino acid residue substituted with
N-[heterocyclic(lower)alkyl]-N-(lower alkyl)carbamoyl-arylcarbamoyl [e.g. N-pyridylmethyl-N-methylcarbamoylphenylcarbamoylglycyl, etc.],

amino acid residue substituted with
10 N-[heterocyclic(lower)alkyl]-N-[lower alkoxy(lower)alkyl]carbamoyl-arylcarbamoyl [e.g. N-pyridylmethyl-N-methoxyethylcarbamoylphenylcarbamoylglycyl, etc.],

amino acid residue substituted with
15 arylcarbamoyl-arylcarbamoyl [e.g. phenylcarbamoylphenylcarbamoylglycyl, etc.],

amino acid residue substituted with lower
20 alkylaminoarylcarbamoyl-arylcarbamoyl [e.g. dimethylamino-phenylcarbamoylphenylcarbamoylglycyl, etc.], etc.,

amino acid residue substituted with arylthiocarbamoyl
25 [e.g. phenylthiocarbamoylglycyl, naphthylthiocarbamoylglycyl, phenylthiocarbamoylalanyl, phenylthiocarbamoylsarcosyl, etc.], amino acid residue substituted with ar(lower)alkylcarbamoyl [e.g. benzylcarbamoylglycyl, benzylcarbamoylsarcosyl, benzylcarbamoylalanyl, etc.], amino acid residue substituted with aroylcarbamoyl [e.g. benzoylcarbamoylglycyl, etc.], amino acid residue substituted with heterocycliccarbamoyl [e.g. pyridylcarbamoylglycyl, pyridylcarbamoylalanyl, pyridylcarbamoylsarcosyl, thienylcarbamoylglycyl, pyrazolylcarbamoylglycyl, pyrimidinylcarbamoylglycyl, quinolylcarbamoylglycyl, isoquinolylcarbamoylglycyl, etc.], amino acid residue substituted with heterocyclic(lower)alkylcarbamoyl [e.g. pyridylmethylcarbamoylglycyl, pyridylethylcarbamoylglycyl,
thienylmethylcarbamoylglycyl, etc.],
amino acid residue substituted with arylaminocarbamoyl
[e.g. phenylaminocarbamoylglycyl, etc.], amino acid
residue substituted with ar(lower)alkenylsulfonfonyl [e.g.
styrylsulfonylglycyl, cinnamylsulfonylglycyl, etc.], amino acid residue substituted with lower alkylsulfonfonyl
[e.g. mesylglycyl, ethylsulfonylglycyl, mesylsarcosyl,
mesylalanyl, etc.], amino acid residue substituted with
phthaloxy [e.g. phthaloxyglycyl, phthaloxyalanyl,
phthaloxy-β-alanyl, etc.], amino acid residue having
unsubstituted amino acid residue [e.g. glycylglycyl,
alanylglycyl, sarcosylglycyl, prolylglycyl,
glycylsarcosyl, prolylsarcosyl, etc.], amino acid residue having substituted amino acid residue [e.g. amino acid
residue having amino acid residue substituted with lower
alkyl (e.g. dimethylglycylglycyl, diethylglycylglycyl,
dimethylglycylsarcosyl, ethylsarcosylglycyl,
isopropylsarcosylglycyl, ethylglycylglycyl,
propylglycylglycyl, isopropylglycylglycyl,
edthylglycylalanyl, dimethylglycylalanyl,
dimethylalanylglycyl, dimethyl-β-alanylglycyl, etc.),
amino acid residue having amino acid residue substituted
with a heterocyclic group (e.g. morpholinoglycylglycyl,
piperidinoglycylglycyl, pyridylglycylglycyl,
piperidinosarcosylglycyl, etc.), amino acid residue having
amino acid residue substituted with
heterocyclic(lower)alkyl (e.g. pyridylmethylglycylglycyl,
imidazolylmethylglycylglycyl, furylmethylglycylglycyl,
thienylmethlysarcosylglycyl, etc.), amino acid residue
having amino acid residue substituted with cycloalkyl
[e.g. cyclopropylglycylglycyl, cyclobutylglycylglycyl,
cyclopentylglycylglycyl, cyclohexylglycylglycyl,
cycloheptylglycylglycyl, cyclooctylglycylglycyl,
adamantylglycylglycyl, cyclohexylsarcosylglycyl,
cycloheptylsarcosylglycyl, cyclohexylglycylsarcosyl,
cyclohexylglycylalanyl, etc.), amino acid residue having amino acid residue substituted with aryl (e.g. phenylglycylglycyl, phenylsarcosylglycyl, etc.), amino acid residue having amino acid residue substituted with acyl (e.g. amino acid residue having amino acid residue substituted with alkanoyl (e.g. acetylglycylglycyl, acetylprolylglycyl, propionylglycylglycyl, acetyllalanylglcycl, etc.), amino acid residue having amino acid residue substituted with lower alkoxy carbonyl (e.g. tert-butoxycarbonylglycyl, tert-butoxycarbonylprolylglycyl, etc.), amino acid residue having amino acid residue substituted with phthaloyl (e.g. phthaloylglycylglycyl, etc.), amino acid residue having amino acid residue substituted with ar(lower)alkyl (e.g. benzylglycylglycyl, etc.), etc., or the like.

Groups of the formulas of the compounds [If] and [Ig]:

\[-(AA)-CO-Y\]  \[-(AA)-CO-Y\]

wherein \(R^{1,2}\), \(R^{1,3}\), \((AA)\), \(Y\) and \(Z\) are each as defined above, are also included within "acyl".

Suitable "acyl having amino" may be unsubstituted amino acid residue, amino acid residue having unsubstituted amino acid residue, or the like, and preferred examples thereof can be referred to those exemplified above.

Suitable "acyl having acylamino" may be amino acid residue substituted with acyl, amino acid residue having amino acid residue substituted with acyl, or the like, and
preferred examples thereof can be referred to those exemplified above.

Suitable "protected or unprotected hydroxy(lower)alkyl" may be hydroxymethyl, hydroxyethyl, hydroxypropyl, benzyloxyethyl, tert-butylphenylsilyloxyethyl or the like.

Suitable "lower alkoxy(lower)alkyl" may be methoxymethyl, methoxyethyl, methoxypropyl, ethoxymethyl, ethoxyethyl, or the like.

Suitable "lower alkylamino(lower)alkyl" may be methylaminomethyl, methylaminoethyl, methylaminopropyl, dimethylaminomethyl, dimethylaminoethyl, dimethylaminopropyl, diethylaminoethyl, or the like.

Suitable "acyl having lower alkylamino" may be amino acid residue substituted with lower alkyl, amino acid residue having amino acid residue substituted with lower alkyl, or the like, and preferred examples thereof can be referred to those exemplified above.

Suitable "acyl having ar(lower)alkylamino" may be amino acid residue substituted with ar(lower)alkyl, amino acid residue having amino acid residue substituted with ar(lower)alkyl, or the like, and preferred examples thereof can be referred to those exemplified above.

Suitable "heterocyclic group" and heterocyclic moiety in the term "heterocyclic(lower)alkyl" may be saturated or unsaturated, monocyclic or polycyclic heterocyclic group containing at least one hetero-atom such as an oxygen, sulfur and/or nitrogen atom such as:

- unsaturated 3 to 8-membered, preferably 5 or 6-membered heteromonocyclic group containing 1 to 4 nitrogen atom(s), for example, pyrrolyl, pyrrolinyl, imidazolyl, pyrazolyl, pyridyl, and its N-oxide, pyrimidinyl, pyrazinyl, pyridazinyl, triazolyl, tetrazolyl, dihydrotiazinyl, etc.;

- saturated 3 to 8-membered, preferably 4 or
6-membered heteromonocyclic group containing 1 to 4 nitrogen atom(s), for example, azetidinyl, pyrrolidinyl, imidazolidinyl, piperidyl, pyrazolidinyl, piperazinyl, etc.;

- unsaturated condensed 7 to 12-membered heterocyclic group containing 1 to 5 nitrogen atom(s), for example, indolyl, isoindolyl, indolizinyl, benzimidazolyl, quinolyl, isoquinolyl, tetrahydroquinolyl, indazole, benzotriazolyl, imidazopyridyl, etc.;

- unsaturated 3 to 8-membered, preferably 5 or 6-membered heteromonocyclic group containing an oxygen atom, for example, furyl, etc.;

- unsaturated condensed 7 to 12-membered heterocyclic group containing 1 to 2 oxygen atom(s), for example, benzofuryl, piperonyl, etc.;

- unsaturated 3 to 8-membered, preferably 5 or 6-membered heteromonocyclic group containing a sulfur atom, for example, thieryl, etc.;

- unsaturated condensed 7 to 12-membered heterocyclic group containing 1 to 2 sulfur atom(s), for example, benzothieryl, etc.;

- unsaturated 3 to 8-membered, preferably 5 or 6-membered heteromonocyclic group containing 1 to 2 oxygen atom(s) and 1 to 3 nitrogen atom(s), for example, oxazolyl, isoxazolyl, oxadiazolyl, etc.;

- saturated 3 to 8-membered, preferably 5 or 6-membered heterocyclic group containing 1 to 2 oxygen atom(s) and 1 to 3 nitrogen atom(s), for example, morpholyl, etc.;

- unsaturated condensed 7 to 12-membered heterocyclic group containing 1 to 2 oxygen atom(s) and 1 to 3 nitrogen atom(s), for example, benzoazolyl, benzoazadiazolyl, etc.;

- unsaturated 3 to 8-membered, preferably 5 or 6-membered heteromonocyclic group containing 1 to 2 sulfur atom(s) and 1 to 3 nitrogen atom(s), for example,
thiazolyl, isothiazolyl, thiazolinyl, thiadiazolyl, etc.;
- saturated 3 to 8-membered, preferably 5 or
6-membered heteromonocyclic group containing 1 to 2 sulfur
atom(s) and 1 to 3 nitrogen atom(s), for example,
thiazolidinyl, etc.;
- unsaturated condensed 7 to 12-membered heterocyclic
group containing 1 to 2 sulfur atom(s) and 1 to 3 nitrogen
atom(s), for example, benzothiazolyl, benzothiadiazolyl,
benzothiazinyl, benzothiazolinyl, etc., or the like.

Suitable substituents in the terms "aryl having
suitable substituent(s)" or "heterocyclic group optionally
having suitable substituent(s)" may be the above-mentioned
halogen; the above-mentioned halo(lower)alkyl; the
above-mentioned lower alkyl; the above-mentioned acyl; the
above-mentioned aryl; aryl substituted with substituent(s)
such as halogen or cyano [e.g. chlorophenyl, cyanophenyl,
etc.]; ar(lower)alkyl substituted with hydroxy [e.g.
hydroxybenzyl, etc.]; lower alkoxy [e.g. methoxy, ethoxy,
proproxy, isoproproxy, butoxy, isobutoxy, tert-butoxy,
pentyloxy, hexyloxy, etc.]; oxo; nitro; amino; amino
substituted with the above-mentioned lower alkyl, the
above-mentioned acyl, ar(lower)alkyl [e.g. benzyl,
phenethyl, trityl, etc.], heterocyclic(lower)alkyl [e.g.
pyridylmethyl, etc.], carboxy(lower)alkyl [e.g.
carboxymethyl, carboxyethyl, etc.], lower
alkylaminomethylene [e.g. dimethylaminomethylene,
diethylaminomethylene, etc.], N-methylpyrrolidinylidene,
etc.; the above-mentioned heterocyclic group; heterocyclic
group substituted with oxo [e.g. pyrrolidinyl, etc.]; or
the like.

Suitable "heterocyclic group" formed by R12, R13 and
the attached nitrogen atom may be morpholino,
thiomorpholino, pyrrolidin-1-yl, piperidino,
1,2,3,6-tetrahydropyridin-1-yl, piperazin-1-yl, or the
like. And said heterocyclic group may be substituted with
suitable substituent(s) such as the above-mentioned lower alkyl, the above-mentioned heterocyclic group, the above-mentioned acyl, lower alkylamino, the above-mentioned aryl, or the like.

Preferred examples of "heterocyclic(lower)alkyl" may be morpholinomethyl, morpholinoethyl, pyridylmethyl, pyridylethyl, thiénylmethyl, piperidinomethyl, imidazolylmethyl, or the like.

Particularly, the preferred embodiments of \( R^1, R^2, R^3, R^4, Q \) and \( \Lambda \) are as follows:

1. \( R^1 \) is halogen such as fluorine, chlorine, bromine and iodine;
2. \( R^2 \) and \( R^3 \) are each hydrogen; lower alkyl such as methyl, ethyl, propyl, isopropyl, butyl, isobutyl, tert-butyl, pentyl and hexyl; halo(lower)alkyl such as fluoromethyl, chloromethyl, bromomethyl and trifluoromethyl; or acyl such as carboxy and esterified carboxy, for example, lower alkoxy carbonyl [e.g. methoxy carbonyl, ethoxycarbonyl, tert-butoxycarbonyl, etc.];
3. \( R^4 \) is phenyl substituted with substituent(s) such as -halogen such as fluorine, chlorine, bromine and iodine,
   -lower alkyl such as methyl, ethyl, propyl and isopropyl,
   -halo(lower)alkyl such as trifluoromethyl, -phenyl,
   -cyanophenyl,
   -hydroxy benzyl,
   -lower alkoxy such as methoxy, ethoxy, propoxy and isopropoxy,
   -nitro,
4. a group of the formula:
5 \( R^a \) is hydrogen; lower alkyl such as methyl, ethyl, propyl and butyl; carboxy(lower)alkyl such as carboxymethyl and carboxyethyl; and acyl such as lower alkanoyl [e.g. formyl, acetyl, propionyl, etc.], carboxy and esterified carboxy [e.g. lower alkoxy carbonyl (e.g. methoxycarbonyl, ethoxycarbonyl, tert-butoxycarbonyl, etc.), etc.],

10 \( R^{10} \) is hydrogen; lower alkyl such as methyl, ethyl, propyl, isopropyl and butyl; ar(lower)alkyl such as benzyl; heterocyclic(lower)alkyl such as pyridyl(lower)alkyl [e.g. pyridylmethyl, pyridylethyl, etc.]; and acyl such as lower alkanoyl [e.g. formyl, acetyl, propionyl, butyryl, isobutyryl, etc.], halo(lower)alkanoyl [e.g. trifluoroacetyl, etc.], carboxy, esterified carboxy [e.g. lower alkoxy carbonyl (e.g. methoxycarbonyl, ethoxycarbonyl, tert-butoxycarbonyl, etc.), etc.], hydroxy(lower)alkanoyl [e.g. glycoloyl, lactoyl, 3-hydroxypropionyl, etc.], lower alkanoyloxy(lower)alkanoyl [e.g. acetoxyacetyl, acetoxypropionyl, etc.], lower alkoxy(lower)alkanoyl [e.g. methoxyacetyl, methoxypropionyl, etc.], benzoyl, toluoyl, benzoyl substituted with lower alkoxy [e.g. methoxybenzoyl, etc.], benzoyl substituted with esterified carboxy [e.g. lower alkoxy carbonylbenzoyl (e.g. methoxycarbonylbenzoyl, tert-butoxycarbonylbenzoyl, etc.), etc.], benzoyl...
substituted with halogen [e.g. chlorobenzoyl, fluorobenzoyl, etc.], phenoxy carbonyl optionally substituted with nitro, lower alkylsulfonyl [e.g. mesyl, ethylsulfonyl, etc.], carbamoyl, lower alkylcarbamoyl [e.g. methylcarbamoyl, ethylcarbamoyl, isopropylcarbamoyl, etc.], halo(lower)alkanoylcarbamoyl [e.g. trichloroacetylcarbamoyl, etc.], phenylcarbamoyl, unsubstituted amino acid residue [e.g. glycyl, sarcosyl, alanyl, β-alanyl, etc.] and substituted amino acid residue [e.g. amino acid residue substituted with lower alkyl (e.g. ethylglycyl, isopropylglycyl, dimethylglycyl, diethylglycyl, ethylsarcosyl, isopropylsarcosyl, methylalanyl, methyl-β-alanyl, etc.), amino acid residue substituted with optionally substituted amidino (e.g. amidinoglycyl, N-ethyl-N'-cyanoaminodinoglycyl, etc.), amino acid residue substituted with acyl (e.g. amino acid residue substituted with alkanoyl (e.g. formylglycyl, acetylglycyl, acetylscarcosyl, acetylalanyl, acetyl-β-alanyl, propionylglycyl, butyrylglycyl, isobutyrylglycyl, valerylglycyl, isovalerylglycyl, pivaloylglycyl, hexanoylglycyl, heptanoylglycyl, etc.), amino acid residue substituted with halo(lower)alkanoyl (e.g. trifluoroacetylglcycl, trifluoroacetylscarcosyl, trifluoroacetylalanyl, bromoacylglcycl, heptafluorobutyrylglycyl, etc.), amino acid residue substituted with hydroxy(lower)alkanoyl (e.g. glycoloylglycyl, glycoloylsarcosyl, lactoylglycyl, lactoylalanyl, etc.), amino acid residue substituted with lower
alkylsulfonyloxy(lower)alkanoyl (e.g. mesyloxyacetylglucyl, ethylsulfonyloxyacetylglucyl, mesyloxyacetylserosyl, etc.), amino acid residue substituted with lower alkoxy(lower)alkanoyl (e.g. methoxyacetylglucyl, ethoxyacetylglucyl, methoxyacetylserosyl, methoxypropionylalanyl, etc.), amino acid residue substituted with aryloxy(lower)alkanoyl (e.g. phenyloxyacetylglucyl, phenyloxypropionylglucyl, phenyloxyacetylserosyl, etc.), amino acid residue substituted with lower alkylthio(lower)alkanoyl (e.g. methylthioacetylglucyl, methylthiopropionylglucyl, etc.), amino acid residue substituted with lower alkylcarbamoyl(lower)alkanoyl (e.g. methylcarbamoylpropionylglucyl, methylcarbamoylpropionylalanyl, etc.), amino acid residue substituted with lower alkanoyloxy(lower)alkanoyl (e.g. acetyloxyacetylglucyl, acetyloxyacetylserosyl, propionyoxyacetylglucyl, acetyloxypropionylalanyl, etc.), amino acid residue substituted with carboxy(lower)alkanoyl (e.g. carboxyacetylglucyl, carboxypropionylglucyl, carboxypropionylserosyl, carboxyacetylalanyl, etc.), amino acid residue substituted with lower alkoxy carbonyl(lower)alkanoyl (e.g. methoxycarbonylacetylglucyl, ethoxycarbonylacetylglucyl, methoxycarbonylpropionylglucyl, methoxycarbonylacetylserosyl, etc.), amino acid residue substituted with ar(lower)alkanoyl (e.g.
phenylacetylglycyl, phenylpropionylglycyl, phenylbutyrylglycyl, phenylacetylsarcosyl, phenylpropionylalanyl, naphthylacetylglycyl, etc.), amino acid residue substituted with optionally substituted heterocyclic(lower)alkanoyl (e.g. morpholinoacetylglycyl, pyridylacetylglycyl, morpholinopropionylalanyl, imidazolylacetylglycyl, piperidinoacetylglycyl, pyrrolidinylacetylglycyl, hexamethyleneminoacetylglycyl, methylpiperazinylacetylglycyl, pyridylpiperazinylacetylglycyl, thiomorpholinoacetylglycyl, its oxide or dioxide, etc.), amino acid residue substituted with lower alkenoyl (e.g. acryloylglycyl, crotonoylglycyl, 3-pentenoylglycyl, 3-butenoylglycyl, 4-pentenoylglycyl, 3-butenoylsarcosyl, etc.), amino acid residue substituted with ar(lower)-alkenoyl (e.g. cinnamoylglycyl, α-methylcinnamoylglycyl, 4-methylcinnamoylglycyl, etc.), amino acid residue substituted with lower alkoxy-ar(lower)alkenoyl (e.g. methoxycinnamoylglycyl, ethoxycinnamoylglycyl, dimethoxycinnamoylglycyl, etc.), amino acid residue substituted with lower alkylenedioxy-ar(lower)alkenoyl (e.g. methylenedioxyacetylglycyl, ethylenedioxyacetylglycyl, etc.), amino acid residue substituted with nitro-ar(lower)alkenoyl (e.g. nitrocinnamoylglycyl, etc.), amino acid residue substituted with halo-ar(lower)alkenoyl (e.g. chlorocinnamoylglycyl, fluorocinnamoylglycyl, etc.), amino acid residue substituted with
hydroxy-ar(lower)alkenoyl (e.g.
hydroxycinnamoylglycyl, etc.),
amino acid residue substituted with
hydroxy(lower)alkoxy-ar(lower)alkenoyl (e.g.
hydroxymethoxycinnamoylglycyl,
hydroxyethoxycinnamoylglycyl, etc.),
amino acid residue substituted with amino(lower)-
alkoxy-ar(lower)alkenoyl (e.g.
aminoethoxycinnamoylglycyl, etc.),
amino acid residue substituted with lower
alkylamino(lower)alkoxy-ar(lower)alkenoyl (e.g.
methylaminomethoxycinnamoylglycyl,
dimethylaminoethoxycinnamoylglycyl, etc.),
amino acid residue substituted with
heterocyclic(lower)alkoxy-ar(lower)alkenoyl (e.g.
pyridylmethoxycinnamoylglycyl, etc.),
amino acid residue substituted with optionally
substituted heterocyclic-ar(lower)alkenoyl (e.g.
morpholinocinnamoylglycyl, methylpiperazinylcinn-
amoylglycyl, pyrrolidinylcinnamoylglycyl,
oxopyrrolidinylcinnamoylglycyl,
oxopiperidinocinnamoylglycyl,
dioxopyrrolidinylcinnamoylglycyl,
oxooxazolidinylcinnamoylglycyl,
pyrolylcinnamoylglycyl, etc.),
amino acid residue substituted with
amino-ar(lower)alkenoyl (e.g.
aminocinnamoylglycyl, etc.),
amino acid residue substituted with lower
alkylamino-ar(lower)alkenoyl (e.g.
methylaminocinnamoylglycyl,
dimethylaminocinnamoylglycyl, etc.),
amino acid residue substituted with lower
alkanoylamino-ar(lower)alkenoyl (e.g.
acetylaminocinnamoylglycyl,
propionylaminocinnamoylglycyl, etc.),
amino acid residue substituted with
hydroxy(lower)alkanoylaminoo-ar(lower)alkenoyl
(e.g. hydroxyacetylaminoacinnamoylglycyl,
hydroxypropionylaminocinnamoylglycyl, etc.),
amino acid residue substituted with lower
alkoxy(lower)alkanoylaminoo-ar(lower)alkenoyl
(e.g. methoxyacetylaminoacinnamoylglycyl,
methoxypropionylaminocinnamoylglycyl, etc.),
amino acid residue substituted with
halo(lower)alkanoylaminoo-ar(lower)alkenoyl (e.g.
chloroacetylaminoacinnamoylglycyl,
bromobutyrlyaminocinnamoylglycyl, etc.),
amino acid residue substituted with
amino(lower)alkanoylaminoo-ar(lower)alkenoyl (e.g.
aminoacetylaminoacinnamoylglycyl,
aminoaminopropionylaminocinnamoylglycyl, etc.),
amino acid residue substituted with lower
alkylamino(lower)alkanoylaminoo-ar(lower)alkenoyl
(e.g. methylaminooacetylaminoacinnamoylglycyl,
dimethylaminooacetylaminoacinnamoylglycyl, etc.),
amino acid residue substituted with lower
alkanoylaminoo-ar(lower)alkanoylaminoo-ar
(lower)alkenoyl (e.g.
acetylaminoacetylaminoacinnamoylglycyl,
acetylaminoaminopropionylaminocinnamoylglycyl, etc.),
amino acid residue substituted with
carboxy(lower)alkanoylaminoo-ar(lower)alkenoyl
(e.g. carboxyacetylaminoacinnamoylglycyl,
carboxypropionylaminocinnamoylglycyl, etc.),
amino acid residue substituted with lower
alkoxy carbonyl(lower)alkanoylaminoo-ar(lower)alkenoyl (e.g. ethoxycarbonyl-
acetylaminoacinnamoylglycyl, ethoxycarbonyl-
propionylaminocinnamoylglycyl, etc.),
amino acid residue substituted with lower alkoxy carbonyl(lower) alkenoylamino-
ar(lower) alkenoyl (e.g. ethoxycarbonyl-
acryloylamino cinnamoylglycyl, etc.),
amino acid residue substituted with
halo(lower) alkoxy carbonyl-
ar(lower) alkenoyl (e.g. chloro ethoxycarbonyl-
amino cinnamoylglycyl, etc.),
amino acid residue substituted with heterocyclic-
(lower) alkanoylamino-
ar(lower) alkenoyl (e.g. pyridylacetylaminocinnamoylglycyl, etc.),
amino acid residue substituted with
arylamino-
ar(lower) alkenoyl (e.g. benzoylamino cinnamoylglycyl, etc.),
amino acid residue substituted with heterocycliccarbonylamino-
ar(lower) alkenoyl (e.g. nicotinoylamino cinnamoylglycyl,
isonicotinoylamino cinnamoylglycyl,
morpholinocarbonylamino cinnamoylglycyl, etc.),
amino acid residue substituted with lower alkyl sulfonamino-
ar(lower) alkenoyl (e.g. mesylaminocinnamoylglycyl,
ethyl sulfonamino cinnamoylglycyl, etc.),
amino acid residue substituted with
N-(lower alkanoyl)-N-(lower alkyl) amino-
ar(lower) alkenoyl (e.g. N-acetyl-
N-methy lamino cinnamoylglycyl,
N-acetyl-
N-ethylaminocinnamoylglycyl,
N-propionyl-
N-methy lamino cinnamoylglycyl, etc.),
amino acid residue substituted with N-[lower alkoxy(lower)-
lower alkenoyl]-N-(lower alkyl) amino-
ar(lower) alkenoyl (e.g. N-methoxyacetyl-
N-methy lamino cinnamoylglycyl,
N-methoxypropionyl-
N-methy lamino cinnamoylglycyl, etc.), amino acid residue substituted with
N-(lower alkanoyl)-N-[heterocyclic(lower)alkyl]-amino-ar(lower)alkenoyl (e.g. N-acetyl-N-pyridylmethylaminocinnamoylglycyl, etc.), amino acid residue substituted with
ureido-ar(lower)alkenoyl (e.g. ureidocinnamoylglycyl, etc.), amino acid residue substituted with lower alkylureido-ar(lower)alkenoyl (e.g. methylureidocinnamoylglycyl, ethylureidocinnamoylglycyl, dimethylureidocinnamoylglycyl, etc.), amino acid residue substituted with heterocyclicureido-ar(lower)alkenoyl (e.g. pyridylureidocinnamoylglycyl, pyrimidinylureidocinnamoylglycyl, thierylureidocinnamoylglycyl, etc.), amino acid residue substituted with lower alkanoyl-ar(lower)alkenoyl (e.g. formylcinnamoylglycyl, acetylcinnamoylglycyl, propionylcinnamoylglycyl, etc.), amino acid residue substituted with carboxy-ar(lower)alkenoyl (e.g. carboxycinnamoylglycyl, etc.), amino acid residue substituted with lower alkoxy carbonyl-ar(lower)alkenoyl (e.g. methoxycarbonylcinnamoylglycyl, ethoxycarbonylcinnamoylglycyl, etc.), amino acid residue substituted with carbamoyl-ar(lower)alkenoyl (e.g. carbamoylcinnamoylglycyl, etc.), amino acid residue substituted with lower alkyl carbamoyl-ar(lower)alkenoyl (e.g. methyl carbamoylcinnamoylglycyl, ethyl carbamoylcinnamoylglycyl, dimethyl carbamoylcinnamoylglycyl,
propylcarbamoylcinnamoylglycyl,  
isopropylcarbamoylcinnamoylglycyl,  
diethylcarbamoylcinnamoylglycyl, N-methyl-N-ethylcarbamoylcinnamoylglycyl, etc.),  
amino acid residue substituted with hydroxy-(lower)alkylcarbamoyl-ar(lower)alkenoyl (e.g.  
hydroxyethylcarbamoylcinnamoylglycyl,  
bis(hydroxyethyl)carbamoylcinnamoylglycyl, etc.),  
amino acid residue substituted with  
N-[hydroxy(lower)alkyl]-N-(lower alkyl)carbamoyl-ar(lower)alkenoyl (e.g. N-hydroxyethyl-N-methylcarbamoylcinnamoylglycyl, etc.),  
amino acid residue substituted with lower alkoxy(lower)alkylcarbamoyl-ar(lower)alkenoyl (e.g. methoxymethylcarbamoylcinnamoylglycyl, methoxyethylcarbamoylcinnamoylglycyl, bis(methoxyethyl)carbamoylcinnamoylglycyl, ethoxyethylcarbamoylcinnamoylglycyl, methoxypropylcarbamoylcinnamoylglycyl, bis(ethoxyethyl)carbamoylcinnamoylglycyl, etc.),  
amino acid residue substituted with N-[lower alkoxy(lower)alkyl]-N-(lower alkyl)carbamoyl-ar(lower)alkenoyl (e.g. N-methoxyethyl-N-methylcarbamoylcinnamoylglycyl, etc.),  
amino acid residue substituted with heterocycliccarbamoyl-ar(lower)alkenoyl (e.g. morpholinylcarbamoylcinnamoylglycyl, thienylcarbamoylcinnamoylglycyl, pyridylcarbamoylcinnamoylglycyl, pyrimidinylcarbamoylcinnamoylglycyl, etc.),  
amino acid residue substituted with heterocycliccarbonyl-ar(lower)alkenoyl (e.g. morpholinocarbonylcinnamoylglycyl, pyrrolidinylcarbonylcinnamoylglycyl, piperidinocarbonylcinnamoylglycyl, etc.),
amino acid residue substituted with ar(lower)-alkynoyl (e.g. phenylpropioloylglycyl, etc.),
amino acid residue substituted with heterocyclic(lower)alkenoyl (e.g. morpholinylacryloylglycyl, pyridylacryloylglycyl, thienylacryloylglycyl, etc.),
amino acid residue substituted with amino-heterocyclic(lower)alkenoyl (e.g. aminopyridylacryloylglycyl, etc.)
amino acid residue substituted with lower alkylamino-heterocyclic(lower)alkenoyl (e.g. methylaminopyridylacryloylglycyl, dimethylaminopyridylacryloylglycyl, etc.),
amino acid residue substituted with lower alkanoylamino-heterocyclic(lower)alkenoyl (e.g. acetylaminopyridylacryloylglycyl, propionylaminopyridylacryloylglycyl, etc.),
amino acid residue substituted with lower alkylureido-heterocyclic(lower)alkenoyl (e.g. methylureidopyridylacryloylglycyl, etc.),
amino acid residue substituted with carboxy-heterocyclic(lower)alkenoyl (e.g. carboxypyridylacryloylglycyl, etc.),
amino acid residue substituted with lower alkoxy carbonyl-heterocyclic(lower)alkenoyl (e.g. ethoxycarbonylpyridylacryloylglycyl, etc.),
amino acid residue substituted with lower alkylcarbamoyl-heterocyclic(lower)alkenoyl (e.g. methylcarbamoylpyridylacryloylglycyl, ethylcarbamoylpyridylacryloylglycyl, dimethylcarbamoylpyridylacryloylglycyl, diethylcarbamoylpyridylacryloylglycyl, isopropylcarbamoylpyridylacryloylglycyl, etc.),
amino acid residue substituted with lower alkoxy(lower)alkylcarbamoyl-
heterocyclic(lower)alkenoyl (e.g. methoxymethylcarbamoylpyridylacryloylglycyl, methoxyethylcarbamoylpyridylacryloylglycyl, methoxypropylcarbamoylpyridylacryloylglycyl, ethoxyethylcarbamoylpyridylacryloylglycyl, bis(methoxyethyl)carbamoylpyridylacryloylglycyl, etc.), amino acid residue substituted with hydroxy(lower)alkylcarbamoyl-
heterocyclic(lower)alkenoyl (e.g. hydroxymethylcarbamoylpyridylacryloylglycyl, hydroxyethylcarbamoylpyridylacryloylglycyl, bis(hydroxyethyl)carbamoylpyridylacryloylglycyl, etc.), amino acid residue substituted with heterocyclicthio(lower)alkanoyl (e.g. pyridylthioacetyl glycyl, pyrimidinylthioacetyl glycyl, imidazolylthiopropionyl glycyl, etc.), amino acid residue substituted with optionally substituted heterocycliccarbonyl (e.g. morpholinocarbonylglycyl, indolylcarbonylglycyl, 4-methyl-1-piperazinylcarbonylglycyl, etc.), amino acid residue substituted with cyclo(lower)alkylcarbonyl (e.g. cyclopropylcarbonylglycyl, cyclopentylcarbonylglycyl, cyclohexylcarbonylglycyl, cyclohexylcarbonylsarcosyl, etc.), amino acid residue substituted with lower alkoxy carbonyl (e.g. methoxycarbonylglycyl, tert-butoxycarbonylglycyl, tert-butoxycarbonylsarcosyl, tert-butoxycarborny1alanyl, etc.), amino acid residue substituted with aryloxy carbonyl (e.g. phenoxycarbonylglycyl, etc.), amino acid residue substituted with
aroyl(lower)alkanoyl (e.g. phenyloxalylglycyl, benzoilpropionylglycyl, etc.), amino acid residue substituted with aroyl (e.g. benzoilglycyl, benzoilsarcosyl, naphthoylglycyl, benzoylalanyl, etc.), amino acid residue substituted with nitro-aryloxycarbonyl (e.g. nitrophenyloxycarbonylglycyl, etc.), amino acid residue substituted with carbamoyl (e.g. carbamoylglycyl, carbamoylalanyl, carbamoylsarcosyl, carbamoyl-β-alanyl, etc.), amino acid residue substituted with lower alkylcarbamoyl (e.g. methylcarbamoylglycyl, ethylcarbamoylglycyl, propylcarbamoylglycyl, isopropylcarbamoylglycyl, pentylcramooylglycyl, methylcarbamoylsarcosyl, ethylcarbamoylalanyl, isopropylcarbamoyl-β-alanyl, etc.), amino acid residue substituted with lower alkoxy-carbonyl(lower)alkylcarbamoyl (e.g. methoxycarbonylmethylcarbamoylglycyl, ethoxycarbonylmethylcarbamoylglycyl, etc.), amino acid residue substituted with lower alkenylcarbamoyl (e.g. vinylcarbamoylglycyl, allylcarbamoylglycyl, allylcarbamoylsarcosyl, etc.), amino acid residue substituted with cyclo(lower)alkylcarbamoyl (e.g. cyclopentylcarbamoylglycyl, cyclohexylcarbamoylglycyl, cyclohexylcarbamoylsarcosyl, etc.), amino acid residue substituted with arylcarbamoyl (e.g. phenylcarbamoylglycyl, naphthylcarbamoylglycyl, tolylcarbamoylglycyl, ethylphenylcarbamoylglycyl, phenylcarbamoylalanyl, phenylcarbamoylsarcosyl, etc.), amino acid residue substituted with lower alkoxy-arylcramooyl (e.g.
methoxyphenylcarbamoylglycyl,
ethoxyphenylcarbamoylglycyl,
methoxyphenylcarbamoylalanyl, etc.), amino acid residue substituted with
halo(lower)alkyl-arylcarbamoyl (e.g.
trifluoromethylphenylcarbamoylglycyl,
trifluoromethylphenylcarbamoylalanyl,
trifluoromethylphenylcarbamoylsarcosyl, etc.), amino acid residue substituted with
halo-arylcarbamoyl (e.g.
chlorophenylcarbamoylglycyl,
fluorophenylcarbamoylglycyl,
fluorophenylcarbamoylalanyl, etc.), amino acid residue substituted with lower
alkanoyl-arylcarbamoyl (e.g.
acetylphenylcarbamoylglycyl,
propionylphenylcarbamoylalanyl, etc.), amino acid residue substituted with
hydroxy(lower)alkyl-arylcarbamoyl (e.g.
hydroxymethylphenylcarbamoylglycyl,
hydroxyethylphenylcarbamoylglycyl,
hydroxyethylphenylcarbamoylalanyl, etc.), amino acid residue substituted with
heterocycliccarbonyl-arylcarbamoyl (e.g.
morpholinocarbonylphenylcarbamoylglycyl,
piperidinocarbonylphenylcarbamoylglycyl,
thiomorpholinocarbonylphenylcarbamoylalanyl,
piperazinylcarbonylphenylcarbamoylglycyl,
pyrrolidinylcarbonylphenylcarbamoylglycyl,
1,2,3,6-tetrahydropyridylcarbonylphenylcarbamoylglycyl, etc.),
amino acid residue substituted with
carboxy-arylcarbamoyl (e.g.
carboxyphenylcarbamoylglycyl, etc.), amino acid residue substituted with lower
alkoxycarbonyl-arylcarbamoyl (e.g. methoxycarbonylphenylcarbamoylglycyl, ethoxycarbonylphenylcarbamoylglycyl, etc.), amino acid residue substituted with lower alkylcarbamoyl-arylcarbamoyl (e.g. methylcarbamoylphenylcarbamoylglycyl, dimethylcarbamoylphenylcarbamoylglycyl, diethylcarbamoylphenylcarbamoylglycyl, N-ethyl-N-methylcarbamoylphenylcarbamoylglycyl, N-isopropyl-N-methylcarbamoylphenylcarbamoylglycyl, etc.), amino acid residue substituted with nitro-arylcarbamoyl (e.g. nitrophenylcarbamoylglycyl, etc.), amino acid residue substituted with cyano-arylcarbamoyl (e.g. cyanophenylcarbamoylglycyl, etc.), amino acid residue substituted with amino-arylcarbamoyl (e.g. aminophenylcarbamoylglycyl, etc.), amino acid residue substituted with lower alkylamino-arylcarbamoyl (e.g. methylaminophenylcarbamoylglycyl, ethylaminophenylcarbamoylglycyl, dimethylaminophenylcarbamoylglycyl, etc.), amino acid residue substituted with lower alkanoylamino-arylcarbamoyl (e.g. acetylaminophenylcarbamoylglycyl, propionylaminophenylcarbamoylglycyl, etc.), amino acid residue substituted with N-(lower alkanoyl)-N-(lower alkyl)amino-arylcarbamoyl (e.g. N-acetyl-N-methylaminophenylcarbamoylglycyl, N-propionyl-N-methylaminophenylcarbamoylglycyl, etc.), amino acid residue substituted with lower alkoxy(lower)alkanoylamino-arylcarbamoyl (e.g. methoxyacetylaminophenylcarbamoylglycyl, methoxy-
propionylaminophenylcarbamoylglycyl, etc.), amino acid residue substituted with lower alkoxy carbonyl(lower) alkanoylamino-arylcarbamoyl (e.g. ethoxy carbonylacetaminophenyl-carbamoylglycyl, methoxycarbonylpropionylaminophenylcarbamoylglycyl, etc.), amino acid residue substituted with carboxyamino-arylcarbamoyl (e.g. carboxyaminophenylcarbamoylglycyl, etc.), amino acid residue substituted with lower alkoxy carbonylamino-arylcarbamoyl (e.g. ethoxycarbonylaminophenylcarbamoylglycyl, etc.), amino acid residue substituted with ureido-arylcarbamoyl (e.g. ureidophenylcarbamoylglycyl, etc.), amino acid residue substituted with lower alkylureido-arylcarbamoyl (e.g. methylureidophenylcarbamoylglycyl, ethylureidophenylcarbamoylglycyl, etc.), amino acid residue substituted with hydroxyimino(lower) alkyl-arylcarbamoyl (e.g. hydroxyiminoethylphenylcarbamoylglycyl, etc.), amino acid residue substituted with lower alkoxyimino(lower) alkyl-arylcarbamoyl (e.g. methoxyiminoethylphenylcarbamoylglycyl, etc.), amino acid residue substituted with lower alkylhydrazono(lower) alkyl-arylcarbamoyl (e.g. methylhydrazonoethylphenylcarbamoylglycyl, dimethylhydrazonoethylphenylcarbamoylglycyl, etc.), amino acid residue substituted with optionally substituted heterocyclic-arylcarbamoyl (e.g. oxopyrrolidinylphenylcarbamoylglycyl, oxopiperidinophenylcarbamoylglycyl, dioxopyrrolidinylphenylcarbamoylglycyl, oxooxazolidinylphenylcarbamoylglycyl,
pyrrolylphenylcarbamoylglycyl, etc.), amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having lower alkyl (e.g. methylpiperazinylcarbonylphenylcarbamoylglycyl, ethylpiperazinyl-carbonylphenylcarbamoylglycyl, etc.), amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having aryl (e.g. phenylpiperazinylcarbonylphenylcarbamoylglycyl, etc.), amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having a heterocyclic group (e.g. pyridylpiperazinylcarbonylphenylcarbamoylglycyl, etc.), amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having lower alkanoyl (e.g. acetylpiperazinylcarbonylphenylcarbamoylglycyl, etc.), amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having lower alkoxy carbonyl (e.g. ethoxycarbonylpiperazinylcarbonylphenylcarbamoylglycyl, etc.), amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having lower alkylamino (e.g. methylaminopiperazinylcarbonylphenylcarbamoylglycyl, dimethylaminopiperidino-carbonylphenylcarbamoylglycyl, etc.), amino acid residue substituted with heterocycliccarbonyl-arylcarbamoyl having lower alkylcarbamoyl (e.g. methylcarbamoylpiperazinylcarbonylphenylcarbamoylglycyl, etc.), amino acid residue substituted with hydroxy(lower)alkylcarbamoyl-arylcarbamoyl (e.g. hydroxymethylcarbamoylphenylcarbamoylglycyl, hydroxyethylcarbamoylphenylcarbamoylglycyl, bis(hydroxyethyl)carbamoylphenylcarbamoylglycyl,
etc.), amino acid residue substituted with N-[hydroxy(lower)alkyl]-N-(lower alkyl)carbamoyl-arylcarbamoyl (e.g. N-(hydroxyethyl)-N-methylcarbamoylphenylcarbamoylglycyl, etc.), amino acid residue substituted with lower alkoxy(lower)alkylcarbamoyl-arylcarbamoyl (e.g. methoxymethylcarbamoylphenylcarbamoylglycyl, methoxyethylcarbamoylphenylcarbamoylglycyl, bis(methoxyethyl)carbamoylphenylcarbamoylglycyl, bis(ethoxyethyl)carbamoylphenylcarbamoylglycyl, etc.), amino acid residue substituted with N-[lower alkoxy(lower)alkyl]-N-(lower alkyl)carbamoylarylcarbamoyl (e.g. N-(methoxyethyl)-N-methylcarbamoylphenylcarbamoylglycyl, N-(methoxypropyl)-N-methylcarbamoylphenylcarbamoylglycyl, etc.), amino acid residue substituted with lower alkylamino(lower)alkylcarbamoyl-arylcarbamoyl (e.g. methylaminoethylcarbamoylphenylcarbamoylglycyl, dimethylaminoethylcarbamoylphenylcarbamoylglycyl, etc.), amino acid residue substituted with N-[lower alkylamino(lower)alkyl]-N-(lower alkyl)carbamoylarylcarbamoyl (e.g. N-(dimethylaminoethyl)-N-methylcarbamoylphenylcarbamoylglycyl, N-(dimethylaminopropyl)-N-methylcarbamoylphenylcarbamoylglycyl, etc.), amino acid residue substituted with heterocycliccarbamoyl-arylcarbamoyl (e.g. morpholinylcarbamoylphenylcarbamoylglycyl, thienylcarbamoylphenylcarbamoylglycyl, pyridylcarbamoylphenylcarbamoylglycyl, pyrimidinylcarbamoylphenylcarbamoylglycyl, etc.).
amino acid residue substituted with
N-(heterocyclic)-N-(lower alkyl)carbamoylarylcarbamoyl (e.g. N-pyridyl-N-methylcarbamoylphenylcarbamoylglycyl, etc.),
amino acid residue substituted with heterocyclic-(lower)alkylcarbamoyl-arylcarbamoyl (e.g. pyridylmethylcarbamoylphenylcarbamoylglycyl, pyridylethylcarbamoylphenylcarbamoylglycyl, thienylmethylcarbamoylphenylcarbamoylglycyl, etc.), amino acid residue substituted with
N-[heterocyclic(lower)alkyl]-N-(lower alkyl)-carbamoyl-arylcarbamoyl (e.g. N-pyridylmethyl-N-methylcarbamoylphenylcarbamoylglycyl, etc.),
amino acid residue substituted with
N-[heterocyclic(lower)alkyl]-N-[lower alkoxy(lower)alkyl]carbamoyl-arylcarbamoyl (e.g. N-pyridylmethyl-N-methoxyethylcarbamoylphenylcarbamoylglycyl, etc.),
amino acid residue substituted with
arylcarbamoyl-arylcarbamoyl (e.g. phenylcarbamoylphenylcarbamoylglycyl, etc.),
amino acid residue substituted with lower alkylaminoarylcarbamoyl-arylcarbamoyl (e.g. dimethylaminophenylcarbamoylphenylcarbamoylglycyl, etc.),
amino acid residue substituted with
arylthiocarbamoyl (e.g. phenylthiocarbamoylglycyl, naphthylthiocarbamoylglycyl, phenylthiocarbamoylalanyl, phenylthiocarbamoylsarcosyl, etc.), amino acid residue substituted with ar(lower)alkylcarbamoyl (e.g. benzylcarbamoylglycyl, benzylcarbamoylsarcosyl, benzylcarbamoylalanyl, etc.), amino acid
residue substituted with aroylcarbamoyl (e.g. benzoylcarbamoylglycyl, etc.), amino acid residue substituted with heterocycliccarbamoyl (e.g. pyridylcarbamoylglycyl, pyridylcarbamoylalanyl, pyridylcarbamoylsarcosyl, thienylcarbamoylglycyl, pyrazolylcarbamoylglycyl, pyrimidinylcarbamoylglycyl, quinolylcarbamoylglycyl, isoquinolylcarbamoylglycyl, etc.), amino acid residue substituted with heterocyclic(lower)alkylcarbamoyl (e.g. pyridylmethylcarbamoylglycyl, pyridylethylcarbamoylglycyl, thienylmethylcarbamoylglycyl, etc.), amino acid residue substituted with arylaminocarbamoyl (e.g. phenylaminocarbamoylglycyl, etc.), amino acid residue substituted with ar(lower)alkenylsulfonyl (e.g. styrylsulfonylglycyl, cinnamylsulfonylglycyl, etc.), amino acid residue substituted with lower alkylsulfonyl (e.g. mesylglycyl, ethylsulfonylglycyl, mesylsarcosyl, mesylalanyl, etc.), amino acid residue substituted with phthaloyl (e.g. phthaloylglycyl, phthaloylalanyl, phthaloyl-β-alanyl, etc.), amino acid residue having unsubstituted amino acid residue (e.g. glycylglycyl, alanylglycyl, sarcosylglycyl, prolylglycyl, glycylsarcosyl, prolylsarcosyl, etc.), amino acid residue having substituted amino acid residue such as amino acid residue having amino acid residue substituted with lower alkyl (e.g. dimethylglycylglycyl, diethylglycylglycyl, dimethylglycylsarcosyl, ethylsarcosylglycyl, isopropylsarcosylglycyl, ...
ethylglycylglycyl, propylglycylglycyl, isopropylglycylglycyl, ethylglycylalanyl, dimethylglycylalanyl, dimethyllalanylglycyl, dimethyl-β-alanylglycyl, etc.), amino acid residue having amino acid residue substituted with a heterocyclic group (e.g. morpholinoglycylglycyl, piperidinoglycylglycyl, pyridylglycylglycyl, piperidinosarcosylglycyl, etc.), amino acid residue having amino acid residue substituted with heterocyclic(lower)alkyl (e.g. pyridylmethylglycylglycyl, imidazolylmethylglycylglycyl, furylmethylglycylglycyl, thienylmethylsarcosylglycyl, etc.), amino acid residue having amino acid residue substituted with cycloalkyl (e.g. cyclopropylglycylglycyl, cyclobutylglycylglycyl, cyclopentylglycylglycyl, cyclohexylglycylglycyl, cycloheptylglycylglycyl, adamantylglycylglycyl, cyclohexylsarcosylglycyl, cyclohexylglycylalanyl, cyclohexylglycylsarcosyl, cyclohexylglycylalanylglycyl, etc.), amino acid residue having amino acid residue substituted with aryl (e.g. phenylglycylglycyl, phenylsarcosylglycyl, etc.), amino acid residue having amino acid residue substituted with lower alkanoyl (e.g. acetylglycylglycyl, acetylprolylglycyl, propionylglycylglycyl, acetyllalanylglycyl, etc.), amino acid residue having amino acid residue substituted with lower alkoxy carbonyl (e.g. tert-butoxycarbonylglycylglycyl, tert-butoxycarbonylprolylglycyl, etc.), amino acid residue having amino acid residue substituted with ar(lower)alkyl (e.g.
benzylglycylglycyl, etc.) and amino acid residue having amino acid residue substituted with phthaloyl (e.g. phthaloylglycylglycyl, etc.), etc.];

-acyl such as carboxy, esterified carboxy [e.g. lower alkoxy carbonyl (e.g. methoxycarbonyl, ethoxycarbonyl, tert-butoxycarbonyl, etc.), etc.], aroyl [e.g. benzoyl, etc.] and heterocyclic carbonyl which may be substituted lower alkyl [e.g. morpholinocarbonyl, 4-methyl-1-piperazinyl carbonyl, etc.], lower alkylaminomethylene amino \( \equiv \) as dimethylaminomethylene amino and
diethylaminomethylene amino, heterocyclic group such as pyrrolidinyl, heterocyclic group substituted with oxo such as pyrrolidonyl, and/or

-N-methylpyrrolidinylidene amino;
or heterocyclic group such as thienyl, benzofuryl, benzothiazolinyl, benzothiazinyl and piperonyl, each of which may be substituted with substituent(s) such as halogen [e.g. fluorine, chlorine, bromine and iodine], lower alkyl [e.g. methyl, ethyl, propyl, isopropyl, etc.], haloaryl [e.g. chlorophenyl, etc.] and/or oxo;

Q is O or N-R\(^{11}\), in which R\(^{11}\) is hydrogen; or acyl such as lower alkanoyl [e.g. formyl, acetyl, propionyl, butyryl, etc.];

A is lower alkylene such as methylene, ethylene, methylmethylene and propylene.
Suitable "a leaving group" may be a conventional acid residue such as halogen [e.g. fluoro, chloro, bromo and iodo], arenesulfonyloxy [e.g. benzenesulfonyloxy, tosylxy, etc.], alkanesulfonyloxy [e.g. mesyloxy, ethanesulfonyloxy, etc.], and the like.

Suitable pharmaceutically acceptable salts of the object compound [I] are conventional non-toxic salts and include a metal salt such as an alkali metal salt [e.g. sodium salt, potassium salt, etc.] and an alkaline earth metal salt [e.g. calcium salt, magnesium salt, etc.], an ammonium salt, an organic base salt [e.g. trimethylamine salt, triethylamine salt, pyridine salt, picoline salt, dicyclohexylamine salt, N,N'-dibenzylethlenediamine salt, etc.], an organic acid addition salt [e.g. formate, acetate, trifluoroacetate, maleate, tartrate, oxalate, methanesulfonate, benzenesulfonate, toluenesulfonate, etc.], an inorganic acid addition salt [e.g. hydrochloride, hydrobromide, sulfate, phosphate, etc.], a salt with an amino acid [e.g. arginine salt, aspartic acid salt, glutamic acid salt, etc.], an intramolecular salt and the like.

With respect to the salts of the compounds [Ia] to [Ig] in the Processes 3 to 6, it is to be noted that these compounds are included within the scope of the compound [I], and accordingly the suitable examples of the salts of these compounds are to be referred to those as exemplified for the object compound [I].

The processes for preparing the object compound [I] are explained in detail in the following.

Process 1

The compound [I] or its salt can be prepared by halogenating a compound [II] or its salt.

Suitable salts of the compound [II] may be the same
as those exemplified for the compound [I].

The halogenation is carried out in the presence of a halogenating agent.

Suitable halogenating agents of this reaction may include conventional ones such as N-halosuccinimide [e.g. N-chlorosuccinimide, N-bromosuccinimide, etc.], and the like. These halogenating agents may be selected according to the kind of the starting compound [II] to be used.

This reaction is usually carried out in a conventional solvent such as chloroform, methylene chloride, carbon tetrachloride, dimethylformamide, methanol, ethanol, dioxane, or the like.

The reaction temperature is not critical, and the reaction is usually carried out at ambient temperature or under warming or heating.

Process 2

The object compound [I] or its salt can be prepared by reacting a compound [III] or its salt with a compound [IV] or its salt.

Suitable salts of the compounds [III] and [IV] may be the same as those exemplified for the compound [I].

The reaction is preferably carried out in the presence of a base such as alkali metal [e.g. lithium, sodium, potassium, etc.], the hydroxide or carbonate or bicarbonate thereof [e.g. sodium hydroxide, potassium carbonate, potassium bicarbonate, etc.], alkali metal alkoxide [e.g. sodium methoxide, sodium ethoxide, potassium tert-butoxide, etc.], or the like.

This reaction is usually carried out in a conventional solvent such as tetrahydrofuran, dioxane, N,N-dimethylformamide, acetone, or the like.

The reaction temperature is not critical, and the reaction is usually carried out under cooling to heating.
Process 3

The object compound [Ib] or its salt can be prepared by acylating a compound [Ia] or its salt.

The acylation is carried out in the presence of an acylating agent.

Suitable acylating agents are the corresponding carboxylic acid or sulfonic acid compounds, which are represented by the formula: R-OH wherein R is acyl, and reactive derivatives thereof, and the corresponding isocyanate or isothiocyanate compounds.

As suitable said reactive derivatives, there may be mentioned acid halides, acid anhydrides, active amides and active esters. Suitable examples are acid halides such as acid chloride and acid bromide, mixed acid anhydrides with various acids [e.g. substituted phosphoric acid such as dialkyl phosphoric acid, sulfuric acid, aliphatic carboxylic acid, aromatic carboxylic acid, etc.], symmetric acid anhydrides, active amides with various imidazoles, and active esters such as p-nitrophenyl ester and N-hydroxysuccinimide ester. The kind of such reactive derivatives can be selected depending on the kind of acyl group to be introduced.

The reaction is usually carried out in a conventional solvent, such as methylene chloride, chloroform, pyridine, dioxane, tetrahydrofuran, N,N-dimethylformamide, or the like. In case that the acylating agent is liquid, it can also be used as a solvent. In case that the carboxylic acid or sulfonic acid compounds are used as acylating agent in the free acid form or salt form, it is preferable to carry out the reaction in the presence of a conventional condensing agent such as N,N'-dicyclohexylcarbodiimide or the like.

The reaction temperature is not critical and the reaction can be carried out under cooling, at ambient temperature, or under heating.
This reaction is preferably carried out in the presence of a conventional inorganic base or in the presence of a conventional organic base.

**Process 4**

The object compound [Id] or its salt can be prepared by acylating a compound [Ic] or its salt.

This reaction can be carried out in substantially the same manner as **Process 3**, and therefore the reaction mode and reaction condition of this reaction are to be referred to those explained in **Process 3**.

**Process 5**

The object compound [Ie] or its salt can be prepared by alkylating a compound [Ic] or its salt.

This alklylation is carried out in the presence of an alkylating agent.

Suitable alkylating agents to be used in this reaction may be halide compounds such as lower alkyl halide [e.g. methyl iodide, ethyl iodide, propyl iodide, butyl iodide, etc.] or ar(lower)alkyl halide [e.g. benzylbromide, etc.], aldehyde compounds, ketone compounds, or the like.

When halide compounds are used as alkylating agents, the reaction is usually carried out in the presence of a base such as an alkali metal [e.g. sodium, potassium, etc.], an alkaline earth metal [e.g. magnesium, calcium, etc.], the hydride or hydroxide thereof, alkali metal alkoxide [e.g. sodium methoxide, sodium ethoxide, potassium tert-butoxide, etc.], or the like.

When aldehyde compounds or ketone compounds are used as alkylating agents, the reaction is usually carried out under acidic condition in the presence of reducing agent such as formic acid, sodium cyanoborohydride, or the like.

The reaction of this process is usually carried out
in a conventional solvent such as methanol, ethanol, tetrahydrofuran, dioxane, N,N-dimethylformamide, dimethyl sulfoxide, or the like. And in case that the above-mentioned alkylating agent is in liquid, it can be also used as a solvent.

The reaction temperature is not critical, and the reaction is usually carried out under cooling, at ambient temperature or under warming or heating.

**Process 6**

The object compound [Ig] or its salt can be prepared by reacting a compound [If] or its reactive derivative at the carboxy group or a salt thereof with a compound [IX] or its reactive derivative at the amino group or a salt thereof.

Suitable reactive derivative at the carboxy group of the compound [If] may include an acid halide, an acid anhydride, an activated amide, an activated ester, and the like. Suitable examples of the reactive derivatives may be an acid chloride; an acid azide; a mixed acid anhydride with an acid such as dialkylphosphoric acid, sulfuric acid aliphatic carboxylic acid or aromatic carboxylic acid; a symmetrical acid anhydride; an activated amide with imidazole; or an activated ester [e.g. p-nitrophenyl ester, etc.]. These reactive derivatives can optionally be selected from them according to the kind of the compound [If] to be used.

Suitable reactive derivative at the amino group of the compound [IX] may be a silyl derivative formed by the reaction of the compound [IX] with a silyl compound such as bis(trimethylsilyl)acetamide or mono(trimethylsilyl)acetamide, or the like.

Suitable salts of the compound [IX] and its reactive derivative can be referred to the organic or inorganic acid addition salts as exemplified for the compound [I].
This reaction can be carried out in substantially the same manner as Process 3, and therefore the reaction mode and reaction condition of this reaction are to be referred to those explained in Process 3.

The starting compounds [II] or [III] or salts thereof can be prepared by the following reaction schemes.

**Process A**

\[ \begin{align*}
R^3 & \quad \text{[V]} \\
Q \text{-} A \text{-} R^4 & \quad \text{or its salt}
\end{align*} \]

\[ \begin{align*}
R^2 \text{-CO-CH}_2 \text{X} & \quad \text{[VI]} \\
\text{or its salt}
\end{align*} \]

\[ \begin{align*}
R^3 & \quad \text{[II]} \\
Q \text{-} A \text{-} R^4 & \quad \text{or its salt}
\end{align*} \]

**Process B**

\[ \begin{align*}
R^3 & \quad \text{[VII]} \\
Q \text{H} & \quad \text{or its salt}
\end{align*} \]

\[ \begin{align*}
X \text{-A-} R^4 & \quad \text{[IV]} \\
\text{or its salt}
\end{align*} \]

\[ \begin{align*}
R^3 & \quad \text{[V]} \\
Q \text{-} A \text{-} R^4 & \quad \text{or its salt}
\end{align*} \]
Process C

\[
\begin{align*}
\text{R}^3 & \quad \text{N} \quad \text{N} \quad \text{R}^2 \\
\text{QH} & \quad \text{X-A-R}^4 \quad \text{[IV]} \quad \text{or its salt} \quad \text{R}^3 & \quad \text{N} \quad \text{N} \quad \text{R}^2 \\
\text{[VIII]} & \quad \text{or its salt} & \quad \text{Q-A-R}^4 & \quad \text{[II]} \quad \text{or its salt}
\end{align*}
\]

Process D

\[
\begin{align*}
\text{R}^3 & \quad \text{N} \quad \text{N} \quad \text{R}^2 \\
\text{QH} & \quad \text{Halogenation} \quad \text{R}^3 & \quad \text{N} \quad \text{N} \quad \text{R}^2 \\
\text{[VIII]} & \quad \text{or its salt} & \quad \text{QH} & \quad \text{[III]} \quad \text{or its salt}
\end{align*}
\]

wherein R\(^1\), R\(^2\), R\(^3\), R\(^4\), Q, A and X are each as defined above.

The above-mentioned processes for preparing the starting compounds are explained in detail in the following.
Process A
The compound [II] or its salt can be prepared by reacting a compound [V] or its salt with a compound [VI] or its salt.

Suitable salts of the compounds [V] and [VI] may be the same as those exemplified for the compound [I].

This reaction is usually carried out in a conventional solvent such as methanol, ethanol, acetonitrile, N,N-dimethylformamide, dimethyl sulfoxide or the like.

The reaction temperature is not critical, and the reaction is usually carried out at ambient temperature or under warming or heating.

Process B
The compound [V] or its salt can be prepared by reacting a compound [VII] or its salt with a compound [IV] or its salt.

Suitable salts of the compound [VII] are the same as those exemplified for the compound [I].

This reaction can be carried out in substantially the same manner as Process 2, and therefore the reaction mode and reaction condition of this reaction are to be referred to those explained in Process 2.

Process C
The compound [II] or its salt can be prepared by reacting a compound [VIII] or its salt with a compound [IV] or its salt.

Suitable salts of the compound [VIII] are the same as those exemplified for the compound [I].

This reaction can be carried out in substantially the same manner as Process 2, and therefore the reaction mode and reaction condition of this reaction are to be referred to those explained in Process 2.
Process D

The compound [III] or its salt can be prepared by halogenating a compound [VIII] or its salt.

This reaction can be carried out in substantially the same manner as Process 1, and therefore the reaction mode and reaction condition of this reaction are to be referred to those explained in Process 1.

The object compound [I] and the starting compounds can also be prepared by the methods of Examples and Preparations mentioned below or similar manners thereto or conventional manners.

The compounds obtained by the above processes can be isolated and purified by a conventional method such as pulverization, recrystallization, chromatography, reprecipitation or the like.

It is to be noted that the compound [I] and the other compounds may include one or more stereoisomers and geometrical isomers due to asymmetric carbon atoms and double bonds, and all of such isomers and mixture thereof are included within the scope of this invention.

The object compound [I] and pharmaceutically acceptable salts thereof possess strong activities as bradykinin antagonists, and are useful for the treatment and/or the prevention of bradykinin or its analogues mediated diseases such as allergy, inflammation, autoimmune disease, shock, pain, or the like, and more particularly for the prevention and/or the treatment of asthma, cough, bronchitis, rhinitis, rhinorrhea, obstructive pulmonary disease [e.g. pulmonary emphysema, etc.], expectoration, pneumonitis, systemic inflammatory response syndrome (SIRS), septic shock, endotoxin shock, anaphylactic shock, adult respiratory distress syndrome, disseminated intravascular coagulopathy, arthritis, rheumatism, osteoarthritis, lumbago, inflammation-induced bone resorption, conjunctivitis, vernal conjunctivitis,
uveitis, iritis, iridocyclitis, headache, migraine, toothache, backache, superficial pain, cancerous pain, postoperative pain, tenalgia, trauma [e.g. wound, burn, etc.], rash, erythema, eczema or dermatitis [e.g. contact dermatitis, atopic dermatitis, etc.], urticaria, herpes, itching, psoriasis, lichen, inflammatory bowel disease [e.g. ulcerative colitis, Crohn's disease, etc.], diarrhea, hepatitis, pancreatitis, gastritis, esophagitis, food allergy, ulcer, irritable bowel syndrome, nephritis, angina, periodontitis, edema, hereditary angioneurotic edema, cerebral edema, low blood pressure, thrombosis, myocardial infarction, cerebral vasospasm, congestion, coagulation, gout, central nervous system injury, premature labor, arteriosclerosis, postgastrectomy dumping syndrome, carcinoid syndrome, altered sperm mobility, diabetic neuropathy, neuralgia, graft rejection in transplantation, or the like, in human being or animals.

And further, it is known that bradykinin relates to the release of mediators such as prostaglandins, leukotrienes, tachykinins, histamine, thromboxanes, or the like, so the compound [I] is expected to be useful for the prevention and/or the treatment of such mediators mediated diseases.

In order to illustrate the usefulness of the object compound [I], the pharmacological test data of some representative compounds of the compound [I] are shown in the following.

\[ ^3 \text{H-Bradykinin receptor binding} \]

(i) Test Method:

(a) Crude ileum membrane preparation

Male Hartly strain guinea pigs were sacrificed by decapitation. The ileum was removed and homogenized in
buffer (50 mM trimethylaminoethanesulfonic acid (TES), 1 mM 1,10-phenanthroline pH 6.8). The homogenate was centrifuged (1000 xg, 20 minutes) to remove tissue clumps and the supernatant was centrifuged (100,000 xg, 60 minutes) to yield a pellet. The pellet was resuspended in buffer (50 mM TES, 1 mM 1,10-phenanthroline, 140 mg/£ bacitracin, 1 mM dithiothreitol, 0.1 % bovine serum albumin pH 6.8) and homogenized with a glass-teflon homogenizer to yield suspension which was referred to as crude membrane suspension. The obtained membrane suspension was stored at -80°C until use.

(b) $^3$H-Bradykinin binding to the membrane

The frozen crude membrane suspension was thawed. In binding assays, $^3$H-Bradykinin (0.06 nM) and drug (1 x $10^{-5}$M) were incubated with 50 µl of the membrane suspension at room temperature for 60 minutes in a final volume of 250 µl. Separation of receptor-bound from free $^3$H-Bradykinin is achieved by immediate filtration under vacuum and washed three times with 5 ml of ice-cold buffer (50 mM Tris-HCl pH 7.5). Non-specific binding was defined as binding in the presence of 0.1 µM Bradykinin. The radioactivity retained on rinsed filters was determined by a liquid-scintillation counter.
(ii) Test Results

<table>
<thead>
<tr>
<th>Test Compound (Example No.)</th>
<th>Inhibition % of $^3$H-Bradykinin binding (concentration: $1 \times 10^{-5}$M)</th>
</tr>
</thead>
<tbody>
<tr>
<td>35</td>
<td>99</td>
</tr>
<tr>
<td>40-(21)</td>
<td>97</td>
</tr>
<tr>
<td>47</td>
<td>100</td>
</tr>
<tr>
<td>49</td>
<td>100</td>
</tr>
<tr>
<td>60</td>
<td>100</td>
</tr>
<tr>
<td>61-(1)</td>
<td>95</td>
</tr>
<tr>
<td>61-(5)</td>
<td>99</td>
</tr>
<tr>
<td>61-(7)</td>
<td>100</td>
</tr>
<tr>
<td>61-(11)</td>
<td>98</td>
</tr>
<tr>
<td>61-(13)</td>
<td>98</td>
</tr>
<tr>
<td>61-(16)</td>
<td>100</td>
</tr>
<tr>
<td>61-(18)</td>
<td>97</td>
</tr>
<tr>
<td>61-(23)</td>
<td>100</td>
</tr>
<tr>
<td>61-(28)</td>
<td>99</td>
</tr>
<tr>
<td>61-(41)</td>
<td>100</td>
</tr>
<tr>
<td>61-(51)</td>
<td>100</td>
</tr>
<tr>
<td>61-(54)</td>
<td>100</td>
</tr>
</tbody>
</table>

The effects of the compound [I] on bradykinin-induced bronchoconstriction and carrageenan-induced paw edema were measured according to similar manners described in British Journal of Pharmacology, **102**, 774-777 (1991).

For therapeutic purpose, the compound [I] and a pharmaceutically acceptable salt thereof of the present invention can be used in a form of pharmaceutical preparation containing one of said compounds, as an active
ingredient, in admixture with a pharmaceutically acceptable carrier such as an organic or inorganic solid, semi-solid or liquid excipient suitable for oral, parenteral such as intravenous, intramuscular, subcutaneous or intraarticular, external such as topical, enteral, intrarectal, transvaginal, inhalant, ophthalmic, nasal or hypoglossal administration. The pharmaceutical preparations may be capsules, tablets, dragees, granules, suppositories, solution, lotion, suspension, emulsion, ointment, gel, cream, or the like. If desired, there may be included in these preparations, auxiliary substances, stabilizing agents, wetting or emulsifying agents, buffers and other commonly used additives.

While the dosage of the compound [I] will vary depending upon the age and condition of the patient, an average single dose of about 0.1 mg, 1 mg, 10 mg, 50 mg, 100 mg, 250 mg, 500 mg and 1000 mg of the compound [I] may be effective for preventing and/or treating the above-mentioned diseases. In general, amounts between 0.1 mg/body and about 1,000 mg/body may be administered per day.

The following Preparations and Examples are given for the purpose of illustrating this invention.

Preparation 1

Concentrated sulfuric acid (2.8 ml) was added dropwise to 70% nicinic acid (4.0 ml) in an ice-water bath. This mixture was added to a solution of 2,4,6-trichlorobenzoic acid (5.13 g) in concentrated sulfuric acid (23 ml) dropwise for 20 minutes in an ice-water bath. The mixture was stirred for 16 hours at ambient temperature and poured into ice-water (300 ml) slowly. This mixture was stirred for 1 hour at ambient temperature. The precipitate was collected by vacuum filtration and washed with water to give 2,4,6-trichloro-3-nitrobenzoic acid (5.26 g) as colorless fine crystals.
Preparation 2
To a solution of 2,6-dichloro-3-nitrotoluene (50 g) in acetic acid (400 ml) and ethanol (200 ml) was added iron (67.8 g) and the mixture was refluxed for 1.5 hours under nitrogen atmosphere. The insoluble material was filtered off and the filtrate was concentrated. To the residue was added a saturated aqueous solution of sodium bicarbonate and the mixture was extracted with ethyl acetate three times. The combined organic layers were washed with a saturated aqueous solution of sodium bicarbonate three times and brine, dried over magnesium sulfate, and concentrated in vacuo to give 2,4-dichloro-3-methylaniline (41 g).

\[
\begin{align*}
\text{mp} & : 54-56\degree C \\
\text{NMR (CDCl}_3, \delta) & : 2.45 (3H, s), 4.06 (2H, br s), 6.58 (1H, d, J=9Hz), 7.07 (1H, d, J=9Hz)
\end{align*}
\]

Preparation 3
(1) To a mixture of 2,4-dichloro-6-mercapto-3-methylaniline (861 mg), sodium hydrogen carbonate (1.16 g), tetrahydrofuran (9 ml) and water (9 ml) was added chloroacetyl chloride (0.36 ml) dropwise under an ice-water bath cooling. The mixture was stirred under an ice-water bath cooling for 15 minutes and then heated under reflux for 1 hour. The reaction mixture was cooled and poured into a mixture of dichloromethane and brine.

The aqueous layer and insoluble precipitate was extracted twice with a mixture of dichloromethane and methanol (4:1, V/V). The organic layer and extracts were combined, dried over magnesium sulfate and evaporated in vacuo to give an yellow powder of 2H-5,7-dichloro-6-methyl-1,4-benzothiazin-3(4H)-one (801 mg). This powder was used for
the next step without further purification.

(2) To a solution of 2H-5,7-dichloro-6-methyl-1,4-benzothiazin-3(4H)-one (846 mg) in N,N-dimethylformamide (17 ml) was added sodium hydride (40% in oil, 150 mg) under an ice-water bath cooling. The mixture was stirred at ambient temperature for 30 minutes and then, methyl iodide (0.4 ml) was added thereto. After stirring for 30 minutes, the mixture was partitioned between ethyl acetate and brine. The aqueous layer was extracted with ethyl acetate. The organic layers were combined, dried over magnesium sulfate and evaporated in vacuo. The residue was purified by silica gel column chromatography (dichloromethane-n-hexane) followed by crystallization from n-hexane to give 2H-5,7-dichloro-4,6-dimethyl-1,4-benzothiazin-3(4H)-one (590 mg) as pale yellow crystals.

mp : 98-99°C  
NMR (CDCl₃, δ) : 2.50 (3H, s), 3.35 (2H, s), 3.40 (3H, s), 7.39 (1H, s)

Preparation 4
To a solution of 2,4-dichloro-6-mercapto-3-methylaniline (2.08 g) in dichloromethane (40 ml) was added 1,1'-carbonyldiimidazole (1.78 g) at ambient temperature. The mixture was stirred for one hour at the same temperature. The separated precipitate was collected by filtration, washed with dichloromethane, and dried to give 4,6-dichloro-5-methyl-2-benzothiazolinone (1.40 g).

mp : >250°C  
NMR (DMSO-d₆, δ) : 2.42 (3H, s), 7.26 (1H, s)

Preparation 5
The following compounds were obtained according to a similar manner to that of Preparation 3-(2).
Preparation 6

The following compounds were obtained according to similar manners to those of Example 14 or .5 mentioned below.

(1) 4,6-Dichloro-3.5-dimethyl-2-benzothiazolinone

mp : 120-122°C
NMR (CDCl₃, δ) : 2.52 (3H, s), 3.87 (3H, s), 7.33 (1H, s)

(2) 4,6-Dichloro-3-ethyl-5-methyl-2-benzothiazolinone

mp : 101-103°C
NMR (CDCl₃, δ) : 1.39 (3H, t, J=7.5Hz), 2.53 (3H, s), 4.49 (2H, q, J=7.5Hz), 7.36 (1H, s)

(3) 4,6-Dichloro-3-(phthalimidoacetyl)aminotoluene

mp : 245-246°C
The following compounds were obtained according to a similar manner to that of Example 23 mentioned below.

(1) 3-(N-Acetyl-N-methylamino)-2,6-dichlorotoluene
   mp: 118-119°C
   NMR (CDCl₃, δ): 2.48 (3H, s), 4.59 (2H, s), 7.27 (1H, d, J=9Hz), 7.70-7.96 (4H), 8.00 (1H, br s), 8.12 (1H, d, J=9Hz)

(2) 3-(N-Acetoxyacetyl-N-methylamino)-2,6-dichlorotoluene
   mp: 107-108°C
   NMR (CDCl₃, δ): 2.13 (3H, s), 2.55 (3H, s), 3.18 (3H, s), 4.16, 4.44 (each 1H, d, J=15Hz), 7.19, 7.40 (each 1H, d, J=9Hz)

(3) 2,6-Dichloro-3-[N-(phthalimidoacetyl)-N-methylamino]toluene
   mp: 193-194°C
   NMR (CDCl₃, δ): 2.58 (3H, s), 3.21 (3H, s), 4.10 (2H, s), 7.30 (1H, d, J=9Hz), 7.42 (1H, d, J=9Hz), 7.65-7.91 (4H)

Preparation 8
To a solution of 3-(4-chlorobutyryl)amino-2,6-dichlorotoluene (2.80 g) in N,N-dimethylformamide (30 ml) was added sodium hydride (60% oil dispersion, 440 mg) in one portion at 5°C. The mixture was stirred for 10 minutes at 5°C and then for 2 hours at 60°C. The cooled mixture was poured into ice water. The separated oil was extracted with dichloromethane. The organic layer was
washed with water three times, dried, and concentrated in vacuo. The residue was purified by flash chromatography on silica gel to give 1-(2,4-dichloro-3-methylphenyl)-2-pyrrolidinone (1.95 g).

\[ \text{mp: 77-82°C} \]

NMR (CDCl$_3$, $\delta$): 2.25 (2H, m), 2.50 (3H, s), 2.58 (2H, t, $J=8\text{Hz}$), 3.75 (2H, t, $J=8\text{Hz}$), 7.08 (1H, d, $J=8\text{Hz}$), 7.33 (1H, d, $J=8\text{Hz}$)

**Preparation 9**

A mixture of 4,6-dichloro-3,5-dimethyl-2-benzothiazolinone (1.30 g), N-bromosuccinimide (1.03 g), 2,2'-azobis(2,4-dimethyl-4-methoxyvaleronitrile) (65 mg) and dichloromethane (26 ml) was heated under reflux for 2 hours. N-Bromosuccinimide (500 mg) was added therein and the mixture was heated under reflux for additional 4 hours. The reaction mixture was washed with water twice and brine, dried over magnesium sulfate and evaporated in vacuo. The residue was crystallized from diethyl ether to give 5-bromomethyl-4,6-dichloro-3-methyl-2-benzothiazolinone (1.20 g) as crystals.

\[ \text{mp: 142-143°C} \]

NMR (CDCl$_3$, $\delta$): 3.89 (3H, s), 4.82 (2H, s), 7.40 (1H, s)

**Preparation 10**

The following compounds were obtained according to a similar manner to that of Preparation 9.

(1) 3-Bromomethyl-2,4-dichloro-N-trifluoroacetyl-N-methylaniline

NMR (CDCl$_3$, $\delta$): 3.31 (3H, s), 4.78 (2H, s), 7.23 (1H, d, $J=9\text{Hz}$), 7.43 (1H, d, $J=9\text{Hz}$)

(2) N-Acetyl-3-bromomethyl-2,4-dichloro-N-methylaniline
mp : 123°C (dec.)
NMR (CDCl₃, δ) : 1.81 (3H, s), 3.19 (3H, s), 4.79 (2H, s), 7.22 (1H, d, J=9Hz), 7.43 (1H, d, J=9Hz)

(3) 1-(3-Bromomethyl-2,4-dichlorophenyl)-2-pyrrolidinone
mp : 106-109°C
NMR (CDCl₃, δ) : 2.17-2.38 (2H, m), 2.59 (2H, t, J=8Hz), 3.79 (2H, t, J=6Hz), 4.77 (2H, s), 7.21, 7.40 (each 1H, d, J=9Hz)

(4) N-Acetoxyacetyl-3-bromomethyl-2,4-dichloro-N-methylaniline
mp : 92-93°C
NMR (CDCl₃, δ) : 2.12 (3H, s), 3.21 (3H, s), 4.16, 4.44 (each 1H, d, J=15Hz), 4.79 (2H, s), 7.31, 7.48 (each 1H, d, J=9Hz)

(5) 3-Bromomethyl-2,4-dichloro-N-methyl-N-(phthalimidoacetyl)aniline
mp : 211°C (dec.)
NMR (CDCl₃, δ) : 3.24 (3H, s), 4.09 (2H, s), 4.81 (2H, s), 7.44 (1H, d, J=9Hz), 7.51 (1H, d, J=9Hz), 7.68-7.91 (4H)

(6) N,N-Di-(tert-butoxycarbonyl)-2,4-dichloro-3-bromomethylaniline
NMR (CDCl₃, δ) : 1.40 (18H, s), 4.79 (2H, s), 7.13 (1H, d, J=9Hz), 7.35 (1H, d, J=9Hz)

(7) 5-Bromomethyl-4,6-dichloro-3-ethyl-2-benzothiazolinone
mp : 120-126°C
NMR (CDCl₃, δ) : 1.41 (3H, t, J=7.5Hz), 4.50 (2H, q, J=7.5Hz), 4.85 (2H, s), 7.42 (1H, s)
(8) 2H-6-Bromomethyl-5,7-dichloro-4-methyl-1,4-
benzothiazin-3(4H)-one
NMR (CDCl₃, δ): 3.38 (2H, s), 3.42 (3H, s),
4.78 (2H, s), 7.44 (1H, s)

Preparation 11
To a solution of 2,4-dichloro-1-isopropoxybenzene
(3.53 g) in tetrahydrofuran (35 ml) was added n-butyl
lithium solution (1.6 Mol solution in n-hexane, 11 ml)
through a syringe at -78°C. After stirring for one hour
at -78°C, the mixture was poured into dry diethyl ether
(50 ml) containing pulverized dry ice in a few minutes.
After warming to ambient temperature, the mixture was
concentrated in vacuo. The residue was partitioned
between diethyl ether (50 ml) and aqueous 10% sodium
hydroxide solution (100 ml). The aqueous layer was
adjusted to pH 2 with 10% hydrochloric acid. The
separated oil was extracted with dichloromethane. The
extract was washed with water, dried, and evaporated under
reduced pressure. The residue was crystallized from
n-hexane to give 2,6-dichloro-3-isopropoxybenzoic acid
(3.0 g).
mp: 133-138°C
NMR (CDCl₃, δ): 1.39 (6H, d, J=6Hz), 4.55 (1H, m),
6.95 (1H, d, J=10Hz), 7.29 (1H, d, J=10Hz)

Preparation 12
To a solution of 2,6-dichloro-3-isopropoxybenzoic
acid (2.49 g) in tetrahydrofuran (30 ml) was added
borane-methyl sulfide complex (2 ml, 10 Mol solution)
through a syringe at ambient temperature. The mixture was
refluxed for half an hour, cooled, and quenched with
aqueous saturated ammonium chloride solution. The organic
layer was dried and concentrated in vacuo. The residue
was purified by flash chromatography on silica gel to give
2,6-dichloro-3-isopropoxybenzyl alcohol as a colorless oil (1.66 g).

\[
\begin{align*}
\text{NMR (CDCl}_3, \delta) : & \quad 1.38 (6H, d, J=5Hz), 2.15 (1H, t, J=7.5Hz), 4.52 (1H, m), 4.97 (2H, d, J=7.5Hz), \\
& \quad 6.87 (1H, d, J=10Hz), 7.25 (1H, d, J=10Hz)
\end{align*}
\]

Preparation 13

The following compounds were obtained according to a similar manner to that of Preparation 12.

1. **2, 6-Dichloro-3-methoxybenzyl alcohol**
   - mp: 99-101°C
   - NMR (CDCl\(_3\), \delta): 2.12 (1H, br s), 3.90 (3H, s), 4.98 (2H, s), 6.86 (1H, d, J=9Hz), 7.30 (1H, d, J=9Hz)

2. **2, 4, 6-Trichloro-3-nitrobenzyl alcohol**
   - mp: 121-124°C
   - NMR (CDCl\(_3\), \delta): 2.08 (1H, t, J=6Hz), 4.98 (2H, d, J=6Hz), 7.57 (1H, s)

3. **2, 6-Dichloro-3-nitrobenzyl alcohol**
   - mp: 99-100°C
   - NMR (CDCl\(_3\), \delta): 5.03 (2H, s), 7.50 (1H, d, J=8Hz), 7.72 (1H, d, J=8Hz)

Preparation 14

To a mixture of 2,6-dichloro-3-isopropoxybenzyl alcohol (1.65 g) and triethylamine (808 mg) in dichloromethane was added methanesulfonyl chloride (884 mg) in a few minutes at 5°C. After stirring for half an hour at 5°C, the mixture was washed with diluted hydrochloric acid and then with aqueous sodium bicarbonate solution, dried, and concentrated in vacuo. The residue was crystallized from n-hexane to give 2,6-dichloro-3-
isopropoxybenzyl methanesulfonate (2.07 g).

\[ \text{mp: 78-81°C} \]

\[ \text{NMR (CDCl}_3, \delta): 1.49 (6H, d, J=5Hz), 3.10 (3H, s), 4.56 (1H, m), 5.54 (2H, s), 6.98 (1H, d, J=10Hz), 7.31 (1H, d, J=10Hz) \]

**Preparation 15**

The following compounds were obtained according to a similar manner to that of Preparation 14.

1. **2,6-Dichloro-3-methoxybenzyl methanesulfonate**
   
   \[ \text{NMR (CDCl}_3, \delta): 3.10 (3H, s), 3.91 (3H, s), 5.53 (2H, s), 6.98 (1H, d, J=9Hz), 7.46 (1H, d, J=9Hz) \]

2. **2,4,6-Trichloro-3-nitrobenzyl methanesulfonate**
   
   \[ \text{mp: 113-114°C} \]
   \[ \text{NMR (CDCl}_3, \delta): 3.12 (3H, s), 5.50 (2H, s), 7.63 (1H, s) \]

3. **2,6-Dichloro-3-nitrobenzyl methanesulfonate**
   
   \[ \text{mp: 78-80°C} \]
   \[ \text{NMR (CDCl}_3, \delta): 3.13 (3H, s), 5.60 (2H, s), 7.57 (1H, d, J=8Hz), 7.85 (1H, d, J=8Hz) \]

**Preparation 16**

A mixture of 8-hydroxy-2-methylimidazo[1,2-a]pyridine (207 mg), 2,6-dichloro-3-nitrobenzyl bromide (400 mg) and potassium carbonate (580 mg) in N,N-dimethylformamide (8 ml) was stirred for 2 hours at 60°C. The mixture was cooled and diluted with water. The separated oil was extracted with dichloromethane. The organic layer was washed with water, dried, and concentrated in vacuo. The residue was purified by flash chromatography on silica gel to give 8-(2,6-dichloro-3-nitrobenzyloxy)-2-methylimidazo-
[1,2-a]pyridine (120 mg) as crystals.

mp : 183-185°C

NMR (CDCl₃, δ): 2.43 (3H, s), 5.50 (2H, s), 6.59 (1H, d, J=7.5Hz), 6.66 (1H, t, J=7.5Hz), 7.32 (1H, s), 7.52 (1H, d, J=9Hz), 7.75 (1H, d, J=7.5Hz), 7.79 (1H, d, J=9Hz)

Preparation 17

A mixture of 8-hydroxy-2-methylimidazo[1,2-a]pyridine (296 mg), 1,3-dichloro-2-(2-mesyloxyethyl)benzene (645 mg) and potassium carbonate (828 mg) in N,N-dimethylformamide (12 ml) was stirred for 6 hours at 70°C. The mixture was treated according to a similar manner to that of Preparation 16 to give 8-[(2-(2,6-dichlorophenyl)ethyloxy]-2-methylimidazo[1,2-a]pyridine (150 mg).

mp : 92-94°C

NMR (CDCl₃, δ): 2.48 (3H, s), 3.69 (2H, m), 4.29 (2H, m), 6.49 (1H, d, J=7.5Hz), 6.61 (1H, t, J=7.5Hz), 7.15 (1H, dd, J=7Hz and 5Hz), 7.29-7.35 (3H, m), 7.69 (1H, d, J=7.5Hz)

Preparation 18

The following compounds were obtained according to similar manners to those of Preparation 16 or 17.

(1) 2-Amino-3-(2,6-dichlorobenzyloxy)-6-methylpyridine

mp : 140-141°C

NMR (CDCl₃, δ): 2.35 (3H, s), 4.61 (2H, br s), 5.28 (2H, s), 6.49 (1H, d, J=8Hz), 7.08 (1H, d, J=8Hz), 7.20-7.41 (3H)

(2) 8-[1-(2,6-Dichlorophenyl)ethyloxy]-2-methylimidazo-[1,2-a]pyridine

mp : 173-174°C

NMR (CDCl₃, δ): 1.93 (3H, d, J=7Hz), 2.48 (3H, s),
6.06-6.22 (2H), 6.42 (1H, t, J=7Hz), 7.06-7.32 (4H), 7.59 (1H, d, J=7Hz)

(3) 8-[2,6-Dichloro-3-(N-methyl-N-acetylamino)-benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 159-161°C
NMR (CDCl₃, δ) : 1.86 (3H, s), 2.44 (3H, s), 3.20 (3H, s), 5.50 (2H, s), 6.55-6.75 (2H), 7.29 (1H, d, J=9Hz), 7.33 (1H, s), 7.46 (1H, d, J=9Hz), 7.74 (1H, d, J=7Hz)

(4) 8-[2,6-Dichloro-3-(N-methyl-N-trifluoroacetylamino)-benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.42 (3H, s), 3.31 (3H, s), 5.48 (2H, s), 6.57-6.70 (2H, m), 7.30 (1H, d, J=10Hz), 7.33 (1H, s), 7.46 (1H, d, J=10Hz), 7.74 (1H, dd, J=7.5Hz and 2Hz)

(5) 8-[2,6-Dichloro-3-(N-methyl-N-tert-butoxycarbonylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine

(6) 8-[2,6-Dichloro-3-(N-methyl-N-mesylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 190-191°C

NMR (CDCl₃, δ) : 2.13-2.35 (2H, m), 2.42 (3H, s), 2.59 (2H, t, J=9Hz), 3.76 (2H, t, J=9Hz), 5.43 (2H, s), 6.54-6.75 (2H, m), 7.21-7.48 (3H, m), 7.73 (1H, d, J=6Hz)
(8) 8-(2,6-Dichloro-3-methoxybenzyloxy)-2-methylimidazo[1,2-a]pyridine

mp : 179-180°C

NMR (CDCl₃, δ) : 2.42 (3H, s), 3.91 (3H, s), 5.46 (2H, s), 6.55-6.72 (2H), 6.92 (1H, d, J=9Hz), 7.24-7.36 (2H), 7.71 (1H, d, J=7Hz)

(9) 8-(2,6-Dichloro-3-isopropoxybenzyloxy)-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 1.40 (6H, d, J=5Hz), 2.45 (3H, s), 4.57 (1H, m), 5.45 (2H, s), 6.59-6.70 (2H, m), 6.92 (1H, d, J=10Hz), 7.25-7.31 (2H, m), 7.72 (1H, dd, J=7.5Hz and 1.5Hz)

(10) 8-(2,4,6-Trichloro-3-nitrobenzyloxy)-2-methylimidazo[1,2-a]pyridine

mp : 184-186°C

NMR (CDCl₃, δ) : 2.44 (3H, s), 5.47 (2H, s), 6.57 (1H, d, J=8Hz), 6.67 (1H, t, J=8Hz), 7.33 (1H, s), 7.58 (1H, s), 7.76 (1H, d, J=8Hz)

(11) 8-[2-Chloro-5-(N-methyl-N-acetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.49 (3H, s), 3.21 (3H, s), 5.43 (2H, s), 6.40 (1H, d, J=7.5Hz), 6.58 (1H, t, J=7.5Hz), 7.10 (1H, dd, J=10Hz and 1.5Hz), 7.37 (1H, s), 7.46 (1H, d, J=10Hz), 7.49 (1H, d, J=1.5Hz), 7.74 (1H, d, J=7.5Hz)

(12) 8-[3-(N-Acetoxyacetyl-N-methylamino)-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 118-119°C

NMR (CDCl₃, δ) : 2.14 (3H, s), 2.42 (3H, s), 3.21 (3H, s), 4.19, 4.48 (each 1H, d, J=15Hz), 5.49 (1H, s), 6.54-6.72 (2H), 7.32 (1H, s), 7.40-7.49
(each 1H, d, J=9Hz), 7.74 (1H, d, J=7Hz)

(13) 8-[2,6-Dichloro-3-(N-methyl-N-phthalimidoacetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 221-223°C
NMR (CDCl₃, δ) : 2.42 (3H, s), 3.22 (3H, s), 4.11 (2H, s), 5.52 (2H, s), 6.59-6.73 (2H), 7.32 (1H, s), 7.52 (2H, s), 7.67-7.92 (4H)

(14) 2-Methyl-8-(2-trifluoromethylbenzyloxy)imidazo[1,2-a]pyridine
mp : 131-132°C
NMR (CDCl₃, δ) : 2.48 (3H, s), 5.38 (2H, s), 6.35 (1H, d, J=7Hz), 6.55 (1H, t, J=7Hz), 7.32 (1H, s), 7.61 (5H, s), 7.69 (1H, d, J=7Hz)

(15) 8-(2-Methoxycarbonylbenzyloxy)-2-methylimidazo[1,2-a]pyridine
mp : 88-89°C
NMR (CDCl₃, δ) : 2.49 (3H, s), 3.92 (3H, s), 5.77 (2H, s), 6.38 (1H, d, J=7.5Hz), 6.55 (1H, d, J=7.5Hz), 7.33 (1H, s), 7.37 (1H, t, J=7.5Hz), 7.53 (1H, t, J=7.5Hz), 7.67 (1H, d, J=7.5Hz), 7.88 (1H, d, J=7.5Hz), 8.05 (1H, d, J=7.5Hz)

(16) 8-(2-Phenylbenzyloxy)-2-methylimidazo[1,2-a]pyridine
mp : 90-92°C
NMR (CDCl₃, δ) : 2.48 (3H, s), 5.19 (2H, s), 6.14 (1H, d, J=7.5Hz), 6.48 (1H, t, J=7.5Hz), 7.31-7.45 (9H, m), 7.63 (1H, d, J=7.5Hz), 7.75 (1H, m)

(17) 8-(2,6-Difluorobenzoyloxy)-2-methylimidazo[1,2-a]pyridine
mp : 114-116°C
NMR (CDCl₃, δ) : 2.45 (3H, s), 5.31 (2H, s), 6.56-6.67 (2H, m), 6.86-6.99 (2H, m), 7.25-7.41 (1H, m), 7.32 (1H, s), 7.70 (1H, d, J=7.5Hz)

15
(18) 8-(2,6-Dibromobenzyl)oxy)-2-methylimidazo[1,2-a]-pyridine
mp : 159-162°C
NMR (CDCl₃, δ) : 2.43 (3H, s), 5.49 (2H, s), 6.59 (1H, d, J=7Hz), 6.67 (1H, t, J=7Hz), 7.08 (1H, t, J=8Hz), 7.30 (1H, s), 7.58 (2H, d, J=8Hz), 7.71 (1H, d, J=7Hz)

15
(19) 8-(2-Chloro-6-fluorobenzyl)oxy)-2-methylimidazo[1,2-a]-pyridine
mp : 150-151°C
NMR (CDCl₃, δ) : 2.44 (3H, s), 5.38 (2H, s), 6.39 (1H, d, J=7Hz), 6.63 (1H, t, J=7Hz), 7.02 (1H, t, J=8Hz), 7.20-7.38 (3H), 7.70 (1H, d, J=7Hz)

20
(20) 8-(4-Bromo-2-fluorobenzyl)oxy)-2-methylimidazo[1,2-a]-pyridine
mp : 150-151°C
NMR (CDCl₃, δ) : 2.48 (3H, s), 5.31 (2H, s), 6.42 (1H, d, J=7.5Hz), 6.58 (1H, t, J=7.5Hz), 7.23-7.33 (3H, m), 7.49 (1H, t, J=7.5Hz), 7.69 (1H, d, J=7.5Hz)

25
(21) 8-(2-Chloro-5-nitrobenzyl)oxy)-2-methylimidazo[1,2-a]-pyridine
mp : 134-135°C
NMR (CDCl₃, δ) : 2.49 (3H, s), 5.48 (2H, s), 6.41 (1H, d, J=7Hz), 6.60 (1H, t, J=7Hz), 7.36 (1H, s), 7.60 (1H, d, J=9Hz), 7.74 (1H, d, J=7Hz), 8.16 (1H, dd, J=9Hz and 2Hz), 8.55 (1H, d, J=2Hz)
(22) 8-[2-Chloro-6-(N-methyl-N-acetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 1.90 (3H, s), 2.41 (3H, s), 3.23 (3H, s), 5.19 (1H, d, J=10Hz), 5.26 (1H, d, J=10Hz), 6.55 (1H, d, J=7.5Hz), 6.67 (1H, t, J=7.5Hz), 7.18 (1H, dd, J=7.5Hz and 2Hz), 7.33 (1H, s), 7.48-7.54 (2H, m), 7.74 (1H, dd, J=7.5Hz and 2Hz)

(23) 8-(2-Chloro-6-nitrobenzyloxy)-2-methylimidazo[1,2-a]pyridine
mp: 162-163°C
NMR (CDCl₃, δ): 2.41 (3H, s), 5.62 (2H, s), 6.51-6.69 (2H), 7.30 (1H, s), 7.48 (1H, t, J=8Hz), 7.63-7.75 (2H), 7.85 (1H, d, J=7Hz)

(24) 8-[3-(N,N-Di-tert-butoxycarbonylamino)-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine
mp: 180-181°C
NMR (CDCl₃, δ): 1.41 (18H, s), 2.42 (3H, s), 5.53 (2H, s), 6.51-6.68 (2H), 7.20 (1H, d, J=9Hz), 7.30 (1H, s), 7.38 (1H, d, J=9Hz), 7.70 (1H, d, J=7Hz)

(25) 2-Methyl-8-(2,4,6-trichlorobenzyloxy)imidazo[1,2-a]-pyridine
mp: 187-189°C
NMR (CDCl₃, δ): 2.42 (3H, s), 5.40 (2H, s), 6.58 (1H, d, J=7Hz), 6.67 (1H, t, J=7Hz), 7.31 (1H, s), 7.49 (2H, s), 7.71 (1H, d, J=7Hz)

(26) 8-(2,3,6-Trichlorobenzyloxy)-2-methylimidazo[1,2-a]-pyridine
mp: 148-149°C
NMR (CDCl₃, δ): 2.42 (3H, s), 5.48 (2H, s),
6.52-6.71 (2H), 7.30 (1H, d, J=9Hz), 7.31 (1H, s), 7.45 (1H, d, J=9Hz), 7.72 (1H, d, J=7Hz)

(27) 8-(2,6-Dibromo-4-methoxycarbonylbenzyloxy)-2-methylimidazo[1,2-a]pyridine
mp :  153-154°C
NMR (CDCl₃, δ) :  2.42 (3H, s), 3.97 (3H, s), 5.51 (2H, s), 6.58 (1H, d, J=7Hz), 6.68 (1H, t, J=7Hz), 7.31 (1H, s), 7.74 (1H, d, J=7Hz), 8.21 (2H, s)

(28) 8-(4-Benzoyl-2-chlorobenzyloxy)-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) :  2.49 (3H, s), 5.49 (2H, s), 6.40 (1H, d, J=8Hz), 6.60 (1H, d, J=8Hz), 7.35 (1H, s), 7.44-7.88 (9H)

(29) 8-(2,6-Dichloro-4-benzoylbenzyloxy)-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) :  2.45 (3H, s), 5.50 (2H, s), 6.57-6.72 (2H), 7.28-7.92 (9H)

(30) 8-[4-(2-Cyanophenyl)benzoyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) :  2.49 (3H, s), 5.38 (2H, s), 6.44 (1H, d, J=8Hz), 6.58 (1H, t, J=8Hz), 7.32 (1H, s), 7.38-7.82 (9H)

(31) 8-(6-Chloropiperonyl)methoxy-2-methylimidazo[1,2-a]pyridine
mp :  141-142°C
NMR (CDCl₃, δ) :  2.49 (3H, s), 5.31 (2H, s), 5.96 (2H, s), 6.38 (1H, d, J=7Hz), 6.58 (1H, t, J=7Hz), 6.86 (1H, s), 7.11 (1H, s), 7.31 (1H, s), 7.68 (1H, d, J=7Hz)
(32) 8-(2-Bromothiophen-4-yl)methoxy-2-methylimidazo-[1,2-a]pyridine  
mp : 127-128°C  
NMR (CDCl₃, δ) : 2.46 (3H, s), 5.42 (2H, s),  
6.48 (1H, d, J=7Hz), 6.59 (1H, t, J=7Hz), 7.09  
(1H, s), 7.16-7.35 (2H), 7.69 (1H, d, J=7Hz)  

(33) 8-(3-Chlorobenzofuran-2-yl)methoxy-2-methylimidazo-[1,2-a]pyridine  
mp : 141-143°C  
NMR (CDCl₃, δ) : 2.45 (3H, s), 5.44 (2H, s),  
6.56-6.66 (2H, m), 7.29-7.71 (6H, m)  

(34) 8-[3-(4-Chlorophenyl)-5-methylbenzofuran-2-yl]-methoxy-2-methylimidazo[1,2-a]pyridine  
mp : 140-141°C  
NMR (CDCl₃, δ) : 2.45 (6H, s), 5.33 (2H, s), 6.49  
(1H, d, J=7Hz), 6.58 (1H, t, J=7Hz), 7.12-7.56  
(8H), 7.70 (1H, d, J=7Hz)  

(35) 8-(4,6-Dichloro-3-methyl-2-benzothiazolinon-5-yl)-methoxy-2-methylimidazo[1,2-a]pyridine  
mp : 236-237°C  
NMR (CDCl₃, δ) : 2.42 (3H, s), 3.87 (3H, s), 5.50  
(2H, s), 6.56-6.72 (2H, m), 7.32 (1H, s), 7.42  
(1H, s), 7.72 (1H, d, J=7Hz)  

(36) 8-(4,6-Dichloro-3-ethyl-2-benzothiazolinon-5-yl)-methoxy-2-methylimidazo[1,2-a]pyridine  
mp : 202-205°C  
NMR (CDCl₃, δ) : 1.41 (3H, t, J=7.5Hz), 2.45 (3H,  
s), 4.49 (2H, q, J=7.5Hz), 5.53 (2H, s),  
6.59-6.72 (2H, m), 7.31 (1H, s), 7.44 (1H, s),  
7.74 (1H, dd, J=7.5Hz and 1.5Hz)
(37) 8-(5,7-Dichloro-4-methyl-1,4-benzothiazin-3(4H)-on-6-yl)methoxy-2-methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, δ) : 2.44 (3H, s), 3.40 (2H, s), 3.43 (3H, s), 5.45 (2H, s), 6.58 (1H, d, J=8Hz), 6.67 (1H, t, J=8Hz), 7.32 (1H, s), 7.47 (1H, s), 7.73 (1H, d, J=8Hz)

Preparation 19
A mixture of 2-amino-3-(2,6-dichlorobenzyloxy)-10-pyridine (1.345 g) and 3-bromo-1,1,1-trifluoroacetone (1.147 g) in ethanol (26 ml) was refluxed for 5 hours, cooled, and concentrated in vacuo. The residue was partitioned between ethyl acetate and aqueous sodium bicarbonate solution. The organic layer was washed with water, dried, and concentrated under reduced pressure to give a solid, which was purified by flash chromatography on silica gel to give 8-(2,6-dichlorobenzyloxy)-2-trifluoromethylimidazo[1,2-a]pyridine as a white solid (803 mg).

mp : 184-185°C
NMR (CDCl$_3$, δ) : 5.50 (2H, s), 6.75 (1H, d, J=7.5Hz), 6.85 (1H, t, J=7.5Hz), 7.23-7.38 (3H, m), 7.83 (1H, d, J=7.5Hz), 7.88 (1H, s)

Preparation 20
The following compounds were obtained according to a similar manner to that of Preparation 19.

(1) 8-(2,6-Dichlorobenzyloxy)-2-ethylimidazo[1,2-a]-pyridine
mp : 153-154°C
NMR (CDCl$_3$, δ) : 1.30 (3H, t, J=7Hz), 2.84 (2H, q, J=7Hz), 5.46 (2H, s), 6.55-6.71 (2H), 7.19-7.42 (4H), 7.73 (1H, d, J=7Hz)
(2) Ethyl 8-(2,6-dichlorobenzyloxy)imidazo[1,2-a]-pyridine-2-carboxylate
mp : 176-178°C
NMR (CDCl₃, δ) : 1.40 (3H, t, J=7Hz), 4.42 (2H, q, J=7Hz), 5.49 (2H, s), 6.69 (1H, d, J=7Hz), 6.81 (1H, t, J=7Hz), 7.21-7.43 (3H), 7.81 (1H, d, J=7Hz), 8.18 (1H, s)

(3) 8-(2,6-Dichlorobenzyloxy)-2,5-dimethylimidazo[1,2-a]-pyridine
mp : 140-141°C
NMR (CDCl₃, δ) : 2.47 (6H, s), 5.44 (2H, s), 6.45 (1H, d, J=7Hz), 6.60 (1H, d, J=7Hz), 7.15-7.40 (4H)

(4) 8-Amino-2,7-dimethylimidazo[1,2-a]pyridine dihydrochloride
mp : >250°C
NMR (DMSO-d₆, δ) : 2.25 (3H, s), 2.46 (3H, s), 6.82 (2H, br s), 7.10 (1H, d, J=7Hz), 7.90 (1H, s), 8.03 (1H, d, J=7Hz)

Preparation 21
8-(2,6-Dichlorophenyl)methylamino-2,7-dimethylimidazo[1,2-a]pyridine (100 mg) was obtained by reacting 8-amino-2,7-dimethylimidazo[1,2-a]pyridine (100 mg) with 2,6-dichlorobenzaldehyde (191 mg) according to a similar manner to that of Example 10 mentioned below.
mp : 87-88°C
NMR (CDCl₃, δ) : 2.26 (3H, s), 2.40 (3H, s), 4.46 (1H, br s), 4.99 (2H, d, J=5Hz), 6.41 (1H, d, J=7Hz), 7.07-7.40 (4H), 7.53 (1H, d, J=7Hz)

Preparation 22
8-(2,6-Dichlorophenyl)methylamino-2-methylimidazo-
[1,2-a]pyridine was obtained according to a similar manner to that of Preparation 21.

mp : 119-121°C

NMR (CDCl₃, δ) : 2.38 (3H, s), 4.69 (2H, d, J=4Hz), 5.19 (1H, t, J=4Hz), 6.29 (1H, d, J=7.5Hz), 6.63 (1H, t, J=7.5Hz), 7.15-7.36 (3H, m), 7.32 (1H, s), 7.47 (1H, d, J=7.5Hz)

Preparation 23

To a suspension of sodium hydride (60% oil dispersion, 17 mg) was added 8-acetylamino-2-methylimidazo[1,2-a]pyridine (73 mg), and the mixture was stirred for 30 minutes, 2,6-dichlorobenzyl bromide (97 mg) was added thereto, and the mixture was stirred for 1 hour. Water was added thereto, and the mixture was extracted with methylene chloride three times. The combined organic layer was washed with water four times and brine, dried over magnesium sulfate, and concentrated in vacuo. The residue was crystallized with diethyl ether to give

8-[N-(2,6-dichlorophenyl)methyl-N-acetylamino]-2-methylimidazo[1,2-a]pyridine (85 mg).

mp : 177-178°C

NMR (CDCl₃, δ) : 1.95 (3H, s), 2.48 (3H, s), 4.92 (1H, br d, J=15Hz), 6.00 (1H, br d, J=15Hz), 6.36 (1H, d, J=7Hz), 6.45 (1H, t, J=7Hz), 7.00-7.21 (3H), 7.38 (1H, s), 7.98 (1H, d, J=7Hz)

Preparation 24

To a solution of 8-[2,6-dichloro-3-(2-pyrroldin-1-yl)benzyl]-2-methylimidazo[1,2-a]pyridine (120 mg) in tetrahydrofuran (3 ml) was added lithium aluminum hydride (19 mg) under ice-cooling, and the mixture was stirred for 2 hours. A saturated aqueous ammonium chloride solution was added thereto, and insoluble material was filtered
off. The filtrate was concentrated, and the residue was purified by preparative thin-layer chromatography (5% solution of methanol in methylene chloride) to give 8-[2, 6-dichloro-3-(1-pyrrolidinyl)benzylxoy]-2-methylimidazo[1, 2-a]pyridine (23 mg).

\[ \text{NMR (CDCl}_3, \delta) : 1.86-2.01 (4H, m), 2.41 (3H, s), 3.26-3.40 (4H, m), 5.44 (2H, s), 6.57-6.72 (2H, m), 6.90, 7.19 (each 1H, d, J=9Hz), 7.31 (1H, s), 7.72 (1H, d, J=7Hz) \]

**Preparation 25**

8-(3-Amino-2, 6-dichlorobenzylxoy)-2-methylimidazo-[1, 2-a]pyridine was obtained according to a similar manner to that of Example 4 mentioned below.

\[ \text{NMR (DMSO-d}_6, \delta) : 2.27 (3H, s), 5.30 (2H, s), 5.70 (2H, s), 6.64-7.00 (3H), 7.24 (1H, d, J=8Hz), 7.65 (1H, s), 8.09 (1H, d, J=7Hz) \]

**Preparation 26**

8-(2, 6-Dichloro-3-methylaminobenzylxoy)-2-methylimidazo[1, 2-a]pyridine was obtained according to a similar manner to that of Example 7 mentioned below.

mp : 167-168°C

\[ \text{NMR (CDCl}_3, \delta) : 2.40 (3H, s), 2.91 (3H, d, J=5Hz), 4.41 (1H, br d, J=5Hz), 5.48 (2H, s), 6.56-6.78 (3H), 7.18 (1H, d, J=9Hz), 7.33 (1H, br s), 7.80 (1H, br d, J=6Hz) \]

**Preparation 27**

8-[2, 6-Dichloro-3-(N-methyl-N-acetylamino)benzylxoy]-2-methylimidazo[1, 2-a]pyridine was obtained according to a similar manner to that of Example 14 mentioned below.

mp : 159-161°C

\[ \text{NMR (CDCl}_3, \delta) : 1.86 (3H, s), 2.44 (3H, s), 3.20 (3H, s), 5.50 (2H, s), 6.55-6.75 (2H), 7.29 (1H, s) \]
Preparation 28

8-[2,6-Dichloro-3-(N-methoxycarbonyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Example 18 mentioned below.

NMR (CDCl₃, δ): 2.40 (3H, s), 3.2 (3H, s), 3.68 (3H, s), 5.45 (2H, s), 6.59-6.72 (2H), 7.21-7.49 (3H), 7.81 (1H, d, J=6Hz)

Preparation 29

Pivaloyl chloride (0.22 ml) was added dropwise to a mixture of acetylsarcosine (264 mg), N-methylmorpholine (0.22 ml) and N-methylpyrrolidone (30 ml) under a dry ice-tetrachloromethane bath cooling. This mixture was stirred for 10 minutes under ice-cooling, and 8-[3-amino-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine (500 mg) was added thereto under a dry ice-tetrachloromethane bath cooling. The mixture was stirred for 22 hours at ambient temperature. The mixture was partitioned between ethyl acetate and a saturated aqueous sodium bicarbonate solution, and the aqueous layer was extracted with ethyl acetate twice. The combined organic layer was dried over magnesium sulfate and concentrated in vacuo, and the residue was purified by flash column chromatography (methylene chloride:methanol = 50:1, V/V) to give 8-[3-(acetylsarcosylamino)-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine (260 mg).

NMR (CDCl₃, δ): 2.18 (3H, s), 2.42 (3H, s), 3.09 (0.3H, s), 3.19 (2.7H, s), 4.19 (2H, s), 5.45 (2H, s), 6.55-6.72 (2H), 7.30 (1H, s), 7.34 (1H, d, J=9Hz), 7.72 (1H, d, J=8Hz), 8.38 (1H, d, J=9Hz), 8.81 (1H, br s)
Preparation 30

8-[2,6-Dichloro-3-(N-ethyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Example 26 mentioned below.

NMR (DMSO-d$_6$, $\delta$) : 1.10 (3H, t, J=7Hz), 2.43 (3H, s), 2.75 (3H, s), 3.05 (2H, q, J=7Hz), 5.59 (2H, s), 7.28-7.62 (3H), 7.71 (1H, d, J=7Hz), 8.16 (1H, br s), 8.55 (1H, d, J=7Hz)

Preparation 31

(1) 3,5'-Dichlorobenzanilide was obtained according to a similar manner to that of Example 15 from 3,5-dichloroaniline and benzoyl chloride.

mp : 148-149°C

NMR (CDCl$_3$, $\delta$) : 7.15 (1H, t, J=0.5Hz), 7.44-7.60 (3H, m), 7.51 (2H, d, J=0.5Hz), 7.76-7.97 (3H, m)

(2) 3',5'-Dichloro-N-methylbenzanilide was obtained according to a similar manner to that of Example 23.

NMR (CDCl$_3$, $\delta$) : 3.46 (3H, s), 6.94 (2H, d, J=0.5Hz), 7.13 (1H, t, J=0.5Hz), 7.20-7.41 (5H, m)

(3) To a stirred solution of 3',5'-dichloro-N-methylbenzanilide (2.317 g) in tetrahydrofuran was added lithium aluminum hydride (380 mg) in an ice-bath and the resulting suspension was stirred at the same temperature for one hour. The reaction mixture was quenched with saturated ammonium chloride and filtered through celite pad. The inorganic material on the celite was washed with ethyl acetate. Methanol was added to the filtrate and stirred at ambient temperature for half an hour. The organic
layer was washed with water, dried over anhydrous magnesium sulfate and concentrated in vacuo. The residue was purified by flash chromatography eluting with ethyl acetate – n-hexane (1:9, V/V) to afford 3,5-dichloro-N-methylaniline (1.38 g) as a colorless oil.

NMR (CDCl₃, δ) : 2.81 (3H, s), 3.87 (1H, br s), 6.94 (2H, d, J=0.5Hz), 6.66 (1H, t, J=0.5Hz)

The mixture of 3,5-dichloro-N-methylaniline (345 mg) and excess amount of benzyl bromide and triethylamine in acetonitrile (7 ml) was refluxed for 4 hours. The reaction mixture was separated with ethyl acetate and water, and the organic layer was washed with water, dried and concentrated in vacuo to give N-benzyl-3,5-dichloro-N-methylaniline (532 mg).

NMR (CDCl₃, δ) : 3.01 (3H, s), 4.50 (2H, s), 6.57 (2H, d, J=0.5Hz), 6.67 (2H, t, J=0.5Hz), 7.10-7.46 (5H, m)

Phosphoryl chloride (12.7 ml) was dropwise added to N,N-dimethylformamide (70 ml), and the mixture was stirred for 30 minutes at ambient temperature. A solution of N-benzyl-3,5-dichloro-N-methylaniline (7.26 g) in N,N-dimethylformamide (30 ml) was dropwise added thereto, and the mixture was stirred for 30 minutes at ambient temperature and then for 30 minutes at 50°C. The reaction mixture was neutralized with 1N aqueous sodium hydroxide solution and extracted with ethyl acetate. The organic solution was washed with water and brine, dried over magnesium sulfate and concentrated in vacuo. The residue was purified by a flash chromatography (ethyl acetate : n-hexane = 1:6, V/V) to give N-benzyl-3,5-dichloro-4-formyl-N-methylaniline (5.70 g).
mp: 66-70°C  
NMR (CDCl$_3$, $\delta$): 3.10 (3H, s), 4.60 (2H, s), 6.65 (2H, s), 6.98-7.42 (5H, m), 10.30 (1H, s)

(6) To a solution of N-benzyl-3,5-dichloro-4-formyl-N-methylaniline (820 mg) in ethyl acetate (10 ml) was added palladium hydroxide (80 mg) under nitrogen atmosphere. This mixture was stirred under hydrogen atmosphere under atmospheric pressure at ambient temperature for one and half an hour. The precipitate was dissolved into chloroform and filtered through celite and the filtrate was concentrated in vacuo. The residual solid was suspended in diisopropyl ether and warmed at 90°C. After being stirred and cooled, the solid was collected by filtration to afford 3,5-dichloro-4-formyl-N-methylaniline (470 mg) as a pale brown solid.  
mp: 172-174°C  
NMR (CDCl$_3$, $\delta$): 2.90 (3H, t, J=5Hz), 4.50 (1H, br s), 6.50 (2H, s), 10.32 (1H, s)

(7) To a solution of 3,5-dichloro-4-formyl-N-methylaniline (242 mg) in methanol (3 ml) and tetrahydrofuran (3 ml) was added sodium borohydride (45 mg) and the mixture was stirred for 30 minutes at ambient temperature. The reaction mixture was quenched with aqueous saturated ammonium chloride solution and ethyl acetate was added thereto. The separated organic layer was washed with aqueous saturated ammonium chloride solution, dried and concentrated in vacuo to give 3,5-dichloro-4-hydroxymethyl-N-methylaniline (246 mg).  
mp: 108-111°C  
NMR (CDCl$_3$, $\delta$): 1.85 (1H, t, J=6Hz), 2.81 (3H, d,
3.5-Dichloro-4-hydroxymethyl-N-methylacetanilide was obtained according to a similar manner to that of Example 14.

NMR (CDCl₃, δ): 1.97 (3H, br s), 2.18 (1H, t, J=7Hz), 3.26 (3H, s), 4.96 (2H, d, J=7Hz), 7.23 (2H, s).

8-[2,6-Dichloro-4-(N-acetyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Preparation 14 and then Preparation 17.

NMR (CDCl₃, δ): 2.04 (3H, br s), 2.44 (3H, s), 3.27 (3H, s), 5.42 (2H, s), 6.60 (1H, dd, J=7Hz and 0.5Hz), 6.68 (1H, t, J=7Hz), 7.24 (2H, s), 7.33 (1H, s), 7.73 (1H, dd, J=7Hz and 0.5Hz)

Preparation 32
8-[3-(N-Glycyl-N-methylamino)-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Example 34.

mp: 144-147°C

NMR (CDCl₃, δ): 2.42 (3H, s), 3.00 (1H, d, J=17Hz), 3.12 (1H, d, J=17Hz), 3.22 (3H, s), 5.49 (2H, s), 6.56-6.72 (2H), 7.25 (1H, d, J=9Hz), 7.31 (1H, s), 7.45 (1H, d, J=9Hz), 7.72 (1H, d, J=7Hz)

Preparation 33
8-[2,6-Dichloro-3-[N-[N-(N,N-dimethylglycyl)glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Example 69 mentioned below.
Example 1

To a solution of 8-(2,6-dichloro-3-nitrobenzyloxy)-2-methylimidazo[1,2-a]pyridine (85 mg) in a mixture of ethanol (1 ml) and 1,4-dioxane (1 ml) was added in one portion N-bromosuccinimide (43 mg) at ambient temperature. After stirring for one hour at the same temperature, the mixture was filtered to give 3-bromo-8-(2,6-dichloro-3-nitrobenzyloxy)-2-methylimidazo[1,2-a]pyridine (85 mg) as a yellow solid.

mp : 217-219°C
NMR (DMSO-d$_6$, $\delta$): 2.31 (3H, s), 5.50 (2H, s)
7.0-7.04 (2H, m), 7.89-7.98 (2H, m),
8.23 (1H, d, $J$=9Hz)

Example 2

To a solution of 8-(2,6-dichlorobenzylxoxy)-2-methylimidazo[1,2-a]pyridine (100 mg) in ethanol (2 ml) was added in one portion N-chlorosuccinimide (65.3 mg) at ambient temperature. After stirring for 1 hour at the same temperature, water was added thereto, and the mixture was extracted with methylene chloride. The organic layer was washed with brine, dried and concentrated in vacuo. The residue was subjected to a column chromatography on silica gel eluting with 1% solution of methanol in methylene chloride. The desired residue was recrystallized with a mixture of benzene and n-hexane to give 3-chloro-8-(2,6-dichlorobenzylxoxy)-2-methylimidazo[1,2-a]pyridine (63 mg).
mp : 185-186°C
NMR (CDCl₃, δ) : 2.43 (3H, s), 5.48 (2H, s), 6.69 (1H, d, J=7Hz), 6.81 (1H, t, J=7Hz), 7.19-7.40 (3H, m), 7.69 (1H, d, J=7Hz)

Example 3
The following compounds were obtained according to similar manners to those of Examples 1 or 2.

(1) 3-Bromo-8-(2,6-dichlorobenzyloxy)-2-methylimidazo-[1,2-a]pyridine
mp : 173-174°C
NMR (CDCl₃, δ) : 2.44 (3H, s), 5.48 (2H, s), 6.70 (1H, d, J=7Hz), 6.82 (1H, t, J=7Hz), 7.19-7.41 (3H, m), 7.75 (1H, d, J=7Hz)

(2) 3-Chloro-8-(2,6-dichlorobenzyloxy)-2-trifluoromethylimidazo[1,2-a]pyridine
mp : 213-5°C
NMR (CDCl₃-CD₃OD, δ) : 5.53 (2H, s), 6.98 (1H, d, J=7.5Hz), 7.12 (1H, t, J=7.5Hz), 7.31-7.46 (3H, m), 7.93 (1H, d, J=7.5Hz)

(3) 3-Chloro-8-(2,6-dichlorobenzyloxy)-2-ethylimidazo-[1,2-a]pyridine
mp : 169-170°C
NMR (CDCl₃, δ) : 1.30 (3H, t, J=7Hz), 2.81 (2H, q, J=7Hz), 5.49 (2H, s), 6.69 (1H, d, J=7Hz), 6.72 (1H, t, J=7Hz), 7.19-7.41 (3H, m), 7.71 (1H, d, J=7Hz)

(4) Ethyl 3-chloro-8-(2,6-dichlorobenzyloxy)imidazo-[1,2-a]pyridine-2-carboxylate
mp : 207-208°C
NMR (CDCl₃, δ) : 1.41 (3H, t, J=7Hz), 4.47 (2H, q,
3-Chloro-8-(2,6-dichlorobenzyl oxy)-2,5-dimethylimidazo[1,2-a]pyridine
mp : 181-182°C
NMR (CDCl₃, δ) : 2.40 (3H, s), 2.89 (3H, s), 5.40 (2H, s), 6.47 (1H, d, J=7Hz), 6.53 (1H, d, J=7Hz), 7.21-7.44 (3H, m), 7.82 (1H, d, J=7Hz)

3-Bromo-8-[1-(2,6-dichlorophenyl)ethoxy]-2-methylimidazo[1,2-a]pyridine
mp : 135-136°C
NMR (CDCl₃, δ) : 1.93 (3H, d, J=7Hz), 2.49 (3H, s), 6.11-6.27 (2H), 6.40 (1H, t, J=7Hz), 7.13 (1H), 7.22-7.32 (2H), 7.61 (1H, d, J=7Hz)

3-Chloro-8-(2,6-dichlorobenzylamino)-2,7-dimethylimidazo[1,2-a]pyridine
mp : 144-145°C
NMR (CDCl₃, δ) : 2.38 (3H, s), 2.40 (3H, s), 4.44 (1H, br s), 5.00 (2H, d, J=5Hz), 6.59 (1H, d, J=7Hz), 7.09-7.42 (3H, m), 7.51 (1H, d, J=7Hz)

3-Bromo-8-[N-(2,6-dichlorobenzyl)-N-acetylamino]-2-methylimidazo[1,2-a]pyridine
mp : 141-142°C
NMR (CDCl₃, δ) : 1.90 (3H, s), 2.49 (3H, s), 4.90 (1H, br d, J=14Hz), 5.99 (1H, br d, J=14Hz), 6.45 (1H, d, J=7Hz), 6.61 (1H, t, J=7Hz), 7.01-7.21 (3H, m), 7.98 (1H, d, J=7Hz)

3-Chloro-8-[2-(2,6-dichlorophenyl)ethoxy]-2-methylimidazo[1,2-a]pyridine
mp : 131-134°C
NMR (CDCl₃, δ) : 2.48 (3H, s), 3.66 (2H, m), 4.33 (2H, m), 6.58 (1H, d, J=7.5Hz), 6.78 (1H, s, J=7.5Hz), 7.15 (1H, dd, J=7Hz and 5Hz), 7.32 (2H, d, J=7.5Hz), 7.66 (1H, d, J=7.5Hz)

(10) 3-Chloro-8-(2,6-dichloro-3-nitrobenzyloxy)-2-methylimidazo[1,2-a]pyridine
mp : 203-205°C
NMR (CDCl₃, δ) : 2.43 (3H, s), 5.55 (2H, s), 6.70 (1H, d, J=7.5Hz), 6.85 (1H, s, J=7.5Hz), 7.54 (1H, d, J=9Hz), 7.74 (1H, d, J=7.5Hz), 7.82 (1H, d, J=9Hz)

(11) 3-Chloro-8-[2,6-dichloro-3-(N-methyl-N-acetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 212-213°C
NMR (CDCl₃, δ) : 1.87 (3H, s), 2.48 (3H, s), 3.20 (3H, s), 5.51 (2H, s), 6.74 (1H, br d, J=7Hz), 6.90 (1H, br t, J=7Hz), 7.29 (1H, d, J=9Hz), 7.45 (1H, d, J=9Hz), 7.76 (1H, d, J=7Hz)

(12) 3-Bromo-8-[2,6-dichloro-3-(N-methyl-N-trifluoroacetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 175-176°C
NMR (CDCl₃, δ) : 2.45 (3H, s), 3.33 (3H, s), 5.52 (2H, s), 6.71 (1H, d, J=7.5Hz), 6.85 (1H, t, J=7.5Hz), 7.33 (1H, d, J=8Hz), 7.46 (1H, d, J=8Hz), 7.79 (1H, dd, J=7.5Hz and 1.5Hz)

(13) 3-Bromo-8-[2,6-dichloro-3-(N-methoxycarbonyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 178-179°C
NMR (CDCl₃, δ) : 2.48 (3H, s), 3.22 (3H, s),
3.65 (3H, s), 5.49 (2H, s), 6.78 (1H, br d, J=7Hz), 6.90 (1H, br t, J=7Hz), 7.25 (1H, d, J=9Hz), 7.39 (1H, d, J=9Hz), 7.79 (1H, d, J=7Hz)

(14) 3-Bromo-8-[2,6-dichloro-3-(N-methyl-N-tert-butoxy carbonylamino)benzyloxy]-2-methylimidazo[1,2-a]-pyridine
mp : 122-123°C
Mass (M+1) : 516

(15) 3-Bromo-8-[2,6-dichloro-3-(N-methyl-N-mesylamino)-
benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 161-164°C
NMR (CDCl₃, δ) : 2.45 (3H, s), 3.07 (3H, s), 3.30 (3H, s), 5.48 (2H, m), 6.71 (1H, d, J=7.5Hz), 6.85 (1H, t, J=7.5Hz), 7.52 (1H, d, J=10Hz), 7.60 (1H, d, J=10Hz), 7.77 (1H, d, J=7.5Hz and 1.5Hz)

(16) 3-Bromo-8-[2,6-dichloro-3-(N-ethyl-N-methylamino)-
benzyloxy]-2-methylimidazo[1,2-a]pyridine
dihydrochloride
mp : 128-130°C
NMR (CDCl₃ + CD₃OD, δ) : 1.30 (3H, t, J=7Hz), 2.64 (3H, s), 3.39 (3H, s), 3.80 (2H, q, J=7Hz), 5.69 (2H, s), 7.49 (2H, d, J=4Hz), 7.65 (1H, d, J=9Hz), 8.01-8.21 (2H)

(17) 3-Bromo-8-[2,6-dichloro-3-(2-pyrrolidinon-1-yl)-
benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 219-220°C
NMR (CDCl₃, δ) : 2.16-2.35 (2H, m), 2.42 (3H, s), 2.58 (2H, t, J=8Hz), 3.78 (2H, t, J=7Hz), 5.47 (2H, s), 6.71 (1H, d, J=7Hz), 6.85 (1H, t, J=7Hz), 7.28, 7.42 (each 1H, d, J=9Hz), 7.76
(1H, d, J=7Hz)

(18) 3-Bromo-8-[2,6-dichloro-3-(1-pyrrolidinyl)benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 113-114°C
NMR (CDCl₃, δ) : 1.88-2.02 (4H, m), 2.42 (3H, s),
3.25-3.41 (4H, m), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.82 (1H, t, J=7Hz), 6.90, 7.20 (each 1H, d, J=9Hz), 7.72 (1H, d, J=7Hz)

(19) 3-Bromo-8-(2,6-dichloro-3-methoxybenzyloxy)-2-methylimidazo[1,2-a]pyridine
mp : 175-176°C
NMR (CDCl₃, δ) : 2.43 (3H, s), 3.91 (3H, s), 5.48 (2H, s), 6.70 (1H, d, J=7Hz), 6.82 (1H, t, J=7Hz), 6.92 (1H, d, J=9Hz), 7.31 (1H, d, J=9Hz), 7.72 (1H, d, J=7Hz)

(20) 3-Bromo-8-(2,6-dichloro-3-isopropoxybenzyloxy)-2-methylimidazo[1,2-a]pyridine
mp : 115-117°C
NMR (CDCl₃, δ) : 1.40 (6H, d, J=5Hz), 2.44 (3H, s), 4.53 (1H, m), 5.45 (2H, s), 6.70 (1H, d, J=7.5Hz), 6.81 (1H, t, J=7.5Hz), 6.93 (1H, d, J=10Hz), 7.27 (1H, d, J=10Hz), 7.73 (1H, dd, J=7.5Hz and 1.5Hz)

(21) 3-Bromo-8-(2,4,6-trichloro-3-nitrobenzyloxy)-2-methylimidazo[1,2-a]pyridine
mp : 167-168°C
NMR (CDCl₃, δ) : 2.45 (3H, s), 5.48 (2H, s), 6.68 (1H, d, J=8Hz), 6.83 (1H, t, J=8Hz), 7.60 (1H, s), 7.78 (1H, d, J=8Hz)

(22) 3-Bromo-8-[2-chloro-5-(N-methyl-N-acetylamino)-
benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp: 141-145°C
NMR (CDCl₃, δ): 1.78 (3H, s), 2.50 (3H, s), 3.20 (3H, s), 5.45 (2H, s), 6.50 (1H, d, J=7.5Hz), 6.77 (1H, t, J=7.5Hz), 7.13 (1H, dd, J=10Hz and 1.5Hz), 7.42-7.52 (2H, m), 7.75 (1H, d, J=10Hz)

(23) 3-Bromo-8-[3'(N-acetoxyacetyl-N-methylamino)-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine
mp: 229-230°C
NMR (CDCl₃, δ): 2.14 (3H, s), 2.43 (3H, s), 3.21 (2H, s), 4.18, 4.49 (each 1H, d, J=15Hz), 5.50 (2H, s), 6.70 (1H, d, J=7Hz), 6.83 (1H, t, J=7Hz), 7.39, 7.49 (each 1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(24) 3-Bromo-8-[2,6-dichloro-3-(N-methyl-N-phthalimido-acetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp: 229-230°C
NMR (CDCl₃, δ): 2.43 (3H, s), 3.26 (3H, s), 4.12 (2H, s), 5.53 (2H, s), 6.72 (1H, d, J=7Hz), 6.87 (1H, d, J=7Hz), 7.52 (2H, s), 7.68-7.92 (5H)

(25) 8-[3-(Acetylsarcosyl)amino-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine
mp: 164-165°C
NMR (CDCl₃, δ): 2.20 (3H, s), 2.42 (3H, s), 3.19 (3H, s), 4.19 (2H, s), 5.48 (2H, s), 6.70 (1H, d, J=8Hz), 6.83 (1H, t, J=8Hz), 7.35 (1H, d, J=9Hz), 7.75 (1H, d, J=8Hz), 8.37 (1H, d, J=9Hz), 8.82 (1H, br s)

(26) 3-Chloro-8-(2-trifluromethylbenzyloxy)-2-methylimidazo[1,2-a]pyridine
mp: 164-165°C
NMR (CDCl₃, δ): 2.48 (3H, s), 5.40 (2H, s), 6.44 (1H, d, J=7Hz), 6.71 (1H, t, J=7Hz), 7.55-7.71 (5H)
(27) 3-Chloro-8-{2-methoxycarbonylbenzyloxy}-2-methylimidazo[1,2-a]pyridine
mp : 141-144°C
NMR (CDCl₃, δ) : 2.48 (3H, s), 3.92 (3H, s), 5.79 (2H, s), 6.49 (1H, d, J=7.5Hz), 6.72 (1H, t, J=7.5Hz), 7.39 (1H, t, J=7.5Hz), 7.67 (1H, d, J=7.5Hz), 7.85 (1H, d, J=7.5Hz)

(28) 8-(2-Phenylbenzyloxy)-3-chloro-2-methylimidazo-15····[1,2-a]pyridine
mp : 121-122°C
NMR (CDCl₃, δ) : 2.45 (3H, s), 5.19 (2H, s), 6.21 (1H, d, J=7.5Hz), 6.63 (1H, t, J=7.5Hz), 7.30-7.41 (8H, m), 7.71 (1H, d, J=7.5Hz), 7.72 (1H, m)

(29) 3-Chloro-8-(2,6-difluorobenzyloxy)-2-methylimidazo-1,2-a]pyridine
mp : 168-170°C
NMR (CDCl₃, δ) : 2.45 (3H, s), 5.33 (2H, s), 6.68 (1H, d, J=7.5Hz), 6.80 (1H, t, J=7.5Hz), 6.87-7.00 (2H, m), 7.28-7.41 (1H, m), 7.70 (1H, d, J=7.5Hz)

(30) 3-Chloro-8-(2,6-dibromobenzyloxy)-2-methylimidazo-1,2-a]pyridine
mp : 157-158°C
NMR (CDCl₃, δ) : 2.44 (3H, s), 5.51 (2H, s), 6.69 (1H, d, J=7Hz), 6.82 (1H, t, J=7Hz), 7.08 (1H, t, J=8Hz), 7.57 (2H, d, J=8Hz), 7.70 (1H, d, J=7Hz)

(31) 3-Bromo-8-(2-chloro-6-fluorobenzyloxy)-2-methylimidazo[1,2-a]pyridine
(32) 8-(4-Bromo-2-fluorobenzyl)oxy)-3-chloro-2-methylimidazo[1,2-a]pyridine

mp : 182-123°C
NMR (CDCl₃, δ) : 2.47 (3H, s), 5.32 (2H, s), 6.51 (1H, d, J=7.5Hz), 6.75 (1H, t, J=7.5Hz), 7.29 (2H, d, J=7.5Hz), 7.48 (1H, t, J=7.5Hz), 7.66 (1H, d, J=7.5Hz)

(33) 3-Bromo-8-(2-chloro-5-nitrobenzyloxy)-2-methylimidazo[1,2-a]pyridine

mp : 155-156°C
NMR (CDCl₃, δ) : 2.50 (3H, s), 5.48 (2H, s), 6.53 (1H, d, J=7Hz), 6.78 (1H, t, J=7Hz), 7.60 (1H, d, J=7Hz), 7.78 (1H, d, J=7Hz), 8.15 (1H, dd, J=9Hz and 2Hz), 8.52 (1H, d, J=2Hz)

(34) 3-Bromo-8-[2-chloro-6-(N-methyl-N-acetylamino)-benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 135-136°C
NMR (CDCl₃, δ) : 1.89 (3H, s), 2.41 (3H, s), 3.23 (3H, s), 5.21 (1H, d, J=10Hz), 5.26 (1H, d, J=10Hz), 6.68 (1H, d, J=7.5Hz), 6.82 (1H, t, J=7.5Hz), 7.18 (1H, d, J=7.5Hz), 7.38-7.53 (2H, m), 7.74 (1H, d, J=7.5Hz)

(35) 3-Bromo-8-(2-chloro-6-nitrobenzyloxy)-2-methylimidazo[1,2-a]pyridine

mp : 157-158°C
NMR (CDCl₃, δ) : 2.42 (3H, s), 5.66 (2H, s), 6.67 (1H, d, J=7Hz), 6.82 (1H, t, J=7Hz), 7.49 (1H,
t, J=8Hz), 7.72 (2H, t, J=8Hz), 7.85 (1H, d, J=7Hz)

(36) 3-Bromo-8-[3-(N,N-di-tert-butoxycarbonylamino)-2,6-dichlorobenzoyloxy]-2-methylimidazo[1,2-a]pyridine
mp: 154-155°C
NMR (CDCl₃, δ): 1.40 (18H, s), 2.43 (3H, s), 5.58 (2H, s), 6.68 (1H, d, J=7Hz), 6.79 (1H, t, J=7Hz), 7.20 (1H, d, J=8Hz), 7.38 (1H, d, J=8Hz), 7.72 (1H, d, J=7Hz)

(37) 3-Bromo-8-(2,4,6-trichlorobenzoyloxy)-2-methylimidazo[1,2-a]pyridine
mp: 170-172°C
NMR (CDCl₃, δ): 2.44 (3H, s), 5.42 (2H, s), 6.68 (1H, d, J=7Hz), 6.81 (1H, t, J=7Hz), 7.49 (2H, s), 7.76 (1H, d, J=7Hz)

(38) 3-Bromo-8-(2,3,6-trichlorobenzoyloxy)-2-methylimidazo[1,2-a]pyridine
mp: 183-184°C
NMR (CDCl₃, δ): 2.44 (3H, s), 5.50 (2H, s), 6.70 (1H, d, J=7Hz), 6.83 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.46 (1H, d, J=9Hz), 7.76 (1H, d, J=7Hz)

(39) 3-Chloro-8-(2,4,6-trimethylbenzoyloxy)-2-methylimidazo[1,2-a]pyridine
mp: 154-155°C
NMR (CDCl₃, δ): 2.29 (3H, s), 2.38 (6H, s), 2.43 (3H, s), 5.21 (2H, s), 6.63 (1H, d, J=7Hz), 6.75-6.90 (3H), 7.69 (1H, d, J=7Hz)

(40) 3-Bromo-8-(2,6-dibromo-4-methoxycarbonylbenzoyloxy)-2-methylimidazo[1,2-a]pyridine
mp: 189-190°C
NMR (CDCl₃, δ): 2.44 (3H, s), 3.96 (3H, s), 5.52 (2H, s), 6.69 (1H, d, J=7Hz), 6.82 (1H, t, J=7Hz), 7.76 (1H, d, J=7Hz), 8.21 (2H, s)

(41) 3-Bromo-8-(4-benzoyl-2-chlorobenzyloxy)-2-methylimidazolo[1,2-a]pyridine
NMR (CDCl₃, δ): 2.50 (3H, s), 5.50 (2H, s), 6.46 (1H, d, J=8Hz), 6.78 (1H, t, J=8Hz), 7.34 (1H, d, J=9Hz), 7.42-7.91 (9H)

(42) 3-Bromo-8-(2,6-dichloro-4-benzoylbenzyloxy)-2-methylimidazolo[1,2-a]pyridine
mp: 129-130°C
NMR (CDCl₃, δ): 2.46 (3H, s), 5.50 (2H, s), 6.71 (1H, d, J=8Hz), 6.83 (1H, t, J=8Hz), 7.34 (1H, d, J=9Hz), 7.42-7.91 (7H)

(43) 3-Bromo-8-[4-(2-cyanophenyl)benzyloxy]-2-methylimidazolo[1,2-a]pyridine
mp: 138-140°C
NMR (CDCl₃, δ): 2.49 (3H, s), 5.40 (2H, s), 6.55 (1H, d, J=8Hz), 6.75 (1H, t, J=8Hz), 7.40-7.81 (9H)

(44) 3-Bromo-8-(6-chloropiperonylmethoxy)-2-methylimidazolo[1,2-a]pyridine
mp: 185-186°C
NMR (CDCl₃, δ): 2.49 (3H, s), 5.32 (2H, s), 5.96 (2H, s), 6.49 (1H, d, J=7Hz), 6.74 (1H, t, J=7Hz), 6.88 (1H, s), 7.09 (1H, s), 7.70 (1H, d, J=7Hz)

(45) 8-(2-Bromothiophen-4-yl)methoxy-3-chloro-2-methylimidazolo[1,2-a]pyridine
mp: 124-125°C
NMR (CDCl$_3$, δ): 2.48 (3H, s), 5.45 (2H, s), 6.56 (1H, d, J=7Hz), 6.75 (1H, t, J=7Hz), 7.09 (1H, s), 7.20 (1H, s), 7.69 (2H, d, J=7Hz)

(46) 3-Chloro-8-(3-chlorobenzofuran-2-yl)methoxy-2-methylimidazo[1,2-a]pyridine
mp: 132-134°C
NMR (CDCl$_3$, δ): 2.46 (3H, s), 5.48 (2H, s), 6.71-6.82 (2H, m), 7.31-7.71 (5H, m)

(47) 3-Bromo-8-[3-(4-chlorophenyl)-5-methylbenzofuran-2-yl)methoxy-2-methylimidazo[1,2-a]pyridine
mp: 167-170°C
NMR (CDCl$_3$, δ): 2.44 (6H, s), 5.38 (2H, s), 6.59 (1H, d, J=7Hz), 6.72 (1H, t, J=7Hz), 7.18 (1H, br d, J=8Hz), 7.32-7.52 (6H), 7.71 (1H, d, J=7Hz)

(48) 3-Bromo-8-(4,6-dichloro-3-methyl-2-benzothiazolinon-5-yl)methoxy-2-methylimidazo[1,2-a]pyridine
mp: 225-227°C
NMR (CDCl$_3$, δ): 2.45 (3H, s), 3.87 (3H, s), 5.51 (2H, s), 6.71 (1H, d, J=7Hz), 6.83 (1H, t, J=7Hz), 7.42 (1H, s), 7.78 (1H, d, J=7Hz)

(49) 3-Bromo-8-(4,6-dichloro-3-ethyl-2-benzothiazolinon-5-yl)methoxy-2-methylimidazo[1,2-a]pyridine
mp: 219-220°C
NMR (DMSO-d$_6$, δ): 1.35 (3H, t, J=7.5Hz), 2.35 (3H, s), 4.43 (2H, q, J=7.5Hz), 5.52 (2H, s), 6.93-7.02 (2H, m), 7.83-7.89 (2H, m)

(50) 3-Bromo-8-(5,7-dichloro-4-methyl-1,4-benzothiazin-3(4H)-on-6-yl)methoxy-2-methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, δ): 2.44 (3H, s), 3.39 (2H, s), 5.45 (2H, s), 6.75 (1H, t, J=7Hz), 7.09 (1H, s), 7.20 (1H, s), 7.69 (2H, d, J=7Hz)
3.41 (3H, s), 5.47 (2H, s), 6.70 (1H, d, J=8Hz),
6.84 (1H, t, J=8Hz), 7.48 (1H, s), 7.77 (1H, d, J=8Hz)

Example 4

A suspension of 3-bromo-8-(2,6-dichloro-3-
nitrobenzyl)oxy)-2-methylimidazo[1,2-a]pyridine (215 mg)
and iron (powder, 84 mg) in a mixture of conc.
hydrochloric acid (1 ml) and methanol (1 ml) was refluxed
for half an hour. The cooled mixture was poured into an
ice water (15 ml). The precipitates were collected and
washed with water to give 8-(3-amino-2,6-
dichlorobenzyl)oxy)-3-bromo-2-methylimidazo[1,2-a]pyridine
dihydrochloride (140 mg) as an off-white solid.

mp : 181-183°C

NMR (DMSO-\(d_6\), \(\delta\)) : 2.42 (3H, s), 5.48 (2H, s), 6.95
(1H, d, J=7.5Hz), 7.24 (1H, d, J=7.5Hz), 7.42
(1H, t, J=7.5Hz), 7.62 (1H, d, J=7.5Hz), 8.25
(1H, d, J=7.5Hz)

Example 5

To a mixture of 3-bromo-8-(2-chloro-6-
nitrobenzyl)oxy)-2-methylimidazo[1,2-a]pyridine (300 mg),
nickel(II) chloride hexahydrate (360 mg) and methanol (9
ml) was added sodium borohydride (114 mg) portionwise
under an ice-water bath cooling for 20 minutes. After
stirring for 30 minutes, the mixture was concentrated in
vacuo and diluted with water. The pH value of the mixture
was adjusted to 3 and the separated crystals were
collected by filtration. The crystals were dissolved in
dichloromethane, and the solution was washed with
saturated aqueous solution of sodium hydrogencarbonate,
dried over magnesium sulfate and evaporated in vacuo. The
residue was purified by silica gel column chromatography
(3% solution of methanol in dichloromethane) and
preparative thin-layer chromatography (3% solution of methanol in dichloromethane) followed by crystallization from diethyl ether to give 8-(2-amino-6-chlorobenzyloxy)-3-bromo-2-methylimidazo[1,2-a]pyridine (40 mg).

mp: 177-178°C
NMR (CDCl₃, δ): 2.44 (3H, s), 4.64 (2H, br s), 5.51 (2H, s), 6.57 (1H, d, J=7Hz), 6.71-6.85 (3H), 7.02 (1H, t, J=7Hz), 7.71 (1H, m)

Example 6

The following compounds were obtained according to similar manners to those of Examples 4 or 5.

1. 8-(3-Amino-2,4,6-trichlorobenzyloxy)-3-bromo-2-methylimidazo[1,2-a]pyridine
   mp: 217-219°C
   NMR (DMSO-d₆, δ): 2.30 (3H, s), 5.35 (2H, s), 6.99 (2H, d, J=8Hz), 7.58 (1H, s), 7.92 (1H, t, J=8Hz)

2. 8-(3-Amino-2,6-dichlorobenzyloxy)-3-bromo-2-methylimidazo[1,2-a]pyridine
   NMR (DMSO-d₆, δ): 2.29 (3H, s), 5.31 (2H, s), 5.70 (2H, s, D₂O exchangeable), 6.84-7.04 (3H), 7.23 (1H, d, J=8Hz), 7.90 (1H, d, J=5Hz)

3. 8-(5-Amino-2-chlorobenzyloxy)-3-bromo-2-methylimidazo[1,2-a]pyridine dihydrochloride
   mp: >250°C (dec.)
   NMR (DMSO-d₆, δ): 2.45 (3H, s), 4.21 (2H, br s), 5.41 (2H, s), 7.03 (1H, d, J=9Hz), 7.29-7.52 (4H), 8.22 (1H, d, J=6Hz)

Example 7

To a suspension of 3-bromo-8-[2,6-dichloro-3-(N-
methyl-N-trifluoroacetylamino)benzyloxy]-2-methylimidazo-[1,2-a]pyridine (560 mg) in methanol (6 ml) was added 28% sodium methoxide in methanol (1.93 g). The mixture was refluxed for one hour and cooled. The precipitated solid was filtered, washed with methanol, and dried to give 3-bromo-8-(2,6-dichloro-3-methylaminobenzyloxy)-2-methylimidazo[1,2-a]pyridine (350 mg) as crystals.

mp : 184-187°C
NMR (CDCl₃, δ) : 2.44 (3H, s), 2.91 (2H, d, J=6Hz), 4.46 (1H, m), 5.46 (2H, s), 6.69 (1H, d, J=8.5Hz), 6.71 (1H, d, J=7.5Hz), 6.83 (1H, t, J=7.5Hz), 7.24 (1H, d, J=8.5Hz), 7.73 (1H, d, J=7.5Hz)

Example 8

The following compounds were obtained according to a similar manner to that of Example 7.

(1) 3-Bromo-8-(2-chloro-5-methylaminobenzyloxy)-2-methylimidazo[1,2-a]pyridine
mp : 156-159°C
NMR (CDCl₃, δ) : 2.48 (3H, s), 2.76 (3H, s), 5.38 (2H, s), 6.43-6.52 (2H, m), 6.73 (1H, t, J=7.5Hz), 6.85 (1H, d, J=2Hz), 7.19 (1H, d, J=9Hz), 7.69 (1H, d, J=7.5Hz)

(2) 3-Chloro-8-(2,6-dichloro-3-methylaminobenzyloxy)-2-methylimidazo[1,2-a]pyridine
mp : 187-188°C
NMR (CDCl₃, δ) : 2.43 (3H, s), 2.90 (3H, d, J=6Hz), 4.46 (1H, br q, J=6Hz), 5.45 (2H, s), 6.62 (1H, d, J=9Hz), 6.68 (1H, d, J=8Hz), 6.82 (1H, t, J=8Hz), 7.24 (1H, d, J=9Hz), 7.69 (1H, d, J=8Hz)
Example 9
A mixture of 8-(3-amino-2,6-dichlorobenzyloxy)-3-bromo-2-methylimidazo[1,2-a]pyridine dihydrochloride (100 mg) and formaldehyde (200 mg, 37% wt % solution in water) in 90% aqueous formic acid (1 ml) was refluxed for 40 minutes. The cooled mixture was concentrated in vacuo. The residue was partitioned between ethyl acetate and aqueous sodium bicarbonate solution. The organic layer was dried and concentrated in vacuo to give an oil, which was purified by preparation thin-layer chromatography on silica gel (ethyl acetate:n-hexane = 2:1, V/V) to give 3-bromo-8-(2,6-dichloro-3-dimethylaminobenzyloxy)-2-methylimidazo[1,2-a]pyridine as a white solid (31 mg).
mp: 120-122°C
NMR (DMSO-d$_6$, δ): 2.29 (3H, s), 2.76 (6H, s), 5.41 (2H, s), 6.97-7.03 (2H, m), 7.33 (1H, d, J=9Hz), 7.52 (1H, d, J=9Hz), 7.90 (1H, m)

Example 10
To a mixture of 8-(3-amino-2,6-dichlorobenzyloxy)-3-bromo-2-methylimidazo[1,2-a]pyridine dihydrochloride (100 mg) and acetone (116 mg) in 3 M HCl solution in ethanol (2 ml) was added sodium cyanoborohydride (25 mg) in one portion. The mixture was stirred for 2 hours at ambient temperature and then concentrated in vacuo. The residue was partitioned between ethyl acetate and aqueous sodium bicarbonate solution. The organic layer was dried and concentrated in vacuo to give an oil, which was purified by preparative thin-layer chromatography on silica gel (ethyl acetate:n-hexane = 1:2, V/V), to give 3-bromo-8-(2,6-dichloro-3-isopropylaminobenzyloxy)-2-methylimidazo[1,2-a]pyridine as a white solid (50 mg).
mp: 133-134°C
NMR (CDCl$_3$, δ): 1.26 (6H, d, J=5Hz), 2.45 (3H, s), 3.66 (1H, m), 4.22 (1H, d, J=7Hz), 5.42 (2H, s),
6.63 (1H, d, J=7.5Hz), 6.70 (1H, d, J=7.5Hz),
6.83 (1H, t, J=7.5Hz), 7.19 (1H, d, J=7.5Hz),
7.73 (1H, d, J=7.5Hz)

Example 11
The following compounds were obtained according to a similar manner to that of Example 10.

(1) 8-(3-Benzylamino-2,6-dichlorobenzyloxy)-3-bromo-2-methylimidazo[1,2-a]pyridine
mp : 126-127°C
NMR (CDCl₃, δ) : 2.45 (3H, s), 4.43 (2H, d, J=5Hz),
4.89 (1H, t, J=5Hz), 5.46 (2H, s), 6.58 (1H, d,
J=7.5Hz), 6.71 (1H, d, J=7.5Hz), 6.83 (1H, t,
J=7.5Hz), 7.15 (1H, d, J=7.5Hz), 7.29-7.43 (5H,
m), 7.72 (1H, d, J=7.5Hz)

(2) 3-Bromo-8-[2,6-dichloro-3-(4-pyridylmethyl)amino-benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 214-215°C
NMR (CDCl₃, δ) : 2.44 (3H, s), 4.47 (2H, d, J=6Hz),
5.05 (1H, t, J=5Hz), 5.46 (2H, s), 6.41 (1H, d,
J=7.5Hz), 6.71 (1H, d, J=7.5Hz), 6.83 (1H, t,
J=7.5Hz), 7.12 (1H, d, J=7.5Hz), 7.25-7.29 (2H,
m), 7.74 (1H, d, J=7.5Hz), 8.59 (2H, m)

Example 12
To a suspension of 8-(3-amino-2,6-dichlorobenzyloxy)-
3-bromo-2-methylimidazo[1,2-a]pyridine dihydrochloride
(100 mg) in a mixture of pyridine (0.5 ml) and
N,N-diethylformamide (1.5 ml) was added methanesulfonyl
chloride (27 mg) in one portion. The mixture was stirred
at 60-70°C for one and half hours, cooled, and poured into
ice water. The separated oil was extracted with
dichloromethane. The extract was washed with water,
dried, and concentrated in vacuo to give a brown oil, which was purified by preparative thin-layer chromatography on silica gel (5% solution of methanol in dichloromethane) to give 3-bromo-8-[2,6-dichloro-3-[(N,N-diethylaminomethylene)amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (65 mg) as crystals.

mp : 163-165°C
NMR (CDCl₃, δ) : 1.25 (3H, t, J=7Hz), 2.44 (2H, s), 3.18-3.61 (4H, m), 5.49 (2H, s), 6.68-6.85 (3H, m), 7.23 (1H, d, J=7.5Hz), 7.38 (1H, s), 7.71 (1H, d, J=7.5Hz)

Example 13

The following compounds were obtained according to a similar manner to that of Example 12.

(1) 3-Bromo-8-[2,6-dichloro-3-[(N,N-dimethylamino)methylene]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 165-167°C
NMR (CDCl₃, δ) : 1.25 (3H, t, J=7Hz), 2.44 (2H, s), 3.18-3.61 (4H, m), 5.49 (2H, s), 6.68-6.85 (3H, m), 7.23 (1H, d, J=7.5Hz), 7.38 (1H, s), 7.71 (1H, d, J=7.5Hz)

(2) 3-Bromo-8-[2,6-dichloro-3-(1-methyl-2-pyrrolidinylideneamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 178-179°C
NMR (CDCl₃, δ) : 1.97 (2H, m), 2.32 (2H, t, J=7.5Hz), 2.45 (3H, s), 2.98 (3H, s), 3.40 (2H, t, J=7.5Hz), 5.47 (2H, s), 6.69-6.82 (2H, m), 6.83 (1H, d, J=7.5Hz), 7.19 (1H, d, J=7.5Hz), 7.72 (1H, d, J=7.5Hz)
Example 14

A mixture of 8-(3-amino-2,6-dichlorobenzyloxy)-3-
bromo-2-methylimidazo[1,2-a]pyridine dihydrochloride (100 mg) and acetic anhydride (100 mg) in dry pyridine (2 ml) was stirred at 90°C for one hour. The cooled mixture was poured into an ice water (10 ml). The precipitated solid was collected, washed with water, and dried under reduced pressure to give 8-(3-acetylamino-2,6-dichlorobenzyloxy)-3-
bromo-2-methylimidazo[1,2-a]pyridine (65 mg) as a solid.

mp : 210-212°C

NMR (DMSO-d$_6$, δ) : 2.13 (3H, s), 2.29 (3H, s), 5.44 (2H, s), 6.99-7.04 (2H, m), 7.59 (1H, d, J=7.5Hz), 7.84 (1H, d, J=7.5Hz), 7.92 (1H, m), 9.72 (1H, s)

Example 15

To a suspension of 8-(3-amino-2,6-dichlorobenzyloxy)-3-
bromo-2-methylimidazo[1,2-a]pyridine dihydrochloride (200 mg), pyridine (1 ml) and N-methylpyrrolidone (3 ml) was added propionyl chloride (78 mg) and the mixture was stirred at 60°C for 1 hour. To the cooled mixture was added water, and the precipitate was collected by filtration and dissolved in chloroform. This organic solution was washed with water and brine respectively, dried with magnesium sulfate and concentrated in vacuo. The residue was collected by filtration and washed with ethyl acetate to give 3-bromo-8-(2,6-dichloro-3-
propionylaminobenzyloxy)-2-methylimidazo[1,2-a]pyridine (158 mg).

mp : 205-207°C

NMR (CDCl$_3$, δ) : 1.28 (3H, t, J=7Hz), 2.44 (3H, s), 2.48 (2H, q, J=7Hz), 5.49 (2H, s), 6.70 (1H, d, J=7Hz), 6.82 (1H, t, J=7Hz), 7.37 (1H, d, J=9Hz), 7.69 (1H, br s), 7.76 (1H, d, J=7Hz), 8.42 (1H, d, J=9Hz)
Example 16

The following compounds were obtained according to similar manners to those of Examples 14 or 15.

(1) 3-Bromo-8-[2,6-dichloro-3-(4-methylbenzoylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp: 222-223°C
NMR (CDCl₃, δ): 2.43 (6H, s), 5.51 (2H, s), 6.71 (1H, d, J=7Hz), 6.82 (1H, t, J=7Hz), 7.31 (2H, d, J=9Hz), 7.41 (1H, d, J=9Hz), 7.70-7.85 (3H), 8.44 (1H, br s), 8.60 (1H, d, J=9Hz)

(2) 3-Bromo-8-[2,6-dichloro-3-(4-methoxybenzoylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp: 211-212°C
NMR (CDCl₃, δ): 2.44 (3H, s), 3.89 (3H, s), 5.51 (2H, s), 6.70 (1H, d, J=7Hz), 6.81 (1H, t, J=7Hz), 7.00 (2H, d, J=9Hz), 7.41 (1H, d, J=9Hz), 7.75 (1H, d, J=7Hz), 7.88 (2H, d, J=9Hz), 8.40 (1H, br s), 8.59 (1H, d, J=9Hz)

(3) 8-(3-Benzoylamino-2,6-dichlorobenzyloxy)-3-bromo-2-methylimidazo[1,2-a]pyridine

mp: 208-210°C
NMR (CDCl₃, δ): 2.44 (3H, s), 5.52 (2H, s), 6.71 (1H, d, J=7Hz), 6.83 (1H, t, J=7Hz), 7.40-7.66 (4H), 7.75 (1H, d, J=7Hz), 7.87-7.96 (2H), 8.49 (1H, br s), 8.60 (1H, d, J=9Hz)

(4) 3-Bromo-8-[2,6-dichloro-3-(4-methoxycarbonylbenzoylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp: 238-239°C
NMR (DMSO-δ₆, δ): 2.31 (3H, s), 3.92 (3H, s), 5.49 (2H, s), 6.97-7.08 (2H, m), 7.69 (1H, d, J=8.5Hz), 7.77 (1H, d, J=8.5Hz), 7.95 (1H, dd,
(5) 3-Bromo-8-[3-(4-chlorobenzoylamino)-2,6-dichlorobenzylxoy]-2-methylimidazo[1,2-a]pyridine

mp : 201-213°C

NMR (CDCl₃, δ) : 2.34 (6H, s), 2.45 (3H, s), 5.48 (2H, s), 6.69 (1H, d, J=8Hz), 6.84 (1H, t, J=7.5Hz), 7.43 (1H, d, J=8.5Hz), 7.52 (2H, d, J=8.5Hz), 7.76 (1H, d, J=8.5Hz), 8.41 (1H, br s), 8.57 (1H, J=0.5Hz)

(6) 8-(3-Diacetylamino-2,4,6-trichlorobenzylxoy)-3-bromo-2-methylimidazo[1,2-a]pyridine

mp : 141-142°C

NMR (CDCl₃, δ) : 1.97 (2H, q, J=7.5Hz), 2.50 (3H, s), 3.19 (3H, s), 5.45 (2H, s), 6.50 (1H, d, J=7.5Hz), 6.76 (1H, t, J=7.5Hz), 7.11 (1H, dd, J=9Hz and 2Hz), 7.45 (1H, s), 7.48 (1H, d, J=9Hz), 7.75 (1H, d, J=7.5Hz)

(7) 3-Bromo-8-[2-chloro-5-(N-methyl-N-trifluoroacetylaminobenzylxoy]-2-methylimidazo[1,2-a]pyridine

mp : 142-143°C

NMR (CDCl₃, δ) : 2.50 (3H, s), 3.31 (3H, s), 5.48 (2H, s), 6.46 (1H, d, J=7.5Hz), 6.75 (1H, d, J=7.5Hz), 7.18 (1H, d, J=7.5Hz), 7.42-7.53 (1H, m), 7.76 (1H, d, J=7.5Hz)

(8) 3-Bromo-8-[2-chloro-5-(N-methyl-N-propionylaminobenzylxoy]-2-methylimidazo[1,2-a]pyridine

mp : 133-135°C

NMR (CDCl₃, δ) : 0.96 (3H, t, J=7.5Hz), 1.97 (2H, q, J=7.5Hz), 2.50 (3H, s), 3.19 (3H, s), 5.45 (2H, s), 6.50 (1H, d, J=7.5Hz), 6.76 (1H, t, J=7.5Hz), 7.11 (1H, dd, J=9Hz and 2Hz), 7.45 (1H, s), 7.48 (1H, d, J=9Hz), 7.75 (1H, d, J=7.5Hz)
(9) 3-Bromo-8-[2,6-dichloro-3-(phthalimidoacetylamino)-benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 228-230°C
NMR (CDCl₃, δ) : 2.42 (1.2H, s), 2.46 (1.8H, s), 4.60 (1.8H, s), 4.71 (1.2H, s), 5.47 (1.8Hz), 5.54 (1.2H, s), 6.68 (1H, d, J=8Hz), 6.81 (1H, t, J=8Hz), 7.32 (0.6H, d, J=8Hz), 7.56 (0.4H, d, J=5Hz), 7.70-7.98 (5.6H), 8.28-8.40 (1.4H)

(10) 3-Bromo-8-[2,6-dichloro-3-(2-phthalimidopropionyl-amino)benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (DMSO-d₆, δ) : 1.62 (3H, d, J=6Hz), 2.28 (3H, s), 5.01 (1H, q, J=6Hz), 5.43 (2H, s), 6.93-7.04 (2H), 7.60 (2H, s), 7.82-7.97 (5H), 9.92 (1H, s)

(11) 3-Bromo-8-[2,6-dichloro-3-(3-phthalimidopropionyl-amino)benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 207-209°C
NMR (CDCl₃, δ) : 2.41 (3H, s), 2.90 (2H, t, J=6Hz), 4.12 (2H, t, J=6Hz), 5.45 (2H, s), 6.69 (1H, d, J=7Hz), 6.82 (1H, t, J=7Hz), 7.35 (1H, d, J=9Hz), 7.61-7.94 (6H), 8.31 (1H, d, J=7Hz)

(12) 3-Chloro-8-[2,6-dichloro-3-[N-methyl-N-(phthalimido-acetyl)amino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine
mp : 139-141°C
NMR (CDCl₃, δ): 2.42 (3H, s), 3.25 (3H, s), 4.12 (2H, s), 5.53 (2H, s), 6.71 (1H, d, J=7Hz), 6.86 (1H, d, J=7Hz), 7.52 (2H, s), 7.68-7.92 (5H)

Example 17
To a solution of 3-bromo-8-(2,6-dichloro-3-methylaminobenzyloxy-2-methylimidazo[1,2-a]pyridine (70 mg) in formic acid (1 ml) was added acetic anhydride (35 mg) at
ambient temperature. The mixture was stirred for half an hour at the same temperature and then concentrated in vacuo. The residue was partitioned between aqueous sodium bicarbonate solution and ethyl acetate. The organic layer was dried and evaporated under reduced pressure to give 3-bromo-8-[2,6-dichloro-3-(N-methyl-N-formylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine (29 mg).

mp : 207-209°C
NMR (CDCl₃, δ) : 2.45 (3H, s), 3.23 (3H, s), 5.50 (2H, s), 6.71 (1H, d, J=7.5Hz), 6.85 (1H, t, J=7.5Hz), 7.26 (1H, d, J=10Hz), 7.45 (1H, d, J=10Hz), 7.76 (1H, d, J=7.5Hz), 8.15 (1H, s)

Example 18
To a solution of 3-bromo-8-(2,6-dichloro-3-methylaminobenzyloxy)-2-methylimidazo[1,2-a]pyridine (68 mg) and pyridine (0.5 ml) in dichloromethane (2 ml) was added 4-nitrophenyl chloroformate (40 mg) at ambient temperature. After stirring for 1 hour, the mixture was partitioned between dichloromethane and water. The aqueous layer was extracted with dichloromethane twice. The organic layers were combined, washed with water twice and brine, dried over magnesium sulfate and evaporated in vacuo. The residue was crystallized from diethyl ether to give 3-bromo-8-[2,6-dichloro-3-[N-methyl-N-(4-nitrophenoxycarbonyl)amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (84 mg) as crystals.

mp : 229-230°C
NMR (CDCl₃-CD₃OD, δ) : 2.41 (3H, s), 3.32 (3H, s), 5.49 (1H, d, J=10Hz), 5.57 (1H, d, J=10Hz), 6.73 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.19-7.57 (4H), 7.78 (1H, d, J=7Hz), 8.20 (2H, d, J=10Hz)

Example 19
To a solution of 3-bromo-8-(2-chloro-5-methylamino-
benzyloxy)-2-methylimidazo[1,2-a]pyridine (60 mg), pyridine (16 mg) and 4-dimethylaminopyridine (10 mg) in methylene chloride (2 ml) was added mesyl chloride (22 mg) in one portion at 5°C, and the mixture was stirred for 9 hours at ambient temperature. The precipitate was filtered off and the residue was washed with methylene chloride and diethyl ether respectively. The filtrate was concentrated in vacuo and the residue was purified by preparative thin-layer chromatography on silica gel (methanol:methylene chloride = 1:10, V/V) to give 3-bromo-8-[2-chloro-5-(N-methyl-N-methylsulfonylamino)-benzyloxy]-2-methylimidazo[1,2-a]pyridine (25 mg).

\[ \text{mp : 158-162°C} \]

\[ \text{NMR (CDCl}_3, \delta) : 2.49 (3H, s), 2.74 (3H, s), 3.26 (3H, s), 5.44 (2H, s), 6.53 (1H, d, J=7.5Hz), 6.78 (1H, t, J=7.5Hz), 7.32-7.46 (2H, m), 7.60 (1H, d, J=2Hz), 7.73 (1H, d, J=7.5Hz) \]

**Example 20**

To a mixture of 8-(3-amino-2,6-dichlorobenzyloxy)-3-bromo-2-methylimidazo[1,2-a]pyridine dihydrochloride (100 mg), pyridine (0.5 ml) and N-methylpyrrolidone (1.5 ml) was added ethyl isocyanate (0.10 ml). The mixture was stirred at 60°C for 6 hours. The insoluble material was filtered off and washed with water. The filtrate and washings were combined and separated precipitate was collected by filtration. The precipitate was purified by preparative thin-layer chromatography (20% solution of methanol in dichloromethane) followed by recrystallization from n-hexane to give 3-bromo-8-[2,6-dichloro-3-(N'-ethylureido)benzyloxy]-2-methylimidazo[1,2-a]pyridine (13 mg) as crystals.

\[ \text{mp : 238-239°C} \]

\[ \text{NMR (CDCl}_3, \delta) : 1.00 (3H, t, J=7Hz), 2.38 (3H, s), 3.16-3.31 (2H), 5.32 (2H, s), 6.67 (1H, d, } \]
Example 21

A mixture of 3-bromo-8-[2,6-dichloro-3-[N-methyl-N-(4-nitrophenyloxycarbonyl)amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (63 mg) and 30% solution of methylamine in methanol (2 ml) was heated under reflux for 3 hours. After addition of 30% solution of methylamine in methanol (1 ml), the mixture was heated under reflux for additional 1 hour. The mixture was evaporated in vacuo and the residue was extracted with ethyl acetate. The extract was evaporated in vacuo and the residue was purified by preparative thin-layer chromatography (5% solution of methanol in dichloromethane) followed by crystallization from diethyl ether to give 3-bromo-8-[2,6-dichloro-3-(N-methyl-N'-methylureido)benzyloxy]-2-methylimidazo[1,2-a]pyridine (28 mg) as crystals.

mp: 192-193°C

NMR (CDCl₃, δ): 2.42 (3H, s), 2.79 (3H, d, J=5Hz), 3.20 (3H, s), 4.20 (1H, br d, J=5Hz), 5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.85 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.42 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

Example 22

The following compounds were obtained according to similar manners to those of Examples 20 or 21.

(1) 3-Bromo-8-[2,6-dichloro-3-(N'-phenylureido)-benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp: >250°C

NMR (DMSO-d₆, δ): 2.30 (3H, s), 5.45 (2H, s), 6.91-7.10 (3H), 7.31 (2H, t, J=9Hz), 7.41-7.62
(2) 3-Bromo-8-[2,6-dichloro-3-(N-methyl-N'-
5
trichloroacetylureido)benzyloxy]-2-methylimidazo-
[1,2-a]pyridine
mp: 160-164°C (dec.)
NMR (CDCl₃, δ): 2.43 (3H, s), 3.33 (3H, s), 5.54
(2H, s), 6.72 (1H, d, J=7.5Hz), 6.85 (1H, t,
J=7.5Hz), 7.41 (1H, d, J=10Hz), 7.52 (1H, d,
J=10Hz), 7.81 (1H, d, J=7.5Hz)

Example 23

To a solution of 8-(3-acetylamino-2,6-
dichlorobenzyloxy)-3-bromo-2-methylimidazo[1,2-a]pyridine
(222 mg) in N,N-dimethylformamide (2 ml) was added sodium
hydride (24 mg, 60% oil dispersion) in one portion at
ambient temperature. The mixture was stirred for half an
hour at the same temperature and iodomethane (142 mg) was
added thereto. After stirring for half an hour, the
mixture was poured into water. The separated oil was
extracted with ethyl acetate. The extracts were washed
with water, dried, and evaporated under reduced pressure.
The residue was purified by flash chromatography on silica
gel to give 3-bromo-8-[2,6-dichloro-3-(N-acetyl-N-
methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine (135
mg) as a solid.

mp: 201-204°C
NMR (CDCl₃, δ): 1.88 (3H, s), 2.46 (3H, s), 3.19
(3H, s), 5.52 (2H, s), 6.72 (1H, d, J=7.5Hz),
6.87 (1H, t, J=7.5Hz), 7.31 (1H, d, J=8Hz),
7.48 (1H, d, J=8Hz), 7.80 (1H, d, J=7.5Hz)

Example 24

The following compounds were obtained according to a
similar manner to that of Example 23.

(1) 3-Bromo-8-[2,6-dichloro-3-(N-ethyl-N-acetylamino)-benzyloxy]-2-methylimidazo[1,2-a]pyridine  
 mp : 161°C  
 NMR (CDCl₃, δ) : 1.15 (3H, t, J=7.5Hz), 1.83 (3H, s), 2.45 (3H, s), 3.30 (1H, m), 4.12 (1H, m),  
 5.51 (2H, s), 6.71 (1H, d, J=7.5Hz), 6.85 (1H, t, J=7.5Hz), 7.25 (1H, d, J=7.5Hz), 7.46 (1H, d,  
 J=7.5Hz), 7.78 (1H, d, J=7.5Hz)

(2) 3-Bromo-8-[2,6-dichloro-3-(N-propyl-N-acetylamino)-benzyloxy]-2-methylimidazo[1,2-a]pyridine  
 mp : 142-144°C  
 NMR (CDCl₃, δ) : 0.91 (3H, t, J=7Hz), 1.41-1.68 (2H), 1.81 (3H, s), 2.45 (3H, s), 3.14 (1H, dt,  
 J=9Hz and 7Hz), 4.03 (1H, dt, J=9Hz and 7Hz), 5.51 (2H, s), 6.70 (1H, d, J=7Hz), 6.85 (1H, t,  
 J=7Hz), 7.24 (1H, d, J=9Hz), 7.44 (1H, d, J=9Hz), 7.79 (1H, d, J=7Hz)

(3) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-(4-methoxybenzoyl)amino]benzyloxy]-2-methylimidazo-  
 [1,2-a]pyridine  
 mp : 168-169°C  
 NMR (CDCl₃, δ) : 2.45 (3H, s), 3.35 (3H, s), 3.78 (3H, s), 5.45 (2H, s), 6.59-6.86 (4H), 7.03-7.40  
 (4H), 7.76 (1H, d, J=7Hz)

(4) 3-Bromo-8-[2,6-dichloro-3-(N-methyl-N-propionyl-amino)benzyloxy]-2-methylimidazo[1,2-a]pyridine  
 mp : 170-171°C  
 NMR (CDCl₃, δ) : 1.08 (3H, t, J=7Hz), 2.01 (2H, q, J=7Hz), 2.43 (3H, s), 3.20 (3H, s), 5.50 (2H,  
 s), 6.70 (1H, d, J=7Hz), 6.84 (1H, t, J=7Hz),
7.28 (1H, d, J=9Hz), 7.44 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(5) 3-Bromo-8-[2,4,6-trichloro-3-(N-methyl-N-acetyl-
    amino)benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 1.84 (2.7H, s), 2.30 (0.3H, s),
    2.44 (3H, s), 3.15 (2.7H, s), 3.28 (0.3H, s),
    5.41 (0.2H, s), 5.47 (1.8H, s), 6.70 (1H, d, J=8Hz), 6.84 (1H, t, J=8Hz), 7.53 (0.1H, s),
    7.59 (0.9H, s), 7.77 (1H, d, J=8Hz)

(6) 3-Bromo-8-[2,6-dichloro-3-(N-methyl-N-phthalimido-
    acetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 229-230°C

(7) 8-[3-[N-(Acetysarcosyl)-N-methylamino]-2,6-dichloro-
    benzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 1.98 (0.5H, s), 2.11 (2.5H, s),
    2.43 (3H, s), 2.90 (0.5H, s), 3.07 (2.5H, s),
    3.21 (2.5H, s), 3.24 (0.5H, s), 3.30 (1H, d, J=16Hz), 4.27 (1H, d, J=16Hz), 5.50 (2H, s),
    6.70 (1H, d, J=8Hz), 6.84 (1H, t, J=8Hz), 7.33 (0.17H, d, J=8Hz), 7.48 (1.6H, s), 7.51 (0.17H, d, J=8Hz), 7.78 (1H, d, J=8Hz)

(8) 3-Bromo-8-[2,6-dichloro-3-(N-ethyl-N-(phthalimido-
    acetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 185-188°C

NMR (DMSO-d₆, δ) : 1.08 (3H, t, J=7Hz), 2.29 (3H, s), 3.88 (1H, d, J=16Hz), 4.11 (1H, d, J=16Hz),
    5.50 (2H, s), 6.95-7.09 (3H), 7.80-8.00 (6H)

(9) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-(2-phthalimidono-
    propionyl)amino]benzyloxy]-2-methylimidazo[1,2-a]-
    pyridine
NMR (CDCl₃, δ): 1.51-1.65 (6H), 2.41 (0.9H, s), 2.42 (2.1H, s), 3.22 (3H, s), 4.78 (0.3H, d, J=10Hz), 4.83 (1H, q, J=6Hz), 5.17 (0.3H, d, J=10Hz), 5.51 (1.4H, s), 6.46 (0.3H, d, J=8Hz), 6.70-6.90 (1.7H), 7.00 (1H, d, J=8Hz), 7.08 (1H, d, J=8Hz), 7.31 (0.3H, d, J=8Hz), 7.48 (0.3H, d, J=8Hz), 7.67-7.82 (5H)

(10) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-(3-phthalimido-propionyl)amino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine

mp : 213-215°C
NMR (CDCl₃, δ): 2.28-2.58 (2H), 2.41 (3H, s), 3.20 (3H, s), 3.82-4.12 (2H), 5.41 (1H, d, J=9Hz), 5.50 (1H, d, J=9Hz), 6.69 (2H, d, J=7Hz), 6.84 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.43 (1H, d, J=9Hz), 7.61-7.90 (5H)

**Example 25**

To a solution of 3-bromo-8-(2,6-dichloro-3-acetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine (222 mg) in N,N-dimethylformamide (2 ml) was added sodium hydride (24 mg, 60% oil dispersion) at ambient temperature. The mixture was stirred for half an hour at the same temperature and then ethyl bromoacetate (100 mg) was added thereto in one portion. The mixture was stirred for 2 hours at the same temperature and poured into water. The separated oil was extracted with dichloromethane. The extract was washed with water, dried, and concentrated in vacuo. The residue was purified by flash chromatography on silica gel to give an oil, which was dissolved in a mixture of ethanol (10 ml) and 1N sodium hydroxide solution (2 ml). The solution was refluxed for one and half hour. The organic solvent was removed under reduced pressure. The aqueous layer was adjusted to pH 4 with
diluted hydrochloric acid to give 3-bromo-8-[2,6-
dichloro-3-(N-carboxymethyl-N-acetylamino)benzyloxy]-2-
methylimidazo[1,2-a]pyridine (150 mg) as a white solid.

\[
\text{mp: 225-227°C (dec.)}
\]

\[
\begin{align*}
\text{NMR (DMSO-}d_6, \delta) &: 1.79 (3H, s), 2.31 (3H, s), \\
& 3.78 (1H, d, J=17Hz), 4.64 (1H, d, J=17Hz), \\
& 5.45 (2H, s), 6.96-7.06 (2H, m), 7.76 (2H, s), \\
& 7.95 (1H, m)
\end{align*}
\]

**Example 26**

To a solution of 3-bromo-8-[2-chloro-6-(N-acetyl-N-
methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine (103 mg) in tetrahydrofuran (2 ml) was added lithium aluminum hydride (16 mg) in several portions at 5°C. After the addition, the mixture was stirred for 2 hours at 5°C and then quenched with aqueous saturated ammonium chloride solution. The separated organic layer was washed with aqueous saturated ammonium chloride solution, dried, and concentrated in vacuo. The residue was purified by preparative thin-layer chromatography (ethyl acetate:n-hexane = 1:2, V/V) to give 3-bromo-8-[2-
chloro-6-(N-ethyl-N-methylamino)benzyloxy]-2-methyl-
imidazo[1,2-a]pyridine (41 mg).

\[
\text{mp: 124-126°C}
\]

\[
\begin{align*}
\text{NMR (CDCl}_3, \delta) &: 1.02 (3H, t, J=7.5Hz), 2.44 (3H, s), \\
& 2.71 (3H, s), 3.03 (2H, q, J=7.5Hz), 5.41 \\
& (2H, s), 6.73 (1H, d, J=7.5Hz), 6.81 (1H, t, \\
& J=7.5Hz), 7.01-7.28 (3H, m), 7.71 (1H, dd, J=7Hz \\
& \text{and 1.5Hz})
\end{align*}
\]

**Example 27**

A mixture of 3-bromo-8-[3-(N-acetoxyacetyl-N-methyl-
amino)-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]-
pyridine (1.42 g), potassium carbonate (761 mg), methanol 
(7 ml) and tetrahydrofuran (7 ml) was stirred at ambient
temperature for 1 hour. The mixture was partitioned between dichloromethane and water. The aqueous layer was extracted with dichloromethane twice. The combined organic layers were washed with brine, dried over magnesium sulfate and evaporated in vacuo. The residue was crystallized from ethanol to give 3-bromo-8-[2,6-dichloro-3-(N-glycoloyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine (1.16 g) as crystals.

mp : 217-218°C

NMR (CDCl₃, δ) : 2.45 (3H, s), 3.19-3.32 (4H), 3.69, 3.82 (each 1H, d, J=15Hz), 5.50 (2H, s), 6.70 (1H, d, J=7Hz), 6.83 (1H, t, J=7Hz), 7.29 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

Example 28

3-Bromo-8-[2,6-dichloro-3-(N-(glycoloylglycyl)-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Example 27.

mp : 163-165°C

NMR (CDCl₃, δ) : 2.40 (3H, s), 3.21 (3H, s), 3.29 (1H, dd, J=17Hz and 4Hz), 3.94 (2H, s), 4.32 (1H, dd, J=17Hz and 9Hz), 5.31 (1H, d, J=10Hz), 5.59 (1H, d, J=10Hz), 6.30 (1H, br s), 6.73 (1H, d, J=7Hz), 6.90 (1H, t, J=7Hz), 7.10 (1H, br s), 7.32 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.29 (1H, d, J=7Hz)

Example 29

To a suspension of sodium hydride (60% in oil, 9 mg) in N,N-dimethylformamide (1 ml) was added 3-bromo-8-[2,6-dichloro-3-(N-glycoloyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine (100 mg). After stirring for 10 minutes, methyl iodide (36 mg) was added thereto and the mixture was stirred at ambient temperature for 1 hour. To this mixture were added methyl iodide (36 mg) and N,N-dimethylformamide (1 ml). The mixture was stirred at
60°C for additional 2 hours. The reaction mixture was poured into water and extracted with ethyl acetate 3 times. The combined organic layers were washed with water 4 times and brine, dried over magnesium sulfate and evaporated in vacuo. The residue was purified by preparative thin-layer chromatography (dichloromethane: methanol = 20:1, V/V) followed by crystallization from ethanol-diethyl ether to give 3-bromo-8-[3-(N-methoxyacetyl-N-methylamino)-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine (20 mg) as crystals.

mp: 145-146°C

NMR (CDCl₃, δ): 2.43 (3H, s), 3.21 (3H, s), 3.35 (3H, s), 3.68, 3.32 (each 1H, d, J=15Hz), 5.50 (2H, s), 6.70 (1H, d, J=7Hz), 6.85 (1H, t, J=7Hz), 7.29, 7.46 (each 1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

Example 31
To a solution of 3-bromo-8-[2,6-dichloro-3-[N-(glycoloylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (200 mg) and triethylamine (76 mg) in methylene chloride (2 ml) was dropwise added mesyl chloride (52 mg) under ice-cooling, and the mixture was stirred for 1 hour at the same temperature. The mixture was washed with water twice and brine, dried over magnesium sulfate and concentrated in vacuo to give 3-bromo-8-[2,6-dichloro-3-[N-(mesyloxyacetylglucyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (221 mg).

NMR (CDCl₃, δ): 2.48 (3H, s), 3.19 (3H, s), 3.28 (3H, s), 3.61 (1H, dd, J=17Hz and 5Hz), 4.86 (1H, dd, J=17Hz and 5Hz), 4.18 (2H, s), 5.50 (2H, s), 6.78 (1H, d, J=7Hz), 6.91 (1H, t, J=7Hz), 7.25 (1H, br s), 7.32 (1H, d, J=9Hz), 7.51 (1H, d, J=9Hz), 7.80 (1H, d, J=7Hz)
mesyloxyacetylglucyl)-N-methylamino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine (100 mg) in methanol (2 ml)
was added sodium methoxide (317 mg), and the mixture was
stirred for 1 hour. Sodium methoxide (200 mg) was added
thereto, and the mixture was stirred for 3 hours. To the
reaction mixture was added water, and the mixture was
extracted with methylene chloride 3 times. The extracts
were combined, washed with brine, dried over magnesium
sulfate and concentrated in vacuo. The residue was
purified by preparative thin-layer chromatography
(methylene chloride:methanol = 10:1, V/V) to give 3-bromo-
8-[2,6-dichloro-3-[N-(mesyloxyacetylglucyl)-N-methylamino]-
benzyloxy]-2-methylimidazo[1,2-a]pyridine (60 mg).

NMR (CDCl₃, δ) : 2.44 (3H, s), 3.27 (3H, s), 3.59
(1H, dd, J=17Hz and 4Hz), 3.78-3.95 (3H), 5.48
(1H, d, J=9Hz), 5.52 (1H, d, J=9Hz), 6.71 (1H,
d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.31 (1H, d,
J=9Hz), 7.39 (1H, br s), 7.49 (1H, d, J=9Hz),
7.78 (1H, d, J=7Hz)

Example 32

To a solution of 3-bromo-8-[2,6-dichloro-3-[N-
glycoloyl-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-
pyridine (700 mg) and triethylamine (269 mg) in methylene
chloride (14 ml) was added mesyl chloride (224 mg) under
ice-cooling, and the mixture was stirred for 1 hour at
ambient temperature. The reaction mixture was washed with
water twice and brine, dried over magnesium sulfate and
concentrated in vacuo to give the residue of 3-bromo-8-
[2,6-dichloro-3-[N-mesyloxyacetyl-N-methylamino]benzylo-
xy]-2-methylimidazo[1,2-a]pyridine. The residue was
dissolved in N,N-dimethylformamide (7 ml) and potassium
phthalimide (362 mg) was added thereto at ambient
temperature under N₂ atmosphere. The mixture was stirred
for 1 day, and water was added thereto. The precipitate
was collected by filtration and dried to give 3-bromo-8-[2,6-dichloro-3-(N-methyl-N-phthalimidoacetamido)benzyloxy]-2-methylimidazo[1,2-a]pyridine (629 mg).

mp: 229-230°C  
NMR (CDCl₃, δ): 2.43 (3H, s), 3.26 (3H, s), 4.12 (2H, s), 5.53 (2H, s), 6.72 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.52 (2H, s), 7.68-7.92 (5H, m)

Example 33

3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-(phthalimidoacetylglucyl)amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Example 32.

mp: 148-150°C  
NMR (CDCl₃, δ): 2.42 (3H, s), 3.22 (3H, s), 3.53 (1H, dd, J=17Hz and 4Hz), 3.80 (1H, dd, J=17Hz and 5Hz), 4.39 (2H, s), 5.48 (2H, s), 6.70 (1H, d, J=7Hz), 6.78 (1H), 6.84 (1H, t, J=7Hz), 7.29 (1H, d, J=9Hz), 7.45 (1H, d, J=9Hz), 7.62-7.92 (5H)

Example 34

A mixture of 3-bromo-8-[2,6-dichloro-3-(N-methyl-N-phthalimidoacetamido)benzyloxy]-2-methylimidazo[1,2-a]pyridine (450 mg) and hydrazine hydrate (41 mg) in methanol (5 ml) were refluxed for 30 minutes, and hydrazine hydrate (41 mg) was added thereto, and further, the mixture was refluxed for 1.5 hours. The resulting precipitates were filtered off and the filtrate was concentrated in vacuo. The residue was dissolved in chloroform and the solution was washed with water twice and brine, dried over magnesium sulfate and concentrated in vacuo. The residue was crystallized with diethyl ether to give 8-[3-(N-glycyl-N-methylamino)-2,6-dichlorobenzylxy]-3-bromo-2-methylimidazo[1,2-a]pyridine (338 mg).
mp : 175-178°C

NMR (CDCl₃, δ) : 2.43 (3H, s), 2.92-3.20 (2H), 3.23 (3H, s), 5.49 (2H, s), 6.70 (1H, d, J=7Hz), 6.83 (1H, t, J=7Hz), 7.29, 7.47 (each 1H, d, J=9Hz), 7.79 (1H, d, J=7Hz)

Example 35

The following compounds were obtained according to a similar manner to that of Example 34.

(1) 8-[3-[N-(Glycylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.41 (3H, s), 3.22 (3H, s), 3.41 (3H, s), 3.60 (1H, dd, J=17Hz and 4Hz), 3.84 (1H, dd, J=17Hz and 4Hz), 5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.34 (1H, d, J=9Hz)

(2) 8-[3-(N-Glycyl-N-ethylamino)-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine

mp : 169-171°C

NMR (CDCl₃, δ) : 1.16 (3H, t, J=7Hz), 2.45 (3H, s), 2.92-3.40 (3H), 4.18 (1H), 5.50 (2H, s), 6.70 (1H, d, J=7Hz), 6.83 (1H, t, J=7Hz), 7.22 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(3) 3-Bromo-8-[2,6-dichloro-3-(N-DL-alanyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 1.11 (1.8H, d, J=6Hz), 1.15 (1.2H, d, J=6Hz), 2.45 (3H, s), 3.20 (0.4H, q, J=6Hz), 3.22 (3H, s), 3.36 (0.6H, q, J=6Hz), 5.43-5.56 (2H), 6.70 (1H, d, J=8Hz), 6.84 (1H, t, J=8Hz), 7.30 (0.6Hz, d, J=8Hz), 7.33 (0.4H, d, J=8Hz), 7.45 (0.4Hz, d, J=8Hz), 7.48 (0.6Hz, d, J=8Hz), 7.77 (1H, d, J=8Hz)
(4) 8-[3-(N-β-Alanyl-N-methylamino)-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine  
mp : 163-165°C  
NMR (CDCl₃, δ) : 2.10-2.25 (2H), 2.41 (3H, s), 2.85-2.99 (2H), 3.20 (3H, s), 5.50 (2H, s), 6.72 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(5) 3-Chloro-8-[2,6-dichloro-3-(N-glycyl-N-methylamino)-benzyloxy]-2-methylimidazo[1,2-a]pyridine  
mp : 184-186°C  
NMR (CDCl₃, δ) : 2.43 (3H, s), 3.01 (1H, d, J=17Hz), 3.12 (1H, d, J=17Hz), 3.22 (3H, s), 5.50 (2H, s), 6.70 (1H, d, J=7Hz) 6.86 (1H, t, J=7Hz), 7.29 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.72 (1H, d, J=7Hz)

Example 36

To a solution of 3-bromo-8-[2,6-dichloro-3-(N-glycyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine (100 mg) in methylene chloride (2 ml) were added pyridine (17 mg), 4-dimethylaminopyridine (10 mg) and acetic anhydride (32 mg), and the mixture was stirred for 1.5 hours. The mixture was washed with water twice and brine, dried over magnesium sulfate and concentrated in vacuo. The residue was purified by silica gel column chromatography (methylene chloride:methanol = 5:1, V/V) to give 8-[3-[N-(acetylglycyl)-N-methylamino]-2,6-dichlorobenzyl-3-bromo-2-methylimidazo[1,2-a]pyridine (74 mg).  
mp : 187-189°C  
NMR (CDCl₃, δ) : 2.01 (3H, s), 2.43 (3H, s), 3.24 (3H, s), 3.51, 3.78 (each 1H, dd, J=17Hz and 4Hz), 5.49 (2H, s), 6.45 (1H, br s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.30, 7.49 (each
Example 37

To a mixture of 3-bromo-8-[2,6-dichloro-3-(N-glycyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine (100 mg), pyridine (25 mg) and methylene chloride (2 ml) was added propionyl chloride (0.02 ml) at ambient temperature and the mixture was stirred at the same temperature for 30 minutes. The mixture was partitioned between methylene chloride and water, and the organic layer was washed with water, dried over magnesium sulfate and concentrated in vacuo to give 3-bromo-8-[2,6-dichloro-3-[N-methyl-N-(propionylglycyl)amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (111 mg).

NMR (CDCl₃, δ) : 1.25 (3H, t, J=6Hz), 2.27 (2H, q, J=6Hz), 2.43 (2.8H, s), 3.30 (0.2H, s), 3.52 (1H, dd, J=17Hz and 4Hz), 3.80 (1H, dd, J=17Hz and 4Hz), 5.48 (2H, s), 6.43 (1H, t like), 6.73 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.30 (1H, d, J=8Hz), 7.47 (1H, d, J=8Hz), 7.76 (1H, d, J=7Hz)

Example 38

Pivaloyl chloride (0.04 ml) was added dropwise to a mixture of tert-butoxycarbonyl-L-proline (77 mg), N-methylmorpholine (0.04 ml) and dichloromethane (3 ml) under a dry ice-tetrachloromethane bath cooling. This mixture was stirred for 5 minutes under an ice-water bath cooling and cooled under a dry ice-tetrachloromethane bath cooling. To the mixture was added a solution of 8-[3-(N-glycyl-N-methylamino)-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine (139 mg) in dichloromethane (4 ml). The mixture was stirred for 30 minutes at ambient temperature and washed with saturated aqueous solution of sodium hydrogen carbonate and water. The organic layer was dried over magnesium sulfate and
evaporated in vacuo. The residue was purified by preparative thin-layer chromatography (dichloromethane:methanol = 19:1, V/V) to give 3-bromo-8-[3-[N-(tert-butoxycarbonyl-L-prolylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]-pyridine (72 mg) (Compound A), and 3-bromo-8-[2,6-dichloro-3-(N-methyl-N-pivaloylglycylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine (93 mg) (Compound B).

Compound A:

NMR (CDCl$_3$, δ): 1.45 (9H, br s), 1.78-2.22 (4H), 2.43 (3H, s), 3.23 (3H, s), 3.32-3.90 (4H), 4.25 (1H, m), 5.48 (2H, s), 6.72 (1H, d, J=8Hz), 6.86 (1H, t, J=8Hz), 7.31 (1H, d, J=9Hz), 7.47 (1H, d, J=9Hz), 7.77 (1H, d, J=8Hz)

Compound B:

NMR (CDCl$_3$, δ): 1.20 (9H, s), 2.44 (3H, s), 3.26 (3H, s), 3.49 (1H, dd, J=16Hz and 4Hz), 3.77 (1H, dd, J=16Hz and 4Hz), 5.48 (2H, s), 6.64 (1H, t, like), 6.72 (1H, d, J=8Hz), 6.85 (1H, t, J=8Hz), 7.31 (1H, d, J=9Hz), 7.47 (1H, d, J=9Hz), 7.77 (1H, d, J=8Hz)

Example 39

A mixture of 3-bromo-8-[2,6-dichloro-3-(N-glycyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine (150 mg), 4-pentenoic acid (33 mg), N-ethyl-N'-(3-dimethylaminopropyl)carbodiimide hydrochloride (77 mg), 1-hydroxybenzotriazole (64 mg) and N,N-dimethylformamide (1.5 ml) was stirred for 2 hours at ambient temperature. Water was added thereto, and the mixture was extracted with methylene chloride three times. The organic layers were combined, washed with water four times and brine, dried over magnesium sulfate and concentrated in vacuo.
The residue was purified by silica gel column chromatography (methylene chloride:methanol = 30:1, V/V) to give 3-bromo-8-[2,6-dichloro-3-[N-(4-pentenoylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (172 mg).

NMR (CDCl$_3$, $\delta$): 2.24-2.42 (4H), 2.44 (3H, s), 3.26 (3H, s), 3.52 (1H, dd, J=17Hz and 4Hz), 3.80 (1H, dd, J=17 and 4Hz), 4.96-5.16 (2H), 5.49 (2H, s), 6.46 (1H, br s), 6.72 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

Example 40

The following compounds were obtained according to similar manners to those of Examples 36 to 39.

(1) 3-Bromo-8-[3-[N-(butyrylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 0.94 (3H, t, J=7Hz), 1.65 (2H, m), 2.20 (2H, t, J=7Hz), 2.44 (3H, s), 3.26 (3H, s), 3.52 (1H, dd, J=16Hz and 4Hz), 3.80 (1H, dd, J=16Hz and 4Hz), 5.49 (2H, s), 6.41 (1H, t like), 6.72 (1H, d, J=8Hz), 6.86 (1H, t, J=8Hz), 7.31 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.77 (1H, d, J=8Hz)

(2) 3-Bromo-8-[3-[N-(isobutyrylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 1.14 (6H, d, J=6Hz), 2.42 (1H, m), 2.44 (3H, s), 3.25 (3H, s), 3.50 (1H, dd, J=16Hz and 4Hz), 3.79 (1H, dd, J=16Hz and 5Hz), 5.48 (2H, s), 6.45 (1H, t like), 6.72 (1H, d, J=8Hz), 6.86 (1H, t, J=8Hz), 7.31 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.77 (1H, d, J=8Hz)
(3) 3-Bromo-8-[2,6-dichloro-3-[N-(cyclopropylcarbonyl-
glycyl)-N-methylamino]benzyloxy]-2-methylimidazo-
[1,2-a]pyridine
NMR (CDCl₃, δ): 0.70-1.00 (4H), 1.47 (1H, m), 2.44
(3H, s), 3.25 (3H, s), 3.54 (1H, dd, J=16Hz and
4Hz), 3.81 (1H, dd, J=16Hz and 4Hz), 5.48 (2H,
s), 6.58 (1H, t like), 6.7 (1H, d, J=8Hz), 6.86
(1H, t, J=8Hz), 7.30 (1H, d, J=9Hz), 7.46 (1H,
d, J=9Hz), 7.77 (1H, d, J=8Hz)

(4) 3-Bromo-8-[2,6-dichloro-3-[N-(trifluoroacetylglycyl)-
N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-
pyridine
NMR (CDCl₃, δ): 2.42 (3H, s), 3.29 (3H, s), 3.61
(1H, dd, J=18Hz and 4Hz), 3.83 (1H, dd, J=18Hz
and 4Hz), 5.48 (1H, d, J=7Hz), 5.53 (1H, d,
J=7Hz), 6.71 (1H, d, J=7Hz), 6.88 (1H, t,
J=7Hz), 7.31 (1H, d, J=9Hz), 7.41 (1H, br s),
7.50 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(5) 8-[3-[N-(Benzoylglycyl)-N-methylamino]-2,6-dichloro-
benzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine
mp : 126°C (dec.)
NMR (CDCl₃, δ): 2.44 (3H, s), 3.29 (3H, s), 3.73
(1H, dd, J=18Hz and 4Hz), 3.99 (1H, dd, J=18Hz
and 4Hz), 5.48 (1H, d, J=7Hz), 5.54 (1H, d,
J=7Hz), 6.73 (1H, d, J=7Hz), 6.88 (1H, t,
J=7Hz), 7.18 (1H, br s), 7.31-7.59 (5H),
7.72-7.88 (3H)

(6) 3-Bromo-8-[2,6-dichloro-3-[N-(cyclohexylcarbonyl-
glycyl)-N-methylamino]benzyloxy]-2-methylimidazo-
[1,2-a]pyridine
NMR (CDCl₃, δ): 1.14-1.94 (10H), 2.14 (1H, m), 2.44
(3H, s), 3.26 (3H, s), 3.50 (1H, dd, J=18Hz and
(7) 3-Bromo-8-[2,6-dichloro-3-[N-(phenylacetylglucyl)-N-
-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 2.43 (3H, s), 3.21 (3H, s), 3.49
(1H, dd, J=17Hz and 4Hz), 3.59 (2H, s), 3.75
(1H, dd, J=17Hz and 5Hz), 5.48 (2H, s), 6.40
(1H, t like), 6.72 (1H, d, J=8Hz), 6.86 (1H, t,
J=8Hz), 7.21-7.43 (6H), 7.46 (1H, d, J=8Hz),
7.77 (1H, d, J=8Hz)

(8) 3-Bromo-8-[3-[N-(tert-butoxycarbonyl-D-prolylglycyl)-
-N-methylamino]-2,6-dichlorobenzyloxy]-2-methyl-
imidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 1.45 (9H, br s), 1.78-2.22 (4H),
2.43 (3H, s), 3.23 (3H, s), 3.32-3.90 (4H), 4.25
(1H, m), 5.48 (2H, s), 6.72 (1H, d, J=8Hz), 6.86
(1H, t, J=8Hz), 7.31 (1H, d, J=9Hz), 7.47 (1H,
d, J=9Hz), 7.77 (1H, d, J=8Hz)

(9) 8-[3-[N-(Acetoxyacetylglucyl)-N-methylamino]-2,6-
dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]-
pyridine
NMR (CDCl₃, δ) : 2.20 (3H, s), 2.44 (3H, s), 3.26
(3H, s), 3.59 (1H, dd, J=17Hz and 4Hz), 3.82
(1H, dd, J=17Hz and 4Hz), 4.59 (2H, s), 5.46
(1H, d, J=10Hz), 5.52 (1H, d, J=10Hz), 6.71 (1H,
d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.11 (1H, br s),
7.31 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.78
(1H, d, J=7Hz)

(10) 8-[3-[N-(Acetylglucyl)-N-ethylamino]-2,6-dichloro-
benzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 1.19 (3H, t, J=7Hz), 2.00 (3H, s),
2.42 (3H, s), 3.29 (1H), 3.48 (1H, dd, J=17Hz
and 5Hz), 3.72 (1H, dd, J=17Hz and 5Hz), 4.21
(1H), 5.49 (2H, s), 6.46 (1H, br s), 6.71 (1H,
d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.22 (1H, d,
J=9Hz), 7.48 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(11) 8-[3-[N-(Acetyl-DL-alanyl)-N-methylamino]-2,6-
dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]-
pyridine
NMR (CDCl₃, δ) : 1.18 (1.5H, d, J=6Hz), 1.20 (1.5H,
d, J=6Hz), 1.95 (1.5H, s), 1.99 (1.5H, s), 2.43
(3H, s), 3.22 (3H, s), 4.26-4.51 (1H), 5.47 (1H,
s), 5.51 (1H, s), 6.15 (0.5H, d, J=8Hz), 6.43
(1H, d, J=8Hz), 6.65-6.90 (2H), 7.30 (0.5H, d,
J=8Hz), 7.47 (0.5H, d, J=8Hz), 7.50 (1H, s),
7.76 (1H, d, J=8Hz)

(12) 8-[3-[N-(Acetyl-β-alanyl)-N-methylamino]-2,6-
dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]-
pyridine
NMR (CDCl₃, δ) : 1.94 (3H, s), 2.10-2.30 (2H), 2.42
(3H, s), 3.20 (3H, s), 3.39-3.56 (2H), 5.49 (2H,
s), 6.39 (1H, br s), 6.70 (1H, d, J=7Hz), 6.85
(1H, t, J=7Hz), 7.26 (1H, d, J=9Hz), 7.45 (1H,
d, J=9Hz), 7.78 (1H, d, J=7Hz)

(13) 3-Bromo-8-[2,6-dichloro-3-[N-methoxycarbonylglycyl]-
N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-
pyridine
NMR (CDCl₃, δ) : 2.42 (3H, s), 3.22 (3H, s), 3.49
(1H, dd, J=17Hz and 5Hz), 3.67 (3H, s), 3.72
(1H, dd, J=17Hz and 5Hz), 5.48 (2H, s), 5.54
(1H, br s), 6.71 (1H, d, J=7Hz), 6.82 (1H, t,
J=7Hz), 7.31 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(14) 3-Bromo-8-[2,6-dichloro-3-[N-(valerylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 0.91 (3H, t, J=7Hz), 1.34 (2H, m),
1.61 (2H, m), 2.22 (2H, t, J=7Hz), 2.45 (3H, s),
3.26 (3H, s), 3.52 (1H, dd, J=16Hz and 4Hz),
3.80 (1H, dd, J=16Hz and 4Hz), 5.48 (2H, s),
6.41 (1H, t like), 6.71 (1H, d, J=8Hz), 6.86 (1H, dd, J=8Hz and 7Hz), 7.30 (1H, d, J=9Hz),
7.48 (1H, d, J=9Hz), 7.77 (1H, d, J=7Hz)

(15) 3-Bromo-8-[2,6-dichloro-3-[N-(isovalerylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 0.96 (6H, d, J=7Hz), 2.02-2.17 (3H),
2.44 (3H, s), 3.26 (3H, s), 3.52 (1H, dd, J=16Hz and 4Hz),
3.80 (1H, dd, J=16Hz and 4Hz), 5.49 (2H, s),
6.39 (1H, t like), 6.72 (1H, d, J=8Hz),
6.86 (1H, dd, J=8Hz and 7Hz), 7.30 (1H, d, J=9Hz),
7.48 (1H, d, J=9Hz), 7.77 (1H, d, J=7Hz)

(16) 3-Bromo-8-[2,6-dichloro-3-[N-(2-pyridylacetylglucyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 2.45 (3H, s), 3.22 (3H, s), 3.54 (1H, dd, J=18Hz and 5Hz),
3.76 (2H, s), 3.81 (1H, dd, J=18Hz and 5Hz), 5.48 (2H, s),
6.71 (1H, d, J=7Hz), 6.85 (1H, t, J=7Hz), 7.16-7.32 (3H),
7.47 (1H, d, J=9Hz), 7.68 (1H, t, J=7Hz),
7.77 (1H, d, J=7Hz), 7.93 (1H, br t, J=5Hz)

(17) 3-Bromo-8-[2,6-dichloro-3-[N-(3-pyridylacetylglucyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ):  2.43 (3H, s), 3.22 (3H, s), 3.51 (1H, dd, J=17Hz and 5Hz), 3.58 (2H, s), 3.79 (1H, dd, J=17Hz and 5Hz), 5.48 (2H, s), 6.50 (1H, br t, J=5Hz), 6.70 (1H, d, J=7Hz), 6.85 (1H, t, J=7Hz), 7.21-7.32 (2H), 7.48 (1H, d, J=9Hz), 7.66 (1H, d, J=8Hz), 7.78 (1H, d, J=7Hz), 8.49-8.59 (2H)

(18) 3-Bromo-8-[2,6-dichloro-3-[N-(3-ethoxycarbonylpropionylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ):  1.26 (3H, t, J=7Hz), 2.44 (3H, s), 2.49-2.69 (4H), 3.25 (3H, s), 3.53 (1H, dd, J=16Hz and 4Hz), 3.79 (1H, dd, J=16Hz and 4Hz), 4.3 (2H, q, J=7Hz), 5.50 (2H, s), 6.56 (1H, t like), 6.71 (1H, d, J=8Hz), 6.86 (1H, dd, J=8Hz and 6Hz), 7.30 (1H, d, J=10Hz), 7.48 (1H, d, J=10Hz), 7.77 (1H, d, J=6Hz)

(19) 3-Bromo-8-[2,6-dichloro-3-[N-(bromoacetylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ):  2.44 (3H, s), 3.28 (3H, s), 3.57 (1H, dd, J=18Hz and 4Hz), 3.80 (1H, dd, J=18Hz and 4Hz), 3.88 (2H, s), 5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.39 (1H, br t, J=4Hz), 7.50 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(20) 3-Bromo-8-[2,6-dichloro-3-[N-(phthalimidoacetylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp: 211-213°C

NMR (CDCl₃, δ):  2.43 (3H, s), 3.22 (3H, s), 3.56 (1H, dd, J=18Hz and 5Hz), 3.82 (1H, dd, J=18Hz and 5Hz), 4.40 (2H, s), 5.49 (2H, s), 6.69 (1H, d, J=7Hz), 6.78 (1H, br t, J=5Hz), 6.82 (1H, t,
(21) 3-Bromo-8-[2,6-dichloro-3-[N-(4-nitrophenoxycarbonyl- 
glycyl)-N-methylamino]benzyloxy]-2-methylimidazo- 
[1,2-a]pyridine

mp : 166-168°C

NMR (CDCl₃, δ) : 2.43 (3H, s), 3.29 (3H, s), 3.60
(1H, dd, J=17Hz and 5Hz), 3.81 (1H, dd, J=17Hz
and 5Hz), 5.50 (2H, s), 6.05 (1H, br t, J=5Hz),
6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz),
7.22-7.38 (3H), 7.49 (1H, d, J=9Hz), 7.79 (1H,
d, J=7Hz), 8.22 (2H, d, J=9Hz)

(22) 3-Chloro-8-[2,6-dichloro-3-[N-(phenylacetylglycyl)- 
N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]- 
pyridine

mp : 198-200°C

NMR (CDCl₃, δ) : 2.42 (3H, s), 3.21 (3H, s), 3.49
(1H, dd, J=15Hz and 4Hz), 3.60 (2H, s), 3.76
(1H, dd, J=15Hz and 4Hz), 5.46 (2H, s), 6.40
(1H, t like), 6.69 (1H, d, J=8Hz), 6.84 (1H, dd,
J=8Hz and 7Hz), 7.20-7.50 (7H), 7.71 (1H, d,
J=6Hz)

(23) 3-Chloro-8-[2,6-dichloro-3-[N-(butyrylglycyl)-N- 
methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 103-104°C

NMR (CDCl₃, δ) : 0.93 (3H, t, J=7Hz), 1.54-1.77 (2H),
2.20 (2H, t, J=7Hz), 2.42 (3H, s), 3.23 (3H, s),
3.51 (1H, dd, J=17Hz and 4Hz), 3.79 (1H, dd,
J=17Hz and 4Hz), 5.49 (2H, s), 6.41 (1H, br t,
J=4Hz), 6.70 (1H, d, J=7Hz), 6.86 (1H, t,
J=7Hz), 7.31 (1H, d, J=9Hz), 7.47 (1H, d,
J=9Hz), 7.71 (1H, d, J=7Hz)
Example 41

To a suspension of 3-bromo-8-[2,6-dichloro-3-(N-glycyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine (150 mg) in formic acid (1 ml) was added acetic anhydride (65 mg), and the mixture was stirred for 1.5 hours. Acetic anhydride (40 mg) was added thereto, and the mixture was stirred for 3 hours. The mixture was concentrated in vacuo, and a saturated aqueous solution of sodium bicarbonate was added thereto, and the mixture was extracted with methylene chloride three times. The organic layers were combined, washed with brine, dried over magnesium sulfate, and concentrated in vacuo. The residue was purified by preparative thin-layer chromatography (methylene chloride:methanol = 5:1, V/V) to give 3-bromo-8-[2,6-dichloro-3-[N-(formylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (102 mg).

mp : 211-212°C

NMR (CDCl₃, δ) : 2.43 (3H, s), 3.26 (3H, s), 3.58 (1H, dd, J=17Hz and 5Hz), 3.82 (1H, dd, J=17Hz and 5Hz), 5.47 (1H, d, J=7Hz), 5.52 (1H, d, J=7Hz), 6.61 (1H, br s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.50 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 8.20 (1H, s)

Example 42

To a solution of 3-bromo-8-[2,6-dichloro-3-(N-glycyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine (98 mg) in methylene chloride (2 ml) was added methyl isocyanate (0.02 ml), and the mixture was stirred for 1 hour at ambient temperature. The mixture was concentrated in vacuo to give 3-bromo-8-[2,6-dichloro-3-[N-methyl-N-(N'-methylureidoacetyl)amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (9.5 mg).
NMR (CDCl₃, δ): 2.44 (3H, s), 2.74 (3H, d, J=5Hz), 3.23 (3H, s), 3.58 (1H, dd, J=16Hz and 5Hz), 4.76 (1H, br s), 5.35 (1H, t like), 5.49 (2H, s), 6.74 (1H, d, J=8Hz), 6.88 (1H, t, J=8Hz), 7.35 (1H, d, J=8Hz), 7.48 (1H, d, J=8Hz), 7.78 (1H, d, J=8Hz)

Example 43
A mixture of 3-bromo-8-[2,6-dichloro-3-[N-(4-nitrophenoxycarbonylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (100 mg), 3-aminopyridine (18 mg) and dioxane (1 ml) was refluxed for 1.5 hours under nitrogen atmosphere. The reaction mixture was washed with water four times and brine, dried over magnesium sulfate, and concentrated in vacuo. The residue was purified by preparative thin-layer chromatography (methylene chloride:methanol = 10:1, V/V) to give 3-bromo-8-[2,6-dichloro-3-[N-[N'-((3-pyridyl)ureidoacetyl)-N-methylamino]-benzyloxy]-2-methylimidazo[1,2-a]pyridine (58 mg).

NMR (CDCl₃, δ): 2.40 (3H, s), 3.22 (3H, s), 3.82 (2H, d, J=5Hz), 5.42 (1H, d, J=10Hz), 5.52 (1H, d, J=10Hz), 6.07 (1H, br t, J=5Hz), 6.72 (1H, d, J=7Hz), 6.89 (1H, t, J=7Hz), 7.11 (1H, dd, J=9Hz and 5Hz), 7.34 (1H, d, J=9Hz), 7.41 (1H, d, J=9Hz), 7.79 (1H, d, J=7Hz), 7.86 (1H, d, J=9Hz), 8.18 (1H, d, J=5Hz), 8.31 (1H, br s), 8.41 (1H, br s)

Example 44
The following compounds were obtained according to similar manners to those of Examples 42 or 43.

(1) 3-Bromo-8-[2,6-dichloro-3-[N-(N'-ethylureidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 126-127°C

NMR (CDCl₃, δ) : 1.20 (3H, t, J=7Hz), 2.42 (3H, s), 3.09-3.33 (5H), 3.56 (1H, dd, J=17Hz and 5Hz), 3.80 (1H, dd, J=17Hz and 5Hz), 4.71 (1H, br t, J=5Hz), 5.32 (1H, br t, J=5Hz), 5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.46 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

10 (2) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-(N'-phenylureidoacetyl)amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 144-147°C

NMR (CDCl₃, δ) : 2.42 (3H, s), 3.22 (3H, s), 3.70 (1H, dd, J=17Hz and 4Hz), 3.86 (1H, dd, J=17Hz and 4Hz), 5.49 (2H, s), 5.79 (1H, br t, J=4Hz), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.02 (1H), 7.19-7.30 (5H), 7.32 (1H, d, J=9Hz), 7.42 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

15 (3) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-cyclohexylureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 0.98-2.01 (10H), 2.43 (3H, s), 3.22 (3H, s), 3.42 (1H, m), 3.52 (1H, dd, J=17Hz and 5Hz), 3.80 (1H, dd, J=17Hz and 5Hz), 4.56 (1H, d, J=8Hz), 5.26 (1H, br t, J=5Hz), 5.48 (2H, s), 6.71 (1H, d, J=8Hz), 6.87 (1H, t, J=8Hz), 7.32 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.78 (1H, d, J=8Hz)

20 (4) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-(a-naphthyl)ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 148-151°C
NMR (CDCl₃, δ) : 2.41 (3H, s), 3.16 (3H, s), 3.61 (1H, dd, J=18Hz and 4Hz), 3.87 (1H, dd, J=18Hz and 4Hz), 5.42 (2H, s), 6.01 (1H, br t, J=4Hz), 6.69 (1H, d, J=8Hz), 6.85 (1H, t, J=8Hz), 7.18 (1H, br s), 7.25 (1H, d, J=9Hz), 7.40 (1H, d, J=9Hz), 7.42-7.56 (4H), 7.62-7.92 (3H), 7.99-8.10 (1H, m)

(5) 3-Bromo-8-[2,6-dichloro-3-[N-(N'-benzyleidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.42 (3H, s), 3.11 (3H, s), 3.57 (1H, dd, J=17Hz and 5Hz), 3.82 (1H, dd, J=17Hz and 5Hz), 4.32 (2H, d, J=5Hz), 5.30 (1H, br t, J=5Hz), 5.48 (2H, s), 5.58 (1H, br t, J=5Hz), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.18-7.39 (6H), 7.46 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(6) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-(m-tolyl)ureido]-2-acetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.29 (3H, s), 2.42 (3H, s), 3.22 (3H, s), 3.65 (1H, dd, J=17Hz and 5Hz), 3.85 (1H, dd, J=17Hz and 5Hz), 5.47 (2H, s), 5.96 (1H, br t, J=5Hz), 6.71 (1H, d, J=7Hz), 6.80-6.91 (2H), 7.00-7.39 (4H), 7.32 (1H, d, J=9Hz), 7.43 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(7) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-(p-tolyl)ureido]-2-acetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.30 (3H, s), 2.42 (3H, s), 3.21 (3H, s), 3.62 (1H, dd, J=17Hz and 5Hz), 3.85 (1H, dd, J=17Hz and 5Hz), 5.47 (2H, s), 5.91
(1H, br t, J=5Hz), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.01-7.21 (5H), 7.31 (1H, d, J=9Hz), 7.43 (1H, d, J=9Hz), 7.3 (1H, d, J=7Hz)

5 (8) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-{(4-methoxyphenyl)-
ureidoacetyl]-N-methylamino]benzyloxy]-2-methyl-
imidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.42 (3H, s), 3.21 (3H, s), 3.62 (1H, dd, J=17Hz and 5Hz), 3.78 (3H, s), 3.83 (1H, dd, J=17Hz and 5Hz), 5.48 (2H, s), 5.78 (1H, br t, J=5Hz), 6.68-6.96 (5H), 7.12-7.25 (2H), 7.32 (1H, d, J=9Hz), 7.45 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

10 (9) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-{(4-fluorophenyl)ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.41 (3H, s), 3.22 (3H, s), 3.72 (1H, dd, J=17Hz and 5Hz), 3.86 (1H, dd, J=17Hz and 5Hz), 5.43 (1H, d, J=9Hz), 5.51 (1H, d, J=9Hz), 6.18 (1H, br t, J=5Hz), 6.72 (1H, d, J=7Hz), 6.89 (1H, t, J=7Hz), 7.26-7.45 (6H), 7.79 (1H, d, J=7Hz), 8.42 (1H, br s)

20 (10) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-{(4-fluorophenyl)-
ureidoacetyl]-N-methylamino]benzyloxy]-2-methyl-
imidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.42 (3H, s), 3.21 (3H, s), 3.69 (1H, dd, J=17Hz and 5Hz), 3.83 (1H, dd, J=17Hz and 5Hz), 5.48 (2H, s), 5.91 (1H, br t, J=5Hz), 6.71 (1H, d, J=7Hz), 6.81-7.00 (3H), 7.16-7.29 (2H), 7.34 (1H, d, J=9Hz), 7.42 (1H, d, J=9Hz), 7.58 (1H, br s), 7.78 (1H, d, J=7Hz)

35 (11) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-n-propylureido-
acetyl)-N-methylamino]benzyloxy]-2-methylimidazo-
[1,2-a]pyridine

NMR (CDCl₃, δ): 0.90 (3H, t, J=7Hz), 1.39-1.59
(2H), 2.43 (3H, s), 3.10 (2H, q, J=6Hz), 3.22
(3H, s), 3.54 (1H, dd, J=17Hz and 4Hz), 3.80
(1H, dd, J=17Hz and 4Hz), 4.73 (1H, br t,
J=6Hz), 5.32 (1H, br t, J=4Hz), 5.49 (2H, s),
6.71 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.32
(1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.77 (1H,
d, J=7Hz)

(12) 3-Bromo-8-[2,6-dichloro-3-[N-(N'-isopropylureido-
acetyl)-N-methylamino]benzyloxy]-2-methylimidazo-
[1,2-a]pyridine

NMR (CDCl₃, δ): 1.12 (6H, d, J=7Hz), 2.45 (3H, s),
3.52 (1H, dd, J=17Hz and 4Hz), 3.70-3.88 (2H),
4.51 (1H, br d, J=7Hz), 5.23 (1H, br t, J=4Hz),
5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t,
J=7Hz), 7.32 (1H, d, J=9Hz), 7.48 (1H, d,
J=9Hz), 7.78 (1H, d, J=7Hz)

(13) 3-Bromo-8-[2,6-dichloro-3-[N-(N'-allylureidoacetyl)]-
N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-
pyridine

NMR (CDCl₃, δ): 2.43 (3H, s), 3.22 (3H, s), 3.57
(1H, dd, J=17Hz and 5Hz), 3.70-3.89 (3H), 4.94
(1H, br t, J=5Hz), 5.04-5.28 (2H), 5.42-5.55
(3H), 5.83 (1H, m), 6.71 (1H, d, J=7Hz), 6.87
(1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.48 (1H,
d, J=9Hz), 7.78 (1H, d, J=7Hz)

(14) 3-Bromo-8-[2,6-dichloro-3-[N-(N'-4-pyridyl)u-
reidoacetyl]-N-methylamino]benzyloxy]-2-methyl-
imidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 2.41 (3H, s), 3.20 (3H, s),
3.80 (2H, d, J=5Hz), 5.41 (1H, d, J=10Hz), 5.51 (1H, d, J=10Hz), 6.19 (1H, br t, J=5Hz), 6.76 (1H, d, J=7Hz), 6.90 (1H, t, J=7Hz), 7.19 (2H, d, J=6Hz), 7.32 (1H, d, J=9Hz), 7.40 (1H, d, J=9Hz), 7.79 (1H, d, J=7Hz), 8.28 (2H, d, J=6Hz), 8.95 (1H, br s)

(15) 3-Chloro-8-[2,6-dichloro-3-[N-(N'-phenylureidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 230-231°C
NMR (CDCl₃, δ) : 2.42 (3H, s), 3.23 (3H, s), 3.69 (1H, dd, J=16Hz and 4Hz), 3.85 (1H, dd, J=16Hz and 4Hz), 5.49 (2H, s), 5.93 (1H, t like), 6.71 (1H, d, J=8Hz), 6.86 (1H, dd, J=8Hz and 7Hz), 7.20-7.37 (7H), 7.44 (1H, d, J=9Hz), 7.74 (1H, d, J=7Hz)

(16) 3-Chloro-8-[2,6-dichloro-3-[N-(N'-cyclohexylureidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 137-140°C
NMR (CDCl₃, δ) : 0.98-1.47 (6H), 1.50-1.78 (2H), 1.81-2.00 (2H), 2.43 (3H, s), 3.23 (3H, s), 3.43 (1H, m), 3.52 (1H, dd, J=16Hz and 4Hz), 3.79 (1H, dd, J=16Hz and 4Hz), 4.50 (1H, d, J=10Hz), 5.21 (1H, t like), 5.48 (2H, s), 6.70 (1H, d, J=8Hz), 6.86 (1H, dd, J=8Hz and 7Hz), 7.34 (1H, d, J=8Hz), 7.47 (1H, d, J=8Hz), 7.72 (1H, d, J=7Hz)

(17) 3-Chloro-8-[2,6-dichloro-3-[N-(N'-ethylureidoacetetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 1.11 (3H, t, J=6Hz), 2.42 (3H, s),
Example 45

To a mixture of 8-[3-(N-glycyl-N-methylamino)-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine (100 mg), acetic acid (0.5 ml) and water (1.0 ml) was added an aqueous solution (1 ml) of sodium cyanate (138 mg) at 40°C. The mixture was stirred at the same temperature for 1 hour and evaporated in vacuo. The residue was partitioned into dichloromethane and an saturated aqueous solution of sodium hydrogen carbonate and the organic layer was separated. The aqueous layer was extracted with dichloromethane twice. The combined organic layers were washed with brine, dried over magnesium sulfate and evaporated in vacuo. The residue was purified by preparative thin-layer chromatography (dichloromethane:methanol = 10:1, V/V) followed by crystallization from ethyl acetate - diethyl ether to give 3-bromo-8-[2,6-dichloro-3-[N-methyl-N-(ureidoacetyl)amino]benzyl-oxy]-2-methylimidazo[1,2-a]pyridine (72 mg) as crystals.

mp : 147-149°C

NMR (CDCl₃, δ) : 2.44 (3H, s), 3.23 (3H, s), 3.52 (1H, dd, J=17Hz and 5Hz), 3.80 (1H, dd, J=17Hz and 5Hz), 4.76 (2H, br s), 5.49 (2H, s), 5.82 (1H, t, J=5Hz), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.33 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.79 (1H, d, J=7Hz)

Example 46

To a solution of 3-bromo-8-[2,6-dichloro-3-
(N-glycyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]-pyridine (150 mg) in methylene chloride (3 ml) was added phenylisothiocyanate (86 mg), and the mixture was refluxed for 1 hour under nitrogen atmosphere. The mixture was concentrated in vacuo, and the residue was purified by silica gel column chromatography (methylene chloride:methanol = 40:1, V/V) to give 3-bromo-8-[2,6-dichloro-3-[N-(N'-phenylthioureaidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (190 mg).

NMR (CDCl₃, δ) : 2.42 (3H, s), 3.21 (3H, s), 3.82 (1H, dd, J=18Hz and 4Hz), 4.21 (1H, dd, J=18Hz and 5Hz), 5.49 (2H, s), 6.71 (1H, d, J=8Hz), 6.88 (1H, t, J=8Hz), 7.16 (1H, br s), 7.21-7.56 (7H), 7.71-7.87 (2H)

Example 47
3-Chloro-8-[2,6-dichloro-3-[N-(N'-phenylthioureaidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine was obtained according to a similar manner to that of Example 46.
mp : 135-137°C
NMR (CDCl₃, δ) : 2.43 (3H, s), 3.22 (3H, s), 3.85 (1H, dd, J=16Hz and 4Hz), 4.22 (1H, dd, J=16Hz and 4Hz), 5.51 (2H, s), 6.70 (1H, d, J=8Hz), 6.87 (1H, dd, J=8Hz and 7Hz), 7.16 (1H, t like), 7.21-7.54 (6H), 7.68-7.80 (2H)

Example 48
3-Bromo-8-[3-[N-(tert-butoxycarbonyl-L-prolylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine (67 mg) was dissolved in 4N solution of hydrogen chloride in ethyl acetate (3 ml). The solution was evaporated in vacuo to give 3-bromo-8-[2,6-dichloro-
3-[N-(L-prolylglycyl)-N-methylamino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine dihydrochloride (64 mg) as colorless glass.

NMR (CDCl$_3$-CD$_3$OD, δ): 2.00-2.20 (4H), 2.56 (3H, s), 3.26 (2.5H, s), 3.31-3.97 (4.5H), 4.38 (1H, t-like), 5.70 (2H, s), 7.41-7.71 (4H), 8.23 (1H, d, J=8Hz)

Example 49

3-Bromo-8-[2,6-dichloro-3-[N-(D-prolylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride was obtained according to a similar manner to that of Example 48.

NMR (CDCl$_3$-CD$_3$OD, δ): 2.00-2.20 (4H), 2.54 (3H, s), 3.26 (2.5H, s), 3.31-3.95 (4.5H), 4.38 (1H, t-like), 5.72 (2H, s), 7.49-7.71 (4H), 8.29 (1H, d, J=8Hz)

Example 50

A mixture of 3-bromo-8-[2,6-dichloro-3-[N-(L-prolylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo-[1,2-a]pyridine dihydrochloride (40 mg), acetic anhydride (0.01 ml), pyridine (0.03 ml) and methylene chloride (2 ml) was stirred for 2 hours at ambient temperature. The mixture was washed with water twice, dried over magnesium sulfate, and concentrated in vacuo to give 8-[3-[N-(acetyl-L-prolylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine (31 mg).

NMR (CDCl$_3$, δ): 1.82-2.37 (7H), 2.44 (3H, s), 3.24 (3H, s), 3.34-3.89 (4H), 4.58 (1H, br s), 5.48 (2H, s), 6.71 (1H, d, J=8Hz), 6.85 (1H, t, J=8Hz), 7.29-7.53 (2H), 7.77 (1H, d, J=8Hz)

Example 51

The following compounds were obtained according to a similar manner to that of Example 50.
Example 52

The following compounds were obtained according to similar manners to those of Examples 9 or 10.

(1) 8-[3-[N-(Acetyl-D-prolylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]-pyridine
NMR (CDCl₃, δ): 1.82-2.37 (7H), 2.44 (3H, s), 3.24 (3H, s), 3.34-3.89 (4H), 4.58 (1H, br s), 5.48 (2H, s), 6.71 (1H, d, J=8Hz), 6.85 (1H, t, J=8Hz), 7.29-7.53 (2H), 7.77 (1H, d, J=8Hz)

(2) 8-[3-[N-(Acetylglycylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]-pyridine
mp: 220-222°C
NMR (CDCl₃, δ): 2.04 (3H, s), 2.41 (3H, s), 3.22 (3H, s), 3.53 (1H, dd, J=17Hz and 5Hz), 3.84 (1H, dd, J=17Hz and 5Hz), 3.38-4.00 (2H), 5.51 (2H, s), 6.72 (1H, d, J=7Hz), 6.90 (1H, t, J=7Hz), 7.40 (1H, d, J=9Hz), 7.52 (1H, d, J=9Hz), 7.79 (1H, d, J=7Hz)

(1) 3-Bromo-8-[2,6-dichloro-3-[N-(N,N-dimethylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine
NMR (CDCl₃, δ): 2.27 (6H, s), 2.45 (3H, s), 2.72 (1H, d, J=15Hz), 2.91 (1H, d, J=15Hz), 3.20 (3H, s), 5.49 (1H, d, J=7Hz), 5.55 (1H, d, J=7Hz), 6.71 (1H, d, J=7Hz), 6.85 (1H, t, J=7Hz), 7.79 (1H, d, J=9Hz), 7.45 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(2) 3-Bromo-8-[2,6-dichloro-3-[N-(isopropylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 0.98-1.09 (6H), 2.43 (3H, s), 2.72 (1H), 3.00 (1H, d, J=16Hz), 3.18 (1H, d, J=16Hz), 3.22 (3H, s), 5.51 (2H, s), 6.71 (1H, d, J=7Hz), 6.85 (1H, t, J=7Hz), 7.30 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.78 (7H, d, J=7Hz)

(3) 3-Bromo-8-[2,6-dichloro-3-[N-(isopropylglycylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 1.10 (6H, d, J=6Hz), 2.43 (3H, s), 3.26 (3H, s), 3.29 (2H, s), 3.56 (1H, dd, J=17Hz and 5Hz), 3.82 (1H, dd, J=17Hz and 5Hz), 5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.76 (1H, d, J=7Hz), 8.08 (1H, br t, J=5Hz)

(4) 3-Bromo-8-[2,6-dichloro-3-[N-(N,N-dimethylglycyl)glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 2.32 (6H, s), 2.43 (3H, s), 2.96 (2H, s), 3.25 (3H, s), 3.55 (1H, dd, J=18Hz and 4Hz), 3.85 (1H, dd, J=18Hz and 4Hz), 5.00 (2H, s), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 7.89 (1H, br t, J=4Hz)

(5) 3-Bromo-8-[2,6-dichloro-3-[N-(benzylglycylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 2.43 (3H, s), 3.28 (3H, s), 3.31 (2H, s), 3.58 (1H, dd, J=17Hz and 5Hz), 3.80 (2H, s), 3.84 (1H, dd, J=17Hz and 5Hz), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.19-7.40 (7H), 7.48 (1H, d, J=9Hz), 7.27 (1H, d, J=7Hz), 7.97 (1H, br t, J=5Hz)
Example 53

A mixture of 3-bromo-8-[2,6-dichloro-3-[N-(bromo-acetylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (150 mg) and 30% solution of methylamine in methanol (2 ml) was stirred for 1 hour at ambient temperature. The mixture was concentrated in vacuo, and the residue was purified by silica gel column chromatography (methylene chloride:methanol = 10:1, V/V) to give 3-bromo-8-[2,6-dichloro-3-[N-sarcosylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (103 mg).

NMR (CDCl₃, δ) : 2.45 (3H, s), 2.48 (3H, s), 3.26 (5H, s), 3.59 (1H, dd, J=17Hz and 5Hz), 3.83 (1H, dd, J=17.5Hz), 5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 7.89 (1H, br t, J=5Hz)

Example 54

The following compounds were obtained according to a similar manner to that of Example 53.

(i) 3-Bromo-8-[2,6-dichloro-3-[N-(ethylglycylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine

NMR (CDCl₃, δ) : 1.12 (3H, t, J=6Hz), 2.43 (3H, s), 2.69 (2H, g, J=6Hz), 3.26 (3H, s), 3.30 (2H, s), 3.58 (1H, dd, J=17Hz and 5Hz), 3.82 (1H, dd, J=17Hz and 5Hz), 5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 7.99 (1H, br t, J=5Hz)

(ii) 3-Bromo-8-[2,6-dichloro-3-[N-(phenylglycyl)-glycyl]-N-methylamino]benzyloxy]-2-methylimidazo-
[1,2-a]pyridine
NMR (CDCl$_3$, $\delta$): 2.42 (3H, s), 3.21 (3H, s), 3.60 (1H, dd, $J$=18Hz and 5Hz), 3.80 (2H, s), 3.83 (1H, dd, $J$=18Hz and 5Hz), 4.39 (1H, br s), 5.49 (2H, s), 6.60 (2H, d, $J$=6Hz), 6.68-6.90 (3H), 7.12-7.41 (4H), 7.48 (1H, d, $J$=9Hz), 7.78 (1H, d, $J$=7Hz)

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Example 55
To a mixture of 3-bromo-8-(2,6-dichloro-4-benzoylbenzyloxy)-2-methylimidazo[1,2-a]pyridine (80 mg), sodium borohydride (18.5 mg) and ethanol (2 ml) was stirred at ambient temperature for 1 hour. The reaction mixture was partitioned between dichloromethane and water and the aqueous layer was extracted with dichloromethane twice. The combined organic layers were washed with brine, dried over magnesium sulfate and evaporated in vacuo. The residue was crystallized from diethyl ether to give 3-bromo-8-[2,6-dichloro-4-(a-hydroxybenzyl)benzyloxy]-2-methylimidazo[1,2-a]pyridine (73 mg) as crystals.

mp: 178-179°C

NMR (CDCl$_3$, $\delta$): 2.40 (3H, s), 2.73 (1H, br s), 5.45 (2H, dd, $J$=10Hz and 9Hz), 6.28 (1H, s), 6.69 (1H, d, $J$=7Hz), 6.80 (1H, t, $J$=7Hz), 7.23-7.48 (6H), 7.60-7.80 (2H)
Example 56

3-Bromo-8-[2,6-dichloro-3-[N-(mesylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Example 19.

NMR (CDCl₃, δ): 2.45 (3H, s), 2.97 (2.7Hz, s), 3.02 (0.3H, s), 3.27 (2.7H, s), 3.30 (0.3H, s), 3.50 (1H, dd, J=16Hz and 5Hz), 3.67 (1H, dd, J=16Hz and 5Hz), 5.18 (1H, t like), 5.51 (2H, s), 6.71 (1H, d, J=8Hz), 6.87 (1H, t, J=8Hz), 7.32 (1H, d, J=8Hz), 7.51 (1H, d, J=8Hz), 7.78 (1H, d, J=8Hz)

Example 57

To a mixture of 3-bromo-8-(2,6-dibromo-4-methoxycarbonylbenzyloxy)-2-methylimidazo[1,2-a]pyridine (550 mg), methanol (10 ml) and tetrahydrofuran (5 ml) was added 1N aqueous solution of sodium hydroxide (1.135 ml), and the mixture was stirred for 1 hour at 60°C. The reaction mixture was adjusted pH 4 with 1N hydrochloric acid, and water was added thereto. The precipitate was collected by filtration to give 3-bromo-8-(2,6-dibromo-4-carboxybenzyloxy)-2-methylimidazo[1,2-a]pyridine (518 mg).

mp: 241-242°C

NMR (DMSO-d₆, δ): 2.41 (3H, s), 5.62 (2H, s), 7.54 (1H, t, J=7Hz), 7.78 (1H, d, J=7Hz), 8.19 (2H, s), 8.32 (1H, d, J=7Hz)

Example 58

The following compounds were obtained according to a similar manner to that of Example 57.

(1) 3-Chloro-8-(2,6-dichlorobenzyloxy)imidazo[1,2-a]pyridine-2-carboxylic acid

mp: 212-213°C
NMR (CDCl₃ + CD₃OD, δ):  5.50 (2H, s), 6.82 (1H, d, J=7Hz), 7.02 (1H, t, J=7Hz), 7.22-7.48 (3H), 7.86 (1H, d, J=7Hz)

3-Bromo-8-[2,6-dichloro-3-[N-(3-carboxypropionyl-glycyl)-N-methylamino]benzyloxy]-2-methylimidazo-[1,2-a]pyridine
NMR (CDCl₃-CD₃OD, δ):  2.30 (3H, s), 2.44 (4H, s), 3.12 (3H, s), 3.68-3.86 (2H), 5.40 (2H, s), 6.69 (1H, d, J=8Hz), 6.81 (1H, dd, J=8Hz and 6Hz), 7.46 (2H, s), 7.69 (1H, d, J=6Hz)

Example 59
To a mixture of 3-bromo-8-(2,6-dibromo-4-carboxy-benzyloxy)-2-methylimidazo[1,2-a]pyridine (100 mg), methylene chloride (2 ml) and N,N-dimethylformamide (1 drop) was added oxalyl chloride (49 mg), and the mixture was stirred for 30 minutes and evaporated. The residue was dissolved in methylene chloride, triethylamine (0.5 ml) and 1-methylpiperazine (23 mg) were added thereto, and the mixture was stirred for 1 hour. Water was added thereto, the mixture was extracted with methylene chloride 3 times. The combined organic layer was washed with brine, dried over magnesium sulfate, and concentrated in vacuo. The residue was purified by silica gel column chromatography (5% solution of methanol in methylene chloride) to give 3-bromo-8-[2,6-dibromo-4-(N-methylpiperazinylcarbonyl)benzyloxy]-2-methylimidazo-[1,2-a]pyridine (102 mg).
mp: 188-190°C
NMR (CDCl₃, δ):  2.45 (3H, s), 2.48 (3H, s), 2.49-2.79 (4H), 3.56-4.02 (4H), 5.50 (2H, s), 6.70 (1H, d, J=7Hz), 6.85 (1H, t, J=7Hz), 7.61 (2H, s), 7.77 (1H, d, J=7Hz)
Example 60

To a suspension of 3-bromo-8-(2,6-dichloro-3-nitrobenzyloxy)-2-methylimidazo[1,2-a]pyridine (200 mg) in ethanol (2 ml) was added hydrogen chloride in ethanol (3.5 Mol solution, 1 ml). The solution was concentrated to half volume under reduced pressure. The separated precipitates were collected by filtration and washed with ethanol to give 3-bromo-8-(2,6-dichloro-3-nitrobenzyloxy)-2-methylimidazo[1,2-a]pyridine hydrochloride (175 mg) as an off-white solid.

mp : 195-197°C
NMR (DMSO-δ6, δ) : 2.39 (3H, s), 5.60 (2H, s), 7.66 (1H, t, J=7.5Hz), 7.56 (1H, d, J=7.5Hz), 7.42 (1H, d, J=7.5Hz), 8.25 (2H, d, J=7.5Hz)

Example 61

The following compounds were obtained according to a similar manner to that of Example 60.

(1) 3-Bromo-8-[2,6-dichloro-3-(N-acetyl-N-methylamino)-benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

mp : 148-150°C
NMR (CDCl3 + CD3OD, δ) : 1.88 (3H, s), 2.68 (3H, s), 3.22 (3H, s), 5.62 (1H, d, J=9Hz), 5.70 (1H, d, J=9Hz), 7.32-7.61 (4H), 8.09 (1H, d, J=6Hz)

(2) 3-Chloro-8-[2,6-dichloro-3-(N-acetyl-N-methylamino)-benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

mp : 145-149°C
NMR (CDCl3 + CD3OD, δ) : 1.89 (3H, s), 2.70 (3H, s), 3.22 (3H, s), 5.61 (1H, d, J=10Hz), 5.70 (1H, d, J=10Hz), 7.25-7.59 (4H), 8.01 (1H, d, J=7Hz)
(3) 3-Bromo-8-[2,6-dichloro-3-(N-propionyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
mp : 146-148°C
NMR (CDCl₃, δ) : 1.08 (3H, t, J=7Hz), 2.09 (2H, q, J=7Hz), 2.74 (3H, s), 3.26 (3H, s), 5.66 (2H, br t, J=12Hz), 7.16-7.58 (4H), 8.00 (1H)

(4) 3-Bromo-8-[2,4,6-trichloro-3-(N-acetyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
mp : 116-120°C
NMR (CDCl₃, δ) : 1.90 (2.6H, s), 2.29 (0.4H, s),
2.70 (2.6H, s), 2.72 (0.4H, s), 3.22 (2.6H, s),
3.43 (0.4H, s), 5.50-5.70 (2H), 7.18 (1H, d, J=8Hz), 7.30 (1H, t, J=8Hz), 7.52 (0.13H, s), 7.62 (0.87H, s), 7.98 (1H, d, J=8Hz)

(5) 8-[3-[N-(Acetylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine hydrochloride
mp : 175-176°C
NMR (DMSO-d₆, δ) : 1.83, 2.40, 3.12 (each 3H, q),
3.37, 3.67 (each 1H, dd, J=16Hz and 5Hz), . . . 59
(2H, s), 7.31-7.83 (4H), 8.10 (1H, t, J=5Hz),
8.25 (1H, d, J=7Hz)

(6) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-(propionylglycyl)amino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine hydrochloride
NMR (CDCl₃, δ) : 1.13 (3H, t, J=6Hz), 2.23 (2H, q, J=6Hz),
2.71 (3H, s), 3.33 (2.8H, s), 3.47
(0.2H, s), 3.66 (1H, dd, J=16Hz and 4Hz), 3.80
(1H, dd, J=16Hz and 4Hz), 5.61 (1H, d, J=10Hz),
5.68 (1H, d, J=10Hz), 6.74 (1H, t like),
7.20-7.44 (3H), 7.50 (1H, d, J=8Hz), 8.00 (1H, d, J=7Hz)

(7) 3-Bromo-8-[3-[N-(butyrylglycyl)-N-methylamino]-
2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (DMSO-d$_6$, $\delta$) : 0.84 (3H, t, J=7Hz), 1.49 (2H, m), 2.10 (2H, t, J=7Hz), 2.40 (3H, s), 3.13 (2.4H, s), 3.30 (0.6H, s), 3.36 (1H, dd, J=16Hz and 5Hz), 3.68 (1H, dd, J=16Hz and 5Hz), 5.60 (2H, s), 7.37-7.86 (4H), 8.03 (1H, t like), 8.28 (1H, d, J=6Hz)

(8) 3-Bromo-8-[3-[N-(isobutyrylglycyl)-N-methylamino]-
2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (DMSO-d$_6$, $\delta$) : 0.97 (6H, d, J=6Hz), 2.40 (3H, s), 2.45 (1H, m), 3.12 (2.3H, s), 3.28 (0.7H, s), 3.34 (1H, dd, J=16Hz and 5Hz), 3.67 (1H, dd, J=16Hz and 5Hz), 5.58 (2H, s), 7.36-7.84 (4H), 7.99 (1H, t like), 8.27 (1H, d, J=7Hz)

(9) 3-Bromo-8-[2,6-dichloro-3-[N-(cyclopropylcarbonyl-
glycyl)-N-methylamino]benzyloxy]-2-methylimidazo-
[1,2-a]pyridine hydrochloride

NMR (DMSO-d$_6$, $\delta$) : 0.53-0.72 (4H), 1.68 (1H, m), 2.39 (3H, s), 3.12 (2.5H, s), 3.28 (0.5H, s), 3.40 (1H, dd, J=16Hz and 5Hz), 3.70 (1H, dd, J=16Hz and 5Hz), 5.59 (2H, s), 7.33-7.89 (4H), 8.26 (1H, d, J=7Hz), 8.34 (1H, t, J=6Hz)

(10) 3-Bromo-8-[2,6-dichloro-3-[N-(trifluoroacetyl-
glycyl)-N-methylamino]benzyloxy]-2-methylimidazo-
[1,2-a]pyridine hydrochloride

mp : 194-195°C
NMR (DMSO-d$_6$, δ): 2.39 (3H, s), 3.14 (3H, s), 3.51 (1H, dd, J=17Hz and 5Hz), 3.78 (1H, dd, J=17Hz and 5Hz), 5.58 (2H, s), 7.39 (1H, t, J=6Hz), 7.59 (1H), 7.81 (2H, s), 8.25 (1H, d, J=6Hz), 9.71 (1H, t, J=6Hz)

(11) 8-[3-[N-(Benzoylglucycl)-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]-pyridine hydrochloride

mp: 138°C (dec.)

NMR (DMSO-d$_6$, δ): 2.39 (3H, s), 3.15 (3H, s), 3.58 (1H, dd, J=16Hz and 6Hz), 3.89 (1H, dd, J=16Hz and 6Hz), 5.59 (2H, s), 7.34-7.64 (5H), 7.77-7.93 (4H), 8.28 (1H, d, J=6Hz), 8.72 (1H, t, J=6Hz)

(12) 3-Bromo-8-[2,6-dichloro-3-[N-(cyclohexylcarbonylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (DMSO-d$_6$, δ): 1.02-1.79 (10H), 2.18 (1H, m), 2.39 (3H, s), 3.11 (2.6H, s), 3.27 (0.4H, s), 3.31 (1H, dd, J=16Hz and 5Hz), 3.65 (1H, dd, J=16Hz and 5Hz), 5.58 (2H, s), 7.31-7.82 (4H), 7.92 (1H, t, J=5Hz), 8.26 (1H, d, J=7Hz)

(13) 3-Bromo-8-[2,6-dichloro-3-[N-(phenylacetylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (DMSO-d$_6$, δ): 2.38 (3H, s), 3.13 (2.5H, s), 3.27 (0.5H, s), 3.30-3.76 (4H), 5.56 (2H, s), 7.14-7.85 (9H), 8.24 (1H, d, J=7Hz), 8.32 (1H, t, J=5Hz)

(14) 8-[3-[N-(Acetoxyacetylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]-
pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$): 2.21 (3H, s), 2.61 (3H, s),
3.25 (3H, s), 3.59-3.88 (2H), 4.09 (1H, br s),
4.59 (1H, br s), 5.69 (2H, br s), 7.35-7.68 (4H), 8.11 (1H, br s)

(15) 3-Bromo-8-[2,6-dichloro-3-[N-(glycoloylglycyl)-N-
methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
hydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$): 2.59 (3H, s), 3.28 (3H, s),
3.78 (2H, s), 4.09 (2H, s), 5.54-5.81 (2H),
7.47-7.68 (4H), 8.14 (1H, br s)

(16) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-(methoxyacetylglycyl)-
acetyl-N-methylamino]benzyloxy]-2-methylimidazo-
[1,2-a]pyridine hydrochloride
mp: 175-177°C
NMR (DMSO-d$_6$, $\delta$): 2.40 (3H, s), 3.12 (3H, s), 3.32 (3H, s),
3.42 (1H, dd, J=17Hz and 5Hz), 3.70 (1H, dd, J=17Hz and 5Hz), 3.82 (2H, s), 5.59 (2H, s), 7.34-7.79 (5H), 8.26 (1H, d, J=7Hz)

(17) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-(phthalimido-
acetylglycyl)amino]benzyloxy]-2-methylimidazo-
[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$): 2.57 (3H, s), 3.26 (3H, s),
3.65 (1H, d, J=15Hz), 3.78 (1H, d, J=15Hz), 4.32 (1H, d, J=15Hz), 4.46 (1H, d, J=15Hz), 5.66 (2H, s), 7.35-7.64 (4H), 7.70-7.92 (4H), 8.08 (1H, d, J=5Hz)

(18) 8-[3-[N-(Acetylglycylglycyl)-N-methylamino]-2,6-
dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]-pyridine hydrochloride
NMR (CDCl$_3$ + CD$_3$OD, $\delta$): 2.01 (3H, s), 2.52 (3H, s),
3.22 (3H, s), 3.56-3.70 (2H), 3.84 (2H, s), 5.70
(2H, br s), 7.51-7.72 (5H), 8.29 (1H, d, J=6Hz)

(19) 8-[3-[N-(Acetylglycyl)-N-ethylamino]-2,6-dichloro-
benzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine
hydrochloride
NMR (CDCl₃, δ): 1.20 (3H, t, J=7Hz), 2.00 (3H, s),
2.71 (3H, s), 3.43-3.88 (3H), 4.14 (1H), 5.68
(2H, s), 6.88 (1H, br s), 7.18-7.58 (3H), 8.00
(1H, br s)

(20) 8-[3-[N-(Acetyl-DL-alanyl)-N-methylamino]-2,6-
dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]-pyridine
hydrochloride
NMR (CDCl₃, δ): 1.11 (3H, d, J=6Hz), 1.94 (3H, s),
2.68 (3H, s), 3.30 (3H, s), 4.36 (1H, m),
5.53-5.76 (2H), 6.60-6.80 (1H), 7.13-7.60 (4H),
7.93-8.06 (1H)

(21) 8-[3-[N-(Acetyl-β-alanyl)-N-methylamino]-2,6-
dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]-pyridine
hydrochloride
mp: 188-190°C
NMR (CDCl₃ + CD₃OD, δ): 2.01 (3H, s), 2.12-2.72
(2H), 3.63 (3H, s), 3.20 (3H, s), 3.30-3.68
(2H), 5.58 (1H, d, J=10Hz), 5.73 (1H, d,
J=10Hz), 7.32-7.61 (4H), 8.11 (1H, d, J=5Hz)

(22) 3-Bromo-8-[2,6-dichloro-3-[N-(N,N-dimethylglycyl)]-
N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine
dihydrochloride
mp: 162-164°C
NMR (CDCl₃ + CD₃OD, δ): 2.66 (3H, s), 3.00-3.19
(6H), 3.28 (3H, s), 4.01 (1H, d, J=17Hz), 4.36
(1H, d, J=17Hz), 5.49 (1H, d, J=9Hz), 5.75 (1H,
d, J=9Hz), 7.32-7.56 (2H), 7.61 (1H, d, J=9Hz), 7.75 (1H, d, J=9Hz), 8.09 (1H, d, J=5Hz)

(23) 3-Bromo-8-[2,6-dichloro-3-[N-(isopropylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃ + CD₃OD, δ) : 1.41 (6H, br s), 2.67 (3H, br s), 3.29 (3H, br s), 3.60-3.98 (3H), 5.51 (1H, br s), 5.76 (1H, br s), 7.32-8.14 (5H)

(24) 3-Bromo-8-[2,6-dichloro-3-[N-(methoxycarbonyl-glycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
mp : 176-177°C
NMR (CDCl₃ + CD₃OD, δ) : 2.68 (3H, s), 3.29 (3H, s), 3.50-3.79 (5H), 5.68 (2H, s), 7.31-7.60 (4H), 8.02 (1H, d, J=6Hz)

(25) 3-Bromo-8-[2,6-dichloro-3-[N-(mesylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
mp : 174-176°C
NMR (CDCl₃-CD₃OD, δ) : 2.63 (3H, s), 3.03 (3H, s), 3.27 (3H, s), 3.60 (1H, d, J=16Hz), 3.70 (1H, d, J=16Hz), 5.62 (1H, d, J=12Hz), 5.70 (1H, d, J=12Hz), 7.33-7.51 (3H), 7.58 (1H, d, J=8Hz), 8.06 (1H, d, J=6Hz)

(26) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-(ureidoacetyl)-amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (DMSO-d₆, δ) : 2.40 (3H, s), 3.12 (3H, s), 3.39 (1H, d, J=17Hz), 3.61 (1H, d, J=17Hz), 5.57 (1H, d, J=8Hz), 5.67 (1H, d, J=8Hz), 7.49 (1H, t, J=7Hz), 7.69 (1H, d, J=7Hz), 7.81 (2H, s), 8.31 (1H, d, J=7Hz)
(27) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-(N'--methylureidoacetyl)amino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine hydrochloride
mp : 153-155°C
NMR (CDCl₃, δ) : 2.63 (3H, s), 2.74 (3H, s), 3.23 (3H, s), 3.85 (1H, d, J=16Hz), 3.98 (1H, d, J=16Hz), 5.57 (1H, d, J=10Hz), 5.67 (1H, d, J=10Hz), 7.30-7.60 (4H), 8.04 (1H, d, J=7Hz)

(28) 3-Bromo-8-[2,6-dichloro-3-[N-(N'-ethylureidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine hydrochloride
its dihydrochloride
mp : 173-175°C
NMR (DMSO-d₆, δ) : 0.96 (3H, t, J=7Hz), 2.41 (3H, s), 2.98 (2H, q, J=7Hz), 3.11 (3H, s), 3.30 (1H, d, J=17Hz), 3.61 (1H, d, J=17Hz), 5.56 (1H, d, J=9Hz), 5.67 (1H, d, J=9Hz), 7.50 (1H, t, J=7Hz), 7.70 (1H, d, J=7Hz), 7.81 (2H, s), 8.31 (1H, d, J=7Hz)

(29) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-(N'-phenylureidoacetyl)amino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine hydrochloride
NMR (DMSO-d₆, δ) : 2.38 (3H, s), 3.16 (2.5H, s), 3.29 (0.5H, s), 3.42 (1H, d, J=16Hz), 3.70 (1H, d, J=16Hz), 5.58 (2H, s), 6.45 (1H, br s), 6.89 (1H, t, J=7Hz), 7.13-7.90 (8H), 8.26 (1H, d, J=7Hz), 8.97 (1H, s)

(30) 3-Bromo-8-[2,6-dichloro-3-[N-(4-pentenoylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.20-2.32 (4H), 2.46 (3H, s), 3.17 (3H, s), 3.56 (1H, s), 3.59 (1H, s), 4.83-5.06 (2H), 5.56-5.84 (3H), 7.42-7.62 (4H), 8.19 (1H, d, J=6Hz)
(31) 3-Bromo-8-[2,6-dichloro-3-[N-(valerylglucyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$, $\delta$) : 0.90 (3H, t, $J=7$Hz), 1.32 (2H, m),
1.60 (2H, m), 2.22 (2H, t, $J=7$Hz), 2.72 (3H, s),
3.33 (3H, s), 3.58-3.87 (2H), 5.60 (1H, d,
$J=11$Hz), 5.68 (1H, d, $J=11$Hz), 6.76 (1H, br s),
7.21-7.45 (3H), 7.50 (1H, d, $J=9$Hz), 8.00 (1H,
d, $J=6$Hz)

(32) 3-Bromo-8-[2,6-dichloro-3-[N-(isovalerylglucyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$, $\delta$) : 0.93 (6H, d, $J=6$Hz), 2.00-2.20
(3H), 2.73 (3H, s), 3.33 (3H, s), 3.60-3.88
(2H), 5.65 (2H, s), 6.77 (1H, br s), 7.20-7.44
(3H), 7.50 (1H, d, $J=9$Hz), 8.00 (1H, d, $J=6$Hz)

(33) 3-Bromo-8-[2,6-dichloro-3-[N-(2-pyridylacetylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 2.53 (3H, s), 3.25 (3H, s),
3.35 (2H, s), 3.62 (1H, d, $J=17$Hz), 3.82 (1H, d,
$J=17$Hz), 5.72 (2H, s), 7.48-7.72 (4H), 7.87-8.03
(2H), 8.29 (1H, d, $J=6$Hz), 8.48 (1H, t, $J=7$Hz),
8.75 (1H, d, $J=6$Hz)

(34) 3-Bromo-8-[2,6-dichloro-3-[N-(3-pyridylacetylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 2.56 (3H, s), 3.25 (3H, s),
3.60 (1H, d, $J=17$Hz), 3.80 (1H, d, $J=17$Hz), 3.93
(2H, s), 5.72 (2H, s), 7.52-7.73 (4H), 8.07 (1H,
dd, $J=7$Hz and 5Hz), 8.29 (1H, d, $J=7$Hz), 8.60
(1H, d, $J=7$Hz), 8.75 (1H, d, $J=5$Hz), 8.89 (1H, br s)
(35) 3-Bromo-8-[2,6-dichloro-3-[N-[(3-ethoxycarbonylpropionyl)glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃, δ): 1.23 (3H, t, J=7Hz), 2.44-2.66 (4H), 2.70 (3H, s), 3.30 (3H, s), 3.63 (1H, dd, J=15Hz and 4Hz), 3.79 (1H, dd, J=15Hz and 4Hz), 4.10 (2H, q, J=7Hz), 5.58 (1H, d, J=11Hz), 5.67 (1H, d, J=11Hz), 6.89 (1H, br s), 7.09-7.43 (3H), 7.51 (1H, d, J=9Hz), 7.96 (1H, d, J=5Hz)

(36) 3-Bromo-8-[2,6-dichloro-3-[N-(sarcosylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.28 (3H, s), 2.46 (3H, s), 2.98 (3H, s), 3.34 (1H, d, J=16Hz), 3.52 (2H, s), 3.59 (1H, d, J=16Hz), 5.43 (2H, s), 7.22-7.42 (4H), 8.00 (1H, d, J=6Hz)

(37) 3-Bromo-8-[2,6-dichloro-3-[N-[(ethylglycyl)glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ): 1.38 (3H, t, J=6Hz), 2.56 (3H, s), 3.10 (2H, q, J=6Hz), 3.27 (3H, s), 3.61 (1H, d, J=16Hz), 3.82 (2H, s), 3.88 (1H, d, J=16Hz), 5.72 (2H, s), 7.50-7.74 (4H), 8.29 (1H, d, J=6Hz)

(38) 3-Bromo-8-[2,6-dichloro-3-[N-[(N-phenylglycyl)glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.59 (3H, s), 3.26 (3H, s), 3.61-3.78 (2H), 4.00 (2H, s), 5.71 (2H, br s), 7.02-7.20 (3H), 7.30-7.41 (2H), 7.53-7.70 (4H), 8.26 (1H, d, J=6Hz)
(39) 3-Bromo-8-[2,6-dichloro-3-[N-(morpholinoacetyl-glycyl)-N-methylamino]benzyloxy]-2-methylimidazo-[1,2-a]pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 2.53 (3H, s), 3.21 (3H, s), 3.30-3.48 (4H), 3.59 (1H, d, $J=16$Hz), 3.81 (1H, d, $J=16$Hz), 3.85-4.07 (6H), 5.69 (2H, s), 7.50-7.69 (4H), 8.24 (1H, d, $J=6$Hz)

(40) 3-Bromo-8-[2,6-dichloro-3-[N-(isopropylglycylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 1.38 (6H, d, $J=6$Hz), 2.56 (3H, s), 3.28 (3H, s), 3.81-3.48 (3H), 3.82 (2H, s), 5.72 (2H, s), 7.49-7.73 (4H), 8.30 (1H, d, $J=6$Hz)

(41) 3-Bromo-8-[2,6-dichloro-3-[N-(N,N-dimethylglycyl)-glycyl]-N-methylamino]benzyloxy]-2-methylimidazo-[1,2-a]pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 2.29 (3H, s), 2.68 (6H, s), 2.97 (3H, s), 3.35 (1H, d, $J=17$Hz), 3.58 (1H, d, $J=17$Hz), 3.71 (2H, s), 5.42 (2H, s), 7.25-7.44 (4H), 7.99 (1H, d, $J=6$Hz)

(42) 3-Bromo-8-[2,6-dichloro-3-[N-(benzylglycylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 2.58 (3H, s), 3.25 (3H, s), 3.62 (1H, d, $J=17$Hz), 3.70-3.92 (3H), 4.22 (2H, s), 5.72 (2H, s), 7.40-7.75 (9H), 8.30 (1H, d, $J=6$Hz)

(43) 3-Bromo-8-[2,6-dichloro-3-[N-(N'-cyclohexylureido-acetyl)-N-methylamino]benzyloxy]-2-methylimidazo-[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 1.03-1.92 (10H), 2.58 (3H, s), 3.27 (3H, s), 3.44 (1H, m), 3.60 (1H, d, J=17Hz), 3.71 (1H, d, J=17Hz), 5.68 (1H, d, J=10Hz), 5.79 (1H, d, J=10Hz), 7.53-7.72 (4H), 8.79 (1H, d, J=6Hz)

(44) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[(α-naphthyl)ureido]acetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 1.99 (3H, s), 3.00 (3H, s), 3.45 (1H, d, J=17Hz), 3.62 (1H, d, J=17Hz), 5.40 (2H, s), 7.02-7.50 (10H), 7.69 (1H, d, J=9Hz), 7.91 (1H, d, J=5Hz)

(45) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-benzyureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 2.48 (3H, s), 3.28 (3H, s), 3.70 (2H, s), 4.30 (2H, s), 5.68 (1H, d, J=10Hz), 6.79 (1H, d, J=10Hz), 7.16-7.35 (5H), 7.54-7.72 (4H), 8.29 (1H, d, J=6Hz)

(46) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-(m-tolyl)ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 2.30 (3H, s), 2.52 (3H, s), 3.29 (3H, s), 3.72 (2H, s), 5.67 (1H, d, J=10Hz), 5.79 (1H, d, J=10Hz), 6.82 (1H, m), 7.09-7.19 (3H), 7.51-7.71 (4H), 8.26 (1H, d, J=5Hz)

(47) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-(p-tolyl)ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 2.18 (3H, s), 2.42 (3H, s),
3.19 (3H, s), 3.62 (2H, s), 5.59 (1H, br d, J=10Hz), 5.69 (1H, br d, J=10Hz), 6.93 (2H, d, J=9Hz), 7.10 (2H, d, J=9Hz), 7.43-7.62 (4H), 8.18 (1H, d, J=6Hz)

(48) 3-Bromo-8-[2,6-dichloro-3-[N-[N'(4-methoxyphenyl)-ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.52 (3H, s), 3.29 (3H, s), 3.72 (2H, s), 3.79 (3H, s), 5.69 (1H, d, J=10Hz), 5.79 (1H, d, J=10Hz), 6.80 (2H, d, J=10Hz), 7.22 (2H, d, J=10Hz), 7.52-7.73 (4H), 8.29 (1H, d, J=6Hz)

(49) 3-Bromo-8-[2,6-dichloro-3-[N-[N'(4-trifluoromethylphenyl)ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.54 (3H, s), 3.29 (3H, s), 3.66 (1H, d, J=17Hz), 3.76 (1H, d, J=17Hz), 5.68 (1H, d, J=10Hz), 5.76 (1H, d, J=10Hz), 7.49-7.71 (8H), 8.28 (1H, d, J=6Hz)

(50) 3-Bromo-8-[2,6-dichloro-3-[N-[N'(4-fluorophenyl)ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.52 (3H, s), 3.28 (3H, s), 3.68 (1H, d, J=16Hz), 3.78 (1H, d, J=16Hz), 5.70 (1H, d, J=10Hz), 5.79 (1H, d, J=10Hz), 6.89-7.02 (2H), 7.22-7.36 (2H), 7.55-7.75 (4H), 8.80 (1H, d, J=5Hz)

(51) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-n-propylureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
mp : 168-171°C
NMR (CDCl$_3$-CD$_3$OD, δ) : 0.92 (3H, t, J=7Hz),
1.40-1.60 (2H), 2.55 (3H, s), 3.08 (2H, t, J=6Hz), 3.22 (3H, s), 3.58 (1H, d, J=17Hz), 3.71 (1H, d, J=17Hz), 5.69 (1H, d, J=10Hz), 5.79 (1H, d, J=10Hz), 7.53-7.77 (4H), 8.31 (1H, d, J=6Hz)

(52) 3-Bromo-8-[2,6-dichloro-3-[N-(N'-isopropylureidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ) : 1.12 (6H, d, J=6Hz), 2.56 (3H, s), 3.25 (3H, s), 3.57-3.32 (3H), 5.65 (1H, d, J=10Hz), 5.78 (1H, d, J=10Hz), 7.51-7.71 (4H), 8.28 (1H, d, J=6Hz)

(53) 3-Bromo-8-[2,6-dichloro-3-[N-(N'-allylureidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ) : 2.51 (3H, s), 3.20 (3H, s), 3.57-3.79 (4H), 5.00-5.21 (2H), 5.57-2.90 (3H), 7.50-7.68 (4H), 8.22 (1H, d, J=6Hz)

(54) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-(3-pyridyl)ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ) : 2.48 (3H, s), 3.19 (3H, s), 3.52 (1H, d, J=17Hz), 3.73 (1H, d, J=17Hz), 5.62 (2H, s), 7.44-7.62 (4H), 7.81 (1H, m), 8.15-8.29 (3H), 9.12 (1H, s)

(55) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-(4-pyridyl)ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ) : 2.55 (3H, s), 3.28 (3H, s), 3.62 (1H, d, J=17Hz), 3.84 (1H, d, J=17Hz), 5.72 (2H, s), 7.50-7.72 (4H), 7.98 (2H, d, J=6Hz),
8.26 (1H, d, J=6Hz), 8.39 (2H, d, J=6Hz)

(56) 3-Bromo-8-[2,6-dichloro-3-[N-(N'-phenylthioureidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ):  2.58 (3H, s), 3.26 (3H, s),
  3.92 (1H, d, J=17Hz), 4.18 (1H, d, J=17Hz), 5.74
  (2H, s), 7.20-7.72 (9H), 8.27 (1H, d, J=6Hz)

(57) 3-Chloro-8-[2,6-dichloro-3-[N-(N'-phenylureidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
mp: 157-162°C
NMR (DMSO-d₆, δ):  2.40 (3H, s), 3.16 (3H, s), 3.42
  (1H, d, J=16Hz), 3.70 (1H, d, J=16Hz), 5.60 (2H,
  s), 6.48 (1H, br s), 6.89 (1H, t, J=8Hz),
  7.12-7.90 (8H), 8.29 (1H, d, J=6Hz), 9.00 (1H,
  s)

(58) 3-Chloro-8-[2,6-dichloro-3-[N-(N'-cyclohexylureidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
mp: 170-172°C
NMR (DMSO-d₆, δ):  0.91-1.80 (10H), 2.43 (3H, s),
  3.12 (3H, s), 3.20-3.40 (2H), 3.62 (1H, d,
  J=16Hz), 5.58 (1H, d, J=10Hz), 5.66 (1H, dd,
  J=10Hz), 7.45-7.85 (4H), 8.37 (1H, d, J=6Hz)

(59) 3-Chloro-8-[2,6-dichloro-3-[N-(N'-ethylureidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ):  1.10 (3H, t, J=6Hz), 2.57
  (3H, s), 3.15 (2H, q, J=6Hz), 3.27 (3H, s), 3.66
  (2H, s), 5.68 (1H, d, J=10Hz), 5.79 (1H, d,
  J=10Hz), 7.54-7.72 (4H), 8.29 (1H, d, J=6Hz)
(60) 3-Chloro-8-[2,6-dichloro-3-[N-(phenylacetylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine hydrochloride
NMR (DMSO-d$_6$, δ) : 2.39 (3H, s), 3.12 (2.2H, s), 3.27 (0.8H, s), 3.48 (1H, dd, J=16Hz and 6Hz), 3.47 (2H, s), 3.69 (1H, dd, J=16Hz and 6Hz), 5.58 (2H, s), 7.14-7.84 (9H), 8.23-8.37 (2H)

(61) 3-Chloro-8-[2,6-dichloro-3-[N-(butyrylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, δ) : 0.98 (3H, t, J=6Hz), 1.53-1.75 (2H), 2.23 (2H, t, J=6Hz), 2.57 (3H, s), 3.27 (3H, s), 3.68 (1H, s), 3.70 (1H, s), 5.70 (1H, d, J=10Hz), 5.78 (1H, d, J=10Hz), 7.56-7.75 (4H), 8.30 (1H, d, J=6Hz)

(62) 3-Chloro-8-[2,6-dichloro-3-[N'-phenylthioureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
mp : 150-155°C
NMR (DMSO-d$_6$, δ) : 2.43 (3H, s), 3.18 (3H, s), 3.80 (1H, dd, J=16Hz and 4Hz), 4.23 (1H, dd, J=16Hz and 4Hz), 5.61 (2H, s), 7.11 (1H, t, J=8Hz), 7.27-7.92 (8H), 7.97 (1H, br s), 8.28 (1H, d, J=6Hz), 10.15 (1H, br s)

(63) 3-Bromo-8-[2,6-dichloro-3-[N-(glycylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, δ) : 2.56 (3H, s), 3.25 (3H, s), 3.69 (1H, d, J=17Hz), 3.72 (2H, s), 3.86 (1H, d, J=17Hz), 5.72 (2H, s), 7.50-7.78 (4H), 8.31 (1H, d, J=6Hz)
Example 62

The following compounds were obtained according to similar manners to those of Examples 1 or 2 using N-iodosuccinimide instead of N-bromosuccinimide or N-chlorosuccinimide.

1) 8-[2,6-Dichloro-3-[N-[N-(N,N-dimethylglycyl)glycyl]-N-methylamino]benzyloxy]-3-iodo-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 2.31 (6H, s), 2.48 (3H, s), 2.96 (2H, s), 3.25 (3H, s), 3.55 (1H, dd, J=18Hz and 5Hz), 3.85 (1H, dd, J=18Hz and 5Hz), 5.50 (2H, s), 6.72 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.79 (1H, d, J=7Hz), 7.89 (1H, br s)

Example 63

The following compounds were obtained according to similar manners to those of Example 1 or 2.

1) 3-Bromo-8-[2,6-dichloro-4-(N-acetyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp: 82-88°C

NMR (CDCl₃, δ): 2.06 (3H, br s), 2.45 (3H, s), 3.28 (3H, s), 5.45 (2H, s), 6.73 (1H, d, J=7.5Hz), 6.86 (1H, t, J=7.5Hz), 7.25 (2H, s), 7.77 (1H, d, J=7.5Hz)
(2) 3-Chloro-8-[2,6-dichloro-4-(N-acetyl-N-methylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine
   mp : 74-78°C
   NMR (CDCl₃, δ) : 2.04 (3H, br s), 2.44 (3H, s), 3.28 (3H, s), 4.54 (2H, s), 6.70 (1H, d, J=7Hz), 6.84 (1H, t, J=7.0Hz), 7.25 (2H, s), 7.71 (1H, d, J=7Hz)

Example 64

The following compounds were obtained according to similar manners to those of Example 42 or 43.

(1) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-(3-methoxyphenyl)ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
   NMR (CDCl₃, δ) : 2.41 (3H, s), 3.21 (3H, s), 3.61-3.91 (5H), 5.47 (2H, s), 5.95 (1H, br t, J=4Hz), 6.58 (1H, dd, J=7Hz and 1Hz), 6.58-6.91 (3H), 7.01 (1H, t, J=1Hz), 7.13 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.42 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(2) 3-Bromo-8-[3-[N-[N'-(3-chlorophenyl)ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine
   mp : 151-152°C
   NMR (CDCl₃, δ) : 2.42 (3H, s), 3.21 (3H, s), 3.88 (1H, dd, J=18Hz and 4Hz), 4.21 (1H, dd, J=18Hz and 4Hz), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.18-7.42 (6H), 7.50 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 8.01 (1H, br s)

(3) 3-Bromo-8-[2,6-dichloro-3-[N'-{(3-trifluoromethylphenyl)ureidoacetyl]-N-methylamino]benzyloxy]–2-methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, $\delta$) : 2.40 (3H, s), 3.22 (3H, s), 3.82 (2H, br d, $J=5$Hz), 5.42 (1H, d, $J=10$Hz), 5.52 (1H, d, $J=10$Hz), 6.07 (1H, br t, $J=5$Hz), 6.73 (1H, d, $J=7$Hz), 6.89 (1H, t, $J=7$Hz), 7.11-7.46 (5H), 7.65 (1H, br s), 7.79 (1H, d, $J=7$Hz), 8.22 (1H, br s)

(4) 8-[3-[N-[N'-[(3-cyanophenyl)ureidoacetyl]ureidoacetyl]-N-methylamino]-2,6-dichlorobenzylcxy]-3-bromo-2-methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, $\delta$) : 2.40 (3H, s), 2.54 (3H, s), 3.37 (3H, s), 3.76 (1H, dd, $J=18$Hz and 5Hz), 3.89 (1H, d, $J=18$Hz and 5Hz), 5.44 (1H, d, $J=10$Hz), 5.52 (1H, d, $J=10$Hz), 6.09 (1H, br t, $J=5$Hz), 6.72 (1H, d, $J=7$Hz), 6.88 (1H, t, $J=7$Hz), 7.28 (1H, t, $J=7$Hz), 7.39 (1H, d, $J=9$Hz), 7.43 (1H, d, $J=9$Hz), 7.54 (2H, d, $J=7$Hz), 7.79 (1H, d, $J=7$Hz), 7.86 (1H, br s), 8.00 (1H, br s)

(5) 3-Bromo-8-[3-[N-[N'-[(3-cyanophenyl)ureidoacetyl]ureidoacetyl]-N-methylamino]-2,6-dichlorobenzylcxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, $\delta$) : 2.40 (3H, s), 3.21 (3H, s), 3.81 (2H, d, $J=5$Hz), 5.42 (1H, d, $J=10$Hz), 5.53 (1H, d, $J=10$Hz), 6.08 (1H, br t, $J=5$Hz), 6.76 (1H, d, $J=7$Hz), 6.90 (1H, t, $J=7$Hz), 7.11-7.26 (2H), 7.31-7.42 (3H), 7.86 (1H, br s), 7.80 (1H, d, $J=7$Hz), 8.63 (1H, br s)

(6) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[(o-tolyl)ureidoacetyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, $\delta$) : 2.25 (3H, s), 2.42 (3H, s), 3.21 (3H, s), 3.58 (1H, dd, $J=18$Hz and 5Hz), 3.85 (1H, dd, $J=18$Hz and 5Hz), 5.48 (2H, s), 5.72
(1H, br s), 6.37 (1H, br s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.05-7.51 (6H), 7.78 (1H, d, J=7Hz)

5  (7) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-((3-fluorophenyl)ureidoacetyl]-N-methylamino)benzoylox]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 2.41 (3H, s), 3.21 (3H, s), 3.79 (2H, br t, J=5Hz), 5.43 (1H, d, J=10Hz), 5.51 (1H, d, J=10Hz), 6.02 (1H, br t, J=5Hz), 6.58-6.94 (4H), 7.03-7.28 (2H), 7.34 (1H, d, J=9Hz), 7.41 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 7.98 (1H, br s)

10 (8) 3-Bromo-8-[3-[N-[N'-((3-ethylphenyl)ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 1.20 (3H, t, J=7Hz), 2.42 (3H, s), 2.60 (2H, q, J=7Hz), 3.22 (3H, s), 3.68 (1H, dd, J=18Hz and 5Hz), 3.87 (1H, dd, J=18Hz and 5Hz), 5.48 (2H, s), 5.91 (1H, br t, J=5Hz), 6.71 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.01-7.26 (4H), 7.32 (1H, d, J=9Hz), 7.45 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

20 (8) 8-[3-[N-(N'-Benzyolureidoacetyl)-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine
mp : 187-190°C (dec.)
NMR (CDCl₃, δ) : 2.42 (3H, s), 3.28 (3H, s), 3.71 (1H, dd, J=18Hz and 5Hz), 3.96 (1H, dd, J=18Hz and 5Hz), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.36 (1H, d, J=9Hz), 7.40-7.62 (4H), 7.77 (1H, d, J=7Hz), 7.83 (2H, d, J=8Hz), 8.45 (1H, br s), 9.23 (1H, br t, J=5Hz)
(10) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-\(3\)-nitrophenyl]ureidoacetyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 160-168°C (dec.)

NMR (CDCl\(_3\), \(\delta\)) : 2.40 (3H, s), 3.25 (3H, s), 3.83 (2H, d, \(J=5\)Hz), 5.42 (1H, d, \(J=10\)Hz), 5.52 (1H, d, \(J=7\)Hz), 6.16 (1H, br t, \(J=5\)Hz), 6.77 (1H, d, \(J=8\)Hz), 7.42 (1H, t, \(J=8\)Hz), 7.70 (1H, dd, \(J=8\)Hz and 1Hz), 7.79 (1H, d, \(J=7\)Hz), 8.14 (1H, t, \(J=1\)Hz), 8.79 (1H, s)

(11) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-\(3\)-ethoxycarbonylphenyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl\(_3\), \(\delta\)) : 1.35 (3H, t, \(J=7\)Hz), 2.41 (3H, s), 3.25 (3H, s), 3.80 (2H, dd, \(J=7\)Hz and 5Hz), 4.32 (2H, q, \(J=7\)Hz), 5.44 (1H, d, \(J=10\)Hz), 5.51 (1H, d, \(J=10\)Hz), 6.01 (1H, br t, \(J=5\)Hz), 6.72 (1H, d, \(J=7\)Hz), 6.88 (1H, t, \(J=7\)Hz), 7.36 (1H, d, \(J=9\)Hz), 7.42 (1H, d, \(J=9\)Hz), 7.56-7.69 (2H), 7.72-7.89 (3H)

(12) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-\(3\)-ethoxycarbonylmethylureidoacetyl]]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 194-196°C

NMR (CDCl\(_3\), \(\delta\)) : 1.28 (3H, t, \(J=7\)Hz), 2.42 (3H, s), 3.22 (3H, s), 3.57 (1H, dd, \(J=18\)Hz and 5Hz), 3.80 (1H, dd, \(J=18\)Hz and 5Hz), 3.92 (2H, d, \(J=5\)Hz), 4.20 (2H, q, \(J=7\)Hz), 5.38 (1H, br t, \(J=5\)Hz), 5.49 (2H, s), 5.62 (1H, br t, \(J=5\)Hz), 6.71 (1H, d, \(J=7\)Hz), 6.86 (1H, t, \(J=7\)Hz), 7.32 (1H, d, \(J=9\)Hz), 7.47 (1H, d, \(J=9\)Hz), 7.78 (1H, d, \(J=7\)Hz)
Example 65

A mixture of 2-thiophenecarboxylic acid (60 mg), triethylamine (52 mg) and diphenylphosphoryl azide (135 mg) in dry toluene (0.6 ml) was refluxed. After 1 hour, to the cooled mixture was added a solution of 8-[3-(N-glycyl-N-methylamino)-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine (200 mg) in dry dichloromethane (1 ml), and the mixture was stirred at ambient temperature. After 1 hour, to the mixture was added 2-thiophenecarboxylic acid (60 mg), triethylamine (52 mg) and diphenylphosphoryl azide (135 mg). The mixture was refluxed for 1 hour. The reaction mixture was washed with water twice and brine. The organic layer was dried over magnesium sulfate and evaporated in vacuo. The residue was purified by silica gel column chromatography (ethyl acetate:methanol = 50:1, V/V) followed by preparative thin layer chromatography (dichloromethane: methanol = 10:1, V/V) to give 3-bromo-8-[2,6-dichloro-3-[N-[N'-\(\text{N}''\)-2-thienyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (201 mg) as amorphous.

NMR (CDCl₃, δ) : 2.42 (3H, s), 3.29 (3H, s), 3.71 (1H, dd, J=18Hz and 4Hz), 3.94 (1H, dd, J=18Hz and 5Hz), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.00 (1H, br s), 7.09 (1H, dd, J=5Hz and 4Hz), 7.34 (1H, d, J=9Hz), 7.42-7.60 (3H), 7.78 (1H, d, J=7Hz)

Example 66

3-Bromo-8-[3-[N-[N'-\(\text{N}''\)-3-carboxyphenyl]ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Example 57.

mp : 248-250°C

NMR (DMSO-d₆, δ) : 2.30 (3H, s), 3.18 (3H, s), 3.43 (1H, dd, J=18Hz and 5Hz), 3.69 (1H, dd, J=18Hz
Example 67

To a solution of 3-bromo-8-[3-[N-[N'-((3-carboxyphenyl)ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine (200 mg) and dimethylamine hydrochloride (31 mg) in N,N-dimethylformamide (2 ml) were added N-ethyl-N'-(3-dimethylaminopropyl)carbodiimide (64 mg) and 1-hydroxybenzotriazole (64 mg), and the mixture was stirred for 1 hour at ambient temperature. Water was added thereto, and the mixture was extracted with ethyl acetate three times. The organic layer was washed with water for times and brine, dried over magnesium sulfate and concentrated in vacuo. The residue was purified by a silica gel column chromatography (methylene chloride:methanol = 20:1, V/V) to give 3-bromo-8-[3-[N-[N'-[3-(N,N-dimethylcarbamoyl)phenyl]ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine (189 mg).

NMR (CDCl₃, δ) : 2.41 (3H, s), 2.95 (3H, br s), 3.09 (3H, br s), 3.22 (3H, s), 3.64 (1H, dd, J=17Hz and 5Hz), 3.82 (1H, dd, J=17Hz and 5Hz), 5.48 (2H, s), 6.06 (1H, br t, J=5Hz), 6.72 (1H, d, 7Hz), 6.88 (1H, t, J=7Hz), 6.99 (1H, d, J=8Hz); 7.20 (1H, t, J=8Hz), 7.30-7.49 (4H), 7.78 (1H, d, J=7Hz), 8.11 (1H, s)

Example 68

3-Bromo-8-[2,6-dichloro-3-[N-[N-[3-(N-methylcarbamoyl)propionyl]glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Example 67.
NMR (CDCl$_3$, $\delta$): 2.40-2.63 (7H), 2.79 (3H, d, J=5Hz), 3.25 (3H, s), 3.52, 3.80 (each 1H, dd, J=18Hz and 5Hz), 5.48 (2H, s), 6.01 (1H, br s), 6.62-6.76 (2H), 6.86 (1H, t, J=7Hz), 7.31, 7.48 (each 1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

Example 69

To a mixture of 8-[3-(N-glycyl-N-methylamino)-2,6-dichlorobenzylloxy]-3-chloro-2-methylimidazo[1,2-a]pyridine (200 mg), triethylamine (0.1 ml) and dichloromethane (2 ml) was added bromoacetyl chloride (0.042 ml) in a dry ice-acetone bath. After 20 minutes, to the mixture was added 50% aqueous solution of dimethylamine (0.42 ml). The mixture was stirred for 1 hour at ambient temperature. The reaction mixture was washed with aqueous sodium bicarbonate solution, water and brine. The organic layer was dried over magnesium sulfate and evaporated in vacuo. The residue was purified by a silica gel column chromatography (dichloromethane:methanol = 20:1, V/V) to yield colorless crystals (191 mg) of 3-chloro-8-[2,6-dichloro-3-[N-[N-(N,N-dimethylglycyl)glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine.

mp: 171-172°C

NMR (CDCl$_3$, $\delta$): 2.31 (6H, s), 2.44 (3H, s), 2.96 (2H, s), 3.23 (3H, s), 3.56 (1H, dd, J=18Hz, and 4Hz), 3.85 (1H, dd, J=18Hz and 4Hz), 5.49 (2H, s), 6.70 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.72 (1H, d, J=7Hz), 7.89 (1H, br s)

Example 70

The following compounds were obtained according to similar manners to those of Examples 53 or 69.

(1) 3-Bromo-8-[2,6-dichloro-3-[N-[N-(N-ethyl-N-methyl-
glycyl)glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp : 168-170°C
NMR (CDCl₃, δ) : 1.09 (3H, t, J=7Hz), 2.30 (3H, s), 2.43 (3H, s), 2.50 (2H, q, J=7Hz), 2.99 (2H, s), 3.25 (3H, s), 3.56 (1H, dd, J=17Hz and 5Hz), 3.85 (1H, dd, J=17Hz and 5Hz), 5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 8.01 (1H, br s)

(2) 3-Bromo-8-[3-[N-(N-cyclopropylglycyl)glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 0.39-0.53 (4H), 2.22 (1H, m), 2.42 (3H), 3.25 (3H, s), 3.38 (2H, s), 3.55 (1H, dd, J=18Hz and 4Hz), 3.82 (1H, dd, J=18Hz and 4Hz), 5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.68 (1H, br s), 7.78 (1H, d, J=7Hz)

(3) 3-Bromo-8-[3-[N-(N-cyclohexylglycyl)glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 182-184°C
NMR (CDCl₃, δ) : 0.98-1.31 (5H), 1.52-1.95 (5H), 2.39 (1H, m), 2.43 (3H, s), 3.24 (3H, s), 3.30 (2H, s), 3.56 (1H, dd, J=18Hz and 4Hz), 3.82 (1H, dd, J=18Hz and 4Hz), 5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.47 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 8.11 (1H, br s)

(4) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N-(1-pyrrolidinylacetyl)glycyl]amino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 1.72-1.90 (4H), 2.42 (3H, s), 2.57-2.70 (4H), 3.16 (2H, s), 3.25 (3H, s), 3.57 (1H, dd, J=18Hz and 5Hz), 3.87 (1H, dd, J=18Hz and 5Hz), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 7.89 (1H, br s)

(5) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N-(piperidinoacetyl)glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 1.38-1.72 (6H), 2.39-2.55 (7H), 2.94 (2H, s), 3.27 (3H, s), 3.55 (1H, dd, J=18Hz and 4Hz), 3.83 (1H, dd, J=18Hz and 4Hz), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 8.08 (1H, br s)

(6) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N-[[4-((4-pyridyl)-1-piperazinyl)acetyl]glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 2.43 (3H, s), 2.61-2.78 (4H), 3.09 (2H, s), 3.26 (3H, s), 3.34-3.49 (4H), 3.58 (1H, dd, J=18Hz and 4Hz), 3.88 (1H, dd, J=18Hz and 4Hz), 5.48 (1H, d, J=10Hz), 5.52 (1H, d, J=10Hz), 6.61-6.78 (3H), 6.86 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.50 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 7.92 (1H, br t, J=4Hz), 8.29 (2H, br d, J=6Hz)

(7) 3-Bromo-8-[2,6-dichloro-3-[N-[(N-isopropyl-N-methylglycyl)glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp: 150-151°C
NMR (CDCl$_3$, $\delta$): 1.02 (6H, d, $J=6$Hz), 2.38 (3H, s), 2.42 (3H, s), 2.85 (1H, m), 3.00 (2H, s), 3.25 (3H, s), 3.55 (1H, dd, $J=18$Hz and 5Hz), 3.85 (1H, dd, $J=18$Hz and 5Hz), 5.49 (2H, s), 6.71 (1H, d, $J=7$Hz), 6.86 (1H, t, $J=7$Hz), 7.32 (1H, d, $J=9$Hz), 7.49 (1H, d, $J=9$Hz), 7.76 (1H, d, $J=7$Hz), 8.12 (1H, br s)

(8) 3-Bromo-8-[3-[N-[N-(N-cyclohexyl-N-methylglycyl)-glycyl] -N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 1.00-1.33 (6H), 1.70-1.90 (4H), 2.31 (3H, s), 2.38 (1H, m), 2.42 (3H, s), 3.02 (2H, s), 3.25 (3H, s), 3.56 (1H, dd, $J=18$Hz and 5Hz), 3.83 (1H, dd, $J=18$Hz and 5Hz), 5.49 (2H, s), 6.71 (1H, d, $J=7$Hz), 6.06 (1H, t, $J=7$Hz), 7.32 (1H, d, $J=9$Hz), 7.48 (1H, d, $J=9$Hz), 7.78 (1H, d, $J=7$Hz), 8.19 (1H, br s)

(9) 3-Bromo-8-[3-[N-[N-(N-cycloheptylglycyl)glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 1.29-1.92 (12H), 2.43 (3H, s), 2.62 (1H, m), 3.24 (3H, s), 3.28 (2H, s), 3.58 (1H, dd, $J=18$Hz and 5Hz), 3.82 (1H, dd, $J=18$Hz and 5Hz), 5.49 (2H, s), 6.71 (1H, d, $J=7$Hz), 6.86 (1H, t, $J=7$Hz), 7.31 (1H, d, $J=9$Hz), 7.48 (1H, d, $J=9$Hz), 7.78 (1H, d, $J=7$Hz), 8.10 (1H, br s)

(10) 3-Bromo-8-[3-[N-[N-(N-cyclopentylglycyl)glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 1.23-1.90 (8H), 2.42 (3H, s), 3.09 (1H, m), 3.24 (3H, s), 3.28 (2H, s), 3.56 (1H, dd, $J=18$Hz and 5Hz), 3.82 (1H, dd, $J=18$Hz and
5Hz), 5.49 (2H, s), 6.72 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 8.00 (1H, br s)

(11) 3-Bromo-8-[3-[N-[N-(N-cycloctylglycyl)glycyl]-N-methylamino]-2,6-dichlobenzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, $\delta$): 1.35-1.85 (14H), 2.42 (3H, s), 2.65 (1H, m), 3.27 (5H, s), 3.58 (1H, dd, J=18Hz and 10Hz), 3.82 (1H, dd, J=18Hz and 5Hz), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.79 (1H, d, J=7Hz), 8.10 (1H, br s)

(12) 3-Bromo-8-[3-[N-[N-(cycloheptyl-N-methylglycyl)-glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, $\delta$): 1.28-1.92 (12H), 2.29 (3H, s), 2.15 (3H, s), 2.60 (1H, m), 3.01 (2H, s), 3.25 (3H, s), 3.56 (1H, dd, J=18Hz and 5Hz), 3.85 (1H, dd, J=18Hz and 5Hz), 5.50 (2H, s), 6.72 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.79 (1H, d, J=7Hz), 8.18 (1H, br s)

(13) 3-Bromo-8-[2,6-dichloro-3-[N-[N-(hexamethyleneimino-acetyl)glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, $\delta$): 1.58-1.78 (8H), 2.42 (3H, s), 2.63-2.73 (4H), 3.13 (2H, s), 3.28 (3H, s), 3.58 (1H, dd, J=18Hz and 5Hz), 3.83 (1H, dd, J=18Hz and 5Hz), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 8.12 (1H, br s)
(14) 8-[3-[N-[N-[1-Adamantyl]glycyl]glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 1.49-1.74 (12H), 1.99-2.12 (3H), 2.43 (3H, s), 3.25 (5H, s), 3.56 (1H, dd, $J$=18Hz and 5Hz), 3.82 (1H, dd, $J$=18Hz and 5Hz), 5.50 (2H, s), 6.71 (1H, d, $J$=7Hz), 6.86 (1H, t, $J$=7Hz), 7.32 (1H, d, $J$=9Hz), 7.49 (1H, d, $J$=9Hz), 7.78 (1H, d, $J$=7Hz), 8.25 (1H, br s)

(15) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N-[N-2-furylmethyl]-glycyl]glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 242-243°C

NMR (CDCl$_3$-CD$_3$OD, $\delta$): 1.50-1.90 (4H), 2.06-2.34 (2H), 2.42 (3H, s), 2.66 (1H, m), 3.22 (3H, s), 3.52-3.87 (5H), 4.53 (2H, s), 5.50 (2H, s), 6.73 (1H, d, $J$=7Hz), 6.89 (1H, t, $J$=7Hz), 7.44 (1H, d, $J$=9Hz), 7.52 (1H, d, $J$=9Hz), 7.79 (1H, d, $J$=7Hz)

(16) 3-Bromo-8-[2,6-dichloro-3-[N-[N-[2-furylmethyl]-glycyl]glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 2.42 (3H, s), 3.25 (3H, s), 3.30 (2H, s), 3.56 (1H, dd, $J$=18Hz and 5Hz), 3.72-3.91 (3H), 5.50 (2H, s), 6.20 (1H, d, $J$=2Hz), 6.30 (1H, m), 6.71 (1H, d, $J$=7Hz), 6.86 (1H, t, $J$=7Hz), 7.29-7.39 (2H), 7.48 (1H, d, $J$=9Hz), 7.77 (1H, d, $J$=7Hz), 7.88 (1H, br s)

(17) 3-Bromo-8-[2,6-dichloro-3-[N-[N,N-diethylglycyl]-glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp : 159-160°C
NMR (CDCl$_3$, $\delta$): 1.06 (6H, t, $J=7$Hz), 2.43 (3H, s), 2.58 (4H, q, $J=7$Hz), 3.02 (2H, s), 3.26 (3H, s), 3.55 (1H, dd, $J=18$Hz and 4Hz), 3.84 (1H, dd, $J=18$Hz and 4Hz), 5.49 (2H, s), 6.71 (1H, d, $J=7$Hz), 6.86 (1H, t, $J=7$Hz), 7.31 (1H, d, $J=9$Hz), 7.48 (1H, d, $J=9$Hz), 7.76 (1H, d, $J=7$Hz), 8.18 (1H, br s)

Example 71

To a solution of 3-bromo-8-[2,6-dichloro-3-[N-[N-(1-imidazolylacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (200 mg) in N,N-dimethylformamide (2 ml) was added sodium methanethiolate (35 mg), and the mixture was stirred for 5 hours at ambient temperature under nitrogen atmosphere. The reaction mixture was concentrated in vacuo, the residue was dissolved in methylene chloride. The solution was washed with water four times and brine, dried over magnesium sulfate and concentrated in vacuo. The residue was purified by a silica gel column chromatography (methylene chloride:methanol = 40:1, V/V) to give 3-bromo-8-[2,6-dichloro-3-[N-methyl-N-[N-(methylthioacetyl)glycyl]amino]-benzyloxy]-2-methylimidazo[1,2-a]pyridine (105 mg).

NMR (CDCl$_3$, $\delta$): 2.19 (3H, s), 2.43 (3H, s),
3.21 (2H, s), 3.28 (3H, s), 3.59 (1H, dd, J=18Hz and 4Hz), 3.84 (1H, dd, J=18Hz and 5Hz), 5.50 (2H, s), 6.72 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.61 (1H, br s), 7.78 (1H, d, J=7Hz)

Example 72

The following compounds were obtained according to similar manners to those of Examples 36 to 39.

(1) 3-Bromo-8-[2,6-dichloro-3-[N-(N-hexanoylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

mp: 131-132°C

NMR (CDCl₃, δ): 0.89 (3H, t, J=6Hz), 1.19-1.40 (4H), 1.51-1.71 (2H), 2.20 (2H, t, J=6Hz), 2.42 (3H, s), 3.25 (3H, s), 3.51 (1H, dd, J=18Hz and 4Hz), 3.79 (1H, dd, J=18Hz and 5Hz), 5.49 (2H, s), 6.40 (1H, br s), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.30 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(2) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N-(3-phenylpropionyl)glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 2.42 (3H, s), 2.88-3.02 (4H), 3.23 (3H, s), 3.50 (1H, dd, J=18Hz and 4Hz), 3.79 (1H, dd, J=18Hz and 5Hz), 5.49 (2H, s), 6.45 (1H, br s), 6.71 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.11-7.33 (5H), 7.46 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(3) 3-Bromo-8-[2,6-dichloro-3-[N-[N-(N-methyl-N-phenylglycyl)glycyl]N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
Example 73

The following compounds were obtained according to a similar manner to that of Example 60.

1. 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N-
(methylthioacetyl)glycyl]amino]benzyl]oxy]-2-methyl-
imidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.19 (3H, s), 2.53 (3H, s),
3.21 (2H, s), 3.27 (3H, s), 3.66 (1H, d,
J=18Hz), 3.79 (1H, d, J=18Hz), 5.72 (2H, s),
7.49-7.73 (4H), 8.29 (1H, d, J=6Hz)

2. 3-Bromo-8-[2,6-dichloro-3-[N-[N-(1-imidazolylacetyl)]-
glycyl]N-methylamino]benzyl]oxy]-2-methylimidazo-
[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.52 (3H, s), 3.26 (3H, s),
3.63 (1H, d, J=18Hz), 3.83 (1H, d, J=18Hz), 5.12
(2H, s), 5.71 (2H, s), 7.48-7.72 (6H), 8.29 (1H,
d, J=6Hz), 9.00 (1H, s)

3. 3-Bromo-8-[2,6-dichloro-3-[N-[N-hexanoylglycyl]-N-
methylamino]benzyl]oxy]-2-methylimidazo[1,2-a]pyridine
hydrochloride

NMR (CDCl₃-CD₃OD, δ): 0.90 (3H, t, J=6Hz),
1.21-1.42 (4H), 1.52-1.71 (2H), 2.25 (2H, t,
J=6Hz), 2.56 (3H, s), 3.27 (3H, s), 3.62 (1H, d,
J=16Hz), 3.74 (1H, d, J=16Hz), 5.70 (1H, d,
J=10Hz), 5.78 (1H, d, J=10Hz), 7.53-7.76 (4H),
8.30 (1H, d, J=6Hz)
(4) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N-(3-phenylpropionyl)glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.53 (3H, s), 2.83-3.00 (4H), 3.22 (3H, s), 3.18 (2H, s), 5.71 (2H, s), 7.10-7.30 (5H), 7.55-7.76 (4H), 8.30 (1H, d, J=6Hz)

(5) 3-Bromo-8-[2,6-dichloro-3-[N-[N-(N,N-diethylglycyl)glycyl]N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ): 1.40 (6H, t, J=7Hz), 2.63 (3H, s), 3.25 (3H, s), 3.28-3.49 (4H), 3.60-4.09 (4H), 5.61 (1H, d, J=10Hz), 5.70 (1H, d, J=10Hz), 7.40-7.62 (4H), 8.11 (1H, d, J=6Hz)

(6) 3-Bromo-8-[2,6-dichloro-3-[N-[N-(N-ethyl-N-methylglycyl)glycyl]N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ): 0.84-1.03 (4H), 2.59 (3H, s), 2.79 (1H, m), 3.25 (3H, s), 3.58-3.71 (2H), 3.80-4.00 (3H), 5.73 (2H, s), 7.56-7.75 (4H), 8.30 (1H, d, J=6Hz)

(7) 3-Bromo-8-[3-[N-[N-(N-cyclopropylglycyl)glycyl]N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ): 0.84-1.03 (4H), 2.59 (3H, s), 2.79 (1H, m), 3.25 (3H, s), 3.58-3.71 (2H), 3.80-4.00 (3H), 5.73 (2H, s), 7.56-7.75 (4H), 8.30 (1H, d, J=6Hz)

(8) 3-Bromo-8-[3-[N-[N-(N-cyclohexylglycyl)glycyl]N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
1.95

NMR (CDCl₃-CD₃OD, δ):  1.16-1.51 (5H), 1.68-2.20 (5H), 2.58 (3H, s), 3.08 (1H, m), 3.26 (3H, s), 3.56-3.71 (2H), 3.80-3.92 (3H), 5.72 (2H, s), 7.48-7.74 (4H), 8.30 (1H, d, J=6Hz)

(9) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(1-pyrrolidinylacetyl)glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ):  1.97-2.29 (4H), 2.58 (3H, s), 3.09-3.30 (5H), 3.53-3.91 (4H), 4.09 (2H, s), 5.71 (2H, s), 7.53-7.76 (4H), 8.30 (1H, d, J=6Hz)

(10) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(N-(piperidino-acetyl)glycyl)amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ):  1.42-2.09 (6H), 2.58 (3H, s), 2.98-3.19 (2H), 3.28 (3H, s), 3.50-3.71 (4H), 3.80-4.04 (2H), 5.72 (2H, s), 7.55-7.76 (4H), 8.30 (1H, d, J=6Hz)

(11) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[[N-(4-pyridyl)-1-piperazinylacetyl)glycyl]amino]-benzyloxy]-2-methylimidazo[1,2-a]pyridine tetrahydrochloride

NMR (CDCl₃-CD₃OD, δ):  2.59 (3H, s), 3.28 (3H, s), 3.58-3.72 (5H), 3.89 (1H, d, J=18Hz), 4.02-4.25 (4H), 5.72 (2H, s), 7.30 (2H, d, J=7Hz), 7.55-7.76 (4H), 8.21 (2H, d, J=7Hz), 8.30 (1H, d, J=6Hz)

(12) 3-Bromo-8-[2,6-dichloro-3-[N-[(N-isopropyl-N-methylglycyl)glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ):  1.40 (6H, d, J=6Hz),
2.56 (3H, s), 2.85 (3H, s), 3.25 (3H, s),
2.59-3.71 (3H), 3.90 (1H, d, J=18Hz), 4.01 (1H, m), 5.72 (2H, s), 7.48-7.78 (4H), 8.30 (1H, d, J=6Hz)

(13) 3-Bromo-8-[3-[N-[N-(N-cyclohexyl-N-methylglycyl)-glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, δ) : 1.43-1.90 (8H), 2.01-2.19 (2H), 2.56 (3H, s), 3.26 (3H, s), 3.31-3.41 (4H), 3.62 (1H, d, J=18Hz), 3.80-3.96 (3H), 5.72 (2H, s), 7.50-7.72 (4H), 8.29 (1H, d, J=6Hz)

(14) 3-Bromo-8-[3-[N-[N-(N-cycloheptylglycyl)glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, δ) : 1.25-2.20 (12H), 2.59 (3H, s), 2.89 (3H, s), 3.26 (3H, s), 3.31-3.41 (2H), 3.62 (1H, d, J=18Hz), 3.90 (1H, d, J=18Hz), 4.09 (1H, m), 5.73 (2H, s), 7.56-7.75 (4H), 8.30 (1H, d, J=6Hz)

(15) 3-Bromo-8-[3-[N-[N-(N-cyclopentylglycyl)glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, δ) : 1.59-1.93 (6H), 2.00-2.23 (2H), 2.56 (3H, s), 3.26 (3H, s), 3.49-3.67 (2H), 3.80-3.92 (3H), 5.72 (2H, s), 7.50-7.74 (4H), 8.30 (1H, d, J=6Hz)

(16) 3-Bromo-8-[3-[N-[N-(N-cyclooctylglycyl)glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, δ) : 1.31-2.10 (14H), 2.63 (3H, s), 3.23 (3H, s), 3.30 (1H, m), 3.71 (1H, d,
J=18Hz), 3.81 (2H, s), 3.90 (1H, d, J=18Hz), 5.65 (2H, s), 7.40-7.66 (4H), 8.09 (1H, d, J=6Hz)

(17) 3-Bromo-8-[3-[N-[N-(N-cycloheptyl-N-methylglycyl)glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ): 1.42-1.90 (10H), 2.05-2.30 (2H), 2.65 (3H, s), 2.88 (3H, s), 3.24 (3H, s), 3.50-3.72 (2H), 3.80-4.10 (3H), 5.61 (1H, d, J=10Hz), 5.71 (1H, d, J=10Hz), 7.42-7.62 (4H), 8.10 (1H, d, J=6Hz)

(18) 3-Bromo-6-[2,6-dichloro-3-[N-[N-[hexamethyleneiminoacetyl]glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ): 1.42-2.20 (8H), 2.66 (3H, s), 3.22 (3H, s), 3.30-3.82 (6H), 3.90-4.08 (2H), 5.56 (1H, d, J=10Hz), 5.69 (1H, d, J=10Hz), 7.20-7.50 (3H), 7.59 (1H, d, J=9Hz), 8.00 (1H, d, J=6Hz)

(19) 8-[3-[N-[N-[N-[1-Adamantyl]glycyl]glycyl]glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ): 1.56-2.35 (15H), 2.65 (3H, s), 3.22 (3H, s), 3.66-3.99 (4H), 5.60 (1H, d, J=10Hz), 5.70 (1H, d, J=10Hz), 7.31-7.54 (3H), 7.62 (1H, d, J=9Hz), 8.02 (1H, d, J=6Hz)

(20) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N-[N-(piperidino)glycyl]glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine trihydrochloride
NMR (CDCl₃-CD₃OD, δ): 1.52-2.40 (6H), 2.68 (3H, s), 3.22 (3H, s), 3.51-3.99 (6H), 4.59 (1H, d,
J=11Hz), 4.70 (1H, d, J=11Hz), 5.67 (2H, s),
7.31-7.50 (2H), 7.55 (1H, d, J=9Hz), 7.67 (1H,
d, J=9Hz), 8.06 (1H, d, J=6Hz)

(21) 3-Bromo-8-[2,6-dichloro-3-[N-[N-(2-furylmethyl)glycyl]glycyl]-N-methylamino]benzyloxy]-2-methyl-
imidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.62 (3H, s), 3.22 (3H, s),
3.70-3.99 (4H), 4.30 (2H, s), 5.60 (1H, d,
J=10Hz), 5.79 (1H, d, J=10Hz), 6.39 (1H, m),
6.69 (1H, d, J=3Hz), 7.37-7.62 (5H), 8.08 (1H,
d, J=6Hz)

(22) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-methoxyphenyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-methyl-
imidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.51 (3H, s), 3.28 (3H, s),
3.71 (2H, s), 3.76 (3H, s), 5.70 (1H, d,
J=10Hz), 5.79 (1H, d, J=10Hz), 6.56 (1H, dd,
J=9Hz and 1Hz), 6.81 (1H, dd, J=9Hz and 1Hz),
7.06 (1H, t, J=1Hz), 7.12 (1H, t, J=9Hz),
7.50-7.78 (4H), 8.30 (1H, d, J=6Hz)

(23) 3-Bromo-8-[3-[N-[N'-[3-chlorophenyl]ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methyl-
imidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.55 (3H, s), 3.29 (3H, s),
3.95 (1H, d, J=18Hz), 4.16 (1H, d, J=18Hz), 5.77
(2H, s), 7.18 (1H, m), 7.31 (2H, d, J=5Hz),
7.50-7.72 (5H), 8.29 (1H, d, J=6Hz)

(24) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-trifluoromethylphenyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-
imidazol[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.52 (3H, s), 3.30 (3H, s),
(25) 8-[3-[N-[N'-{(3-Acetylphenyl)ureidoacetyl]}-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.54 (3H, s), 2.60 (3H, s), 3.30 (3H, s), 3.72 (2H, br s), 5.70 (1H, d, J=10Hz), 5.80 (1H, d, J=10Hz), 7.49-7.72 (6H), 8.04 (1H, br s), 8.29 (1H, d, J=6Hz)

(26) 3-Bromo-8-[3-[N-[N'-{(3-cyanophenyl)ureidoacetyl]}-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.58 (3H, s), 3.29 (3H, s), 3.72 (2H, s), 5.69 (1H, d, J=10Hz), 5.79 (1H, d, J=10Hz), 7.25-7.72 (7H), 7.97 (1H, br s), 8.28 (1H, d, J=6Hz)

(27) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-{(o-tolyl)ureidoacetyl]}-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.25 (3H, s), 2.49 (3H, s), 3.28 (3H, s), 3.73 (2H, s), 5.68 (1H, d, J=10Hz), 5.78 (1H, d, J=10Hz), 6.95-7.20 (3H), 7.40-7.71 (5H), 8.29 (1H, d, J=6Hz)

(28) 3-Bromo-8-[3-[N-[N'-{(3-fluorophenyl)ureidoacetyl]}-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.55 (3H, s), 3.29 (3H, s), 3.71 (2H, s), 5.67 (1H, d, J=10Hz), 5.78 (1H, d, J=6Hz)
J=10Hz), 6.68 (1H, dt, J=8Hz and 1Hz), 6.93 (1H, br d, J=8Hz), 7.20 (1H, dt, J=8Hz and 6Hz), 7.32 (1H, dt, J=10Hz and 1Hz), 7.48-7.71 (4H), 8.24 (1H, d, J=6Hz)

(29) 3-Bromo-8-[3-[N-[N'-(3-ethylphenyl)ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 1.21 (3H, t, J=7Hz), 2.52 (3H, s), 2.60 (2H, q, J=7Hz), 3.28 (3H, s), 3.71 (2H, s), 5.65 (1H, d, J=10Hz), 5.79 (1H, d, J=10Hz), 6.87 (1H, m), 7.10-7.22 (3H), 7.52-7.71 (4H), 8.25 (1H, d, J=6Hz)

(30) 8-[3-[N'-Benzoylureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.62 (3H, s), 3.30 (3H, s), 3.75 (1H, d, J=18Hz), 3.95 (1H, d, J=18Hz), 5.68 (2H, s), 7.38-7.66 (7H), 7.89 (2H, d, J=8Hz), 8.09 (1H, d, J=6Hz)

(31) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-3-nitrophenyl)ureidoacetyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.63 (3H, s), 3.25 (3H, s), 3.92 (2H, s), 5.59 (1H, d, J=10Hz), 5.68 (1H, d, J=10Hz), 7.29-7.58 (5H), 7.70 (1H, dd, J=7Hz and 1Hz), 7.79 (1H, d, J=7Hz), 8.10 (1H, d, J=6Hz), 8.19 (1H, br s)

(32) 3-Bromo-8-[2,6-dichloro-3-[N'-[3-ethoxycarbonylphenyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 1.30 (3H, t, J=7Hz),
2.62 (3H, s), 3.21 (3H, s), 3.99 (2H, s), 4.20 (2H, q, J=7Hz), 5.58 (2H, s), 7.12-7.52 (6H), 7.62-7.80 (2H), 8.01 (1H, d, J=6Hz)

(33) 3-Bromo-8-[2,6-dichloro-3-[N-(N'-ethoxycarbonylmethylureidoacetyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃, δ): 2.57 (3H, s), 3.28 (3H, s), 3.68 (2H, s), 3.89 (2H, s), 5.68 (1H, d, J=10Hz), 5.78 (1H, d, J=10Hz), 7.57-7.72 (4H), 8.29 (1H, d, J=6Hz)

(34) 3-Bromo-8-[3-[N-[N'-[3-(N,N-dimethylcarbamoyl)phenyl]ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.56 (3H, s), 3.01 (3H, br s), 3.11 (3H, br s), 3.30 (3H, s), 3.73 (2H, s), 5.68 (1H, d, J=10Hz), 5.79 (1H, d, J=10Hz), 7.00 (1H, d, J=8Hz), 7.30 (1H, t, J=8Hz), 7.38-7.70 (6H), 8.24 (1H, d, J=6Hz)

(35) 3-Bromo-8-[2,6-dichloro-3-[N-[N-[N-methylcarbamoyl]propionyl]glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.25-2.70 (5H), 2.80 (3H, s), 3.26 (3H, s), 3.72 (1H, d, J=18Hz), 3.89 (1H, d, J=18Hz), 5.60 (1H, d, J=10Hz), 5.70 (1H, d, J=10Hz), 7.38-7.61 (4H), 8.09 (1H, d, J=6Hz)

(36) 3-Bromo-8-[2,6-dichloro-3-[N-[N-methyl-N-phenylglycyl]glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.69 (3H, s), 3.21 (3H, s), 3.29 (3H, s), 3.64 (1H, d, J=17Hz), 3.81 (1H, d, J=17Hz), 4.20 (1H, d, J=16Hz), 4.32 (1H, d,
J=16Hz), 5.62 (2H, s), 7.20-7.61 (9H), 8.06 (1H, d, J=6Hz)

(37) 3-Bromo-8-[2,6-dichloro-3-[N-[N-[(2-thienyl)ureidoacetyl]]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, δ) : 2.63 (3H, s), 3.30 (3H, s), 3.81 (1H, d, J=17Hz), 3.93 (1H, d, J=17Hz), 5.62 (1H, d, J=10Hz), 5.72 (1H, d, J=10Hz), 7.09 (1H, dd, J=5Hz and 4Hz), 7.45-7.59 (5H), 7.69 (1H, d, J=4Hz), 8.06 (1H, dd, J=5Hz and 1Hz)

(38) 3-Bromo-8-[3-[N-[N'-((3-carboxyphenyl)ureidoacetyl)]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, δ) : 2.52 (3H, s), 3.29 (3H, s), 3.75 (2H, s), 5.68 (1H, d, J=10Hz), 5.79 (1H, d, J=10Hz), 7.32 (1H, t, J=8Hz), 7.49-7.72 (6H), 8.05 (1H, br s), 8.28 (1H, d, J=6Hz)

(39) 3-Chloro-8-[2,6-dichloro-3-[N-[N-(N,N-dimethylglycyl)glycyl]]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, δ) : 2.65 (3H, s), 2.96 (3H, s), 2.99 (3H, s), 3.22 (3H, s), 3.62-4.04 (4H), 5.68 (2H, s), 7.38-7.68 (4H), 8.08 (1H, d, J=6Hz)

(40) 8-[2,6-Dichloro-3-[N-[N-(N,N-dimethylglycyl)glycyl]]-N-methylamino]benzyloxy]-3-iodo-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, δ) : 2.58 (3H, s), 2.97 (6H, s), 3.27 (3H, s), 3.61 (1H, d, J=18Hz), 3.88 (1H, d, J=18Hz), 4.01 (2H, s), 5.72 (2H, s), 7.50-7.75 (4H), 8.28 (1H, d, J=6Hz)
(41) 3-Bromo-8-[2,6-dichloro-4-(N-acetyl-N-methylamino)-benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
mp: 118-121°C
NMR (DMSO-d$_6$, δ): 2.03 (3H, br s), 2.38 (3H, s), 3.25 (3H, s), 5.51 (2H, s), 7.36 (1H, t, J=7Hz), 7.50 (1H, d, J=7Hz), 7.71 (2H, s), 8.20 (1H, d, J=7Hz)

(42) 3-Chloro-8-[2,6-dichloro-4-(N-acetyl-N-methylamino)-benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (DMSO-d$_6$, δ): 2.01 (3H, br s), 2.38 (3H, s), 3.25 (3H, s), 5.50 (2H, s), 7.35 (1H, t, J=7Hz), 7.50 (1H, d, J=7Hz), 7.72 (2H, s), 8.22 (1H, d, J=7Hz)
Preparation 34

To a stirred two-phase solution of 3-nitrobenzoyl chloride (9.3 g) in a mixture of diethyl ether (50 ml) and saturated sodium bicarbonate solution (50 ml) was added 3-aminomethylpyridine (5.4 g) in an ice-cooled bath. The mixture was stirred vigorously at ambient temperature for 30 minutes. The reaction mixture was filtered, and the resulting solid was washed with water. The solid was further solidified with diisopropyl ether to afford 3-nitro-N-(3-pyridylmethyl)pyridine (5.91 g) as a pale yellow amorphous solid.

NMR (CDCl₃, δ) : 4.70 (2H, d, J=5Hz), 7.05 (1H, br s), 7.30 (1H, dd, J=7, 5Hz), 7.68 (1H, t, J=9Hz), 7.76 (1H, dt, J=8, 0.5Hz), 8.22 (1H, d, J=8Hz), 8.39 (1H, m), 8.54 (1H, dd, J=5, 0.5Hz), 8.60 (1H, d, J=0.5Hz), 8.65 (1H, t, J=0.5Hz)

Preparation 35

To a solution of N,N-bis(2-methoxyethyl)amine (2.40 g) and triethylamine (2.27 g) in dichloromethane (30 ml) was added 3-nitrobenzoyl chloride (2.78 g) in an ice-water bath. The mixture was stirred at ambient temperature for 1 hour. The reaction mixture was washed with saturated sodium bicarbonate solution, water and brine, dried over anhydrous magnesium sulfate, and evaporated in vacuo. The residue was purified with column chromatography eluting with dichloromethane-methanol to give N,N-bis(2-methoxyethyl)-3-nitrobenzamide (4.12 g) as an oil.

NMR (CDCl₃, δ) : 3.22-3.88 (14H), 7.59 (1H, t, J=8Hz), 7.80 (1H, dt, J=8, 1Hz), 8.26 (1H, dt, J=8, 1Hz), 8.39 (1H, t, J=8Hz)

Preparation 36

The following compounds were obtained according to similar manners to those of Preparations 34 or 35.
(1) 3-Nitro-N-(4-pyridyl)benzamide
mp : >250°C
NMR (DMSO-d₆, δ) : 7.80 (2H, d, J=6Hz), 7.89 (1H, t, J=7Hz), 8.38-8.58 (4H), 8.80 (1H, t, J=1Hz)

(2) 4-Methyl-1-(3-nitrobenzoyl)piperazine
mp : 97-98°C
NMR (CDCl₃, δ) : 2.31-2.66 (7H), 3.38-3.97 (4H), 7.62 (1H, dt, J=8, 1Hz), 7.78 (1H, dt, J=1, 8Hz), 8.25-8.34 (2H)

(3) 1-(3-Nitrobenzoyl)pyrrolidine
mp : 67°C
NMR (CDCl₃, δ) : 1.85-2.10 (4H, m), 3.45 (2H, t, J=6Hz), 3.69 (2H, t, J=6Hz), 7.61 (1H, t, J=8Hz), 7.89 (1H, dif-dd, J=8Hz), 8.29 (1H, dif-dd, J=8Hz), 8.40 (1H, dif-dd)

Preparation 37
To a stirred solution of 3-nitro-N-(3-pyridylmethyl)benzamide (2.00 g) in tetrahydrofuran (40 ml) was added potassium tert-butoxide (917 mg) in one portion in an ice-cooled bath. The stirring was continued for 40 minutes and then iodomethane (0.53 ml) was added thereto. The reaction mixture was stirred at 0°C for one hour, then at ambient temperature for five hours. Saturated sodium bicarbonate solution was added thereto and the mixture was extracted with ethyl acetate. The organic layer was washed with water and saturated sodium chloride solution. After dried over anhydrous magnesium sulfate and filtered, the solvent was removed in vacuo and the residue was purified by flash chromatography (methanol-chloroform 3%, V/V) to afford 3-nitro-N-methyl-N-(3-pyridylmethyl)benzamide (1.8 g) as a yellow oil.
NMR (CDCl₃, δ) : 2.80-3.22 (3H, m), 4.40-4.93 (2H,
m), 7.30-7.42 (1H, m), 7.44-7.90 (3H, m), 8.24-8.37 (2H, m), 8.40-8.75 (2H, m)

Preparation 38

Ethyl 4-[N-(3-pyridymethyl)acetamido]cinnamate was obtained by reacting ethyl 4-acetamidocinnamate with 3-pyridylmethyl chloride according to a similar manner to that of Preparation 37.

NMR (CDCl₃, δ) : 1.34 (3H, t, J=7Hz), 1.92 (3H, s), 4.29 (2H, q, J=7Hz), 4.90 (2H, s), 6.41 (1H, d, J=15Hz), 7.02 (2H, d, J=7Hz), 7.24 (1H, m), 7.51 (2H, d, J=7Hz), 7.60-7.70 (2H), 8.38 (1H, br s), 8.51 (1H, d, J=3Hz)

Preparation 39

A mixture of N,N-bis(2-methoxyethyl)-3-nitrobenzamide (4.11 g) and palladium on charcoal (411 mg) in ethyl acetate (41 ml) was hydrogenated under 1 atmospheric pressure of hydrogen for 1 hour at ambient temperature. The catalyst was removed by filtration and washed with ethyl acetate, and the volatiles were removed in vacuo. The residue was purified with column chromatography eluting with dichloromethane-methanol to give 3-amino-N,N-bis(2-methoxyethyl)benzamide (3.62 g) as an oil.

NMR (CDCl₃, δ) : 3.19-3.86 (16H), 6.62-6.79 (3H), 7.16 (1H, dt, J=8, 1Hz)

Preparation 40

The following compounds were obtained according to a similar manner to that of Preparation 39.

(1) 3-Amino-N-methyl-N-(3-pyridymethyl)benzamide

NMR (CDCl₃, δ) : 2.87 (3H, br s), 3.75 (1H, or 2H, br s), 4.41-4.88 (2H, m), 6.55-6.84 (3H, m), 7.03-7.40 (2H, m), 7.42-7.84 (1H, m)
8.35-8.70 (2H, m)

(2) 3-Amino-N-(4-pyridyl)benzamide

mp : 232-234°C

NMR (DMSO-d$_6$, δ) : 5.39 (2H, br s), 6.79 (1H, br d, J=8Hz), 7.02-7.11 (2H), 7.19 (1H, t, J=8Hz), 7.78 (2H, d, J=7Hz), 8.46 (2H, d, J=7Hz)

(3) 1-(3-Aminobenzoyl)-4-methylpiperazine

mp : 114-116°C

NMR (CDCl$_3$, δ) : 2.28-2.60 (7H), 3.38-3.90 (6H), 6.68-6.79 (3H), 7.68 (1H, t, J=8Hz)

(4) 1-(3-Aminobenzoyl)pyrrolidine

NMR (CDCl$_3$, δ) : 1.75-2.05 (4H, m), 3.40 (2H, t, J=6Hz), 3.60 (2H, t, J=6Hz), 3.72 (2H, br s), 6.71 (1H, dif-ddd, J=8Hz), 6.78-6.89 (2H, m), 7.10 (1H, t, J=8Hz)

(5) 3-Amino-N-(3-pyridylmethyl)benzamide

NMR (CDCl$_3$-CD$_3$OD, δ) : 4.60 (2H, s), 6.82 (1H, dt, J=8, 1Hz), 7.10-7.39 (4H), 7.79 (1H, dt, J=9, 1Hz), 8.45 (1H, dd, J=5, 1Hz), 8.52 (1H, d, J=1Hz)

Preparation 41

To a stirred solution of 3-amino-N,N-bis(2-methoxyethyl)benzamide (1.01 g) in 1,4-dioxane (10 ml) was added 1N sodium hydroxide solution (5.2 ml) and phenyl chloroformate (0.55 ml) successively in an ice-cooled bath. The bath was removed and the reaction mixture was stirred vigorously for 1 hour, during which time phenyl chloroformate (0.25 ml) was further added. The mixture was extracted with dichloromethane and the organic layer was washed with water twice and brine, dried over
anhydrous magnesium sulfate, and evaporated in vacuo. The residue was crystallized from diisopropyl ether to give phenyl 3-[N,N-bis(2-methoxyethyl)carbamoyl]phenylcarbamate (1.30 g) as a colorless powder.

\[ \text{mp: 116-118°C} \]
\[ \text{NMR (CDCl}_3, \delta): 3.19-3.82 (14H), 7.10-7.57 (10H) \]

**Preparation 42**

The following compounds were obtained according to a similar manner to that of Preparation 41.

1. Phenyl 3-[N-(4-pyridyl)carbamoyl]phenylcarbamate
   - mp: 204-206°C
   - NMR (DMSO-d6, δ): 5.39 (1H, br s), 6.71-6.82 (2H), 7.02-7.33 (4H), 7.40-7.81 (4H), 8.09 (1H, br s), 8.41-8.51 (2H), 9.32 (1H, br s)

2. Phenyl 3-(4-methyl-1-piperazinylcarbonyl)phenylcarbamate
   - mp: 152-154°C
   - NMR (CDCl3, δ): 2.27-2.56 (7H), 3.38-3.91 (4H), 7.10-7.60 (9H)

3. Phenyl 3-(1-pyrrolidinylcarbonyl)phenylcarbamate
   - mp: 135-140°C
   - NMR (CDCl3, δ): 1.74-2.00 (4H, m), 3.45 (2H, t, J=6Hz), 3.63 (2H, t, J=6Hz), 7.07-7.53 (9H, m), 7.72 (1H, br s)

4. Phenyl 3-[N-methyl-N-(3-pyridylmethyl)carbamoyl]-phenylcarbamate
   - NMR (CDCl3, δ): 2.90-3.08 (3H), 4.58 (0.5H, br s), 4.76 (1.5H, br s), 7.15-7.80 (13H), 8.58 (1H, d, J=5Hz)
(5) Phenyl 3-[N-(3-pyridylmethyl)carbamoyl]-phenylcarbamate  
mp : 185-188°C  
NMR (CDCl₃-CD₃OD, δ) : 4.62 (2H, s), 7.11-7.49 (7H), 7.56 (1H, dt, J=8, 1Hz), 7.63-7.80 (2H), 7.84 (1H, t, J=1Hz), 8.49 (1H, dd, J=5, 1Hz), 8.55 (1H, d, J=1Hz)

(6) Phenyl 3-(N,N-dimethylamino)phenylcarbamate  
mp : 226-228°C  
NMR (CDCl₃, δ) : 2.93 (6H, s), 6.49 (1H, d, J=7Hz), 6.65 (1H, d, J=7Hz), 6.87 (1H, br s), 7.03 (1H, br s), 7.12-7.29 (4H), 7.32-7.42 (2H)

Preparation 43

(1) Ethyl 4-(phenoxycarbonylamino)cinnamate was obtained by reacting ethyl 4-aminocinnamate with phenyl chloroformate according to a similar manner to that of Preparation 42.  
mp : 136-138°C  
NMR (CDCl₃, δ) : 1.33 (3H, t, J=7Hz), 4.27 (2H, q, J=7Hz), 6.39 (1H, d, J=15Hz), 7.09 (1H, br s), 7.15-7.58 (9H), 7.65 (1H, d, J=15Hz)

(2) A solution of ethyl 4-(phenoxycarbonylamino)cinnamate (500 mg) 3-aminopyridine (154 mg) and triethylamine (325 mg) in N,N-dimethylformamide (5 ml) was stirred for 2 hours at 80°C. Water was added thereto, and the resulting precipitate was collected by filtration to give ethyl 4-[3-(3-pyridyl)ureido]cinnamate (307 mg) as a colorless powder.  
mp : 188-189°C  
NMR (DMSO-d₆, δ) : 1.26 (3H, t, J=7Hz), 4.19 (2H, q, J=7Hz), 6.50 (1H, d, J=15Hz), 7.34 (1H, dd, J=9, 5Hz), 7.46-7.72 (5H), 7.96 (1H, dt, J=9, 1Hz),
8.21 (1H, dd, J=9, 1Hz), 8.62 (1H, d, J=1Hz),
8.98 (1H, br s), 9.10 (1H, m)

Preparation 44
To a mixture of ethyl 4-aminocinnamate (300 mg),
triethylamine (167 mg) and dichloromethane (3 ml) was
added a solution of propionyl chloride (182 mg) in
dichloromethane (1 ml) in an ice-water bath, and the
mixture was stirred for 1 hour at the same temperature.
To the reaction mixture was added 4 drops of N,N-
dimethylpropanediamine, and the mixture was further
stirred for 5 minutes. The reaction mixture was washed
with water, dried over magnesium sulfate, and evaporated
in vacuo. The residue was crystallized from diisopropyl
ether to give ethyl 4-propionamidocinnamate (341 mg) as a
colorless powder.
mp : 138°C
NMR (CDCl₃, δ) : 1.26 (3H, t, J=8Hz), 1.34 (3H, t,
J=8Hz), 2.42 (2H, q, J=8Hz), 4.26 (2H, q,
J=8Hz), 6.37 (1H, d, J=16Hz), 7.21 (1H, br s),
7.49 (2H, d, J=8Hz), 7.58 (2H, d, J=8Hz), 7.68
(1H, d, J=16Hz)

Preparation 45
To a solution of ethyl 4-aminocinnamate (2.00 g) and
methoxyacetic acid (1.04 ml) in N,N-dimethylformamide (20
ml) were added 1-ethyl-3-(3-
dimethylaminopropyl)carbodiimide hydrochloride (2.61 g)
and 1-hydroxybenzotriazole (2.12 g) at ambient
temperature, and the mixture was stirred for 1 hour at the
same temperature. The reaction mixture was poured into
water, and extracted with dichloromethane. The organic
layer was washed with aqueous sodium bicarbonate solution
and water, dried over magnesium sulfate, and evaporated in
vacuo. The residue was crystallized from diisopropyl
ether to give ethyl 4-(methoxyacetamido)cinnamate (4 g) as a pale yellow powder.

mp : 92.2°C

NMR (CDCl₃, δ) : 1.34 (3H, t, J=7.5Hz), 3.52 (3H, s), 4.03 (2H, s), 4.26 (2H, q, J=7.5Hz), 6.88 (1H, d, J=16Hz), 7.50 (2H, d, J=9Hz), 7.62 (2H, d, J=9Hz), 7.65 (1H, d, J=16Hz), 8.34 (1H, br s)

Preparation 46

The following compounds were obtained according to similar manners to those of Preparations 44 or 45.

(1) Ethyl 4-(4-bromobutyramido)cinnamate

mp : 119-124°C

NMR (CDCl₃, δ) : 1.32 (3H, t, J=7.5Hz), 2.21 (2H, quint, J=6Hz), 2.59 (2H, t, J=6Hz), 3.66 (2H, t, J=6Hz), 4.25 (2H, q, J=7.5Hz), 6.34 (1H, d, J=16Hz), 7.47 (2H, d, J=8Hz), 7.55 (2H, d, J=8Hz), 7.61 (1H, d, J=16Hz)

(2) Ethyl 4-(methoxyacetamido)cinnamate

mp : 87-92°C

NMR (CDCl₃, δ) : 1.34 (3H, t, J=7.5Hz), 3.53 (3H, s), 4.03 (2H, s), 4.26 (2H, q, J=7.5Hz), 6.37 (1H, d, J=16Hz), 7.50 (2H, d, J=8Hz), 7.63 (2H, d, J=8Hz), 7.65 (1H, d, J=16Hz), 8.35 (1H, br s)

(3) Ethyl 4-(isonicotinoylamino)cinnamate

mp : 179-188°C

NMR (CDCl₃, δ) : 1.34 (3H, t, J=7.5Hz), 4.26 (2H, q, J=7.5Hz), 6.40 (1H, d, J=16Hz), 7.52 (2H, d, J=9Hz), 7.57 (1H, d, J=16Hz), 7.65-7.78 (4H), 8.19 (1H, br s), 8.31 (2H, dd, J=6, 0.5Hz)

(4) Ethyl 4-(morpholinocarbonylamino)cinnamate
mp : 170-173°C

NMR (CDCl₃, δ): 1.33 (3H, t, J=7Hz), 3.43-3.56 (4H), 3.70-3.81 (4H), 4.28 (2H, q, J=7Hz), 6.35 (1H, d, J=15Hz), 6.49 (1H, br s), 7.40 (2H, d, J=9Hz), 7.48 (2H, d, J=9Hz), 7.63 (1H, d, J=15Hz)

(5) Ethyl 4-((5-bromovaleramido)cinnamate

mp : 124.0°C-124.7°C

NMR (CDCl₃, δ): 1.33 (3H, t, J=7.5Hz), 1.79-2.06 (4H, m), 2.42 (2H, t, J=6Hz), 3.44 (2H, t, J=6Hz), 4.26 (2H, q, J=7.5Hz), 6.36 (1H, d, J=16Hz), 7.30 (1H, br s), 7.49 (2H, d, J=8Hz), 7.57 (2H, d, J=8Hz), 7.64 (1H, d, J=16Hz)

Preparation 47

To a solution of methyl 4-carboxycinnamate (160 mg) in methylene chloride was added methylamine hydrochloride (58 mg) and 1-ethyl-3-(3-dimethylaminopropyl)carbodiimide (140 mg) at ambient temperature, and the mixture was stirred for 2 hours. To this suspension was added 1-hydroxybenzotriazole (137 mg) and dimethylformamide (2 ml), and the mixture was stirred for 14 hours at same temperature. The reaction mixture was poured into water, and extracted with dichloromethane. The organic layer was washed with aqueous sodium bicarbonate solution and water, dried over magnesium sulfate, and evaporated in vacuo. The residue was crystallized from diisopropyl ether to give methyl 4-((methylcarbamoyl)cinnamate (82 mg) as a colorless powder.

mp : 210.5°C

NMR (DMSO-d₆, δ): 2.79 (3H, d, J=5Hz), 3.74 (3H, s), 6.74 (1H, d, J=16Hz), 7.69 (1H, d, J=16Hz), 7.80 (2H, d, J=8Hz), 7.87 (2H, d, J=8Hz), 8.51 (1H, q-like)
Preparation 48
To a stirred suspension of methyl 4-carboxycinnamate (400 mg) in thionyl chloride (1.4 ml) was added one drop of N,N-dimethylformamide. The mixture was refluxed for 20 minutes. The solvent was removed in vacuo. To the residue was added toluene (2 ml) and the mixture was evaporated in vacuo twice. The residue was dissolved with dichloromethane (4 ml), and 4-aminopyridine (201 mg) and triethylamine (0.81 ml) were added thereto in an ice-water bath. After 10 minutes the mixture was stirred at ambient temperature. After 3 hours, to the reaction mixture was added water and the mixture was extracted with dichloromethane-methanol (5:1, V/V). The organic layer was washed with saturated sodium bicarbonate solution, water and brine, and dried over anhydrous magnesium sulfate. The solvent was evaporated in vacuo. The residue was crystallized from ethyl acetate to give methyl 4-[N-(4-pyridyl)carbamoyl]cinnamate (555 mg) as a colorless powder.

mp : 209-211°C
NMR (DMSO-d$_6$, δ) : 3.76 (3H, s), 6.82 (1H, d, J=15Hz), 7.69-7.83 (3H), 7.92 (2H, d, J=9Hz), 8.01 (2H, d, J=9Hz), 8.50 (2H, d, J=7Hz)

Preparation 49
The following compounds were obtained according to similar manners to those of Preparations 47 or 48.

(1) Methyl 4-(N,N-dimethylcarbamoyl)cinnamate
mp : 130°C
NMR (CDCl$_3$, δ) : 3.00 (3H, s), 3.12 (3H, s), 3.83 (3H, s), 6.49 (1H, d, J=16Hz), 7.45 (2H, d, J=8Hz), 7.58 (2H, d, J=8Hz), 7.70 (1H, d, J=16Hz)
(2) Methyl 4-\([N-(2\text{-}\text{methoxyethyl})\text{carbamoyl}]\text{cinnamate}\)

mp : 122-124°C

NMR (CDCl\(_3\), \(\delta\)) : 3.40 (3H, s), 3.53-3.72 (4H), 3.83 (3H, s), 6.45-6.60 (3H), 7.58 (2H, d, \(J=8\text{Hz}\)), 7.71 (1H, d, \(J=15\text{Hz}\)), 7.80 (2H, d, \(J=8\text{Hz}\))

(3) Methyl 4-\([N,N\text{-}\text{bis}(2\text{-}\text{methoxyethyl})\text{carbamoyl}]\text{cinnamate}\)

NMR (CDCl\(_3\), \(\delta\)) : 3.21-3.86 (17H), 6.48 (1H, d, \(J=15\text{Hz}\)), 7.44 (1H, d, \(J=9\text{Hz}\)), 7.57 (1H, d, \(J=9\text{Hz}\)), 7.70 (1H, d, \(J=15\text{Hz}\))

Preparation 50

To a solution of ethyl 4-aminocinnamate (150 mg) in methylene chloride were added methyl isocyanate (0.06 ml) under nitrogen atmosphere at ambient temperature, and the mixture was stirred for 2 hours at the same temperature. The reaction mixture was poured into the mixture of ethyl acetate and water. The organic layer was washed with water twice, dried over magnesium sulfate, and concentrated in vacuo. The residue was crystallized from diisopropyl ether to give ethyl 4-(3-methylureido)cinnamate (136 mg).

mp : 166°C

NMR (DMSO-\(d_6\), \(\delta\)) : 1.25 (3H, t, \(J=7.5\text{Hz}\)), 2.64 (3H, d, \(J=5\text{Hz}\)), 4.17 (2H, q, \(J=7.5\text{Hz}\)), 6.12 (1H, q, \(J=5\text{Hz}\)), 6.43 (1H, d, \(J=16\text{Hz}\)), 7.45 (2H, d, \(J=8\text{Hz}\)), 7.56 (1H, d, \(J=16\text{Hz}\)), 7.59 (2H, d, \(J=8\text{Hz}\)), 8.81 (1H, s)

Preparation 51

To a stirred solution of ethyl 4-(4-bromobutyramido)cinnamate (420 mg) in N,N-dimethylformamide (5 ml) was added potassium carbonate (552 mg) at ambient temperature and the resulting mixture was warmed at 50°C for three hours. The reaction mixture
was diluted with ethyl acetate and washed with water and brine. The organic phase was dried over anhydrous magnesium sulfate, and concentrated in vacuo. The residue was purified by flash column chromatography eluting with chloroform to afford ethyl 4-(2-oxo-1-pyrrolidinyl)cinnamate (281 mg) as a pale yellow solid.

\[ \text{mp : } 134^\circ C \]

NMR (CDCl\(_3\), \(\delta\)):
- 1.34 (3H, t, \(J=7.7\) Hz), 2.19 (2H, quint, \(J=7.7\) Hz), 2.63 (2H, t, \(J=7.7\) Hz), 3.88 (2H, t, \(J=7.7\) Hz), 4.26 (2H, q, \(J=7.7\) Hz), 6.38 (1H, d, \(J=16\) Hz), 7.53 (2H, d, \(J=8\) Hz), 7.64 (1H, d, \(J=16\) Hz), 7.68 (2H, d, \(J=8\) Hz)

**Preparation 52**

Ethyl 4-(2-oxopiperidino)cinnamate was obtained according to a similar manner to that of Preparation 51.

\[ \text{mp : } 120.2^\circ C \]

NMR (CDCl\(_3\), \(\delta\)):
- 1.34 (3H, t, \(J=7.5\) Hz), 1.83-2.05 (4H, m), 2.56 (2H, dif-t), 3.65 (2H, dif-d), 4.26 (2H, q, \(J=7.5\) Hz), 6.40 (1H, d, \(J=16\) Hz), 7.29 (2H, d, \(J=8\) Hz), 7.54 (2H, d, \(J=8\) Hz), 7.65 (1H, d, \(J=16\) Hz)

**Preparation 53**

A solution of ethyl 4-aminocinnamate (1 g) and 2,5-dimethoxytetrahydrofuran (0.677 ml) in toluene (3 ml) and acetic acid (3 ml) was refluxed for 5 hours with removing methanol. After cooling, the mixture was washed with water twice and saturated sodium bicarbonate solution, dried over anhydrous magnesium sulfate. The solvent was evaporated in vacuo. The residue was purified with column chromatography eluting with n-hexane - ethyl acetate to give ethyl 4-(1-pyrolyl)cinnamate (740 mg) as colorless crystals.

\[ \text{mp : } 86-87^\circ C \]
NMR (CDCl₃, δ): 1.37 (3H, t, J=7Hz), 4.29 (2H, q, J=7Hz), 6.32-6.49 (2H), 7.12 (2H, t, J=1Hz), 7.41 (2H, d, J=9Hz), 7.60 (2H, d, J=9Hz), 7.69 (1H, d, J=16Hz)

Preparation 54

Ethyl 4-(N-methyl-2-methoxyacetamido)cinnamate was obtained according to a similar manner to that of Preparation 37.

NMR (CDCl₃, δ): 1.35 (3H, t, J=7.5Hz), 3.29 (3H, s), 3.35 (3H, s), 3.85 (2H, br s), 4.27 (2H, q, J=7.5Hz), 6.46 (1H, d, J=16Hz), 7.24 (2H, d, J=8Hz), 7.57 (2H, d, J=8Hz), 7.67 (1H, d, J=16Hz)

Preparation 55

To a solution of ethyl 4-propionamidocinnamate (160 mg) in ethanol (5 ml) was added 1N aqueous sodium hydroxide solution (1.5 ml) at ambient temperature. The mixture was stirred at same temperature for 14 hours, and then at 40°C for 2 hours. 1N-hydrochloric acid (1.5 ml) was added to the reaction mixture and evaporated in vacuo. The residue was diluted with 10% methanol-dichloromethane, washed with water, dried over magnesium sulfate and evaporated in vacuo. The residue was crystallized from diisopropyl ether to give 4-propionamidocinnamic acid (115 mg) as a colorless powder.

mp: 243°C

NMR (DMSO-d₆, δ): 1.08 (3H, t, J=8Hz), 2.34 (2H, q, J=8Hz), 6.39 (1H, d, J=16Hz), 7.51 (1H, d, J=16Hz), 7.62 (4H, s-like), 10.07 (1H, s)

Preparation 56

The following compounds were obtained according to a similar manner to that of Preparation 55.
(1) 4-(Methylcarbamoyl)cinnamic acid
mp: >250°C
NMR (DMSO-d$_6$, δ): 2.78 (3H, d, J=5Hz), 6.62 (1H, d, J=16Hz), 7.61 (1H, d, J=16Hz), 7.77 (2H, d, J=8Hz), 7.85 (2H, d, J=8Hz), 8.51 (1H, q-like)

(2) 4-(N,N-Dimethylcarbamoyl)cinnamic acid
mp: 82°C
NMR (DMSO-d$_6$, δ): 2.93 (3H, s), 2.97 (3H, s), 6.59 (1H, d, J=16Hz), 7.43 (2H, d, J=8Hz), 7.61 (1H, d, J=16Hz), 7.75 (2H, d, J=8Hz)

(3) 4-(3-Methylureido)cinnamic acid
mp: 234°C
NMR (DMSO-d$_6$, δ): 2.64 (3H, d, J=5Hz), 6.12 (1H, q, J=5Hz), 6.33 (1H, d, J=16Hz), 7.44 (2H, d, J=8Hz), 7.51 (1H, d, J=16Hz), 7.55 (2H, d, J=8Hz), 8.78 (1H, s)

(4) 4-[N-(2-Methoxyethyl)carbamoyl]cinnamic acid
mp: 207-209°C
NMR (DMSO-d$_6$, δ): 3.20-3.50 (7H), 6.63 (1H, d, J=15Hz), 7.62 (1H, d, J=15Hz), 7.79 (2H, d, J=8Hz), 7.89 (2H, d, J=8Hz), 8.61 (1H, br s)

(5) 4-[N,N-Bis(2-methoxyethyl)carbamoyl]cinnamic acid
NMR (CDCl$_3$, δ): 3.21-3.86 (17H), 6.48 (1H, d, J=15Hz), 7.44 (2H, d, J=9Hz), 7.57 (2H, d, J=9Hz), 7.70 (1H, d, J=15Hz)

(6) 4-[N-(4-Pyridyl)carbamoyl]cinnamic acid
mp: >250°C
NMR (DMSO-d$_6$, δ): 6.69 (1H, d, J=16Hz), 7.52-8.08 (7H), 8.49 (2H, d, J=6Hz)
(7) 4-(2-Oxo-1-pyrrolidinyl)cinnamic acid
   mp : >250°C
   NMR (DMSO-d$_6$, δ) : 2.06 (2H, quint, J=8Hz), 3.86 (2H, t, J=8Hz), 6.46 (1H, d, J=16Hz), 7.55 (1H, d, J=16Hz), 7.65-7.76 (4H, m)

(8) 4-(Methoxyacetamido)cinnamic acid
   mp : 201.5-229°C
   NMR (CDCl$_3$, δ) : 4.02 (2H, s), 6.43 (1H, d, J=16Hz), 7.52 (1H, d, J=16Hz), 7.63 (2H, d, J=8Hz), 7.74 (2H, d, J=8Hz), 9.97 (1H, s)

(9) 4-[N-(3-Pyridylmethyl)acetamido]cinnamic acid
   mp : 184-186°C
   NMR (DMSO-d$_6$, δ) : 1.90 (3H, s), 4.91 (2H, s), 6.52 (1H, d, J=15Hz), 7.21-7.39 (3H), 7.50-7.79 (4H), 8.39 (1H, d, J=2Hz), 8.43 (1H, dd, J=5, 2Hz)

(10) 4-[(Isonicotinoylamino)cinnamic acid
   mp : 283-290°C
   NMR (DMSO-d$_6$, δ) : 6.46 (1H, d, J=16.2Hz), 7.53 (1H, d, J=16.2Hz), 7.68 (2H, d, J=9Hz), 7.79-7.91 (4H), 8.80 (2H, dd, J=6, 0.5Hz)

(11) 4-[(3-(3-Pyridyl)ureido)cinnamic acid
   mp : 219-221°C
   NMR (DMSO-d$_6$, δ) : 6.40 (1H, d, J=15Hz), 7.37 (1H, dd, J=9, 5Hz), 7.47-7.70 (5H), 7.98 (1H, dt, J=9, 1Hz), 8.21 (1H, br d, J=5Hz), 8.62 (1H, d, J=1Hz), 9.03 (1H, s), 9.16 (1H, s)

(12) 4-(Morpholinocarbonylamino)cinnamic acid
   mp : 219-221°C
   NMR (DMSO-d$_6$, δ) : 3.39-3.49 (4H), 3.58-3.68 (4H), 6.37 (1H, d, J=15Hz), 7.46-7.60 (5H),
8.76 (1H, br s)

(13) 4-(2-Oxopiperidino)cinnamic acid

mp : 235.7-243.2°C

NMR (DMSO-d$_6$, δ) : 1.73-1.97 (4H, m), 2.39 (2H, dif-t), 3.61 (2H, dif-t), 6.51 (1H, d, J=16Hz), 7.34 (2H, d, J=8Hz), 7.59 (1H, d, J=16Hz), 7.70 (2H, d, J=8Hz)

(14) 4-(N-Methyl-2-methoxyacetamido)cinnamic acid

mp : 182.1°C

NMR (DMSO-d$_6$, δ) : 3.17 (3H, s), 3.20 (3H, s), 3.87 (2H, br s), 6.57 (1H, d, J=15Hz), 7.39 (2H, d, J=9Hz), 7.60 (1H, d, J=15Hz), 7.74 (2H, d, J=9Hz)

(15) 4-(1-Pyrrolyl)cinnamic acid

mp : 236-240°C

NMR (DMSO-d$_6$, δ) : 6.30 (2H, t, J=1Hz), 6.54 (1H, d, J=16Hz), 7.48 (2H, t, J=1Hz), 7.53-7.71 (3H), 7.80 (2H, d, J=9Hz)

Preparation 57

To a solution of ethyl 4-aminocinnamate (150 mg) and triethylamine (94 mg) in methylene chloride (3 ml) was added mesyl chloride (0.08 ml) under ice-cooling under nitrogen atmosphere, and the mixture was stirred at ambient temperature for 2 hours. The reaction mixture was poured into water, and extracted with methylene chloride twice. The combined organic layer was washed with water, dried over magnesium sulfate and concentrated to give a residue including ethyl 4-mesylaminocinnamate and ethyl 4-(N,N-dimesylamino)cinnamate. The residue was dissolved in ethanol, and 1N aqueous sodium hydroxide solution (1.5 ml) was added thereto at 40°C. The mixture was stirred at
ambient temperature for 2 days, and 1N hydrochloric acid (1.5 ml) was added thereto. The mixture was concentrated in vacuo, and the residue was partitioned between 10% methanol-methylene chloride and water. The organic layer was washed with water, dried over magnesium sulfate and concentrated in vacuo. The residue was purified by preparative thin-layer chromatography (methylene chloride - methanol, 10:1, V/V) to give 4-mesylaminocinnamic acid (49.3 mg).

mp : 218°C
NMR (DMSO-d$_6$, 5) : 3.05 (3H, s), 6.44 (1H, d, J=16Hz), 7.21 (2H, d, J=8Hz), 7.53 (1H, d, J=16Hz), 7.66 (2H, d, J=8Hz)

Preparation 58

4-(N,N-Dimethylcarbamoyl)benzaldehyde was obtained by reacting 4-formylbenzoic acid with dimethylamine hydrochloride according to a similar manner to that of Preparation 47.

mp : 60-67°C
NMR (CDCl$_3$, 8) : 2.97 (3H, s), 3.15 (3H, s), 7.59 (2H, d, J=7.5Hz), 7.94 (2H, d, J=7.5Hz), 10.3 (1H, s)

Preparation 59

A mixture of 2-acetylamino-5-formylpyridine (241 mg) and malonic acid (168 mg) in pyridine (0.12 ml) and ethanol (0.36 ml) was refluxed for 2 hours. After cooling the mixture, the precipitate was collected by filtration, and washed ethyl acetate to give (E)-3-(6-acetylamino-3-pyridyl)acrylic acid (248 mg) as a colorless powder.

mp : 291-292°C
NMR (DMSO-d$_6$, 8) : 2.10 (3H, s), 6.55 (1H, d, J=16Hz), 7.58 (1H, d, J=16Hz), 8.07-8.21 (2H), 8.59 (1H, br s)
Preparation 60

The following compounds were obtained according to a similar manner to that of Preparation 59.

(1) (E)-3-(6-Ethoxycarbonyl-3-pyridyl)acrylic acid (from ethyl 5-formyl-2-pyridinecarboxylate)
mp : 201-202°C
NMR (DMSO-d$_6$, δ) : 1.33 (3H, t, J=7Hz), 4.36 (2H, q, J=7Hz), 6.80 (1H, d, J=16Hz), 7.69 (1H, d, J=16Hz), 8.07 (1H, d, J=9Hz), 8.33 (1H, dd, J=9, 2Hz), 9.00 (1H, d, J=2Hz)

(2) 4-((N,N-Dimethylcarbamoyl)cinnamic acid
NMR (CDCl$_3$, δ) : 2.99 (3H, s), 3.11 (3H, s), 6.49 (1H, d, J=15Hz), 7.46 (2H, d, J=8Hz), 7.59 (2H, d, J=8Hz), 7.76 (1H, d, J=15Hz)

Preparation 61

To a suspension of sodium hydride (60% active, 124 mg) in dimethylformamide (2 ml) at ambient temperature was added ethyl 4-hydroxycinnamate (500 mg) under nitrogen, and the mixture was stirred for 1 hour. 2-Bromoethyl acetate (522 mg) was added to the mixture at same temperature, and the mixture was allowed to stand for 19 hours. The reaction mixture was poured into water and extracted with ethyl acetate. The organic layer was washed with water and brine, dried over magnesium sulfate and evaporated in vacuo. The residue was chromatographed on silica gel eluting with chloroform to give ethyl 4-(2-acetoxyethoxy)cinnamate (716 mg) as an oil.

NMR (CDCl$_3$, δ) : 1.33 (3H, t, J=7.5Hz), 2.11 (3H, s), 4.19 (2H, t, J=6Hz), 4.25 (2H, q, J=7.5Hz), 4.44 (2H, t, J=6Hz), 6.31 (1H, d, J=16Hz), 6.94 (2H, d, J=8Hz), 7.49 (2H, d, J=8Hz), 7.64 (1H, d, J=16Hz)
Preparation 62

4-(2-Hydroxyethoxy)cinnamic acid was obtained from ethyl 4-(2-acetoxyethoxy)cinnamate according to a similar manner to that of Preparation 55.

mp : 194°C

NMR (DMSO-d$_6$, $\delta$): 3.64-3.79 (2H, br peak), 4.02 (2H, t, $J=6$Hz), 4.90 (1H, br peak), 6.37 (1H, d, $J=16$Hz), 6.98 (2H, d, $J=8$Hz), 7.54 (1H, d, $J=16$Hz), 7.63 (2H, d, $J=8$Hz)
Example 74

The following compounds were obtained according to similar manners to those of Examples 53 or 69.

5  (1) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-
[(thiomorpholinoacetyl)glycyl]amino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.44 (3H, s), 2.66-2.88 (8H), 3.01 (2H, s), 3.28 (3H, s), 3.55 (1H, dd, J=18, 5Hz), 3.82 (1H, dd, J=18, 5Hz), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.50 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz), 7.90 (1H, br s)

15  (2) 3-Bromo-8-[2,6-dichloro-3-[N-[N,N-dimethyl-β-
alanyl]glycyl]N-methylamino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.39-2.56 (11H), 2.78 (2H, t, J=6Hz), 3.24 (3H, s), 3.55 (1H, dd, J=17, 4Hz), 3.82 (1H, dd, J=17, 5Hz), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.34 (1H, d, J=9Hz), 7.49 (1H, d, J=9Hz), 7.79 (1H, d, J=7Hz), 8.49 (1H, br s)

Example 75

The following compounds were obtained according to similar manners to those of Examples 36 to 39.

(1) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(2-
pyrimidinylthio)acetyl]glycyl]amino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine

mp : 212-212.5°C

NMR (CDCl₃, δ) : 2.42 (3H, s), 3.21 (3H, s), 3.52 (1H, dd, J=18, 5Hz), 3.70-3.89 (3H), 5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.07 (1H, t, J=5Hz), 7.28 (1H, d, J=9Hz), 7.47
(1H, d, J=9Hz), 7.77 (1H, d, J=7Hz), 8.03 (1H, br s), 8.62 (2H, d, J=5Hz)

(2) 3-Bromo-8-\{2,6-dichloro-3-\{N-methyl-N-[\(\text{phenoxyacetyl}\)glycyl]amino\}benzyloxy\}-2-methylimidazo[1,2-a]pyridine

NMR (CDCl\textsubscript{3}, δ): 2.43 (3H, s), 3.28 (3H, s), 3.62 (1H, dd, J=17, 5Hz), 3.89 (1H, dd, J=17, 5Hz), 4.50 (2H, s), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.80-7.09 (4H), 7.22-7.39 (3H), 7.46-7.60 (2H), 7.78 (1H, d, J=7Hz)

(3) 3-Bromo-8-\{2,6-dichloro-3-\{N-\{heptafluorobutanoyl\}glycyl\}N-methylamino\}benzyloxy\}-2-methylimidazo[1,2-a]pyridine

NMR (CDCl\textsubscript{3}, δ): 0.88 (3H, t, J=7Hz), 1.16-1.40 (6H), 1.50-1.77 (2H), 2.21 (2H, t, J=7Hz), 2.42 (3H, s), 3.25 (3H, s), 3.52 (1H, dd, J=18, 4Hz), 3.80 (1H, dd, J=18, 5Hz), 5.49 (2H, s), 6.41 (1H, br t, J=5Hz), 6.72 (1H, d, J=7Hz), 6.85 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(4) 3-Bromo-8-\{2,6-dichloro-3-\{N-\{heptanoylglycyl\}N-methylamino\}benzyloxy\}-2-methylimidazo[1,2-a]pyridine

NMR (CDCl\textsubscript{3}, δ): 0.88 (3H, t, J=7Hz), 1.16-1.40 (6H), 1.50-1.77 (2H), 2.21 (2H, t, J=7Hz), 2.42 (3H, s), 3.25 (3H, s), 3.52 (1H, dd, J=18, 4Hz), 3.80 (1H, dd, J=18, 5Hz), 5.49 (2H, s), 6.41 (1H, br t, J=5Hz), 6.72 (1H, d, J=7Hz), 6.85 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(5) 3-Bromo-8-\{2,6-dichloro-3-\{N-methyl-N-(cinnamoylglycyl)amino\}benzyloxy\}-2-methylimidazo[1,2-a]pyridine

NMR (CDCl\textsubscript{3}, δ): 2.43 (3H, s), 3.29 (3H, s), 3.69
(1H, dd, J=17, 5Hz), 3.92 (1H, dd, J=17, 5Hz),
5.50 (2H, s), 6.49 (1H, d, J=15Hz), 6.62 (1H, br
s), 6.72 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz),
7.30-7.64 (8H), 7.78 (1H, d, J=7Hz)

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(6) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(trans-3-
pentenoyl)glycyl]amino]benzyloxy]-2-
methyllumidazo[1,2-a]pyridine
NMR (CDCl3, δ): 1.72 (3H, d, J=5Hz), 2.43 (3H, s),
2.95 (2H, d, J=5Hz), 3.25 (3H, s), 3.51 (1H, dd,
J=18, 4Hz), 3.79 (1H, dd, J=18, 5Hz), 5.43-5.79
(4H), 6.60 (1H, br s), 6.71 (1H, d, J=7Hz), 6.87
(1H, t, J=7Hz), 7.30 (1H, d, J=9Hz), 7.49 (1H,
d, J=9Hz), 7.78 (1H, d, J=7Hz)

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(7) 3-Bromo-8-[3-[N-[(3-butenoyl)glycyl]-N-methylamino]-
2,6-dichlorobenzyloxy]-2-methyllumidazo[1,2-a]
pyridine
NMR (CDCl3, δ): 2.43 (3H, s), 3.02 (2H, d, J=7Hz),
3.25 (3H, s), 3.52 (1H, dd, J=18, 4Hz), 3.79
(1H, dd, J=18, 5Hz), 5.19-5.32 (2H), 5.49 (2H,
s), 6.94 (1H, m), 6.59 (1H, br s), 6.72 (1H, d,
J=7Hz), 6.87 (1H, t, J=7Hz), 7.30 (1H, d,
J=9Hz), 7.48 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

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(8) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(4-
phenylbutanoyl)glycyl]amino]benzyloxy]-2-
methyllumidazo[1,2-a]pyridine
NMR (CDCl3, δ): 1.86-2.06 (2H), 2.22 (2H, t,
J=8Hz), 2.42 (3H, s), 2.63 (2H, t, J=8Hz), 3.25
(3H, s), 3.51 (1H, dd, J=17, 5Hz), 3.79 (1H, dd,
J=17, 5Hz), 5.49 (2H, s), 6.39 (1H, br s), 6.71
(1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.11-7.34
(6H), 7.48 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

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(9) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-
...
(methylcarbamoyl)cinnamoylglycyl]amino)benzyloxy)-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 2.45 (3H, s), 3.03 (3H, d, $J$=5Hz), 3.28 (3H, s), 3.66 (1H, dd, $J$=4, 18Hz), 3.92 (1H, dd, $J$=4, 18Hz), 5.46 (1H, d, $J$=10Hz), 5.53 (1H, d, $J$=10Hz), 6.16 (1H, q-like), 6.53 (1H, d, $J$=16Hz), 6.62-6.78 (2H, m), 6.85 (1H, t, $J$=7.5Hz), 7.33 (1H, d, $J$=8Hz), 7.45-7.68 (4H, m), 7.70-7.82 (3H, m)

(10) 3-Bromo-8-[2,6-dichloro-3-[N-[4-[(dimethylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 2.45 (3H, s), 2.99 (3H, br s), 3.10 (3H, br s), 3.29 (3H, s), 3.69 (1H, dd, $J$=17, 4Hz), 3.91 (1H, dd, $J$=17, 5Hz), 5.47 (1H, d, $J$=10Hz), 5.52 (1H, d, $J$=10Hz), 6.50 (1H, d, $J$=15Hz), 6.65 (1H, br t, $J$=4Hz), 6.71 (1H, d, $J$=7Hz), 6.86 (1H, t, $J$=7Hz), 7.29-7.62 (7H), 7.78 (1H, d, $J$=7Hz)

(11) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(2-methoxyethylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 2.44 (3H, s), 3.29 (3H, s), 3.40 (3H, s), 3.52-3.77 (5H), 3.91 (1H, dd, $J$=17, 5Hz), 5.50 (2H, s), 6.48-6.61 (2H), 6.63-6.79 (2H), 6.88 (1H, t, $J$=7Hz), 7.33 (1H, d, $J$=9Hz), 7.46-7.65 (4H), 7.72-7.83 (3H)

(12) 8-[3-[N-[4-[N,N-Bis(2-methoxyethyl)carbamoyl]-cinnamoylglycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 2.44 (3H, s), 3.21-3.82 (15H), 3.92
(1H, dd, J=17, 5Hz), 5.48 (1H, d, J=10Hz), 5.56 (1H, d, J=10Hz), 6.50 (1H, d, J=15Hz), 6.65 (1H, br t, J=4Hz), 6.73 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.30-7.62 (7H), 7.78 (1H, d, J=7Hz)

(13) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(4-pyridylcarbamoyl)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.40 (3H, s), 3.25 (3H, s), 3.65 (1H, dd, J=17, 4Hz), 3.90 (1H, dd, J=17, 5Hz), 5.42 (1H, d, J=10Hz), 5.51 (1H, d, J=10Hz), 6.51 (1H, d, J=15Hz), 6.69-6.80 (2H), 6.87 (1H, t, J=7Hz), 7.30 (1H, d, J=9Hz), 7.41-7.69 (6H), 7.7 (1H, d, J=7Hz), 7.87 (2H, d, J=8Hz), 8.43-8.59 (3H)

(14) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(2-oxo-1-pyrrolidinyl)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.15 (1H, quint, J=7.5Hz), 2.41 (3H, s), 2.63 (2H, t, J=7.5Hz), 3.26 (3H, s), 3.65 (1H, dd, J=4, 18Hz), 3.80-3.99 (3H, m), 5.43-5.57 (2H, m), 6.43 (1H, d, J=16Hz), 6.60 (1H, t-like), 6.73 (1H, d, J=8Hz), 6.85 (1H, t, J=8Hz), 7.32 (1H, d, J=8Hz), 7.45-7.61 (4H, m), 7.67 (2H, d, J=8Hz), 7.78 (1H, d, J=6Hz)

(15) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(methoxyacetamido)-cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.43 (3H, s), 3.26 (3H, s), 3.50 (3H, s), 3.65 (1H, dd, J=4, 18Hz), 3.90 (1H, dd, J=4, 18Hz), 4.01 (2H, s), 5.42-5.57 (2H, m), 6.40 (1H, d, J=16Hz), 6.59 (1H, t-like), 6.72 (1H, d, J=7.5Hz), 6.85 (1H, t, J=7.5Hz), 7.32
(16) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-
(propionamido)cinnamoylglycyl]amino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, δ): 1.24 (3H, t, J=7.5Hz), 2.39 (2H, q,
J=7.5Hz), 2.43 (3H, s), 3.26 (3H, s), 3.66 (1H, dd,
J=16, 5Hz), 3.88 (1H, dd, J=16, 6Hz), 5.45
(1H, d, J=9Hz), 5.50 (1H, d, J=9Hz), 6.39 (1H,
d, J=15Hz), 6.60 (1H, br t, J=5Hz), 6.73 (1H, d,
J=7Hz), 6.86 (1H, t, J=7Hz), 7.32 (1H, d,
J=9Hz), 7.40-7.60 (6H, m), 7.76 (1H, d, J=7Hz)

(17) 8-[3-[N-[4-(Acetamido)cinnamoylglycyl]-N-
methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-
methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, δ): 2.18 (3H, br s), 2.43 (3H, s), 3.27
(3H, s), 3.66 (1H, dd, J=4, 18Hz), 3.90 (1H, dd,
J=4, 18Hz), 5.41-5.55 (2H, m), 6.40 (1H, d,
J=16Hz), 6.59 (1H, t-like), 6.73 (1H, d, J=7Hz),
6.87 (1H, t, J=7Hz), 7.33 (1H, d, J=8Hz), 7.39-
7.61 (7H, m), 7.78 (1H, d, J=6Hz)

(18) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(N-methylacetamido)-
cinnamoylglycyl]-N-methylamino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, δ): 1.91 (3H, br s), 2.43 (3H, s), 3.29
(6H, s), 3.68 (1H, dd, J=17, 4Hz), 3.92 (1H, dd,
J=17, 5Hz), 5.47 (1H, d, J=10Hz), 5.53 (1H, d,
J=10Hz), 6.49 (1H, d, J=15Hz), 6.65 (1H, br t,
J=4Hz), 6.72 (1H, d, J=7Hz), 6.88 (1H, t,
J=7Hz), 7.15-7.39 (3H), 7.48-7.62 (4H), 7.79
(1H, d, J=7Hz)
(19) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-[N-(3-
pyridylmethyl)acetamido]cinnamoylglycyl]amino]-
benzyl oxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 1.91 (3H, s), 2.42 (3H, s), 3.29
(3H, s), 3.67 (1H, dd, J=17, 4Hz), 3.91 (1H, dd,
J=17, 5Hz), 4.90 (2H, s), 5.48 (1H, d, J=10Hz),
5.53 (1H, d, J=10Hz), 6.46 (1H, d, J=15Hz), 6.67
(1H, br t, J=4Hz), 6.72 (1H, d, J=7Hz), 6.88
(1H, t, J=7Hz), 7.01 (2H, d, J=8Hz), 7.19-7.68
(7H), 7.79 (1H, d, J=7Hz), 7.39 (1H, d, J=1Hz),
8.51 (1H, dd, J=5, 1Hz)

(20) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-
(isonicotinoylamino)cinnamoylglycyl]amino]benzyl oxy]-
2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.45 (3H, s), 3.29 (3H, s), 3.69
(1H, dd, J=17Hz), 3.92 (1H, d, J=17Hz), 5.45-
5.58 (2H), 6.47 (1H, d, J=15Hz), 6.65 (1H, br
s), 6.78 (1H, d, J=7Hz), 6.90 (1H, t, J=7Hz),
7.31-7.83 (10H), 8.71 (1H, s), 8.82 (2H, d;
J=7Hz)

(21) 8-[3-[N-(4-Aminocinnamoylglycyl)-N-methylamino]-2,6-
dichlorobenzyl oxy]-3-bromo-2-methylimidazo[1,2-
a]pyridine

NMR (CDCl₃, δ) : 2.43 (3H, s), 3.26 (3H, s), 3.66
(1H, dd, J=18, 4Hz), 3.66 (2H, br s), 5.45 (1H, d,
J=9Hz), 5.51 (1H, d, J=9Hz), 6.26 (1H, d,
J=16Hz), 6.50 (1H, br t, J=5Hz), 6.63 (2H, d,
J=9Hz), 6.72 (1H, d, J=8Hz), 6.87 (1H, t,
J=8Hz), 7.31 (2H, d, J=9Hz), 7.46 (1H, d,
J=5Hz), 7.47 (1H, d, J=16Hz), 7.51 (1H, d,
J=5Hz), 7.77 (1H, d, J=8Hz)

(22) 3-Bromo-8-[2,6-dichloro-3-[N-[4-
(methanesulfonamido)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 2.43 (3H, s), 3.03 (3H, s), 3.27 (3H, s), 3.65 (1H, dd, $J=4$, 7Hz), 3.91 (1H, dd, $J=4$, 17Hz), 5.42-5.58 (2H, m), 6.40 (1H, d, $J=16$Hz), 6.55-6.77 (3H, m), 6.86 (1H, dd, $J=8$, 6Hz), 7.13-7.30 (2H, m), 7.33 (1H, d, $J=8$Hz), 7.42-7.59 (4H, m), 7.78 (1H, d, $J=6$Hz)

(23) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(3-methoxycarbonyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 2.42 (3H, s), 2.80 (3H, d, $J=5$Hz), 3.23 (3H, s), 3.65 (1H, dd, $J=4$, 18Hz), 3.88 (1H, dd, $J=4$, 18Hz), 5.05-5.18 (1H, m), 5.39-5.53 (2H, m), 6.34 (1H, d, $J=16$Hz), 6.63-6.77 (2H, m), 6.88 (1H, t, $J=8$Hz), 7.23-7.57 (8H, m), 7.77 (1H, d, $J=8$Hz)

(24) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(methoxycarbonyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 2.43 (3H, s), 3.28 (3H, s), 3.69 (1H, dd, $J=17$, 4Hz), 3.91 (1H, dd, $J=17$, 5Hz), 3.93 (3H, s), 5.46 (1H, d, $J=9$Hz), 5.51 (1H, d, $J=9$Hz), 6.56 (1H, d, $J=16$Hz), 6.70 (1H, m), 6.73 (1H, d, $J=8$Hz), 6.86 (1H, t, $J=8$Hz), 7.34 (1H, d, $J=9$Hz), 7.50 (1H, d, $J=9$Hz), 7.50 (1H, d, $J=9$Hz), 7.56 (2H, d, $J=9$Hz), 7.61 (1H, d, $J=16$Hz), 7.77 (1H, d, $J=8$Hz), 8.02 (2H, d, $J=9$Hz)

(25) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-[3-(3-pyridyl)ureido]cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃-CD₃OD, δ): 2.41 (3H, s), 3.25 (3H, s), 3.63 (1H, d, J=18Hz), 3.93 (1H, d, J=18Hz), 5.50 (2H, s), 6.41 (1H, d, J=15Hz), 6.77 (1H, d, J=7Hz), 6.90 (1H, t, J=7Hz), 7.22-7.59 (8H), 7.80 (1H, d, J=7Hz), 8.14-8.32 (3H)

(26) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(morpholinocarbonylamino)cinnamoylglycyl]amino]-benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 2.42 (3H, s), 3.26 (3H, s), 3.43-3.56 (4H), 3.59-3.80 (5H), 3.91 (1H, dd, J=17, 5Hz), 5.50 (2H, s), 6.38 (1H, d, J=15Hz), 6.51-6.62 (2H), 6.72 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.24-7.59 (7H), 7.78 (1H, d, J=7Hz)

(27) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(2-oxopiperidino)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 1.86-2.03 (4H, m), 2.44 (3H, s), 2.57 (3H, dif-t), 3.28 (3H, s), 3.57-3.75 (3H, m), 3.91 (1H, dd, J=4, 18Hz), 5.43-5.56 (2H, m), 6.43 (1H, d, J=16Hz), 6.61 (1H, t-like), 6.71 (1H, d, J=7Hz), 6.85 (1H, t, J=7Hz), 7.21-7.38 (3H, m), 7.44-7.63 (4H, m), 7.78 (1H, d, J=6Hz)

(28) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-[(N-methyl-2-methoxyacetamido)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 2.43 (3H, s), 3.28 (3H, s), 3.30 (3H, s), 3.36 (3H, s), 3.68 (1H, dd, J=18, 4Hz), 3.83 (2H, br s), 3.91 (1H, dd, J=18, 5Hz), 5.48 (1H, d, J=9Hz), 5.52 (1H, d, J=9Hz), 6.48 (1H, d, J=15Hz), 6.66 (1H, br t, J=4Hz), 6.73 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.20 (2H, d, J=9Hz), 7.34 (1H, d, J=7Hz), 7.50 (1H, d,
\[
\begin{align*}
J &= 7\text{Hz}, \quad 7.55 \ (2\text{H}, \ d, \ J=9\text{Hz}), \quad 7.59 \ (1\text{H}, \ d, \\
& \quad \ J=15\text{Hz}), \quad 7.78 \ (1\text{H}, \ d, \ J=7\text{Hz}) \\
\end{align*}
\]

(29) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(1-pyrrolyl)cinnamoylglycyl]amino]benzyl]oxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl\textsubscript{3}, \(\delta\)) : 2.43 (3H, s), 3.29 (3H, s), 3.69 (1H, dd, \(J=18, 4\text{Hz}\)), 3.92 (1H, dd, \(J=18, 5\text{Hz}\)), 5.50 (2H, s), 6.38 (2H, t, \(J=1\text{Hz}\)), 6.47 (1H, d, \(J=16\text{Hz}\)), 6.65 (1H, br t, \(J=4\text{Hz}\)), 6.72 (1H, d, \(J=7\text{Hz}\)), 6.88 (1H, t, \(J=7\text{Hz}\)), 7.12 (2H, t, \(J=1\text{Hz}\)), 7.30-7.64 (7H), 7.78 (1H, d, \(J=7\text{Hz}\))

(30) 3-Chloro-8-[2,6-dichloro-3-[N-[4-(dimethylcarbamoyl)-cinnamoylglycyl]N-methylamino]benzyl]oxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl\textsubscript{3}, \(\delta\)) : 2.41 (3H, s), 3.00 (3H, br s), 3.12 (3H, br s), 3.29 (3H, s), 3.69 (1H, dd, \(J=17, 4\text{Hz}\)), 3.92 (1H, dd, \(J=17, 5\text{Hz}\)), 5.47 (1H, d, \(J=10\text{Hz}\)), 5.53 (1H, d, \(J=10\text{Hz}\)), 6.50 (1H, d, \(J=16\text{Hz}\)), 6.61-6.78 (2H), 6.88 (1H, t, \(J=7\text{Hz}\)), 7.30-7.64 (7H), 7.73 (1H, d, \(J=7\text{Hz}\))

(31) 3-Chloro-8-[2,6-dichloro-3-[N-methyl-N-[4-(methylcarbamoyl)cinnamoylglycyl]amino]benzyl]oxy]-2-methylimidazo[1,2-a]pyridine

NMR (DMSO-d\textsubscript{6}, \(\delta\)) : 2.30 (3H, s), 2.79 (3H, d, \(J=5\text{Hz}\)), 3.17 (3H, s), 3.53 (1H, dd, \(J=4, 18\text{Hz}\)), 3.81 (1H, dd, \(J=4, 18\text{Hz}\)), 5.48 (2H, s), 6.86 (1H, d, \(J=16\text{Hz}\)), 6.93-7.05 (2H, m), 7.42 (1H, d, \(J=16\text{Hz}\)), 7.58-7.70 (2H, m), 7.75-7.90 (4H, m), 7.90-8.00 (1H, m), 8.35 (1H, t-like), 8.50 (1H, q-like)

(32) 3-Chloro-8-[2,6-dichloro-3-[N-[4-(2-
methoxyethylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 2.42 (3H, s), 3.28 (3H, s), 3.40 (3H, s), 3.54-3.60 (2H), 3.63-3.73 (3H), 3.91 (1H, dd, J=18, 5Hz), 5.49 (1H, d, J=10Hz), 5.52 (1H, d, J=10Hz), 6.50-6.60 (2H), 6.68-6.74 (2H), 6.87 (1H, t, J=7.5Hz), 7.34 (1H, d, J=8Hz), 7.50 (1H, d, J=8Hz), 7.52-7.63 (3H), 7.72 (1H, d, J=7.5Hz), 7.79 (2H, d, J=8Hz)

(33) 3-Chloro-8-[2,6-dichloro-3-[N-[4-(methoxycarbonyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 2.44 (3H, s), 3.29 (3H, s), 3.68 (1H, dd, J=4, 18Hz), 3.85-3.96 (4H, m), 5.49 (1H, d, J=10Hz), 5.63 (1H, d, J=10Hz), 6.57 (1H, d, J=16Hz), 6.66-6.74 (2H, m), 6.86 (1H, t, J=7.7Hz), 7.34 (1H, d, J=8Hz), 7.50 (1H, d, J=8Hz), 7.53-7.64 (3H, m), 7.73 (1H, d, J=6Hz), 7.99-8.06 (2H, m)

(34) 3-Chloro-8-[2,6-dichloro-3-[N-methyl-N-[4-(2-oxo-1-pyrrolidinyl)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 2.19 (2H, quint, J=8Hz), 2.44 (3H, s), 2.63 (3H, t, J=8Hz), 3.29 (3H, s), 3.66 (1H, dd, J=4, 17.5Hz), 3.81-3.99 (3H, m), 5.43-5.58 (2H, m), 6.42 (1H, d, J=16Hz), 6.60 (1H, t-like), 6.70 (1H, d, J=7.5Hz), 6.86 (1H, t, J=7.5Hz), 7.33 (1H, d, J=8Hz), 7.45-7.60 (4H, m), 7.60-7.76 (3H, m)

(35) 8-[3-[N-[4-(Acetamido)cinnamoylglycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-chloro-2-
methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.18 (3H, s), 2.42 (3H, s), 3.26 (3H, s), 3.65 (1H, dd, J=4, 18Hz), 3.90 (1H, dd, J=4, 18Hz), 5.41-5.56 (2H, m), 6.39 (16H, d),
6.59 (1H, t-like), 6.70 (7.5H, d), 6.85 (7.5H, t), 7.31 (8H, d), 7.39-7.59 (7H, m), 7.72 (6H, d)

(36) 3-Chloro-8-[2,6-dichloro-3-[N-methyl-N-[4-(3-methylureido)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.40 (3H, s), 2.78 (3H, d, J=4Hz),
3.21 (3H, s), 3.77 (1H, dd, J=18, 4Hz), 3.87 (1H, dd, J=18, 5Hz), 5.35 (1H, br d, J=4Hz),
5.41 (1H, d, J=10Hz), 5.49 (1H, d, J=10Hz), 6.33 (1H, br t, J=16Hz), 6.71 (1H, d, J=7.5Hz), 6.79 (1H, br t, J=7.5Hz), 7.26-7.37 (5H), 7.40 (1H, d, J=8Hz), 7.49 (1H, d, J=16Hz), 7.67 (1H, br s), 7.74 (1H, d, J=7.5Hz)

(37) 3-Chloro-8-[2,6-dichloro-3-[N-[4-(methoxyacetamido)cinnamoylglycyl]N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.43 (3H, s), 3.29 (3H, s), 3.51 (3H, s), 3.68 (1H, dd, J=18, 4Hz), 3.91 (1H, dd, J=18, 5Hz), 4.02 (2H, s), 5.48 (1H, d, J=10Hz),
5.52 (1H, d, J=10Hz), 6.41 (1H, d, J=15Hz), 6.60 (1H, br s), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.32 (1H, d, J=8Hz), 7.46-7.64 (6H),
7.72 (1H, d, J=7Hz), 8.32 (1H, br s)

(38) 3-Chloro-8-[2,6-dichloro-3-[N-methyl-N-[4-(propionamido)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

30

35
NMR (CDCl₃, δ) : 1.25 (3H, t, J=7.5Hz), 3.34-3.45 (5H, m), 3.26 (3H, s), 3.66 (1H, dd, J=4, 18Hz), 3.90 (1H, dd, J=4, 18Hz), 5.43-5.55 (2H, m), 6.40 (1H, d, J=16Hz), 6.69 (1H, t-like), 6.71 (1H, d, J=7.5Hz), 7.30-7.36 (2H, m), 7.43-7.57 (5H, m), 7.72 (1H, d, J=6Hz)

(39) 8-[3-[N-(4-Aminocinnamoylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-3-chloro-2-methylimidazo-[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.43 (3H, s), 3.26 (3H, s), 3.65 (1H, dd, J=4, 18Hz), 3.83-3.95 (3H, m), 5.42-5.54 (2H, m), 6.25 (1H, d, J=16Hz), 6.50 (1H, t-like), 6.63 (2H, d, J=8Hz), 6.70 (1H, d, J=7.7Hz), 6.85 (1H, t, J=7.7Hz), 7.29-7.36 (3H, m), 7.43-7.52 (2H, m), 7.71 (1H, d, J=6Hz)

(40) 3-Bromo-8-[2,6-dichloro-3-[N-[(E)-3-(6-ethoxycarbonyl-3-pyridyl)acryloylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 1.47 (3H, t, J=7Hz), 2.44 (3H, s), 3.29 (3H, s), 3.70 (1H, dd, J=18, 4Hz), 3.92 (1H, dd, J=18, 5Hz), 4.50 (2H, q, J=7Hz), 5.51 (2H, s), 6.64 (1H, d, J=16Hz), 6.70-6.80 (2H), 6.88 (1H, t, J=7Hz), 7.35 (1H, d, J=9Hz), 7.51 (1H, d, J=9Hz), 7.62 (1H, d, J=16Hz), 7.78 (1H, d, J=7Hz), 7.94 (1H, dd, J=8, 1Hz), 8.16 (1H, d, J=8Hz), 8.88 (1H, d, J=1Hz)

(41) 8-[3-[N-[(E)-3-(6-Acetylamino-3-pyridyl)acryloylglycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.21 (3H, s), 2.44 (3H, s), 3.29 (3H, s), 3.69 (1H, dd, J=18, 4Hz), 3.92 (1H, dd, J=18, 5Hz), 5.50 (2H, s), 6.48 (1H, d, J=16Hz),
6.65-6.78 (2H), 6.87 (1H, t, J=7Hz), 7.35 (1H, d, J=9Hz), 7.50 (1H, d, J=9Hz), 7.55 (1H, d, J=16Hz), 7.78 (1H, d, J=7Hz), 7.85 (1H, dd, J=9, 1Hz), 8.10 (1H, br s), 8.22 (1H, d, J=9Hz), 8.37 (1H, d, J=1Hz)

(42) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(2-hydroxyethoxy)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 2.00 (1H, t, J=5Hz), 2.43 (3H, s), 3.27 (3H, s), 3.65 (1H, dd, J=4, 18Hz), 3.83-4.04 (3H, m), 4.11 (2H, t), 5.47 (1H, d, J=10Hz), 5.53 (1H, d, J=10Hz), 6.35 (1H, d, J=16Hz), 6.56 (1H, t-like), 6.71 (1H, d, J=7Hz), 6.79-6.97 (3H, m), 7.33 (1H, d, J=8Hz), 7.40-7.61 (4H, m), 7.77 (1H, d, J=6Hz)

(43) 3-Bromo-8-[2,6-dichloro-3-[N-[3,4-dimethoxycinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 2.44 (3H, s), 3.29 (3H, s), 3.67 (1H, dd, J=4, 18Hz), 3.84-4.00 (7H, m), 5.43-5.58 (2H, m), 6.37 (1H, d, J=16Hz), 6.59 (1H, t-like), 6.73 (1H, d, J=7.5Hz), 6.81-6.92 (2H, m), 7.00-7.15 (2H, m), 7.34 (1H, d, J=8Hz), 7.49 (1H, d, J=1Hz), 7.54 (1H, d, J=8Hz), 7.78 (1H, d, J=6Hz)

(44) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[3,4-(methyleneedioxy)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 2.44 (3H, s), 3.28 (3H, s), 3.65 (1H, dd, J=4, 18Hz), 3.92 (1H, dd, J=4, 18Hz), 5.42-5.57 (2H, m), 6.00 (2H, s), 6.30 (1H, d, J=16Hz), 6.57 (1H, t-like), 6.69-6.92 (3H, m),
6.92-7.04 (2H, m), 7.33 (2H, d, J=8Hz), 7.42-7.55 (2H, m), 7.78 (1H, d, J=6Hz)

(45) 3-Bromo-8-[2,6-dichloro-3-[N-{(1-indolylcarbonyl)glycyl}-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 2.44 (3H, s), 3.31 (3H, s), 3.79 (1H, dd, J=17, 4Hz), 4.01 (1H, dd, J=17, 5Hz), 5.51 (2H, s), 6.62 (1H, d, J=3Hz), 6.68-6.78 (2H), 6.88 (1H, t, J=7Hz), 7.19-7.62 (6H), 7.78 (1H, d, J=7Hz), 8.11 (1H, d, J=8Hz)

(46) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(morpholinocarbonyl)glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 2.42 (3H, s), 3.24 (3H, s), 3.30-3.34 (4H), 3.51 (1H, dd, J=17, 4Hz), 3.62-3.88 (5H), 5.40-5.60 (3H), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.32 (1H, d, J=9Hz), 7.48 (1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

**Example 76**

A mixture of 8-[3-(N-glycyl-N-methylamino)-2,6-dichlorobenzyloxy]-3-chloro-2-methylimidazo[1,2-a]pyridine (120 mg), phenyl 3-[N,N-bis(2-methoxyethyl)carbamoyl]-phenylcarbamate (110 mg) and triethylamine (57 mg) in N,N-dimethylformamide (1.2 ml) was stirred at 80°C for 1.5 hours. The mixture was extracted with dichloromethane and washed with water. After dried over magnesium sulfate, the solvent was removed in vacuo. The residue was purified by preparative thin layer chromatography eluting with dichloromethane-methanol to give 8-[3-[N-[N'-[3-[N,N-bis(2-methoxyethyl)carbamoyl]phenyl]ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-chloro-2-methylimidazo[1,2-a]pyridine (162 mg) as amorphous.
NMR (CDCl₃, δ): 2.41 (3H, s), 3.13-3.90 (19H), 5.47 (2H, s), 5.98 (1H, br t, J=4Hz), 6.71 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.00 (1H, br d, J=7Hz), 7.14-7.49 (5H), 7.72 (1H, d, J=7Hz), 7.89 (1H, br s)

Example 77

The following compounds were obtained according to similar manners to those of Examples 42, 43, 65 or 76.

(1) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-{(5-

-isoquinolyl)ureidoacetyl]-N-methylamino}benzyloxy]-2-

methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 2.40 (3H, s), 3.19 (3H, s), 3.71

(1H, dd, J=17, 5Hz), 3.91 (1H, dd, J=17, 5Hz),

5.46 (2H, s), 6.29 (1H, br t, J=5Hz), 6.71 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.41 (1H, d, J=9Hz), 7.54 (1H, t, J=7Hz), 7.66-7.81 (4H), 7.99 (1H, d, J=7Hz),

8.41 (1H, d, J=5Hz), 9.21 (1H, br s)

(2) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N-

'-{(3-

-pyrazolyl)ureidoacetyl]amino}benzyloxy]-2-

methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 2.44 (3H, s), 3.28 (3H, s), 3.68

(1H, dd, J=17.4Hz), 3.81-3.99 (2H), 5.50 (2H, s), 5.79 (1H, d, J=3Hz), 6.71 (1H, d, J=7Hz),

6.85 (1H, t, J=7Hz), 7.34 (1H, d, J=9Hz), 7.42-

7.60 (2H), 7.78 (1H, d, J=7Hz), 7.89 (1H, d, J=3Hz)

(3) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-{(4-

-pyrimidinyl)ureidoacetyl]amino}benzyloxy]-2-

methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 2.42 (3H, s), 3.28 (3H, s), 3.80
(1H, dd, J=17, 5Hz), 4.03 (1H, dd, J=17, 5Hz),
5.49 (2H, s), 6.71 (1H, d, J=7Hz), 6.81-6.95
(2H), 7.36 (1H, d, J=9Hz), 7.45 (1H, d, J=9Hz),
7.78 (1H, d, J=7Hz), 8.40 (1H, d, J=6Hz), 8.76-
8.85 (2H), 9.35 (1H, br s)

(4) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(6-quinolyl)ureidoacetyl]amino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.41 (3H, s), 3.24 (3H, s), 3.80
(1H, d, J=5Hz), 3.87 (1H, d, J=5Hz), 5.43 (1H,
d, J=10Hz), 5.52 (1H, d, J=10Hz), 6.15 (1H, br
t, J=5Hz), 6.72 (1H, d, J=7Hz), 6.88 (1H, t,
J=7Hz), 7.19-7.46 (4H), 7.70-8.00 (4H), 8.35
(1H, s), 8.70 (1H, d, J=5Hz)

(5) 3-Bromo-8-[3-[N-[N'-n-butylureidoacetyl]-N-
methylamino]-2,6-dichlorobenzyloxy]-2-
methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 0.90 (3H, t, J=6Hz), 1.21-1.54
(4H), 2.42 (3H, s), 3.12 (1H, q, J=6Hz), 3.23
(3H, s), 3.52 (2H, dd, J=17, 5Hz), 3.80 (1H, dd,
J=17, 5Hz), 4.68 (1H, br t, J=6Hz), 5.30 (1H, br
t, J=5Hz), 5.49 (2H, s), 6.71 (1H, d, J=7Hz),
6.88 (1H, t, J=7Hz), 7.33 (1H, d, J=9Hz), 7.48
(1H, d, J=9Hz), 7.78 (1H, d, J=7Hz)

(6) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(3-
quinolyl)ureidoacetyl]amino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ) : 2.41 (3H, s), 3.22 (3H, s), 3.86
(1H, dd, J=17, 5Hz), 3.99 (1H, dd, J=17, 5Hz),
5.42 (1H, d, J=10Hz), 5.52 (1H, d, J=10Hz), 6.11
(1H, br t, J=5Hz), 6.71 (1H, d, J=7Hz), 6.89
(1H, t, J=7Hz), 7.31-7.55 (4H), 7.61 (1H, d,
(7) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-[3-(4-pyridylcarbamoyl)phenyl]ureidoacetyl]amino]-benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 2.42 (3H, s), 3.25 (3H, s), 3.90-4.03 (2H, m), 5.11 (1H, d, J=10Hz), 5.38 (1H, d, J=10Hz), 6.54-6.69 (2H, m), 6.81 (1H, br s), 6.90 (1H, t, J=7.5Hz), 7.01 (1H, t, J=8Hz), 7.20-7.40 (3H, m), 7.50 (1H, d, J=8Hz), 7.78 (1H, d, J=6Hz), 7.94 (2H, d, J=6Hz), 8.28 (1H, br s), 8.57 (2H, d, J=6Hz)

(8) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-N'-[3-(1-pyrrolidinylcarbonyl)phenyl]ureidoacetyl]amino]-benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 1.76-1.99 (4H, m), 2.40 (3H, s), 3.20 (3H, s), 3.40 (2H, t, J=7Hz), 3.55-3.74 (3H, m), 3.81 (1H, dd, J=4, 17Hz), 5.36-5.52 (2H, m), 5.90-6.00 (1H, m), 6.72 (1H, d, J=8Hz), 6.85 (1H, t, J=8Hz), 7.10 (1H, d, J=6Hz), 7.18-7.28 (1H, m), 7.28-7.48 (4H, m), 7.77 (1H, d, J=6Hz), 7.93 (1H, br s)

(9) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-N'-[3-[N-methyl-N-(3-pyrrolidinylmethyl)carbamoyl]phenyl]ureidoacetyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 2.41 (3H, s), 2.82-3.02 (3H), 3.21 (3H, s), 3.69 (1H, dd, J=18, 5Hz), 3.80 (1H, dd, J=18, 5Hz), 4.52 (0.5H, br s), 4.73 (1.5H, br s), 5.42 (1H, d, J=10Hz), 5.51 (1H, d, J=10Hz), 5.96 (1H, br t, J=5Hz), 6.72 (1H, d, J=7Hz),

J=9Hz), 7.79 (1H, d, J=7Hz), 7.91 (1H, d, J=9Hz), 8.35 (1H, d, J=3Hz), 8.51 (1H, d, J=3Hz), 8.80 (1H, br s)
(10) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-[3-(4-methyl-1-piperazinylcarbonyl)phenyl]ureidoacetyl]-amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, 6): 2.21-2.55 (10H), 3.21 (3H, s), 3.33-3.53 (2H), 3.60-3.90 (4H), 5.43 (1H, d, J=10Hz), 5.51 (1H, d, J=10Hz), 5.96 (1H, br t, J=5Hz), 6.72 (1H, d, J=7Hz), 6.86 (1H, t, J=5Hz), 7.18-7.48 (5H), 7.79 (1H, d, J=7Hz), 8.00 (1H, br s)

(11) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-[3-(3-pyridylmethylcarbamoyl)phenyl]ureidoacetyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, 6): 2.40 (3H, s), 3.19 (3H, s), 3.62 (1H, dd, J=17, 5Hz), 4.57 (2H, d, J=5Hz), 5.49 (2H, s), 6.21 (1H, br t, J=5Hz), 6.72 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.01 (1H, t, J=8Hz), 7.18-7.39 (5H), 7.45 (1H, d, J=9Hz), 7.62-7.80 (3H), 8.25 (1H, br s), 8.49 (1H, dd, J=5, 1Hz), 8.60 (1H, br s)

(12) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-(dimethylamino)phenyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, 6): 2.42 (3H, s), 2.92 (6H, s), 3.22 (3H, s), 3.63 (1H, dd, J=17, 5Hz), 3.82 (1H, dd, J=17, 5Hz), 5.47 (2H, s), 5.91 (1H, br t, J=5Hz), 6.46 (1H, dd, J=8, 2Hz), 6.69-6.80 (3H), 6.96 (1H, t, J=7Hz), 7.13 (1H, t, J=8Hz), 7.31 (1H, d, J=8Hz), 7.44 (1H, d, J=8Hz), 7.77 (1H, d, J=7Hz)
Example 78
The following compounds were obtained according to a similar manner to that of Example 57.

(13) 8-[3-[N-[N'-[3-[N,N-Bis(2-methoxyethyl)carbamoyl]-phenyl]ureidoacetyl]-N-methylamino]-2,6-dichloro-benzyl]oxy]-3-bromo-2-methylimidazo[1,2-a]pyridine

\[
\text{NMR (CDCl}_3, \delta): \ 2.41 \ (3H, s), \ 3.22-3.83 \ (19H, m), \ 5.45 \ (1H, d, J=10Hz), \ 5.53 \ (1H, d, J=10Hz), \ 5.82 \ (1H, t-like), \ 6.73 \ (1H, d, J=8Hz), \ 6.88 \ (1H, dd, J=6, 8Hz), \ 7.01 \ (1H, d, J=8Hz), \ 7.17-7.48 \ (5H, m), \ 7.58 \ (1H, br s), \ 7.78 \ (1H, d, J=6Hz)
\]

(14) 3-Chloro-8-[2,6-dichloro-3-[N-methyl-N-[N'-[3-(4-pyridyl)carbamoyl]phenyl]ureidoacetyl]amino]-benzyl]oxy]-2-methylimidazo[1,2-a]pyridine

\[
\text{NMR (CDCl}_3, \delta): \ 2.40 \ (3H, s), \ 3.25 \ (3H, s), \ 3.81-4.06 \ (2H, m), \ 5.10 \ (1H, br d, J=9Hz), \ 5.36 \ (1H, d, J=9Hz), \ 6.55-6.64 \ (2H, m), \ 6.80 \ (1H, br s), \ 6.90 \ (1H, t, J=7Hz), \ 7.01 \ (1H, t, J=7Hz), \ 7.24-7.32 \ (2H, m), \ 7.36 \ (1H, d, J=8Hz), \ 7.50 \ (1H, br d, J=9Hz), \ 7.84 \ (2H, d, J=8Hz), \ 8.31 \ (1H, br s), \ 8.58 \ (2H, d, J=8Hz), \ 9.76 \ (1H, br s)
\]

(1) 3-Bromo-8-[3-[N-(4-carboxycinnamoylglycyl)]-N-methylamino]-2,6-dichlorobenzyl]oxy]-2-methylimidazo[1,2-a]pyridine

\[
\text{mp: 230°C (dec.)}
\]

\[
\text{NMR (DMSO-d}_6, \delta): \ 2.29 \ (3H, s), \ 3.15 \ (3H, s), \ 3.54 \ (1H, dd, J=16, 5Hz), \ 3.81 \ (1H, dd, J=16, 6Hz), \ 5.49 \ (2H, s), \ 6.90 \ (1H, d, J=16Hz), \ 6.31-7.73 \ (2H, m), \ 7.79 \ (1H, d, J=7Hz), \ 7.84 \ (1H, d, J=7Hz), \ 7.88-8.01 \ (3H, m), \ 8.41 \ (1H, t, J=5Hz)
\]

(2) 3-Chloro-8-[3-[N-(4-carboxycinnamoylglycyl)]-N-
methylamino]-2,6-dichlorobenzyloxy]-2-methyylimidazo[1,2-a]pyridine

mp : >250°C

NMR (DMSO-\(d_6\), \(\delta\)) : 2.30 (3H, s), 3.17 (3H, s), 3.82 (1H, dd, \(J=4, 18\)Hz), 5.50 (2H, s), 6.80-7.06 (3H, m), 7.74-7.88 (2H, m), 7.88-8.03 (3H, m), 8.33-8.48 (1H, m)

Example 79

The following compounds were obtained according to a similar manner to that of Example 67.

(1) 3-Bromo-8-[3-(E)-3-(6-carboxy-3-pyridyl)acyrloylglycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methyylimidazo[1,2-a]pyridine

mp : 226-228°C (dec.)

NMR (DMSO-\(d_6\), \(\delta\)) : 2.30 (3H, s), 3.16 (3H, s), 3.45-3.62 (1H, overlapped with \(H_2O\)), 3.82 (1H, dd, \(J=18, 5\)Hz), 5.49 (2H, s), 6.93-7.12 (3H), 7.51 (1H, d, \(J=16\)Hz), 7.79 (1H, d, \(J=9\)Hz), 7.85 (1H, d, \(J=9\)Hz), 7.93 (1H, dd, \(J=5, 3\)Hz), 8.01-8.20 (2H), 8.44 (1H, br t, \(J=5\)Hz), 8.89 (1H, br s)

(2) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(ethylcarbamoyl)cinnamoylglycyl]-N-
3-Bromo-8-[2,6-dichloro-3-[N-[4-(N-ethyl-N-methylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, δ) : 1.28 (3H, t, $J$=7Hz), 2.44 (3H, s), 3.29 (3H, s), 3.42-3.60 (2H), 3.68 (1H, dd, $J$=18, 4Hz), 3.91 (1H, dd, $J$=18, 5Hz), 5.50 (2H, s), 6.13 (1H, br t, $J$=5Hz), 6.52 (1H, d, $J$=15Hz), 6.67-6.78 (2H), 6.87 (1H, t, $J$=7Hz), 7.34 (1H, d, $J$=8Hz), 7.48-7.66 (4H), 7.71-7.81 (3H)

(3) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(N-ethyl-N-methylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, δ) : 1.08-1.31 (3H), 2.45 (3H, s), 2.90-3.12 (3H), 3.20-3.39 (4H), 3.51-3.77 (2H), 3.91 (1H, dd, $J$=18, 5Hz), 5.50 (2H, s), 6.5 (1H, d, $J$=15Hz), 6.68 (1H, br t, $J$=5Hz), 6.72 (1H, d, $J$=7Hz), 6.88 (1H, t, $J$=7Hz), 7.30-7.64 (7H), 7.78 (1H, d, $J$=7Hz)

(4) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(1-pyrrolidinylcarbonyl)cinnamoylglycyl]amino]-benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, δ) : 1.80-2.07 (4H, m), 2.44 (3H, s), 3.28 (3H, s), 3.43 (2H, t, $J$=6Hz), 3.57-3.74 (3H, m), 3.91 (1H, dd, $J$=4, 18Hz), 5.42-5.56 (2H, m), 6.49 (1H, d, $J$=16Hz), 6.63 (1H, t-like), 6.71 (1H, d, $J$=7.5Hz), 6.85 (1H, t, $J$=7.5Hz), 7.33 (1H, d, $J$=8Hz), 7.44-7.66 (6H, m), 7.71 (1H, d, $J$=6Hz)

(5) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(morpholinocarbonyl)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, δ) : 2.44 (3H, s), 3.27 (3H, s), 3.35-3.87 (9H, m), 3.91 (1H, dd, $J$=4, 18Hz), 5.41-
5.53 (2H, m), 6.50 (1H, d, J=16Hz), 6.65 (1H, t-like), 6.72 (1H, d, J=7.5Hz), 6.85 (1H, t, J=7.5Hz), 7.33 (1H, d, J=8Hz), 7.41 (2H, d, J=8Hz), 7.47-7.64 (4H, m), 7.77 (1H, d, J=6Hz)

(6) 3-Bromo-8-[2,6-dichloro-3-[N-[4-[N-(2-methoxyethyl)-N-methylcarbamoyl]cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 2.43 (3H, s), 3.00-3.15 (3H, m), 3.23-3.50 (8H, m), 3.50-3.78 (3H, m), 3.93 (1H, dd, J=4, 18Hz), 5.44-5.56 (2H, m), 6.50 (1H, d, J=16Hz), 6.64 (1H, t-like), 6.72 (1H, d, J=7.5Hz), 6.86 (1H, t, J=7.5Hz), 7.34 (1H, d, J=8Hz), 7.39-7.65 (6H, m), 7.76 (1H, d, J=6Hz)

(7) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(dimethylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 2.45 (3H, s), 2.99 (3H, br s), 3.10 (3H, br s), 3.29 (3H, s), 3.69 (1H, dd, J=17, 4Hz), 3.91 (1H, dd, J=17, 5Hz), 5.47 (1H, d, J=10Hz), 5.52 (1H, d, J=10Hz), 6.50 (1H, d, J=15Hz), 6.65 (1H, br t, J=4Hz), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.29-7.62 (7H), 7.78 (1H, d, J=7Hz)

(8) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(isopropylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp: 238°C (dec.)
NMR (CDCl₃, δ): 1.27 (6H, d, J=6Hz), 2.44 (3H, s), 3.28 (3H, s), 3.67 (1H, dd, J=4, 18Hz), 3.91 (1H, dd, J=4, 18Hz), 4.31 (1H, m), 5.43-5.57 (2H, m), 5.92 (1H, d, J=6Hz), 6.53 (1H, d,
5 (9) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(n-propylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) :  1.00 (3H, t, J=7.5Hz), 1.63 (2H, m), 2.44 (3H, s), 3.28 (3H, s), 3.43 (2H, q, J=7.5Hz), 3.67 (1H, dd, J=4, 18Hz), 3.92 (1H, dd, J=4, 18Hz), 5.43-5.56 (2H, m), 6.13 (1H, t-like), 6.54 (1H, d, J=16Hz), 6.64-6.77 (2H, m), 6.85 (1H, t, J=7.5Hz), 7.35 (1H, d, J=8Hz), 7.44-7.66 (4H, m), 7.69-7.81 (1H, m)

10 (10) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(3-methoxypropylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃-CD₃OD, δ) :  1.23 (3H, t, J=7Hz), 2.43 (3H, s), 3.28 (3H, s), 3.46-3.77 (7H, m), 3.93 (1H, dd, J=4, 18Hz), 5.42-5.56 (2H, m), 6.46-6.61 (2H, m), 6.65-6.78 (2H, m), 6.87 (1H, t, J=7 5Hz), 7.35 (8H, d), 7.46-7.64 (4H, m), 7.72-7.83 (3H, m)

15 (11) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(2-ethoxyethylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃-CD₃OD, δ) :  1.23 (3H, t, J=7Hz), 2.43 (3H, s), 3.28 (3H, s), 3.46-3.77 (7H, m), 3.93 (1H, dd, J=4, 18Hz), 5.42-5.56 (2H, m), 6.46-6.61 (2H, m), 6.65-6.78 (2H, m), 6.87 (1H, t, J=7 5Hz), 7.35 (8H, d), 7.46-7.64 (4H, m), 7.72-7.83 (3H, m)
(12) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(2-
hydroxyethylcarbamoyl)cinnamoylglycyl]-N-
methy lamino]benzyloxy]-2-methylimida zo[1,2-a]pyridine
NMR (CDCl₃-CD₃OD, δ): 2.41 (3H, s), 3.26 (3H, s),
3.34-3.46 (1H, m), 3.57 (2H, q, J=5Hz), 3.65-
3.81 (3H, m), 3.95 (1H, d, J=17.5Hz), 5.50 (2H,
s), 6.58 (1H, d, J=16Hz), 6.75 (1H, d, J=7.5Hz),
6.89 (1H, t, J=7.5Hz), 7.42 (1H, d, J=8Hz),
7.48-7.63 (4H, m), 7.73-7.87 (3H, m)

(13) 3-Bromo-8-[2,6-dichloro-3-[N-[4-
diethylcarbamoyl)cinnamoylglycyl]-N-methylamino]-
benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 1.00-1.35 (6H, m), 2.43 (3H, s),
3.26 (3H, s), 3.40-3.75 (5H, m), 3.91 (1H, dd,
J=4, 18Hz), 5.47 (1H, d, J=10Hz), 5.53 (1H, d,
J=10Hz), 6.48 (1H, d, J=16Hz), 6.65 (1H,
t-like), 6.71 (1H, d, J=7.5Hz), 6.86 (1H, t,
J=7.5Hz), 7.30-7.45 (3H, m), 7.45-7.64 (4H, m),
7.76 (6H, d)

(14) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-(2-
 methoxyethylcarbamoyl)phenyl]ureidoacetyl]-N-
methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 2.42 (3H, s), 3.23 (3H, s), 3.35
(3H, s), 3.49-3.79 (5H, m), 3.87 (1H, dd, J=18,
5Hz), 5.44 (1H, d, J=10Hz), 5.52 (1H, d,
J=10Hz), 6.07 (1H, t-like), 6.72 (1H, d, J=8Hz),
6.80-6.93 (2H, m), 7.15-7.60 (6H, m), 7.78 (1H,
d, J=6Hz), 7.92 (1H, s)

(15) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-[N-(2-
methoxyethyl)]-N-methylcarbamoyl]phenyl]ureidoacetyl]-
N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]-
pyridine
NMR (CDCl₃, δ): 2.42 (3H, s), 2.96-3.14 (3H, m),
3.20-3.49 (8H, m), 5.43 (1H, d, J=10Hz),
5.51 (1H, d, J=10Hz), 5.88 (1H, t-like),
6.72 (1H, d, J=7.5Hz), 6.86 (1H, t, J=7.5Hz),
7.00 (1H, dif-dd, J=7.5Hz), 7.18-7.47 (5H, m),
7.72 (1H, s), 7.77 (1H, d, J=6Hz)

(16) 3-Bromo-8-[2,6-dichloro-3-[N'-[3-[N,N-bis(2-
ethoxyethyl)carbamoyl]phenyl]ureidoacetyl]-N-
methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 1.07-1.28 (6H, m),
2.40 (3H, s), 3.23 (3H, s),
3.26-4.01 (12H, m), 5.39-5.55 (2H, m), 6.01 (1H, t-like),
6.71 (1H, d, J=7.5Hz), 6.86 (1H, t, J=7.5Hz),
7.02 (1H, d, J=7.5Hz), 7.10-7.53 (3H, m),
7.58 (1H, s), 7.79 (1H, d, J=6Hz)

(17) 3-Bromo-8-[2,6-dichloro-3-[N'-[3-[N,N-bis(2-
hydroxyethyl)carbamoyl]phenyl]ureidoacetyl]-N-
methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 1.05-1.32 (3H, m), 2.43 (3H, s),
2.88-3.13 (3H, m), 3.20-3.37 (4H, m), 3.47-3.76 (2H, m), 3.93 (1H, dd, J=4, 18Hz), 5.41-5.57 (2H, m), 6.50 (1H, d, J=16Hz), 6.60-6.75 (2H, m), 6.86 (1H, t, J=7.5Hz), 7.30-7.46 (3H, m),

(18) 3-Chloro-8-[2,6-dichloro-3-[N-[4-(N-ethyl-N-
methylcarbamoyl)cinnamoylglycyl]-N-
methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ): 1.05-1.32 (3H, m), 2.43 (3H, s),
2.88-3.13 (3H, m), 3.20-3.37 (4H, m), 3.47-3.76 (2H, m), 3.93 (1H, dd, J=4, 18Hz), 5.41-5.57 (2H, m), 6.50 (1H, d, J=16Hz), 6.60-6.75 (2H, m), 6.86 (1H, t, J=7.5Hz), 7.30-7.46 (3H, m),
7.46-7.64 (4H, m), 7.73 (1H, d, J=6Hz)

(19) 3-Chloro-8-[2,6-dichloro-3-[N-[4-(ethylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 1.24 (3H, t, J=7.5Hz), 2.41 (3H, s), 3.26 (3H, s), 3.50 (2H, quint, J=7.5Hz), 3.65 (1H, dd, J=4, 18Hz), 3.90 (1H, dd, J=4, 18Hz), 5.43-5.55 (2H, m), 6.10 (1H, t-like), 6.53 (1H, d, J=16Hz), 6.63-6.75 (2H, m), 6.85 (1H, t, J=7.5Hz), 7.33 (1H, d, J=8Hz). 7.44-7.64 (4H, m), 7.67-7.80 (3H, m)

(20) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(E)-3-(6-dimethylcarbamoyl-3-pyridyl)acryloylglycyl]-amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 2.45 (3H, s), 3.05 (3H, d, J=5Hz), 3.29 (3H, s), 3.70 (1H, dd, J=18, 4Hz), 3.92 (1H, dd, J=18, 4Hz), 5.49 (1H, d, J=10Hz), 5.52 (1H, d, J=10Hz), 6.61 (1H, d, J=16Hz), 6.70-6.79 (2H), 6.88 (1H, t, J=7.5Hz), 7.34 (1H, d, J=7.5Hz), 7.50 (1H, d, J=7.5Hz), 7.61 (1H, d, J=16Hz), 7.78 (1H, d, J=7.5Hz), 7.92-8.02 (2H), 8.20 (1H, d, J=7.5Hz), 8.62 (1H, br s)

(21) 3-Bromo-8-[2,6-dichloro-3-[N-[(E)-3-(6-dimethylcarbamoyl-3-pyridyl)acryloylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 2.43 (3H, s), 3.11 (3H, s), 3.16 (3H, s), 3.29 (3H, s), 3.69 (1H, dd, J=18, 5Hz), 3.91 (1H, dd, J=18, 5Hz), 5.49 (1H, d, J=10Hz), 5.52 (1H, d, J=10Hz), 6.60 (1H, d, J=16Hz), 6.70-6.78 (2H), 6.87 (1H, t, J=7.5Hz), 7.35 (1H, d, J=8Hz), 7.50 (1H, d, J=8Hz), 7.60 (1H, d, J=16Hz), 7.69 (1H, d, J=8Hz), 7.78 (1H, d,
J=7.5Hz), 7.91 (1H, dd, J=8, 2Hz), 8.69 (1H, br s)

(22) 3-Bromo-8-[2,6-dichloro-3-[N-[(E)-3-(6-
ethylcarbamoyl-3-pyridyl)acryloylglycyl]-N-
methy lamino]benzyloxy]-2-methylimidazo[1,2-
a]pyridine
NMR (CDCl₃, δ) : 1.29 (3H, t, J=7Hz), 2.45 (3H, s), 3.29 (3H, s), 3.48-3.59 (2H), 3.70 (1H, dd, J=18, 4Hz), 3.92 (1H, dd, J=18, 5Hz), 5.49 (1H, d, J=19Hz), 6.53 (1H, d, J=10Hz), 6.61 (1H, d, J=16Hz), 7.34 (1H, d, J=8Hz), 7.51 (1H, d, J=8Hz), 7.62 (1H, d, J=16Hz), 7.68 (1H, d, J=7Hz), 7.92-8.01 (2H), 8.20 (1H, d, J=7.5Hz), 8.63 (1H, br s)

(23) 3-Bromo-8-[2,6-dichloro-3-[N-[(E)-3-[6-(3-
ethoxypropylcarbamoyl)-3-pyridyl)acryloylglycyl]-N-
methylamino]benzyloxy]-2-methylimidazo[1,2-
a]pyridine
NMR (CDCl₃, δ) : 1.91 (2H, quint, J=6.5Hz), 2.45 (3H, s), 3.28 (3H, s), 3.38 (3H, s), 3.50-3.63 (4H, m), 3.68 (1H, dd, J=4, 18Hz), 3.93 (1H, dd, J=4, 18Hz), 5.48 (1H, d, J=10Hz), 5.53 (1H, d, J=10Hz), 6.61 (1H, d, J=16Hz), 6.70-7.76 (2H, m), 6.86 (1H, t, J=7.3Hz), 7.35 (1H, d, J=8.7Hz), 7.50 (1H, d, J=8.3Hz), 7.62 (1H, d, J=16Hz), 7.78 (1H, d, J=7.5Hz), 7.95 (1H, dd, J=8.3, 2.2Hz), 8.20 (1H, d, J=8.3Hz), 8.29 (1H, t-like), 8.65 (1H, d, J=2.0Hz)

Example 80
To a suspension of 3-bromo-8-[2,6-dichloro-3-[N-
methyl-N-[N'-[(3-nitrophenyl)ureidoacetyl]amino]benzyloxy]-
2-methylimidazo[1,2-a]pyridine (800 mg) in ethanol (8 ml) was added tin(II) chloride (954 mg) at ambient temperature. The mixture was refluxed for 1.5 hours.
After cooling, the mixture was adjusted to pH 10 with 1N sodium hydroxide solution. To this mixture was added dichloromethane (10 ml) and the precipitate was removed by filtration. The filtrate was extracted with dichloromethane twice. The organic layer was washed with saturated sodium bicarbonate, water and brine. After dried over magnesium sulfate, the solvent was removed in vacuo. The residue was purified by column chromatography eluting with dichloromethane-methanol to give 8-[3-[N-[N'-
(3-aminophenyl)ureidoacetyl]-N-methylamino]-2,6-
dichlorobenzyl]oxy]-3-bromo-2-methylimidazo[1,2-a]pyridine (539 mg) as amorphous.

NMR (CDCl₃, δ): 2.42 (3H, s), 3.22 (3H, s), 3.56-
3.75 (3H), 3.82 (1H, dd, J=18, 5Hz), 5.48 (2H,
s), 5.92 (1H, br t, J=4Hz), 6.88 (1H, dd, J=8,
1Hz), 6.53 (1H, br d, J=7Hz), 6.72 (1H, d,
J=7Hz), 6.79-6.91 (2H), 7.01 (1H, t, J=8Hz),
7.09 (1H, br s), 7.33 (1H, d, J=9Hz), 7.45 (1H,
d, J=9Hz), 7.78 (1H, d, J=7Hz)

Example 81

3-Bromo-8-[2,6-dichloro-3-[N-[4-
(ethoxycarbonylacetyl)cinnamonylglycyl]-N-
methylamino]benzyl]oxy]-2-methylimidazo[1,2-a]pyridine was obtained by reacting 8-[3-[N-[4-aminoacetylglycyl]-N-
methylamino]-2,6-dichlorobenzyl]oxy]-3-bromo-2-
methylimidazo[1,2-a]pyridine with ethyl chloroformylacetate according to a similar manner to that of Example 37.

NMR (CDCl₃, δ): 1.33 (3H, t, J=7.5Hz), 2.44 (3H,
s), 3.26 (3H, s), 3.47 (2H, s), 3.65 (1H, dd,
J=4, 18Hz), 3.90 (1H, dd, J=4, 18Hz), 4.26 (2H,
g, J=7.5Hz), 5.42-5.55 (2H, m), 6.41 (1H, d,
J=16Hz), 6.60 (1H, t-like), 6.72 (1H, d,
J=7.5Hz), 6.85 (1H, t, J=7.5Hz), 7.33 (1H, d,
Example 82

8-[3-[N-[4-(Benzamido)cinnamoylglycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine was obtained by reacting 8-[3-[N-(4-aminocinnamoylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine with benzoic acid in the presence of N-ethyl-N'-{(3-dimethylaminopropyl)carbodiimide hydrochloride and 1-hydroxybenzotriazole according to a similar manner to that of Example 39.

NMR (CDCl$_3$, $\delta$): 2.41 (3H, s), 3.25 (3H, s), 3.66 (1H, dd, $J= 18$, 4Hz), 3.90 (1H, dd, $J= 18$, 5Hz), 5.49 (2H, br s), 6.42 (1H, d, $J= 16$Hz), 6.60 (1H, br t), 6.72 (1H, d, $J= 7$Hz), 6.86 (1H, t, $J= 7$Hz), 7.18-7.80 (9H, m), 7.86 (2H, d, $J= 9$Hz), 8.01 (1H, br s)

Example 83

The following compounds were obtained according to similar manners to those of Examples 81 or 82.

(1) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(4-pyridylacetamido)cinnamoylglycyl]amino]benzoyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl$_3$, $\delta$): 2.40 (3H, s), 3.23 (3H, s), 3.64 (1H, dd, $J= 18$, 5Hz), 3.67 (2H, br s), 3.86 (1H, dd, $J= 18$, 4Hz), 5.42 (1H, d, $J= 9$Hz), 5.48 (1H, d, $J= 9$Hz), 6.39 (1H, d, $J= 16$Hz), 6.63 (1H, br t, $J= 4$Hz), 6.73 (1H, d, $J= 7.5$Hz), 6.86 (1H, t, $J= 7.5$Hz), 7.15-7.33 (3H, m), 7.35-7.57 (6H, m), 7.78 (1H, d, $J= 7.5$Hz), 8.18 (1H br s), 8.58 (2H, dd, $J= 7$, 2Hz)
(2) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(3-
methoxypropionamido)cinnamoylglycyl]-N-methylamino]-benzyl oxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) 2.45 (3H, s), 2.65 (2H, t, J=6Hz),
3.28 (3H, s), 3.46 (3H, s), 3.59-3.79 (3H, m),
3.90 (1H, dd, J=4, 18Hz), 5.42-5.57 (2H, m),
6.40 (1H, d, J=16Hz), 6.60 (1H, t-like), 6.73
(1H, d, J=7Hz), 6.87 (1H, d, J=7Hz), 7.33 (1H,
d, J=8Hz), 7.40-7.60 (6H, m), 7.77 (1H, d,
J=6Hz), 8.39 (1H, s)

(3) 8-[3-[N-[4- (Acetamidoacetamido)cinnamoylglycyl]-N-
methylamino]-2,6-dichlorobenzyl oxy]-3-bromo-2-
methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 2.10 (3H, s), 2.44 (3H, s), 3.27
(3H, s), 3.64 (1H, dd, J=4, 17.5Hz), 3.92 (1H,
dd, J=4, 17.5Hz), 4.06 (2H, d, J=6Hz), 5.43-5.56
(2H, m), 6.35 (1H, d, J=16Hz), 6.59 (1H,
t-like), 6.66-6.92 (3H, m), 7.30-7.57 (7H, m),
7.76 (1H, d, J=6Hz), 8.86 (1H, s)

(4) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-[2-
dimethylamino)acetamido)cinnamoylglycyl]amino]-
benzyl oxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl₃, δ) : 2.39 (6H, s), 2.43 (3H, s), 3.08
(2H, s), 3.28 (3H, s), 3.68 (1H, dd, J=4,
17.5Hz), 4.92 (1H, dd, J=4, 17.5Hz), 5.42-5.56
(2H, m), 6.42 (1H, d, J=16Hz), 6.59 (1H,
t-like), 6.72 (1H, d, J=7.5Hz), 6.86 (1H, t,
J=7.5Hz), 7.33 (1H, d, J=8Hz), 7.40-7.68 (6H,
m), 7.76 (1H, d, J=6Hz), 9.23 (1H, s)

(5) 3-Bromo-8-[2,6-dichloro-3-[N-[4-[(E)-3-
(ethoxycarbonyl)acrylamido)cinnamoylglycyl]-N-
methylamino]benzyl oxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, $\delta$): 1.30 (3H, t, $J=7.5$Hz), 2.41 (3H, s), 3.25 (3H, s), 3.64 (1H, dd, $J=17$, 4Hz), 3.88 (1H, dd, $J=17$, 5Hz), 4.24 (2H, q, $J=7.5$Hz), 5.42 (1H, d, $J=9$Hz), 5.49 (1H, d, $J=9$Hz), 6.40 (1H, d, $J=16$Hz), 6.67 (1H, br t, $J=4$Hz), 6.73 (1H, d, $J=7$Hz), 7.10 (1H, d, $J=15$Hz), 7.20 (2H, d, $J=9$Hz), 7.36-7.52 (3H, m), 7.58 (1H, d, $J=11$Hz), 7.63 (1H, d, $J=7$Hz), 7.78 (1H, d, $J=7$Hz), 8.82 (1H, br s)

(6) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(hydroxyacetamido)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, $\delta$): 2.39 (3H, s), 3.24 (3H, s), 3.67 (1H, dd, $J=17$, 4Hz), 3.94 (1H, dd, $J=17$, 5Hz), 4.11 (2H, s), 5.48 (2H, s), 6.30 (1H, d, $J=15$Hz), 6.66-6.90 (2H, m), 6.86 (1H, t, $J=7$Hz), 7.25-7.60 (7H, m), 7.76 (1H, d, $J=7$Hz), 8.78 (1H, br s)

(7) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(2-chloroethoxycarbonylamino)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
NMR (CDCl$_3$, $\delta$): 2.44 (3H, s), 3.27 (3H, s), 3.57-3.80 (3H, m), 3.91 (1H, dd, $J=4$, 18Hz), 4.45 (2H, t, $J=6$Hz), 5.50 (2H, s), 6.40 (1H, d, $J=16$Hz), 6.60 (1H, t-like), 6.72 (1H, d, $J=7.5$Hz), 6.80-6.91 (2H, m), 7.22-7.60 (7H, m), 7.78 (1H, d, $J=6$Hz)

(8) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-(methoxyacetamido)phenyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
mp: 238-239°C
NMR (CDCl$_3$-CD$_3$OD, $\delta$): 2.40 (3H, s), 3.25 (3H, s),
3.45-3.68 (4H, overlapped with H2O), 3.89 (1H, d, J=18Hz), 4.00 (2H, s), 5.50 (2H, s), 6.78 (1H, d, J=7Hz), 6.91 (1H, t, J=7Hz), 7.08-7.38 (3H), 7.44 (1H, d, J=9Hz), 7.52 (1H, d, J=9Hz), 7.59 (1H, br s), 7.80 (1H, d, J=7Hz)

(9) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-ethoxycarbonylacetamido]phenyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 1.27 (3H, t, J=7Hz), 2.40 (3H, s), 3.19 (3H, s), 3.41 (2H, s), 3.59 (1H, br d, J=18Hz), 3.89 (1H, br d, J=18Hz), 4.19 (2H, q, J=7Hz), 5.40 (2H, s), 6.22 (1H, br s), 6.72 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.00-7.52 (6H), 7.78 (1H, d, J=7Hz), 8.18 (1H, br s), 9.62 (1H, br s)

(10) 3-Chloro-8-[2,6-dichloro-3-[N-[4-(3-methoxypropionamido)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine

NMR (CDCl₃, δ): 2.09 (3H, s), 2.42 (3H, s), 3.25 (3H, s), 3.64 (1H, dd, J=4, 18Hz), 3.90 (1H, dd, J=4, 18Hz), 4.05 (2H, d, J=5Hz), 5.43-5.54 (2H, m), 6.39 (1H, d, J=16Hz), 6.62 (1H, t-like),
Example 84

A solution of 8-[3-[N-(4-aminocinnamoylglycyl)-N-methylamino]-2,6-dichlorobenzoyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine (150 mg) and succinic anhydride (26 mg) in dioxane (3 ml) was refluxed for 2 hours. After cooling, the solution was removed in vacuo to give 3-bromo-8-[3-[N-[4-(3-carboxypropionamido)cinnamoylglycyl]-N-methylamino]-2,6-dichlorobenzoyloxy]-2-methylimidazo[1,2-a]pyridine (175 mg) as amorphous.

NMR (CDCl₃-CD₃OD, δ): 2.41 (3H, s), 2.59-2.70 (4H), 3.27 (3H, s), 3.64 (1H, d, J=18Hz), 3.98 (1H, d, J=18Hz), 5.51 (2H, s), 6.41 (1H, d, J=15Hz), 6.76 (1H, d, J=7Hz), 6.90 (1H, t, J=7Hz), 7.38-7.62 (7H), 7.79 (1H, d, J=7Hz)

Example 85

A solution of 8-[3-(N-glycyl-N-methylamino)-2,6-dichlorobenzoyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine (300 mg) and dimethyl cyanodithioiminocarbonate (93 mg) in dimethylformamide (3 ml) was heated at 70°C for 1 hour. After cooling the reacting mixture, to the mixture was added 70% solution of ethylamine in water (0.57 ml) and the mixture was heated at 60°C for 2 hours. To this mixture was added water (3 ml) in an ice-water bath. The precipitates were collected by vacuum filtration and washed with ethyl acetate to give 3-bromo-8-[3-[N-[[2-cyano-3-ethylguanidino]acetyl]-N-methylamino]-2,6-dichlorobenzoyloxy]-2-methylimidazo[1,2-a]pyridine (278 mg) as colorless crystals.

mp: >250°C

NMR (CDCl₃-CD₃OD, δ): 1.20 (3H, t, J=7Hz), 2.39
(3H, s), 3.16-3.31 (5H), 3.61 (1H, d, J=17Hz), 3.73 (1H, d, J=17Hz), 5.47 (1H, d, J=10Hz), 5.57 (1H, d, J=10Hz), 6.79 (1H, d, J=7Hz), 6.91 (1H, t, J=7Hz), 7.42 (1H, d, J=9Hz), 7.56 (1H, d, J=9Hz), 7.80 (1H, d, J=7Hz)

Example 86

A suspension of 3-bromo-8-[3-N-[4-(3-carboxypropionamido)cinnamoylglycyl]-N-methylamino]-2,6-dichlorobenzylolox]-2-methylimidazo[1,2-a]pyridine (110 mg) and anhydrous sodium acetate (16 mg) in acetic anhydride (1.1 ml) was refluxed for 5 hours. After cooling, the solvent was removed in vacuo. The residue was dissolved in dichloromethane (5 ml) and washed with water, saturated sodium bicarbonate twice and brine. After dried over magnesium sulfate, the solvent was removed in vacuo. The residue was purified by preparative thin-layer chromatography (ethyl acetate - methanol) to give 3-bromo-8-[2,6-dichloro-3-[N-[4-(succinimido)cinnamoylglycyl]-N-methylamino]benzylolox]-2-methylimidazo[1,2-a]pyridine (12 mg) as amorphous.

NMR (CDCl₃, δ) : 2.43 (3H, s), 2.91 (4H, s), 3.29 (3H, s), 3.68 (1H, dd, J=18, 4Hz), 3.91 (1H, dd, J=18, 5Hz), 5.48 (1H, d, J=10Hz), 5.53 (1H, d, J=10Hz), 6.50 (1H, d, J=16Hz), 6.68 (1H, br t, J=4Hz), 6.72 (1H, d, J=7Hz), 6.88 (1H, t, J=7Hz), 7.22-7.39 (3H), 7.48-7.67 (4H), 7.78 (1H, d, J=7Hz)

Example 87

To a solution of 3-bromo-8-[2,6-dichloro-3-[N-[4-(2-chloroethoxycarbonylamino)cinnamoylglycyl]-N-methylamino]benzylolox]-2-methylimidazo[1,2-a]pyridine (128 mg) in methanol was added dropwise sodium methanolate (28% in methanol, 37 mg) under nitrogen in ice-water bath and
The mixture was stirred for 1 hour at same temperature and then at ambient temperature for 2 hours. The reaction mixture was poured into water and extracted with dichloromethane. The organic layer was washed with water and brine, dried over magnesium sulfate and evaporated in vacuo. The residue was purified by preparative thin layer chromatography (ethyl acetate) to give 3-bromo-8-[2,6-dichloro-3-[N-[4-(2-oxo-3-oxazolidinyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (77 mg) as an amorphous powder.

NMR (CDCl₃, δ): 2.44 (3H, s), 3.28 (3H, s), 3.66 (1H, dd, J=4, 18Hz), 3.91 (1H, dd, J=4, 18Hz), 4.10 (2H, dd, J=6, 8Hz), 4.53 (2H, dd, J=6, 8Hz), 5.45-5.57 (2H, m), 6.43 (1H, d, J=16Hz), 6.60 (1H, t-like), 6.72 (1H, d, J=7.5Hz), 6.85 (1H, t, J=7.5Hz), 7.34 (1H, d, J=8Hz), 7.45-7.62 (6H, m), 7.76 (6H, d)

**Example 88**

1. 3-Bromo-8-[3-[N-[N'-[3-(4-bromobutyramido)phenyl]ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Example 82.

NMR (CDCl₃, δ): 2.01-2.27 (2H), 2.41 (3H, s), 2.59 (2H, t, J=8Hz), 3.23 (3H, s), 3.63 (1H, dd, J=17, 5Hz), 3.74-3.92 (3H), 4.58 (1H, br t, J=6Hz), 5.46 (2H, s), 5.99 (1H, br t, J=5Hz), 6.72 (1H, d, J=7Hz), 6.88 (1H, t, J=8Hz), 7.02-7.50 (5H), 7.60 (1H, d, J=9Hz), 7.77 (1H, d, J=7Hz)

2. 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-[3-(2-oxo-1-pyrrolidinyl)phenyl]ureidoacetyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Example 87.

NMR (CDCl₃, δ): 2.01-2.27 (2H), 2.41 (3H, s), 2.59
Example 89

A mixture of 3-bromo-8-[2,6-dichloro-3-[N-[4-(3-carboxypropionamido)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (60 mg), ethanol (12 mg), N-ethyl-N'-(3-dimethylaminopropyl)carbodiimide hydrochloride (21 mg), 1-hydroxybenzotriazole (17 mg) and N,N-dimethylformamide (0.6 ml) was stirred for 3 hours at ambient temperature. Ethyl acetate was added thereto, and the mixture was washed with water four times, dried over magnesium sulfate and concentrated in vacuo. The residue was purified by preparative thin layer chromatography (methylene chloride:methanol = 10:1, V/V) to give 3-bromo-8-[2,5-dichloro-3-[N-[4-(3-ethoxycarbonylpropionamido)-cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (7 mg) as an amorphous.

NMR (CDCl₃, δ) : 1.28 (3H, t, J=7Hz), 2.44 (3H, s), 2.60-2.82 (4H), 3.28 (3H, s), 3.66 (1H, dd, J=18, 4Hz), 3.91 (1H, dd, J=18, 5Hz), 4.18 (2H, q, J=7Hz), 5.49 (2H, s), 6.39 (1H, d, J=15Hz), 6.61-6.79 (2H), 6.87 (1H, t, J=7Hz), 7.30-7.60 (7H), 7.78 (1H, d, J=7Hz), 8.12 (1H, br s)

Example 90

To a solution of 8-[3-[N-[N'-(3-acetylpheynyl)ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine (200 mg) in methanol (2 ml) was added sodium borohydride (24 mg) under ice-bath cooling, and the mixture was stirred for 1 hour at ambient temperature. To the reaction mixture was added 1N hydrochloric acid, and the mixture was stirred for 30 minutes. Saturated aqueous solution of sodium bicarbonate was added thereto, and the mixture was extracted with methylene chloride three times. The combined organic layer was washed with water and
brine, dried over magnesium sulfate, and concentrated in vacuo. The residue was purified by silica gel column chromatography (methylene chloride:methanol=30:1, V/V) to give 3-bromo-8-[2,6-dichloro-3-[N-[N'-[3-(1-hydroxyethyl)-phenyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (160 mg) as an amorphous.

NMR (CDCl₃, δ): 1.41 (3H, d, J=6Hz), 2.41 (3H, s), 3.22 (3H, s), 3.61 (1H, br dt, J=17, 3Hz), 3.88 (1H, dd, J=17, 5Hz), 4.79 (1H, m), 5.49 (2H, s), 6.09 (1H, br t, J=5Hz), 6.72 (1H, d, J=7Hz), 6.87 (1H, t, J=7Hz), 6.96 (1H, m), 7.10-7.49 (4H), 7.59 (1H, br d, J=7Hz), 7.78 (1H, d, J=7Hz)

Example 91

To a solution of 3-bromo-8-[2,6-dichloro-3-[N-methyl-N-[thiomorpholinoacetyl]glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (200 mg) in methylene chloride (2 ml) was added m-chloroperbenzoic acid (80% purity, 76 mg) under ice-bath cooling, and the mixture was stirred for 1 hour at the same temperature. The reaction mixture was washed with water twice and brine, dried over magnesium sulfate, and concentrated in vacuo. The residue was purified by preparative thin layer chromatography (methylene chloride:methanol = 5:1, V/V) to give 3-bromo-8-[2,6-dichloro-3-[N-methyl-N-[(1-oxothiomorpholino)acetyl]glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (96 mg) as an amorphous.

NMR (CDCl₃, δ): 2.44 (3H, s), 2.70-3.02 (6H), 3.11 (2H, s), 3.15-3.36 (5H), 3.57 (1H, dd, J=17, 5Hz), 3.85 (1H, dd, J=17, 5Hz), 5.47 (1H, dd, J=10Hz), 5.52 (1H, d, J=10Hz), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.31 (1H, d, J=9Hz), 7.50 (1H, d, J=9Hz), 7.79 (1H, d, J=7Hz), 7.84 (1H, br t, J=5Hz)
Example 92

To a solution of 3-bromo-8-[2,6-dichloro-3-[N-methyl-N-[(thiomorpholinoacetyl)glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (200 mg) in methylene chloride (2 ml) was added m-chloroperbenzoic acid (80% purity, 190 mg) under ice-bath cooling, and the mixture was stirred for 2 hours at the same temperature. The reaction mixture was washed with water twice and brine, dried over magnesium sulfate, and concentrated in vacuo. The residue was purified by preparative thin layer chromatography (methylene chloride:methanol = 3:1, V/V) to give 3-bromo-8-[2,6-dichloro-3-[N-[(1,1-dioxothiomorpholino)acetyl]glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine (99 mg) as an amorphous.

NMR (CDCl$_3$, δ) : 2.43 (3H, s), 2.81-3.01 (2H), 3.21 (3H, s), 3.48-4.20 (10H), 5.50 (2H, s), 6.71 (1H, d, J=7Hz), 6.86 (1H, t, J=7Hz), 7.40 (1H, d, J=9Hz), 7.50 (1H, d, J=9Hz), 7.77 (1H, d, J=7Hz)

Example 93

The following compounds were obtained according to a similar manner to that of Example 60.

(1) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(thiomorpholinoacetyl)glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ) : 2.61 (3H, s), 2.70-4.30 (15H), 5.65 (1H, d, J=10Hz), 5.75 (1H, d, J=10Hz), 7.52-7.70 (4H), 8.22 (1H, dd, J=5, 3Hz)

(2) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(1-oxothiomorpholino)acetyl]glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
(3) 3-Bromo-8-[2,6-dichloro-3-[N-[(1,1-dioxothiomorpholino)acetyl]glycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.63 (3H, s), 3.00-3.19 (2H), 3.24 (3H, s), 3.35-3.57 (2H), 3.72 (1H, d, J=16Hz), 3.93 (1H, d, J=16Hz), 4.20-4.41 (2H), 4.62-4.98 (4H), 5.67 (2H, s), 7.38-7.67 (4H), 8.09 (1H, d, J=6Hz)

(4) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[[2-pyrimidinylthio]acetyl]glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.57 (3H, s), 3.26 (3H, s), 3.67 (2H, s), 3.86 (2H, s), 5.64 (1H, d, J=10Hz), 5.73 (1H, d, J=10Hz), 7.19 (1H, t, J=5Hz), 7.48-7.69 (4H), 8.21 (1H, dd, J=5, 1Hz), 8.66 (2H, d, J=5Hz)

(5) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[[phenoxyacetyl]glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.58 (3H, s), 3.30 (3H, s), 3.78 (2H, s), 4.51 (2H, s), 5.67 (1H, d, J=10Hz), 5.77 (1H, d, J=10Hz), 6.90-7.10 (3H), 7.25-7.42 (2H), 7.51-7.70 (4H), 8.21 (1H, dd, J=5, 2Hz)

(6) 3-Bromo-8-[2,6-dichloro-3-[N-[(heptafluorobutanoyl)-glycyl]-N-methylamino]benzyloxy]-2-methylimidazo-
[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$): 2.62 (3H, s), 3.28 (3H, s), 3.77 (1H, d, $J$=17Hz), 3.89 (1H, d, $J$=17Hz), 5.62 (1H, d, $J$=10Hz), 5.71 (1H, d, $J$=10Hz), 7.39-7.63 (4H), 8.09 (1H, dd, $J$=6, 1Hz)

(7) 3-Bromo-8-[2,6-dichloro-3-[N-(n-heptanoylglycyl)-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$): 0.89 (3H, t, $J$=7Hz), 1.21-1.40 (6H), 1.51-1.70 (2H), 2.26 (2H, t, $J$=7Hz), 2.58 (3H, s), 3.29 (3H, s), 3.68 (2H, s), 5.67 (1H, d, $J$=10Hz), 5.77 (1H, d, $J$=10Hz), 7.52-7.70 (4H), 8.24 (1H, dd, $J$=5, 2Hz)

(8) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-(cinnamoylglycyl)amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$): 2.60 (3H, s), 3.30 (3H, s), 3.81 (2H, s), 5.67 (1H, d, $J$=10Hz), 5.78 (1H, d, $J$=10Hz), 6.60 (1H, d, $J$=15Hz), 7.32-7.69 (10H), 8.20 (1H, dd, $J$=5, 2Hz)

(9) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(trans-3-pentenoyl)glycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$): 1.73 (3H, d, $J$=6Hz), 2.58 (3H, s), 2.99 (2H, d, $J$=6Hz), 3.29 (3H, s), 3.68 (2H, s), 5.40-5.80 (4H), 7.52-7.70 (4H), 8.25 (1H, dd, $J$=5, 1Hz)

(10) 3-Bromo-8-[3-[N-[(3-butenoyl)glycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$): 2.59 (3H, s), 3.06 (2H, d,
(11) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(4-
phenylbutanoyl)glycyl]amino]benzoyloxy]-2-
methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ): 1.83-2.02 (2H), 2.29 (2H, t, J=8Hz), 2.57 (3H, s), 2.66 (2H, t, J=8Hz), 3.28
(3H, s), 3.68 (2H, s), 5.65 (1H, d, J=10Hz), 5.71 (1H, d, J=10Hz), 7.10-7.32 (5H), 7.51-7.69
(4H), 8.22 (1H, dd, J=5, 2Hz)

(12) 3-Bromo-8-[2,6-dichloro-3-[N-[N,N-dimethyl-8-alanyl]-
glycyl]-N-methylamino]benzoyloxy]-2-methylimidazo-
[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.60 (3H, s), 2.79-2.95 (8H),
3.25 (2H, s), 3.36-3.49 (2H), 3.60 (1H, d, J=17Hz), 3.82 (1H, d, J=17Hz), 5.67 (1H, d, J=10Hz),
5.77 (1H, d, J=10Hz), 7.53-7.70 (4H), 8.24 (1H, d, J=6Hz)

(13) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[(5-
isoquinolyl)ureidoacetyl]-N-methylamino]benzoyloxy]-2-
methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.58 (3H, s), 3.29 (3H, s),
3.75 (1H, d, J=17Hz), 3.90 (1H, d, J=17Hz), 5.66
(1H, d, J=10Hz), 5.78 (1H, d, J=10Hz), 7.52-7.66
(4H), 7.91 (1H, t, J=8Hz), 8.10 (1H, d, J=8Hz),
8.20 (1H, br s), 8.50 (1H, d, J=6Hz), 8.60 (1H, d, J=8Hz), 8.88 (1H, d, J=6Hz), 9.60 (1H, s)

(14) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-(3-
pyrazolyl)ureidoacetyl]amino]benzoyloxy]-2-
methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.59 (3H, s), 3.30 (3H, s), 3.79 (1H, d, J=17Hz), 3.93 (1H, d, J=17Hz), 5.67 (1H, d, J=10Hz), 5.76 (1H, d, J=10Hz), 6.40 (1H, d, J=3Hz), 7.50-7.70 (4H), 8.12 (1H, d, J=3Hz), 8.21 (1H, dd, J=5, 3Hz)

(15) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-(4-pyrimidinyl)ureidoacetyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ) : 2.60 (3H, s), 3.29 (3H, s), 3.71 (1H, d, J=18Hz), 3.88 (1H, d, J=18Hz), 5.71 (2H, br s), 7.52-7.71 (4H), 8.07-8.79 (2H), 8.58 (1H, br s), 8.99 (1H, br s)

(16) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-(6-quinolyl)ureidoacetyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ) : 2.59 (3H, s), 3.28 (3H, s), 3.83 (1H, d, J=16Hz), 3.88 (1H, d, J=16Hz), 5.64 (1H, d, J=10Hz), 5.76 (1H, d, J=10Hz), 7.47-7.66 (4H), 7.90 (1H, dd, J=9, 5Hz), 8.04-8.46 (4H), 8.81 (1H, d, J=9Hz), 8.90 (1H, d, J=5Hz)

(17) 3-Bromo-8-[2,6-dichloro-3-[N-{[1-indolylcarbonyl]glycyl}N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

mp : 224°C (dec.)

NMR (CDCl₃-CD₃OD, δ) : 2.58 (3H, s), 3.31 (3H, s), 3.83 (1H, d, J=17Hz), 3.95 (1H, d, J=17Hz), 5.67 (1H, d, J=10Hz), 5.74 (1H, d, J=10Hz), 6.65 (1H, d, J=4Hz), 7.18-7.38 (2H), 7.53-7.70 (6H), 8.12 (1H, d, J=8Hz), 8.21 (1H, m)

(18) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-

[(morpholinocarbonyl)glycyl]amino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.58 (3H, s), 3.27 (3H, s), 3.31-3.48 (4H), 3.61-3.79 (6H), 5.63 (1H, d, J=10Hz), 5.78 (1H, d, J=10Hz), 7.52-7.69 (4H), 8.23 (1H, dd, J=5, 1Hz)

(19) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-[3-(morpholinocarbonyl)phenyl]ureidoacetyl]amino]-benzyl]oxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.55 (3H, s), 3.29 (3H, s), 3.41-3.86 (10H), 5.69 (1H, d, J=10Hz), 5.78 (1H, d, J=10Hz), 7.00 (1H, d, J=7Hz), 7.27-7.71 (7H), 8.26 (1H, d, J=5Hz)

(20) 3-Bromo-8-[3-[N-(N'-n-butylureidoacetyl)-N-methylamino]-2,6-dichlorobenzyl]oxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 0.89 (3H, t, J=6Hz), 1.20-1.41 (2H), 1.48-1.66 (2H), 2.62 (3H, s), 3.11-3.29 (5H), 3.87 (1H, d, J=17Hz), 4.00 (1H, d, J=17Hz), 5.56 (1H, d, J=10Hz), 5.71 (1H, d, J=10Hz), 7.36-7.61 (4H), 8.06 (1H, d, J=6Hz)

(21) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-[(3-quinolyl)ureidoacetyl]amino]benzyl]oxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃, δ): 2.69 (3H, s), 3.23 (3H, s), 3.94 (2H, s), 5.59 (1H, d, J=10Hz), 5.69 (1H, d, J=10Hz), 7.36-7.60 (4H), 7.69-7.79 (2H), 8.00 (1H, d, J=9Hz), 8.11 (1H, d, J=6Hz), 8.41 (1H, d, J=9Hz), 9.07 (1H, br s), 9.28 (1H, br s)

(22) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-(1-hydroxyethyl)phenyl]ureidoacetyl]-N-methylamino]benzyl]oxy]-2-
methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 1.42 (3H, d, J=6Hz), 2.53 (3H, s), 3.28 (3H, s), 3.73 (2H, s), 4.79 (1H, q, J=6Hz), 5.62 (1H, d, J=10Hz), 5.77 (1H, d, J=10Hz), 6.99 (1H, d, J=6Hz), 7.13-7.31 (3H), 7.51-7.67 (4H), 8.20 (1H, dd, J=5, 3Hz)

(23) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(methylcarbamoil)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.52 (3H, s), 3.29 (3H, s), 3.84 (2H, s), 5.64 (1H, d, J=10Hz), 5.70 (1H, d, J=10Hz), 6.65 (1H, d, J=16Hz), 7.28-7.33 (1H, m), 7.45-7.61 (6H, m), 7.77 (2H, d, J=8Hz), 8.07-8.14 (1H, m)

(24) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(dimethylcarbamoil)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.60 (3H, s), 3.02 (3H, br s), 3.12 (3H, br s), 3.30 (3H, s), 3.81 (2H, s), 5.66 (1H, d, J=10Hz), 5.77 (1H, d, J=10Hz), 6.68 (1H, d, J=15Hz), 7.37-7.70 (9H), 8.21 (1H, dd, J=5, 2Hz)

(25) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(2-methoxyethylcarbamoil)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.60 (3H, s), 3.29 (3H, s), 3.42 (3H, s), 3.61 (4H, s), 3.81 (2H, s), 5.67 (1H, d, J=10Hz), 5.77 (1H, d, J=10Hz), 6.69 (1H, d, J=15Hz), 7.50-7.70 (7H), 7.82 (2H, d, J=8Hz), 8.21 (1H, dd, J=5, 3Hz)

(26) 8-[3-[N-[4-[N,N-Bis(2-methoxyethyl)carbamoil]-
cinnamoylglycyl]-N-methylamino]-2,6-
dichlorobenzylolxy]-3-bromo-2-methylimidazo-
[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.60 (3H, s), 3.21-3.87
(16H), 5.68 (1H, d, J=10Hz), 5.77 (1H, d,
J=10Hz), 6.65 (1H, d, J=15Hz), 7.34-7.70 (9H),
8.21 (1H, m)

(27) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(4-
pyridylcarbamoyl)cinnamoylglycyl]amino]benzylolxy]-2-
methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.60 (3H, s), 3.30 (3H, s),
3.84 (2H, br s), 5.67 (1H, d, J=10Hz), 5.77 (1H,
d, J=10Hz), 6.72 (1H, d, J=15Hz), 7.44-7.77
(7H), 8.11 (2H, d, J=8Hz), 8.21 (1H, t, J=4Hz),
8.49-8.61 (4H)

(28) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(2-oxo-1-
pyrrolidinyl)cinnamoylglycyl]amino]benzylolxy]-2-
methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.18 (2H, quint, J=7.5Hz),
2.54-2.68 (5H, m), 3.29 (3H, s), 3.81 (2H, s),
3.89 (2H, t, J=7.5Hz), 5.65 (1H, d, J=10Hz),
5.74 (1H, d, J=10Hz), 6.50 (1H, d, J=16Hz),
7.44-7.68 (9H, m), 8.11 (1H, t, J=4Hz)

(29) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(methoxyacetamido)-
cinnamoylglycyl]-N-methylamino]benzylolxy]-2-
methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.66 (3H, s), 3.29 (3H, s),
3.51 (3H, s), 3.82 (2H, s), 4.03 (2H, s), 5.60-
5.77 (2H, m), 6.50 (1H, d, J=16Hz), 7.40-7.65
(9H, m), 8.04-8.15 (1H, m)

(30) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-
(propionamido)cinnamoylglycylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CD$_3$OD, $\delta$): 1.19 (3H, t, J=7.5Hz), 2.40 (2H, q, J=7.5Hz), 2.50 (3H, s), 3.24 (3H, s), 3.79 (1H, d, J=16Hz), 3.86 (1H, d, J=16Hz), 5.69-5.80 (2H, m), 6.60 (1H, d, J=15Hz), 7.36-7.80 (9H, m)

(31) 8-[3-[N-((4-(Acetamido)cinnamoylglycyl)-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$): 2.18 (3H, s), 2.58 (3H, s), 3.28 (3H, s), 3.76 (1H, d, J=16Hz), 3.90 (1H, d, J=16Hz), 5.65 (2H, s), 6.43 (1H, d, J=16Hz), 7.25-7.39 (2H, m), 7.39-7.64 (7H, m), 8.03-8.16 (1H, m)

(32) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(N-methylacetamido)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$): 1.91 (3H, br s), 2.61 (3H, s), 3.28 (3H, br s), 3.31 (3H, s), 3.81 (2H, br s), 5.66 (1H, br d, J=10Hz), 5.78 (1H, br d, J=10Hz), 6.63 (1H, d, J=15Hz), 7.22 (2H, d, J=9Hz), 7.46-7.70 (7H), 8.20 (1H, br s)

(33) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-[N-(3-pyridyl)methyl]acetamido)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$): 1.99 (3H, s), 2.60 (3H, s), 3.76 (1H, d, J=18Hz), 3.88 (1H, d, J=18Hz), 5.10 (2H, s), 5.68 (1H, d, J=10Hz), 5.78 (1H, d, J=10Hz), 6.70 (1H, d, J=15Hz), 7.20 (2H, d, J=8Hz), 7.47-7.70 (7H), 8.04 (1H, dd, J=8, 6Hz), 8.22 (1H, dd, J=6, 1Hz), 8.51 (1H, br d, J=8Hz),
(34) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(isonicotinoylamino)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ) : 2.60 (3H, s), 3.30 (3H, s), 3.82 (2H, s), 5.66 (1H, d, J=10Hz), 5.75 (1H, d, J=10Hz), 6.59 (1H, d, J=15Hz), 7.44-7.69 (7H), 7.40 (2H, d, J=9Hz), 8.20 (1H, t, J=5Hz), 8.60 (2H, d, J=6Hz), 9.00 (2H, d, J=6Hz)

(35) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(ethoxycarbonylacetamido)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ) : 1.34 (3H, t, J=7.5Hz), 2.55 (3H, s), 3.28 (3H, s), 3.50 (2H, s), 3.77 (1H, d, J=18Hz), 3.89 (1H, d, J=18Hz), 4.26 (2H, q, J=7.5Hz), 5.60 (2H, s), 6.45 (1H, d, J=16Hz), 7.20-7.35 (2H, m), 7.35-7.64 (7H, m), 7.98 (1H, d, J=6Hz)

(36) 8-[3-[N-[4-(Benzamido)cinnamoylglycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ, 3:1 V/V) : 2.60 (3H, s), 3.29 (3H, s), 3.83 (2H, s), 5.66 (1H, d, J=9Hz), 5.74 (1H, d, J=9Hz), 6.56 (1H, d, J=15Hz), 7.36-7.70 (10H, m), 7.78 (2H, d, J=9Hz), 7.93 (2H, dd, J=9, 0.5Hz), 8.21 (1H, m)

(37) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(4-pyridy lacetamido)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ, 3:1 V/V) : 2.59 (3H, s), 3.29 (3H, s), 3.78 (1H, d, J=16Hz), 3.84 (1H, d,
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J=16Hz), 4.19 (2H, s), 5.64 (1H, d, J=10Hz), 5.72 (1H, d, J=10Hz), 6.51 (1H, d, J=15Hz), 7.38-7.77 (9H, m), 8.10-8.29 (3H, m), 8.72 (2H, d, J=7Hz)

(38) 3-Bromo-8-[2,6-dichloro-3-[N-[4-{methanesulfonamido)-cinnamoylglycyl]-N-methylamino]benzyloxy]-2-
methyylimidazolo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.60 (3H, s), 3.00 (3H, s), 3.25 (3H, s), 3.82 (2H, s), 5.63 (1H, d, J=10Hz), 5.73 (1H, d, J=10Hz), 6.51 (1H, d, J=16Hz), 7.22 (2H, d, J=8Hz), 7.38-7.65 (7H, m), 8.07-8.19 (1H, m)

(39) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(methylureido)cinnamoylglycyl]amino]benzyloxy]-2-
methyylimidazolo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.31 (3H, s), 2.80 (3H, s), 3.27 (3H, s), 3.73 (1H, d, J=17Hz), 3.95 (1H, d, J=17Hz), 5.63 (2H, s), 6.34 (1H, d, J=16Hz), 7.19 (2H, d, J=8Hz), 7.24-7.43 (3H, m), 7.43-7.64 (4H, m), 8.05-8.14 (1H, m)

(40) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-[3-(3-
pyridyl)ureido)cinnamoylglycyl]amino]benzyloxy]-2-
methyylimidazolo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.60 (3H, s), 3.30 (3H, s), 3.76 (1H, d, J=17Hz), 3.91 (1H, d, J=17Hz), 5.66 (1H, d, J=10Hz), 5.75 (1H, d, J=10Hz), 6.50 (1H, d, J=15Hz), 7.36-7.70 (9H), 7.89 (1H, dd, J=8, 5Hz), 8.20 (1H, t, J=5Hz), 8.31 (1H, d, J=6Hz), 8.63 (1H, br d, J=8Hz), 9.34 (1H, d, J=1Hz)

(41) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(morpholino-
carbonylamino)cinnamoylglycyl]amino]benzyloxy]-2-

methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.60 (3H, s), 3.30 (3H, s),
3.50-3.60 (4H), 3.70-3.85 (6H), 5.66 (1H, d,
J=10Hz), 5.76 (1H, d, J=10Hz), 6.49 (1H, d,
J=15Hz), 7.39-7.70 (9H), 8.21 (1H, dd, J=5, 3Hz)

(42) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-[3-(4-
pyridylcarbamoyl)phenyl]ureidoacetyl]amino]-
benzyl]oxy]-2-methylimidazo[1,2-a]pyridine
hydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.56 (3H, s), 3.27 (3H, s),
3.83 (2H, s), 5.60 (1H, d, J=10Hz), 5.70 (1H, d,
J=10Hz), 7.22-7.38 (1H, m), 7.38-7.67 (6H, m),
7.80-7.93 (1H, m), 8.05-8.16 (1H, m), 8.45 (2H, d,
J=7.5Hz), 8.53 (2H, d, J=7.5Hz)

(43) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-[3-(1-
pyrrolidinylcarbonyl)phenyl]ureidoacetyl]amino]-
benzyl]oxy]-2-methylimidazo[1,2-a]pyridine
hydrochloride
NMR (DMSO-d₆, δ): 1.71-1.91 (4H, m), 2.39 (3H, s),
3.13 (3H, s), 5.51-5.65 (2H, m), 6.42-6.51 (1H, m),
7.00 (1H, d, J=7Hz), 7.19-7.44 (3H, m),
7.51-7.75 (2H, m), 7.79-7.88 (2H, m), 8.24 (1H, d,
J=6Hz), 9.07-9.16 (1H, m)

(44) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-[3-[N-
methyl-N-(3-pyridylmethyl)carbamoyl]phenyl]-
ureidoacetyl]amino]benzyl]oxy]-2-methylimidazo-
[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.54 (3H, s), 3.10 (3H, s),
3.25 (3H, s), 3.65-3.98 (2H, overlapped with
H₂O), 4.89 (2H, br s), 5.61 (1H, d, J=10Hz),
5.72 (1H, d, J=10Hz), 7.02 (1H, d, J=7Hz), 7.29
(1H, t, J=7Hz), 7.41-7.62 (6H), 8.09 (1H, dd,
J=9, 6Hz), 8.18 (1H, t, J=5Hz), 8.60 (1H, br s),
8.80 (1H, d, J=5Hz), 8.91 (1H, br s)

(45) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-[4-
meptyl-1-piperazinylcarbonyl]phenyl]ureidoacetyl]-
amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 2.54 (3H, s), 2.90 (3H, s),
3.00-3.80 (13H), 5.62 (1H, d, J=10Hz), 5.74 (1H, d, J=10Hz), 7.01 (1H, d, J=7Hz), 7.26-7.68 (7H ), 8.20 (1H, m)

(46) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[N'-[3-(3-
pyridyl)methylcarbamoyl]phenyl]ureidoacetyl]amino]-
benzyloxy]-2-methylimidazo[1,2-a]pyridine
dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 2.51 (3H, s), 3.26 (3H, s),
3.76 (2H, s), 4.76 (2H, s), 5.62 (1H, d, J=10Hz), 5.72 (1H, d, J=10Hz), 7.29 (1H, t, J=8Hz), 7.49-7.74 (7H), 7.99 (1H, dd, J=7, 5Hz),
8.18 (1H, t, J=4Hz), 8.61-8.72 (2H), 8.90 (1H, br s)

(47) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-(dimethylamino)-phenyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl$_3$-CD$_3$OD, $\delta$) : 2.59 (3H, s), 3.21 (6H, s),
3.29 (3H, s), 3.66 (1H, d, J=17Hz), 3.89 (1H, d, J=17Hz), 5.64 (1H, d, J=10Hz), 5.76 (1H, d, J=10Hz), 7.21-7.68 (7H), 7.89 (1H, br s), 8.21 (1H, t, J=4Hz)

(48) 3-Bromo-8-[2,6-dichloro-3-[N-[4-ethylcarbamoyl]-
cinnamoylglycy]a-N-methylamino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 1.27 (3H, t, J=7Hz), 2.60 (3H, s), 3.30 (3H, s), 3.47 (2H, q, J=7Hz), 3.83 (2H, s), 5.68 (1H, d, J=10Hz), 5.77 (1H, d, J=10Hz), 6.70 (1H, d, J=15Hz), 7.49-7.70 (7H), 7.82 (2H, d, J=9Hz), 8.22 (1H, dd, J=5, 3Hz)

(49) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(N-ethyl-N-methylcarbamoyl)cinnamoylglycyl]N-methylamino]-benzylloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ): 1.11-1.33 (3H), 2.60 (3H, s), 2.94-3.12 (3H), 3.23-3.42 (4H), 3.59 (1H, m), 3.81 (2H, s), 5.68 (1H, d, J=10Hz), 5.77 (1H, d, J=10Hz), 6.68 (1H, d, J=15Hz), 7.37-7.69 (9H), 8.21 (1H, dd, J=5, 3Hz)

(50) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(1-pyrrolidinylcarbonyl)cinnamoylglycyl]amino]-benzylloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ): 1.78-2.07 (4H, m), 2.63 (3H, s), 3.28 (3H, s), 3.34-3.51 (2H, m), 3.51-3.70 (2H, m), 3.76 (1H, d, J=18Hz), 3.90 (1H, d, J=18Hz), 5.63 (1H, d, J=10Hz), 5.72 (1H, d, J=10Hz), 6.63 (1H, d, J=16Hz), 7.38-7.67 (9H, m), 8.03-8.17 (1H, m)

(51) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(morpholinocarbonyl)cinnamoylglycyl]amino]benzylloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.61 (3H, s), 3.30 (3H, s), 3.35-3.95 (10H, m), 5.64 (1H, d, J=10Hz), 5.74 (1H, d, J=10Hz), 6.65 (1H, d, J=16Hz), 7.40 (2H, d, J=8Hz), 7.46-7.65 (7H, m), 8.09-8.18 (1H, m)
(52) 3-Bromo-8-[2,6-dichloro-3-[N-[4-[N-(2-methoxyethyl)-N-methylcarbamoyl]cinnamoylglycyl]-N-methylamino]-benzyl]oxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ) : 2.62 (3H, s), 3.00-3.15 (3H, m), 3.60-3.93 (4H, m), 5.63 (1H, d, J=10Hz), 5.73 (1H, d, J=10Hz), 6.61 (1H, d, J=16Hz), 7.41 (2H, d, J=8Hz), 7.47-7.65 (7H, m), 8.08-8.20 (1H, m)

(53) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(isopropylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyl]oxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ) : 1.28 (6H, d, J=6Hz), 2.63 (3H, s), 3.29 (3H, s), 3.77 (1H, d, J=16Hz), 3.90 (1H, d, J=16Hz), 4.25 (1H, m), 5.63 (1H, d, J=16Hz), 5.72 (1H, d, J=10Hz), 6.65 (1H, d, J=16Hz), 7.41-7.63 (7H, m), 7.74 (2H, d, J=8Hz), 8.04-8.17 (1H, m)

(54) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(n-propylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyl]oxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ) : 0.99 (3H, t, J=7.5Hz), 1.65 (2H, m), 2.61 (3H, s), 3.30 (3H, s), 3.34-3.47 (2H, m), 3.84 (2H, s-like), 5.63 (1H, d, J=10Hz), 5.72 (1H, d, J=10Hz), 6.66 (1H, d, J=16Hz), 7.40-7.65 (7H, m), 7.77 (2H, d, J=8Hz), 8.05-8.16 (1H, m)

(55) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(3-methoxypropylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyl]oxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ) : 1.90 (2H, quint, J=6Hz), 2.64 (3H, s), 3.28 (3H, s), 3.40 (3H, s), 3.55 (4H,
(56) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(2-ethoxyethyl-carbamoyl)cinnamoyl]glycyl]-N-methylamino]benzyloxy-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ): 1.22 (3H, t, J=7Hz), 3.63 (3H, s), 3.50-3.71 (6H, m), 3.88 (1H, d, J=18Hz), 3.90 (1H, d, J=18Hz), 5.65 (1H, d, J=10Hz), 5.72 (1H, d, J=10Hz), 6.65 (1H, d, J=16Hz), 7.46-7.65 (7H, m), 7.79 (2H, d, J=8Hz), 8.05-8.14 (1H, m)

(57) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(2-hydroxyethyl-carbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.60 (3H, s), 3.26 (3H, s), 3.57 (2H, t, J=5Hz), 3.78 (2H, t, J=5Hz), 3.85 (2H, s), 5.59-5.74 (2H, m), 6.64 (2H, d, J=16Hz), 7.40-7.64 (7H, m), 7.82 (2H, d, J=8Hz), 8.03-8.15 (1H, m)

(58) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(diethylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ): 1.03-1.33 (6H, m), 2.65 (3H, s), 3.13-3.35 (5H, m), 3.45-3.63 (2H, m), 3.77 (1H, d, J=18Hz), 3.90 (1H, d, J=18Hz), 5.63 (1H, d, J=10Hz), 5.71 (1H, d, J=10Hz), 6.60 (1H, d, J=16Hz), 7.33 (2H, d, J=8Hz), 7.41-7.63 (7H, m), 8.02-8.13 (1H, m)

(59) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(2-oxopiperidino)cinnamoylglycyl]amino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 1.90-2.03 (4H, m), 2.50-2.66 (5H, m), 3.60-3.72 (2H, m), 3.80 (2H, s), 5.65 (1H, d, J=10Hz), 5.75 (1H, d, J=10Hz), 6.55 (1H, d, J=16Hz), 7.27 (2H, d, J=8Hz), 7.45-7.67 (7H, m), 8.09-8.21 (1H, m)

(60) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(N-methyl-2-methoxyacetamido)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ, 3:1 V/V): 2.60 (3H, s), 3.30 (6H, s), 3.35 (3H, s), 3.76-3.96 (4H, m), 5.68 (1H, d, J=10Hz), 5.76 (1H, d, J=9Hz), 6.69 (1H, d, J=15Hz), 7.26 (2H, br d, J=8Hz), 7.48-7.73 (7H, m), 8.24 (1H, br d, J=8Hz)

(61) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-(1-pyrrolyl)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.61 (3H, s), 3.30 (3H, s), 3.81 (2H, s), 5.68 (1H, d, J=10Hz), 5.78 (1H, d, J=9Hz), 6.39 (2H, s), 6.60 (1H, d, J=15Hz), 7.38-7.70 (11H), 8.20 (1H, dd, J=5, 3Hz)

(62) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(3-methoxypropionamido)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 2.55-2.70 (5H, m), 3.30 (3H, s), 3.44 (3H, s), 3.76 (2H, t, J=6Hz), 3.83 (2H, s), 5.67 (2H, s), 6.46 (1H, d, J=16Hz), 7.33-7.63 (9H, m), 8.02-8.14 (1H, m)

(63) 8-[3-[N-[4-(Acetamidoacetamido)cinnamoylglycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-
methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.07 (3H, s), 2.56 (3H, s),
3.27 (3H, s), 3.75 (1H, d, J=17Hz), 3.88 (1H, d,
J=17Hz), 4.00 (2H, s), 5.59-5.75 (2H, m), 6.48
(1H, d, J=16Hz), 7.35-7.45 (2H, m), 7.45-7.67
(7H, m), 8.08-8.19 (1H, m)

(64) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[4-[(dimethyl-
amino)acetamido]cinnamoylglycyl]amino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.58 (3H, s), 3.04 (6H, s),
3.27 (3H, s), 3.83 (2H, br s), 4.21 (2H, br s),
5.64 (2H, br s), 6.46 (1H, d, J=16Hz), 7.26-7.67
(9H, m), 8.06-8.16 (1H, m)

(65) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(succinimido)-
cinnamoylglycyl]-N-methylamino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.60 (3H, s), 2.95 (4H, s),
3.30 (3H, s), 3.81 (2H, s), 5.68 (1H, br d,
J=10Hz), 5.78 (1H, br d, J=10Hz), 6.64 (1H, d,
J=16Hz), 7.35 (2H, d, J=9Hz), 7.50-7.60 (7H),
8.21 (1H, m)

(66) 3-Bromo-8-[2,6-dichloro-3-[N-[4-[(E)-3-(
ethoxycarbonyl)acrylamido]cinnamoylglycyl]-N-
methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
hydrochloride
NMR (CDCl₃-CD₃OD, δ, 3:1, V/V) : 1.36 (3H, t,
J=7.5Hz), 2.49 (3H, s), 3.28 (3H, s), 3.37 (2H,
q, J=7.5Hz), 3.81 (2H, s), 5.66 (1H, d, J=9Hz),
5.75 (1H, d, J=9Hz), 6.56 (1H, d, J=15Hz), 6.91
(1H, d, J=15Hz), 7.17 (1H, d, J=15Hz), 7.36-7.80
(9H, m), 8.20 (1H, br d, J=4Hz)
(67) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(2-oxo-3-oxazolidinyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ) :
- 2.60 (3H, s), 3.28 (3H, s), 3.81 (2H, s), 4.11 (2H, dd, J=6, 8Hz), 4.52 (2H, dd, J=6, 8Hz), 5.64 (1H, d, J=10Hz), 5.74 (1H, d, J=10Hz), 6.54 (1H, d, J=16Hz), 7.37-7.69 (9H, m), 8.07-8.20 (1H, m)

(68) 8-[3-[N-[N'-[3-[N,N-Bis(2-methoxyethyl)carbamoyl]phenyl]ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ) :
- 2.56 (3H, s), 3.22 (3H, s), 3.52-3.73 (4H, m), 3.79 (1H, d, J=18Hz), 4.01 (1H, d, J=18Hz), 5.58 (1H, d, J=10Hz), 5.65 (1H, d, J=10Hz), 6.88 (1H, d, J=7.5Hz), 7.20 (1H, t, J=8Hz), 7.33-7.56 (6H, m), 8.06 (1H, d, J=6Hz)

(69) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-(2-methoxyethyl)carbamoyl]phenyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ) :
- 2.58 (3H, s), 3.24 (3H, s), 3.38 (3H, s), 3.55 (4H, s-like), 3.88 (2H, s), 5.61 (2H, s), 7.19 (1H, t, J=8Hz), 7.25-7.40 (1H, m), 7.40-7.54 (5H, m), 7.60-7.66 (1H, m), 8.07 (1H, d, J=6Hz)

(70) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-[N-(2-methoxyethyl)-N-methylcarbamoyl]phenyl]ureidoacetyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ) :
- 2.53 (3H, s), 2.94-3.07 (3H,
m), 3.55-3.69 (2H, br peak), 3.75 (1H, d, 
J=16Hz), 3.87 (1H, d, J=16Hz), 5.58 (1H, d, 
J=10Hz), 5.69 (1H, d, J=10Hz), 6.92 (1H, d, 
J=7.5Hz), 7.22 (1H, t, J=7.5Hz), 7.28-7.59 (6H, 
m), 8.05-8.15 (1H, m)

(71) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-[N,N-bis(2-
ethoxyethyl)carbamoyl]phenyl]ureidoacetyl]-N-
methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine 
hydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ) : 1.05-1.28 (6H, m), 2.54 (3H, 
s), 3.25 (3H, s), 3.30-3.80 (13H, m), 3.86 (1H, 
d, J=17.5Hz), 5.58 (1H, d, J=10Hz), 5.67 (1H, d, 
J=10Hz), 6.94 (1H, d, J=7.5Hz), 7.15-7.35 (2H, 
m), 7.35-7.60 (5H, m), 8.07 (1H, d, J=6Hz)

(72) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-[N,N-bis(2-
hydroxyethyl)carbamoyl]phenyl]ureidoacetyl]-N-
methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine 
hydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ) : 2.55 (3H, s), 3.23 (3H, s), 
3.53-3.75 (4H, br peak), 3.75-3.94 (4H, br 
peak), 5.57 (1H, d, J=10Hz), 5.67 (1H, d, 
J=10Hz), 6.99 (1H, d, J=7.5Hz), 7.21 (1H, t, 
J=7.5Hz), 7.25-7.60 (6H, m), 8.08 (1H, d, J=6Hz)

(73) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-
(methoxyacetamido)phenyl]ureidoacetyl]-N-
methylamino]benzyloxy]-2-methylimidazo[1,2-a]- 
pyridine hydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ) : 2.54 (3H, s), 3.28 (3H, s), 
3.51 (3H, s), 3.74 (2H, s), 3.92-4.09 (2H, 
overlapped with H$_2$O), 5.63 (1H, d, J=10Hz), 5.74 
(1H, d, J=10Hz), 7.13-7.20 (3H), 7.51-7.66 (5H), 
8.19 (1H, t, J=3Hz)
(74) 3-Bromo-8-[2,6-dichloro-3-[N-[N'-[3-
(ethoxycarbonyl)acetamido]phenyl]ureidoacetyl]-N-
methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
hydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ): 1.31 (3H, t, J=7Hz), 2.52 (3H, s), 3.28 (3H, s), 3.47 (2H, s), 3.75 (2H, br s), 4.23 (2H, q, J=7Hz), 5.63 (1H, br d, J=10Hz), 5.73 (1H, br d, J=10Hz), 7.11-7.20 (3H), 7.49-7.66 (5H), 8.18 (1H, br t, J=4Hz)

(75) 3-Chloro-8-[2,6-dichloro-3-[N-4-(dimethylcarbamoyl)-
cinnamoylglycyl]-N-methylamino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ): 2.60 (3H, s), 3.02 (3H, br s), 3.12 (3H, br s), 3.30 (3H, s), 3.81 (2H, s), 5.68 (1H, d, J=10Hz), 5.78 (1H, d, J=10Hz), 6.69 (1H, d, J=16Hz), 7.41-7.70 (9H), 8.20 (1H, t, J=4Hz)

(76) 3-Chloro-8-[2,6-dichloro-3-[N-methyl-N-[4-
(methylcarbamoyl)cinnamoylglycyl]amino]benzyloxy]-2-
methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ): 2.61 (3H, s), 3.98 (3H, s), 3.29 (3H, s), 3.84 (2H, s), 5.62 (1H, d, J=10Hz), 5.70 (1H, d, J=10Hz), 6.64 (1H, d, J=16Hz), 7.42-7.66 (7H, m), 7.77 (2H, d, J=8Hz), 8.02-8.14 (1H, m)

(77) 3-Chloro-8-[2,6-dichloro-3-[N-[4-[(2-
methoxyethyl)carbamoyl]cinnamoylglycyl]-N-
methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine
hydrochloride

NMR (CDCl$_3$-CD$_3$OD, δ): 2.61 (3H, s), 3.29 (3H, s), 3.41 (3H, s), 3.47-3.73 (4H, overlapped with H$_2$O), 3.82 (2H, br s), 5.65 (1H, d, J=10Hz),
5.74 (1H, d, J=10Hz), 6.68 (1H, d, J=15Hz), 7.48-7.66 (7H), 7.80 (2H, br d, J=9Hz), 8.14 (1H, t, J=4Hz)

(78) 3-Chloro-8-[2,6-dichloro-3-[N-[4-(N-ethyl-N-methylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 1.07-1.33 (3H, m), 2.61 (3H, s), 3.27 (3H, s), 3.44-3.66 (2H, m), 3.76 (1H, d, J=18Hz), 3.88 (1H, d, J=18Hz), 5.63 (1H, d, J=10Hz), 5.73 (1H, d, J=10Hz), 6.61 (1H, d, J=16Hz), 7.23-7.44 (2H, m), 7.44-7.63 (7H, m), 8.01-8.15 (1H, m)

(79) 3-Chloro-8-[2,6-dichloro-3-[N-[4-(ethylcarbamoyl)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ): 1.25 (3H, t, J=7.5Hz), 2.61 (3H, s), 3.26 (3H, s), 3.45 (2H, q, J=7.5Hz), 3.78 (1H, d, J=17.5Hz), 3.90 (1H, d, J=17.5Hz), 5.63 (1H, d, J=10Hz), 5.70 (1H, d, J=10Hz), 6.64 (1H, d, J=16Hz), 7.41-7.64 (7H, m), 7.76 (2H, d, J=8Hz), 8.00-8.13 (1H, m)

(80) 3-Chloro-8-[2,6-dichloro-3-[N-methyl-N-[4-(2-oxo-1-pyrrolidinyl)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃, δ): 2.20 (2H, quint, J=7Hz), 2.55-2.70 (5H, m), 3.28 (3H, s), 3.80 (3H, s), 3.90 (2H, t, J=7.5Hz), 5.65 (1H, d, J=10Hz), 5.74 (1H, d, J=10Hz), 6.51 (1H, d, J=16Hz), 7.40-7.58 (9H, m), 8.04-8.15 (1H, m)

(81) 8-[3-[N-[4-(Acetamido)cinnamoylglycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-chloro-2-
methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.19 (3H, s), 2.60 (3H, s), 3.28 (3H, s), 3.78 (1H, d, J=18Hz), 3.89 (1H, d, J=18Hz), 5.67 (2H, s), 6.46 (1H, d, J=16Hz), 7.24-7.64 (9H, m), 8.02-8.13 (1H, m)

(82) 3-Chloro-8-[2,6-dichloro-3-[N-methyl-N-[4-(3-methylureido)cinnamoylglycyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.58 (3H, s), 2.81 (3H, s), 3.29 (3H, s), 3.74 (1H, d, J=17Hz), 3.83-4.00 (1H, overlapped with H₂O), 5.63 (1H, d, J=10Hz), 5.71 (1H, d, J=10Hz), 6.40 (1H, d, J=16Hz), 7.28-7.46 (5H), 7.51-7.68 (4H), 8.18 (1H, dd, J=5.1Hz)

(83) 3-Chloro-8-[2,6-dichloro-3-[N-[4-(methoxyacetamido)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 1.23 (3H, t, J=7.5Hz), 2.42 (2H, q, J=7.5Hz), 2.59 (3H, s), 3.28 (3H, s), 3.76 (1H, d, J=17.5Hz), 3.88 (1H, d, J=17.5Hz), 5.66 (2H, s), 6.43 (1H, d, J=16Hz), 7.32-7.63 (9H, m), 8.01-8.12 (1H, m)

(85) 3-Chloro-8-[2,6-dichloro-3-[N-[4-(3-
methoxypropionamido)cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.55-2.70 (5H), 3.29 (3H, s), 3.92 (3H, s), 3.70-4.00 (4H, overlapped with H₂O), 5.61-5.69 (2H), 6.50 (1H, d, J=16Hz), 7.39-7.68 (9H), 8.18 (1H, br s)

(86) 8-[3-[N-[4-(Acetamidoacetamido)cinnamoylglycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-chloro-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.09 (3H, s), 2.59 (3H, s), 3.82 (2H, d, J=6Hz), 4.00 (2H, s), 5.68 (2H, s-like), 6.47 (2H, d, J=16Hz), 7.35-7.69 (9H, m), 8.06-8.15 (1H, m)

(87) 8-[3-[N-[N'-[3-[N,N-Bis(2-methoxyethyl)carbamoyl]phenyl]ureidoacetyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-chloro-2-methylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃-CD₃OD, δ) : 2.54 (3H, s), 3.28 (3H, s), 3.31-3.79 (16Hz), 5.63 (1H, d, J=10Hz), 5.75 (1H, d, J=10Hz), 6.99 (1H, d, J=7Hz), 7.20-7.48 (3H), 7.51-7.67 (4H), 8.18 (1H, t, J=4Hz)

(88) 3-Chloro-8-[2,6-dichloro-3-[N-methyl-N-[N'-[3-(4-pyridylcarbamoyl)phenyl]ureidoacetyl]amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride
NMR (CDCl₃-CD₃OD, δ, 3:1 V/V) : 2.55 (3H, s), 3.27 (3H, s), 3.70 (1H, d, J=15Hz), 3.79 (1H, d, J=15Hz), 5.65 (1H, d, J=9Hz), 5.75 (1H, d, J=9Hz), 7.39 (1H, d, J=8Hz), 7.51-7.75 (6H, m), 7.98 (1H, m), 8.15-8.26 (1H, m), 8.50 (2H, d, J=8Hz), 8.56 (1H, d, J=8Hz)
(89) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[(E)-3-(6-methylcarbamoyl-3-pyridyl)acryloylglycyl]amino]-benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.62 (3H, s), 3.05 (3H, s),
3.29 (3H, s), 3.70-3.85 (1H, overlapped with H₂O), 3.91 (1H, d, J=18Hz), 5.64 (1H, d, J=10Hz), 5.73 (1H, d, J=10Hz), 6.90 (1H, d, J=16Hz), 7.50-7.67 (5H), 8.11-8.27 (2H), 8.32 (1H, br d, J=8Hz), 8.82 (1H, br s)

(90) 8-[3-[N-[(E)-3-(6-Acetamido-3-pyridyl)acryloylglycyl]-N-methylamino]-2,6-dichlorobenzyloxy]-3-bromo-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.41 (3H, s), 2.60 (3H, s),
3.28 (3H, s), 3.75 (1H, d, J=18Hz), 3.91 (1H, d, J=18Hz), 5.63 (1H, d, J=10Hz), 5.75 (1H, d, J=10Hz), 6.92 (1H, d, J=16Hz), 7.40-7.65 (5H), 8.00 (1H, d, J=9Hz), 8.20 (1H, t, J=5Hz), 8.46-8.61 (2H)

(91) 3-Bromo-8-[2,6-dichloro-3-[N-[4-(2-hydroxyethoxy)-cinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD): 2.63 (3H, s), 3.29 (3H, s), 3.80 (2H, s), 3.94 (2H, t, J=5Hz), 4.10 (2H, t, J=5Hz), 5.60-5.77 (2H, m), 6.43 (1H, d, J=16Hz), 6.90 (2H, d, J=8Hz), 7.40-7.64 (7H, m), 8.07-8.17 (1H, m)

(92) 3-Bromo-8-[2,6-dichloro-3-[N-[3,4-dimethoxycinnamoylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine hydrochloride

NMR (CDCl₃-CD₃OD, δ): 2.64 (3H, s), 3.30 (3H, s),
3.80 (2H, s), 3.91 (3H, s), 3.94 (3H, s), 5.65
(1H, d, J=10Hz), 5.75 (1H, d, J=10Hz), 6.44 (1H, d, J=16Hz), 6.87 (1H, d, J=8Hz), 7.03-7.15 (2H, m), 7.39-7.65 (5H, m), 8.05-8.19 (1H, m)

(93) 3-Bromo-8-[2,6-dichloro-3-[N-methyl-N-[3,4-
(methyleneedioxy)cinnamoylglycyl]amino]benzyloxy]-2-
methyylimidazo[1,2-a]pyridine hydrochloride
NMR (CDCl₃, δ) :  2.60 (3H, s), 3.80 (3H, s), 5.65
(1H, d, J=10Hz), 5.74 (1H, d, J=16Hz), 6.00 (2H, s), 6.40 (1H, d, J=16Hz), 6.80 (1H, d, J=8Hz),
6.94-7.04 (2H, m), 7.37-7.66 (5H, m), 8.09-8.19
(1H, m)

(94) 3-Bromo-8-[2,6-dichloro-3-[N-[(E)-3-(6-
dimethylcarbamoyl-3-pyridyl)acryloylglycyl]-N-
methylamino]benzyloxy]-2-methyylimidazo[1,2-a]pyridine
dihydrochloride
NMR (CDCl₃-CD₃OD, δ) :  2.53 (3H, s), 3.02 (3H, br
s), 3.10 (3H, br s), 3.20 (3H, s), 3.70 (1H, d, J=17Hz), 3.86 (1H, d, J=17Hz), 5.56 (1H, d, J=10Hz), 5.67 (1H, d, J=10Hz), 6.85 (1H, d, J=16Hz) 7.38-7.59 (5H), 7.71 (1H, br d, J=8Hz),
8.09 (1H, t, J=4Hz), 8.21 (1H, br d, J=8Hz),
8.80 (1H, br s)

(95) 3-Bromo-8-[2,6-dichloro-3-[N-[(E)-3-(6-
ethylcarbamoyl-3-pyridyl)acryloylglycyl]-N-
methylamino]benzyloxy]-2-methyylimidazo[1,2-a]pyridine
dihydrochloride
NMR (CDCl₃-CD₃OD, δ) :  1.22 (3H, t, J=7Hz), 2.57
(3H, s), 3.46 (1H, q, J=7Hz), 3.73 (1H, d,
J=17Hz), 3.89 (1H, d, J=17Hz), 5.58 (1H, d, J=10Hz), 5.68 (1H, d, J=10Hz), 6.81 (1H, d, J=16Hz), 7.38-7.59 (5H), 8.00-8.19 (2H), 8.26
(1H, d, J=8Hz), 8.75 (1H, br s)
(96) 3-Bromo-8-[2,6-dichloro-3-[N-(E)-3-[6-(3-methoxypropylcarbamoyl)]-3-pyridyl]acryloylglycyl]-N-methylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine dihydrochloride

NMR (CDCl₃-CD₃OD, δ) : 1.85 (2H, quint, J=6Hz), 2.58 (3H, s), 3.23 (3H, s), 3.41-3.60 (4H, m), 3.88 (1H, d, J=17.5Hz), 5.67 (1H, d, J=10Hz), 6.77 (1H, d, J=16Hz), 7.37-7.60 (5H, m), 7.98-8.11 (2H, m), 8.17 (1H, d, J=8Hz), 8.66-8.74 (1H, m)

Example 94

3-Chloro-8-(2,6-dichlorophenyl)methylamino-2-methylimidazo[1,2-a]pyridine was obtained according to a similar manner to that of Example 2.

mp : 142.9°C (dec.)

Example 95

Sulfuric acid salt of 3-bromo-8-[2,6-dichloro-3-[N-methyl-N-(butyrylglycyl)amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained by mixing 3-bromo-8-[2,6-dichloro-3-[N-methyl-N-(butyrylglycyl)amino]benzyloxy]-2-methylimidazo[1,2-a]pyridine with sulfuric acid.

mp : 154-156°C

Example 96

Maleic acid salt of 3-chloro-8-[2,6-dichloro-3-[N-methyl-N-acetylamino]benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained by mixing 3-chloro-8-[2,6-dichloro-3-(N-methyl-N-acetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine with maleic acid.

mp : 193-195°C

Example 97

Methanesulfonic acid salt of 3-chloro-8-[2,6-
dichloro-3-(N-methyl-N-acetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained by mixing 3-chloro-8-[2,6-dichloro-3-(N-methyl-N-acetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine with methanesulfonic acid.

mp: 165°C (dec.)

Example 98

Oxalic acid salt of 3-chloro-8-[2,6-dichloro-3-(N-methyl-N-acetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine was obtained by mixing 3-chloro-8-[2,6-dichloro-3-(N-methyl-N-acetylamino)benzyloxy]-2-methylimidazo[1,2-a]pyridine with oxalic acid.

mp: 180-181°C

Throughout this specification and the claims which follow, unless the context requires otherwise, the word "comprise", or variations such as "comprises" or "comprising", will be understood to imply the inclusion of a stated integer or group of integers but not the exclusion of any other integer or group of integers.
THE CLAIMS DEFINING THE INVENTION ARE AS FOLLOWS:

1. The compound of the formula:

\[
\begin{array}{c}
\text{R}^3 \quad \text{N} \\
\text{Q-A-R}^4 \\
\end{array}
\]

wherein \( R^1 \) is halogen,

\( R^2 \) and \( R^3 \) are each hydrogen, lower alkyl, halo (lower) alkyl or acyl,

\( R^4 \) is aryl substituted with substituent(s) selected from the group consisting of halogen; halo (lower) alkyl; acyl; aryl substituted with halogen or cyano; ar (lower) alkyl substituted with hydroxy; lower alkoxy; nitro; amino; amino substituted with substituent(s) selected from the group consisting of lower alkyl, acyl, ar (lower) alkyl, heterocyclic (lower) alkyl, carboxy (lower) alkyl, lower alkylinomethylene and N-methylpyrrolidinylidene; a heterocyclic group and a heterocyclic group substituted with oxo; or a heterocyclic group substituted with substituent(s) selected from the group consisting of halogen; lower alkyl; halo (lower) alkyl; acyl; aryl substituted with halogen or cyano; ar (lower) alkyl substituted with hydroxy; lower alkoxy; oxo; nitro; amino; amino substituted with substituent(s) selected from the group consisting of lower alkyl, acyl, ar (lower) alkyl, heterocyclic (lower) alkyl, carboxy (lower) alkyl, lower alkylinomethylene and N-methylpyrrolidinylidene; a heterocyclic group and a heterocyclic group substituted with oxo,

\( Q \) is \( O \) or \( N-R^{11} \),

in which \( R^{11} \) is hydrogen or acyl, and

\( A \) is lower alkylene,

and pharmaceutically acceptable salts thereof.
2. A compound of claim 1, wherein

\( R^2 \) is lower alkyl,
\( R^3 \) is hydrogen and
\( Q \) is O or NH.

3. A compound of claim 2, wherein

\( R^4 \) is phenyl substituted with substituents selected from the group consisting of halogen, nitro, amino and a group of the formula:

\[
\begin{array}{c}
\text{N} \\
\text{R}^5 \\
\text{R}^10
\end{array}
\]

in which \( R^5 \) is hydrogen or lower alkyl, and
\( R^{10} \) is acyl, and
\( A \) is methylene.

4. A compound of claim 3, wherein

\( R^4 \) is phenyl substituted with one or two halogen(s) and a group of the formula:
in which
\(R^5\) is hydrogen or lower alkyl, and
\(R^\text{e}_{10}\) is an amino acid residue or an amino acid residue substituted with a substituent selected from the group consisting of lower alkyl, alkanoyl, halo(lower)alkanoyl, ar(lower)alkanoyl, aroyl, heterocyclic(lower)alkanoyl, lower alkenoyl, ar(lower)alkenoyl, lower alkoxy-ar(lower)alkenoyl, lower alkylenedioxy-ar(lower)alkenoyl, nitro-ar(lower)alkenoyl, halo-ar(lower)alkenoyl, hydroxy-ar(lower)alkenoyl, hydroxy(lower)alkoxy-ar(lower)alkenoyl, amino(lower)alkoxy-ar(lower)alkenoyl, lower alkylamino(lower)alkoxy-ar(lower)alkenoyl, heterocyclic(lower)alkoxy-ar(lower)alkenoyl, heterocyclic-ar(lower)alkenoyl optionally having oxo, amino-ar(lower)alkenoyl, lower alkylamino-ar(lower)alkenoyl, lower alkanoylamino-ar(lower)alkenoyl, N-(lower alkanoyl)-N-(lower alkyl)amino-ar(lower)-alkenoyl, hydroxy(lower)alkanoylamino-ar(lower)alkenoyl, lower alkoxy(lower)-alkanoylamino-ar(lower)alkenoyl, halo(lower)alkanoylamino-ar(lower)alkenoyl, amino(lower)alkanoylamino-ar(lower)alkenoyl, lower alkylamino(lower)alkanoylamino-ar(lower)alkenoyl, lower alkanoylamino-ar(lower)alkenoyl, lower alkanoylamino-ar(lower)alkenoyl,
carboxy(lower) alkanoylamino-ar(lower) alkenoyl, lower
alkoxycarbonyl(lower) alkanoylamino-ar(lower) alkenoyl, lower
alkoxycarbonyl(lower) alkanoylamino-ar(lower) alkenoyl,
halo(lower) alkoxy carbonylamino-ar(lower) alkenoyl, heterocyclic(lower) alkanoylamino-
ar(lower) alkenoyl,
aroylamino-ar(lower) alkenoyl,
heterocyclic carbonylamino-ar(lower) alkenoyl, lower
alkylsulfonylamino-ar(lower) alkenoyl, N-[lower alkoxylower) alkanoyl]-N-(lower alkyl) amino-ar(lower) alkenoyl, N-(lower alkyl)alnoyloyl]-N-[heterocyclic(lower) alkyl]amino-
ar(lower) alkenoyl, ureido-ar(lower) alkenoyl, lower alkylureido-ar(lower) alkenoyl,
heterocyclicureido-ar(lower) alkenoyl, lower alkanoyl-ar(lower) alkenoyl, carboxy-ar(lower) alkenoyl,
lower alkoxycarbonyl-ar(lower) alkenoyl, carbamoyl-ar(lower) alkenoyl, lower
alkyl carbamoyl-ar(lower) alkenoyl, hydroxy(lower) alkyl carbamoyl-
ar(lower) alkenoyl, N-[hydroxy(lower) alkyl]-N-(lower alkyl) carbamoyl-ar(lower) alkenoyl, lower
alkoxy(lower) alkyl carbamoyl-ar(lower) alkenoyl, N-(lower alk oxy(lower) alkyl)-N-(lower alkyl) carbamoyl-
ar(lower) alkenoyl, heterocyclic carbamoyl-ar(lower) alkenoyl, heterocyclic carbonyl-ar(lower) alkenoyl,
ar(lower) alkynoyl, heterocyclic(lower) alkenoyl, heterocyclic thio(lower) alkanoyl,
amino-heterocyclic(lower)alkenoyl, lower
alkylamino-heterocyclic(lower)alkenoyl, lower
alkanoylamino-heterocyclic(lower)alkenoyl, lower
alkylureido-heterocyclic(lower)alkenoyl, lower
carboxy-heterocyclic(lower)alkenoyl, lower
alkoxycarbonyl-heterocyclic(lower)alkenoyl, lower
alkylcarbamoyl-heterocyclic(lower)alkenoyl, lower
alkoxy(lower)alkylcarbamoyl-heterocyclic-
(lower)alkenoyl, hydroxy(lower)alkylcarbamoyl-
20 heterocyclic(lower)alkenoyl, lower
heterocycliccarbonyl,
cyclo(lower)alkylcarbonyl, lower
alkoxycarbonyl, aryloxy carbonyl, lower
aryloyl(lower)alkanoyl, aroyl,
nitro-aryloxycarbonyl, carbamoyl, lower
alkylcarbamoyl, lower
alkoxycarbonyl(lower)alkylcarbamoyl, lower
alkenylcarbamoyl, cyclo(lower)alkylcarbamoyl,
arylcarbamoyl, lower alkoxy-arylcarbamoyl,
halo(lower)alkyl-arylcarbamoyl,
halo-arylcarbamoyl, lower
alkanoyl-arylcarbamoyl,
hydroxy(lower)alkyl-arylcarbamoyl,
heterocycliccarbonyl-arylcarbamoyl,
carboxy-arylcarbamoyl, lower
alkoxycarbonyl-arylcarbamoyl, lower
arylcarbamoyl-arylcarbamoyl,
nitro-arylcarbamoyl, cyano-arylcarbamoyl,
amino-arylcarbamoyl, lower
alkylamino-arylcarbamoyl, lower
alkanoylamino-arylcarbamoyl, N-(lower
alkanoyl)-N-(lower alkyl)amino-arylcarbamoyl,
lower alkoxy(lower)alkanoylamino-
arylcarbamoyl, lower alkoxy carbonyl(lower)-
alkanoylamino-arylcarbamoyl, carboxyamino-arylcarbamoyl, lower alkoxy carbonylamino-arylcarbamoyl, ureido-arylcarbamoyl, lower alkylureido-arylcarbamoyl, hydroxyimino(lower)alkyl-arylcarbamoyl, lower alkoxyimino(lower)alkyl-arylcarbamoyl, lower alkylhydr azono(lower)alkyl-arylcarbamoyl, heterocyclic-arylcarbamoyl optionally having oxo, heterocycliccarbonylarylcarbamoyl having lower alkyl, heterocycliccarbonylarylcarbamoyl having aryl, heterocycliccarbonylarylcarbamoyl having a heterocyclic group, heterocycliccarbonylarylcarbamoyl having lower alkanoyl, heterocycliccarbonylarylcarbamoyl having lower alkoxy carbonyl, heterocycliccarbonylarylcarbamoyl having lower alkylamino, heterocycliccarbonylarylcarbamoyl having lower alkylcarbamoyl, hydroxy(lower)alkylcarbamoyl-arylcarbamoyl, N-[hydroxy(lower)alkyl]-N-(lower alkyl)carbamoyl-arylcarbamoyl, lower alkoxy(lower)alkylcarbamoyl-arylcarbamoyl, N-[lower alkoxy(lower)alkyl]-N-(lower alkyl)carbamoylarylcarbamoyl, lower alkylamino(lower)alkylcarbamoyl-arylcarbamoyl- N-[lower alkylamino(lower)alkyl]-N-(lower alkyl)carbamoylarylcarbamoyl, heterocycliccarbonylarylcarbamoyl, N-(heterocyclic)-N-(lower alkyl)carbamoylarylcarbamoyl, heterocyclic-(lower)alkylcarbamoylarylcarbamoyl, N-[heterocyclic(lower)alkyl]-N-(lower alkyl)carbamoylarylcarbamoyl, N-[heterocyclic(lower)alkyl]-N-[lower
alkoxy(lower)alkyl]carbamoyl-arylcarbamoyl, arylcarbamoyl-arylcarbamoyl, lower alkylaminoarylcarbamoyl-arylcarbamoyl, arylthiocarbamoyl, ar(lower)alkylcarbamoyl, aroylcarbamoyl, heterocycliccarbamoyl, heterocyclic(lower)alkylcarbamoyl, arylaminocarbamoyl, ar(lower)alkenylsulfonyl, lower alkylsulfonyl, phthaloyl, amino acid residue, amino acid residue substituted with lower alkyl, amino acid residue substituted with a heterocyclic group, amino acid residue substituted with heterocyclic(lower)alkyl, amino acid residue substituted with cycloalkyl, amino acid residue substituted with aryl, amino acid residue substituted with alkanoyl, amino acid residue substituted with lower alkoxy carbonyl, amino acid residue substituted with ar(lower)alkyl and amino acid residue substituted with phthaloyl.

6. A compound of claim 3, wherein

R₄ is a heterocyclic group substituted with substituent(s) selected from the group consisting of halogen, lower alkyl, oxo, acyl, amino and a group of the formula:

\[ \text{R}^5 \text{N}^{-} \text{R}^{10} \text{R} \]

in which R⁵ is hydrogen or lower alkyl, and R¹⁰ is acyl.

7. A process for preparing a compound of the formula:
wherein $R^1$ is halogen,

$R^2$ and $R^3$ are each hydrogen, lower alkyl, halo (lower) alkyl or acyl,

$R^4$ is aryl substituted with substituent(s) selected from the group consisting of halogen; halo (lower) alkyl; acyl; aryl substituted with halogen or cyano; ar (lower) alkyl substituted with hydroxy; lower alkoxy; nitro; amino; amino substituted with substituent(s) selected from the group consisting of lower alkyl, acyl, ar (lower) alkyl, heterocyclic (lower) alkyl, carboxy (lower) alkyl, lower alkyaminomethylene and N-methylpyrroldinylidene; a heterocyclic group and a heterocyclic group substituted with oxo; or a heterocyclic group constituted with a heterocyclic group substituted with oxo; or a heterocyclic group constituted with a heterocyclic group substituted with oxo; or a heterocyclic group constituted with a heterocyclic group substituted with oxo.

$Q$ is O or N-$R^{11}$,

in which $R^{11}$ is hydrogen or acyl, and

$A$ is lower alkylene,

or its salt, which comprises

a) halogenating a compound of the formula:
wherein $R^2$, $R^3$, $R^4$, $Q$ and $A$ are each as defined above,

or its salt to give a compound of the formula:

![Chemical Structure](attachment:image.png)
wherein $R^1$, $R^2$, $R^3$, $R^4$, $Q$ and $A$ are each as defined above,
or its salt, or

b) reacting a compound of the formula:

\[
\begin{align*}
\text{R}^3 & \quad \text{N} \\
\text{N} & \quad \text{R}^1 \\
\text{OH} & \quad \text{R}^2
\end{align*}
\]

wherein $R^1$, $R^2$, $R^3$ and $Q$ are each as defined above,
or its salt with a compound of the formula:

\[
X-A-R^4
\]

wherein $X$ is a leaving group, and
$R^4$ and $A$ are each as defined above,
or its salt to give a compound of the formula:

\[
\begin{align*}
\text{R}^3 & \quad \text{N} \\
\text{N} & \quad \text{R}^1 \\
\text{Q} & \quad \text{A} \\
\text{R}^4 & \quad \text{N}
\end{align*}
\]

wherein $R^1$, $R^2$, $R^3$, $R^4$, $Q$ and $A$ are each as defined above,
or its salt, or

c) acylating a compound of the formula:
wherein \( R^1 \) is hydrogen or lower alkyl,
\( R^6, R^7, R^8 \) and \( R^9 \) are each hydrogen or halogen, and
\( R^1, R^2, R^3, Q \) and \( A \) are each as defined above,
or its salt to give a compound of the formula:

wherein \( R^{10} \) is acyl, and
\( R^1, R^2, R^3, R^5, R^6, R^7, R^8, R^9, Q \) and \( A \) are each as defined above,
or its salt, or

d) acylating a compound of the formula:
wherein $R^{10}$ is acyl having amino, and

$R^1, R^2, R^3, R^5, R^6, R^7, R^8, R^9, Q$ and $A$ are each as defined above,
or its salt to give a compound of the formula:

wherein $R_B^{10}$ is acyl having acylamino, and

$R^1, R^2, R^3, R^5, R^6, R^7, R^8, R^9, Q$ and $A$ are each as defined above,
or its salt, or

e) alkylating a compound of the formula:
wherein $R_1, R_2, R_3, R_5, R_6, R_7, R_8, R_9, R_{10}, Q$ and $A$ are each as defined above, or its salt to give a compound of the formula:

wherein $R_{10}^1$ is acyl having lower alkylamino or acyl having ar(lower)alkylamino, and $R_1, R_2, R_3, R_5, R_6, R_7, R_8, R_9, Q$ and $A$ are each as defined above, or its salt, or

f) reacting a compound of the formula:
wherein (AA) is amino acid residue,
Y is NH or lower alkenylene,
Z is CH or N, and
R\(^1\), R\(^2\), R\(^3\), R\(^5\), R\(^6\), R\(^7\), R\(^8\), R\(^9\), Q and A are
each as defined above,
or its reactive derivative at the carboxy group or
a salt thereof with a compound of the formula:

\[
\begin{align*}
\text{HN} & \quad \text{R}^{12} \\
\text{R}^{13} & \quad \text{HN}
\end{align*}
\]

wherein R\(^{12}\) is hydrogen, lower alkyl, lower
alkoxy(lower)alkyl, lower
alkylamino(lower)alkyl, a
heterocyclic(lower)alkyl, a
heterocyclic group, protected or
unprotected hydroxy(lower)alkyl or
aryl optionally substituted with lower
alkylamino, and
R\(^{13}\) is hydrogen, lower alkyl, lower
alkoxy(lower)alkyl or protected or
unprotected hydroxy(lower)alkyl, or
R\(^{12}\) and R\(^{13}\)' are taken together with the
attached nitrogen atom to form a
heterocyclic group optionally having suitable substituent(s),
or its reactive derivative at the amino group or a salt thereof to give a compound of the formula:

\[
\begin{array}{c}
\text{R}^1 \quad \text{R}^2 \\
\text{R}^3 \\
\text{R}^4 \\
\text{R}^5 \\
\text{R}^6 \\
\text{R}^7 \\
\text{R}^8 \\
\text{R}^9 \\
\text{R}^{10} \\
\text{R}^{11} \\
\text{R}^{12} \\
\text{R}^{13}
\end{array}
\]

wherein \( R^1, R^2, R^3, R^4, R^5, R^6, R^7, R^8, R^9, R^{10}, R^{11}, A, \) 
\( (AA), Q, Y \) and \( Z \) are each as defined above, or its salt.

8. A pharmaceutical composition comprising a compound of claim 1, as an active ingredient, in association with a pharmaceutically acceptable, substantially nontoxic carrier or excipient.

9. A compound of claim 1 for use as a medicament.

10. A method for the prevention and/or the treatment of bradykinin or its analogues mediated diseases which comprises administering a compound of claim 1 to human being or animals.

11. Use of a compound of claim 1 for manufacture of a medicament for the prevention and/or the treatment of bradykinin or its analogues mediated diseases.
11. Compounds of claim 1, methods for their manufacture or pharmaceutical compositions or methods of treatment involving them, substantially as hereinbefore described with reference to the Examples.

12. A compound of claim 4, wherein \( R^4 \) is phenyl substituted with two halogens and a group of the formula:

\[
\begin{array}{c}
\text{N} \\
R^5 \\
R_c \end{array}
\]

in which

- \( R^5 \) is lower alkyl, and
- \( R_c^{10} \) is glycyl substituted with lower alkylcarbamoyl-ar (lower alkenoyl).

13. A compound of claim 11, which is 3-bromo-8-[2,6-dichloro-3-[N-[4-(dimethylcarbamoyl)cinnamoyl]glycyl]-N-methylamino]benzyl-2-methylimidazo[1,2-a]pyridine and its acid addition salt.


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ABSTRACT

A compound of the formula:

\[
\begin{array}{c}
\text{Q-A-R}^4 \\
N-N \\
\text{R}^2 \\
\text{R}^1 \\
\end{array}
\]

wherein \( R^1 \) is halogen,

- \( R^2 \) and \( R^3 \) are each hydrogen, lower alkyl,
- halo(lower)alkyl or acyl,

- \( R^4 \) is aryl having suitable substituent(s), or a heterocyclic group optionally having suitable substituent(s),
- \( Q \) is 0 or \( N-R^{11} \), in which \( R^{11} \) is hydrogen or acyl, and

- \( A \) is lower alkyene,

and pharmaceutically acceptable salts thereof, useful for the prevention and/or treatment of bradykinin or its analogues mediated diseases.