The present invention provides edible compositions comprising a compound of the present invention, food products comprising such edible compositions and methods of preparing such food products. The present invention also provides methods of reducing the amount of NaCl in a food product, methods of reducing the sodium intake in a diet, and methods of reducing bitter taste in a food product.
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Title: COMPOUNDS, COMPOSITIONS, AND METHODS FOR REDUCING OR ELIMINATING BITTER TASTE

Abstract: The present invention provides edible compositions comprising a compound of the present invention, food products comprising said edible compositions and methods of preparing such food products. The present invention also provides methods of reducing the amount of NaCl in a food product, methods of reducing the sodium intake in a diet, and methods of reducing bitter taste in a food product.
COMPOUNDS, COMPOSITIONS, AND METHODS
FOR REDUCING OR ELIMINATING BITTER TASTE

Field of the Invention

[0001] The present invention relates to flavor in edible compositions.

5 Background of the Invention

[0002] The sense of taste, e.g., in humans, can detect at least five traditional tastes: sweet, sour, salty, bitter, and umami (savory). Many nutritious substances including vegetables, foods, food ingredients and nutrients comprise bitter tastants and/or have a bitter taste. In addition, many pharmaceutical substances important to maintain or improve health comprise bitter tastants and/or have a bitter taste. While certain food products and consumer products have desirable bitter tastes, including coffee, beer and dark chocolate, in many contexts, consumers dislike such bitter tastes.

For example, many consumers dislike the perception of certain bitter tastants and/or bitter taste and will avoid food or pharmaceutical products with an undesirable bitter tastant or bitter taste in favor of food or pharmaceutical products that have reduced levels of undesirable bitter tastants or that have reduced or that completely lack bitter taste. This aversion to products containing undesirable bitter tastants and/or having undesirable bitter taste may be caused by perception of bitter tastants and/or bitter taste mediated by activation of bitter receptors present in the oral cavity and/or in the gastrointestinal tract. In many cases, consumer dislike of bitter tastants and/or bitter taste prevents or hampers improvement of the nutritive quality and safety of foods as desired levels of nutrients or preservatives comprising bitter tastants and/or having bitter taste cannot be used. Also, dislike of or aversion to the bitter tastants or bitter taste of some pharmaceutical agents negatively impacts compliance with prescribed regimens for their use.

[0003] For instance, several additives, preservatives, emulsifiers and foodstuffs used in the production of food products comprise bitter tastants and/or have a bitter taste. While these additives, preservatives, emulsifiers and foodstuffs may affect the taste of a food product, they may also be important for improving the shelf life, nutritive quality, or texture of the food product. For
example, the increasing trend of hypertension and cardiovascular disease has been attributed, in part, to the high sodium intake of the Western diet. Accordingly, substitution of sodium chloride with another salty tasting compound is desirable. The most common sodium chloride substitute is potassium chloride, which, to a portion of the population, is perceived as possessing a bitter taste in addition to its salty taste. The bitter taste of potassium chloride limits the extent to which it may be used to replace sodium chloride in foods without causing undesired bitter taste for the portion of the population sensitive to it.

[0004] Another common food additive, sodium lactate, has a broad antimicrobial action, is effective at inhibiting spoilage, and growth of pathogenic bacteria, and is commonly used in food products (e.g., meat and poultry products) to extend shelf life and increase food safety. Due to its sodium content, however, sodium lactate, can be undesirable as a preservative. Potassium lactate, which has similar antimicrobial properties, has been used in lieu of sodium lactate. However, potassium lactate is also associated with a bitter taste which limits the extent to which it may be used to replace sodium lactate in foods without causing undesired bitter taste.

[0005] In addition, the increasing incidence of obesity and diabetes has been attributed, in part, to the high sugar intake of many diets. Accordingly, substitution of sugar with another sweet tasting compound is desirable. Artificial and natural sugar substitutes that may be used to reduce sugar in foods are often associated with bitter taste which again limit the extent to which these may be used to replace sugar in foods without causing adverse bitter taste. For example, a common sugar substitute is Acesulfame K, which also has a bitter taste in addition to its sweet taste.

[0006] Without being limited by theory, bitter, sweet, and umami tastants and compounds typically elicit a taste response via G-protein coupled receptors, while salty and sour tastants and compounds are typically hypothesized to elicit a taste response via ion channels. Bitter taste receptors belong to the T2R (also referred to as TAS2R) family of G-protein coupled receptors that induce intracellular calcium concentration changes in response to a bitter tastant. T2R receptors act via gustducin, a taste-specific G-protein. There are at least twenty-five different members of the T2R family, suggesting that the perception of bitter taste is complex, involving several different tastant-receptor interactions. Compounds capable of modulating the activation and/or signaling of bitter taste receptors in the oral cavity and/or the gastrointestinal tract could be effective to allow desired usage levels of bitter tastants or bitter tasting substances in food and pharmaceutical products without resulting in consumer dislike of such products due to perception of the increased levels of bitter tastants or bitter tastes. In some instances, blockers or modulators of bitter taste receptors and bitter taste may reduce the perception of bitter tastants and/or bitter taste via the bitter taste receptors and/or taste transduction signaling machinery present in the oral cavity and/or the gastrointestinal tract.

[0007] Traditionally in food preparation and pharmaceuticals, bitter taste was masked using sweeteners and other tastants, including salt. In some cases, however, this is undesirable or insufficient because it can alter, mask, or interfere with other tastes/flavors impressions (e.g., non bitter tastes or desired bitter tastes) in the food product. Additionally, this approach has rarely been
able to completely mask the bitter taste present in such food products or pharmaceuticals. For that reason, compounds which reduce bitter taste instead of, or in addition to, masking agents are preferred.

[0008] It is, therefore, desirable to provide compounds that may be added to food products, consumer products and pharmaceuticals comprising bitter tastants or having a bitter taste to eliminate, modulate or reduce the perception of the bitter tastants or bitter taste, or to reduce the corresponding activation of the bitter receptors in the oral cavity and/or the gastrointestinal tract. Similarly, it is desirable to provide food products, consumer products, and pharmaceutical compositions comprising such compounds. It is also desirable to decrease the sodium intake of a subject using such compounds to eliminate, modulate or reduce the perception of bitter taste associated with salt substitutes. It is further desirable to decrease the sugar intake of a subject using such compounds to eliminate, modulate or reduce the perception of bitter taste associated with sugar substitutes.

Summary of the Invention

[0009] The present invention provides compounds that modulate bitter taste, edible compositions comprising such compounds, and methods of preparing such edible compositions. The present invention also provides methods of reducing the amount of sodium or sugar in an edible composition and methods of reducing bitter taste of a food product. The present invention further provides a method of reducing, modulating or eliminating the bitter taste of a food, consumer or pharmaceutical product in a subject. The present invention also provides a method of modulating, particularly reducing the activation of a bitter taste receptor.

Edible compositions

[0010] One aspect of the present invention provides edible compositions for reducing bitter taste of a bitter tastant. In some embodiments, the edible composition comprises a diphenyl-containing compound. In some embodiments, the diphenyl-containing compound is a compound having a molecular weight less than about 1000, 500, or 300 daltons. In certain embodiments, the diphenyl-containing compound is a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb') or Formula (IIIb") or Compounds 1-22 or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof.

[0011] In some embodiments, the edible composition comprises a pyrazole-containing compound. In some embodiments, the pyrazole-containing compound is a compound having a molecular weight less than about 1000, 500, or 300 daltons. In certain embodiments, the pyrazole-containing compound is a compound of Formula (IV), Formula (Va), Formula (Vb), Formula (VIa), Formula (VIb), Formula (VIIa) or Formula (VIIb) or Compounds 23-36 or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof.
[0012] In some embodiments, the edible composition comprises a hydroquinoline compound. In some embodiments, the hydroquinoline compound is a compound having a molecular weight less than about 1000, 500, or 300 daltons. In certain embodiments, the hydroquinoline compound is a compound of Formula (VIII), Formula (IX), or Formula (X) or Compounds 37-43 or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof.

[0013] In some embodiments, the edible composition comprises a quinoline compound. In some embodiments, the quinoline compound is a compound having a molecular weight less than about 1000, 500, or 300 daltons. In certain embodiments, the quinoline compound is a compound of Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIa), or Formula (XIIib) or Compounds 44-48 or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof.

[0014] In some embodiments, the edible composition comprises a N-phenylalkylamide compound. In some embodiments, the N-phenylalkylamide compound is a compound having a molecular weight less than about 1000, 500, or 300 daltons. In certain embodiments, the N-phenylalkylamide compound is a compound of Formula (XIV), Formula (XVa), Formula (XVb), or Formula (XVc) or Compounds 49-58 or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof.

[0015] In some embodiments, the edible composition comprises (a) a compound of the invention; and (b) a bitter tastant. In some embodiments, the compound of the invention is a compound having a molecular weight less than about 1000, 500, or 300 daltons. In certain embodiments, the compound of the invention is a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIb), Formula (IIIb), Formula (IIIb), Formula (IIIb"), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIa), Formula (XIIib), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof.

[0016] In another embodiment, the edible composition comprises (a) any one of Compounds 1-58, or combinations thereof; and (b) a bitter tastant.

[0017] According to the invention, the bitter tastant can be inherent in, e.g., a food product (such as coffee or chocolate) or can be a component of an edible composition (such as a bitter tasting preservative). In some embodiments, the bitter tastant present in the edible composition is a bitter tasting salt. In some embodiments, the bitter tastant present in the edible composition is a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the bitter tastant is a potassium salt. In some embodiments, the bitter tastant present in the edible compositions is KCl. In other embodiments, the bitter tastant present in the edible composition is potassium lactate.

[0018] In some embodiments, the edible composition further comprises a sodium salt. In some embodiments, the edible composition further comprises NaCl. In other embodiments, the edible composition further comprises sodium lactate. In some embodiments, the edible composition further comprises sugar.
[0019] In another aspect of the invention, the edible composition is a food product comprising at least one compound of the invention. In certain embodiments, the compound of the invention is a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Via), Formula (Vib), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof. In another embodiments, the pharmaceutical composition comprises a bitter tasting pharmaceutically active ingredient and any one of Compounds 1-58, or combinations thereof.

[0020] In another aspect of the present invention, the edible composition is a pharmaceutical composition comprising a bitter tasting pharmaceutically active ingredient and a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Via), Formula (Vib), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof. In another embodiments, the pharmaceutical composition comprises a bitter tasting pharmaceutically active ingredient and any one of Compounds 1-58, or combinations thereof.

[0021] In yet other embodiments, the edible composition is a pharmaceutical composition comprising a pharmaceutically active ingredient, a bitter tastant, and a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Via), Formula (Vib), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof. In yet other embodiments, the pharmaceutical composition comprises a pharmaceutically active ingredient, a bitter tastant, and any one of Compounds 1-58, as described herein, or combinations thereof.

[0022] In another aspect of the present invention, the edible composition is a consumer product comprising a bitter tastant and a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Via), Formula (Vib), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof. In another embodiments, the consumer product comprises a bitter tasting ingredient and any one of Compounds 1-58, or combinations thereof.

[0023] Yet another embodiment of the present invention provides a consumer product for reducing bitter taste of a bitter tastant, wherein said consumer product comprises a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Via), Formula (Vib), Formula (VIIa),
Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa),
Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XV$a$),
Formula (XV$b$) or Formula (XV$c$), as described herein, or combinations thereof. In yet other
embodiments, the consumer product for reducing bitter taste of a bitter tastant comprises any one of
Compounds 1-58, as described herein, or combinations thereof.

[0024] In a further aspect, the present invention provides a method of preparing an edible
composition comprising:

(a) providing a comestibly acceptable carrier; and
(b) adding to the comestibly acceptable carrier of (a) a compound of Formula (I),

Formula (I$a$), Formula (I$b$), Formula (III$a$), Formula (III$b$), Formula (III$b''$), Formula (IV),
Formula (Va), Formula (V$b$), Formula (Vla), Formula (V$b$), Formula (VII$a$), Formula (VII$b$),
Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XII$a$), Formula (XIIb),
Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XV$b$) or
Formula (XV$c$), as described herein, or combinations thereof.

[0025] In another embodiment, the method of preparing an edible composition comprises:

(a) providing a comestibly acceptable carrier; and
(b) adding to the comestibly acceptable carrier of (a) any one of Compounds
1-58, or combinations thereof.

[0026] In some embodiments, the edible composition is a food product, a consumer product or a
pharmaceutical composition. In some embodiments, the comestibly acceptable carrier is a
foodstuff, a food product, or a pharmaceutically acceptable carrier.

[0027] In some embodiments, the comestibly acceptable carrier in (a) is inherently bitter. In such
embodiments, the comestibly acceptable carrier may inherently contain a bitter tastant (i.e., the
comestibly acceptable carrier is bitter without addition of a bitter tastant). In some embodiments,
the inherent bitter tastant is a bitter tasting salt. In some embodiments, the inherently bitter
cometstibly acceptable carrier comprises a potassium salt, a magnesium salt, or a calcium salt. In
some embodiments, the inherently bitter comestibly acceptable carrier comprises a potassium salt,
such as KCl.

[0028] In other embodiments, the method of preparing an edible composition further comprises:
(c) adding a bitter tastant. In some embodiments, the bitter tastant used in the methods of preparing
an edible composition is a bitter tasting salt. In some embodiments, the bitter tastant used in the
methods of preparing an edible composition is a potassium salt, a magnesium salt, or a calcium salt.
In some embodiments, the bitter tastant used in the methods of preparing an edible composition is a
potassium salt. In some embodiments, the bitter tastant used in the methods of preparing an edible
composition is KCl. In other embodiments, the bitter tastant used in the methods of preparing an edible
composition is potassium lactate.

[0029] In some embodiments, the edible composition further comprises a sodium salt. In some
embodiments, the edible composition further comprises NaCl. In some embodiments, the edible
composition further comprises sodium lactate. In some embodiments, the edible composition further comprises sugar.

[0030] The present invention also provides a method of reducing the amount of sodium in an edible composition. In some embodiments, such methods comprise:

(a) replacing an amount of one or more sodium salts used to prepare an edible composition with an amount of one or more potassium salts; and

(b) incorporating into the edible composition an effective amount of a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (VIa),

[0031] In another embodiment, the method of reducing the amount of sodium in an edible composition comprises:

(a) replacing an amount of one or more sodium salts used to prepare an edible composition with an amount of one or more potassium salts; and

(b) incorporating into the edible composition an effective amount of any one of Compounds 1-58, or combinations thereof.

[0032] In some embodiments, the edible composition is a food product, a consumer product or a pharmaceutical composition.

[0033] In some embodiments of the present invention, the method of reducing the amount of sodium in an edible composition, comprises adding an amount of the compound in (b) sufficient to permit replacement of up to 25% of the sodium present in an edible composition with potassium. In other embodiments, the amount of the compound added in (b) is sufficient to permit replacement of up to 50% of the sodium present in an edible composition with potassium. In yet other embodiments, the amount of the compound added in (b) is sufficient to permit replacement of up to 75% of the sodium present in an edible composition with potassium. In other embodiments, the amount of the compound added in (b) is sufficient to permit replacement of up to 100% of the sodium present in an edible composition with potassium. In some embodiments, the edible composition maintains a salty flavor.

[0034] The present invention also provides a method of reducing the amount of NaCl in an edible composition. In some embodiments, such methods comprise:

(a) replacing an amount of NaCl used to prepare an edible composition with an amount of KCl; and

(b) incorporating into the edible composition an effective amount of a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (VIa),

Formula (VIb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX),
Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa),
Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as
described herein, or combinations thereof.

In another embodiment, the method of reducing the amount of NaCl in an edible
composition comprises:

(a) replacing an amount of NaCl used to prepare an edible composition with an
amount of KCl; and

(b) incorporating into the edible composition an effective amount of any one of
Compounds 1-58, or combinations thereof.

In some embodiments, the edible composition is a food product, a consumer product or a
pharmaceutical composition.

In some embodiments of the present invention, the method of reducing the amount of
sodium in an edible composition, comprises adding an amount of the compound in (b) sufficient to
permit replacement of up to 25% of the NaCl present in an edible composition with KCl. In other
embodiments, the amount of the compound added in (b) is sufficient to permit replacement of up to
50% of the NaCl present in an edible composition with KCl. In yet other embodiments, the amount of
the compound added in (b) is sufficient to permit replacement of up to 75% of the NaCl present
in an edible composition with KCl. In other embodiments, the amount of the compound added in
(b) is sufficient to permit replacement of up to 100% of the NaCl present in an edible composition
with KCl. In some embodiments, the edible composition maintains a salty flavor.

In another embodiment, the present invention provides a method of reducing the amount of
sodium lactate in an edible composition comprises:

(a) replacing an amount of sodium lactate used to prepare an edible composition
with an amount of potassium lactate; and

(b) incorporating into the edible composition an effective amount of a compound
according to Formula (I), Formula (Iia), Formula (Iib), Formula (IIIb), Formula (IIIb'),
Formula (IIb'), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla),
Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX),
Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa),
Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as
described herein, or combinations thereof.

In another embodiment, the invention provides a method of reducing the amount of
sodium lactate in an edible composition comprising:

(a) replacing an amount of sodium lactate used to prepare an edible composition
with an amount of potassium lactate; and

(b) incorporating into the edible composition an effective amount of any one of
Compounds 1-58, or combinations thereof.

In some embodiments, the edible composition is a food product, a consumer product or a
pharmaceutical composition.
In some embodiments of the present invention, the method of reducing the amount of sodium lactate in an edible composition, comprises adding an amount of the compound in (b) sufficient to permit replacement of up to 25% of the sodium lactate present in an edible composition with potassium lactate. In other embodiments, the amount of the compound added in (b) is sufficient to permit replacement of up to 50% of the sodium lactate present in an edible composition with potassium lactate. In yet other embodiments, the amount of the compound added in (b) is sufficient to permit replacement of up to 75% of the sodium lactate present in an edible composition with potassium lactate. In other embodiments, the amount of the compound added in (b) is sufficient to permit replacement of up to 100% of the sodium lactate present in an edible composition with potassium lactate. In some embodiments, the edible composition has the same shelf life as an edible composition comprising sodium lactate.

In another embodiment, the invention provides a method of reducing the amount of sugar in an edible composition comprising:

(a) replacing an amount of sugar used to prepare an edible composition with an amount of Acesulfame K; and

(b) incorporating into the edible composition an effective amount of a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof.

In another embodiment, the invention provides a method of reducing the amount of sugar in an edible composition comprising:

(a) replacing an amount of sugar used to prepare an edible composition with an amount of Acesulfame K; and

(b) incorporating into the edible composition an effective amount of any one of Compounds 1-58, or combinations thereof.

In some embodiments, the edible composition is a food product, a consumer product or a pharmaceutical composition.

In some embodiments of the present invention, the method of reducing the amount of sugar in an edible composition, comprises adding an amount of the compound in (b) sufficient to permit replacement of up to 25% of the sugar present in an edible composition with Acesulfame K. In other embodiments, the amount of the compound added in (b) is sufficient to permit replacement of up to 50% of the sugar present in an edible composition with Acesulfame K. In yet other embodiments, the amount of the compound added in (b) is sufficient to permit replacement of up to 75% of the sugar present in an edible composition with Acesulfame K. In other embodiments, the amount of the compound added in (b) is sufficient to permit replacement of up to 100% of the sugar
present in an edible composition with Acesulfame K. In some embodiments, the edible composition maintains a sweet flavor.

[0046] The present invention also provides a method of reducing the sodium intake of a subject. Such method comprises:

(a) replacing an amount of NaCl used to prepare an edible composition with an amount of KCl; and

(b) incorporating into the edible composition an effective amount of a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IV), Formula (V), Formula (VI), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof, thereby reducing the sodium intake of the subject.

[0047] In another embodiment, the method of reducing the sodium intake of a subject comprises:

(a) replacing an amount of NaCl used to prepare an edible composition with an amount of KCl; and

(b) incorporating into the edible composition an effective amount of any one of Compounds 1-58, or combinations thereof, thereby reducing the sodium intake of the subject.

[0048] In another embodiment, the method of reducing the sodium intake of a subject comprises:

(a) replacing an amount of sodium lactate used to prepare an edible composition with an amount of potassium lactate; and

(b) incorporating into the edible composition an effective amount of a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IV), Formula (V), Formula (VI), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof, thereby reducing the sodium intake of the subject.

[0049] In another embodiment, the method of reducing the sodium intake of a subject comprises:

(a) replacing an amount of sodium lactate used to prepare an edible composition with an amount of potassium lactate; and

(b) incorporating into the edible composition an effective amount of any one of Compounds 1-58, or combinations thereof, thereby reducing the sodium intake of the subject.

[0050] In some embodiments, the edible composition is a food product, a consumer product or a pharmaceutical composition.

[0051] In some embodiments of the present invention, the methods of reducing the sodium intake of a subject further comprise (c) identifying a subject in need thereof. In some embodiments, the methods of reducing the sodium intake of a subject comprise adding an amount of the compound in (b) sufficient to reduce sodium intake by up to 25% using potassium replacement. In other
embodiments, the amount of compound added in (b) is sufficient to reduce sodium intake by up to 50% using potassium replacement. In yet other embodiments, the amount of compound added in (b) is sufficient to reduce sodium intake by up to 75% using potassium replacement. In other embodiments, the amount of compound added in (b) is sufficient to reduce sodium intake by up to 100% using potassium replacement.

[0052] The present invention also provides a method of reducing sugar intake of a subject comprising:

(a) replacing an amount of sugar used to prepare an edible composition with an amount of Acesulfame K; and

(b) incorporating into the edible composition an effective amount of a compound according to Formula (1), Formula (Ia), Formula (Ib), Formula (IIIb), Formula (IIIb'), Formula (IIIb*), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof, thereby reducing the sugar intake of the subject.

[0053] In another embodiment, the method of reducing the sugar intake of a subject comprises:

(a) replacing an amount of sugar used to prepare an edible composition with an amount of Acesulfame K; and

(b) incorporating into the edible composition an effective amount of any one of Compounds 1-58, or combinations thereof, thereby reducing the sugar intake in the diet or meal of the subject.

[0054] In some embodiments, the edible composition is a food product, a consumer product or a pharmaceutical composition.

[0055] In some embodiments of the present invention, the methods of reducing the sugar intake of a subject further comprises (c) identifying a subject in need thereof. In some embodiments, the methods of reducing the sugar intake of a subject comprise adding an amount of the compound in (b) sufficient to reduce sugar intake by up to 25% using Acesulfame K replacement. In other embodiments, the amount of compound added in (b) is sufficient to reduce sugar intake by up to 50% using Acesulfame K replacement. In yet other embodiments, the amount of compound added in (b) is sufficient to reduce sugar intake by up to 75% using Acesulfame K replacement. In other embodiments, the amount of compound added in (b) is sufficient to reduce sugar intake by up to 100% using Acesulfame K replacement.

[0056] The present invention also provides a method of reducing the bitter taste attributed to a bitter tastant in an edible composition comprising adding an effective amount of a compound according to Formula (1), Formula (Ia), Formula (Ib), Formula (IIIb), Formula (IIIb'), Formula (IIIb*), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV),
Formula (XV\(a\)), Formula (XV\(b\)) or Formula (XV\(c\)), as described herein, or combinations thereof, to the edible composition such that any bitter taste induced by the bitter tastant is reduced. In other embodiments, the compound added to the edible composition is any one of Compounds 1-58, or combinations thereof.

[0057] The present invention further provides a method of reducing the bitter taste attributed to a bitter tastant in an edible composition comprising ingesting an effective amount of a compound according to Formula (I), Formula (Ia), Formula (IIb), Formula (IIIb), Formula (IIIb\(a\)), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIIb), Formula (XIIIa), Formula (XIV), Formula (XV), Formula (XVIIb) or Formula (XVc), as described herein, or combinations thereof, before, along with, or after the edible composition such that any bitter taste induced by the bitter tastant is reduced. In other embodiments, the compound ingested with the edible composition is any one of Compounds 1-58, or combinations thereof.

[0058] In some embodiments, the method reduces the bitter taste induced by the bitter tastant by up to 25%. In some embodiments, the method reduces the bitter taste induced by the bitter tastant by up to 50%. In other embodiments, the bitter taste induced by the bitter tastant is reduced by up to 75%. In yet other embodiments, the bitter taste induced by the bitter tastant is reduced by up to 100%. In some embodiments, the bitter tastant present in the edible composition is a bitter tasting salt. In some embodiments, the bitter tastant present in the edible composition is a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the bitter tastant present in the edible compositions is K\(\text{Cl}\). In other embodiments, the bitter tastant present in the edible composition is potassium lactate.

[0059] In further aspect, the present invention provides a method of preserving an edible composition comprising:

(a) providing an edible composition; and

(b) adding to the edible composition of (a) potassium lactate and an effective amount of a compound of Formula (I), Formula (Ia), Formula (IIb), Formula (IIIb), Formula (IIIb\(a\)), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (VIIb), Formula (VIIa), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIIb), Formula (XIIIa), Formula (XIV), Formula (XV), Formula (XVIIb) or Formula (XVc), as described herein, or combinations thereof.

[0060] In another embodiment, the method of preserving or extending the shelf life of an edible composition comprises:

(a) providing an edible composition; and

(b) adding to the edible composition of (a) potassium lactate and an effective amount of any one of Compounds 1-58, or combinations thereof.
[0061] The present invention also provides a method of reducing the amount of sodium in an edible composition while preserving the edible composition. In some embodiments, such method comprises:

(a) replacing an amount of sodium lactate used to prepare an edible composition with an amount of potassium lactate; and

(b) incorporating into the edible composition an effective amount of a compound according to Formula (I), Formula (Ia), Formula (Ib), Formula (Iib), Formula (Iib‘), Formula (Iib”), Formula (IV), Formula (V), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIia), Formula (XIIib), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof.

[0062] The present invention also provides a method of reducing the amount of sodium in an edible composition while preserving the edible composition. In some embodiments, such method comprises:

(a) replacing an amount of sodium lactate used to prepare an edible composition with an amount of potassium lactate; and

(b) incorporating into the edible composition an effective amount of any one of Compounds 1-58, or combinations thereof.

[0063] In some embodiments, the edible composition is a food product. In some embodiments, the edible composition is a consumer product. In some embodiments, the edible composition is a pharmaceutical composition.

[0064] The present invention also provides a method of reducing or eliminating bitter taste in a subject utilizing an edible composition comprising a compound of Formula (I), Formula (Ia), Formula (Ib), Formula (Iib), Formula (Iib‘), Formula (Iib”), Formula (IV), Formula (V), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIia), Formula (XIIib), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof. In other embodiments, the composition that reduces or eliminates a bitter taste in a subject comprises any one of Compounds 1-58, or combinations thereof.

[0065] In some embodiments, the bitter taste is inherent. In some embodiments, the bitter taste is due to a bitter tasting salt. In some embodiments, the bitter taste is due to a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the bitter taste is due to KCl. In other embodiments, the bitter taste is due to potassium lactate.

[0066] The present invention also provides a method of inhibiting or reducing the activation and/or signaling of a bitter taste receptor, wherein the method comprises contacting a bitter taste receptor with a compound of Formula (I), Formula (Ia), Formula (Ib), Formula (Iib), Formula (Iib‘), Formula (Iib”), Formula (IV), Formula (V), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIia), Formula (XIIib), Formula (XIV),
Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof. In other embodiments, the method comprises contacting a bitter taste receptor with any one of Compounds 1-58, or combinations thereof. In some embodiments, the bitter taste receptor is in the mouth. In other embodiments, the bitter taste receptor is in the gastrointestinal tract, for example, in the stomach. In other embodiments, the bitter taste receptor is in an in vitro assay.

[0067] Particular embodiments of the invention are set forth in the following numbered paragraphs:

1. A composition comprising a compound according to Formula (I):

\[
(R^1)_n \quad (R^2)_m
\]

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

- \( R^1 \), independently for each occurrence, is selected from the group consisting of
  - \( C_{1,10} \) alkyl, \( C_{1,10} \) haloalkyl, \( C_{2,10} \) alkynyl, halo, hydroxy, carboxyl,
  - \( C_{1,10} \) alkoxy carbonyl, \( C_{2,10} \) alkenyloxy carbonyl, \( C_{2,10} \) alkynyl oxycarbonyl, \( C_{1,10} \) acyl,
  - \( C_{1,10} \) acyl-amino, \( C_{1,10} \) acyloxy, \( C_{1,10} \) carbonate, \( C_{1,10} \) alkoxy, \( C_{6,10} \) aryloxy,
  - \( C_{6,10} \) aryl-\( C_{1,10} \) alkoxy, \( C_{1,3} \) heteroaryloxy, \( C_{1,3} \) heteroaryl-\( C_{1,6} \) alkoxy, \( C_{3,10} \) alkenyloxy,
  - \( C_{3,10} \) alkenyloxy, \( C_{1,3} \) heteroaryloxy, \( C_{1,3} \) heteroaryl-\( C_{1,6} \) alkoxy, \( C_{3,10} \) alkenyloxy,
  - \( C_{1,3} \) heteroaryloxy, \( C_{1,3} \) heteroaryl-\( C_{1,6} \) alkoxy, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

- \( R^2 \), independently for each occurrence, is selected from the group consisting of
  - \( C_{1,10} \) alkyl, \( C_{1,10} \) haloalkyl, \( C_{2,10} \) alkynyl, halo, hydroxy, carboxyl,
  - \( C_{1,10} \) alkoxy carbonyl, \( C_{2,10} \) alkenyloxy carbonyl, \( C_{2,10} \) alkynyl oxycarbonyl, \( C_{1,10} \) acyl,
  - \( C_{1,10} \) acyl-amino, \( C_{1,10} \) acyloxy, \( C_{1,10} \) carbonate, \( C_{1,10} \) alkoxy, \( C_{6,10} \) aryloxy,
  - \( C_{6,10} \) aryl-\( C_{1,10} \) alkoxy, \( C_{1,3} \) heteroaryloxy, \( C_{1,3} \) heteroaryl-\( C_{1,6} \) alkoxy, \( C_{3,10} \) alkenyloxy,
  - \( C_{3,10} \) alkenyloxy, \( C_{1,3} \) heteroaryloxy, \( C_{1,3} \) heteroaryl-\( C_{1,6} \) alkoxy, \( C_{3,10} \) alkenyloxy,
  - \( C_{1,3} \) heteroaryloxy, \( C_{1,3} \) heteroaryl-\( C_{1,6} \) alkoxy, \( C_{3,10} \) alkenyloxy,
  - \( C_{1,3} \) heteroaryloxy, \( C_{1,3} \) heteroaryl-\( C_{1,6} \) alkoxy, \( C_{3,10} \) alkenyloxy,
  - \( C_{1,3} \) heteroaryloxy, \( C_{1,3} \) heteroaryl-\( C_{1,6} \) alkoxy, \( C_{3,10} \) alkenyloxy,
  - \( C_{1,3} \) heteroaryloxy, \( C_{1,3} \) heteroaryl-\( C_{1,6} \) alkoxy, \( C_{3,10} \) alkenyloxy,
  - \( C_{1,3} \) heteroaryloxy, \( C_{1,3} \) heteroaryl-\( C_{1,6} \) alkoxy, \( C_{3,10} \) alkenyloxy,
  - \( C_{1,3} \) heteroaryloxy, \( C_{1,3} \) heteroaryl-\( C_{1,6} \) alkoxy, \( C_{3,10} \) alkenyloxy,
  - \( C_{1,3} \) heteroaryloxy, \( C_{1,3} \) heteroaryl-\( C_{1,6} \) alkoxy, \( C_{3,10} \) alkenyloxy,
C_1-10_\text{arrea, cyano, nitro, azido, sulphydryl, C}_{1,10}\text{alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C}_{3,5}\text{-carboxycarbonyl, C}_{3,5}\text{-carboxycarbonyl-C}_{1,8}\text{alkyl, C}_{1,8}\text{heterocyclic, C}_{1,8}\text{heterocyclic-C}_{1,8}\text{alkyl, phenyl, phenyl-C}_{1,8}\text{alkyl, C}_{1,8}\text{heteroaryl, and C}_{1,8}\text{heteroaryl-C}_{1,8}\text{alkyl}, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;}

X is O or NR^2, wherein R^2 is absent or is selected from the group consisting of hydrogen, C_1-10_\text{alkyl, C}_{1-10}\text{haloalkyl, C}_{2-10}\text{alkenyl, C}_{2-10}\text{alkynyl, carboxyl, C}_{1-10}\text{alkoxy-carbonyl, C}_{2,3}\text{hydroxyalkoxy-carbonyl, C}_{2,3}\text{hydroxyalkoxy-carbonyl-1,8-acyl, phosphoryl, phosphonate, phosphinate, cyano, sulfonate, sulfamoyl, sulfonyl, C}_{3,5}\text{carboxycarbonyl, C}_{3,5}\text{carboxycarbonyl-C}_{1,8}\text{alkyl, C}_{1,8}\text{heterocyclic, C}_{1,8}\text{heterocyclic-C}_{1,8}\text{alkyl, phenyl, phenyl-C}_{1,8}\text{alkyl, C}_{1,8}\text{heteroaryl, and C}_{1,8}\text{heteroaryl-C}_{1,8}\text{alkyl}, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;}

wherein any of R^1, R^2, and R^3, independently and independently for each occurrence, is optionally substituted with 1-3 substituents selected from the group consisting of C_1-10_\text{alkyl, C}_{1-10}\text{haloalkyl, halo, hydroxyl, carboxyl, C}_{1-10}\text{alkoxy-carbonyl, C}_{2,3}\text{hydroxyalkoxy-carbonyl, C}_{2,3}\text{hydroxyalkoxy-carbonyl-1,8-acyl, C}_{1-10}\text{aminocarbonyl, C}_{1-10}\text{amino, C}_{1-10}\text{hydroxy-carbonyl, C}_{1-10}\text{amino-carbonyl, C}_{1-10}\text{alkoxy, phenol, phenol, phosphite, phosphonate, phosphinate, amino, diC}_{1-10}\text{alkylamin, monoC}_{1-10}\text{alkylamin, C}_{1-10}\text{amido, C}_{1-10}\text{amido, C}_{1-10}\text{carbamate, C}_{1-10}_\text{arrea, cyano, nitro, azido, sulphydryl, C}_{1-10}_\text{alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C}_{3,5}_\text{carboxycarbonyl, C}_{3,5}_\text{carboxycarbonyl-C}_{1,8}_\text{alkyl, C}_{1,8}_\text{heterocyclic, C}_{1,8}_\text{heterocyclic-C}_{1,8}_\text{alkyl, phenyl, phenyl-C}_{1,8}_\text{alkyl, C}_{1,8}_\text{heteroaryl, and C}_{1,8}_\text{heteroaryl-C}_{1,8}_\text{alkyl}; and wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;}

m is 1-3; and

n is 0-3;

wherein the composition is edible and capable of reducing bitter taste of a bitter tastant.

2. The composition according to paragraph 1, wherein as valence and stability permit:

R^1, independently for each occurrence, is selected from the group consisting of halo; hydroxyl; C_1-6_\text{alkyl; C}_{1-6}\text{haloalkyl, C}_{1-6}\text{hydroxyalkyl, or C}_{1-6}\text{acyxy-C}_{1-6}\text{alkyl; C}_{2,3}\text{alkenyl; C}_{2,3}\text{alkynyl, C}_{1-6}\text{alkoxy, C}_{1-6}\text{alkylthio, and C}_{6,10}\text{aryl-C}_{1,8}\text{alkyloxy optionally substituted by halo, hydroxyl, C}_{1-6}_\text{alkyl, C}_{1-6}_\text{alkoxy, or C}_{1-6}_\text{acyloxy;}

R^2, independently for each occurrence, is selected from the group consisting of halo; hydroxyl; C_1-6_\text{alkyl; C}_{1-6}\text{haloalkyl, C}_{1-6}\text{hydroxyalkyl, or C}_{1-6}\text{acyxy-C}_{1-6}\text{alkyl; C}_{2,3}_\text{alkenyl; C}_{2,3}\text{alkynyl, C}_{1-6}_\text{alkoxy, C}_{1-6}_\text{alkylthio, and C}_{6,10}\text{aryl-C}_{1,8}\text{alkyloxy optionally substituted by halo, hydroxyl, C}_{1-6}_\text{alkyl, C}_{1-6}_\text{alkoxy, or C}_{1-6}_\text{acyloxy;
X is O or NR², wherein R² is absent or is selected from the group consisting of hydrogen and C₁₆₋₅₃-alkyl;

wherein any of R¹, R², and R³, independently and independently for each occurrence, is optionally further substituted as in paragraph 1;

m is 1.3; and

n is 0.3.

3. The composition according to paragraph 1, wherein said compound according to Formula (I) is a compound according to Formula (IIa):

\[
\begin{align*}
\text{(R}_1^n\text{)} & \quad \text{N} \quad \text{\text{\Large \text{\&}}} \quad \text{(R}_2^m\text{)} \\
\end{align*}
\]

Formula (IIa);

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, wherein, as valence and stability permit, R¹, R², m, and n are as defined in paragraph 1.

4. The composition according to paragraph 1, wherein said compound according to Formula (I) is a compound according to Formula (IIb):

\[
\begin{align*}
\text{(R}_1^n\text{)} & \quad \text{N} \quad \text{\text{\Large \text{\&}}} \quad \text{(R}_2^m\text{)} \\
\end{align*}
\]

Formula (IIb);

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, wherein, as valence and stability permit, R¹, R², m, and n are as defined in paragraph 1.

5. The composition according to paragraph 4, wherein said compound according to Formula (IIb) is a compound according to Formula (IIIb):
Formula (IIIb);

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

R¹ and n are as defined in paragraph 1; and

R³ is selected from the group consisting of methyl and ethyl.

6. The composition according to paragraph 4, wherein said compound according to Formula (IIIb) is a compound according to Formula (IIIb°):

Formula (IIIb°);

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

R¹, R², and n are as defined in paragraph 1; and

Ar is C₆₋₁₅aryl optionally substituted by halo, hydroxyl, C₁₋₆alkyl, C₁₋₆alkoxy, or C₁₋₆acyloxy.

7. The composition according to paragraph 4, wherein said compound according to Formula (IIIb) is a compound according to Formula (IIIb°):
or a pharmaceutically or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

\( R^1, R^2, \) and \( m \) are as defined in paragraph 1; and

\( R^3 \) is C1-alkyl, such as methyl.

The composition according to paragraph 1, wherein said compound according to Formula (I) is selected from the group consisting of:

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<tr>
<th>Compound 1</th>
<th>( \text{HO} \text{-} \text{O} \text{-} \text{O} \text{-} \text{CH}_3 )</th>
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Compound 4

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Compound 5

(Chembridge ID No. 5456409),

Compound 6

(Chembridge ID No. 5464866),

Compound 7

(Chembridge ID No. 5531378),
Compound 8

(Chembridge ID No. 5537313),

Compound 9

(Chembridge ID No. 5538324),

Compound 10

(Chembridge ID No. 5539449),

Compound 11

(Chembridge ID No. 5549065),
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<td>20</td>
<td><img src="image" alt="Chemical Structure 20" /></td>
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</table>
comestibly or biologically acceptable derivatives thereof, or an enantiomer or diastereomer thereof.

9. A composition comprising a compound according to Formula (IV):

\[
\begin{align*}
&\text{(R}^1\text{)}_n \text{N} \text{C} \text{O} \text{N} \text{R}^3 \\
&\text{R}^2 \text{N} \text{R}^3 \text{N} \text{C} \text{O} \text{N} \text{(R}^4\text{)}_m
\end{align*}
\]

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

- \( R^1 \), independently for each occurrence, is selected from the group consisting of
- \( C_{1-10} \text{alkyl, } C_{1-10} \text{haloalkyl, } C_{2,10} \text{alkenyl, } C_{2,10} \text{alkynyl, halo, hydroxy, carboxyl,} \)
- \( C_{1,10} \text{alkoxycarbonyl, } C_{2,10} \text{alkenyloxycarbonyl, } C_{2,10} \text{alkynylxoycarbonyl, } C_{1-10} \text{acyl,} \)
- \( C_{1,10} \text{acylamino, } C_{1,10} \text{acyloxy, } C_{1,10} \text{carbonate, } C_{1,10} \text{alkoxy, phenylxoy, phenyl-C}_{1,10} \text{alkyloxyl,} \)
- \( C_{1,3} \text{heteroaryloxyl, } C_{1,3} \text{heteroaryl-C}_{1,10} \text{alkyloxyl, } C_{2,10} \text{alknyloxyl, } C_{2,10} \text{alknyloyxyl,} \)
- \( \text{phosphoryl, phosphate, phosphonate, phosphinate, amino, } diC_{1,10} \text{alkylaminolino,} \)
- \( \text{monoC}_{1,10} \text{alkylaminino, } C_{1,10} \text{amido, } C_{1,10} \text{amino, } C_{1,10} \text{carbamate, } C_{1,10} \text{urea, cyano, nitro,} \)

\text{SUBSTITUTE SHEET (RULE 26)}
azido, sulphydryl, C₁₋₁₀alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₃₋₅carbocycl, C₃₋₅carbocycl-C₁₋₄alkyl, C₁₋₅heterocycl, C₁₋₅heterocycl-C₁₋₄alkyl, phenyl, phenyl-C₁₋₄alkyl, C₁₋₅heteroaryl, and C₁₋₅heteroaryl-C₁₋₄alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R² is selected from the group consisting of hydrogen, C₁₋₁₀alkyl, C₁₋₁₀haloalkyl, C₂₋₁₀alkenyl, C₂₋₁₀alkynyl, carboxyl, C₁₋₁₀alkoxy carbonyl, C₂₋₁₀alkenyl oxy carbonyl, C₂₋₁₀alkyl, phosphoryl, phosphonate, phosphinate, cyan, sulfonate, sulfamoyl, sulfonyl, C₃₋₅carbocycl, C₃₋₅carbocycl-C₁₋₄alkyl, C₁₋₅heterocycl, C₁₋₅heterocycl-C₁₋₄alkyl, phenyl, phenyl-C₁₋₄alkyl, C₁₋₅heteroaryl, and C₁₋₅heteroaryl-C₁₋₄alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R¹ is selected from the group consisting of hydrogen, C₁₋₁₀alkyl, C₁₋₁₀haloalkyl, C₂₋₁₀alkenyl, C₂₋₁₀alkynyl, carboxyl, C₁₋₁₀alkoxy carbonyl, C₂₋₁₀alkenyl oxy carbonyl, C₂₋₁₀alkyl, phosphoryl, phosphonate, phosphinate, cyan, sulfonate, sulfamoyl, sulfonyl, C₃₋₅carbocycl, C₃₋₅carbocycl-C₁₋₄alkyl, C₁₋₅heterocycl, C₁₋₅heterocycl-C₁₋₄alkyl, phenyl, phenyl-C₁₋₄alkyl, C₁₋₅heteroaryl, and C₁₋₅heteroaryl-C₁₋₄alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R³, independently for each occurrence, is selected from the group consisting of C₁₋₁₀alkyl, C₁₋₁₀haloalkyl, C₂₋₁₀alkenyl, C₂₋₁₀alkynyl, halo, hydroxyl, carboxyl, C₁₋₁₀alkoxy carbonyl, C₂₋₁₀alkenyl oxy carbonyl, C₂₋₁₀alkyl, C₁₋₁₀acyl, C₁₋₁₀aclyarnino, C₁₋₁₀acylox, C₁₋₁₀carbonate, C₁₋₁₀alkoxy, phenolxy, phenyl-C₁₋₄alkyl oxy, C₁₋₅heteroaryl, C₁₋₅heteroaryl-C₁₋₄alkyl, C₃₋₅alkenyl oxy, C₁₋₁₀alkynoxy, phosphoryl, phosphate, phosphonate, phosphinite, amino, diC₁₋₁₀alkoxyarnino, monoC₁₋₁₀alkylarnino, C₁₋₁₀amidino, C₁₋₁₀imino, C₁₋₁₀carbamate, C₁₋₁₀urea, cyan, nitro, azido, sulphydryl, C₁₋₁₀alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₃₋₅carbocycl, C₃₋₅carbocycl-C₁₋₄alkyl, C₁₋₅heterocycl, C₁₋₅heterocycl-C₁₋₄alkyl, phenyl, phenyl-C₁₋₄alkyl, C₁₋₅heteroaryl, and C₁₋₅heteroaryl-C₁₋₄alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

wherein any of R¹, R², R³, and R⁴, independently and independently for each occurrence, is optionally substituted with 1-3 substituents selected from the group consisting of C₁₋₁₀alkyl, C₁₋₁₀haloalkyl, halo, hydroxyl, carboxyl, C₁₋₁₀alkoxy carbonyl, C₂₋₁₀alkenyl oxy carbonyl, C₂₋₁₀alkyl, C₁₋₁₀acyl, C₁₋₁₀aclyarnino, C₁₋₁₀acylox, C₁₋₁₀carbonate, C₁₋₁₀alkoxy, phenolxy, phosphoryl, phosphate, phosphonate, phosphinite, amino, diC₁₋₁₀alkoxyarnino, monoC₁₋₁₀alkylarnino, C₁₋₁₀amidino, C₁₋₁₀imino, C₁₋₁₀carbamate, C₁₋₁₀urea, cyan, nitro, azido, sulphydryl, C₁₋₁₀alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₃₋₅carbocycl, C₃₋₅carbocycl-C₁₋₄alkyl, C₁₋₅heterocycl,
C_{1-4} heterocyclic-C_{1-alkyl}, phenyl, phenyl-C_{4-alkyl}, C_{1-3} heteroaryl, and
C_{1-3} heteroaryl-C_{1-alkyl}; and wherein heterocyclic or heteroaromatic rings, independently
for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;
n is 0-2; and
m is 0-3;
wherein the composition is edible and capable of reducing bitter taste of a bitter
tastant.

10. The composition according to paragraph 9, wherein:
R^1, independently for each occurrence, is selected from the group consisting of
halo, C_{1-alkyl}, C_{2-alkenyl}, and C_{2-alkynyl};
R^2 is selected from the group consisting of hydrogen, C_{1-alkyl}, C_{2-alkenyl},
C_{2-alkynyl}, and C_{1-acyl};
R^3 is selected from the group consisting of hydrogen, C_{1-alkyl}, C_{2-alkenyl}, and
C_{2-alkynyl};
R^4, independently for each occurrence, is selected from the group consisting of
halo, C_{1-alkyl}, C_{2-alkenyl}, C_{2-alkynyl}, C_{1-alkoxy}, \text{-C(O)-O-R}^5, and \text{-C(O)-N(R)}^5;
R^5, independently for each occurrence, is selected from the group consisting of
hydrogen, C_{1-alkyl}, C_{2-alkenyl}, and C_{2-alkynyl};
wherein any of R^1, R^2, R^3, and R^4, independently and independently for each
occurrence, is optionally substituted as noted paragraph 9;
n is 0-2; and
m is 0-3.

11. The composition according to paragraph 9, wherein said compound
according to Formula (IV) is a compound according to Formula (Va):

\[
\begin{align*}
\text{Formula (Va):} \\
\text{or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or}
\text{diastereomer thereof,}
\end{align*}
\]

wherein, as valence and stability permit, R^1, R^2, R^3, R^4, and m are as defined in
paragraph 9.
12. The composition according to paragraph 11, wherein said compound according to Formula (Vla) is a compound according to Formula (VIa):

\[
\begin{array}{c}
\text{Formula (VIa);}
\end{array}
\]

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

\( R^1, R^2, R^3 \), and \( R^4 \) are as defined in paragraph 11; and \( o \) is 0-2.

13. The composition according to paragraph 12, wherein said compound according to Formula (VIIa) is a compound according to Formula (VIIa):

\[
\begin{array}{c}
\text{Formula (VIIa);}
\end{array}
\]

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit, \( R^1, R^2, R^3, R^4, R^5 \), and \( o \) are as defined in paragraph 12.

14. The composition according to paragraph 9, wherein said compound according to Formula (IV) is a compound according to Formula (Vb):
or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or
diastereomer thereof,

5 wherein, as valence and stability permit, $R^1$, $R^2$, $R^3$, $R^4$, and $m$ are as defined in
paragraph 9.

15. The composition according to paragraph 14 wherein said compound
according to Formula (Vb) is a compound according to Formula (VIb):

10 Formula (VIb);

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or
diastereomer thereof;

wherein, as valence and stability permit:

$R^1$, $R^2$, $R^3$, $R^4$, are as defined in paragraph 14; and

15 $o$ is 0-2.

16. The composition according to paragraph 15 wherein said compound
according to Formula (VIb) is a compound according to Formula (VIIb):

Formula (VIIb);
and compostibly or biologically acceptable derivatives thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit, \( R_1, R_2, R_3, R_4, R_5 \), and \( o \) are as defined in paragraph 9.

17. The composition according to paragraph 9, wherein said compound according to Formula (IV) is selected from the group consisting of:

<table>
<thead>
<tr>
<th>Compound 23</th>
<th>(Chembridge ID No. 7533235)</th>
</tr>
</thead>
</table>

<table>
<thead>
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<th>Compound 24</th>
<th>(Chembridge ID No. 6741054)</th>
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<tbody>
<tr>
<td>Compound 25</td>
<td>(Chembridge ID No. 7529691)</td>
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<tr>
<td>---------------------------------</td>
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<tr>
<td>![Compound 25 Diagram]</td>
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<table>
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<tr>
<th>Compound 26</th>
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<tbody>
<tr>
<td>![Compound 26 Diagram]</td>
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</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Compound 27</th>
<th>(Chembridge ID No. 9052639)</th>
</tr>
</thead>
<tbody>
<tr>
<td>![Compound 27 Diagram]</td>
<td></td>
</tr>
</tbody>
</table>
Compound 28

Chembridge ID No. 7500589

Compound 29

Chembridge ID No. 7530695

Compound 30

Chembridge ID No. 7903488

Compound 31

Chembridge ID No. 9052883
comestibly or biologically acceptable derivatives thereof, or an enantiomer or diastereomer thereof.

18. A composition comprising a compound according to Formula (VIII):

\[
\begin{align*}
&\text{Formula (VIII),} \\
&\text{or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or} \\
&\text{diastereomer thereof,} \\
&\text{wherein, as valence and stability permit:} \\
&R^1, \text{ independently for each occurrence, is selected from the group consisting of} \\
&\text{C}_{1-10}\text{alkyl, } \text{C}_{1-10}\text{haloalkyl, } \text{C}_{2-10}\text{alkenyl, } \text{C}_{2-10}\text{alkynyl, } \text{halo, hydroxyl, carboxyl,} \\
&\text{C}_{1-10}\text{alkoxycarbonyl, } \text{C}_{2-10}\text{alkenylcarbonyl, } \text{C}_{2-10}\text{alkynylcarbonyl, } \text{C}_{1-10}\text{acyl,} \\
&\text{C}_{1-10}\text{acylamino, } \text{C}_{1-10}\text{acyloxy, } \text{C}_{1-10}\text{carbonate, } \text{C}_{1-10}\text{alkoxy, phenyloxy,} \\
\end{align*}
\]
phenyl-C$_1$-alkoxy, C$_1$-heteroaryloxy, C$_1$-heteroaryl-C$_1$-alkyl, C$_3$-alkenyloxy, C$_3$-alkynloxy, phosphoryl, phosphate, phosphonate, phosphinate, amino, diC$_1$-alkylamino, monoC$_1$-alkylamino, C$_1$-amido, C$_1$-amino, C$_1$-carbamate, C$_1$-arene, cyano, nitro, azido, sulhydryl, C$_1$-alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C$_3$-carbocycleyl, C$_3$-carbocycleyl-C$_1$-alkyl, C$_1$-heterocyclel, C$_1$-hetereocycleyl-C$_1$-alkyl, phenyl, phenyl-C$_1$-alkyl, C$_1$-heteroaryl, and C$_1$-heteroaryl-C$_1$-alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R$^2$, independently for each occurrence, is selected from the group consisting of is selected from the group consisting of C$_1$-alkyl, C$_1$-haloalkyl, C$_2$-alkenyl, C$_2$-alkynyl, halo, hydroxy, carbonyl, C$_2$-alkoxycarbonyl, C$_2$-alkenyloxyacyl, C$_2$-alkynloxyacyl, C$_1$-acetyl, C$_1$-acylamino, C$_1$-acyloxy, C$_1$-carbonate, C$_1$-alkoxy, phenol, phenyl-C$_1$-alkoxy, C$_1$-heteroaryloxy, C$_1$-heteroaryl-C$_1$-alkyl, C$_3$-alkenyloxy, C$_3$-alkynloxy, phosphoryl, phosphate, phosphonate, phosphinate, amino, diC$_1$-alkylamino, monoC$_1$-alkylamino, C$_1$-amido, C$_1$-amino, C$_1$-carbamate, C$_1$-arene, cyano, nitro, azido, sulhydryl, C$_1$-alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C$_3$-carbocycleyl, C$_3$-carbocycleyl-C$_1$-alkyl, phenyl, phenyl-C$_1$-alkyl, C$_1$-heteroaryl, and C$_1$-heteroaryl-C$_1$-alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R$^3$, independently for each occurrence, is selected from the group consisting of C$_1$-alkyl, C$_1$-haloalkyl, C$_2$-alkenyl, C$_2$-alkynyl, halo, hydroxy, carbonyl, C$_2$-alkoxycarbonyl, C$_2$-alkenyloxyacyl, C$_2$-alkynloxyacyl, C$_1$-acetyl, C$_1$-acylamino, C$_1$-acyloxy, C$_1$-carbonate, C$_1$-alkoxy, phenol, phenyl-C$_1$-alkoxy, C$_1$-heteroaryloxy, C$_1$-heteroaryl-C$_1$-alkyl, C$_3$-alkenyloxy, C$_3$-alkynloxy, phosphoryl, phosphate, phosphonate, phosphinate, amino, diC$_1$-alkylamino, monoC$_1$-alkylamino, C$_1$-amido, C$_1$-amino, C$_1$-carbamate, C$_1$-arene, cyano, nitro, azido, sulhydryl, C$_1$-alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C$_3$-carbocycleyl, C$_3$-carbocycleyl-C$_1$-alkyl, C$_1$-heterocyclel, C$_1$-hetereocycleyl-C$_1$-alkyl, phenyl, phenyl-C$_1$-alkyl, C$_1$-heteroaryl, and C$_1$-heteroaryl-C$_1$-alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R$^4$ is selected from the group consisting of hydrogen C$_1$-alkyl, C$_1$-haloalkyl, C$_2$-alkenyl, C$_2$-alkynyl, carbonyl, C$_2$-alkoxycarbonyl, C$_2$-alkenyloxyacyl, C$_2$-alkynloxyacyl, C$_1$-acetyl, phosphoryl, phosphonate, phosphinate, cyano, sulfonate, sulfamoyl, sulfonyl, C$_3$-carbocycleyl, C$_3$-carbocycleyl-C$_1$-alkyl, C$_1$-heterocyclel, C$_1$-hetereocycleyl-C$_1$-alkyl, phenyl, phenyl-C$_1$-alkyl, C$_1$-heteroaryl, and
C₃₋₅ heteroaryl-C₁₋₅ alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

Ar is selected from the group consisting of C₆₋₁₀ aryl and C₅₋₁₀ heteroaryl;

Cy is a 5 to 7-membered carbocyclic or heterocyclic ring, wherein heterocyclic ring comprises 1-4 heteroatoms selected from N, O, and S;

wherein any of R¹, R², R³, and R⁴, independently and independently for each occurrence, is optionally substituted with 1-3 substituents selected from the group consisting of C₁₋₁₀ alkyl, C₁₋₁₀ haloalkyl, halo, hydroxy, carbonyl, C₁₋₁₀ alkoxycarbonyl, C₂₋₁₀ alkenyloxy carbonyl, C₂₋₁₀ alkynlyloxycarbonyl, C₁₋₁₀ acyl, C₁₋₁₀ acylamino, C₁₋₁₀ acyloxy, C₁₋₁₀ carbonate, C₁₋₁₀ alkoxycarbonyl, phenyloxy, phosphoryl, phosphate, phosphonate, phosphinate, amino, diC₁₋₁₀ alkylamino, monoC₁₋₁₀ alkylamino, C₁₋₁₂ amidino, C₁₋₁₂ imino, C₁₋₁₂ carbamate, C₁₋₁₂ urea, cyano, nitro, azido, sulphonyl, C₁₋₁₂ alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₁₋₁₂ carbocyclyl, C₃₋₁₀ carbocyclyl-C₁₋₅ alkyl, C₃₋₁₀ heterocyclyl, C₃₋₁₀ heterocyclyl-C₁₋₅ alkyl, phenyl, phenyl-C₁₋₅ alkyl, C₁₋₅ heteroaryl, and C₁₋₅ heteroaryl-C₁₋₅ alkyl; and wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

m is 1-3;

n is 0-3; and

α is 0-3;

wherein the composition is edible and capable of reducing bitter taste of a bitter tastant.

19. The composition according to paragraph 18, wherein:

R¹, independently for each occurrence, is selected from the group consisting of halo, hydroxy, C₁₋₁₀ alkyl, C₁₋₁₀ haloalkyl, C₂₋₁₀ alkenylnyl, C₂₋₁₀ alkynynyl, C₁₋₁₀ alkoxy, and C₁₋₁₀ acyloxy;

R², independently for each occurrence, is C₁₋₁₀ alkyl;

R³, independently for each occurrence, is selected from the group consisting of halo, C₁₋₁₀ alkyl, C₂₋₁₀ alkenylnyl, C₂₋₁₀ alkynynyl, C(O)-O-R⁴, and C(O)-N(R⁴)₂;

R⁴, independently for each occurrence, is selected from the group consisting of hydrogen, C₁₋₁₀ alkyl, C₂₋₁₀ alkenylnyl, and C₂₋₁₀ alkynynyl;

Ar is selected from the group consisting of C₆₋₁₀ aryl and C₅₋₁₀ heteroaryl;

Cy is a 5 to 7-membered carbocyclic or heterocyclic ring, optionally including one or two carbon-carbon or carbon-nitrogen double bonds in the ring;

wherein any of R¹, R², R³, and R⁴, independently and independently for each occurrence, is optionally substituted as described in paragraph 18;
m is 1-3;
n is 0-3; and
o is 0-3.

20. The composition according to paragraph 18, wherein said compound according to Formula (VIII) is a compound according to Formula (IX):

![Chemical Structure](image)

Formula (IX);
or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, wherein, as valence and stability permit, R₁, R₂, R₃, R₄, m, n, and o are as defined in paragraph 18.
21. The composition according to paragraph 20, wherein said compound according to Formula (IX) is a compound according to Formula (X):

![Chemical Structure](image)

Formula (X):

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomeric thereof,

wherein, as valence and stability permit:

$R^{1}$, $R^{3}$, $R^{4}$, $R^{a}$, and $n$ are as defined in paragraph 20; and

$p$ is 0-2.

5

22. The composition according to paragraph 18, wherein said compound according to Formula (VIII) is selected from the group consisting of:

<table>
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<tr>
<th>Compound 37</th>
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(Chembridge ID No. 5846684),
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<td>40</td>
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<td>(Asinex ID BAS02001668),</td>
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<tr>
<td>(Chembridge ID No. 5580105),</td>
<td></td>
</tr>
</tbody>
</table>
comestibly or biologically acceptable derivatives thereof.

23. A composition comprising a compound according to Formula (XI):

\[
\begin{array}{c}
\text{(R1)}_n \quad \text{N} \\
\text{R}^2 & \text{R}^3 & \text{R}^4
\end{array}
\]

5

Formula (XI);

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

\( R^1 \), independently for each occurrence, is selected from the group consisting of

\( C_{1-10} \)alkyl, \( C_{1-10} \)haloalkyl, \( C_{2-10} \)alkenyl, \( C_{2-10} \)alkynyl, halo, hydroxyl, carboxyl,

\( C_{1-10} \)alkoxycarbonyl, \( C_{2-10} \)alkenylOxy carbonyl, \( C_{2-10} \)alkynylOxy carbonyl, \( C_{1-10} \)aeryl,

\( C_{1-10} \)acylamino, \( C_{1-10} \)acyloxy, \( C_{1-10} \)carbonate, \( C_{1-10} \)alkoxy, phenyloxy, phenyl-\( C_{1-10} \)alkyloxy,
C$_{1-4}$heteroaryloxy, C$_{1-3}$heteroaryl-C$_{1-4}$alkyloxy, C$_{3-10}$alkenyloxy, C$_{3-10}$alkynloxy, phosphoryl, phosphate, phosphonate, phosphate, amino, diC$_{1-4}$alkylamino, monoC$_{1-4}$alkylamino, C$_{1-4}$amido, C$_{1-4}$diamo, C$_{1-4}$carbamate, C$_{1-4}$urea, cyano, nitro, azido, sulfhydryl, C$_{1-4}$alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C$_{3-7}$carbocycl, C$_{3-7}$carbocycl-C$_{1-4}$alkyl, C$_{1-4}$heterocycl, C$_{1-4}$heterocycl-C$_{1-4}$alkyl, phenyl, phenyl-C$_{1-4}$alkyl, C$_{1-4}$heteroary, and C$_{1-4}$heteroaryl-C$_{1-4}$alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R$^2$ is selected from the group consisting of hydrogen, C$_{1-4}$alkyl, Het-C$_{1-4}$alkyl, C$_{1-4}$haloalkyl, C$_{2-10}$alkenyl, C$_{2-10}$alkynyl, halo, hydroxyl, carboxyl, C$_{1-10}$alkoxycarbonyl, C$_{2-10}$alkenoylcarbonyl, C$_{2-10}$alkynlyoxycarbonyl, C$_{1-10}$acetyl, C$_{1-10}$acylamo, C$_{1-10}$acyloxy, C$_{1-10}$carbonyl, C$_{1-10}$alkoxy, phenol, phenyl-C$_{1-4}$alkyl, C$_{1-10}$heteroaryloxy, C$_{1-10}$heteroary, C$_{1-4}$diamo, C$_{1-4}$amino, C$_{1-4}$carbamate, C$_{1-4}$urea, cyano, nitro, azido, sulfhydryl, C$_{1-4}$alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C$_{3-7}$carbocycl, C$_{3-7}$carbocycl-C$_{1-4}$alkyl, phenyl, phenyl-C$_{1-4}$alkyl, C$_{1-4}$heteroary, and C$_{1-4}$heteroaryl-C$_{1-4}$alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R$^3$ is selected from the group consisting of hydrogen, C$_{1-4}$alkyl, Het-C$_{1-4}$alkyl, C$_{1-4}$haloalkyl, C$_{2-10}$alkenyl, C$_{2-10}$alkynyl, halo, hydroxyl, carboxyl, C$_{1-10}$alkoxycarbonyl, C$_{2-10}$alkenoylcarbonyl, C$_{2-10}$alkynlyoxycarbonyl, C$_{1-10}$acetyl, C$_{1-10}$acylamo, C$_{1-10}$acyloxy, C$_{1-10}$carbonyl, C$_{1-10}$alkoxy, phenol, phenyl-C$_{1-4}$alkyl, C$_{1-10}$heteroaryloxy, C$_{1-10}$heteroary, C$_{1-4}$diamo, C$_{1-4}$amino, C$_{1-4}$carbamate, C$_{1-4}$urea, cyano, nitro, azido, sulfhydryl, C$_{1-4}$alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C$_{3-7}$carbocycl, C$_{3-7}$carbocycl-C$_{1-4}$alkyl, phenyl, phenyl-C$_{1-4}$alkyl, C$_{1-4}$heteroary, and C$_{1-4}$heteroaryl-C$_{1-4}$alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;
sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₃₋₇ carboxycyl, C₃₋₇ carboxycyl-C₆₋₁₈ alkyl, C₃₋₇ heterocyclyl, C₁₋₄ heterocyclyl-C₆₋₁₈ alkyl, phenyl, phenyl-C₆₋₁₈ alkyl, C₃₋₇ heteroaryl, and C₃₋₇ heteroaryl-C₆₋₁₈ alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

or R³ and R⁴ together with the atoms to which they are attached form a 5 to 6-membered aryl or heteroaryl ring optionally substituted by 1 to 4 groups selected from the group consisting of Het, C₆₋₁₈ alkyl, C₆₋₁₈ haloalkyl, halo, hydroxyl, carboxyl, C₆₋₁₈ alkoxy carbonyl, C₂₋₁₀ alkenyl oxy carbonyl, C₂₋₁₀ alkylnitroxyl carbonyl, C₁₋₄ acetyl, C₁₋₄ aminocyan, C₁₋₄ acyl oxo, C₁₋₄ carbonoyl, C₁₋₄ alkoxy, phenol, phosphoryl, phosphate, phosphonate, phosphinate, amino, diC₁₋₄ alkylamino, monoC₁₋₄ alkylamino, C₁₋₄ amido, C₁₋₄ imino, C₁₋₄ carbamate, C₁₋₄ urea, cyano, nitro, azido, sulphydryl, C₁₋₄ alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₃₋₇ carboxycyl, C₃₋₇ carboxycyl-C₆₋₁₈ alkyl, C₃₋₇ heterocyclyl, C₁₋₄ heterocyclyl-C₆₋₁₈ alkyl, phenyl, phenyl-C₆₋₁₈ alkyl, C₃₋₇ heteroaryl, and C₃₋₇ heteroaryl-C₆₋₁₈ alkyl; and wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

Het is a C₃₋₇ heterocyclyl including 1-4 heteroatoms in the ring selected from oxygen, sulfur, and nitrogen;

wherein any of R¹, R², R³, R⁴, and Het, independently and independently for each occurrence, is optionally substituted with 1-3 substituents selected from the group consisting of C₆₋₁₈ alkyl, C₆₋₁₈ haloalkyl, halo, hydroxyl, carboxyl, C₆₋₁₈ alkoxy carbonyl, C₂₋₁₀ alkenyl oxy carbonyl, C₂₋₁₀ alkylnitroxyl carbonyl, C₁₋₄ acetyl, C₁₋₄ aminocyan, C₁₋₄ acyl oxo, C₁₋₄ carbonoyl, C₁₋₄ alkoxy, phenol, phosphoryl, phosphate, phosphonate, phosphinate, amino, diC₁₋₄ alkylamino, monoC₁₋₄ alkylamino, C₁₋₄ amido, C₁₋₄ imino, C₁₋₄ carbamate, C₁₋₄ urea, cyano, nitro, azido, sulphydryl, C₁₋₄ alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₃₋₇ carboxycyl, C₃₋₇ carboxycyl-C₆₋₁₈ alkyl, C₁₋₄ heterocyclyl, C₁₋₄ heterocyclyl-C₆₋₁₈ alkyl, C₄₋₁₀ aryly, C₁₋₄ alkyl-C₆₋₁₀ ary, C₆₋₁₀ ary-C₁₋₄ alkyl, C₁₋₄ heteroaryl, and C₁₋₄ heteroaryl-C₆₋₁₈ alkyl; and wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S; and

n is 0-4;

wherein the composition is edible and capable of reducing bitter taste of a bitter tastant.

24. The composition according to paragraph 23, wherein as valence and stability permit:

R¹, independently for each occurrence, is selected from the group consisting of halo, hydroxyl, C₁₋₄ alkyl, C₁₋₄ haloalkyl, C₂₋₁₀ alkenyl, C₂₋₁₀ alkynyl, and C₁₋₄ alkoxy;
R² is selected from the group consisting of hydrogen, halo, hydroxyl, C₁₋₆alkyl, C₁₋₆haloalkyl, Het-C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, and C₁₋₆alkoxy;

R³ is selected from the group consisting of hydrogen, halo, hydroxyl, C₁₋₆alkyl, C₁₋₆haloalkyl, Het-C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, and C₁₋₆alkoxy;

R⁴ is selected from the group consisting of hydrogen, halo, hydroxyl, C₁₋₆alkyl, C₁₋₆haloalkyl, Het-C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, and C₁₋₆alkoxy;

or R³ and R⁴ together with the atoms to which they are attached form a 5 to 6-membered aryl ring optionally substituted by 1 to 4 groups selected from the group consisting of halo, hydroxyl, C₁₋₆alkyl, C₁₋₆haloalkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₁₋₆alkoxy, and Het;

Het is a C₂₋₆heterocycyl including 1-3 heteroatoms in the ring selected from oxygen, sulfur, and nitrogen and is optionally substituted with one or more groups selected from the group consisting of halo, hydroxyl, C₁₋₆alkyl, C₁₋₆haloalkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₁₋₆alkoxy, and C₆₋₁₀aryl optionally substituted by C₁₋₆alkyl;

wherein any of R¹, R², R³, R⁴, and Het, independently and independently for each occurrence, is optionally further substituted as in paragraph 23; and

n is 0-4.

25. The composition according to paragraph 23, wherein said compound according to Formula (XI) is a compound according to Formula (XIIa):

![Formula (XIIa)](image)

Formula (XIIa);

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

R¹, R², Het, and n are as defined in paragraph 23;

R³, independently for each occurrence, is selected from the group consisting of halo, hydroxyl, C₁₋₆alkyl, C₁₋₆haloalkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, and C₁₋₆alkoxy; and

m is 0-3.

26. The composition according to paragraph 25, wherein said compound according to Formula (XIIa) is a compound according to Formula (XIIIa):
Formula (XIIa);
or a comestibly or biologically acceptable salt or derivative thereof, or an
enantiomer or diastereomer thereof, wherein, as valence and stability permit, R¹, R², R⁵,
Het, n, and m are as defined in paragraph 25.

27. The composition according to paragraph 23, wherein said compound
according to Formula (XI) is a compound according to Formula (XIIb):

Formula (XIIb);
or a comestibly or biologically acceptable salt or derivative thereof, or an
enantiomer or diastereomer thereof, wherein, as valence and stability permit, R¹, R², R³,
Het, and n are as defined in paragraph 23.

28. The composition according to paragraph 27, wherein said compound
according to Formula (XIIb) is a compound according to Formula (XIIib):

Formula (XIIib);
or a comestibly or biologically acceptable salt or derivative thereof; or an enantiomer or
diastereomer thereof,
wherein, as valence and stability permit:
R₁, R₂, R₃, Het, and n are as in paragraph 27; and
Ar is C₆₋₁₀aryl, such as phenyl or naphthyl, optionally substituted by C₁₋₆alkyl.

29. The composition according to paragraph 23, wherein said compound according to Formula (XI) is selected from the group consisting of:

<table>
<thead>
<tr>
<th>Compound 44</th>
<th><img src="image" alt="Chembridge ID No. 7728336" /></th>
</tr>
</thead>
<tbody>
<tr>
<td>Compound 45</td>
<td><img src="image" alt="Chembridge ID No. 7733323" /></td>
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<tr>
<td>Compound 46</td>
<td><img src="image" alt="Chembridge ID No. 7726077" /></td>
</tr>
<tr>
<td>Compound 47</td>
<td><img src="image" alt="Chembridge ID No. 9149274" /></td>
</tr>
</tbody>
</table>
comestibly or biologically acceptable derivatives thereof.

30. A composition comprising a compound according to Formula (XIV):

\[
\begin{align*}
\text{Compound 48} & \\
\text{(Chembidge ID No. 9126324), and} & \\
\end{align*}
\]

\[
\begin{align*}
\text{Formula (XIV);} & \\
\end{align*}
\]

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

R\(^{1}\), independently for each occurrence, is selected from the group consisting of

- C\(_{1-10}\)alkyl, C\(_{1-10}\)haloalkyl, C\(_{2-10}\)alkynyl, halo, hydroxy, carboxyl,
- C\(_{1-10}\)alkoxyacarbonyl, C\(_{2-10}\)alkoxyloxyacarbonyl, C\(_{2-10}\)alkynylloxyacarbonyl, C\(_{1-10}\)acyl,
- C\(_{1-10}\)acylamino, C\(_{1-10}\)acyloxy, C\(_{1-10}\)carbonate, C\(_{1-10}\)alkoxy, phenol, phenyl-C\(_{1-10}\)alkoxy,
- C\(_{1-10}\)heteroaryloxy, C\(_{1-10}\)heteroaryl-C\(_{1-10}\)alkoxy, C\(_{1-10}\)alkenylloxy, C\(_{3-10}\)alkynylloxy,
- phosphoryl, phosphate, phosphonate, phosphinate, amino, diC\(_{1-10}\)alkylamino,
- monoC\(_{1-10}\)alkylamino, C\(_{1-10}\)amido, C\(_{1-10}\)diamo, C\(_{1-10}\)carbamate, C\(_{1-10}\)urea, cyano, nitro,
- azido, sulphydryl, C\(_{1-10}\)alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl,
- C\(_{2-10}\)carboxycarbonyl, C\(_{2-10}\)carboxyloxy-C\(_{1-10}\)alkyl, C\(_{2-10}\)heterocyclic-C\(_{1-10}\)alkyl, C\(_{2-10}\)heterocyclic-C\(_{1-10}\)alkyl, phenyl, phenyl-C\(_{1-10}\)alkyl, C\(_{1-4}\)heteroaryl, and C\(_{1-4}\)heteroaryl-C\(_{1-4}\)alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R\(^{2}\) is selected from the group consisting of C\(_{1-10}\)alkyl, C\(_{1-10}\)haloalkyl, C\(_{2-10}\)alkynyl,
- C\(_{2-10}\)alkynyl, hydroxy, C\(_{1-10}\)alkoxy, C\(_{6-10}\)aryloxy, C\(_{6-10}\)aryloxy-C\(_{1-10}\)alkyl,
- C\(_{6-10}\)arylamino-C\(_{1-10}\)alkyl, C\(_{6-10}\)aryl-C\(_{1-10}\)alkyl, C\(_{1-10}\)heteroaryloxy,
- C\(_{3-10}\)heteroaryloxy-C\(_{1-10}\)alkyl, C\(_{3-10}\)heteroarylamino-C\(_{1-10}\)alkyl, C\(_{3-10}\)heteroaryl-C\(_{1-10}\)alkyl,
- C\(_{3-10}\)alkenylloxy, C\(_{3-10}\)alkynylloxy, amino, diC\(_{1-10}\)alkylamino, monoC\(_{1-10}\)alkylamino,
sulphydryl, C_{1-10}alkylthio, C_{3-10}carbocycleyloxy,
C_{3-10}carbocycleyl-C_{1-6}alkyl, C_{3-10}carbocycleyloxy-C_{1-6}alkyl,
C_{3-10}carbocycleylamino-C_{1-6}alkyl,
C_{3-10}heterocycleyl, C_{1-4}heterocycleyl-C_{1-6}alkyl, C_{1-4}heterocycleyloxy-C_{1-6}alkyl,
C_{1-4}heterocyclicamino-C_{1-6}alkyl, C_{6-10}aryl, C_{6-10}arylmethyl-C_{1-6}alkyl, C_{1-6}heteroaryl, and
C_{1-6}heteroarylmethyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R’ is selected from the group consisting of hydrogen, C_{1-10}alkyl, C_{1-10}haloalkyl,
C_{2-10}alkenyl, C_{2-10}alkynyl, carboxyl, C_{1-10}alkoxycarbonyl, C_{2-10}alkenyloxycarbonyl, C_{2-10}alkynloxycarbonyl, C_{1-10}alkyl, phosphoryl, phosphonate, phosphinate, cyanocarbonato, sulfonate, sulfamoyl, sulfonyl, C_{3-7}carbocycleyl, C_{3-7}carbocycleyloxy-C_{1-6}alkyl, C_{1-4}heterocycleyl,
C_{1-4}heterocycleyl-C_{1-6}alkyl, phenyl, phenyl-C_{1-6}alkyl, C_{1-6}heteroaryl, and
C_{1-6}heteroarylmethyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

wherein any of R’, R”, and R’’, independently and independently for each occurrence, is optionally substituted with 1-3 substituents selected from the group consisting of C_{1-10}alkyl, C_{1-10}haloalkyl, halo, hydroxyl, carboxyl, C_{1-10}alkoxycarbonyl, C_{2-10}alkenyloxycarbonyl, C_{2-10}alkynloxycarbonyl, C_{1-10}alkyl, C_{1-10}acylamino, C_{1-10}acyloxy, C_{1-10}carbonate, C_{1-10}alkoxy, C_{6-10}aryloxy, C_{6-10}arylamino, phosphoryl, phosphinate, phosphonate, phosphinate, amino, diC_{1-10}alkylamino, monoc_{1-10}alkylamino, C_{1-6}amido,
C_{1-6}iminio, C_{1-6}carbamate, C_{1-6}urea, cyano, nitro, azido, sulphydryl, C_{1-6}alkylthio,
sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C_{3-7}carbocycleyl,
C_{3-7}carbocycleyl-C_{1-6}alkyl, C_{1-4}heterocycleyl, C_{1-4}heterocycleyl-C_{1-6}alkyl, phenyl, phenyl-C_{1-6}alkyl, C_{1-6}heteroaryl, and C_{1-6}heteroarylmethyl; and wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S; and

n is 0-3;

wherein the composition is edible and capable of reducing bitter taste of a bitter tastant.

The composition according to paragraph 30, wherein as valence and stability permit:

R’, independently for each occurrence, is selected from the group consisting of halo, hydroxyl, C_{1-6}alkyl, C_{1-6}haloalkyl, C_{2-6}alkenyl, C_{2-6}alkynyl, C_{1-6}alkoxy, and
C_{1-6}acyloxy;

R’’ is selected from the group consisting of C_{1-6}alkyl, C_{1-6}alkoxy-substituted
C_{1-6}alkyl, C_{2-6}alkenyl, C_{2-6}alkynyl, C_{6-10}arylmethyl-C_{1-6}alkyl, and
-(CH_{2})_{n}X_{p}-Ar, wherein aryl groups of R’ are optionally substituted by one or more halo, hydroxyl, C_{1-6}alkyl, C_{1-6}haloalkyl, C_{1-6}alkoxy, or C_{1-6}acyloxy;
R' is selected from the group consisting of hydrogen, C₆-alkyl, C₆-alkenyl, and C₃-alkynyl;

X is selected from the group consisting of O, NH, and CH₂;

Ar is selected from the group consisting of C₆-aryl, C₆-heteroaryl,

C₅-10carbocyclic, and C₆-heterocyclic, including fused bicyclic groups, wherein Ar is optionally substituted by one or more halo, hydroxyl, C₆-alkyl, C₆-haloalkyl, C₆-alkoxy, or C₆-acyl oxy;

wherein any of R¹, R², and R³, independently and independently for each occurrence, is optionally further substituted as in paragraph 30;

m is 1-3;

n is 0-3; and

p is 0 or 1.

32. The composition according to paragraph 30, wherein said compound according to Formula (XIV) is a compound according to Formula (XVa):

![Formula (XVa)](image)

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

R¹ and R³ are as defined in paragraph 30; and

R² is C₆-alkyl, such as methyl or ethyl.

33. The composition according to paragraph 30, wherein said compound according to Formula (XIV) is a compound according to Formula (XVb):

![Formula (XVb)](image)
or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, wherein, as valence and stability permit, \( R^1, R^a, X, Ar \), and \( n \) are as defined in paragraph 30.

34. The composition according to paragraph 30, wherein said compound according to Formula (XIV) is a compound according to Formula (XVc):

\[
\begin{align*}
\text{Formula (XVc),} \\
\end{align*}
\]

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, wherein, as valence and stability permit, \( R^1, R^a, Ar \), and \( n \) are as defined in paragraph 30.

35. The composition according to paragraph 30, wherein said compound according to Formula (XIV) is selected from the group consisting of:

<table>
<thead>
<tr>
<th>Compound</th>
<th>Structure</th>
</tr>
</thead>
<tbody>
<tr>
<td>49</td>
<td><img src="image" alt="Structure 49" /></td>
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<td>(Chembridge ID No. 5838356),</td>
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<td></td>
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</tr>
</tbody>
</table>
A composition comprising:

(a) a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIa), Formula (IIIb), Formula (IIIb*), Formula (IV), Formula (Va), Formula (Vb),

Formulas (VIa), Formula (VIb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIA), Formula (XIIIB), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof; and
(b) a bitter tastant,
wherein the composition is edible.

37. The composition according to paragraph 36, wherein the bitter tastant is a
foodstuff.

5 38. The composition according to paragraph 36, wherein the bitter tastant is a
bitter tasting salt.

39. The composition according to paragraph 38, wherein the bitter tasting salt is
a magnesium salt, a calcium salt, or a potassium salt.

40. The composition according to paragraph 40, wherein the potassium
containing salt is KCl or potassium lactate.

41. The composition of any one of paragraphs 1-40, wherein the edible
composition further comprises one or more components selected from the group consisting of:
NaCl, sodium lactate, and sugar.

42. A food product comprising the compositions of any one of paragraphs 1-41.

43. A method of preparing an edible composition comprising:
(a) providing comestibly acceptable carrier; and
(b) adding to the comestibly acceptable carrier of (a) a compound according to
Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''),
Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (Vla),
Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa),
Formula (XIIb), Formula (XIIa), Formula (XIIIb), Formula (XIIIa), Formula (XIV), Formula (XV),
Formula (XVI) or Formula (XVe), as described herein, or combinations thereof, or any one of
Compounds 1-58, as described herein, or combinations thereof.

44. The method according to paragraph 43, wherein said comestibly acceptable
carrier is inherently bitter.

45. The method according to paragraph 44, wherein the comestibly acceptable
carrier comprises a bitter tasting salt.

46. The method according to paragraph 45, wherein the bitter tasting salt is a
magnesium salt, a calcium salt, or a potassium salt.

47. The method according to paragraph 46, wherein the potassium salt is KCl or
potassium lactate.
48. The method according to any one of paragraphs 43-47, wherein the edible composition further comprises one or more components selected from the group consisting of: NaCl, sodium lactate, and sugar.

49. The method according to paragraph 43, wherein the method further comprises:

(c) adding a bitter tastant.

50. The method according to paragraph 49, wherein the bitter tastant is a bitter tasting salt.

51. The method according to paragraph 50, wherein the bitter tasting salt is a magnesium salt, a calcium salt, or a potassium salt.

52. The method according to paragraph 51, wherein the potassium salt is KCl or potassium lactate.

53. The method according to any one of paragraphs 49-52, wherein the edible composition further comprises one or more components selected from the group consisting of: NaCl, sodium lactate, and sugar.

54. A method of reducing the amount of NaCl in an edible composition comprising:

(a) replacing an amount of NaCl present in an edible composition with an amount of KCl; and

(b) adding to the edible composition generated in (a) an effective amount of a compound according to Formula (I), Formula (Ia), Formula (Iib), Formula (IIb), Formula (IIIb), Formula (IIIb*), Formula (IV), Formula (V), Formula (Vb), Formula (Va), Formula (VIa), Formula (VIb), Formula (VIiA), Formula (VIib), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XV), Formula (XVI), or Formula (XVii), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.

55. The method according to paragraph 54, wherein the amount of compound added in (b) is sufficient to permit replacement of the amount of NaCl typically present in the edible composition by up to 25%.

56. The method according to paragraph 54, wherein the amount of compound added in (b) is sufficient to permit replacement of the amount of NaCl typically present in the edible composition by up to 50%.
57. The method according to paragraph 54, wherein the amount of compound added in (b) is sufficient to permit replacement of the amount of NaCl typically present in the edible composition by up to 75%.

58. The method according to paragraph 54, wherein the amount of compound added in (b) is sufficient to permit replacement of the amount of NaCl typically present in the edible composition by up to 100%.

59. The method according to any one of paragraphs 54-58, wherein the edible composition maintains a salty flavor.

60. A method of reducing the amount of sodium lactate in an edible composition comprising:

(a) replacing an amount of sodium lactate present in an edible composition with an amount of potassium lactate; and

(b) adding to the edible composition generated in (a) an effective amount of a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IVb), Formula (Vla), Formula (Vb), Formula (VIa), Formula (VIb), Formula (VIIa), Formula (VIIb), Formula (VIIia), Formula (VIIib), Formula (VIIia), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIIb), Formula (XIIIa), Formula (XIIIb'), Formula (XIV), Formula (XVla), Formula (XVib) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58 as described herein, or combinations thereof.

61. The method according to paragraph 60, wherein the amount of compound added in (b) is sufficient to permit replacement of the amount of sodium lactate typically present in the edible composition by up to 25%.

62. The method according to paragraph 60, wherein the amount of compound added in (b) is sufficient to permit replacement of the amount of sodium lactate typically present in the edible composition by up to 50%.

63. The method according to paragraph 60, wherein the amount of compound added in (b) is sufficient to permit replacement of the amount of sodium lactate typically present in the edible composition by up to 75%.

64. The method according to paragraph 60, wherein the amount of compound added in (b) is sufficient to permit replacement of the amount of sodium lactate typically present in the edible composition by up to 100%.

65. The method according to any one of paragraphs 60-64, wherein the edible composition has the same shelf life as an edible composition comprising sodium lactate.
66. A method of reducing the amount of sugar in an edible composition comprising:

(a) replacing an amount of sugar present an edible composition with an amount of Acesulfame K; and

(b) adding to the edible composition generated in (a) an effective amount of a compound according to Formula (I), Formula (Ia), Formula (IIb), Formula (IIb'), Formula (IIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIa), Formula (XIIIb), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.

67. The method according to paragraph 66, wherein the amount of compound added in (b) is sufficient to permit replacement of the amount of sugar typically present in the edible composition by up to 25%.

68. The method according to paragraph 66, wherein the amount of compound added in (b) is sufficient to permit replacement of the amount of sugar typically present in the edible composition by up to 50%.

69. The method according to paragraph 66, wherein the amount of compound added in (b) is sufficient to permit replacement of the amount of sugar typically present in the edible composition by up to 75%.

70. The method according to paragraph 66, wherein the amount of compound added in (b) is sufficient to permit replacement of the amount of sugar typically present in the edible composition by up to 100%.

71. The method according to any one of paragraphs 66-70, wherein the edible composition maintains a sweet flavor.

72. A method of reducing the sodium intake of a subject, the method comprising:

(a) replacing an amount of a sodium salt present in an edible composition with an amount of a potassium salt; and

(b) adding to the edible composition generated in (a) an effective amount of a compound according to Formula (I), Formula (Ia), Formula (IIb), Formula (IIb'), Formula (IIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.
73. The method according to paragraph 72, wherein the sodium salt is NaCl and the potassium salt is KCl.

74. The method according to paragraph 72, wherein the sodium salt is sodium lactate, and the potassium salt is potassium lactate.

75. The method according to any one of paragraphs 72-74, wherein the method further comprises (c) identifying a subject in need thereof.

76. The method according to any one of paragraphs 72-75, wherein the amount of compound added in (b) is sufficient to reduce sodium intake by up to 25\% by replacement with potassium.

77. The method according to any one of paragraphs 72-75, wherein the amount of compound added in (b) is sufficient to reduce sodium intake by up to 50\% by replacement with potassium.

78. The method according to any one of paragraphs 72-75, wherein the amount of compound added in (b) is sufficient to reduce sodium intake by up to 75\% by replacement with potassium.

79. The method according to any one of paragraphs 72-75, wherein the amount of compound added in (b) is sufficient to reduce sodium intake by up to 100\% by replacement with potassium.

80. A method of reducing the sugar intake of a subject, the method comprising:
   (a) replacing an amount of sugar present in an edible composition with an amount of a Acesulfame K; and
   (b) adding to the edible composition generated in (a) an effective amount of a compound according to Formula (I), Formula (Iia), Formula (Iib), Formula (IIIb), Formula (IIIb\textsuperscript{a}), Formula (IV), Formula (V), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.

81. The method according to paragraph 80, wherein the method further comprises (c) identifying a subject in need thereof.

82. The method according to paragraph 80 or 81, wherein the amount of compound added in (b) is sufficient to reduce sugar intake by up to 25\% by replacement with Acesulfame K.
83. The method according to paragraph 80 or 81, wherein the amount of compound added in (b) is sufficient to reduce sugar intake by up to 50% by replacement with Acesulfame K.

84. The method according to paragraph 80 or 81, wherein the amount of compound added in (b) is sufficient to reduce sugar intake by up to 75% by replacement with Acesulfame K.

85. The method according to paragraph 80 or 81, wherein the amount of compound added in (b) is sufficient to reduce sugar intake by up to 100% by replacement with Acesulfame K.

86. A method of reducing bitter taste attributed to a bitter tastant in an edible composition comprising:

(a) adding an effective amount of a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIlb), Formula (IIIb'), Formula (IIlb"), Formula (IV), Formula (V), Formula (VIa), Formula (VIb), Formula (VIIa), Formula (VIIb), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof, to the edible composition such that any bitter taste induced by the bitter tastant is reduced.

87. A method of reducing bitter taste attributed to a bitter tastant in an edible composition comprising:

(a) ingesting an effective amount of a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIlb), Formula (IIIb'), Formula (IIlb"), Formula (IV), Formula (V), Formula (VIa), Formula (VIb), Formula (VIIa), Formula (VIIb), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof, along with the edible composition such that any bitter taste induced by the bitter tastant is reduced.

88. The method according to any one of paragraphs 43-87 or 89-98, wherein the edible composition is a food product, a consumer product, or a pharmaceutical composition.

89. The method according to any one of paragraphs 86-88, wherein the bitter taste induced by the bitter tastant is reduced by up to 25%
90. The method according to any one of paragraphs 86-88, wherein the bitter taste induced by the bitter tastant is reduced by up to 50%.

91. The method according to any one of paragraphs 86-88, wherein the bitter taste induced by the bitter tastant is reduced by up to 75%.

92. The method according to any one of paragraphs 86-88, wherein the bitter taste induced by the bitter tastant is reduced by up to 100%.

93. The method according to any one of paragraphs 86-92, wherein the bitter tastant is a bitter tasting salt.

94. The method according to paragraph 93, wherein the bitter tasting salt is a magnesium salt, a calcium salt, or a potassium salt.

95. The method according to paragraph 94, wherein the potassium salt is KCl or potassium lactate.

96. The method according to any one of paragraphs 86-95, wherein the edible composition further comprises NaCl, sodium lactate, or sugar.

97. A method of preserving an edible composition comprising:
   (a) providing an edible composition; and
   (b) combining with the edible composition of (a) potassium lactate and a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIb), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.

98. A method of reducing the amount of sodium in an edible composition while preserving the edible composition, the method comprising:
   (a) replacing an amount of sodium lactate present in an edible composition with an amount of potassium lactate; and
   (b) adding to the edible composition generated in (a) an effective amount of a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIb), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.
99. A method of inhibiting, reducing, or eliminating a bitter taste in a subject comprising:
   (a) placing a compound according to Formula (I), Formula (IIa), Formula (IIb),
   Formula (IIIb), Formula (IIIb*), Formula (IV), Formula (Va), Formula (Vb),
   Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb),
   Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.

100. The method according to paragraph 99, wherein the bitter taste is due to a bitter tasting salt.

101. The method according to paragraph 100, wherein the bitter taste is due to a magnesium salt, a calcium salt, or a potassium salt.

102. The method according to paragraph 101, wherein the bitter taste is due to KCl or potassium lactate.

103. A pharmaceutical composition comprising:
   (a) a bitter tasting pharmaceutical active ingredient; and
   (b) a compound according to Formula (I), Formula (IIa), Formula (IIb),
   Formula (IIIb), Formula (IIIb*), Formula (IV), Formula (Va), Formula (Vb),
   Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb),
   Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.

104. A pharmaceutical composition comprising:
   (a) a pharmaceutical active ingredient;
   (b) a bitter tastant; and
   (c) a compound according to Formula (I), Formula (IIa), Formula (IIb),
   Formula (IIIb), Formula (IIIb*), Formula (IV), Formula (Va), Formula (Vb),
   Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb),
   Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.
105. A consumer product comprising:
   (a) a bitter tasting ingredient; and
   (b) a compound according to Formula (I), Formula (IIa), Formula (IIb),
   Formula (IIIb), Formula (IIIb'), Formula (IIIb*'), Formula (IV), Formula (V), Formula (Vb),
   Formula (VIa), Formula (VId), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId),
   Formula (XIV), Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId),
   Formula (XIV), Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   (c) any one of Compounds 1-58, as described herein, or combinations thereof,
   or any one of Compounds 1-58, as described herein, or combinations thereof.

106. A consumer product for reducing bitter taste of a bitter tantant, wherein said consumer product comprises:
   (a) a compound according to Formula (I), Formula (IIa), Formula (IIb),
   Formula (IIIb), Formula (IIIb'), Formula (IIIb*'), Formula (IV), Formula (V), Formula (Vb),
   Formula (VIa), Formula (VId), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   (c) any one of Compounds 1-58, as described herein, or combinations thereof,
   or any one of Compounds 1-58, as described herein, or combinations thereof.

107. A method of inhibiting a bitter taste receptor comprising:
   (a) contacting the bitter taste receptor with a compound according to Formula (I),
   Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb*'), Formula (IV),
   Formula (V), Formula (Vb), Formula (VIa), Formula (VId), Formula (VIIa), Formula (VIIb),
   Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb),
   Formula (XIIId), Formula (XIV), Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIId), Formula (XIV),
   Formula (XVIIa), Formula (XVIIb), Formula (XVIII), Formula (XIX),
   (c) any one of Compounds 1-58, as described herein, or combinations thereof,
   or any one of Compounds 1-58, as described herein, or combinations thereof.

108. The method according to paragraph 107, wherein the bitter taste receptor is in the oral cavity of a subject.

109. The method according to paragraph 107, wherein the bitter taste receptor is in the gastrointestinal tract of a subject.

110. The method according to paragraph 107, wherein the bitter taste receptor is present in an in vitro assay.

Brief Description of the Drawings

[0068] Figures 1A-L disclose exemplary data for solution and foodstuf taste testing of compositions comprising compounds of Formula (I) of the present invention.
[0069] Figures 2A-H disclose exemplary data for solution and foodstuff taste testing of compositions comprising compounds of Formula (IV) of the present invention.

[0070] Figures 3A-D disclose exemplary data for solution and foodstuff taste testing of compositions comprising compounds of Formula (VIII) of the present invention.

[0071] Figures 4A-C disclose exemplary data for solution and foodstuff taste testing of compositions comprising compounds of Formula (XI) of the present invention.

[0072] Figure 5A-E disclose exemplary data for solution and foodstuff taste testing of compositions comprising compounds of Formula (XIV) of the present invention.

Detailed Description of the Invention

[0073] In order that the invention described herein may be fully understood, the following detailed description is set forth.

[0074] Unless defined otherwise, all technical and scientific terms used herein have the same meaning as those commonly understood by one of ordinary skill in the art to which this invention belongs. Although methods and materials similar or equivalent to those described herein can be used in the practice or testing of the present invention, suitable methods and materials are described below. The materials, methods and examples are illustrative only, and are not intended to be limiting. All publications, patents and other documents mentioned herein are incorporated by reference in their entirety.

[0075] Throughout this specification, the word "comprise" or variations such as "comprises" or "comprising" will be understood to imply the inclusion of a stated integer or groups of integers but not the exclusion of any other integer or group of integers.

[0076] The term "acyl" refers to an alkylcarbonyl, alkenylcarbonyl, alkynylcarbonyl or arylcarbonyl substituent, wherein the alky, alkenyl, alkynyl or aryl portion may be optionally substituted. Examples of acyl substituents include, but are not limited to, acetyl, propionyl, butyryl and benzoyl.

[0077] The term "acyloxy" refers to an \(-\text{O-C(O)R}\) substituent, wherein \(R\) is alkyl, alkenyl, alkynyl or aryl, and wherein the alky, alkenyl, alkynyl or aryl portion may be optionally substituted. Examples of acyloxy groups include, but are not limited to, acetoxy, propanoyloxy, butanoyloxy, pentanoyloxy and benzoyloxy.

[0078] The term "aliphatic" refers to straight chain or branched hydrocarbons that are completely saturated or that contain one or more units of unsaturation. For example, aliphatic groups include substituted or unsubstituted linear or branched alkyl, alkenyl and alkynyl groups. Unless indicated otherwise, the term "aliphatic" encompasses both substituted and unsubstituted hydrocarbons.

[0079] The terms "alkylamide," "alkenylamide" and "alkynylamide" refer to amides of the structures alkyl-NR-C(=O)-, alkenyl-NR-C(=O)-, and alkynyl-NR-C(=O)-, wherein \(R\) may be separately defined, or \(R\) is also alkyl, alkenyl or alkynyl.

[0080] The term "alkoxy" refers to \(O\)-alkyl substituent, wherein the alkyl portion may be optionally substituted. Examples of alkoxy substituents include, but are not limited to, methoxy,
ethoxy, n-propoxy, isopropoxy and n-butoxy. Also explicitly included within the scope of the term “alkoxy” are O-alkenyl or O-alkynyl groups. In all cases, the alkyl, alkene and alkyne portions may be optionally substituted.

[0081] The term "alkyl" refers to both straight and branched saturated chains containing, for example, 1-3, 1-6, 1-9, or 1-12 carbon atoms. An alkyl group may be optionally substituted.

[0082] The term "alkythio" refers to an S-alkyl substituent, wherein the alkyl portion may be optionally substituted. Examples of alkythio substituents include, but are not limited to, methylthio, ethylthio and isopropylthio. Also explicitly included within the scope of the term "alkythio" are S-alkenyl or S-alkynyl groups. In all cases, the alkyl, alkene and alkyne portions may be optionally substituted.

[0083] The term "alkenyl" refers to both straight and branched saturated chains containing, for example, 2-3, 2-6, 2-9, or 2-12 carbon atoms, and at least one carbon-carbon double bond. An alkenyl group may be optionally substituted.

[0084] The term "alkynyl" refers to both straight and branched saturated chains containing, for example, 2-3, 2-6, 2-9, or 2-12 carbon atoms, and at least one carbon-carbon triple bond. An alkynyl group may be optionally substituted.

[0085] The term "aralkyl" refers to an alkyl group substituted by an aryl. Also explicitly included within the scope of the term "aralkyl" are alkenyl or alkynyl groups substituted by an aryl. Examples of aralkyl groups include benzyl and phenethyl. An aralkyl group may be optionally substituted.

[0086] The terms "artificial sweetener" and "sugar substitute" refer to a food additive that confers a sweet taste but has less caloric energy than sugar. In some instances, the caloric energy of the "artificial sweetener" or "sugar substitute" is negligible.

[0087] The term "aryl" refers to monocyclic or polycyclic aromatic carbon ring systems having five to fourteen members. Examples of aryl groups include, but are not limited to, phenyl (Ph), 1-naphthyl, 2-naphthyl, 1-anthracyl and 2-anthracyl. An aryl group may be optionally substituted.

[0088] The term "aryllactoxy" refers to a group having the structure –O–R–Ar, where R is alkyl and Ar is an aromatic substituent. Also explicitly included within the scope of the term "aryllactoxy" are –O–R–Ar groups, wherein R is alkenyl or alkynyl. In all cases, the alkyl, alkene, alkyne and aryl portions may be optionally substituted.

[0089] The term "bitter" or "bitter taste" as used herein refers to the perception or gustatory sensation resulting following the detection of a bitter tastant. The following attributes may contribute to bitter taste: astringent, bitter-astringent, metallic, bitter-metallic, as well as off-tastes, aftertastes and undesirable tastes including but not limited to freeze-burn and cardboard taste, and/or any combinations of these. It is noted that, in the art, the term "off-taste" is often synonymous with "bitter taste." Without being limited by theory, the diversity of bitter tastes may reflect the large number of bitter receptors and the differential detection of bitter tastants by these receptors. Bitter taste as used herein includes activation of a bitter taste receptor by a bitter tastant. Bitter taste as used herein also includes activation of a bitter taste receptor by a bitter tastant.
followed by downstream signaling. Bitter taste as used herein also includes activation of a signaling pathway after stimulation by a bitter tastant. Bitter taste as used herein further includes perception resulting from signaling following the detection of a bitter tastant by a bitter taste receptor. Bitter taste as used herein further includes perception resulting from signaling following contacting a bitter taste receptor with a bitter tastant. Bitter taste can be perceived in the brain.

[0090] The term "bitter taste receptor" refers to a receptor, typically a cell surface receptor, to which a bitter tastant can bind. Bitter taste receptors may be present in the oral cavity, and/or throughout the gastrointestinal tract, including the stomach, intestines, and colon. Bitter receptors can also be present in vitro, such as in an assay, including but not limited to a cell based assay or a binding assay.

[0091] The term "bitter tastant," "bitter ligand," or "bitter compound" refers to a compound that activates or that can be detected by a bitter taste receptor and/or confers the perception of a bitter taste in a subject. A "bitter tastant" also refers to a multiplicity of compounds that combine to activate or be detected by a bitter taste receptor and/or confer the perception of a bitter taste in a subject. A "bitter tastant" further refers to a compound that is enzymatically modified upon ingestion by a subject to activate or be detected by a bitter taste receptor and/or confer the perception of a bitter taste in a subject. Because the perception of bitter taste may vary from individual to individual, some individuals may describe a "bitter tastant" as a compound which confers a different kind of bitter taste compared to the kind of bitter taste perceived for the same compound by other individuals. The term bitter tastant also refers to a compound which confers a bitter taste. Those of skill in the art can readily identify and understand what is meant by a bitter tastant. Non-limiting examples of bitter tastants or substances including foods that comprise a bitter tastant and taste bitter include coffee, unsweetened cocoa, marmalade, bitter melon, beer, bitters, citrus peel, dandelion greens, escarole, quinine, magnesium salts, calcium salts, potassium salts, KCl, potassium lactate, Acesulfame K, Brussels sprouts, asparagus, bitter gourd, wild cucumber, celery, hops, kohlrabi, radish leaf, ginseng, pumpkin, collard greens, kale, sparteine, caffeine, atropine, nicotine, urea and strychnine.

[0092] Further examples of bitter tastants include pharmaceuticals. Non-limiting examples of pharmaceuticals as bitter tastants include acetaminophen, ampicillin, azithromycin, chlorpheniramine, cimetidine, dextromethorphan, diphenhydramine, erythromycin, ibuprofen, penicillin, phenylbutazone, pseudoephedrine, ranitidine, spironolactone and theophylline all of which have been associated with bitter taste.

[0093] The term "carboceyl" or "carboceyclic," refers to monocyclic or polycyclic non-aromatic carbon ring systems, which may contain a specified number of carbon atoms, preferably from 3 to 12 carbon atoms, which are completely saturated or which contain one or more units of unsaturation. A carboceyclic ring system may be monocyclic, bicyclic or tricyclic. A carboceyclic ring may be fused to another ring, such as an aryl ring or another carboceyclic ring. Examples of carboceyclic rings could include cyclohexyl, cyclopentyl, cyclobutyl, cyclopenty1, cyclohexenyl, cyclooctenyl, indanyl, tetrahydroanaphthyl and the like. The term "carboceyclic" or "carboceyl,"
whether saturated or unsaturated, also refers to rings that are optionally substituted unless indicated. The term "carbocyclic" or "carbocyclic" also encompasses hybrids of aliphatic and carbocyclic groups, such as (cycloalkyl)alkyl, (cycloalkenyl)alkyl and (cycloalkyl)alkenyl.

[0094] The term "comestibly or biologically acceptable salt" refers to any comestibly or biologically acceptable salt, ester, or salt of such ester, of a compound of the present invention, which, upon ingestion, is capable of providing (directly or indirectly) a compound of the present invention, or a metabolite, residue or portion thereof, characterized by the ability to reduce the perception of a bitter taste attributed to a bitter tastant. Similarly, the term "comestibly or biologically acceptable derivative" refers to any comestibly or biologically acceptable derivative of a compound of the present invention, which, upon ingestion, is capable of providing (directly or indirectly) a compound of the present invention, or a metabolite, residue or portion thereof, characterized by the ability to reduce the perception of a bitter taste attributed to a bitter tastant. A "comestibly product" is a product suitable for oral use, such as eating or drinking. Therefore, a comestibly acceptable compound is an edible compound.

[0095] The term "consumer product" refers to health and beauty products for the personal use and/or consumption by a subject. Consumer products may be present in any form including, but not limited to, liquids, solids, semi-solids, tablets, capsules, lozenges, strips, powders, gels, gums, pastes, slurries, syrups, aerosols and sprays. Non-limiting examples of consumer products include nutriceuticals, nutritional supplements, lipsticks, lip balms, soaps, shampoos, gums, adhesives (e.g., dental adhesives), toothpastes, oral analgesics, breath fresheners, mouthwashes, tooth whiteners, and other dentifrices.

[0096] The term "diet" collectively refers to the food products and/or beverages consumed by a subject. A subject's "diet" also includes any consumer products or pharmaceutical compositions the subject ingests.

[0097] The term "edible composition" refers to a composition suitable for consumption, typically via the oral cavity (although consumption may occur via non-oral means such as inhalation). Edible compositions may be present in any form including, but not limited to, liquids, solids, semi-solids, tablets, lozenges, powders, gels, gums, pastes, slurries, syrups, aerosols and sprays. As used herein, edible compositions include food products, pharmaceutical compositions, and consumer products. The term edible compositions also refers to, for example, dietary and nutritional supplements. As used herein, edible compositions also include compositions that are placed within the oral cavity but not swallowed, including professional dental products, such as dental treatments, fillings, packing materials, molds and polishes. The term "comestible" refers to similar compositions and is generally used as a synonym to the term "edible."

[0098] The term "effective amount" refers to an amount sufficient to produce a desired property or result. For example, an effective amount of a compound of the present invention is an amount capable of reducing the perception of bitter taste associated with a bitter tastant. The term "effective amount" of a compound of the invention also refers to an amount which, when added to an edible composition, reduces the bitter taste of, e.g., a NaCl substitute, thereby allowing for the
maintenance of the perception of a desired salty flavor of a said edible composition. The term "effective amount of a compound" also refers to an amount which, when added to an edible composition, allows for the preservation of a food product, while reducing or eliminating bitter taste associated with a bitter tastant in the preservative. The term "effective amount" also refers to the amount of a compound of the present invention capable of reducing or eliminating the perception of a bitter taste or aftertaste associated with either a bitter tastant in a food product or an inherently bitter food product.

[0099] The term "flavor modifier" refers to a compound or a mixture of compounds that, when added to an edible composition, such as a food product, modifies (e.g., masks, eliminates, decreases, reduces or enhances the perception of) a flavor (e.g., sweet, salty, umami, sour, or bitter taste) present in the edible composition.

[0100] The term "food product" refers to any compositions comprising one or more processed foodstuff. Food products include, but are not limited to, confectionaries, bakery products (including, but not limited to, doughs, breads, cakes, biscuits, crackers, pastries, pies, tarts, quiches, and cookies), ice creams (including but not limited to impulse ice cream, take-home ice cream, frozen yogurt, gelato, sorbet, sherbet and soy, oat, bean and rice-based ice cream), dairy products (including, but not limited to, drinking milk, cheese, yogurt, and sour milk drinks), cheeses (including, but not limited to, natural cheeses and processed cheeses), butter, margarine, sweet and savory snacks (including but not limited to fruit snacks, chips/crisps, tortilla/corn chips, popcorn, pretzels, chocolates, and nuts), hot and cold beverages (including, but not limited to, beverages, beverage mixes, concentrates, juices, carbonated beverages, non-carbonated beverages, alcoholic beverages, non-alcoholic beverages, soft drinks, sports drinks, isotonic drinks, coffees, teas, bottled waters, and beverages prepared from botanicals and botanical extracts (including cold beverages that are prepared with botanical or fungi extracts as ingredients, and drinks that are prepared in various ways, such as infusions, decoctions, or other means of extraction or distillation of various plant parts, including, but not limited to leaves, flowers, stems, fruits, roots, rhizomes, stems, bark, volatile oils, or even the whole plant), snack bars (including, but not limited to granola bars, muesli bars, protein bars, breakfast bars, energy bars, and fruit bars), meal replacement products, ready meals (including, but not limited to canned meals, preserved meals, frozen meals, dried meals, chilled meals, dinner mixes, frozen pizza, chilled pizza, and prepared salads), soups (including but not limited to broth-like soups and cream-based soups), broth, gravy, soy sauce, meats and fish (including raw, cooked, and dried meats), deli products (including but not limited to meats and cheeses suitable for slicing or pre-sliced meats and cheeses, e.g., turkey, chicken, ham, bologna, salami, bierwurst, capicola, chorizo, corned beef, dutch loaf, Serrano, prosciutto, head cheese, liverwurst, meatloaf (including olive loaf, pepper loaf, pimento loaf, and ham and cheese loaf), mortadella, pastrami, pepperoni, roast beef, roast pork, saucisson, smoked meat, summer sausage, tongue, American cheese, blue cheese, cheddar cheese, Colby cheese, Colby-Jack cheese, gouda, Monterey Jack cheese, muenster cheese mozzarella, parmigiano cheese, pepper jack cheese, provolone, romano cheese, string cheese, spray cheese, and swiss cheese), vegetables (including,
but not limited to, raw, pickled, cooked, and dried vegetables, such as french fries), fruits (including raw, cooked, and dried fruits), grains (including, but not limited to, dried cereals and breads), prepared foods (including, but not limited to, dried, canned, or jarred sauces and soups), snack foods, pastas (including, but not limited to, fresh pasta, chilled pasta, frozen pasta, dried pasta), noodles (including, but not limited to, egg noodles, wheat noodles, rice noodles, mung bean noodles, potato noodles, buckwheat noodles, corn noodles, cellophane noodles, chow mein, fettuccini, fusilli, gnocchi, lasagna, linguini, lo mein, macaroni, manicotti, pad thai, penne, ramen, rice vermicelli, rigatoni, soba, spaghetti, spatzle, udon, and ziti), canned foods, frozen foods, dried foods, chilled foods, oils and fats, baby food, spreads, salads, cereals (including, but not limited to, hot and cold cereals), sauces (including, but not limited to, tomato pastes, tomato purees, bouillon cubes, stock cubes, table sauces, boys bases sauces, pasta sauces, cooking sauces, marinades, dry sauces, powder mixes, ketchups, mayonnaise, salad dressings, vinegrettes, mustards, and dips), jellies, jams, preserves, honey, puddings, recipe mixes, syrups, icings, fillings, infused foods, salt-preserved food, marinated foods and condiments (such as ketchup, mustard and steak sauce).

In some embodiments, the food product is animal feed. For example, the food product may be a pet food product, i.e. a food product for consumption by a household pet. In other embodiments, the food product is a livestock food product, i.e. a food product for consumption by livestock.

[0101] The term "foodstuff" refers to an unprocessed ingredient or a basic nutrient or flavor containing element used to prepare a food product. Non-limiting examples of foodstuffs include:

- fruits, vegetables, meats, fishes, grains, milks, eggs, tubers, sugars, sweeteners, oils, herbs, snacks, sauces, spices and salts.

[0102] The term "halo" or "halogen" refers to a fluorine, chlorine, bromine or iodine substituent.

[0103] The term "heteroaryl" refers to monocyclic or polycyclic aromatic systems having five to fourteen members and one or more heteroatoms. One having ordinary skill in the art will recognize that the maximum number of heteroatoms in a stable, chemically feasible heteroaryl ring is determined by the size of the ring and valence. The term "heteroaalkyl" refers to an alkyl group substituted by a heteroaryl. Also explicitly included within the scope of the term "heteroaalkyl" are alkenyl or alkynyl groups substituted by a heteroaryl. In general, a heteroaaryl ring may have one to four heteroatoms. Heteroaaryl groups include, without limitation, 2-furanyl, 3-furanyl,

N-imidazolyl, 2imidazolyl, 4-imidazolyl, 5-imidazolyl, 3-isoxazolyl, 4isoxazolyl, 5-isoxazolyl, 2-oxadiazolyl, 5-oxadiazolyl, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl, 2-pyrrolyl, 3-pyrrolyl, 2-pyrrolidyl, 4-pyrrolidyl, 2-pyrimidyl, 4-pyrimidyl, 5-pyrimidyl, 3-pyridazinyl, 2-thiazolyl, 4thiazolyl, 5-thiazolyl, 5-tetrazolyl, 2-triazolyl, 5-triazolyl, 2-thienyl, and 3-thienyl. The term "heteroaaryl ring", "heteroaaryl group", or "heteroaalkyl" also refers to rings that are optionally substituted.

Examples of fused polycyclic heteroaaryl and aryl ring systems in which a carbocyclic aromatic ring or heteroaaryl ring is fused to one or more other rings include, tetrahydroanaphthyl, benzimidazolyl, benzothienyl, benzofuranyl, indolyl, quinolinyl, benzothiazolyl, benzoazolyl, benzimidazolyl, isoquinolinyl, isindolyl, acridinyl, benzoisoxazolyl, and the like.
The term "heterocyclic" or "heterocyclyl" refers to non-aromatic saturated or unsaturated monocyclic or polycyclic ring systems containing one or more heteroatoms and with a ring size of three to fourteen. One having ordinary skill in the art will recognize that the maximum number of heteroatoms in a stable, chemically feasible heterocyclic ring is determined by the size of the ring, degree of unsaturation, and valence. In general, a heterocyclic ring may have one to four heteroatoms so long as the heterocyclic ring is chemically feasible and stable and may be fused to another ring, such as a carbocyclic, aryl or heteroaryl ring, or to another heterocyclic ring. A heterocyclic ring system may be monocyclic, bicyclic or tricyclic. Also included within the scope of within the scope of the term "heterocyclic" or "heterocyclyl", as used herein, is a group in which one or more carbocyclic rings are fused to a heteroaryl. Examples of heterocyclic rings include, but are not limited to, 3-1H-benzimidazol-2-one, 3-1H-alkyl-benzimidazol-2-one, 2-tetrahydrofuranyl, 3-tetrahydrofuranyl, 2-tetrahydrothiophenyl, 3-tetrahydrothiophenyl, 2-morpholino, 3-morpholino, 4-morpholino, 2-thiomorpholino, 3-thiomorpholino, 4-thiomorpholino, 1-pyrrolidinyl, 2-pyrrolidinyl, 3-pyrrolidinyl, 1-piperazinyl, 2-piperazinyl, 1-piperidinyl, 2-piperidinyl, 3-piperidinyl, 4-piperidinyl, 4-thiazolidinyl, diazolonyl, N-substituted diazolonyl, 1-phthalimidinyl, benzoxyan, benzotriazol-1-yl, benzopyrrolidinyl, benzoazepinidine, benzoazolone, benzoazoline, benzothiophene, azirinyl, oxiranyl, azetidinyl, pyrrolidinyl, diazolonyl, imidazolidinyl, imidazolidinyl, pyrazolonyl, pyrazolonyl, pyranyl, dioxanyl, dihydroxan, thrioxan, quinuclidinyl, oxepanyl, succinimidyl and the like.

The term "isoprene" (also referred to as "isoterpene") refers to 2-methyl-1,3-butadiene and is represented by the formula CH₂=C(CH₃)CH=CH₂.

The terms "parts per million" and "ppm" are used in the food industry to refer to a low concentration of a solution. For example, one gram of solute in 1000 ml of solvent has a concentration of 1000 ppm and one thousandth of a gram (0.001g) of solute in 1000 ml of solvent has a concentration of one ppm. Accordingly, a concentration of one milligram per liter (i.e. 1 mg/L) is equal to 1 ppm.

The terms "perception of a bitter taste," "perception of saltiness," "perception of a flavor" and similar terms, refer to the awareness of a subject of a particular taste or flavor.

The term "pharmaceutically active ingredient" refers to a compound in a pharmaceutical composition which is biologically active.

The term "potassium salt" refers to a salt wherein potassium is the cation. Potassium salts in the context of the present invention are preferably edible potassium salts including, but not limited to, Acetate K (Ace K), aluminum potassium sulfate, dipotassium guanylate, dipotassium inosinate, monopotassium glutamate, potassium acetate, potassium acid tartrate, potassium acid tartrate, potassium adipate, potassium alginate, potassium aluminum silicate, potassium ascorbate, potassium aspartate, potassium benzoate, potassium bicarbonate, potassium bisulfate, potassium bisulfite, potassium bromate, potassium carbonate, potassium chloride, potassium citrate, potassium dihydrogen citrate, potassium dihydrogen phosphate, potassium ferrocyanide, potassium fumarate, potassium giberellate, potassium gluconate, potassium
hydroxide, potassium hydrogen sulfite, potassium iodide, potassium lactate, potassium malate,
potassium metabisulfite, potassium nitrate, potassium nitrite, potassium persulfate, potassium
phosphate (dibasic), potassium phosphate (monobasic), potassium phosphate (tribasic), potassium
polynaphosphate, potassium polyphosphates, potassium pyrophosphate, potassium propionate,
potassium saccharin, potassium sodium tartrate (e.g., potassium sodium L(+)-tartrate), potassium
sorbate, potassium sulfate, potassium sulfite, and potassium tripolyphosphate.

[0110] The term "processed foodstuff" refers to a foodstuff has been subjected to any process
which alters its original state (excluding, e.g., harvesting, slaughtering, and cleaning). Examples of
methods of processing foods include, but are not limited to, removal of unwanted outer layers, such
as potato peeling or the skinning of peaches; chopping or slicing; mincing or macerating;
liquefaction, such as to produce fruit juice; fermentation (e.g. beer); emulsification; cooking, such
as boiling, broiling, frying, heating, steaming or grilling; deep frying; baking; mixing; addition of
gas such as air entrainment for bread or gasification of soft drinks; proofing; seasoning (with, e.g.,
herbs, spices, salts); spray drying; pasteurization; packaging (e.g., canning or boxing); extrusion;
puffing; blending; and preservation (e.g., adding salt, sugar, potassium lactate or other
preservatives).

[0111] The term "replace" or "replacing" refers to substituting one compound for another
compound in or in the preparation of, for example, an edible composition, such as food product. It
includes complete and partial replacements or substitutions.

[0112] The term "salty flavor" refers to the taste elicited by, for example, ions of alkali metals
salts (e.g., Na" and Cl in sodium chloride). Non-limiting examples of compositions eliciting a salty
flavor include table salt (sodium chloride), sea water, sea salt and potassium chloride. The amount
of salty flavor or the saltiness of a composition can be determined by, e.g., taste testing.

[0113] The term "sodium" or "sodium salt" refers to the amount of sodium (i.e., sodium salt)
ingested or otherwise consumed by a subject. In general, "sodium" or a "sodium salt" refers to a
salt or compound wherein sodium is the cation. Sodium salts in the context of the present invention
include, but are not limited to, aluminium sodium sulfate, calcium disodium EDTA, diocetyl sodium
sulfosuccinate, disodium 5'-ribonucleotides, disodium ethylenediaminetetraacetate, disodium
guanilate, disodium inosinate sodium acetate, monosodium glutamate (MSG), potassium sodium
tartrate, sodium acid pyrophosphate, sodium adipate, sodium alginate, sodium aluminosilicate,
sodium aluminum phosphate (acidic), sodium aluminum phosphate (basic), sodium ascorbate,
sodium benzoate, sodium bicarbonate, sodium bisulfate, sodium bisulfite, sodium carbonate,
sodium carboxymethylcellulose, sodium caseinate, sodium chloride, sodium citrate, sodium
cyclamate, sodium dehydroacetate, sodium diacetate, sodium dehydroacetate, sodium dihydrogen
citrate, sodium dihydrogen phosphate, sodium DL-malate, sodium erythorbate, sodium erythorbina,
sodium ethyl para-hydroxybenzoate, sodium ferrocyanide, sodium ferrocyanide, sodium
formate, sodium fumarate, sodium gluconate, sodium hydrogen carbonate, sodium hydrogen DL-
malate, sodium hydrogen acetate, sodium hydrogen sulfite, sodium hydroxide, sodium
hypophosphite, sodium tartrate (e.g., sodium L(+)-tartrate), sodium lactate, sodium laurel sulfate,
sodium malate, sodium metabisulfite, sodium metaphosphate, sodium methyl para-
hydroxybenzoate, sodium nitrate, sodium nitrite, sodium O-phenylphenol, sodium phosphate
(dibasic), sodium phosphate (monobasic), sodium phosphate (tribasic), sodium polyphosphate,
sodium potassium tartrate, sodium propionate, sodium propyl para-hydroxybenzoate, sodium
pyrophosphate, sodium saccharin, sodium sesquicarbonate, sodium stearoyl lactylate, sodium
stearyl fumarate, sodium succinate, sodium sulfate, and starch sodium octenylsuccinate.

[0114] The term "sodium intake" refers to the amount of sodium ingested or otherwise consumed
by a subject.

[0115] The term "stability" or "stable" in the context of a chemical structure refers to the chemical
state when a system is in its lowest energy state, or in chemical equilibrium with its environment.
Thus, a stable compound (or, e.g., a compound containing a number of atoms or substitutions that
are stable) is not particularly reactive in the environment or during normal use, and retains its useful
properties on the timescale of its expected usefulness.

[0116] The term "subject" refers to a mammal. In preferred embodiments, the subject is human.
In some embodiments, a subject is a domestic or laboratory animal, including but not limited to,
household pets, such as dogs, cats, pigs, rabbits, rats, mice, gerbils, hamsters, guinea pigs, and
ferrets. In some embodiments, a subject is a livestock animal. Non-limiting examples of livestock
animals include: alpaca, bison, camel, cattle, deer, pigs, horses, llamas, mules, donkeys, sheep,
goats, rabbits, reindeer, and yak.

[0117] The term "sugar" refers to a simple carbohydrate, such as a monosaccharide or a
disaccharide, that delivers a primary taste sensation of sweetness. Non-limiting examples of sugar
include glucose, fructose, galactose, sucrose, lactose, and maltose.

[0118] The term "sweet flavor" refers to the taste elicited by, for example, sugars. Non-limiting
examples of compositions eliciting a sweet flavor include glucose, sucrose, fructose, saccharin,
cyclamate, aspartame, acesulfame potassium, sucralose, aspartame, and neotame. The amount of
sweet flavor or the sweetness of a composition can be determined by, e.g., taste testing.

[0119] The term "terpenes" refers to compounds comprising repeating units of isoprene. The
basic molecular formula of a terpene is \((C_5H_8)_n\) where \(n\) is the number of linked isoprene units.

[0120] The term "terpenoids" refers to compounds comprising terpenes and derivatives thereof.
Thus, in some embodiments, terpenoids have at least one \(C_5H_8\) hydrocarbon unit with one or more
points of unsaturation. In other embodiments, terpenoids comprise saturated terpene units and
derivatives thereof and have no points of unsaturation.

[0121] An aryl, aralkyl, heteroaryl, or heteroaralkyl group may contain one or more
independently selected substituents. Examples of suitable substituents on the unsaturated carbon
atom of an aryl or heteroaryl group include, but are not limited to, halogen, -CF\(_3\), \(-R^1\), -OR\(_1\), -OH,
-SH, -SR\(_2\), protected OH (such as acyloxy), \(-NO_2\), -CN, -NH\(_2\), -NHR\(_1\), \(-N(R'_2)\), -NHCOR\(_1\), 
-NHCONH\(_2\), -NHCONHR\(_1\), -NHCON(R\(_2\))\(_2\), -NRCOR\(_1\), -NHCO\(_2\)H, -NHCO\(_2\)R\(_1\), -CO\(_2\)R\(_1\), -CO\(_2\)H,
-COR\(_1\), -CONH\(_2\), -CONHR\(_1\), -CON(R\(_2\))\(_2\), -SO\(_2\)H, -SO\(_2\)R\(_1\), -SO\(_2\)R\(_2\), -SO\(_2\)NH\(_2\)
-SO\(_2\)H, -SO\(_2\)R\(_1\), -SO\(_2\)NHR\(_1\), -SO\(_2\)N(R\(_2\))\(_2\), -NHSO\(_2\)H, or \(-NHSO\(_2\)R\(_1\), where R\(_1\) is selected.
from H, aliphatic, carbocyclic, heterocyclic, aryl, aralkyl, heteroaryl, or heteroaralkyl and each R’ is optionally substituted with one or more halogen, nitro, cyano, amino, -NH-(unsubstituted aliphatic), -N-(unsubstituted aliphatic), carboxy, carbamoyl, hydroxy, -O-(unsubstituted aliphatic), -SH, -S-(unsubstituted aliphatic), CF₃, -SO₂NH₂ unsubstituted aliphatic, unsubstituted carbocyclic, unsubstituted heterocyclic, unsubstituted aryl, unsubstituted aralkyl, unsubstituted heteroaryl, or unsubstituted heteroaralkyl.

[0122] An aliphatic group, a carbocyclic ring or a heterocyclic ring may contain one or more substituents. Examples of suitable substituents on a saturated or unsaturated carbon of an aliphatic group, a carbocyclic ring or a heterocyclic ring include, but are not limited to, those listed above for the unsaturated carbon as well as the following: =O, =S, =NNHR', =NN(R')₂, =N-O'R, =NNHCOR', =NNHCO₂R', =NNHSO₃R', =N-CN, or =NR', wherein R' is as defined above. Guided by this specification, the selection of suitable substituents is within the knowledge of one skilled in the art.

[0123] As defined herein, the compounds of the invention are intended to include all stereoisomeric forms of the compound, including geometric isomers (i.e., E, Z) and optical isomers (i.e., R, S). Single stereoisomeric forms as well as enantiomeric and diastereomeric mixtures of the present compounds are within the scope of the invention. Unless otherwise stated, formulas depicted herein are also meant to include compounds which differ only in the presence of one or more isotopically enriched atoms. For example, compounds having the present formulas except for the replacement of a hydrogen by a deuterium or tritium, or the replacement of a carbon by a ¹³C- or ¹⁴C-enriched carbon are within the scope of this invention.

[0124] The present invention provides edible compositions comprising a compound of the present invention, including food products, consumer products, and pharmaceutical compositions comprising said compounds, and methods of preparing such compositions. The present invention also provides methods of reducing the amount of sodium (e.g., NaCl or sodium lactate) or sugar in a food product, a method of reducing the sodium or sugar intake in a diet, a method of reducing bitter taste, and a method of reducing activity of a bitter taste receptor. The present invention also includes reducing the amount of sodium in an edible composition or diet by replacing a sodium containing compound or composition with a potassium containing compound or composition. The present invention also includes reducing the amount of sugar in an edible composition or diet by replacing sugar with a potassium containing sweetener, such as Acesulfame K.

**Edible compositions**

[0125] According to one aspect, the invention provides an edible composition comprising a compound of the invention for reducing bitter taste of a bitter tastant.

**Edible compositions comprising diphenyl-containing compounds**

[0126] The substituent definitions in this section (i.e., R¹, R², R³, X, m, and n) refer to compounds of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb') and Formula (IIIb*).
All stereochemical forms of the compounds disclosed in this and any section herein are specifically contemplated, including geometric isomers (i.e., E, Z) and optical isomers (i.e., R, S). Single stereochemical isomers as well as enantiomeric and diastereomeric mixtures of the compounds disclosed in this and any section herein are also specifically contemplated.

In some embodiments, the present invention provides an edible composition for reducing bitter taste of a bitter tastant, wherein the composition comprises a diphenyl-containing compound. The diphenyl-containing compounds of this invention are capable of reducing or eliminating bitter taste of a bitter tastant. In some embodiments, the diphenyl-containing compound has a molecular weight less than about 1000, 500, or 300 daltons. In certain embodiments, the diphenyl-containing compound is a compound of Formula (I):

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(R^1)_n
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(R^2)_m
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wherein, as valence and stability permit:

R^1, independently for each occurrence, is selected from the group consisting of C_{1-10}alkyl, C_{1-10}haloalkyl, C_{2-10}alkenyl, C_{2-10}alkynyl, halo, hydroxyl, carboxyl, C_{1-10}alkoxy carbonyl, C_{2-10}alkenyl oxy carbonyl, C_{2-10}alkynyl oxy carbonyl, C_{1-10}acyl, C_{1-10}acetylamino, C_{1-10}acyloxy, C_{1-10}carbonate, C_{1-10}alkoxy, C_{1-10}aryloxy,

C_{6-10}aryl-C_{1-10}alkoxy, C_{1-10}heteroaryl oxy, C_{1-10}heteroarylamino, C_{1-10}heteroaryloxy, C_{1-10}alkenyl oxy,

C_{3-10}alkoxy, phosphonyl, phosphate, phosphonate, phosphinate, amino, diC_{1-10}alkyl amine, monoC_{1-10}alkyl amine, C_{1-10}amido, C_{1-10}imino, C_{1-10}carbamate, C_{1-10}urea, cyano, nitro, azido, sulfhydryl, C_{1-10}alkylthio, sulfite, sulfonate, sulfamoyl, sulfonylamido, sulfonyl, C_{1-10}carboxyethyl, C_{1-10}carboxyethoxy-C_{1-10}alkyl, C_{1-10}heterocyclyl,

C_{1-10}heterocyclyl-C_{1-10}alkyl, phenyl, phenyl-C_{1-4}alkyl, C_{1-10}heteroaryl, and C_{1-10}heteroarylamino, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R^2, independently for each occurrence, is selected from the group consisting of C_{1-10}alkyl, C_{1-10}haloalkyl, C_{2-10}alkenyl, C_{2-10}alkynyl, halo, hydroxyl, carboxyl, C_{1-10}alkoxy carbonyl, C_{2-10}alkenyl oxy carbonyl, C_{2-10}alkynyl oxy carbonyl, C_{1-10}acyl, C_{1-10}acetylamino, C_{1-10}acyloxy, C_{1-10}carbonate, C_{1-10}alkoxy, C_{1-10}aryloxy,

C_{6-10}aryl-C_{1-10}alkoxy, C_{1-10}heteroaryl oxy, C_{1-10}heteroarylamino, C_{1-10}heteroaryloxy, C_{1-10}alkenyl oxy,

C_{3-10}alkoxy, phosphonyl, phosphate, phosphonate, phosphinate, amino,
diC_{1:6}alkylamino, monoC_{1:6}alkylamino, C_{1:6}amido, C_{1:6}amine, C_{1:6}carbamate,
C_{1:6}urea, cyano, nitro, azido, sulfhydryl, C_{1:6}alkylthio, sulfate, sulfonate, sulfamoyl,
sulfonamido, sulfonyl, C_{3:7}carboxycarbonyl, C_{3:7}carboxycarbonyl-C_{1:6}alkyl, C_{1:6}heterocyclyl,
C_{1:6}heterocyclyl-C_{1:6}alkyl, phenyl, phenyl-C_{1:6}alkyl, C_{1:6}heteroaryl, and
C_{1:6}heteroaryl-C_{1:6}alkyl, wherein heterocyclic or heteroaromatic rings, independently for
each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

X is O or NR', wherein R' is absent or is selected from the group consisting of
hydrogen, C_{1:6}alkyl, C_{1:6}haloalkyl, C_{2:10}alkenyl, C_{2:10}alkynyl, carboxyl,
C_{1:6}alkoxy-carbonyl, C_{2:10}alkenyl-oxycarbonyl, C_{2:10}alkynyl-oxycarbonyl, C_{1:6}acyl,
phosphoryl, phosphonate, phosphinate, cyano, sulfonate, sulfamoyl, sulfonyl,
C_{3:7}carboxycarbonyl, C_{3:7}carboxycarbonyl-C_{1:6}alkyl, C_{1:6}heterocyclyl, C_{1:6}heterocyclyl-C_{1:6}alkyl,
phenyl, phenyl-C_{1:6}alkyl, C_{1:6}heteroaryl, and C_{1:6}heteroaryl-C_{1:6}alkyl, wherein heterocyclic
or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms
selected from N, O, and S;

wherein any of R', R', and R', independently and independently for each occurrence,
is optionally substituted with 1-3 substituents selected from the group
consisting of C_{1:6}alkyl, C_{1:6}haloalkyl, halo, hydroxyl, carboxyl, C_{1:6}alkoxy-carbonyl,
C_{2:10}alkenyl-oxycarbonyl, C_{2:10}alkynyl-oxycarbonyl, C_{1:6}acyl, C_{1:6}acyloxy,
C_{1:6}carbonate, C_{1:6}alkoxy, phenol, phenyl, phenyl, or C_{1:6}alkyl, C_{1:6}heteroaryl, and
C_{1:6}heteroaryl-C_{1:6}alkyl; and wherein heterocyclic or heteroaromatic rings, independently
for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

m is 1-3; and
n is 0-3.

[0129] According to some embodiments of compounds of Formula I,
as valence and stability permit:

R', independently for each occurrence, is selected from the group consisting of
halo; hydroxyl, C_{1:6}alkyl, C_{1:6}haloalkyl, C_{1:6}hydroxylalkyl, or C_{1:6}acyloxy-C_{1:6}alkyl;
C_{2:10}alkenyl; C_{2:10}alkynyl; C_{1:6}alkoxy, C_{1:6}alkylthio; and C_{1:6}aryl-C_{1:6}alkyl optionally
substituted by halo, hydroxyl, C_{1:6}alkyl, C_{1:6}alkoxy, or C_{1:6}acyloxy;

R', independently for each occurrence, is selected from the group consisting of
halo; hydroxyl, C_{1:6}alkyl, C_{1:6}haloalkyl, C_{1:6}hydroxylalkyl, or C_{1:6}acyloxy-C_{1:6}alkyl;
C_{2:10}alkenyl; C_{2:10}alkynyl; C_{1:6}alkoxy, C_{1:6}alkylthio; and C_{1:6}aryl-C_{1:6}alkyl optionally
substituted by halo, hydroxyl, C_{1:6}alkyl, C_{1:6}alkoxy, or C_{1:6}acyloxy;

X is O or NR', wherein R' is absent or is selected from the group consisting of
hydrogen and C_{1:6}alkyl;
wherein any of \( R^1, R^2, \) and \( R^3 \), independently and independently for each occurrence, is optionally further substituted as noted above;

\[
m \text{ is } 1-3; \text{ and } \\
n \text{ is } 0-3.
\]

5. [0130] According to some embodiments of the compound of Formula (I), \( X = O \). In other embodiments, \( X = NR^4 \), wherein \( R^4 \) is absent. For example, in certain embodiments, the compound of Formula (I) is an imine-containing compound. For instance, in some embodiments, the compound of Formula (I) is a compound of Formula (IIa):

![Chemical Structure](image)

**Formula (IIa);**

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, wherein, as valence and stability permit, \( R^1, R^2, m, \) and \( n \) are as defined above.

[0131] In certain embodiments, one or more occurrences of \( R^1 \) is \( C_{1-6} \)alkyl, such as methyl, one or more occurrences of \( R^1 \) is \( C_{1-6} \)hydroxylalkyl, and/or one or more occurrences of \( R^1 \) is \( C_{1-6} \)alkoxy, such as methoxy.

[0132] In some embodiments, one or more occurrences of \( R^2 \) is \( C_{1-6} \)alkyl, such as methyl, one or more occurrences of \( R^2 \) is \( C_{1-6} \)hydroxylalkyl, and/or one or more occurrences of \( R^2 \) is \( C_{1-6} \)alkoxy, such as methoxy.

20. [0133] In certain embodiments, the compound of Formula (I) or Formula (IIa) is:

![Chemical Structure](image)

(Chembridge ID No. 7993700),
or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof.

[0134] According to some embodiments of the compound of Formula (I), X is NR', wherein R' is hydrogen or C_{1,alkyl}. In particular embodiments, R' is hydrogen. For example, in certain embodiments, the compound of Formula (I) is a benzylamine compound. For instance, in some embodiments, the compound of Formula (I) is a compound of Formula (IIb):

![Formula (IIb)](image)

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, wherein, as valence and stability permit, R^1, R^2, m, and n are as defined above.

[0135] In certain embodiments, one or more occurrences of R^1 is C_{1,alkyl}, such as methyl, one or more occurrences of R^1 is C_{1,alkoxy}, such as methoxy or ethoxy, and/or one or more occurrences of R^1 is C_{1,alkythio}, such as methylthio.

[0136] In some embodiments, one or more occurrences of R^1 is halo, such as fluoro, chloro, or bromo.

[0137] In certain embodiments, one or more occurrences of R^1 is hydroxyl.

[0138] In some embodiments, n is 0. In other embodiments, n is 1. For example, in some embodiments, n is 1 and R^1 is C_{1,alkyl}, such as methyl, or R^1 is C_{1,alkoxy}, such as methoxy. In yet other embodiments, n is 2. For example, in some embodiments, n is 2 and one or both occurrences of R^1 is C_{1,alkyl}, such as methyl, and/or one or both occurrences of R^1 is C_{1,alkoxy}, such as methoxy. In certain embodiments, n is 2 and one occurrence of R^1 is halo, and the other occurrence of R^1 is C_{1,alkyl}, such as methyl.

[0139] In some embodiments, m is 1. For example, in some embodiments, m is 1 and R^2 is C_{1,alkyl}, such as methyl, or R^2 is C_{1,alkoxy}, such as methoxy. In yet other embodiments, m is 2. For example, in some embodiments, m is 2 and one or both occurrences of R^2 is C_{1,alkyl}, such as
methyl, and/or one or both occurrences of R^2 is C_{1,4}alkoxy, such as methoxy or ethoxy. For instance, in some embodiments, the compound of Formula (IIb) is a compound of Formula (IIIb):

![Formula (IIIb)](image)

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

R^1 and n are as defined above; and

R^3 is selected from the group consisting of methyl and ethyl.

[0140] In some embodiments of compounds of Formula (IIb), m is 2 and one or both occurrences of R^2 is C_{1,4}alkyl, such as methyl, one or both occurrences of R^2 is C_{1,4}alkoxy, such as methoxy; and/or one or both occurrences of R^2 is C_6aryl-C_{1,4}alkyloxy, such as phenyl-C_{1,4}alkyloxy, optionally substituted by halo, hydroxyl, C_{1,4}alkyl, C_{1,4}alkoxy, or C_{1,4}acyloxy. For instance, in some embodiments, the compound of Formula (IIb) is a compound of Formula (IIIb):

![Formula (IIIb')] (image)

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

R^1, R^2, and n are as defined above; and

Ar is C_6aryl optionally substituted by halo, hydroxyl, C_{1,4}alkyl, C_{1,4}alkoxy, or C_{1,4}acyloxy.

[0141] In some embodiments, Ar is phenyl, optionally substituted by halo, hydroxyl, C_{1,4}alkyl, C_{1,4}alkoxy, or C_{1,4}acyloxy. In certain embodiments, Ar is substituted by C_{1,4}alkyl, such as methyl.

[0142] In some embodiments for compounds of Formula (IIb), n is 1 and R^1 is C_{1,4}alkyl, such as methyl, R^1 is C_{1,4}alkoxy, such as methoxy, or R^1 is C_{1,4}alkylthio, such as methylthio. In yet other

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embodiments, n is 2. For example, in some embodiments, n is 2 and one or both R¹ is halo (e.g., fluoro, chloro, or bromo), one or both R¹ is C₃₆alkyl, such as methyl, and/or one or both R¹ is C₃₆alkoxy, such as methoxy. For instance, in some embodiments, the compound of Formula (IIb) is a compound of Formula (IIb⁺):

Formula (IIb⁺);

or a comestibly or biologically acceptable salt or derivative thereof; or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

R¹, R², and m are as defined above; and

R³ is C₃₆alkyl, such as methyl.

[0143] In certain embodiments, the compound of Formula (I) is:

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Compound 12

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Compound 13

(Chembridge ID No. 5567336),

Compound 14

(Chembridge ID No. 5572799),

Compound 15

(Chembridge ID No. 5575970),

Compound 16

(Chembridge ID No. 7675788),

Compound 17

(Chembridge ID No. 7684680),
Compound 18

(Chembridge ID No. 5556341),

Compound 19

(Chembridge ID No. 5531571),

Compound 20

(Chembridge ID No. 5453910),

Compound 21

(Chembridge ID No. 7575033),
or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof.

**Edible compositions comprising pyrazole-containing compounds**

5 [0144] The substituent definitions in this section (i.e., $R^1$, $R^2$, $R^3$, $R^4$, $m$, $n$ and $o$) refer to compounds of Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa) and Formula (VIIb).

[0145] All stereochemical forms of the compounds disclosed in this and any section herein are specifically contemplated, including geometric isomers (i.e., E, Z) and optical isomers (i.e., R, S).

10 Single stereochemical isomers as well as enantiomeric and diastereomeric mixtures of the compounds disclosed in this and any section herein are also specifically contemplated.

[0146] In some embodiments, the present invention provides a composition for reducing bitter taste of a bitter tastant, wherein the composition comprises a pyrazole-containing compound. The pyrazole-containing compounds of this invention are capable of reducing or eliminating bitter taste of a bitter tastant. In some embodiments, the composition is an edible composition. In some embodiments, the pyrazole-containing compound has a molecular weight less than about 1000, 500, or 300 daltons. In certain embodiments, the pyrazole-containing compound is a compound of Formula (IV):

![Chembridge ID No. 7678665)
R₁, independently for each occurrence, is selected from the group consisting of
C₁₋₁₀alkyl, C₁₋₁₀haloalkyl, C₂₋₁₀alkenyl, C₂₋₁₀alkynyl, halo, hydroxyl, carboxyl,
C₁₋₁₀alkoxy carbonyl, C₂₋₁₀alkenoyloxy carbonyl, C₂₋₁₀alkynoyloxy carbonyl, C₁₋₁₀acyl,
C₁₋₁₀cyclylaminio, C₁₋₁₀acyloxy, C₁₋₁₀carbonyl, C₁₋₁₀alkoxy, phenolxy, phenyl-C₁₋₁₀alkyloxy,
C₁₋₁₀heteroaryloxy, C₁₋₁₀heteroaryl-C₁₋₁₀alkyloxy, C₁₋₁₀alkenyloxy, C₂₋₁₀alkenyloxy,
phosphoryl, phosphate, phosphonate, phosphinate, amino, diC₁₋₁₀alkylamino,
monoC₁₋₁₀alkylamino, C₁₋₁₀amido, C₁₋₁₀imino, C₁₋₁₀carbamate, C₁₋₁₀urea, cyano, nitro,
azido, sulfhydryl, C₁₋₁₀alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl,
C₃₋₇carbocyclic, C₃₋₇carbocyclyl-C₁₋₁₀alkyl, C₁₋₁₀heterocyclyl-C₁₋₁₀alkyl,
phenyl, phenyl-C₁₋₁₀alkyl, C₁₋₁₀heteroaryl, and C₁₋₁₀heteroaryl-C₁₋₁₀alkyl, wherein heterocyclic
or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms
selected from N, O, or S;

R² is selected from the group consisting of hydrogen, C₁₋₁₀alkyl, C₁₋₁₀haloalkyl,
C₂₋₁₀alkenyl, C₂₋₁₀alkynyl, carboxyl, C₁₋₁₀alkoxy carbonyl, C₂₋₁₀alkenyloxy carbonyl,
C₂₋₁₀alkynylcarbonyl, C₁₋₁₀acyl, phosphoryl, phosphonate, phosphinate, cyano,
sulfonate, sulfamoyl, sulfonyl, C₃₋₇carbocyclic, C₃₋₇carbocyclyl-C₁₋₁₀alkyl, C₁₋₁₀heterocyclyl,
C₁₋₁₀heterocyclic-C₁₋₁₀alkyl, phenyl, phenyl-C₁₋₁₀alkyl, C₁₋₁₀heteroaryl, and
C₁₋₁₀heteroaryl-C₁₋₁₀alkyl, wherein heterocyclic or heteroaromatic rings, independently for
each occurrence, comprise 1-4 heteroatoms selected from N, O, or S;

R³ is selected from the group consisting of hydrogen, C₁₋₁₀alkyl, C₁₋₁₀haloalkyl,
C₂₋₁₀alkenyl, C₂₋₁₀alkynyl, carboxyl, C₁₋₁₀alkoxy carbonyl, C₂₋₁₀alkenyloxy carbonyl,
C₂₋₁₀alkynylcarbonyl, C₁₋₁₀acyl, phosphoryl, phosphonate, phosphinate, cyano,
sulfonate, sulfamoyl, sulfonyl, C₃₋₇carbocyclic, C₃₋₇carbocyclyl-C₁₋₁₀alkyl, C₁₋₁₀heterocyclyl,
C₁₋₁₀heterocyclic-C₁₋₁₀alkyl, phenyl, phenyl-C₁₋₁₀alkyl, C₁₋₁₀heteroaryl, and
C₁₋₁₀heteroaryl-C₁₋₁₀alkyl, wherein heterocyclic or heteroaromatic rings, independently for
each occurrence, comprise 1-4 heteroatoms selected from N, O, or S;

R₄, independently for each occurrence, is selected from the group consisting of
C₁₋₁₀alkyl, C₁₋₁₀haloalkyl, C₂₋₁₀alkenyl, C₂₋₁₀alkynyl, halo, hydroxyl, carboxyl,
C₁₋₁₀alkoxy carbonyl, C₂₋₁₀alkenyloxy carbonyl, C₂₋₁₀alkynylcarbonyl, C₁₋₁₀acyl,
C₁₋₁₀cyclylaminio, C₁₋₁₀acyloxy, C₁₋₁₀carbonyl, C₁₋₁₀alkoxy, phenolxy, phenyl-C₁₋₁₀alkyloxy,
C₁₋₁₀heteroaryloxy, C₁₋₁₀heteroaryl-C₁₋₁₀alkyloxy, C₁₋₁₀alkenyloxy, C₁₋₁₀alkynylloxy,
phosphoryl, phosphate, phosphonate, phosphinate, amino, diC₁₋₁₀alkylamino,
monoC₁₋₁₀alkylamino, C₁₋₁₀amido, C₁₋₁₀imino, C₁₋₁₀carbamate, C₁₋₁₀urea, cyano, nitro,
arido, sulfhydryl, C₁₋₁₀alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl,
C₃₋₇carbocyclic, C₃₋₇carbocyclyl-C₁₋₁₀alkyl, C₁₋₁₀heterocyclyl-C₁₋₁₀alkyl,
phenyl, phenyl-C₁₋₁₀alkyl, C₁₋₁₀heteroaryl, and C₁₋₁₀heteroaryl-C₁₋₁₀alkyl, wherein heterocyclic
or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms
selected from N, O, or S;
wherein any of \(R_1, R_2, R_3,\) and \(R_4\), independently and independently for each occurrence, is optionally substituted with 1-3 substituents selected from the group consisting of \(\text{C}_1\text{-haloalkyl}, \text{C}_1\text{-haloalkyl}, \text{halo}, \text{hydroxyl}, \text{carboxyl}, \text{C}_1\text{-alkoxy-carbonyl}, \text{C}_2\text{-alkynyloxy-carbonyl}, \text{C}_2\text{-alkynyloxy-carbonyl}, \text{C}_1\text{-acyl}, \text{C}_1\text{-acylamino}, \text{C}_1\text{-acyloxy}, \text{C}_1\text{-carbonate, C}_1\text{-alkoxy, phenyl-}, \text{phosphoryl, phosphate, phosphonate, phosphinate, amino, diC}_1\text{-alkylamino, monoc}_{1,10}\text{-alkylamino, C}_1\text{-amido, C}_1\text{-amino, C}_1\text{-carbonate, C}_1\text{-urea, cyano, nitro, azido, sulfhydryl, C}_1\text{-alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C}_3\text{-carbocyclyl, C}_3\text{-carbocyclyl-C}_1\text{-alkyl, C}_1\text{heterocyclyl, C}_1\text{heteroaryl-C}_1\text{alkyl, phenyl, phenyl-C}_1\text{alkyl, C}_1\text{heteroaryl, and C}_1\text{heteroaryl-C}_1\text{alkyl; and wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S; n is 0-2, and m is 0-3.}

[0147] According to some embodiments of compounds of Formula IV,

as valence and stability permit:

\[R_1,\] independently for each occurrence, is selected from the group consisting of halo, \(\text{C}_1\text{-alkyl, C}_2\text{-alkenyl, and C}_2\text{-alkynyl;}
\]

\[R_2\] is selected from the group consisting of hydrogen, \(\text{C}_1\text{-alkyl, C}_2\text{-alkenyl,}
\text{C}_2\text{-alkynyl, and C}_1\text{-acyl;}
\]

\[R_3\] is selected from the group consisting of hydrogen, \(\text{C}_1\text{-alkyl, C}_2\text{-alkenyl, and}
\text{C}_2\text{-alkynyl;}
\]

\[R_4,\] independently for each occurrence, is selected from the group consisting of halo, \(\text{C}_1\text{-alkyl, C}_2\text{-alkenyl, C}_2\text{-alkynyl, C}_1\text{-alkoxy, -C(O)-O-R}_5,\) and \(-\text{C(O)-N(R)}_2,\)

\[R_5,\] independently for each occurrence, is selected from the group consisting of hydrogen, \(\text{C}_1\text{-alkyl, C}_2\text{-alkenyl, and C}_2\text{-alkynyl;}
\]

wherein any of \(R_1, R_2, R_3,\) and \(R_4,\) independently and independently for each occurrence, is optionally substituted as noted above;

\[n\] is 0-2; and
\[m\] is 0-3.

[0148] According to some embodiments, \(n\) is 0. In other embodiments \(n\) is 1 or 2, such as 1. For example, in some embodiments, one or more occurrences of \(R_2\) is halo, such as fluoro, chloro, bromo, or iodo. For instance, in certain embodiments, \(n\) is 1 and \(R_1\) is halo, such as fluoro, chloro, bromo, or iodo.

[0149] In certain embodiments, \(R_2\) is \(\text{C}_1\text{-alkyl, such as methyl or ethyl.}

[0150] According to certain embodiments, one or more occurrences of \(R_3\) is \(-\text{C(O)-O-R}_5\) or \(-\text{C(O)-N(R)}_2\). In some such embodiments, \(R_3\) is \(\text{C}_1\text{-alkyl, such as methyl or ethyl. For example, in some embodiments, m is 1 and R}_4\) is \(-\text{C(O)-O-R}_5\) or \(-\text{C(O)-N(R)}_2\). In other embodiments, \(m\) is 2 and one occurrence of \(R_2\) is \(-\text{C(O)-O-R}_3\) and the other is \(\text{C}_1\text{-alkyl, such as methyl or ethyl, or C}_1\text{-alkoxy, such as methoxy. In other embodiments, m is 3 and one occurrence

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of $R^4$ is -C(=O)-O-$R^5$ and the other two occurrences are, independently, $C_{1-6}$alkyl, such as methyl or ethyl, $C_{1-6}$alkoxy, such as methoxy, or a combination thereof. In some of the above embodiments, $R^5$ is hydrogen or $C_{1-6}$alkyl, such as methyl or ethyl.

[0151] In some embodiments, one or more occurrences of $R^4$ is $C_{1-6}$alkyl, such as methyl or ethyl.

In certain embodiments, one or more occurrences of $R^4$ is $C_{1-6}$alkoxy, such as methoxy. In some embodiments, one or more occurrences of $R^4$ is halo, such as chloro. For example, in some embodiments, $m$ is 2 and both occurrences of $R^4$ are halo, such as chloro.

[0152] In certain embodiments, the compound of Formula (IV) is a compound of Formula (Va):

![Formula (Va)](image)

Formula (Va); or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, wherein, as valence and stability permit, $R^1$, $R^2$, $R^3$, $R^4$, and $m$ are as defined above.

[0153] In certain embodiments, the compound of Formula (IV) or Formula (Va) is a compound of Formula (Vla):

![Formula (Vla)](image)

Formula (Vla); or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

$R^1$, $R^2$, $R^3$, $R^4$, are as defined above; and $o$ is 0-2.

[0154] In certain embodiments, the compound of Formula (IV), Formula (Va), or Formula (Vla) is a compound of Formula (VIIa):
or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or
diastereomer thereof, wherein, as valence and stability permit, $R^1$, $R^2$, $R^3$, $R^4$, and $o$ are as
defined above.

[0155] In certain embodiments, the compound of Formula (IV) is a compound of Formula (Vb):

\[
\begin{array}{c}
\text{Formula (Vb}); \\
\end{array}
\]

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or
diastereomer thereof, wherein, as valence and stability permit, $R^1$, $R^2$, $R^3$, $R^4$, and $m$ are as defined above.

[0156] In certain embodiments, the compound of Formula (IV) or Formula (Vb) is a compound of
Formula (Vlb):

\[
\begin{array}{c}
\text{Formula (Vlb)}; \\
\end{array}
\]

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or
diastereomer thereof,

wherein, as valence and stability permit:

$R^1$, $R^2$, $R^3$, $R^4$, are as defined above; and

$o$ is 0-2.
In certain embodiments, the compound of Formula (IV), Formula (Vb), or Formula (VIIb) is a compound of Formula (VIIb):

\[
\begin{align*}
\text{R}^1 & \quad \text{R}^2 \\
\text{N} & \quad \text{N} \\
\text{R}^3 & \quad \text{R}^4 \\
\text{C(OR)O} & \quad \text{o}
\end{align*}
\]

and cosmetically or biologically acceptable derivatives thereof, wherein, as valence and stability permit, R\textsubscript{1}, R\textsubscript{2}, R\textsubscript{3}, R\textsubscript{4}, R\textsubscript{5}, and o are as defined above.

In certain embodiments, the compound of Formula (IV) is:

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or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof.

**Edible compositions comprising hydroquinoline compounds**

5  [0159] The substituent definitions in this section (i.e., R₁, R₂, R₃, R⁴, Ar, Cy, m, n, o and p) refer to compounds of Formula (VIII), Formula (IX), and Formula (X).

[0160] All stereochemical forms of the compounds disclosed in this and any section herein are specifically contemplated, including geometric isomers (i.e., E, Z) and optical isomers (i.e., R, S). Single stereochemical isomers as well as enantiomeric and diastereomeric mixtures of the compounds disclosed in this and any section herein are also specifically contemplated.

[0161] In some embodiments, the present invention provides an edible composition for reducing bitter taste of a bitter tastant, wherein the composition comprises a hydroquinoline compound. The hydroquinoline compounds of this invention are capable of reducing or eliminating bitter taste of a bitter tastant. In some embodiments, the hydroquinoline compound has a molecular weight less than about 1000, 500, or 300 daltons. In certain embodiments, the hydroquinoline compound is a compound of Formula (VIII):
or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

5

wherein, as valence and stability permit:

R¹, independently for each occurrence, is selected from the group consisting of

C₁-₁₀alkyl, C₁-₁₀haloalkyl, C₂-₁₀alkenyl, C₂-₁₀alkynyl, halo, hydroxyl, carboxyl,
C₁-₁₀alkoxy carbonyl, C₂-₁₀alkenyl oxy carbonyl, C₂-₁₀alkynyl oxy carbonyl, C₁-₁₀acyl,
C₁-₁₀acylamino, C₁-₁₀acyloxy, C₁-₁₀carbonate, C₁-₁₀alkoxy, phenoxy,
phenyl-C₁-₁₀alkoxy, C₁-₁₀heteroaryl, C₁-₁₀heteroaryl-C₁-₁₀alkoxy, C₁-₁₀alkenyl oxy,
C₁-₁₀alkynyl oxy, phosphoryl, phosphate, phosphonate, phosphinate, amino,
diC₁-₁₀alkylamino, monoC₁-₁₀alkylamino, C₁-₁₀amido, C₁-₁₀amino, C₁-₁₀carbamate,
C₁-₁₀urea, cyano, nitro, azido, sulfdryl, C₁-₁₀alkylthio, sulfate, sulfonate, sulfamoyl,
sulfonamido, sulfonyl, C₁-₁₀carbocyclyl, C₁-₁₀carbocyclyl-C₁-₁₀alkyl, C₁-₁₀heterocyclyl,
C₁-₁₀heterocyclyl-C₁-₁₀alkyl, phenyl, phenyl-C₁-₁₀alkyl, C₁-₁₀heteroaryl, and
C₁-₁₀heteroaryl-C₁-₁₀alkyl, wherein heterocyclic or heteroaromatic rings, independently for
each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R², independently for each occurrence, is selected from the group consisting of is
selected from the group consisting of C₁-₁₀alkyl, C₁-₁₀haloalkyl, C₂-₁₀alkenyl, C₂-₁₀alkynyl,
halo, hydroxyl, carboxyl, C₁-₁₀alkoxy carbonyl, C₂-₁₀alkenyl oxy carbonyl,
C₂-₁₀alkynyl oxy carbonyl, C₁-₁₀acyl, C₁-₁₀acylamino, C₁-₁₀acyloxy, C₁-₁₀carbonate,
C₁-₁₀alkoxy, phenoxy, phenyl-C₁-₁₀alkoxy, C₁-₁₀heteroaryl,
C₁-₁₀heteroaryl-C₁-₁₀alkoxy, C₁-₁₀alkenyl oxy, C₁-₁₀alkynyl oxy, phosphoryl, phosphate,
phosphonate, phosphinate, amino, diC₁-₁₀alkylamino, monoC₁-₁₀alkylamino, C₁-₁₀amido,
C₁-₁₀amino, C₁-₁₀carbamate, C₁-₁₀urea, cyano, nitro, azido, sulfdryl, C₁-₁₀alkylthio,
sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₁-₁₀carbocyclyl,
C₁-₁₀carbocyclyl-C₁-₁₀alkyl, C₁-₁₀heterocyclyl, C₁-₁₀heterocyclyl-C₁-₁₀alkyl, phenyl,
phenyl-C₁-₁₀alkyl, C₁-₁₀heteroaryl, and C₁-₁₀heteroaryl-C₁-₁₀alkyl, wherein heterocyclic or
heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms
selected from N, O, and S;
R₁, independently for each occurrence, is selected from the group consisting of
C₁₋₁₀alkyl, C₁₋₁₀haloalkyl, C₂₋₁₀alkenyl, C₂₋₁₀alkynyl, halo, hydroxyl, carboxyl,
C₁₋₁₀alkoxy carbonyl, C₂₋₁₀alkenyloxycarbonyl, C₂₋₁₀alkynyl oxycarbonyl, C₁₋₁₀acyl,
C₁₋₁₀acylamino, C₁₋₁₀acyloxy, C₁₋₁₀carbonate, C₁₋₁₀alkoxy, phenol, phenyl,
phenyl-C₁₋₁₀alkyloxy, C₁₋₁₀heteroaryl-C₁₋₁₀alkyloxy, C₁₋₁₀alkenyloxy, C₁₋₁₀alkynyloxy,
C₁₋₁₀alkynol, C₁₋₁₀alkylamine, C₁₋₁₀amido, C₁₋₁₀dieno, C₁₋₁₀carbamate,
C₁₋₁₀urea, cyano, nitro, azido, sulfhydryl, C₁₋₁₀alkylthio, sulfate, sulfonate, sulfanyl,
sulfonamido, sulfonyl, C₃₋₉ carbocyclyl, C₃₋₉ cyclorcyloyl-C₁₋₁₀alkyl, C₁₋₁₀heterocyclyl,
C₁₋₁₀heterocyclyl-C₁₋₁₀alkyl, phenyl, phenyl-C₁₋₁₀alkyl, C₁₋₁₀heteroaryl, and
C₁₋₁₀heteroaryl-C₁₋₁₀alkyl, wherein heterocyclic or heteroaromatic rings, independently for
each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R² is selected from the group consisting of hydrogen C₁₋₁₀alkyl, C₁₋₁₀haloalkyl,
C₂₋₁₀alkenyl, C₁₋₁₀alkynyl, carboxyl, C₁₋₁₀alkoxy carbonyl, C₂₋₁₀alkenyloxycarbonyl,
C₂₋₁₀alkynyl oxycarbonyl, C₁₋₁₀acyl, phosphoryl, phosphonate, phosphinate, cyano,
sulfonate, sulfamoyl, sulfonyl, C₃₋₉ carbocyclyl, C₃₋₉ cyclorcyloyl-C₁₋₁₀alkyl,
C₁₋₁₀heterocyclyl, C₁₋₁₀heterocyclyl-C₁₋₁₀alkyl, phenyl, phenyl-C₁₋₁₀alkyl, C₁₋₁₀heteroaryl, and
C₁₋₁₀heteroaryl-C₁₋₁₀alkyl, wherein heterocyclic or heteroaromatic rings, independently for
each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

Ar is selected from the group consisting of C₆₋₁₀aryl and C₁₋₁₀heteroaryl;

Cy is n 5 to 7-membered carbocyclic or heterocyclic ring, wherein heterocyclic ring
comprises 1-4 heteroatoms selected from N, O, and S;

wherein any of R₁, R², R³, and R⁴, independently and independently for each occurrence, is optionally substituted with 1-3 substituents selected from the group
consisting of C₁₋₁₀alkyl, C₁₋₁₀haloalkyl, halo, hydroxyl, carboxyl, C₁₋₁₀alkoxy carbonyl,
C₂₋₁₀alkenyloxycarbonyl, C₂₋₁₀alkynyl oxycarbonyl, C₁₋₁₀acyl, C₁₋₁₀acylamino,
C₁₋₁₀acyloxy, C₁₋₁₀carbonate, C₁₋₁₀alkoxy, phenol, phenyl, phosphoryl, phosphonate,
phosphinate, amino, diC₁₋₁₀alkylamino, monoC₁₋₁₀alkylamino, C₁₋₁₀amido,
C₁₋₁₀dieno, C₁₋₁₀carbamate, C₁₋₁₀urea, cyano, nitro, azido, sulfhydryl, C₁₋₁₀alkylthio,
sulfate, sulfonate, sulfamoyl, sulfonyl, C₃₋₉ carbocyclyl, C₃₋₉ cyclorcyloyl-C₁₋₁₀alkyl,
C₁₋₁₀heterocyclyl, C₁₋₁₀heterocyclyl-C₁₋₁₀alkyl, phenyl, phenyl-C₁₋₁₀alkyl, C₁₋₁₀heteroaryl, and
C₁₋₁₀heteroaryl-C₁₋₁₀alkyl; and wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms
selected from N, O, and S;

m is 1-3;

n is 0-3; and

0 is 0-3.

[0162] According to some embodiments of compounds of Formula VIII,
as valence and stability permit:
R¹, independently for each occurrence, is selected from the group consisting of halo, hydroxyl, C₁₋₅alkyl, C₁₋₅haloalkyl, C₂₋₅alkenyl, C₂₋₅alkynyl, C₁₋₅alkoxy, and C₁₋₅acyloxy;

R², independently for each occurrence, is C₁₋₅alkyl;

R³, independently for each occurrence, is selected from the group consisting of halo, C₁₋₅alkyl, C₁₋₅alkenyl, C₂₋₅alkynyl, C(O)-O-R⁴, and C(O)-N(R⁵)₂;

R⁴, independently for each occurrence, is selected from the group consisting of hydrogen, C₁₋₅alkyl, C₁₋₅alkenyl, and C₂₋₅alkynyl;

R⁵ is selected from the group consisting of hydrogen, C₁₋₅alkyl, C₁₋₅alkenyl, and C₂₋₅alkynyl;

Ar is selected from the group consisting of C₆₋₁₀aryl and C₅₋₉heteroaryl;

Cy is a 5 to 7-membered carbocyclic or heterocyclic ring, optionally including one or two carbon-carbon or carbon-nitrogen double bonds in the ring;

wherein any of R¹, R², R³, and R⁵, independently and independently for each occurrence, is optionally substituted as noted above;

m is 1-3;

n is 0-3; and

o is 0-3.

[0163] In some embodiments, n is 0. In other embodiments, n is 1. For example, in certain embodiments, n is 1 and R¹ is halo (such as fluoro, chloro, or bromo) or C₁₋₅acyloxy (such as acetyloxy). In other embodiments, n is 2. For example, in some embodiments, n is 2 and both occurrences of R¹ is halo, such as chloro.

[0164] In some embodiments, m is 1. For example, in certain embodiments, m is 1 and R³ is C(O)-O-R⁴, such as C(O)-OH, C(O)-OMe, or C(O)-OEt. In other embodiments, m is 2. For example, in some embodiments, m is 2 and one occurrence of R³ is C(O)-O-R⁴ and the other occurrence is halo, such as bromo.

[0165] In certain embodiments, o is 0. In other embodiments, o is 1-3.

[0166] In certain embodiments, R² is hydrogen. In other embodiments, R² is C₁₋₅alkyl, such as methyl.

[0167] In certain embodiments, Ar is C₆₋₁₀aryl, such as phenyl. In other embodiments, Ar is C₅₋₉heteroaryl.

[0168] In certain embodiments, Cy is a 5 to 7-membered carbocyclic ring, such as a 5-membered carbocyclic ring, such as a cyclopentyl or cyclopentenyl ring. In some embodiments, Cy includes one carbon-carbon double bond in the ring, such as in a cyclopentenyl ring. For instance, according to one embodiment, the compound of Formula (VIII) is a compound of Formula (IX):
or a comestibly or biologically acceptable salt or derivative thereof, or an enantioomer or diastereomer thereof, wherein, as valence and stability permit, $R^1$, $R^2$, $R^3$, $R^4$, $m$, $n$, and $o$ are as defined above.

[0169] In some embodiments, the compound of Formula (VIII) or Formula (IX) is a compound of Formula (X):

or a comestibly or biologically acceptable salt or derivative thereof, or an enantioomer or diastereomer thereof,

wherein, as valence and stability permit:

$R^1$, $R^2$, $R^3$, $R^4$, and $n$ are as defined above; and

$p$ is 0-2.

[0170] In some embodiments, $p$ is 0. In other embodiments, $p$ is 1 and $R^3$ is halo, such as bromo.
In certain embodiments, the compound of Formula (VIII) is:

**Compound 37**

![Chembridge ID No. 5846684](image)

**Compound 38**

![Chembridge ID No. 6851241](image)

**Compound 39**

![Chembridge ID No. 6527982](image)

**Compound 40**

![Asinex ID BAS02001668](image)
or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or
diastereomer thereof.

**Edible compositions comprising quinoline compounds**

5  [0172] The substituent definitions in this section (i.e., $R^1$, $R^2$, $R^3$, $R^6$, Het, Ar, m and n) refer to compounds of Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), and Formula (XIIIb).

[0173] All stereochemical forms of the compounds disclosed in this and any section herein are specifically contemplated, including geometric isomers (i.e., E, Z) and optical isomers (i.e., R, S).
Single stereochemical isomers as well as enantiomeric and diastereomeric mixtures of the compounds disclosed in this and any section herein are also specifically contemplated.

In some embodiments, the present invention provides a composition for reducing bitter taste of a bitter tastant, wherein the composition comprises a quinoline compound. The quinoline compounds of this invention are capable of reducing or eliminating bitter taste of a bitter tastant. In some embodiments, the composition is an edible composition. In some embodiments, the quinoline compound has a molecular weight less than about 1000, 500, or 300 daltons. In certain embodiments, the quinoline compound is a compound of Formula (XI):

\[
\begin{align*}
&\text{Formula (XI):} \\
&\text{wherein, as valence and stability permit:} \\
&R^1, \text{ independently for each occurrence, is selected from the group consisting of} \\
&C_{1-10} \text{alkyl, } C_{1-10} \text{haloalkyl, } C_{2-10} \text{alkenyl, } C_{2-10} \text{alkynyl, }} \text{halo, hydroxyl, carboxyl,} \\
&C_{1-10} \text{alkoxy carbonyl, } C_{2-10} \text{alkenyl oxo carbonyl, } C_{2-10} \text{alkyl oxo carbonyl, } C_{1-10} \text{acyl,} \\
&C_{1-10} \text{amino, } C_{1-10} \text{amido, } C_{1-10} \text{amino, } C_{1-10} \text{carbamate, } C_{1-10} \text{urea, cyano, nitro,} \\
&\text{azido, sulfonyl, } C_{1-10} \text{alkylthio, sulfone, sulfonate, sulfamoyl, sulfonamido, sulfonyl,} \\
&C_{2-5} \text{carboxylic, } C_{2-5} \text{carboxylic, } C_{2-5} \text{carboxylic, } C_{1-10} \text{alkyl,} \\
&C_{1-10} \text{heterocyclic, and } C_{1-10} \text{heteroaryl, and } C_{1-10} \text{heteroaryl,} \\
&\text{phenyl, phenyl-C}_{1-10} \text{alkyl, } C_{1-10} \text{heteroaryl, and } C_{1-10} \text{heteroaryl,} \\
&\text{wherein heterocyclic, or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms} \\
&\text{selected from N, O, and S;}
\end{align*}
\]

\[
&R^2 \text{ is selected from the group consisting of hydrogen, } C_{1-10} \text{alkyl, } C_{1-10} \text{haloalkyl,} \\
&C_{1-10} \text{alkenyl, } C_{2-10} \text{alkynyl, } C_{2-10} \text{alkenyl, halo, hydroxyl, carboxyl, } C_{1-10} \text{alkoxy carbonyl,} \\
&C_{2-10} \text{alkenyl oxo carbonyl, } C_{2-10} \text{alkenyl oxo carbonyl, } C_{1-10} \text{acyl, } C_{1-10} \text{ami}, \\
&C_{1-10} \text{amido, } C_{1-10} \text{amido, } C_{1-10} \text{carbamate, } C_{1-10} \text{urea, cyano, nitro, azido, sulfonyl,} \\
&C_{1-10} \text{alkylthio, sulfone, sulfonate, sulfamoyl, sulfonamido, sulfonyl,} \\
&C_{2-5} \text{carboxylic, } C_{2-5} \text{carboxylic, } C_{2-5} \text{carboxylic, } C_{1-10} \text{alkyl,} \\
&C_{1-10} \text{heterocyclic, and } C_{1-10} \text{heteroaryl, and } C_{1-10} \text{heteroaryl,} \\
&\text{phenyl, phenyl-C}_{1-10} \text{alkyl, } C_{1-10} \text{heteroaryl, and } C_{1-10} \text{heteroaryl,} \\
&\text{wherein heterocyclic, or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms} \\
&\text{selected from N, O, and S;}
\]

SUBSTITUTE SHEET (RULE 26)
C₃₋₅ carbo cyclic-C₆₅ alkyl, C₆₋₈ heterocyclic, C₉₋₁₀ heterocyclic-C₅₋₆ alkyl, phenyl, phenyl-C₅₋₆ alkyl, C₂₋₅ heteroaryl, and C₆₋₉ heteroaryl-C₆₋₅ alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S,

5 R¹ is selected from the group consisting of hydrogen, C₁₋₅ alkyl, Het-C₆₋₁₀ alkyl, C₁₋₅ haloalkyl, C₂₋₅ alkenyl, C₂₋₅ alkynyl, halo, hydroxy, carboxyl, C₁₋₁₀ alkoxy carbonyl, C₂₋₁₀ alkenyloxy carbonyl, C₂₋₁₀ alkynyl oxy carbonyl, C₁₋₅ acyl, C₁₋ₕ acylamino, C₁₋₆ acyloxy, C₁₋₆ carbonate, C₁₋₁₀ alkoxy, phenol, phenyl-C₁₋₅ alkoxy, C₁₋₅ heteroaryl, C₆₋₉ alkoxy, C₆₋₉ alkenyloxy, C₆₋₉ alkynyl oxy, C₆₋₉ hydroxy, C₂₋₁₀ alkylthio, C₆₋₉ amino, diC₁₋₁₀ alkylamino, monoC₁₋₁₀ alkylamino, C₁₋₅ amido, C₁₋₅ imino, C₁₋₅ carbamate, C₁₋₅ urea, cyano, nitro, azido, sulfhydryl, C₂₋₅ alkythio, sulfite, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₂₋₅ carboxycylyl, C₆₋₉ carbo cyclic-C₆₋₅ alkyl, C₆₋₉ heterocyclic-C₆₋₅ alkyl, and C₁₋₅ heteroaryl-C₁₋₅ alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S.

10 R² is selected from the group consisting of hydrogen, C₁₋₅ alkyl, Het-C₆₋₁₀ alkyl, C₁₋₅ haloalkyl, C₂₋₅ alkenyl, C₂₋₅ alkynyl, halo, hydroxy, carboxyl, C₁₋₁₀ alkoxy carbonyl, C₂₋₁₀ alkenyloxy carbonyl, C₂₋₁₀ alkynyl oxy carbonyl, C₁₋₅ acyl, C₁₋₅ acylamino, C₁₋₆ acyloxy, C₁₋₆ carbonate, C₁₋₁₀ alkoxy, phenol, phenyl-C₁₋₅ alkoxy, C₁₋₅ heteroaryl, C₆₋₉ alkoxy, C₆₋₉ alkenyloxy, C₆₋₉ alkynyl oxy, C₆₋₉ hydroxy, C₂₋₁₀ alkylthio, sulfite, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₂₋₅ carboxycylyl, C₆₋₉ carbo cyclic-C₆₋₅ alkyl, C₆₋₉ heterocyclic-C₆₋₅ alkyl, phenyl, phenyl-C₁₋₅ alkyl, C₁₋₅ heteroaryl, and C₁₋₅ heteroaryl-C₁₋₅ alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S.

15 or R³ and R⁴ together with the atoms to which they are attached form a 5 to 6-membered aryl or heteroaryl ring optionally substituted by 1 to 4 groups selected from the group consisting of Het, C₁₋₅ alkyl, C₁₋₅ haloalkyl, halo, hydroxy, carboxyl, C₁₋₁₀ alkoxy carbonyl, C₂₋₁₀ alkenyloxy carbonyl, C₂₋₁₀ alkynyl oxy carbonyl, C₁₋₅ acyl, C₁₋₅ acylamino, C₁₋₅ acyloxy, C₁₋₁₀ carbonate, C₁₋₅ alkoxy, phenol, phenyl, phosphate, phosphonate, phosphinate, amino, diC₁₋₁₀ alkylamino, monoC₁₋₁₀ alkylamino, C₁₋₅ amido, C₁₋₅ imino, C₁₋₅ carbamate, C₁₋₅ urea, cyano, nitro, azido, sulfhydryl, C₁₋₅ alkythio, sulfite, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₂₋₅ carboxycylyl, C₆₋₉ carbo cyclic-C₆₋₅ alkyl, C₆₋₉ heterocyclic-C₆₋₅ alkyl, phenyl, phenyl-C₁₋₅ alkyl, C₁₋₅ heteroaryl, and C₁₋₅ heteroaryl-C₁₋₅ alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S.
heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

Het is a C_{1-4} heterocyclyl including 1-4 heteroatoms in the ring selected from oxygen, sulfur, and nitrogen;

wherein any of R^1, R^2, R^3, R^4, and Het, independently and independently for each occurrence, is optionally substituted with 1-3 substituents selected from the group consisting of C_{1-10}alkyl, C_{1-10}haloalkyl, halo, hydroxyl, carbonyl, C_{1-10}alkoxy carbonyl, C_{2-10}alkenyl oxycarbonyl, C_{2-10}alkynyl oxycarbonyl, C_{1-10}acyl, C_{1-10}acylamino, C_{1-10}acyloxy, C_{1-10}carbonate, C_{1-10}alkoxy, phenol, phosphoryl, phosphate, phosphonate, phosphinate, amino, diC_{1-10}alkylamino, monoC_{1-10}alkylamino, C_{1-10}amido, C_{1-10}aminocarbonyl, C_{1-10}urea, cyano, nitro, azido, sulfhydryl, C_{1-10}alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfanyl, C_{3-10}carboxyethyl, C_{3-10}carboxyethyl-C_{1-10}alkyl, C_{1-10}heterocyclyl, C_{1-10}heterocyclyl-C_{1-10}alkyl, C_{2-10}arylyl, C_{6-10}arylyl-C_{1-10}alkyl, C_{1-10}heteroaryl, and C_{1-10}heteroaryl-C_{1-10}alkyl; and wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S; and

n is 0-4.

[0175] According to some embodiments of compounds of Formula XI, as valence and stability permit:

R^1, independently for each occurrence, is selected from the group consisting of halo, hydroxyl, C_{1-10}alkyl, C_{1-10}haloalkyl, C_{2-10}alkenyl, C_{2-10}alkynyl, and C_{1-10}alkoxy;

R^2 is selected from the group consisting of hydrogen, halo, hydroxyl, C_{1-10}alkyl, C_{1-10}haloalkyl, Het-C_{1-10}alkyl, C_{2-10}alkenyl, C_{2-10}alkynyl, and C_{1-10}alkoxy;

R^3 is selected from the group consisting of hydrogen, halo, hydroxyl, C_{1-10}alkyl, C_{1-10}haloalkyl, Het-C_{1-10}alkyl, C_{2-10}alkenyl, C_{2-10}alkynyl, and C_{1-10}alkoxy;

R^4 is selected from the group consisting of hydrogen, halo, hydroxyl, C_{1-10}alkyl, C_{1-10}haloalkyl, Het-C_{1-10}alkyl, C_{2-10}alkenyl, C_{2-10}alkynyl, and C_{1-10}alkoxy;

or R^3 and R^4 together with the atoms to which they are attached form a 5 to 6-membered aryl ring optionally substituted by 1 to 4 groups selected from the group consisting of halo, hydroxyl, C_{1-10}alkyl, C_{1-10}haloalkyl, C_{2-10}alkenyl, C_{2-10}alkynyl, C_{1-10}alkoxy, and Het;

Het is a C_{2-4} heterocyclyl including 1-3 heteroatoms in the ring selected from oxygen, sulfur, and nitrogen and is optionally substituted with one or more groups selected from the group consisting of halo, hydroxyl, C_{1-10}alkyl, C_{1-10}haloalkyl, C_{2-10}alkenyl, C_{2-10}alkynyl, C_{1-10}alkoxy, and C_{2-10}arylyl optionally substituted by C_{1-10}alkyl;

wherein any of R^1, R^2, R^3, R^4, and Het, independently and independently for each occurrence, is optionally further substituted as noted above; and

n is 0-4.

[0176] According to some embodiments, one, two, or all of R^2, R^3, or R^4 is not hydrogen.
According to some embodiments of the compound of Formula (XI), \( R^3 \) and \( R^4 \) together with the atoms to which they are attached form a 5 to 6-membered aryl ring, such as a benzo ring, optionally substituted as described above. For example, in certain embodiments, the compound of Formula (XI) is a compound of Formula (XIIa):

﻿

\[
\begin{align*}
& \text{Formula (XIIa);} \\
& \text{or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,} \\
& \text{wherein, as valence and stability permit:} \\
& R^1, R^2, \text{Het}, \text{and } n \text{ are as defined above;} \\
& R^1, \text{independently for each occurrence, is selected from the group consisting of halo, hydroxyl, } C_{1,6}\text{alkyl, } C_{1,6}\text{haloalkyl, } C_{2,6}\text{alkenyl, } C_{2,6}\text{alkynyl, and } C_{1,6}\text{alkoxy;} \text{and } m \text{ is 0-3.}
\end{align*}
\]

According to some embodiments, Het is a nitrogen-containing heterocycle optionally including additional heteroatoms selected from oxygen, sulfur, and nitrogen and optionally substituted as described above. For example, in certain embodiments, the compound of Formula (XIIa) is a compound of Formula (XIIIa):

\[
\begin{align*}
& \text{Formula (XIIIa);} \\
& \text{or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, wherein, as valence and stability permit, } R^1, R^2, R^3, \text{Het, } n, \text{ and } m \text{ are as defined above.}
\end{align*}
\]

In certain embodiments, one or more occurrences of \( R^4 \) is halo, such as fluoro. For example, in some embodiments, for compounds of Formula (XIIa) and Formula (XIIIa), \( m \) is 3 and \( R^3 \) is fluoro for each occurrence.
In some embodiments, Het is a nitrogen-containing heterocycle, such as aziridine, azetidine, diazetidine, pyrrolidine, pyrroline, imidazolidine, pyrazolidine, imidazoline, oxazolidine, isoxazolidines, oxazoline, piperidine, piperazine, morpholine, oxazine, thiazine, azepane, azepine, or diazepine optionally substituted as described above. In particular embodiments, Het is pyrrolidine, piperazine, or morpholine optionally substituted as described above. In certain embodiments, Het is substituted with one or more C$_{1-4}$alkyl, such as methyl.

According to certain embodiments, $n$ is 0 or $n$ is 1 and $R^1$ is C$_{1-4}$alkyl, such as methyl; $R^2$ is C$_{1-4}$alkyl, such as methyl; $m$ is 3 and $R^3$ is fluoro for each occurrence; and Het is pyrrolidine, piperazine, or morpholine optionally substituted with one or more C$_{1-4}$alkyl, such as methyl. For instance, in certain embodiments, the compound of Formula (XI), Formula (XIIa), or Formula (XIIIa) is:

![Chembridge ID No. 7728336](image)

![Chembridge ID No. 7733323](image)

![Chembridge ID No. 7726077](image)

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof.

According to some embodiments of the compound of Formula (XI), $R^3$ and $R^4$, independently, are selected from the group consisting of hydrogen, halo, hydroxyl, C$_{1-4}$alkyl,
C₅₋₁₋₆ haloalkyl, Het-C₅₋₁₋₆ alkyl, C₂₋₆ alkenyl, C₂₋₆ alkynyl, and C₁₋₆ alkoxy. For example, in certain embodiments, R¹ is C₁₋₆ alkyl, such as methyl. In some embodiments, R² is Het-C₁₋₆ alkyl, such as Het-CH₂-. In further embodiments, R³ is C₁₋₆ alkyl, such as methyl, and R⁴ is Het-C₁₋₆ alkyl, such as Het-CH₂-. For example, in certain embodiments, the compound of Formula (XI) is a compound of

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, wherein, as valence and stability permit, R¹, R², R³, Het, and n are as defined above.

According to some embodiments, Het is a nitrogen-containing heterocycle optionally including additional heteroatoms selected from oxygen, sulfur, and nitrogen and optionally substituted as described above. In certain embodiments, Het is substituted by one or more C₆₋₁₅ aryl, such as phenyl or naphthyl, optionally substituted by C₁₋₆ alkyl. For example, in certain embodiments, the compound of Formula (XII) is a compound of Formula (XIII):

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:
R¹, R², R³, Het, and n are as defined above; and
Ar is C₆₋₁₅ aryl, such as phenyl or naphthyl, optionally substituted by C₁₋₆ alkyl.

In some embodiments, Het is a nitrogen-containing heterocycle, such as aziridine, azetidine, diazetidine, pyrrolidine, pyrroline, imidazolidine, pyrazolidine, imidazoline, oxazolidine, isoxazolidines, oxazoline, piperidine, piperazine, morpholine, oxazine, thiazine, azepane, azepine, or diazepine optionally substituted as described above. In specific embodiments, Het is pyrrolidine, piperazine, or morpholine, particularly piperazine, optionally substituted as described above.
[0185] According to certain embodiments, one or more occurrences of R\textsuperscript{1} is C\textsubscript{1-4}alkyl, such as methyl; R\textsuperscript{2} is hydroxyl; R\textsuperscript{3} is C\textsubscript{1-4}alkyl, such as methyl; Het is piperazine; and Ar is phenyl. For instance, in some embodiments, the compound of Formula (XI), Formula (XIIIb), or Formula (XIIIb) is:

<table>
<thead>
<tr>
<th>Compound 47</th>
</tr>
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<tbody>
<tr>
<td><img src="image1.png" alt="Chembridge ID No. 9149274" /></td>
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</table>

[0186] In some embodiments of the above compounds, n is 0, and there are no occurrences of R\textsuperscript{1}. In other occurrences, n is not zero. For example, in some embodiments, one or more occurrences of R\textsuperscript{1} are C\textsubscript{1-4}alkyl, such as methyl. For instance, in certain embodiments, n is 1 and R\textsuperscript{1} is C\textsubscript{1-4}alkyl, such as methyl, optionally in a position para to the nitrogen atom. In another embodiment, n is 2 and R\textsuperscript{1} is C\textsubscript{1-4}alkyl, such as methyl, for both occurrences, with the occurrences of R\textsuperscript{1} optionally in a 1,4-relationship.

[0187] In some embodiments of the above compounds, R\textsuperscript{2} is hydroxyl. In other embodiments, R\textsuperscript{2} is hydroxyl. In other embodiments, R\textsuperscript{2} is C\textsubscript{1-4}alkyl, such as methyl.

15 **Edible compositions comprising N-phenylalkylamide compounds**

[0188] The substituent definitions in this section (i.e., R\textsuperscript{1}, R\textsuperscript{2}, R\textsuperscript{3}, Ar, X, m, n and p) refer to compounds of Formula (XIV), Formula (XV), Formula (XVb), and Formula (XVc).

[0189] All stereochemical forms of the compounds disclosed in this and any section herein are specifically contemplated, including geometric isomers (i.e., E, Z) and optical isomers (i.e., R, S).

[0190] In some embodiments, the present invention provides a composition for reducing bitter taste of a bitter tastant, wherein the composition comprises a N-phenylalkylamide compound. The
N-phenylalkylamide compounds of this invention are capable of reducing or eliminating bitter taste of a bitter tastant. In some embodiments, the composition is an edible composition. In some embodiments, the N-phenylalkylamide compound has a molecular weight less than about 1000, 500, or 300 daltons. In certain embodiments, the N-phenylalkylamide compound is a compound of Formula (XIV):

![Chemical Structure Image]

Formulas (XIV); or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

10 wherein, as valence and stability permit:

R<sup>1</sup>, independently for each occurrence, is selected from the group consisting of C<sub>1</sub>-C<sub>10</sub>alkyl, C<sub>1</sub>-C<sub>10</sub>haloalkyl, C<sub>2</sub>-C<sub>10</sub>alkenyl, C<sub>2</sub>-C<sub>10</sub>alkynyl, halo, hydroxyl, carboxyl, C<sub>2</sub>-C<sub>10</sub>alkoxyoxycarbonyl, C<sub>2</sub>-C<sub>10</sub>alkenylxoyoxycarbonyl, C<sub>2</sub>-C<sub>10</sub>alknyloxyxycarbonyl, C<sub>1</sub>-C<sub>10</sub>acyl, C<sub>1</sub>-C<sub>10</sub>aclylamino, C<sub>1</sub>-C<sub>10</sub>acyloxy, C<sub>1</sub>-C<sub>10</sub>carbonate, C<sub>1</sub>-C<sub>10</sub>alkoxy, phenolxy, phenyl-C<sub>1</sub>-C<sub>10</sub>alkyloxy, C<sub>1</sub>-C<sub>10</sub>heteroarlyoxy, C<sub>1</sub>-C<sub>10</sub>heteroaryl-C<sub>1</sub>-C<sub>10</sub>alkyloxy, C<sub>1</sub>-C<sub>10</sub>alkenylxoy, C<sub>1</sub>-C<sub>10</sub>alknyloxy, C<sub>2</sub>-C<sub>10</sub>alknyloxy, phosphoryl, phosphate, phosphonate, phosphinate, amino, diC<sub>1</sub>-C<sub>10</sub>alkylamino, monoC<sub>1</sub>-C<sub>10</sub>alkylamin, C<sub>1</sub>-C<sub>10</sub>amino, C<sub>1</sub>-C<sub>10</sub>carbamate, C<sub>1</sub>-C<sub>10</sub>urea, cyano, nitro, azido, sulhydryl, C<sub>1</sub>-C<sub>10</sub>alkylthio, sulfate, sulfonate, sulfonamido, sulfonoyl, C<sub>1</sub>-C<sub>10</sub>carbocyclyl, C<sub>1</sub>-C<sub>10</sub>carbocyclyl-C<sub>1</sub>-C<sub>10</sub>alkyl, C<sub>1</sub>-C<sub>10</sub>heterocyclyl, C<sub>1</sub>-C<sub>10</sub>heterocyclyl-C<sub>1</sub>-C<sub>10</sub>alkyl, phenyl, phenyl-C<sub>1</sub>-C<sub>10</sub>alkyl, C<sub>1</sub>-C<sub>10</sub>heteroaryl, and C<sub>1</sub>-C<sub>10</sub>heteroaryl-C<sub>1</sub>-C<sub>10</sub>alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;

R<sup>2</sup> is selected from the group consisting of C<sub>1</sub>-C<sub>10</sub>alkyl, C<sub>1</sub>-C<sub>10</sub>haloalkyl, C<sub>2</sub>-C<sub>10</sub>alkenyl, C<sub>2</sub>-C<sub>10</sub>alkynyl, hydroxyl, C<sub>2</sub>-C<sub>10</sub>alkoxyxoxycarbonyl, C<sub>2</sub>-C<sub>10</sub>alknyloxyxycarbonyl, C<sub>2</sub>-C<sub>10</sub>alkenylxoyoxycarbonyl, C<sub>2</sub>-C<sub>10</sub>alknyloxyxycarbonyl, C<sub>1</sub>-C<sub>10</sub>acyl, C<sub>1</sub>-C<sub>10</sub>aclylamino, C<sub>1</sub>-C<sub>10</sub>acyloxy, C<sub>1</sub>-C<sub>10</sub>carbonate, C<sub>1</sub>-C<sub>10</sub>alkoxy, phenolxy, phenyl-C<sub>1</sub>-C<sub>10</sub>alkyloxy, C<sub>1</sub>-C<sub>10</sub>heteroarlyoxy, C<sub>1</sub>-C<sub>10</sub>heteroaryl-C<sub>1</sub>-C<sub>10</sub>alkyloxy, C<sub>1</sub>-C<sub>10</sub>alkenylxoy, C<sub>1</sub>-C<sub>10</sub>alknyloxy, C<sub>2</sub>-C<sub>10</sub>alknyloxy, phosphoryl, phosphate, phosphonate, phosphinate, amino, diC<sub>1</sub>-C<sub>10</sub>alkylamino, monoC<sub>1</sub>-C<sub>10</sub>alkylamin, C<sub>1</sub>-C<sub>10</sub>amino, C<sub>1</sub>-C<sub>10</sub>carbamate, C<sub>1</sub>-C<sub>10</sub>urea, cyano, nitro, azido, sulhydryl, C<sub>1</sub>-C<sub>10</sub>alkylthio, sulfate, sulfonate, sulfonamido, sulfonoyl, C<sub>1</sub>-C<sub>10</sub>carbocyclyl, C<sub>1</sub>-C<sub>10</sub>carbocyclyl-C<sub>1</sub>-C<sub>10</sub>alkyl, C<sub>1</sub>-C<sub>10</sub>heterocyclyl, C<sub>1</sub>-C<sub>10</sub>heterocyclyl-C<sub>1</sub>-C<sub>10</sub>alkyl, phenyl, phenyl-C<sub>1</sub>-C<sub>10</sub>alkyl, C<sub>1</sub>-C<sub>10</sub>heteroaryl, and C<sub>1</sub>-C<sub>10</sub>heteroaryl-C<sub>1</sub>-C<sub>10</sub>alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;
R¹ is selected from the group consisting of hydrogen, C₁₋₅ alkyl, C₁₋₅ haloalkyl, C₂₋₁₀alkenyl, C₂₋₁₀alkynyl, carboxyl, C₁₋₁₀alkoxy carbonyl, C₂₋₁₀alkoxy carbonyl, C₂₋₁₀alkynyl oxycarbonyl, C₁₋₁₀acyl, phosphoryl, phosphonate, phosphinate, cyano, sulfonate, sulfamoyl, sulfonyl, C₂₋₅carbocyclyl, C₂₋₅heterocyclyl, C₁₋₅alkyl, C₁₋₅heteroaryl, and C₁₋₅heteroaryl-C₁₋₅alkyl, wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S; wherein any of R¹, R², and R³, independently and independently for each occurrence, is optionally substituted with 1-3 substituents selected from the group consisting of C₁₋₅ alkyl, C₁₋₅ haloalkyl, halo, hydroxyl, carboxyl, C₁₋₅ alkoxy carbonyl, C₂₋₁₀alkenyloxycarbonyl, C₂₋₁₀alkynyl oxycarbonyl, C₁₋₁₀acyl, C₁₋₁₀acylamino, C₁₋₁₀acyloxy, C₁₋₁₀carbonate, C₁₋₁₀alkoxy, C₁₋₁₀aryl amine, phosphonate, phosphinate, amino, diC₁₋₁₀alkyl amine, monoC₁₋₁₀alkyl amine, C₁₋₁₀amino, C₁₋₁₀carbamate, C₁₋₁₀urea, cyano, nitro, azido, sulphydryl, C₁₋₁₀alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₂₋₅ carbocyclyl, C₂₋₅heterocyclyl-C₁₋₅alkyl, C₁₋₅heterocyclyl, C₁₋₅heterocyclyl-C₁₋₅alkyl, phenyl, phenyl-C₁₋₅alkyl, C₁₋₅heteroaryl, and C₁₋₅heteroaryl-C₁₋₅alkyl; and wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S; and

n is 0-3.

According to some embodiments of compounds of Formula (XIV), as valence and stability permit:
R¹, independently for each occurrence, is selected from the group consisting of halo, hydroxyl, C₁₋₅ alkyl, C₁₋₅ haloalkyl, C₂₋₁₀alkenyl, C₂₋₁₀alkynyl, C₁₋₅ alkoxy, and C₁₋₅ acyloxy;
R² is selected from the group consisting of C₁₋₅ alkyl, C₁₋₅ alkoxy-substituted C₁₋₅ alkyl, C₁₋₁₀ aryl oxyl substituted C₁₋₅ alkyl, C₂₋₁₀alkenyl, C₂₋₁₀alkynyl, C₁₋₁₀aryl -C₁₋₅ alkyl, and -((CH₂)₃X)₃Ar, wherein aryl groups of R² are optionally substituted by one or more halo, hydroxyl, C₁₋₅ alkyl, C₁₋₅ haloalkyl, C₁₋₅ alkoxy, or C₁₋₅ acyloxy;
R³ is selected from the group consisting of hydrogen, C₁₋₅ alkyl, C₂₋₁₀alkenyl, and C₂₋₁₀alkynyl;
X is selected from the group consisting of O, NH, and CH₂;
Ar is selected from the group consisting of C₆₋₁₀ aryl, C₄₋₅heterocyclyl, C₅₋₁₀ carbocyclyl, and C₂₋₁₀heterocyclyl, including fused bicyclic groups, wherein Ar is optionally substituted by one or more halo, hydroxyl, C₁₋₅ alkyl, C₁₋₅ haloalkyl, C₁₋₅ alkoxy, or C₁₋₅ acyloxy;

wherein any of R¹, R², and R³, independently and independently for each occurrence, is optionally further substituted as noted above;
m is 1-3.
n is 0-3; and
p is 0 or 1.

[0192] In certain embodiments, n is 0. In other embodiments, n is 1-3. For example, in some embodiments, n is 1-3, such as 2, and one or more occurrences of R¹ is C₁₋₄alkoxy, such as methoxy. In some embodiments, n is 1-3, such as 2, and one or more occurrences of R¹ is C₁₋₄alkoxy, such as methoxy, and one or more occurrences of R¹ is halo, such as chloro. In some embodiments, n is 1-3, such as 2, and one or more occurrences of R¹ is C₁₋₄alkyl, such as methyl or ethyl, and one or more occurrences of R¹ is halo, such as chloro. In further embodiments, n is 2-3, such as 2, and two or more occurrences of R¹ is C₁₋₄alkyl, such as methyl or ethyl.

[0193] In some embodiments, R² is C₁₋₄alkyl, such as methyl, ethyl, or propyl. In certain embodiments, R² is C₆₋₁₀aryloxy-substituted C₁₋₄alkyl, such as 2-aryloxyethyl (e.g., 2-phenloxyethyl), optionally substituted as described above. In some embodiments, R² is C₆₋₁₀aryl-C₁₋₄alkyl. For example, in some embodiments, R² is phenyl-C₁₋₄alkyl, such as 2-arylthethyl (e.g., dihydorcinnamyl) or 3-arylpropyl (e.g., 3-phenylpropyl), optionally substituted as described above.

[0194] According to certain embodiments, R³ is hydrogen. In other embodiments, R³ is C₁₋₄alkyl, such as methyl.

[0195] As noted above, in certain embodiments, R² is C₁₋₄alkyl. For instance, in some embodiments, the compound of Formula (XIV) is a compound of Formula (XVa):

```
          R¹
         /\    \
        R²  N   O
         /\    
        R¹  
```

Formula (XVa);
or a pharmaceutically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof,

wherein, as valence and stability permit:

R¹ and R² are as defined above; and
R² is C₁₋₄alkyl, such as methyl or ethyl.

[0196] According to certain embodiments of the compound of Formula (XIV), R² is -((CH₂)ₙX)ₚ-Ar. In certain embodiments, p is 0. In other embodiments, p is 1. For example, in some embodiments, p is 1 and X is O. In some embodiments, p is 1 and X is CH₂. In some embodiments, Ar is C₆₋₁₀aryl, such as phenyl, optionally substituted as described above. In other embodiments, Ar is C₆₋₁₀heterocyclyl (e.g., dioxane), including fused bicyclic groups, such as benzo-fused heterocyclyl (e.g., benzo-fused dioxane), optionally substituted as described above.

[0197] As noted above, according to some embodiments, p is 1. For example, in some embodiments, the compound of Formula (XIV) is a compound of Formula (XVb):
or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, wherein, as valence and stability permit, \( R^1, R^8, X, Ar, \) and \( n \) are as defined above.

As noted above, according to some embodiments, \( p \) is 0. For example, in some embodiments, the compound of Formula (XIV) is a compound of Formula (XVc):

or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, wherein, as valence and stability permit, \( R^1, R^8, Ar, \) and \( n \) are as defined above.

In certain embodiments, the compound of Formula (XIV) is:

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<th>Compound</th>
<th>Structure</th>
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</table>
or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof.

[0200] In some embodiments, the edible compositions of this invention comprise diphenyl-containing compounds, pyrazole-containing compounds, hydroquinoline compounds, quinoline compounds, or N-phenylalkylamide compounds as described herein, or a comestibly or biologically acceptable salt or derivative thereof, or an enantiomer or diastereomer thereof, or mixtures thereof.
If a comestibly or biologically acceptable salt of a compound of the present invention is used, such salt is preferably derived from inorganic or organic acids and bases. Examples of such salts include, but are not limited to, acetate, adipate, alginate, aspartate, benzoate, benzenesulfonate, bisulfate, butyrate, citrate, camphorate, camphorsulfonate, cyclopentanepropionate, dglucoconate, dodecylsulfate, ethanesulfonate, formate, fumarate, glucoheptanoate, glyceroxphosphate, glycolate, hemisulfate, heptanoate, hexanoate, hydrochloride, hydrobromide, hydroiodide, 2-hydroxyethanesulfonate, lactate, maleate, malonate, methanesulfonate, 2-naphthalenesulfonate, nicotinate, nitrate, oxalate, palmitate, pectinate, persulfate, phosphate, picroate, pivalate, propionate, salicylate, succinate, sulfate, tartrate, thiocyanate, tosylate and undecanoate.

Salts derived from appropriate bases include alkali metal (e.g., sodium and potassium), alkaline earth metal (e.g., magnesium), ammonium and $\text{N}^+(C_{1-4}\text{alkyl})_n$ salts. This invention also envisions the quaternization of any basic nitrogen-containing groups of the compounds disclosed herein. Water or oil-soluble or dispersible products may be obtained by such quaternization. In some embodiments, the compounds of the present invention are present as sodium, potassium or citrate salts.

Another aspect of the present invention provides edible compositions comprising a) a compound of the invention; and b) a bitter tastant. In some embodiments, the compound of the invention is a compound having a molecular weight less than about 1000, 500, or 300 daltons. In certain embodiments, the compound of the invention is a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIib), Formula (IIib"), Formula (IIIb"), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIla), Formula (VIlb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIia), Formula (XIIib), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof. In some embodiments, the compound of the invention is a compound selected from Compounds 1-58 or combinations thereof.

In some embodiments, the bitter tastant present in the edible composition is a bitter tasting salt. In some embodiments, the bitter tastant present in the edible composition is a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the bitter tastant present in the edible composition is a potassium salt. In some embodiments, the bitter tastant present in the edible compositions is KCl. In other embodiments, the bitter tastant present in the edible composition is potassium lactate.

In another embodiment, the edible compositions comprise a) a compound of the invention; and b) a potassium salt. In some embodiments, the potassium salt is KCl or potassium lactate. In specific embodiments, the potassium salt is KCl. In certain embodiments, the compound of the invention is a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIib), Formula (IIib"), Formula (IV), Formula (Va), Formula (Vb), Formula (VIa), Formula (VIb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof.
In some embodiments, the compound of the invention is a compound selected from Compounds 1-58 or combinations thereof.

[0205] In some embodiments, the edible composition further comprises a sodium salt. In some embodiments, the edible compositions further comprise NaCl. In some embodiments, the edible compositions further comprise sodium lactate. In some embodiments, the edible compositions further comprise sugar.

[0206] In some embodiments, the edible composition further comprises one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or additional flavor modifiers, which may lack an inherent flavor.

[0207] In some embodiments, the edible composition further comprises one or more emulsifiers. Sodium and potassium based emulsifiers are commonly used as emulsifiers in the food art. Sodium-based emulsifiers include, e.g., sodium salts of fatty acids, sodium alginate, sodium aluminum phosphate, sodium caseinate, sodium metaphosphate, sodium phosphate (dibasic), sodium phosphate (monobasic), sodium phosphate (tribasic), sodium polyphosphate, sodium pyrophosphate, and sodium stearoyl lactylate. Potassium-based emulsifiers include, e.g., potassium salts of fatty acids, potassium alginate, potassium citrate, potassium phosphate (dibasic), potassium phosphate (monobasic), potassium phosphate (tribasic), potassium polyphosphate, potassium polymetaphosphate, and potassium pyrophosphate. Accordingly, some embodiments of the present invention include replacing a sodium-based emulsifier with a potassium based emulsifier and adding a compound of the present invention.

[0208] In some embodiments, the edible composition further comprises a surfactant to increase or decrease the effectiveness of the compounds of the present invention. Suitable surfactants include, but are not limited to, non-ionic surfactants (e.g., mono and diglycerides, fatty acid esters, sorbitan esters, propylene glycol esters, and lactylate esters) anionic surfactants (e.g., sulfosuccinates and lecithin) and cationic surfactants (e.g., quaternary ammonium salts).

[0209] In some embodiments wherein the edible compositions further comprises a preservative, the preservative improves the shelf life of the edible composition. Suitable preservatives include, but are not limited to, ascorbic acid, benzoic acid, butyl p-hydroxybenzoate, calcium benzoate, calcium disodium EDTA, calcium hydrogen sulfate, calcium propionate, calcium sorbate, chitosan, cupric sulfate, dehydroacetic acid, diethyl pyrocarbonate, dimethyl dicarbonate, disodium EDTA, E-polylysine glycine, erythorbic acid, ethyl p-hydroxybenzoate, formic acid, gum guaiac, heptylparaben, hinokitiol, isobutyl paraoxybenzoate, Japanese styrax benzoin extract, methylparaben, milt protein extract, natamycin, nisin, peptin extract, 2-phenylphenol, pimaricin, potassium acetate, potassium benzoate, potassium lactate, potassium metabisulfite, potassium nitrate, potassium nitrite, potassium pyrosulfite, potassium sorbate, potassium sulfate, propionic acid, propyl p-hydroxybenzoate, propyl p-oxybenzoate, propylene oxide, propylparaben, sodium benzoate, sodium bisulfite, sodium dehydroacetate, sodium diacetate, sodium erythorbate, sodium hydrogen sulfate, sodium hypophosphate, sodium hyposulfite, sodium metabisulfite, sodium nitrate, sodium nitrite, sodium o-phenylphenol, sodium propionate, sodium pyrosulfite, sodium sulfite,
sodium thiocyanate, sorbic acid and sulfur dioxide. In some embodiments, the preservative has a bitterness.

[0210] In some embodiments, the composition may further comprise one or more additional components selected from the group consisting of flow agents, processing agents, sugars, amino acids, other nucleotides, and sodium or potassium salts of organic acids such as citrate and tartarate. Such additional ingredients may add flavor, or aid in blending, processing or flow properties of the edible composition.

[0211] In some embodiments, the rate of release of the compound of the present invention is regulated. The release rate of the compound of the present invention can be altered by, for example, varying its solubility in water. Rapid release can be achieved by encapsulating the compound of the present invention with a material with high water solubility. Delayed release of the compound of the present invention can be achieved by encapsulating the compound of the present invention with a material with low water solubility. The compound of the present invention can be co-encapsulated with carbohydrates or masking agents such as sweeteners. The rate of release of the compound of the present invention can also be regulated by the degree of encapsulation. In some embodiments, the compound of the present invention is fully encapsulated. In other embodiments, the compounds of the present invention are partially encapsulated. In some embodiments, the rate of release is regulated so as to release with the bitter taste.

[0212] The edible compositions of this invention are prepared according to techniques well-known in the art. In general, an edible composition of the invention is prepared by mixing a component or ingredient of the edible composition with a compound of the invention. Alternatively, a compound of the invention can be added directly to the edible composition. In some embodiments, a bitter tastant is added simultaneously or sequentially with a compound of the invention. If sequentially, the bitter tastant may be added before or after the compound of the invention. In some embodiments, the edible composition is a food product. In some embodiments, the edible composition is a pharmaceutical composition. In some embodiments, the edible composition is a consumer product.

[0213] The amount of both a compound of the present invention and a bitter tastant used in an edible composition depends upon a variety of factors, including the desired or acceptable perception of bitterness, saltiness, or sweetness. The amount may depend on the nature of the edible composition, the particular compound added, the bitter tastant, other compounds present in the composition, the method of preparation (including amount of heat used), and the pH of the edible composition. It will be understood that those of skill in the art will know how to determine the amounts needed to produce the desired taste(s).

[0214] In general, a compound of the present invention in an edible composition may be present at a concentration between about 0.001 ppm and 1000 ppm. In some embodiments, the edible composition comprises between about 0.005 to 500 ppm; 0.01 to 100 ppm; 0.05 to 50 ppm; 0.1 to 5 ppm; 0.1 to 10 ppm; 1 to 10 ppm; 1 to 30 ppm; 1 to 50 ppm; 10 to 30 ppm; 10 to 50 ppm; or 30 to 50 ppm of a compound of the present invention. In yet other embodiments, the edible composition
comprises about 0.1 to 30 ppm, 1 to 30 ppm or 1 to 50 ppm of a compound of the present invention. In additional embodiments, the edible composition comprises about 0.1 to 5 ppm; 0.1 to 4 ppm; 0.1 to 3 ppm; 0.1 to 2 ppm; 0.1 to 1 ppm; 0.5 to 5 ppm; 0.5 to 4 ppm; 0.5 to 3 ppm; 0.5 to 2 ppm; 0.5 to 1.5 ppm; 0.5 to 1 ppm; 5 to 15 ppm; 6 to 14 ppm; 7 to 13 ppm; 8 to 12 ppm; 9 to 11 ppm; 25 to 35 ppm; 26 to 34 ppm; 27 to 33 ppm; 28 to 32 ppm; or 29 to 31 ppm.

[0215] In yet other embodiments, the edible composition comprises about 0.1 ppm, about 0.5 ppm, about 1 ppm, about 2 ppm, about 3 ppm, about 4 ppm, about 5 ppm, about 6 ppm, about 7 ppm, about 8 ppm, about 9 ppm, or about 10 ppm of a compound of the present invention. In other embodiments, the edible composition comprises about 11 ppm, about 12 ppm, about 13 ppm, about 14 ppm, about 15 ppm, about 16 ppm, about 17 ppm, about 18 ppm, about 19 ppm, about 20 ppm, about 21 ppm, about 22 ppm, about 23 ppm, about 24 ppm, about 25 ppm, about 26 ppm, about 27 ppm, about 28 ppm, about 29 ppm, or about 30 ppm of a compound of the present invention.

[0216] In still other embodiments, the edible composition comprises about 31 ppm, about 32 ppm, about 33 ppm, about 34 ppm, about 35 ppm, about 36 ppm, about 37 ppm, about 38 ppm, about 39 ppm, about 40 ppm, about 41 ppm, about 42 ppm, about 43 ppm, about 44 ppm, about 45 ppm, about 46 ppm, about 47 ppm, about 48 ppm, about 49 ppm, or about 50 ppm of a compound of the present invention.

[0217] In other embodiments, the edible composition comprises more than about 0.5 ppm, 1 ppm, 5 ppm, 10 ppm, 15 ppm, 20 ppm, 25 ppm, or 30 ppm of a compound of the present invention, up to, for example, about 30 ppm or 50 ppm. In additional embodiments, the edible composition comprises less than about 50 ppm, 30 ppm, 25 ppm, 20 ppm, 15 ppm, 10 ppm, 5 ppm, 1 ppm, or 0.5 ppm of a compound of the present invention. In yet additional embodiments, the edible composition comprises less than about 30 ppm, 10 ppm, or 1 ppm of a compound of the present invention.

[0218] When the edible composition comprises KCl, the amount of KCl will vary depending on the nature of the edible composition, the amount of perceived saltiness desired and the presence of other compounds in the composition. In some embodiments, KCl is present at a concentration between about 0.001-5% w/w; 0.01-5% w/w; 0.1-5% w/w; 0.5-4.8% w/w; 0.5-4% w/w; 0.5-3% w/w; 0.75-3% w/w; 1-2.5% w/w; or 1-2% w/w. In some embodiments, KCl is present at a concentration of about 0.5% w/w, about 1% w/w, about 1.5% w/w, about 2% w/w, about 2.5% w/w, about 3% w/w, about 3.5% w/w, about 4% w/w, about 4.5% w/w, or about 5% w/w. In some embodiments, KCl is present at a concentration of up to about 0.5% w/w, up to about 1% w/w, up to about 1.5% w/w, up to about 2% w/w, up to about 2.5% w/w, up to about 3% w/w, up to about 3.5% w/w, up to about 4% w/w, up to about 4.5% w/w, or up to about 5% w/w. In some embodiments, KCl is present at a concentration of about 2% w/w.

[0219] In some embodiments, KCl is added to the edible composition as a salt substitute in an amount sufficient to replace NaCl. For example, the amount of KCl in the edible composition may range from about 0.5 to about 1.5 times the replaced NaCl depending upon the application, e.g., if
about 0.5 mg of NaCl is replaced, about 0.25 to about 0.75 mg of KCl is added. Typically, KCl is added in the same weight amount as the NaCl being replaced.

[0220] Similarly, when the edible composition comprises potassium lactate, the amount of potassium lactate added varies depending on the nature of the edible composition, the amount of preservation required and the presence of other compounds in the composition. Potassium lactate may be present at a concentration between about 0.001-5% w/w; 0.01-5% w/w; 0.1-5% w/w; 0.5-4.8% w/w; 0.5-4% w/w; 0.5-3% w/w; 0.75-3% w/w; 1-2.5% w/w; or 1-2% w/w.

[0221] In some embodiments, potassium lactate is added to the edible composition in an amount sufficient to replace sodium lactate. For example, the amount of potassium lactate in the food or beverage after the sodium lactate substitute is added may range from about 0.5 to about 1.5 times the replaced sodium lactate depending upon the application, e.g., if about 0.5 mg of sodium lactate is replaced, about 0.25 to about 0.75 mg of potassium lactate is added. Typically, potassium lactate will be added in the same weight amount as the sodium lactate being replaced.

[0222] Further, when the edible composition comprises an artificial sweetener, such as Acesulfame K, the amount of the sweetener added varies depending on the nature of the edible composition, the amount of sweetness required and the presence of other compounds in the composition. Acesulfame K, for example, may be present at a concentration between about 1-200 ppm, 10-200 ppm, 50-150 ppm, 50-125 ppm, 75-125 ppm, and 75-100 ppm, preferably about 75 ppm.

[0223] In some embodiments, an artificial sweetener is added to the edible composition in an amount sufficient to replace sugar. In some embodiments, the artificial sweetener has a bitter taste or aftertaste. In some embodiments, the artificial sweetener is Acesulfame K. For example, the amount of Acesulfame K in the edible composition may range from about 0.001 to about 0.01 times the replaced sugar depending upon the application, e.g., if about 100 mg of sugar is replaced, about 0.1 to about 1 mg of Acesulfame K is added. Typically, Acesulfame K will be added in about 0.005 times the amount of sugar being replaced.

[0224] In some embodiments, the edible compositions are included in a package. In some embodiments, the edible composition is packaged in bulk, in which the package contains more of the compositions than would typically be used for a single dish or serving of food or beverage. Such bulk packages can be in the form of paper, plastic, or cloth bags or cardboard boxes or drums. Such bulk packages may be fitted with plastic or metal spouts to facilitate the dispensing of the edible composition.

[0225] In some embodiments, the package contains an edible composition comprising a compound of the present invention and a bitter taste. In some embodiments, the package contains an edible composition comprising a compound of the present invention and bitter tasting salt. In some embodiments, the package contains an edible composition comprising a compound of the present invention and a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the package contains an edible composition comprising a compound of the present invention and a potassium salt. In some embodiments, the package contains an edible composition comprising a
compound of the present invention and KCl. In other embodiments, the package contains an edible composition comprising a compound of the present invention and potassium lactate. In some embodiments, the package contains an edible composition comprising a compound of the present invention and potassium salt, and a sodium salt. In other embodiments, the package contains an edible composition comprising a compound of the present invention, KCl and NaCl. In yet other embodiments, the package contains an edible composition comprising a compound of the present invention, potassium lactate and sodium lactate. In other embodiments, the package contains an edible composition comprising a compound of the present invention and Acesulfame K and sugar. In other embodiments, the package contains an edible composition comprising a compound of the present invention, potassium lactate, KCl and NaCl.

[0226] In some embodiments, the edible compositions of the present invention are compositions suitable to be used as seasonings, as ingredients in food products or as condiments. In such embodiments, the edible composition may or may not contain a bitter tastant. For example, the edible composition may be used in, e.g., a seasoning which comprises a bitter tastant such as, e.g., KCl. Such seasonings can be used in the place of table salt (i.e., NaCl) to season prepared food products. Alternatively, the edible composition may be used in, e.g., a seasoning which does not contain a bitter tastant. Such seasonings can be used to season prepared food products which contain a bitter tastant (either inherently present or added during preparation) in order to reduce the bitter taste associated with the bitter tastant. In some embodiments, the edible composition is a seasoning comprising KCl and a compound of the invention. In some embodiments, the edible composition is a seasoning comprising KCl, NaCl and a compound of the invention. In some embodiments the seasoning further comprises a spice or a blend of spices.

[0227] Alternatively, the edible compositions may be used for medicinal or hygienic purposes, for example, in soaps, shampoos, mouthwash, medicines, pharmaceuticals, cough syrup, nasal sprays, toothpaste, dental adhesives, tooth whiteners, glues (e.g., on stamps and envelopes), and toxins used in insect and rodent control.

**Food product**

[0228] In some embodiments, the edible composition is a food product. According to such embodiments, the food product comprises (a) a food stuff; and (b) a compound of Formula (I), Formula (Ia), Formula (Ib), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb'), Formula (IV), Formula (Va), Formula (Vb), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVIb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof.

[0229] In some embodiments, the food product further comprises a bitter tastant, as described herein. In some embodiments, the bitter tastant is a potassium salt, such as KCl or potassium lactate. In specific embodiments, the potassium salt is KCl.
[0230] In some embodiments, the food product further comprises one or more additional flavor modifiers.

[0231] In some embodiments, the food product further comprises one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or additional flavor modifiers, which may lack an inherent flavor.

Pharmaceutical Composition

[0232] In some embodiments, the edible composition is a pharmaceutical composition. According to such embodiments, the pharmaceutical composition comprises (a) a bitter tasting pharmaceutically active ingredient; and (b) a compound of Formula (I), Formula (IIa), Formula (I Ib), Formula (IIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (V a), Formula (V b), Formula (V I a), Formula (V I b), Formula (VII a), Formula (VII b), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XII a), Formula (XII b), Formula (XIII a), Formula (XIII b), Formula (XIV), Formula (X V a), Formula (X V b) or Formula (X V c), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof.

[0233] According to some embodiments, the pharmaceutical composition can comprise any bitter tasting pharmaceutically active ingredient. Non-limiting examples of bitter pharmaceutical compounds include: acetaminophen, ampicillin, azithromycin, chlorpheniramine, cimetidine, dextromethorphan, diphenhydramine, erythromycin, ibuprofen, penicillin, phenylbutazone, pseudoephedrine, ranitidine, spironolactone statins (including, but not limited to, atorvastatin, cerivastatin, fluvastatin, lovastatin, mevastatin, pitavastatin, pravastatin, rosuvastatin, and simvastatin) and theophylline.

[0234] In other embodiments, the invention provides a pharmaceutical composition comprising (a) a pharmaceutically active ingredient; (b) a compound of Formula (I), Formula (II a), Formula (II b), Formula (III b'), Formula (III b''), Formula (IV), Formula (V a), Formula (V b), Formula (V I a), Formula (V I b), Formula (VII a), Formula (VII b), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XII a), Formula (XII b), Formula (XIII a), Formula (XIII b), Formula (XIV), Formula (X V a), Formula (X V b) or Formula (X V c), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof; and (c) a bitter tastant. In such embodiments, the pharmaceutical compositions may comprise any pharmaceutically active ingredient.

[0235] In other embodiments, the invention provides a pharmaceutical composition comprising (a) a pharmaceutically active ingredient; (b) a compound of Formula (I), Formula (II a), Formula (II b), Formula (III b'), Formula (III b''), Formula (IV), Formula (V a), Formula (V b), Formula (V I a), Formula (V I b), Formula (VII a), Formula (VII b), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XII a), Formula (XII b), Formula (XIII a), Formula (XIII b), Formula (XIV), Formula (X V a), Formula (X V b) or Formula (X V c), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or
combinations thereof; and (c) a potassium salt. In some embodiments, the potassium salt is KCl or potassium lactate. In some embodiments, the potassium salt is KCl.

[0236] In some embodiments, the pharmaceutical composition further comprises one or more additional flavor modifiers.

[0237] In some embodiments, the pharmaceutical composition further comprises one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or additional flavor modifiers, which may lack an inherent flavor.

**Consumer product**

[0238] In some embodiments, the edible compositions is a consumer product. According to such embodiments, the consumer product comprises (a) a bitter tastant and (b) a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb*), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof.

[0239] According to another embodiment, the invention provides a consumer product comprising (a) a potassium salt; and (b) a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb*), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof. In some embodiments, the potassium salt is KCl or potassium lactate. In some embodiments, the potassium salt is KCl.

[0240] In other embodiments, the invention provides a consumer product for reducing bitter taste of a bitter tastant, wherein said consumer product comprises a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb*), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof. In some embodiments, the bitter tastant is a potassium salt. In some embodiments, the potassium salt is KCl or potassium lactate. In some embodiments, the bitter tastant is KCl.

[0241] In some embodiments, the consumer product further comprises one or more additional flavor modifiers.
[0242] In some embodiments, the consumer product further comprises one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or additional flavor modifiers, which may lack an inherent flavor.

Method of preparing an edible composition

[0243] According to another aspect, the invention provides a method of preparing an edible composition. The method comprises: (a) providing a comestibly acceptable carrier; and (b) adding to the comestibly acceptable carrier of (a) a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb"), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof, with the comestibly acceptable carrier. In some embodiments, the compound of the invention has been dissolved in a solvent prior to the addition step (b).

[0244] In some embodiments, the comestibly acceptable carrier in (a) is inherently bitter. In such embodiments, the comestibly acceptable carrier may inherently contain a bitter tantant. In some embodiments, the inherent bitter tantant is a bitter tasting salt. In some embodiments, the inherent bitter tantant is a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the inherent bitter tantant is a potassium salt. In some embodiments, the inherent bitter tantant is KCl. In other embodiments, the inherent bitter tantant is potassium lactate.

[0245] In some embodiments, the method of preparing a edible composition further comprises: (c) adding a bitter tantant. In some embodiments, the bitter tantant is a potassium salt. In some embodiments, the potassium salt is KCl or potassium lactate. In specific embodiments, the potassium salt is KCl. In some embodiments, the bitter tantant is added before the compound of the present invention. In other embodiments, the bitter tantant is added after the compound of the present invention. In some embodiments, the compounds of the present invention are combined with the bitter tantant and then combined with the comestibly acceptable carrier. In other embodiments, the compound of the present invention is combined sequentially with the comestibly acceptable carrier and then the bitter tantant. In yet other embodiments, the compounds of the present invention are combined with a mixture of the bitter tantant and the comestibly acceptable carrier.

[0246] In some embodiments, a compound of the invention and the bitter tantant, if present, are mixed with the comestibly acceptable carrier. In other embodiments, the compound and the bitter tantant, if present, are sprayed onto or coat the comestibly acceptable carrier. In some embodiments, the compound of the invention is plated on a carbohydrate or salt, encapsulated on a salt or a carbohydrate (spray dried), or co-crystallized with a potassium salt to create a "topping" salt.
[0247] In some embodiments, the bitter tastant is a bitter tasting salt. In some embodiments, the bitter tastant is a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the bitter tastant is a potassium salt. In some embodiments, the bitter tastant is KCl. In other embodiments, the bitter tastant is potassium lactate.

[0248] In some embodiments, the edible composition further comprises a sodium salt. In some embodiments, the edible composition further comprises NaCl. In other embodiments, the edible composition further comprises sodium lactate. In further embodiments, the edible composition further comprises sugar.

[0249] In some embodiments, the methods of preparing an edible composition further comprise adding one or more additional components selected from the group consisting of preservatives, nutritive, flavorants or flavor modifiers, which may lack an inherent flavor. In some embodiments, the methods of preparing an edible composition further comprise adding one or more additional flavor modifiers.

[0250] In some embodiments, the edible composition is a consumer product.

Method of preparing a food product

[0251] According to another aspect, the invention provides a method of preparing an edible composition, wherein the edible composition is a food product. The method comprises: (a) providing a foodstuff; and (b) adding to the foodstuff of (a) a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIb), Formula (IIb'), Formula (IIb*), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIa), Formula (XIIb), Formula (XIV), Formula (XV), Formula (XVI) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof. In some embodiments, the compound of the invention is added in the form of an edible composition comprising the compound of the invention.

[0252] In some embodiments, the foodstuff in (a) is inherently bitter. In such embodiments, the foodstuff may inherently contain a bitter tastant. In some embodiments, the inherent bitter tastant is a bitter tasting salt. In some embodiments, the inherent bitter tastant is a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the inherent bitter tastant is a potassium salt. In some embodiments, the inherent bitter tastant is KCl. In other embodiments, the inherent bitter tastant is potassium lactate.

[0253] In some embodiments, the method comprises: (a) providing a food product; and (b) adding to the food product of (a) an edible composition comprising compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIb), Formula (IIb'), Formula (IIb*), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIa), Formula (XIIb), Formula (XIV), Formula (XV), Formula (XVI) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as
described above, or combinations thereof. In some embodiments, the compound of the invention is added in the form of an edible composition comprising the compound of the invention.

[0254] In some embodiments, the food product in (a) comprises a bitter tastant. In some embodiments, the bitter tastant is a bitter tasting salt. In some embodiments, the bitter tastant is a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the bitter tastant is a potassium salt. In some embodiments, the bitter tastant is KCl. In other embodiments, the bitter tastant is potassium lactate.

[0255] In some embodiments, the method of preparing a food product further comprises: (c) adding a bitter tastant. In some embodiments, the bitter tastant is a potassium salt, such as KCl or potassium lactate. In specific embodiments, the potassium salt is KCl. In some embodiments, the bitter tastant is added before the compound of the present invention. In other embodiments, the bitter tastant is added after the compound of the present invention. In some embodiments, the compound of the invention is added with the bitter tastant. In some embodiments, the compound of the present invention is combined with the bitter tastant and then combined with the foodstuff or food product. In other embodiments, the compound of the present invention is combined sequentially with the foodstuff or food product and then the bitter tastant. In yet other embodiments, the compound of the present invention is combined with a mixture of the bitter tastant and the foodstuff or food product.

[0256] In some embodiments, the compound and the bitter tastant, if present, are mixed with the foodstuff. In other embodiments, the compound and the bitter tastant, if present, are sprayed onto or coat the foodstuff. In some embodiments, the compound of the invention is plated on a carbohydrate or salt, encapsulated on a salt or a carbohydrate (spray dried), or co-crystallized with a potassium salt to create a "topping" salt.

[0257] In some embodiments, the bitter tastant is a bitter tasting salt. In some embodiments, the bitter tastant is a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the bitter tastant is a potassium salt. In some embodiments, the bitter tastant is KCl. In other embodiments, the bitter tastant is potassium lactate.

[0258] In some embodiments, the food product further comprises a sodium salt. In some embodiments, the food product further comprises NaCl. In other embodiments, the food product further comprises sodium lactate. In further embodiments, the food product further comprises sugar.

[0259] In some embodiments, the methods of preparing a food product further comprise adding one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or flavor modifiers, which may lack an inherent flavor.

Method of preparing a pharmaceutical composition

[0260] According to another aspect, the invention provides a method of preparing an edible composition, wherein the edible composition is a pharmaceutical composition. The method comprises: (a) providing a pharmaceutically active ingredient; and (b) adding to the
pharmaceutically active ingredient of (a) a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb*), Formula (IIIb**), Formula (IV), Formula (Va), Formula (Vb), Formula (VIa), Formula (VIb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XV), Formula (XVa) or Formula (XVe), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof, with the pharmaceutically active ingredient. In some embodiments, the compound of the invention is added in the form of an edible composition comprising the compound of the invention.

[0261] In some embodiments, the pharmaceutically active ingredient in (a) is inherently bitter. In such embodiments, the pharmaceutically active ingredient may inherently contain a bitter tastant. In some embodiments, the inherent bitter tastant is a bitter tasting salt. In some embodiments, the inherent bitter tastant is a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the inherent bitter tastant is a potassium salt.

[0262] In some embodiments, the method of preparing a pharmaceutical composition further comprises: (c) adding a bitter tastant. In some embodiments, the bitter tastant is a potassium salt. In some embodiments, the potassium salt is KCl or potassium lactate. In specific embodiments, the potassium salt is KCl. In some embodiments, the bitter tastant is added before the compound of the present invention. In other embodiments, the bitter tastant is added after the compound of the present invention. In some embodiments, the bitter tastant is added with the compound of the invention. In some embodiments, the compound of the present invention is combined with the bitter tastant and then combined with the pharmaceutically active ingredient. In other embodiments, the compound of the present invention is combined sequentially with the pharmaceutically active ingredient and then the bitter tastant. In yet other embodiments, the compound of the present invention is combined with a mixture of the bitter tastant and the pharmaceutically active ingredient.

[0263] In some embodiments, the compound and the bitter tastant, if present, are mixed with the pharmaceutically active ingredient. In other embodiments, the compound and the bitter tastant, if present, are sprayed onto or coat the pharmaceutical composition. In some embodiments, the compound of the invention is encapsulated with the pharmaceutically active ingredient. In some embodiments, the compound of the invention is in a form such that the rate of release is regulated vis a vis the rate of release of the bitter tastant, which in some embodiments is the pharmaceutically active ingredient.

[0264] In some embodiments, the bitter tastant is a bitter tasting salt. In some embodiments, the bitter tastant is a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the bitter tastant is a potassium salt. In some embodiments, the bitter tastant is KCl. In other embodiments, the bitter tastant is potassium lactate.

[0265] In some embodiments, the pharmaceutical composition further comprises a sodium salt. In some embodiments, the pharmaceutical composition further comprises NaCl. In other
embodiments, the pharmaceutical composition further comprises sodium lactate. In further embodiments, the pharmaceutical composition further comprises sugar.

[0266] In some embodiments, the pharmaceutical composition further comprises a pharmaceutically acceptable carrier. Pharmaceutically acceptable carriers that may be used in these compositions include, but are not limited to, ion exchangers, alumina, aluminum stearate, lecithin, serum proteins such as human serum albumin, buffer substances such as phosphates, glycine, sorbic acid, potassium sorbate, partial glyceride mixtures of saturated vegetable fatty acids, water, salts or electrolytes such as protamine sulfate, disodium hydrogen phosphate, potassium hydrogen phosphate, sodium chloride, zinc salts, colloidal silica, magnesium trisilicate, polyvinyl pyrrolidone, cellulose-based substances, polyethylene glycol, sodium carboxymethylcellulose, polyacrylates, waxes, polyethylene-polyoxypropylene-block polymers, polyethylene glycol and wool fat.

[0267] In some embodiments, the methods of preparing a pharmaceutical composition further comprise adding one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or flavor modifiers, which may lack an inherent flavor.

Method of reducing or eliminating the perception of bitter taste in a subject

[0268] According to another aspect, the invention provides a method of reducing or eliminating the perception of bitter taste in a subject. The method comprises the use of an edible composition comprising a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb*), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof.

[0269] The method can be used to reduce or eliminate bitter taste in any edible composition, including a foodstuff, food product, pharmaceutical composition or consumer product. The edible composition may be in any form. In some embodiments, the composition is in the form of, for example, a gum, lozenge, sauce, condiment, meat matrix, meat slurry, paste, suspension, spread, coating, a liquid, a gel, an emulsion, granules, or seasoning.

[0270] In some embodiments the edible composition is utilized by, for example, placement in the oral cavity or by ingestion. In some embodiments, the edible composition is placed in the oral cavity or ingested before a bitter foodstuff, food product, pharmaceutical composition or consumer product. In some embodiments, the edible composition is placed in the oral cavity or ingested concurrently with a bitter foodstuff, food product, pharmaceutical composition or consumer product, either as a separate edible composition or by incorporation in the bitter foodstuff, food product, pharmaceutical composition or consumer product. In some embodiments, the edible composition is placed in the oral cavity or ingested after a bitter foodstuff, food product, pharmaceutical composition or consumer product. For example, a compound of the invention can
be combined with foodstuffs or food products to reduce the bitter taste of a food product. Alternatively, a compound of the invention can be used, for example, in a lozenge or gum for use after exposure to a bitter food stuff, food product, pharmaceutical composition or consumer product (e.g., to reduce or eliminate a bitter aftertaste).

5 **Method of reducing the amount of sodium in an edible composition**

[0271] According to another embodiment, the invention provides a method of reducing the amount of sodium in an edible composition, such as a food product, a pharmaceutical composition or a consumer product. In some embodiments, the invention provides a method of reducing the amount of a sodium containing compound in an edible composition, such as a food product, a pharmaceutical composition or a consumer product. In another embodiment, the invention provides a method of reducing the amount of NaCl in an edible composition, such as a food product, a pharmaceutical composition or a consumer product. In another embodiment, the invention provides a method of reducing the amount of sodium lactate in an edible composition, such as a food product, a pharmaceutical composition or a consumer product. In some embodiments, the sodium salt is replaced with a non-sodium salt. In some embodiments, the non-sodium salt is a calcium salt, a magnesium salt, or a potassium salt. In some embodiments, the non-sodium salt is a potassium salt.

[0272] In some embodiments, the method comprises: (a) replacing an amount of a sodium salt present in an edible composition with an amount of a potassium salt; and (b) incorporating into the edible composition an effective amount of a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb′), Formula (IV), Formula (Va), Formula (Vb), Formula (Va′), Formula (VIIb), Formula (VIIIa), Formula (VIIIb), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof. In some embodiments, the compound of the invention is added in the form of an edible composition comprising the compound of the invention.

[0273] In some embodiments, the method of reducing the amount of sodium in an edible composition comprises the steps of: (a) ingesting a first edible composition, in which an amount of a sodium salt has been replaced with an amount of a potassium salt; and (b) ingesting a second edible compound, which comprises a compound of the invention. In some embodiments, the first edible composition is ingested before the second edible composition. In some embodiments, the first edible composition is ingested after the second edible composition. In some embodiments, the first edible composition is ingested concurrently with the second edible composition.

[0274] In some embodiments, the edible composition is a food product. In some embodiments, the edible composition is a pharmaceutical composition. In some embodiments, the edible composition is a consumer product.
In some embodiments, the potassium salt is added to the edible composition prior to addition of an effective amount of a compound of the invention. In some embodiments, the potassium salt is added to the edible composition subsequent to addition of an effective amount of a compound of the invention. In some embodiments, the potassium salt is added to the edible composition concurrent with addition of an effective amount of a compound of the invention. In some embodiments, the amount of sodium replaced in the edible composition in step (a) is an amount sufficient to maintain or restore the health of a subject. In some embodiments, the amount of sodium replaced in the edible composition is an amount sufficient to decrease hypertension in a subject. In some embodiments, the amount of sodium replaced by potassium in the edible composition is an amount sufficient to change the texture or freezing point of the edible composition. In some embodiments, the amount of sodium replaced is up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70% 75%, 80%, 85%, 90%, 95% or 100%. These amounts are not meant to be limiting, and increments between the recited percentages are specifically envisioned as part of the invention. In some embodiments, the amount of compound added in step (b) reduces the perception of bitter taste in the subject. The bitter taste is completely reduced or partially reduced. In some embodiments, the perception of salty taste is maintained. In some embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70% 75%, 80%, 85%, 90%, 95% or 100% of the amount of sodium present in the edible composition with potassium. These amounts are not meant to be limiting, and increments between the recited percentages are specifically envisioned as part of the invention. In some embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 25% of the amount of sodium present in the edible composition with potassium. In other embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 50% of the amount of sodium present in the edible composition with potassium. In other embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 75% of the amount of sodium present in the edible composition with potassium. In yet other embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 100% of the amount of sodium present in the edible composition with potassium. In some embodiments, the method of reducing the amount of sodium in an edible composition further comprises adding one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or flavor modifiers, which may lack an inherent flavor. In some embodiments, the method comprises: (a) replacing an amount of NaCl present in an edible composition with an amount of KCl; and (b) incorporating into the edible composition an effective amount of a compound of Formula (I), Formula (Ia), Formula (Ib), Formula (IIb), Formula (IIIb), Formula (IIb'), Formula (IIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (VIa),
Formula (VIIb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X),
Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV),
Formula (XV), Formula (XVI), Formula (XV), as described herein, or combinations thereof, or
any one of Compounds 1-58, as described above, or combinations thereof.

[0281] In some embodiments, the method of reducing the amount of sodium in an edible
composition comprises the steps of: (a) ingesting a first edible composition, in which an amount of
NaCl has been replaced with an amount of KCl; and (b) ingesting a second edible compound, which
comprises a compound of the invention. In some embodiments, the first edible composition is
ingested before the second edible composition. In some embodiments, the first edible composition
is ingested after the second edible composition. In some embodiments, the first edible composition
is ingested concurrently with the second edible composition.

[0282] In some embodiments, the edible composition is a food product. In some embodiments,
the edible composition is a pharmaceutical composition. In some embodiments, the edible
composition is a consumer product.

[0283] In some embodiments, the KCl is added to the edible composition prior to addition of an
effective amount of a compound of the invention. In some embodiments, the KCl is added to the
edible composition subsequent to addition of an effective amount of a compound of the invention.
In some embodiments, the KCl is added to the edible composition concurrent with addition of an
effective amount of a compound of the invention.

[0284] In some embodiments, the amount of NaCl replaced by KCl in the edible composition in
step (a) is an amount sufficient to maintain or restore the health of a subject. In some embodiments,
the amount of NaCl replaced by KCl in the edible composition is an amount sufficient to decrease
hypertension in a subject. In some embodiments, the amount of NaCl replaced by KCl in the edible
composition is an amount sufficient to change the texture or freezing point of the edible
composition. In some embodiments, the amount of NaCl replaced by KCl is up to 1%, 2%, 3%,
4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%,
70%, 75%, 80%, 85%, 90%, 95% or 100%. These amounts are not meant to be limiting, and
increments between the recited percentages are specifically envisioned as part of the invention.

[0285] In some embodiments, the amount of compound added in step (b) reduces the perception
of bitter taste in the subject. The bitter taste is completely reduced or partially reduced. In some
embodiments, the perception of salty taste is maintained.

[0286] In some embodiments, the amount of compound added in step (b) is sufficient to permit
replacement of up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%,
40%, 45%, 50%, 55%, 60%, 65%, 70%, 75%, 80%, 85%, 90%, 95% or 100% of the amount of
NaCl present in the edible composition with KCl. These amounts are not meant to be limiting, and
increments between the recited percentages are specifically envisioned as part of the invention. In
some embodiments, the amount of compound added in step (b) is sufficient to permit replacement
of up to 25% of the amount of NaCl present in the edible composition with KCl. In other
embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up
to 50% of the amount of NaCl present in the edible composition with KCl. In other embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 75% of the amount of NaCl present in the edible composition with KCl. In yet other embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 100% of the amount of NaCl present in the edible composition with KCl.

[0287] In some embodiments, the method of reducing the amount of NaCl in an edible composition or food product comprises maintaining a salty flavor.

[0288] In some embodiments, the method of reducing the amount of NaCl in an edible composition further comprises adding one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or flavor modifiers, which may lack an inherent flavor.

[0289] In other embodiments, the method of reducing the amount of sodium in an edible composition or food product comprises: (a) replacing an amount of sodium lactate present in a food product with an amount of potassium lactate; and (b) incorporating into the edible composition an effective amount of a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (VIa), Formula (VIb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof.

[0290] In some embodiments, the method of reducing the amount of sodium in an edible composition comprises the steps of: (a) ingesting a first edible composition, in which an amount of sodium lactate has been replaced with an amount of potassium lactate; and (b) ingesting a second edible compound, which comprises a compound of the invention. In some embodiments, the first edible composition is ingested before the second edible composition. In some embodiments, the first edible composition is ingested after the second edible composition. In some embodiments, the first edible composition is ingested concurrently with the second edible composition.

[0291] In some embodiments, the edible composition is a food product. In some embodiments, the edible composition is a pharmaceutical composition. In some embodiments, the edible composition is a consumer product.

[0292] In some embodiments, the potassium lactate is added to the edible composition prior to addition of an effective amount of a compound of the invention. In some embodiments, the potassium lactate is added to the edible composition subsequent to addition of an effective amount of a compound of the invention. In some embodiments, the potassium lactate is added to the edible composition concurrent with addition of an effective amount of a compound of the invention.

[0293] In some embodiments, the amount of sodium lactate replaced by potassium lactate in the edible composition in step (a) is an amount sufficient to maintain or restore the health of a subject. In some embodiments, the amount of sodium lactate replaced by potassium lactate in the edible composition is an amount sufficient to decrease hypertension in a subject. In some embodiments,
the amount of sodium lactate replaced by potassium lactate in the edible composition is an amount to sufficient to change the texture or freezing point of the edible composition. In some embodiments, the amount of sodium lactate replaced by potassium lactate is up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70%, 75%, 80%, 85%, 90%, 95% or 100%. These amounts are not meant to be limiting, and increments between the recited percentages are specifically envisioned as part of the invention.

[0294] In some embodiments, the amount of compound added in step (b) reduces the perception of bitter taste in the subject. The bitter taste is completely reduced or partially reduced. In some embodiments, the perception of salty taste is maintained.

[0295] In some embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70%, 75%, 80%, 85%, 90%, 95% or 100% of the amount of sodium lactate present in the edible composition with potassium lactate. These amounts are not meant to be limiting, and increments between the recited percentages are specifically envisioned as part of the invention. In some embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 25% of the amount of sodium lactate present in the edible composition with potassium lactate. In other embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 50% of the amount of sodium lactate present in the edible composition with potassium lactate. In other embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 75% of the amount of sodium lactate present in the edible composition with potassium lactate. In yet other embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 100% of the amount of sodium lactate present in the edible composition with potassium lactate.

[0296] In some embodiments, the method of reducing the amount of sodium lactate in an edible composition or food product comprises maintaining the preservation of the food product.

[0297] In some embodiments, the method of reducing the amount of sodium lactate in an edible composition further comprises adding one or more additional components selected from the group consisting of preservatives, nutritive, flavorants or flavor modifiers, which may lack an inherent flavor.

[0298] Method of reducing the amount of sugar in an edible composition

According to another embodiment, the invention provides a method of reducing the amount of sugar in an edible composition. In some embodiments, the method comprises: (a) replacing an amount of sugar present in an edible composition with an amount of Acesulfame K; and (b) incorporating into the edible composition an effective amount of a compound of Formula (I), Formula (Ia), Formula (Ib), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa),
Formula (XVb) or Formula (XVe), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof.

[0299] In some embodiments, the edible composition is a food product. In some embodiments, the edible composition is a pharmaceutical composition. In some embodiments, the edible composition is a consumer product.

[0300] In some embodiments, the Acesulfame K is added to the edible composition prior to addition of an effective amount of a compound of the invention. In some embodiments, the Acesulfame K is added to the edible composition subsequent to addition of an effective amount of a compound of the invention. In some embodiments, the Acesulfame K is added to the edible composition concurrent with addition of an effective amount of a compound of the invention.

[0301] In some embodiments, the amount of sugar replaced in the edible composition in (a) is an amount sufficient to maintain or restore the health of a subject. In some embodiments, the amount of sugar replaced in the edible composition is an amount sufficient to result in weight loss in a subject. In some embodiments, the amount of sugar replaced by Acesulfame K in the edible composition is an amount to sufficient to alleviate the effects of, or treat, a disease associated with sugar consumption or excessive weight of the subject (e.g., diabetes). In some embodiments, the amount of sugar replaced by Acesulfame K is up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70%, 75%, 80%, 85%, 90%, 95% or 100%. These amounts are not meant to be limiting, and increments between the recited percentages are specifically envisioned as part of the invention.

[0302] In some embodiments, the amount of compound added in (b) reduces the perception of bitter taste in the subject. The bitter taste is completely reduced or partially reduced. In some embodiments, the perception of sweet taste is maintained.

[0303] In some embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70%, 75%, 80%, 85%, 90%, 95% or 100% of the amount of sugar present in the edible composition with Acesulfame K. These amounts are not meant to be limiting, and increments between the recited percentages are specifically envisioned as part of the invention. In some embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 25% of the amount of sugar present in the edible composition with Acesulfame K. In other embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 50% of the amount of sugar present in the edible composition with Acesulfame K. In other embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 75% of the amount of sugar present in the edible composition with Acesulfame K. In yet other embodiments, the amount of compound added in step (b) is sufficient to permit replacement of up to 100% of the amount of sugar present in the edible composition with Acesulfame K.

[0304] In some embodiments, the method of reducing the amount of sugar in an edible composition comprises maintaining a sweet flavor.
[0305] In some embodiments, the method of reducing the amount of sugar in an edible composition or food product further comprises adding one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or flavor modifiers, which may lack an inherent flavor.

5 Method of reducing sodium intake of a subject

[0306] According to another embodiment, the invention provides a method of reducing sodium intake of a subject. In some embodiments, the method comprises the step of providing an edible composition of the present invention to the subject, wherein all or a portion of the sodium salts in the edible composition is replaced with one or more non-sodium salts, and wherein the edible composition comprises a compound of the present invention. In some embodiments, the non-sodium salt is a calcium salt, a magnesium salt, or a potassium salt. In some embodiments, the non-sodium salt is a potassium salt. In some embodiments, the edible composition is a food product. In some embodiments, the edible composition is a pharmaceutical composition. In some embodiments, the edible composition is a consumer product. In some embodiments the sodium salt is NaCl and the potassium salt is KCl. In some embodiments, the sodium salt is sodium lactate and the potassium salt is potassium lactate.

[0307] In some embodiments, the methods of reducing sodium intake of a subject further comprise the step of identifying a subject in need thereof. The skilled worker would be able to identify a subject in need of reducing sodium intake. Non-limiting examples of such subjects include subjects that suffer from any one or more of the following disorders: hypernatremia, hypertension, cardiovascular disease, edema, seizures due to cerebral edema, dehydration (due to excess sweating, diarrhea, urinary tract disorders or diuretics), diabetes insipidus, Conn's syndrome, and Cushing's syndrome.

[0308] In some embodiments, the amount of the sodium salt replaced by a potassium salt in the edible composition is an amount sufficient to maintain or restore the health of a subject. In some embodiments, the amount of the sodium salt replaced by a potassium salt in the edible composition is an amount sufficient to decrease hypertension in a subject. In some embodiments, the amount of the sodium salt replaced by a potassium salt in the edible composition is up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70%, 75%, 80%, 85%, 90%, 95% or 100%. These amounts are not meant to be limiting, and increments between the recited percentages are specifically envisioned as part of the invention. In some embodiments, a subject's daily sodium intake is less than 2500 mg/day, less than 2000 mg/day, less than 1500 mg/day, less than 1000 mg/day, or less than 500 mg/day, where desirable.

[0309] In some embodiments, the amount of the compound of the invention added to the edible composition is sufficient to reduce a subject's sodium intake by up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70%, 75%, 80%, 85%, 90%, 95% or 100%. These amounts are not meant to be limiting, and increments between the recited percentages are specifically envisioned as part of the invention. In some embodiments, the
amount of compound of the invention added to the edible composition is sufficient to permit reduction of the subject's sodium intake by up to 25%. In other embodiments, the amount of compound of the invention added to the edible composition is sufficient to permit reduction of the subject's sodium intake by up to 50%. In other embodiments, the amount of compound of the invention added to the edible composition is sufficient to permit reduction of the subject's sodium intake by up to 75%. In yet other embodiments, the amount of compound of the invention added to the edible composition is sufficient to permit reduction of the subject's sodium intake by up to 100%.

[0310] In some embodiments, the method of reducing sodium intake of a subject further comprises adding one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or flavor modifiers, which may lack an inherent flavor.

Method of reducing sugar intake of a subject

[0311] According to another embodiment, the invention provides a method of reducing sugar intake of a subject. In some embodiments, the method comprises the step of providing an edible composition of the present invention to the subject, wherein all or a portion of the sugar in the edible composition is replaced with Acesulfame K, and wherein the edible composition comprises a compound of the present invention. In some embodiments, the edible composition is a food product. In some embodiments, the edible composition is a pharmaceutical composition. In some embodiments, the edible composition is a consumer product.

[0312] In some embodiments, the methods of reducing sugar intake of a subject further comprise the step of identifying a subject in need thereof. The skilled worker would be able to identify a subject in need of reducing sugar intake. Non-limiting examples of such subjects include subjects that suffer from any one or more of the following disorders: diabetes, pre-diabetes, insulin resistance, obesity, excessive weight, and hyperglycemia.

[0313] In some embodiments, the amount of sugar replaced by Acesulfame K in the edible composition is an amount sufficient to maintain or restore the health of a subject. In some embodiments, the amount of sugar replaced by Acesulfame K in the edible composition is an amount sufficient to result in weight loss in a subject. In some embodiments, the amount of sugar replaced by Acesulfame K in the edible composition is an amount to sufficient to alleviate the effects of, or treat, a disease associated with sugar consumption or excessive weight of the subject (e.g., diabetes). In some embodiments, the amount of sugar replaced by Acesulfame K in the edible composition is up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70%, 75%, 80%, 85%, 90%, 95% or 100%. These amounts are not meant to be limiting, and increments between the recited percentages are specifically envisioned as part of the invention. In some embodiments, the subject's daily sugar intake is less than 250 g/day, less than 200 g/day, less than 175 g/day, less than 150 g/day, less than 125 g/day, less than 100 g/day, less than 75 g/day, less than 50 g/day or less than 25 g/day.
[0314] In some embodiments, the amount of compound of the invention added to the edible composition is sufficient to permit reduction of a subject's sugar intake by up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70%, 75%, 80%, 85%, 90%, 95% or 100%. These amounts are not meant to be limiting, and increments between the recited percentages are specifically envisioned as part of the invention. In some embodiments, the amount of compound of the invention added to the edible composition is sufficient to permit reduction of a subject's sugar intake by up to 25%. In other embodiments, the amount of compound of the invention added to the edible composition is sufficient to permit reduction of a subject's sugar intake by up to 50%. In other embodiments, the amount of compound of the invention added to the edible composition is sufficient to permit reduction of a subject's sugar intake by up to 75%. In yet other embodiments, the amount of compound of the invention added to the edible composition is sufficient to permit reduction of a subject's sugar intake by up to 100%.

[0315] In some embodiments, the method of reducing sugar intake of a subject further comprises adding one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or flavor modifiers, which may lack an inherent flavor.

Method of reducing bitter taste of an edible composition

[0316] According to another embodiment, the invention provides methods of reducing the bitter taste in an edible composition. In some embodiments, the edible composition is a food product. In some embodiments, the edible composition is a pharmaceutical composition. In some embodiments, the edible composition is a consumer product.

[0317] In one embodiment, the method comprises: (a) adding an effective amount of a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (Xb) or Formula (XVe), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof, to an edible composition such that bitter taste is reduced.

[0318] In alternate embodiments, the method comprises: (a) ingesting an effective amount of a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (Xb) or Formula (XVe), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof, before, along with, or after the edible composition such that bitter taste is reduced.

[0319] In some embodiments, the bitter tastant is a bitter tasting salt. In some embodiments, the bitter tastant is a potassium salt, a magnesium salt, or a calcium salt. In some embodiments, the
bitter tastant is a potassium salt. In some embodiments, the bitter tastant is KCl. In other embodiments, the bitter tastant is potassium lactate. In some embodiments, the bitter tastant is inherent in the edible composition, such as in an inherently bitter foodstuff.

[0320] In some embodiments, the bitter taste is reduced by up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70%, 75%, 80%, 85%, 90%, 95% or 100%. These amounts are not meant to be limiting, and increments between the recited percentages are specifically envisioned as part of the invention. In some embodiments, the bitter taste is reduced by up to 25%. In other embodiments, the bitter taste is reduced by up to 50%. In other embodiments, the bitter taste is reduced by up to 75%. In other embodiments, the bitter taste is reduced by up to 100%.

[0321] In some embodiments, the method of reducing the bitter taste attributed to a bitter tastant in an edible composition further comprises adding one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or flavor modifiers, which may lack an inherent flavor.

Method of preserving an edible composition

[0322] According to another embodiment, the invention provides a method of preserving an edible composition or extending the shelf life of an edible composition comprising:

(a) providing an edible composition; and

(b) combining with the edible composition of (a) a preservative and an effective amount of compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb"), Formula (IV), Formula (Va), Formula (Vb), Formula (VIa), Formula (VIb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIa), Formula (XIIb), Formula (XIIa), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XV a), Formula (XV b) or Formula (XVc), as described herein, or combinations thereof.

[0323] In another embodiment, the method of preserving or extending the shelf life of an edible composition comprises:

(a) providing an edible composition; and

(b) combining with the edible composition of (a) a preservative and an effective amount of any one of Compounds 1-58, or combinations thereof.

[0324] According to the invention, the preservative can be any bitter-tasting preservative. In some embodiments, the preservative in (a) is a potassium salt. In some embodiments, the preservative in (a) is potassium lactate.

[0325] In some embodiments, the edible composition is a food product. In some embodiments, the edible composition is a pharmaceutical composition. In some embodiments, the edible composition is a consumer product.

[0326] In some embodiments, the method of preserving an edible composition further comprises adding one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or flavor modifiers, which may lack an inherent flavor.
Method of reducing the amount of sodium in an edible composition while preserving the edible composition

[0327] According to another embodiment, the invention provides a method of reducing the amount of sodium in an edible composition while preserving the edible composition. In some embodiments, the method comprises replacing an amount of sodium containing preservative present in an edible composition with an amount of potassium containing preservative and adding an effective amount of a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vib), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVIb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof.

[0328] In some embodiments, the method comprises replacing an amount of sodium lactate present in an edible composition with an amount of potassium lactate and adding an effective amount of a compound of Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vib), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVIb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof.

[0329] In some embodiments, the edible composition is a food product. In some embodiments, the edible composition is a pharmaceutical composition. In some embodiments, the edible composition is a consumer product.

[0330] In some embodiments, the effective amount of the compound is sufficient to permit reduction of the amount of sodium lactate typically present in an edible composition by up to 1%, 2%, 3%, 4%, 5%, 6%, 7%, 8%, 9%, 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70%, 75%, 80%, 85%, 90%, 95% or 100%. These amounts are not meant to be limiting, and increments between the recited percentages are specifically envisioned as part of the invention. In some embodiments, the effective amount of the compound is sufficient to permit reduction of the amount of sodium lactate typically present in an edible composition by up to 25%. In other embodiments, the effective amount of the compound is sufficient to permit reduction of the amount of sodium lactate typically present in an edible composition by up to 50%. In other embodiments, the effective amount of the compound is sufficient to permit reduction of the amount of sodium lactate typically present in an edible composition by up to 75%. In yet other embodiments, the effective amount of the compound is sufficient to permit reduction of the amount of sodium lactate typically present in an edible composition by up to 100%.

[0331] In some embodiments, the method of reducing the bitter taste attributed to a bitter tastant in an edible composition further comprises adding one or more additional components selected from the group consisting of preservatives, nutritives, flavorants or flavor modifiers, which may
lack an inherent flavor. In some embodiments, the method of reducing the amount of sodium lactate in an edible composition while preserving the food product further comprises adding one or more additional flavor modifiers.

**Method of inhibiting a bitter taste receptor**

[0332] According to another embodiment, the invention provides a method of inhibiting or reducing activation and/or signaling of a bitter taste receptor. In some embodiments, the method comprises contacting a bitter taste receptor with a compound according to Formula (I), Formula (Ia), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVIIa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof.

[0333] In some embodiments, the method comprises contacting a bitter taste receptor with an edible composition comprising a compound according to Formula (I), Formula (Ia), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVIIa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described above, or combinations thereof.

[0334] In some embodiments, the edible composition is a food product. In some embodiments, the edible composition is a pharmaceutical composition. In some embodiments, the edible composition is a consumer product.

[0335] In some embodiments, the bitter taste receptor is an *ex vivo* receptor present in, for example, an assay. In some embodiments, the bitter taste receptor is an *in vitro* receptor present in, for example, an assay. In other embodiments, the bitter taste receptor is an *in vivo* receptor present in a subject. In some embodiments, the bitter taste receptor is present in the oral cavity or gastrointestinal tract of a subject. In some embodiments, the bitter receptor is in the oral cavity of a human. In some embodiments, the bitter receptor is in the oral cavity of a non-human animal. In some embodiments, the bitter receptor is in the oral cavity of an animal model.

**Preparation of the Compounds of the Invention**

[0336] In some embodiments, one or more of the compounds of Formula (I), Formula (Ia), Formula (IIb), Formula (IIIb), Formula (IIIb'), or Formula (IIIb''), as described herein, is commercially available, for example from commercial sources such as ChemBridge Corporation of San Diego, California, USA; Sigma-Aldrich® of St. Louis, Missouri, USA; TCI America, Portland, Oregon, USA; and Acros Organics, Geel, Belgium; among others.
In other embodiments, one or more of the compounds of Formula (I), Formula (Ia), Formula (IIb), Formula (IIIb'), or Formula (IIIb'') is prepared from commercially available reagents by routine methods in synthetic organic chemistry.

In one embodiment, one or more compounds of Formula (I) or Formula (Ia), wherein X is O, is prepared by nucleophilic displacement of leaving group LG of A2 with the phenoxide anion of A1, generated under basic conditions, to give ether product P1 (Scheme I):

![Scheme I](image)

Suitable leaving groups include those recognized in the art, such as halide (e.g., chloro, bromo, iodo), triflate, mesylate, tosylate, and the like. Suitable bases include those recognized in the art for such reactions, and include but are not limited to alkaline and alkaline earth metal hydroxides (such as NaOH, LiOH, etc.), carbonates (such as Na₂CO₃, K₂CO₃, CaCO₃, etc.), and bicarbonates (such as NaHCO₃, KHCO₃, etc.). Other suitable bases include amine bases, such as ammonia, ammonium hydroxide, triethylamine, pyridine, piperidine, pyrrolidine, 2,6-lutidine, 1,8-diazabicycloundec-7-ene (DBU), 4-(dimethylamino)-pyridine, etc.

In another embodiment, one or more compounds of Formula (I) or Formula (Ia), wherein X is NR°° and R°° is absent, is prepared by imine formation between phenylamine A3 and aldehyde A4, under conditions known in the art to give product P2, for example, conditions employing dehydrating agents, such as molecular sieves (Scheme II):

![Scheme II](image)

In certain embodiments, one or more compounds of Formula (I) or Formula (Ib) is prepared by reduction of imine P2 to generate amine P3 (Scheme III):

![Scheme III](image)
Suitable reducing conditions include those known in the art for reducing imines and iminium ions, such as hydrogenolysis with hydrogen and palladium, such as palladium on carbon. Another suitable source of hydrogen includes formic acid.

In further embodiments, one or more compounds of Formula (I), wherein X is NR and R is not absent, is prepared by reductive alkylation of amine P3 in the presence of the corresponding aldehyde RCHO to form product P4, wherein R is –CH(R) (Scheme IV):

```
R
O

P2

\[ \text{RCHO} \rightarrow \text{P4} \text{ (red.)} \]
```

Suitable reductive alkylation conditions include those known in the art for reducing imines and iminium ions, such as hydrogenolysis with hydrogen and palladium, such as palladium on carbon. Another suitable source of hydrogen includes formic acid.

In some embodiments, one or more of the compounds of Formula (IV), Formula (Va), Formula (Vib), Formula (VIIa), Formula (Vib), Formula (VIIb), or Formula (VIIb), as described herein, is commercially available, for example from commercial sources such as ChemBridge Corporation of San Diego, California, USA; Sigma-Aldrich® of St. Louis, Missouri, USA; TCI America, Portland, Oregon, USA; and Aeros Organics, Geel, Belgium; among others.

In other embodiments, one or more of the compounds of Formula (IV) is prepared from commercially available reagents by routine methods in synthetic organic chemistry.

In one embodiment, one or more compounds of Formula (IV), is prepared by acylation of amine A12 with acyl compound A11 bearing leaving group LG to afford amide P11 (Scheme V):

```
(\text{R}^1_n)\text{N} = \text{O} \quad + \quad \text{R}^3\text{HN}(\text{R}^4_m)\text{N} \rightarrow (\text{R}^1_n)\text{N} = \text{O} \quad + \quad \text{R}^3\text{HN}(\text{R}^4_m)\text{N}
```

Suitable leaving groups include those recognized in the art for acylation reactions, such as halide (e.g., chloro, bromo, iodo), alkoxy, aryloxy, activated leaving groups, and the like. In some embodiments, acylation conditions also employ an inorganic or organic base. Suitable bases include those recognized in the art for such reactions, and include but are not limited to alkaline and alkaline earth metal carbonates (such as Na₂CO₃, K₂CO₃, CaCO₃, etc.) and bicarbonates (such as NaHCO₃, KHCO₃, etc.). Other suitable bases include aprotic amine bases, such as triethylamine, pyridine, 2,6-lutidine, 1,8-diazabicyclocoundec-7-ene (DBU), 4-(dimethylamino)-pyridine, etc.

In one particular embodiment, compound A11 is an acid halide, such as an acid chloride or bromide, and the acylation reaction proceeds in the presence of an aprotic amine base, such as...
triethylamine, pyridine, 2,6-lutidine, 1,8-diazacycloundecene-7-ene (DBU), 4-(dimethylamino)-pyridine.

[0350] Compound A11 can be prepared from the corresponding carboxylic acid using routine methods known in the art.

[0351] In some embodiments, one or more of the compounds of Formula (VIII), Formula (IX), or Formula (X), as described herein, is commercially available, for example from commercial sources such as ChemBridge Corporation of San Diego, California, USA; Sigma-Aldrich® of St. Louis, Missouri, USA; TCI America, Portland, Oregon, USA; and Acros Organics, Geel, Belgium; among others.

[0352] In other embodiments, one or more of the compounds of Formula (VIII), Formula (IX), or Formula (X) is prepared from commercially available reagents by routine methods in synthetic organic chemistry.

[0353] In one embodiment, one or more compounds of Formula (VIII), Formula (IX), or Formula (X) is prepared by a multi-step sequence beginning with condensing amine A21 and aryl or heteroaryl aldehyde A22 to afford imine (when R₂ is H in A21) or iminium ion (when R₂ is not H in A21) P21, which then undergoes [4+2] cycloaddition with cyclic alkene A23 followed by rearomatization to afford fused tricyclic system P22 (Scheme VI):

![Chemical structure diagram]

Scheme VI

[0354] Suitable conditions for imine or iminium ion formation may employ dehydrating agents, such as molecular sieves.

[0355] Suitable cycloaddition conditions may include heating, for example, up to at least about 50, 75, 100, 120, 150 °C or greater. In some embodiments, cycloaddition conditions include the use of Lewis acids, for example boron compounds (e.g., Bu₃BOTf or BF₃·Et₂O), titanium compounds (e.g., TiCl₄ or titanium alkoxides), aluminum compounds (e.g., AlCl₃ or aluminum alkoxides), silicon compounds (e.g., trialkyldisilyl triflates, such as TMS-OTf, trialkylsilyl halides, etc.), and the like, particularly if ring Cy includes an electron withdrawing group (e.g., esters, ketones, aldehydes, cyan, nitro, etc.) in conjugation with the olefin of Cy.

[0356] In certain instances, rearomatization is assisted by the use of a base. Suitable bases include those recognized in the art for such reactions, and include but are not limited to alkaline and alkaline earth metal hydroxides (such as NaOH, LiOH, etc.), carbonates (such as Na₂CO₃, K₂CO₃, CaCO₃, etc.), and bicarbonates (such as NaHCO₃, KHCO₃, etc.). Other suitable bases include amine bases, such as ammonia, ammonium hydroxide, triethylamine, pyridine, piperidine, pyrrolidine, 2,6-lutidine, 1,8-diazabicycloundecene-7-ene (DBU), 4-(dimethylamino)-pyridine, etc.
[0357] In one particular embodiment, one or more compounds of Formula (VIII), Formula (IX), or Formula (X) is prepared first by formation of imine or iminium ion P21, as noted above, followed by [4+2] cycloaddition with cyclopentadiene A24 and then rearomatization, as noted above, to afford fused tricyclic system P23 (Scheme VII):

```
P21  \[\text{[R3]}_{m} N\text{Ar} \rightarrow \text{P23} \[\text{[R2]}_{o} \text{Ar} (\text{R1})_{n} \text{}] \]
```

Scheme VII

[0358] In some embodiments, one or more of the compounds of Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), or Formula (XIIIb), as described herein, is commercially available, for example from commercial sources such as ChemBridge Corporation of San Diego, California, USA; Sigma-Aldrich® of St. Louis, Missouri, USA; TCI America, Portland, Oregon, USA; and Acros Organics, Geel, Belgium; among others.

[0359] In other embodiments, one or more of the compounds of Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), or Formula (XIIIb) is prepared from commercially available reagents by routine methods in synthetic organic chemistry.

[0360] In one embodiment, one or more compounds of Formula (XI), Formula (XIIa), or Formula (XIIIa) is prepared by displacement of the leaving group LG of arylamine A31 with a nucleophilic group of heterocyclic compound A32 to afford product P31 (Scheme VIII):

```
A31 + :Het \[\text{[R5]}_{m} \text{R2} \rightarrow \text{P31} \[\text{[R1]}_{n} \text{L} \text{G} \text{Het} \]
```

Scheme VIII

[0361] Suitable leaving groups include those recognized in the art for such displacement reactions, such as halide (e.g., fluoro, chloro, bromo, iodo), alkoxy, arylxoy, triflate, mesylate, tosylate, and the like. In some embodiments, the displacement conditions also employ an inorganic or organic base. Suitable bases include those recognized in the art for such reactions, and include but are not limited to alkaline and alkaline earth metal hydroxides (such as NaOH, LiOH, etc.), carbonates (such as Na₂CO₃, K₂CO₃, CaCO₃, etc.), and bicarbonates (such as NaHCO₃, KHCO₃, etc.). Other suitable bases include amine bases, such as ammonia, ammonium hydroxide, triethylamine, pyridine, piperidine, pyrrolidine, 2,6-lutidine, 1,8-diazabicycloundec-7-ene (DBU), 4-(dimethylamino)-pyridine, etc. In some instances, suitable bases include strong bases such as
alkoxides (such as sodium or potassium tert-butoxide), lithium disopropyl amide (LDA), lithium bis(trimethylsilyl)amide (LiHMDS), sodium bis(trimethylsilyl)amide (NaHMDS), and the like.

In one particular embodiment, one or more compounds of Formula (XI), Formula (XIIa), or Formula (XIIIa) is prepared by displacement of the leaving group LG of arylamine A33 with an amine of heterocyclic compound A34 under strongly basic conditions to afford product P32 (Scheme IX):

![Scheme IX](attachment:image.png)

Suitable leaving groups include those recognized in the art for such displacement reactions, such as halide (e.g., fluoro, chloro, bromo, iodo), alkoxy, aryloxy, triflate, mesylate, tosylate, and the like. In some embodiments, suitable basic conditions employ an inorganic or organic base. Suitable strong bases include alkoxides (such as sodium or potassium tert-butoxide), lithium disopropyl amide (LDA), lithium bis(trimethylsilyl)amide (LiHMDS), sodium bis(trimethylsilyl)amide (NaHMDS), and the like.

In another embodiment, one or more compounds of Formula (XI), Formula (XIIb), or Formula (XIIIb) is prepared by displacement of the leaving group LG of arylamine A35 with a nucleophilic group of heterocyclic compound A32 to afford product P33 (Scheme X):

![Scheme X](attachment:image.png)

Suitable leaving groups include those recognized in the art for such displacement reactions, such as halide (e.g., chloro, bromo, iodo), alkoxy, aryloxy, triflate, mesylate, tosylate, and the like. In some embodiments, the displacement conditions also employ an inorganic or organic base. Suitable bases include those recognized in the art for such reactions, and include but are not limited to alkaline and alkaline earth metal hydroxides (such as NaOH, LiOH, etc.), carbonates (such as Na₂CO₃, K₂CO₃, CaCO₃, etc.), and bicarbonates (such as NaHCO₃, KHCO₃, etc.). Other suitable bases include amine bases, such as ammonia, ammonium hydroxide, triethylamine, pyridine, piperidine, pyrrolidine, 2,6-lutidine, 1,8-diazabicyclo[5.4.0]undec-7-ene (DBU), 4-(dimethylamino)-pyridine, etc. In some instances, suitable bases include strong bases such as alkoxides (such as sodium or potassium tert-butoxide), lithium disopropyl amide (LDA), lithium bis(trimethylsilyl)amide (LiHMDS), sodium bis(trimethylsilyl)amide (NaHMDS), and the like.
In a particular embodiment, one or more compounds of Formula (XI), Formula (XIIb), or Formula (XIIIb) is prepared by displacement of the leaving group LG of arylamine A35 with an amine of heterocyclic compound A36 under basic conditions to afford product P34 (Scheme XI):

![Scheme XI]

Suitable leaving groups include those recognized in the art for such displacement reactions, such as halide (e.g., chloro, bromo, iodo), alkoxy, aryloxy, triflate, mesylate, tosylate, and the like. In some embodiments, suitable basic conditions employ an inorganic or organic base. Suitable bases include those recognized in the art for such reactions, and include but are not limited to alkaline and alkaline earth metal hydroxides (such as NaOH, LiOH, etc.), carbonates (such as Na₂CO₃, K₂CO₃, CaCO₃, etc.), and bicarbonates (such as NaHCO₃, KHCO₃, etc.). Other suitable bases include amine bases, such as ammonia, ammonium hydroxide, triethylamine, pyridine, piperidine, pyrrolidine, 2,6-lutidine, 1,8-diazabicycloundec-7-ene (DBU), 4-(dimethylamino)-pyridine, etc. In some instances, suitable bases include strong bases such as alkoxides (such as sodium or potassium tert-butoxide), lithium diisopropyl amide (LDA), lithium bis(trimethylsilyl)amide (LiHMDS), sodium bis(trimethylsilyl)amide (NaHMDS), and the like.

In some embodiments, one or more compounds of Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), or even compounds A31, A33, or A35 is prepared by a transition metal-catalyzed coupling reaction of compound A37, bearing leaving group LG, with metallo- or boro-compound A38, wherein M is a metal or boron group, under the appropriate coupling conditions known in the art, to afford coupling product P35 (Scheme XII):

![Scheme XII]

Alternatively, one or more compounds of Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), or even compounds A31, A33, or A35 is prepared by a transition metal-catalyzed coupling reaction of metallo- or boro-compound A39, wherein M is a metal or boron group, with compound A310, bearing leaving group LG, under the appropriate coupling conditions known in the art, to afford coupling product P35 (Scheme XIII):

![Scheme XIII]
[0370] Suitable transition metal catalysts include those derived from palladium, such as Pd(PPh3)4, or other noble transition metals. Suitable leaving groups include those recognized in the art, such as halides (e.g., chloro, bromo, iodo), triflates, mesylates, tosylates, and the like. Suitable metal groups include tin, zinc, magnesium, copper, or other metals known to undergo transmetallation with palladium or other noble transition metals.

[0371] In certain embodiments, substituted quinoline compounds, such as A35, A37, A39, P33, P34, and P35 may be prepared by methods known in the art, for example, such as those described in U.S. Patent No. 6,297,258, which is incorporated by reference herein.

[0372] In some embodiments, one or more of the compounds of Formula (XIV), Formula (XVa), Formula (XVb), or Formula (XVc), as described herein, is commercially available, for example from commercial sources such as ChemBridge Corporation of San Diego, California, USA; Sigma-Aldrich® of St. Louis, Missouri, USA; TCI America, Portland, Oregon, USA; and Acros Organics, Geel, Belgium; among others.

[0373] In other embodiments, one or more of the compounds of Formula (XIV), Formula (XVa), Formula (XVb), or Formula (XVc) is prepared from commercially available reagents by routine methods in synthetic organic chemistry.

[0374] In one embodiment, one or more compounds of Formula (XIV) is prepared by acylation of amine A41 with acyl compound A42 bearing leaving group LG to afford product P41 (Scheme I):

\[
\begin{align*}
\text{A41} & \quad \text{L}^* \quad \text{R}^2 \\
\begin{array}{c}
\text{(R^1)_n} \\
\text{NH}
\end{array} & \quad \begin{array}{c}
\text{C}
\end{array} & \quad \begin{array}{c}
\text{O}
\end{array} & \quad \begin{array}{c}
\text{R}^2
\end{array}
\end{align*}
\]

Scheme XIV

[0375] Suitable leaving groups include those recognized in the art for acylation reactions, such as halide (e.g., chloro, bromo, iodo), alkoxy, aryloxy, leaving groups associated with activated esters (e.g., N-succinimide), and the like. In certain embodiments, acyl compound A42 is an acid anhydride; that is LG is -OC(O)R'. In some embodiments, acylation conditions also employ an inorganic or organic base. Suitable bases include those recognized in the art for such reactions, and include but are not limited to alkali and alkaline earth metal hydroxides (such as NaOH, LiOH, etc.), carbonates (such as Na2CO3, K2CO3, CaCO3, etc.), and bicarbonates (such as NaHCO3, KHCO3, etc.). Other suitable bases include amine bases, such as ammonia, ammonium hydroxide, triethylamine, pyridine, pipieridine, pyrroldine, 2,6-lutidine, 1,8-diazabicyclo[5.4.0]undec-7-ene (DBU), 4-(dimethylamino)-pyridine, etc.

[0376] In one particular embodiment, compound A42 is an acid halide, such as an acid chloride or bromide, and the acylation reaction proceeds in the presence of an amine base, such as triethylamine, pyridine, pipieridine, pyrroldine, 2,6-lutidine, 1,8-diazabicyclo[5.4.0]undec-7-ene (DBU), 4-(dimethylamino)-pyridine, etc.

[0377] In another embodiment, compound A42 is an activated ester and acylation proceeds under mild conditions that do not result in the generation of strong acids.
Compound A42 can be prepared from the corresponding carboxylic acid using routine methods known in the art.

In certain embodiments, compound A41, wherein R is -CH2R, is prepared in a two-step sequence, first by imine formation between amine A43 and aldehyde A44 to give imine P42, second by reduction of imine P42 to give A41 (Scheme XV):

Scheme XV

Suitable conditions for imine formation may employ dehydrating agents, such as molecular sieves. Suitable reducing conditions include those known in the art for reducing imines and inminimum ions, such as hydrogenolysis with hydrogen and palladium, such as palladium on carbon. Another suitable source of hydrogen includes formic acid.

The skilled artisan will appreciate that aryl and/or heteroaryl, alkynyl, alkynyl, aralkyl, heteroaralkyl, allyl, and propargyl moieties herein may be readily coupled directly using Stille, Suzuki, Heck, Negishi, Sonogashira, Kumada, Glaser, or other related reactions, such as palladium-mediated cross-coupling reactions. Aryl and/or heteroaryl moieties herein may also be readily coupled through a heteroatom, e.g., using reactions such as the Ullmann reaction, any of various palladium-mediated reactions developed by S. Buchwald and others, by nucleophilic aromatic substitution, or other such reactions. Similarly, amines, alcohols, thiols, and other such heteroatom-bearing compounds herein may be coupled to aryl and/or heteroaryl moieties using palladium-mediated reactions developed by S. Buchwald and others, nucleophilic aromatic substitution, etc. Aryl and/or heteroaryl moieties linked by substituted or unsubstituted hydrocarbon chains herein may also be prepared by Stille, Suzuki, Heck, Friedel-Crafts, and other reactions as will be apparent to those of skill in the art.

It will be understood that the various substituents on the compounds in the above syntheses can be protected from the reaction conditions as necessary using the proper protecting groups, such as those disclosed in Greene, T.W.; Wuts, P.G.M. Greene's Protective Groups in Organic Synthesis, 4th ed.; Wiley-Interscience: New York, 2006.

Examples

In order that this invention be more fully understood, the following examples are set forth. These examples are for the purpose of illustration only and are not to be construed as limiting the scope of the invention in any way.

The test compounds used in the following examples were obtained from commercial vendors for synthetic and natural compounds, including VitasM, ChemDiv, ChemBridge, Chromadex, Sigma Aldrich, Penta, Spectrum Chemical, Vigon, and Indofine.
The taste test panelists used in the following examples were screened based upon and selected for their ability to perceive the bitter taste associated with potassium chloride. Only panelists capable of perceiving bitter taste participated in the following taste tests.

Due to the complex nature of taste perception in subjects and the inherently subjective nature of the following experiments, individual taste test trials may yield different results for a given compound. The data presented in the following Examples is illustrative of the taste testing results observed. It is noted that the data presented in the Figures represents a subset of the data presented in the Examples below.

The taste testing experiments below were conducted with panels of varying size (i.e., panels comprising varying numbers of panelists).

**Example 1**  Generation of KCl Test Solutions.

Edible KCl solution compositions ("KCl test solutions") were prepared by first dissolving varying amounts of the test compounds in an amount of ethanol or water (depending on the solubility of the compound) to create a 5mg/mL stock compound solution. An amount of this stock compound solution is then added to an aqueous KCl solution. Enough EtOH is then added to the resulting stock compound/KCl solution so that the final KCl test solution contains 1% EtOH. KCl solution standards were similarly prepared by dissolving various amounts of KCl in water and ethanol without adding any test compound. NaCl solution standards were similarly prepared by dissolving various amounts of NaCl in water and ethanol without adding any test compound (NaCl solution standards did not contain any KCl).

**Table 1. KCl Taste Test Solutions**

<table>
<thead>
<tr>
<th>Compound No.</th>
<th>Conc. Of KCl</th>
<th>Conc. of Compound Tested (ppm)</th>
<th>Conc. Where Decrease in Bitter Taste Was Discerned (ppm)</th>
<th>Conc. Where Decrease in Bitter Taste Discerned and p&lt;0.1 (ppm)</th>
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<td>1; 10</td>
<td>1; 10</td>
<td>1; 10</td>
</tr>
<tr>
<td>2</td>
<td>4.85 g/L</td>
<td>1; 10</td>
<td>1; 10</td>
<td></td>
</tr>
<tr>
<td>3</td>
<td>4.85 g/L</td>
<td>2</td>
<td>2</td>
<td></td>
</tr>
<tr>
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<td>4.85 g/L</td>
<td>1; 10</td>
<td>1; 10</td>
<td>10</td>
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<td>10</td>
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<td>1; 10</td>
<td>1</td>
<td></td>
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<td>1; 10</td>
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<td>1</td>
<td></td>
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<td>1; 10</td>
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</tr>
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<td>4.85 g/L</td>
<td>1; 10</td>
<td>--</td>
<td></td>
</tr>
<tr>
<td>Compound No.</td>
<td>Conc. Of KCl (g/L)</td>
<td>Conc. of Compound Tested (ppm)</td>
<td>Conc. Where Decrease in Bitter Taste Was Discerned (ppm)</td>
<td>Conc. Where Decrease in Bitter Taste Discerned and p&lt;0.1 (ppm)</td>
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<td>--------------------------------------------------------</td>
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<td>1; 10</td>
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**Example 2**  
**Effect of test compounds on the perception of bitter taste of aqueous KCl solutions in humans.**

[0389] The effect of the test compounds on the perception of the bitter taste of an aqueous solution of KCl in humans was evaluated using a "sip and spit" test as follows.

[0390] A set of KCl solution standards was developed and each standard solution was assigned a bitterness taste score of 0-15 (corresponding to aqueous KCl concentrations of 0 mM-120 mM). Panelists were trained to recognize these standards. In addition, before each day of testing,
panelists were tested to see if they could determine differences in taste between the standard solutions. If a panelist was unable to recognize a change in KCl concentration, they were excluded from the panel for that day.

[0391] In a blind taste test, panelists were asked to compare the bitter taste of a small quantity (e.g., 8 ml) of each of the KCl Test Solutions to the taste of a KCl solution standard, without swallowing (see, e.g., Table 1). Specifically, panelists were asked to rate the bitterness of each KCl Test Solution on a scale of 0-15 using the same scale developed for the KCl solution standards. Each sample was tested in 2-4 discrete taste test experiments. Panelists were asked to rinse with water, eat a cracker, and wait approximately 10 minutes between samples.

[0392] Illustrative results of the aqueous solution testing are presented in Figures 1-5 and Table 1.

Example 3  Generation of Potassium Lactate Test Solutions.

[0393] Edible potassium lactate solution compositions ("potassium lactate test solutions") were prepared by first dissolving varying amounts of the test compounds in an amount of ethanol or water (depending on the solubility of the compound) to create a 5mg/mL stock compound solution. An amount of this stock compound solution is then added to an aqueous potassium lactate solution. Enough EtOH is then added to the resulting stock compound/potassium lactate solution so that the final potassium lactate test solution contains 1% EtOH. Potassium lactate solution standards were similarly prepared by dissolving various amounts of potassium lactate in water and ethanol without adding any test compound. Sodium lactate solution standards were similarly prepared by dissolving various amounts of sodium lactate in water and ethanol without adding any test compound (sodium lactate solution standards did not contain any potassium lactate).

Table 2. Potassium Lactate Taste Test Solutions

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<th>Compound No.</th>
<th>Conc. Of KLac</th>
<th>Conc. of Compound Tested (ppm)</th>
<th>Conc. Where Decrease in Bitter Taste Was Discerned (ppm)</th>
<th>Conc. Where Decrease in Bitter Taste Discerned and p≤0.1 (ppm)</th>
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**Example 4**  Effect of test compounds on the perception of bitter taste of aqueous potassium lactate solutions in humans.

The effect of the test compounds on the perception of the bitter taste of an aqueous solution of potassium lactate in humans was evaluated using the "sip and spit" test described in Example 2.

Illustrative results of the aqueous solution testing are presented in Figures 1-5 and in Table 2.

**Example 5**  Generation of KCl Test Foodstuff Slurries.

Edible KCl food compositions ("KCl test foodstuff slurries") were prepared as follows. Dehydrated, salt-free turkey powder was weighed and mixed with various amounts of KCl and/or NaCl and then solubilized with boiling water to create a homogenized solubilized turkey slurry. Varying amounts of the test compounds were dissolved in an amount of ethanol or water (depending on the solubility of the compound) to create a 5mg/mL stock compound solution. An amount of this stock compound solution was then added to the turkey slurry. Enough EtOH is then added to the resulting stock compound/turkey slurry so that the slurry contains 1% EtOH. The slurry was again homogenized by boiling and mixing and allowed to cool to yield the final KCl test foodstuff slurry for taste testing. KCl foodstuff slurry standards were similarly prepared without any test compound. NaCl foodstuff slurry standards were similarly prepared without adding any test compound (NaCl foodstuff slurry standards did not contain any KCl).
Table 3. KCI Foodstuff Slurry Compositions

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</table>
Example 6  Effect of test compounds on the perception of bitter taste of KCl foodstuff slurries in humans using a two-alternative forced choice method (2AFC).

[0397] The effect of the test compounds on the perception of the bitter taste of KCl foodstuff slurries in humans was evaluated using a two-alternative-forced-choice "sip and spit" test as follows.

[0398] In a blind taste test, panelists received two portions of turkey slurry – one portion being the KCl foodstuff slurry standard and the other being one of the KCl test foodstuff slurries (each prepared as described in Example 5). The panelists tasted each of the portions by sipping and spitting. Each sample was tested in 2-4 discrete taste test experiments. Panelists were asked to rinse with water, eat a cracker, and wait about 10 minutes between samples. In each case, the panelists were asked to compare the bitter taste of the two turkey samples to each other (i.e. panelists were asked to indicate which sample was less bitter).

[0399] Illustrative results of the foodstuff testing are presented in Figures 1-5 and in Table 3.

Example 7  Generation of Potassium Lactate Test Foodstuff Slurries.

[0400] Edible potassium lactate food compositions ("potassium lactate test foodstuff slurries") were prepared as follows. Dehydrated, salt-free turkey powder was weighed and mixed with various amounts of potassium lactate and/or sodium lactate and then solubilized with boiling water to create a homogenized solubilized turkey slurry. Varying amounts of the test compounds were dissolved in an amount of ethanol or water (depending on the solubility of the compound) to create a 5mg/mL stock compound solution. An amount of this stock compound solution was then added to the turkey slurry. Enough EtOH is then added to the resulting stock compound/turkey slurry so that the final slurry contains 1% EtOH. The final slurry was again homogenized by boiling and mixing and allowed to cool to yield the final slurry for taste testing. Potassium lactate foodstuff slurry standards were similarly prepared without any test compound. Sodium lactate foodstuff slurry standards were similarly prepared without adding any test compound (sodium lactate foodstuff slurry standards did not contain any potassium lactate).

Table 4. Potassium Lactate Foodstuff Slurry Compositions

<table>
<thead>
<tr>
<th>Compound No.</th>
<th>Conc. Of KLac</th>
<th>Conc. of Compound Tested (ppm)</th>
<th>Conc. at Which At Least 50% of Panelist Discerned Decrease in Bitter Taste (ppm)</th>
<th>Conc. at Which At Least 50% of Panelist Discerned Decrease in Bitter Taste and $p \leq 0.1$ (ppm)</th>
</tr>
</thead>
<tbody>
<tr>
<td>3</td>
<td>4.5%</td>
<td>1; 10; 30</td>
<td>1; 10; 30</td>
<td>--</td>
</tr>
<tr>
<td>23</td>
<td>4.5%</td>
<td>1; 10; 30</td>
<td>1; 30</td>
<td>1</td>
</tr>
<tr>
<td>24</td>
<td>4.5%</td>
<td>1; 10; 30</td>
<td>1; 10; 30</td>
<td>30</td>
</tr>
<tr>
<td>25</td>
<td>4.5%</td>
<td>1; 10; 30</td>
<td>1</td>
<td>--</td>
</tr>
<tr>
<td>26</td>
<td>4.5%</td>
<td>1; 10; 30</td>
<td>30</td>
<td>--</td>
</tr>
<tr>
<td>27</td>
<td>4.5%</td>
<td>1; 10; 30</td>
<td>1; 30</td>
<td>--</td>
</tr>
<tr>
<td>37</td>
<td>4.5%</td>
<td>1; 10; 30</td>
<td>1; 10; 30</td>
<td>10</td>
</tr>
<tr>
<td>38</td>
<td>4.5%</td>
<td>1; 10; 30</td>
<td>30</td>
<td>--</td>
</tr>
</tbody>
</table>

SUBSTITUTE SHEET (RULE 26)
<table>
<thead>
<tr>
<th>Compound No.</th>
<th>Conc. Of KLac</th>
<th>Conc. of Compound Tested (ppm)</th>
<th>Conc. at Which At Least 50% of Panelist Discerned Decrease in Bitter Taste (ppm)</th>
<th>Conc. at Which At Least 50% of Panelist Discerned Decrease in Bitter Taste and p≤0.1 (ppm)</th>
</tr>
</thead>
<tbody>
<tr>
<td>49</td>
<td>4.5%</td>
<td>30</td>
<td>30</td>
<td>30</td>
</tr>
<tr>
<td>51</td>
<td>4.5%</td>
<td>1; 10; 30</td>
<td>10</td>
<td>--</td>
</tr>
<tr>
<td>52</td>
<td>4.5%</td>
<td>1; 10; 30</td>
<td>10; 30</td>
<td>--</td>
</tr>
<tr>
<td>53</td>
<td>4.5%</td>
<td>30</td>
<td>30</td>
<td>--</td>
</tr>
<tr>
<td>55</td>
<td>4.5%</td>
<td>1; 10; 30</td>
<td>10</td>
<td>--</td>
</tr>
<tr>
<td>56</td>
<td>4.5%</td>
<td>1; 10; 30</td>
<td>1</td>
<td>1</td>
</tr>
<tr>
<td>Standard</td>
<td>4.5%</td>
<td>--</td>
<td>--</td>
<td>--</td>
</tr>
</tbody>
</table>

**Example 8**

Effect of test compounds on the perception of bitter taste of potassium lactate foodstuff slurries in humans using a two-alternative forced choice method (2AFC).

5

[0401] The effect of the test compounds on the perception of the bitter taste of potassium lactate foodstuffs in humans was evaluated using the two-alternative-forced-choice "sip and spit" test described in Example 6.

[0402] Illustrative results of the foodstuff testing are presented in Figures 1-5 and in Table 4.
We Claim:

1. A composition comprising a compound according to a formula selected from the group consisting of:
   
   (a) Formula (I):

   \[
   \begin{align*}
   (\text{R}^1)_n \text{X} \rightarrow \text{R}^2_m
   \end{align*}
   \]

   Formula (I);

   or a comestibly or biologically acceptable salt or derivative thereof,

   wherein, as valence and stability permit:

   \[
   \text{R}^1, \text{independently for each occurrence, is selected from the group consisting of halo; hydroxyl, C}_1\text{aalkyl, C}_1\text{haloalkyl, C}_1\text{hydroxyalkyl, or C}_1\text{acycloxy-C}_1\text{alkyl; C}_2\text{alkenyl; C}_2\text{alkynyl; C}_1\text{alkoxy; C}_1\text{alkylthio; and C}_6\text{10aryl-C}_1\text{alkyloxoy optionally substituted with halo, hydroxyl, C}_1\text{alkyl, C}_1\text{alkoxy, or C}_1\text{acycloxy;}
   \]

   \[
   \text{R}^2, \text{independently for each occurrence, is selected from the group consisting of halo; hydroxyl, C}_1\text{aalkyl, C}_1\text{haloalkyl, C}_1\text{hydroxyalkyl, or C}_1\text{acycloxy-C}_1\text{alkyl; C}_2\text{alkenyl; C}_2\text{alkynyl; C}_1\text{alkoxy; C}_1\text{alkylthio; and C}_6\text{10aryl-C}_1\text{alkyloxoy optionally substituted with halo, hydroxyl, C}_1\text{alkyl, C}_1\text{alkoxy, or C}_1\text{acycloxy;}
   \]

   \[
   \text{X is O or NR}^3, \text{wherein R}^3 \text{is absent or is selected from the group consisting of hydrogen and C}_1\text{alkyl;}
   \]

   wherein any of \text{R}^1, \text{R}^2, \text{and R}^3, \text{independently and independently for each occurrence, is optionally substituted with 1-3 substituents selected from the group consisting of C}_1\text{alcohol, C}_1\text{haloalkyl, halo, hydroxyl, carboxyl, C}_1\text{alkoxycarbonyl, C}_2\text{alkenylxoxycarbonyl, C}_2\text{alkynylxoxycarbonyl, C}_1\text{acycloxy, C}_1\text{acylamino, C}_1\text{acycloxy, C}_1\text{carbonate, C}_1\text{alkoxy, phenoxyl, phosphoryl, phosphat, phosphonate, phosphinate, amino, diC}_1\text{alkylamino, monoC}_1\text{alkylamino, C}_1\text{amido, C}_1\text{iminio, C}_1\text{carbamate, C}_1\text{urea, cyano, nitro, azido, sulfhydryl, C}_1\text{alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C}_3\text{carbocyclic, C}_3\text{carbocycle-C}_1\text{alkyl, C}_1\text{heterocyclic, C}_1\text{heterocycle-C}_1\text{alkyl, phenyl, phenyl-C}_1\text{alkyl, C}_1\text{heteroaryl, and C}_1\text{heteroaryl-C}_1\text{alkyl, and wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;}
   \]

   \[
   m \text{ is 1-3;}
   \]

   \[
   n \text{ is 0-3; and}
   \]
wherein the composition is edible and capable of reducing bitter taste of a bitter tastant

(b) Formula (IV):

\[
\begin{align*}
\text{Formula (IV);} & \\
\text{or a comestibly or biologically acceptable salt or derivative thereof,} & \\
\text{wherein, as valence and stability permit:} & \\
R^1, \text{ independently for each occurrence, is selected from the group consisting of} & \\
\text{halo, C}_1^1\text{-alkyl, C}_2^2\text{-alkenyl, and C}_2^2\text{-alkynyl;} & \\
R^2 & \text{is selected from the group consisting of hydrogen, C}_1^1\text{-alkyl, C}_2^2\text{-alkenyl,} & \\
& \text{C}_2^2\text{-alkynyl, and C}_1^1\text{-acyl;} & \\
R^3 & \text{is selected from the group consisting of hydrogen, C}_1^1\text{-alkyl, C}_2^2\text{-alkenyl, and} & \\
& \text{C}_2^2\text{-alkynyl;} & \\
R^4, \text{ independently for each occurrence, is selected from the group consisting of} & \\
\text{halo, C}_1^1\text{-alkyl, C}_2^2\text{-alkenyl, C}_2^2\text{-alkynyl, C}_1^1\text{-alkoxy, -C(O)-O-R}, & \\
& \text{and -C(O)-N(R')}_2; & \\
R^5, \text{ independently for each occurrence, is selected from the group consisting of} & \\
\text{hydrogen, C}_1^1\text{-alkyl, C}_2^2\text{-alkenyl, and C}_2^2\text{-alkynyl;} & \\
\text{wherein any of R}^1, R^2, R^3, \text{ and R}^4, \text{ independently and independently for each} & \\
\text{occurrence, is optionally substituted with 1-3 substituents selected from the group} & \\
\text{consisting of C}_1^1\text{-alkyl, C}_1^1\text{-haloalkyl, halo, hydroxyl, carboxyl, C}_1^1\text{-alkoxycarbonyl,} & \\
\text{C}_2^2\text{-alkenylxoxycarbonyl, C}_2^2\text{-alkynylxoxycarbonyl, C}_1^1\text{-alkyl, C}_1^1\text{-acylamino, C}_1^1\text{-acyloxy,} & \\
\text{C}_1^1\text{-carbonate, C}_1^1\text{-alkoxy, phenolxy, phosphonyl, phosphate, phosphonate, phosphinate,} & \\
\text{amino, diC}_1^1\text{-alkylamino, monoC}_1^1\text{-alkylamino, C}_1^1\text{-amino, C}_1^1\text{-carbamate,} & \\
\text{C}_1^1\text{-urea, cyano, nitro, azido, sulfhydryl, C}_1^1\text{-alkylthio, sulfate, sulfonate, sulfamoyl,} & \\
\text{sulfonamide, sulfonyl, C}_3^3\text{-carbocyclyl, C}_3^3\text{-carbocyclyl-C}_1^1\text{-alkyl, C}_1^1\text{-heterocyclyl,} & \\
\text{C}_1^1\text{-heterocyclyl-C}_1^1\text{-alkyl, phenyl, phenyl-C}_1^1\text{-alkyl, C}_1^1\text{-heteroaryl, and} & \\
\text{C}_1^1\text{-heteroaryl-C}_1^1\text{-alkyl; and wherein heterocyclic or heteroaromatic rings, independently} & \\
\text{for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;} & \\
\text{n is 0-2;} & \\
\text{m is 0-3; and} & \\
\text{wherein the composition is edible and capable of reducing bitter taste of a bitter} & \\
tastant. & 
\end{align*}
\]
(c) Formula (VIII):

or a comestibly or biologically acceptable salt or derivative thereof,

wherein, as valence and stability permit:

R¹, independently for each occurrence, is selected from the group consisting of halo, hydroxyl, C₁₋₃alkyl, C₁₋₂haloalkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₁₋₃alkoxy, and C₁₋₂acyloxy;

R², independently for each occurrence, is selected from the group consisting of halo, C₁₋₃alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C(O)-O-R⁴, and C(O)-N(R⁴)₂;

R³, independently for each occurrence, is selected from the group consisting of hydrogen, C₁₋₃alkyl, C₁₋₃alkenyl, and C₂₋₆alkynyl;

R⁴ is selected from the group consisting of hydrogen, C₁₋₃alkyl, C₂₋₆alkenyl, and C₂₋₆alkynyl;

Ar is selected from the group consisting of C₆₋₁₀ary1 and C₃₋₉heteroaryl;

Cy is a 5 to 7-membered carboyclic or heterocyclic ring, optionally including one or two carbon-carbon or carbon-nitrogen double bonds in the ring;

wherein any of R¹, R², R³, and R⁴, independently and independently for each occurrence, is optionally substituted with 1-3 substituents selected from the group consisting of C₁₋₁₀alkyl, C₁₋₁₀haloalkyl, halo, hydroxyl, carboxyl, C₁₋₁₀alkoxycarbonyl, C₁₋₁₀alkenylxoyxycarbonyl, C₂₋₁₀alkynloyxycarbonyl, C₁₋₁₀acyl, C₁₋₁₀cyclylmino, C₁₋₁₀acyloxy, C₁₋₁₀alkoxy, phenolxy, phosphoril, phosphate, phosphonate, phosphinate, amino, diC₁₋₁₀alkylamino, monoC₁₋₁₀alkylamino, C₁₋₁₀amido, C₁₋₁₀thioso, C₁₋₁₀carbamate, C₁₋₁₀urea, cyano, nitro, azido, sulhydryl, C₁₋₁₀alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C₃₋₅carbocyclyl, C₃₋₅carbocyclyl-C₁₋₃alkyl, C₁₋₃heterocyclyl, C₁₋₃heterocyclyl-C₁₋₃alkyl, phenyl, phenyl-C₁₋₃alkyl, C₁₋₃heteroaryl, and C₁₋₃heteroaryl-C₁₋₃alkyl; and wherein heterocyclic or
heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms
selected from N, O, and S;

\[ m \] is 1-3;
\[ n \] is 0-3;
\[ o \] is 0-3; and

wherein the composition is edible and capable of reducing bitter taste of a bitter
tastant

(d) Formula (XI):

![Chemical Structure](image)

or a comestibly or biologically acceptable salt or derivative thereof,

wherein, as valence and stability permit:

\[ R^1 \] is independently for each occurrence, is selected from the group consisting of
halo, hydroxyl, C\(_1\)alkyl, C\(_1\)haloalkyl, C\(_2\)alkenyl, C\(_2\)alkynyl, and C\(_1\)alkoxy;

\[ R^2 \] is selected from the group consisting of hydrogen, halo, hydroxyl, C\(_1\)alkyl,
C\(_1\)haloalkyl, Het-C\(_1\)alkyl, C\(_2\)alkenyl, C\(_2\)alkynyl, and C\(_1\)alkoxy;

\[ R^3 \] is selected from the group consisting of hydrogen, halo, hydroxyl, C\(_1\)alkyl,
C\(_1\)haloalkyl, Het-C\(_1\)alkyl, C\(_2\)alkenyl, C\(_2\)alkynyl, and C\(_1\)alkoxy;

\[ R^4 \] is selected from the group consisting of hydrogen, halo, hydroxyl, C\(_1\)alkyl,
C\(_1\)haloalkyl, Het-C\(_1\)alkyl, C\(_2\)alkenyl, C\(_2\)alkynyl, and C\(_1\)alkoxy;

or \[ R^3 \] and \[ R^4 \] together with the atoms to which they are attached form a 5 to
6-membered aryl ring optionally substituted with 1 to 4 groups selected from the group
consisting of halo, hydroxyl, C\(_1\)alkyl, C\(_1\)haloalkyl, C\(_2\)alkenyl, C\(_2\)alkynyl, C\(_1\)alkoxy,
and Het;

Het is a C\(_2\)heterocyclyl including 1-3 heteroatoms in the ring selected from
oxygen, sulfur, and nitrogen and is optionally substituted with one or more groups selected
from the group consisting of halo, hydroxyl, C\(_1\)alkyl, C\(_1\)haloalkyl, C\(_2\)alkenyl,
C\(_2\)alkynyl, C\(_1\)alkoxy, and C\(_6\)aryl optionally substituted with C\(_1\)alkyl;

wherein any of \[ R^1 \], \[ R^2 \], \[ R^3 \], \[ R^4 \], and Het, independently and independently for each
occurrence, is optionally further substituted with 1-3 substituents selected from the group
consisting of C\(_1\)-alkyl, C\(_1\)-haloalkyl, halo, hydroxyl, carboxyl, C\(_1\)-alkoxycarbonyl,
C_{2-10}alkenylxoycarbonyl, C_{2-10}alknyloxyccarbonyl, C_{1-10}acyl, C_{1-10}aclylamino, C_{1-10}acyloxy, C_{1-10}Carboneate, C_{1-10}alkoxy, phenolxy, phosphoryl, phosphaty, phosphonate, phosphinate, amino, diC_{1-10}alkylamino, monoC_{1-10}alkylaminoto, C_{1-10}amido, C_{1-10}imino, C_{1-10}carbamate, C_{1-10}aminoxy, cyano, nitro, azido, sulhydryl, C_{1-10}alkylthio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C_{3-7}carboxyclyl, C_{3-7}carboxyclyl-C_{1-4}alkyl, C_{1-4}heterocycl, C_{1-4}heterocycl-C_{1-4}alkyl, C_{6-10}aryl, C_{6-10}aryl-C_{1-4}alkyl, C_{6-10}aryl-C_{1-4}alkyl, C_{1-4}heteroaryl, and C_{1-4}heteroaryl-C_{1-4}alkyl; and wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteratoms selected from N, O, and S; and

n is 0-4; and

wherein the composition is edible and capable of reducing bitter taste of a bitter tastant; and

(c) Formula (XIV):

![Diagram](attachment:formula_xiv.png)

Formula (XIV);

or a comestibly or biologically acceptable salt or derivative thereof,

wherein, as valence and stability permit:

- R', independently for each occurrence, is selected from the group consisting of halo, hydroxyl, C_{1-6}alkyl, C_{1-6}haloalkyl, C_{2-6}alkenyl, C_{2-6}alkynyl, C_{1-6}alkoxy, and C_{1-6}acyloxy;

- R^2 is selected from the group consisting of C_{1-6}alkyl, C_{6-10}aryloxy-substituted C_{1-6}alkyl, C_{1-6}alkylkenyl, C_{2-6}alkynyl, C_{6-10}aryl-C_{1-6}alkyl, and -(CH_2)_{n+1}Ar, wherein aryl groups of R^2 are optionally substituted with one or more halo, hydroxyl, C_{1-6}alkyl, C_{1-6}haloalkyl, C_{1-6}alkoxy, or C_{1-6}acyloxy;

- R^2 is selected from the group consisting of hydrogen, C_{1-6}alkyl, C_{2-6}alkenyl, and C_{2-6}alkynyl;

- X is selected from the group consisting of O, NH, and CH_2;

- Ar is selected from the group consisting of C_{6-10}aryl, C_{6-10}heteroaryl, C_{5-10}carboxycyl, and C_{4-9}heterocycl, including fused bicyclic groups, wherein Ar is optionally substituted with one or more halo, hydroxyl, C_{1-6}alkyl, C_{1-6}haloalkyl, C_{1-6}alkoxy, or C_{1-6}acyloxy;

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wherein any of \( R^1, R^2, \) and \( R^3 \), independently and independently for each occurrence, is optionally further substituted with 1-3 substituents selected from the group consisting of \( C_{1-10} \text{alkyl}, C_{1-10} \text{haloalkyl}, \text{halo, hydroxyl, carboxyl, C}_{1-10} \text{alkoxycarbonyl, C}_{2-10} \text{alkenyl} \text{oxycarbonyl, C}_{2-10} \text{alkynyl} \text{oxycarbonyl, C}_{1-10} \text{acyl, C}_{1-10} \text{acylamino, C}_{1-10} \text{cyloxy, C}_{1-10} \text{carbonate, C}_{1-10} \text{alkoxy, C}_{6-10} \text{aryloxy, C}_{6-10} \text{arylamino, phosphoryl, phosphate, phosphonate, phosphinate, amino, diC}_{1-10} \text{alkylamino, monoC}_{1-10} \text{alkylamino, C}_{1-10} \text{amido, C}_{1-10} \text{amino, C}_{1-10} \text{carbamate, C}_{1-10} \text{urea, cyano, nitro, sulfhydryl, C}_{1-10} \text{alkythio, sulfate, sulfonate, sulfamoyl, sulfonamido, sulfonyl, C}_{1-7} \text{carbocyclic, C}_{3,5} \text{carbocyclic-C}_{1-6} \text{alkyl, C}_{1-8} \text{heterocyclic, C}_{1,2} \text{heterocyclic-C}_{1,2} \text{aryl, phenyl, phenyl-C}_{1-6} \text{alkyl, C}_{1,2} \text{heteroaryl, and C}_{1,2} \text{heteroaryl-C}_{1,2} \text{alkyl}; and wherein heterocyclic or heteroaromatic rings, independently for each occurrence, comprise 1-4 heteroatoms selected from N, O, and S;}

\begin{align*}
m & \text{is 1-3;} \\
n & \text{is 0-3;} \\
p & \text{is 0 or 1; and}
\end{align*}

wherein the composition is edible and capable of reducing bitter taste of a bitter taste.

2. The composition according to claim 1, wherein said compound according to Formula (I) is a compound selected from:

(a) a compound of Formula (IIa):

\[
\begin{array}{c}
\text{N} \\
\text{(R\text{\textsuperscript{1}})}_n \\
\text{(R\text{\textsuperscript{2}})}_m
\end{array}
\]

Formula (IIa):

or a comestibly or biologically acceptable salt or derivative thereof,

wherein, as valence and stability permit, \( R^1, R^2, m, \) and \( n \) are as defined for the compound of Formula (I) in claim 1; or

(b) a compound of Formula (IIb):
Formula (IIb);
or a comestibly or biologically acceptable salt or derivative thereof,
wherein, as valence and stability permit, $R^1$, $R^2$, $m$, and $n$ are as defined for the compound of Formula (I) in claim 1.

3. The composition according to claim 1, wherein said compound according to Formula (I) is a compound selected from:

(a) a compound of Formula (IIIb):

Formula (IIIb);
or a comestibly or biologically acceptable salt or derivative thereof,
wherein, as valence and stability permit:
$R^1$ and $n$ are as defined for the compound of Formula (I) in claim 1; and
$R^1$ is selected from the group consisting of methyl and ethyl;

(b) a compound of Formula (IIIb'):

Formula (IIIb');
or a comestibly or biologically acceptable salt or derivative thereof,
wherein, as valence and stability permit:

R¹, R², and n are as defined for the compound of Formula (I) in claim 1; and

Ar is C₉₋₁₀aryl optionally substituted with halo, hydroxyl, C₁₋₆alkyl, C₁₋₆alkoxy, or

C₁₋₆acyloxy; or

(c) a compound of Formula (IIIb⁰⁰):

\[
\begin{array}{c}
\text{Formula (IIIb⁰⁰),}
\end{array}
\]

or a comestibly or biologically acceptable salt or derivative thereof,

wherein, as valence and stability permit:

R¹, R², and m are as defined for the compound of Formula (I) in claim 1; and

R¹ is C₁₋₆alkyl.

4. The composition according to claim 1, wherein said compound according to

Formula (IV) is a compound selected from:

(a) a compound of Formula (Va):

\[
\begin{array}{c}
\text{Formula (Va),}
\end{array}
\]

or a comestibly or biologically acceptable salt or derivative thereof,

wherein, as valence and stability permit, R¹, R², R³, R⁴, and m are as defined for

the compound of Formula (IV) in claim 1; or

(b) a compound of Formula (Vb):
or a comestibly or biologically acceptable salt or derivative thereof,
wherein, as valence and stability permit, R₁, R₂, R₃, R₄, and m are as defined for
the compound of Formula (IV) in claim 1.

5. The composition according to claim 1, wherein said compound according to
Formula (IV) is a compound selected from:

(a) a compound of Formula (VIA):

10

Formula (VIA);
or a comestibly or biologically acceptable salt or derivative thereof,
wherein, as valence and stability permit:
R₁, R₂, R₃, R₄, are as defined for the compound of Formula (IV) in claim 1; and
ο is 0-2; or

15 (b) a compound of Formula (VIB):

Formula (VIB);
or a comestibly or biologically acceptable salt or derivative thereof,
wherein, as valence and stability permit:

R^1, R^2, R^3, R^4 are as defined for the compound of Formula (IV) in claim 1; and

o is 0-2.

6. The composition according to claim 1, wherein said compound according to Formula (IV) is a compound selected from:

(a) a compound of Formula (VIIa):

```
[Diagram of Formula (VIIa)]
```

or a comestibly or biologically acceptable salt or derivative thereof,

wherein, as valence and stability permit:

R^1, R^2, R^3, R^4 and R^5 are as defined for the compound of Formula (IV) in claim 1; and

o is 0-2; or

(b) a compound of Formula (VIIb):

```
[Diagram of Formula (VIIb)]
```

and comestibly or biologically acceptable derivatives thereof,

wherein, as valence and stability permit, R^1, R^2, R^3, R^4 and R^5 are as defined for the compound of Formula (IV) in claim 1; and

o is 0-2.

7. The composition according to claim 1, wherein said compound according to Formula (VII) is a compound of Formula (IX):
Formula (IX);

or a comestibly or biologically acceptable salt or derivative thereof,

wherein, as valence and stability permit:

$$\begin{align*}
R^1, R^2, R^3, R^4, m, n, \text{ and } o & \text{ are as defined for the compound of Formula (VIII) in claim 1.}
\end{align*}$$

8. The composition according to claim 1, wherein said compound according to Formula (VIII) is a compound of Formula (X):

Formula (X);

or a comestibly or biologically acceptable salt or derivative thereof,

wherein, as valence and stability permit:

$$\begin{align*}
R^1, R^2, R^3, \text{ and } n & \text{ are as defined for the compound of Formula (VIII) in claim 1; and}
\end{align*}$$
p is 0-2.

9. The composition according to claim 1, wherein said compound according to Formula (XI) is a compound selected from:

(a) a compound of Formula (XIIa):

Formula (XIIa):

or a comestibly or biologically acceptable salt or derivative thereof,

wherein, as valence and stability permit:

$R_1$, $R_2$, Het, and n are as defined for the compound of Formula (XI) in claim 1;

$R_3$, independently for each occurrence, is selected from the group consisting of halo, hydroxyl, $C_{1-4}$alkyl, $C_{1-4}$haloalkyl, $C_{2-6}$alkenyl, $C_{2-6}$alkynyl, and $C_{1-4}$alkoxy; and $m$ is 0-3; and

(b) a compound of Formula (XIIb):

Formula (XIIb):

or a comestibly or biologically acceptable salt or derivative thereof,

wherein, as valence and stability permit, $R_1$, $R_2$, $R_3$, Het, and n are as defined for the compound of Formula (XI) in claim 1.

10. The composition according to claim 1, wherein said compound according to Formula (XI) is a compound selected from:

(a) a compound of Formula (XIIIa):

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Formula (XIIIa); or a comestibly or biologically acceptable salt or derivative thereof,
wherein, as valence and stability permit, $R^1$, $R^2$, Het, $n$, and $m$ are as defined for the compound of Formula (XI) in claim 1; and

$R^3$ is, independently for each occurrence, is selected from the group consisting of halo, hydroxyl, $C_{1-6}$alkyl, $C_{1-6}$haloalkyl, $C_{2-6}$alkenyl, $C_{2-6}$alkynyl, and $C_{1-6}$alkoxy; and $m$ is 0-3; or

(b) a compound of Formula (XIIIb):

Formula (XIIIb); or a comestibly or biologically acceptable salt or derivative thereof,
wherein, as valence and stability permit:

$R^1$, $R^2$, $R^3$, Het, and $n$ are as defined for the compound of Formula (XI) in claim 1; and

$Ar$ is $C_{6-10}$aryl, optionally substituted with $C_{1-4}$alkyl.

11. The composition according to claim 1, wherein said compound according to Formula (XIV) is a compound selected from:

(a) a compound of Formula (XVa):
or a comestibly or biologically acceptable salt or derivative thereof,
wherein, as valence and stability permit:

5  \( R^1 \) and \( R^a \) are as defined for the compound of Formula (XIV) in claim 1; and
\( R^2 \) is \( C_{1-4} \) alkyl;

(b) a compound of Formula (XVb):

\[
\begin{align*}
(R^1)_n & \quad N \quad \text{Ar} \\
\text{Ar} & \quad X \quad Ar
\end{align*}
\]

Formula (XVb);

or a comestibly or biologically acceptable salt or derivative thereof,
wherein, as valence and stability permit:
\( R^1, R^a, X, \text{Ar}, \) and \( n \) are as defined for the compound of Formula (XIV) in claim 1; or

(e) a compound of Formula (XVc):

\[
\begin{align*}
(R^1)_n & \quad N \quad \text{Ar} \\
\text{Ar} & \quad \text{Ar}
\end{align*}
\]

Formula (XVc);

or a comestibly or biologically acceptable salt or derivative thereof, wherein, as
valence and stability permit, \( R^1, R^a, \text{Ar}, \) and \( n \) are as defined for the compound of Formula (XIV) in
claim 1.
12. The composition according to claim 1, wherein said compound is selected from the group consisting of Compounds 1-58 and having the structure:

<table>
<thead>
<tr>
<th>Compound 1</th>
</tr>
</thead>
<tbody>
<tr>
<td><img src="image1.png" alt="Compound 1" /></td>
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<table>
<thead>
<tr>
<th>Compound 2</th>
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<tbody>
<tr>
<td><img src="image2.png" alt="Compound 2" /></td>
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<tbody>
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Compound 11

Compound 12

Compound 13

Compound 14

Compound 15
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<td>18</td>
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Compound 26

Compound 27

Compound 28

Compound 29
Compound 38

Compound 39

Compound 40

Compound 41
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<td>Compound</td>
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<td>53</td>
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<td>54</td>
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<tr>
<td>55</td>
</tr>
</tbody>
</table>
or comestibly or biologically acceptable derivatives thereof.

13. The composition according to any one of claims 1-12 further comprising a bitter tastant.

14. The composition according to claim 13, wherein the bitter tastant is KCl or, potassium lactate.

15. The composition of any one of claims 1-14, wherein the composition further comprises one or more component selected from the group consisting of: NaCl, sodium lactate, and sugar.

16. A food product comprising the composition of any one of claims 1-15.

17. A method of preparing an edible composition comprising:
   (a) providing comestibly acceptable carrier; and
   (b) adding to said comestibly acceptable carrier a compound according to
       Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb), Formula (IIIb'), Formula (IIIb''),
       Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa),
       Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa),
       Formula (XIIb), Formula (XIIla), Formula (XIIIb), Formula (XIV), Formula (XVa),
       Formula (XVb) or Formula (XVe), as described herein, or combinations thereof, or any one of
       Compounds 1-58, as described herein, or combinations thereof.

18. The method according to claim 17, wherein said comestibly acceptable carrier is inherently bitter.
19. The method according to claim 18, wherein the compostibly acceptable carrier comprises a bitter tasting potassium salt.

20. The method according to claim 19, wherein the potassium salt is KCl or potassium lactate.

21. The method according to any one of claims 17-20, wherein the edible composition further comprises one or more component selected from the group consisting of: NaCl, sodium lactate, and sugar.

22. The method according to claim 17, wherein the method further comprises:
   (c) adding a bitter tastant to said compostibly acceptable carrier, wherein said bitter tastant is a potassium salt.

23. The method according to claim 22, wherein the potassium salt is KCl or potassium lactate.

24. The method according to claim 22 or 23, wherein the edible composition further comprises one or more component selected from the group consisting of: NaCl, sodium lactate, and sugar.

25. A method of reducing the amount of NaCl in an edible composition comprising:
   (a) replacing an amount of NaCl used in preparing said edible composition with an amount of KCl; and
   (b) incorporating into the an edible composition an effective amount of a compound according to Formula (I), Formula (Ia), Formula (IIb), Formula (IIIb), Formula (IIB*), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof to produce an edible composition with reduced NaCl.

26. The method according to claim 25, wherein the amount of compound incorporated into the edible composition is sufficient to permit replacement of the amount of NaCl present in the edible composition by up to 25%, 50%, 75% or 100%.

27. The method according to claim 25 or 26, wherein the edible composition with reduced NaCl maintains a salty flavor.
28. A method of reducing the amount of sodium lactate in an edible composition comprising:

   (a) replacing an amount of sodium lactate used in preparing said edible composition with an amount of potassium lactate; and

   (b) incorporating into said edible composition an effective amount of a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIb'), Formula (IIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVIII), Formula (XVIIla) or Formula (XVIIce), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof to produce a edible composition with reduced NaCl.

29. The method according to claim 28, wherein the amount of compound incorporated into the edible composition is sufficient to permit replacement of the amount of sodium lactate typically present in the edible composition by up to 25%, 50%, 75% or 100%.

30. The method according to claim 28 or 29, wherein the edible composition with reduced sodium lactate has the same shelf life as an edible composition comprising the full amount sodium lactate.

31. A method of reducing the amount of sugar in an edible composition comprising:

   (a) replacing an amount of sugar used in preparing an edible composition with an amount of Acesulfame K; and

   (b) incorporating into said edible composition an effective amount of a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIb'), Formula (IIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVIII), Formula (XVIIla) or Formula (XVIIce), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof to produce an edible composition with reduced sugar.

32. The method according to claim 31, wherein the amount of compound incorporated into the edible composition is sufficient to permit replacement of the amount of sugar present in the edible composition by up to 25%, 50%, 75% or 100%.

33. The method according to claim 31 or 32, wherein the edible composition with reduced sugar maintains a sweet flavor.
34. A method of reducing the sodium intake of a subject, the method comprising:
(a) replacing an amount of a sodium salt used in preparing an edible
composition with an amount of a potassium salt; and
(b) incorporating into said edible composition an effective amount of a
compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb),
Formula (IIIb*), Formula (IV), Formula (Va), Formula (Vb), Formula (Via), Formula (Vib),
Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI),
Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV),
Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof, or
any one of Compounds 1-58, as described herein, or combinations thereof.

35. The method according to claim 34, wherein the sodium salt is NaCl and the
potassium salt is KCl.

36. The method according to claim 34, wherein the sodium salt is sodium lactate,
and the potassium salt is potassium lactate.

37. The method according to any one of claims 34-36, wherein the method
further comprises: (c) identifying a subject in need thereof.

38. The method according to any one of claims 43-46, wherein the amount of
compound incorporated into the edible composition is sufficient to reduce sodium intake by up to
25%, 50%, 75% or 100%.

39. A method of reducing the sugar intake of a subject, the method comprising:
(a) replacing an amount of sugar used in preparing an edible composition
with an amount of a Acctusulfame K; and
(b) incorporating into said edible composition an effective amount of a
compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb),
Formula (IIIb*), Formula (IV), Formula (Va), Formula (Vb), Formula (Via), Formula (Vib),
Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX), Formula (X), Formula (XI),
Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb), Formula (XIV),
Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof, or
any one of Compounds 1-58, as described herein, or combinations thereof.

40. The method according to claim 39, wherein the method further comprises (c)
identifying a subject in need thereof.

41. The method according to claim 39 or 40, wherein the amount of compound
incorporated into the edible composition is sufficient to reduce sugar intake by up to 25%, 50%,
75% or 100%.
42. A method of reducing bitter taste attributed to a bitter tastant in an edible composition, wherein said bitter tastant is a potassium salt, said method comprising:

(a) providing an edible composition comprising said bitter tastant; and
(b) adding an effective amount of a compound according to Formula (I),

Formula (Ia), Formula (Ib), Formula (IIIb), Formula (IIIb’), Formula (IIIb”), Formula (IV),
Formula (Va), Formula (Vb), Formula (Via), Formula (Vb), Formula (Vla), Formula (Vib),
Formula (VIIa), Formula (VIIb), Formula (IXa), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb),
Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or
Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as
described herein, or combinations thereof, to the edible composition generated in (a) such that any
bitter taste induced by the bitter tastant is reduced.

43. A method of reducing bitter taste attributed to a bitter tastant in an edible composition, wherein said bitter tastant is a potassium salt, said method comprising:

(a) ingesting an effective amount of a compound according to Formula (I),

Formula (Ia), Formula (Ib), Formula (IIIb), Formula (IIIb’), Formula (IIIb”), Formula (IV),
Formula (Va), Formula (Vb), Formula (Via), Formula (Vb), Formula (Vla), Formula (Vib),
Formula (VIIa), Formula (VIIb), Formula (IXa), Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb),
Formula (XIIIa), Formula (XIIIb), Formula (XIV), Formula (XVa), Formula (XVb) or
Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as
described herein, or combinations thereof, before, along with, or after the edible composition such
that any bitter taste induced by the bitter tastant is reduced.

44. The method according to claim 42 or 43, wherein the bitter taste induced by
the bitter tastant is reduced by up to 25%, 50%, 75% or 100%

45. The method according to any one of claims 42-44, wherein the potassium salt
is KCl or potassium lactate.

46. The method according to any one of claims 42-45, wherein the edible
composition further comprises NaCl, sodium lactate, or sugar.

47. A method of preserving an edible composition comprising:

(a) providing an edible composition; and
(b) incorporating into said edible composition potassium lactate and an
effective amount of a compound according to Formula (I), Formula (Ia), Formula (Ib),
Formula (IIIb), Formula (IIIb’), Formula (IIIb”), Formula (IV), Formula (Va), Formula (Vb),
Formula (Via), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX),
Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb),
Formulas (XIV), (XVa), (XVb) or (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.

48. A method of reducing the amount of sodium in an edible composition while preserving the edible composition, the method comprising:

(a) replacing an amount of sodium lactate used in preparing said edible composition with an amount of potassium lactate; and

(b) incorporating into said edible composition an effective amount of a compound according to Formula (I), Formula (IIa), Formula (IIb), Formula (IIIb'), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb), Formula (Vla), Formula (Vlb),

Formula (VIIa), Formula (VIIb), Formula (VII), Formula (IX), Formula (X), Formula (XI),

Formula (XIIa), Formula (XIIb), Formula (XII), Formula (XIIIa), Formula (XIIIb), Formula (XIV),

Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.

49. The method according to any one of claims 17-48, wherein the edible composition is selected from the group consisting of a food product, a consumer product, and a pharmaceutical composition.

50. A method of inhibiting, reducing, or eliminating the perception of a bitter taste in a subject, wherein said bitter tastant is a potassium salt, said method comprising:

(a) placing a compound according to Formula (I), Formula (IIa), Formula (IIb),

Formula (IIIb), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb),

Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VII), Formula (IX),

Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb),

Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVc), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof in the oral cavity of the subject.

51. The method according to claim 50, wherein the bitter taste is due to KCl or potassium lactate.

52. A pharmaceutical composition comprising:

(a) a bitter tasting pharmaceutical active ingredient, wherein said pharmaceutical active ingredient is a potassium salt; and

(b) a compound according to Formula (I), Formula (IIa), Formula (IIb),

Formula (IIIb), Formula (IIIb''), Formula (IV), Formula (Va), Formula (Vb),

Formula (Vla), Formula (Vlb), Formula (VIIa), Formula (VIIb), Formula (VII), Formula (IX),

Formula (X), Formula (XI), Formula (XIIa), Formula (XIIIb), Formula (XIII), Formula (XIIIb),
Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.

53. A pharmaceutical composition comprising:
   (a) a pharmaceutical active ingredient;
   (b) a bitter tastant, wherein said bitter tastant is a potassium salt; and
   (c) a compound according to Formula (I), Formula (Ia), Formula (Ib),
       Formula (IIb), Formula (IIb’), Formula (IIb”), Formula (IV), Formula (Va), Formula (Vb),
       Formula (Via), Formula (Vib), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX),
       Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb),
       Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or
       combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.

54. A consumer product comprising:
   (a) a bitter tasting ingredient, wherein said bitter tasting ingredient is a potassium salt; and
   (b) a compound according to Formula (I), Formula (Ia), Formula (Ib),
       Formula (IIb), Formula (IIb’), Formula (IIb”), Formula (IV), Formula (Va), Formula (Vb),
       Formula (Via), Formula (Vib), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX),
       Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb),
       Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or
       combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.

55. A consumer product for reducing bitter taste of a bitter tastant, wherein said bitter tastant is a potassium salt, and wherein said consumer product comprises:
   (a) a compound according to Formula (I), Formula (Ia), Formula (Ib),
       Formula (IIb), Formula (IIb’), Formula (IIb”), Formula (IV), Formula (Va), Formula (Vb),
       Formula (Via), Formula (Vib), Formula (VIIa), Formula (VIIb), Formula (VIII), Formula (IX),
       Formula (X), Formula (XI), Formula (XIIa), Formula (XIIb), Formula (XIIIa), Formula (XIIIb),
       Formula (XIV), Formula (XVa), Formula (XVb) or Formula (XVe), as described herein, or
       combinations thereof, or any one of Compounds 1-58, as described herein, or combinations thereof.
**Solution Testing** — The left data point in the solution charts represents the bitterness or metallic taste/impression score of the KCl/potassium lactate standard. The right data point in the solution charts represents the bitterness or metallic taste/impression score of the Test Solution. The concentration of the Test Compound used in each experiment is recited below the chart. In addition, the statistical significance of the Solution Testing data, determined using a paired T-test analysis, is presented wherein "a" represents p< 0.1; "b" represents p< 0.05; "c" represents p< 0.01; and "d" represents p> 0.1 (data not shown).

**Foodstuff Testing** — The fraction represents the number of tasters that discerned a decrease in the bitterness or metallic taste/impression of the Test Foodstuff. In addition, the concentration of the Test Compound used in each experiment is recited. Further, the statistical significance of the Foodstuff Testing data, determined using binomial distribution analysis, is presented wherein "a" represents p< 0.1; "b" represents p< 0.05; "c" represents p< 0.01; and "d" represents p> 0.1 (data not shown).

"--" denotes that the solution or foodstuff was not tested.

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<th>No.</th>
<th>Test Compound Structure</th>
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<th>KCl foodstuff</th>
<th>Potassium lactate solution</th>
<th>Potassium lactate foodstuff</th>
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<td><img src="image" alt="Chart A" /></td>
<td>19/28 10 ppm b</td>
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**Table:**

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<th>Potassium lactate foodstuff</th>
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- 1 ppm
- n=7
- d
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<th>No.</th>
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<tbody>
<tr>
<td>I</td>
<td>![Compound I]</td>
<td>![Graph I]</td>
<td>![Graph II]</td>
</tr>
<tr>
<td>II</td>
<td>![Compound II]</td>
<td>![Graph III]</td>
<td>![Graph IV]</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Potassium lactate foodstuff</th>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Potassium lactate solution</td>
<td></td>
<td></td>
</tr>
<tr>
<td>KCl foodstuff</td>
<td>9/18 ppm d</td>
<td></td>
</tr>
<tr>
<td>KCl solution</td>
<td>![Graph V]</td>
<td>![Graph VI]</td>
</tr>
</tbody>
</table>

**FIGURE 1F**
<table>
<thead>
<tr>
<th>No.</th>
<th>Test Compound Structure</th>
</tr>
</thead>
<tbody>
<tr>
<td>12</td>
<td><img src="image" alt="Compound 12" /></td>
</tr>
<tr>
<td>13</td>
<td><img src="image" alt="Compound 13" /></td>
</tr>
</tbody>
</table>

<table>
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<tr>
<th>Substance</th>
<th>Description</th>
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</thead>
<tbody>
<tr>
<td>KCl</td>
<td>foodstuffs</td>
</tr>
<tr>
<td>KCl solution</td>
<td>n=7</td>
</tr>
<tr>
<td>Potassium lactate</td>
<td>n=7</td>
</tr>
</tbody>
</table>

FIGURE 1G
<table>
<thead>
<tr>
<th>No.</th>
<th>Test Compound Structure</th>
</tr>
</thead>
<tbody>
<tr>
<td>14</td>
<td><img src="image" alt="Compound Structure 14" /></td>
</tr>
<tr>
<td>15</td>
<td><img src="image" alt="Compound Structure 15" /></td>
</tr>
</tbody>
</table>

**Potassium lactate solution**

<p>| | | |</p>
<table>
<thead>
<tr>
<th></th>
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</tr>
</thead>
<tbody>
<tr>
<td>n=6</td>
<td>5 ppm</td>
<td>d</td>
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<table>
<thead>
<tr>
<th>KCl foodstuff</th>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>14:28</td>
<td>1 ppm</td>
<td>d</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>KCl solution</th>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>n=8</td>
<td>1 ppm</td>
<td>b</td>
</tr>
</tbody>
</table>

<p>| | | |</p>
<table>
<thead>
<tr>
<th></th>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>n=8</td>
<td>10 ppm</td>
<td>c</td>
</tr>
</tbody>
</table>

**FIGURE IH**
**Solution Testing** – The left data point in the solution charts represents the bitterness or metallic taste/impression score of the KCl/potassium lactate standard. The right data point in the solution charts represents the bitterness or metallic taste/impression score of the Test Solution. The concentration of the Test Compound used in each experiment is recited below the chart. In addition, the statistical significance of the Solution Testing data, determined using a paired T-test analysis, is presented wherein "a" represents \( p < 0.1 \); "b" represents \( p < 0.05 \); "c" represents \( p < 0.01 \); and "d" represents \( p > 0.1 \) (data not shown).

**Foodstuff Testing** – The fraction represents the number of tasters that discerned a decrease in the bitterness or metallic taste/impression of the Test Foodstuff. In addition, the concentration of the Test Compound used in each experiment is recited. Further, the statistical significance of the Foodstuff Testing data, determined using binomial distribution analysis, is presented wherein "a" represents \( p < 0.1 \); "b" represents \( p < 0.05 \); "c" represents \( p < 0.01 \); and "d" represents \( p > 0.1 \) (data not shown).

"--" denotes that the solution or foodstuff was not tested.

<table>
<thead>
<tr>
<th>No.</th>
<th>Test Compound Structure</th>
<th>KCl Solution</th>
<th>KCl Foodstuff</th>
<th>Potassium Lactate Solution</th>
<th>Potassium Lactate Foodstuff</th>
</tr>
</thead>
<tbody>
<tr>
<td>23</td>
<td><img src="image" alt="Image" /></td>
<td><img src="image" alt="Image" /></td>
<td><img src="image" alt="Image" /></td>
<td><img src="image" alt="Image" /></td>
<td><img src="image" alt="Image" /></td>
</tr>
</tbody>
</table>

**Legend:**
- \( n = 8 \) for solution data
- \( n = 15 \) for foodstuff data
- \( 10 \) ppm for solution data
- \( 1 \) ppm for foodstuff data
- \( a \) for statistically significant
- \( b \) for non-significant
<table>
<thead>
<tr>
<th>No</th>
<th>Test Compound Structure</th>
</tr>
</thead>
<tbody>
<tr>
<td>24</td>
<td><img src="image" alt="Chemical Structure 24" /></td>
</tr>
<tr>
<td>25</td>
<td><img src="image" alt="Chemical Structure 25" /></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Potassium lactate foodstuff</th>
<th>19/30</th>
<th>1 ppm</th>
</tr>
</thead>
<tbody>
<tr>
<td>Potassium lactate solution</td>
<td></td>
<td></td>
</tr>
<tr>
<td>KCl foodstuff</td>
<td>28/34</td>
<td>10 ppm</td>
</tr>
<tr>
<td>KCl solution</td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>25/34</td>
<td>10 ppm</td>
</tr>
</tbody>
</table>

**FIGURE 2B**
<table>
<thead>
<tr>
<th>Potassium lactate foodstuff</th>
<th>18:32</th>
<th>30 ppm</th>
<th>19:32</th>
<th>30 ppm</th>
</tr>
</thead>
<tbody>
<tr>
<td>Potassium lactate solution</td>
<td><img src="image1" alt="Diagram A" /></td>
<td><img src="image2" alt="Diagram B" /></td>
<td><img src="image3" alt="Diagram C" /></td>
<td><img src="image4" alt="Diagram D" /></td>
</tr>
<tr>
<td><img src="image5" alt="Diagram E" /></td>
<td><img src="image6" alt="Diagram F" /></td>
<td><img src="image7" alt="Diagram G" /></td>
<td><img src="image8" alt="Diagram H" /></td>
<td></td>
</tr>
<tr>
<td>KCl foodstuff</td>
<td>18:28</td>
<td>1 ppm</td>
<td>22:38</td>
<td>30 ppm</td>
</tr>
<tr>
<td>KCl solution</td>
<td><img src="image9" alt="Diagram I" /></td>
<td><img src="image10" alt="Diagram J" /></td>
<td><img src="image11" alt="Diagram K" /></td>
<td><img src="image12" alt="Diagram L" /></td>
</tr>
<tr>
<td><img src="image13" alt="Diagram M" /></td>
<td><img src="image14" alt="Diagram N" /></td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

**Figure 2C**

<table>
<thead>
<tr>
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</tr>
</thead>
<tbody>
<tr>
<td>26</td>
<td><img src="image15" alt="Compound 26" /></td>
</tr>
<tr>
<td>27</td>
<td><img src="image16" alt="Compound 27" /></td>
</tr>
<tr>
<td>No</td>
<td>Test Compound Structure</td>
</tr>
<tr>
<td>-----</td>
<td>-------------------------</td>
</tr>
<tr>
<td>28</td>
<td><img src="image" alt="Compound 28" /></td>
</tr>
<tr>
<td>29</td>
<td><img src="image" alt="Compound 29" /></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Potassium lactate foodstuff</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Potassium lactate solution</td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>KCl foodstuff</th>
</tr>
</thead>
<tbody>
<tr>
<td>12/20</td>
</tr>
<tr>
<td>10 ppm</td>
</tr>
<tr>
<td>d</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>KCl solution</th>
</tr>
</thead>
<tbody>
<tr>
<td>17/26</td>
</tr>
<tr>
<td>1 ppm</td>
</tr>
<tr>
<td>a</td>
</tr>
</tbody>
</table>

![Graph](image)
<table>
<thead>
<tr>
<th>No</th>
<th>Test Compound Structure</th>
</tr>
</thead>
<tbody>
<tr>
<td>30</td>
<td><img src="image1" alt="Compound 30" /></td>
</tr>
<tr>
<td>31</td>
<td><img src="image2" alt="Compound 31" /></td>
</tr>
<tr>
<td>No</td>
<td>Test Compound Structure</td>
</tr>
<tr>
<td>----</td>
<td>--------------------------------------</td>
</tr>
<tr>
<td>No</td>
<td></td>
</tr>
<tr>
<td>KCl solution</td>
<td></td>
</tr>
<tr>
<td>Potassium lactate foodstuff</td>
<td></td>
</tr>
<tr>
<td>Potassium lactate solution</td>
<td></td>
</tr>
<tr>
<td>No</td>
<td>Test Compound Structure</td>
</tr>
<tr>
<td>----</td>
<td>-------------------------</td>
</tr>
<tr>
<td>34</td>
<td><img src="image1" alt="Structure 1" /></td>
</tr>
<tr>
<td>35</td>
<td><img src="image3" alt="Structure 3" /></td>
</tr>
</tbody>
</table>

**Figure 2G**
<table>
<thead>
<tr>
<th>Potassium lactate foodstuff</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>KCl foodstuff</td>
<td>9/16 ppm</td>
</tr>
<tr>
<td>KCl solution</td>
<td>10 ppm</td>
</tr>
</tbody>
</table>

**FIGURE 2H**

Test Compound Structure

<table>
<thead>
<tr>
<th>No</th>
<th>36</th>
<th><img src="image" alt="Chemical Structure" /></th>
</tr>
</thead>
</table>
**Solution Testing** – The left data point in the solution charts represents the bitterness or metallic taste/impression score of the KCl/potassium lactate standard. The right data point in the solution charts represents the bitterness or metallic taste/impression score of the Test Solution. The concentration of the Test Compound used in each experiment is recited below the chart. In addition, the statistical significance of the Solution Testing data, determined using a paired T-test analysis, is presented wherein "a" represents \( p < 0.1 \); "b" represents \( p < 0.05 \); "c" represents \( p < 0.01 \); and "d" represents \( p > 0.1 \) (data not shown).

**Foodstuff Testing** – The fraction represents the number of tasters that discerned a decrease in the bitterness or metallic taste/impression of the Test Foodstuff. In addition, the concentration of the Test Compound used in each experiment is recited. Further, the statistical significance of the Foodstuff Testing data, determined using binomial distribution analysis, is presented wherein "a" represents \( p < 0.1 \); "b" represents \( p < 0.05 \); "c" represents \( p < 0.01 \); and "d" represents \( p > 0.1 \) (data not shown).

"--" denotes that the solution or foodstuff was not tested.

<table>
<thead>
<tr>
<th>No.</th>
<th>Test Compound Structure</th>
<th>KCl solution</th>
<th>KCl foodstuff</th>
<th>Potassium lactate solution</th>
<th>Potassium lactate foodstuff</th>
</tr>
</thead>
<tbody>
<tr>
<td>37</td>
<td>![Structure Image]</td>
<td>![Chart Image]</td>
<td>29/34</td>
<td></td>
<td>26/39</td>
</tr>
</tbody>
</table>

1 ppm \( b \)                           1 ppm \( a \)                       | \( n=7 \)  
10 ppm \( c \)                           |                                      |
<table>
<thead>
<tr>
<th>Test Compound Structure</th>
<th>No.</th>
<th>38</th>
<th>39</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td></td>
<td><img src="image1" alt="Image" /></td>
<td><img src="image2" alt="Image" /></td>
</tr>
<tr>
<td>Test Compound Structure</td>
<td>No.</td>
<td></td>
<td></td>
</tr>
<tr>
<td>-------------------------</td>
<td>-----</td>
<td></td>
<td></td>
</tr>
<tr>
<td><img src="image1" alt="Structure" /></td>
<td>42</td>
<td></td>
<td></td>
</tr>
<tr>
<td><img src="image2" alt="Structure" /></td>
<td>43</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>
Solution Testing — The left data point in the solution charts represents the bitterness or metallic taste/impression score of the KCl/potassium lactate standard. The right data point in the solution charts represents the bitterness or metallic taste/impression score of the Test Solution. The concentration of the Test Compound used in each experiment is recited below the chart. In addition, the statistical significance of the Solution Testing data, determined using a paired T-test analysis, is presented wherein "a" represents p < 0.1; "b" represents p < 0.05; "c" represents p < 0.01; and "d" represents p > 0.1 (data not shown).

Foodstuff Testing — The fraction represents the number of tasters that discerned a decrease in the bitterness or metallic taste/impression of the Test Foodstuff. In addition, the concentration of the Test Compound used in each experiment is recited. Further, the statistical significance of the Foodstuff Testing data, determined using binomial distribution analysis, is presented wherein "a" represents p < 0.1; "b" represents p < 0.05; "c" represents p < 0.01; and "d" represents p > 0.1 (data not shown).

"--" denotes that the solution or foodstuff was not tested.

<table>
<thead>
<tr>
<th>No.</th>
<th>Test Compound Structure</th>
<th>KCl solution</th>
<th>KCl foodstuff</th>
<th>Potassium lactate solution</th>
<th>Potassium lactate foodstuff</th>
</tr>
</thead>
<tbody>
<tr>
<td>44</td>
<td><img src="image" alt="Compound Structure" /></td>
<td><img src="image" alt="KCl Solution Chart" /></td>
<td>14/26 10 ppm d</td>
<td><img src="image" alt="Potassium Lactate Solution Chart" /></td>
<td>--</td>
</tr>
<tr>
<td>No.</td>
<td>Test Compound Structure</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>-----</td>
<td>-------------------------</td>
<td>---</td>
<td>---</td>
<td>---</td>
<td>---</td>
</tr>
<tr>
<td>47</td>
<td><img src="image1.png" alt="Image of compound 47" /></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>48</td>
<td><img src="image2.png" alt="Image of compound 48" /></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

**Potassium Lactate Foodstuff**

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>KCl Foodstuff</strong></td>
<td>12/16 10 ppm</td>
</tr>
<tr>
<td><strong>KCl Solution</strong></td>
<td>17/38 10 ppm</td>
</tr>
</tbody>
</table>

**Test Compound Structures**

**Figure 4C**
**Solution Testing** — The left data point in the solution charts represents the bitterness or metallic taste/impression score of the KCl/potassium lactate standard. The right data point in the solution charts represents the bitterness or metallic taste/impression score of the Test Solution. The concentration of the Test Compound used in each experiment is recited below the chart. In addition, the statistical significance of the Solution Testing data, determined using a paired T-test analysis, is presented wherein "a" represents p<0.1; "b" represents p<0.05; "c" represents p<0.01; and "d" represents p>0.1 (data not shown).

**Foodstuff Testing** — The fraction represents the number of tasters that discerned a decrease in the bitterness or metallic taste/impression of the Test Foodstuff. In addition, the concentration of the Test Compound used in each experiment is recited. Further, the statistical significance of the Foodstuff Testing data, determined using binomial distribution analysis, is presented wherein "a" represents p<0.1; "b" represents p<0.05; "c" represents p<0.01; and "d" represents p>0.1 (data not shown).

"--" denotes that the solution or foodstuff was not tested.

<table>
<thead>
<tr>
<th>No.</th>
<th>Test Compound Structure</th>
<th>KCl solution</th>
<th>KCl foodstuff</th>
<th>Potassium lactate solution</th>
<th>Potassium lactate foodstuff</th>
</tr>
</thead>
<tbody>
<tr>
<td>49</td>
<td><img src="image" alt="Structure" /></td>
<td><img src="image" alt="Chart" /></td>
<td>18/34 1 ppm d</td>
<td>30 ppm n=7 a</td>
<td>26/38 30 ppm b</td>
</tr>
<tr>
<td>No.</td>
<td>Test Compound Structure</td>
<td>19/38 ppm</td>
<td>10 ppm</td>
<td>10/16 ppm</td>
<td>15 ppm</td>
</tr>
<tr>
<td>------</td>
<td>-----------------------------------------</td>
<td>-----------</td>
<td>--------</td>
<td>-----------</td>
<td>--------</td>
</tr>
<tr>
<td>50</td>
<td><img src="image1" alt="Compound Structure" /></td>
<td>d</td>
<td>d</td>
<td>d</td>
<td>d</td>
</tr>
<tr>
<td>51</td>
<td><img src="image2" alt="Compound Structure" /></td>
<td></td>
<td></td>
<td></td>
<td>b</td>
</tr>
</tbody>
</table>

*Potassium lactate foodstuff*

*Potassium lactate solution*

*KCl foodstuff*

*KCl solution*
<table>
<thead>
<tr>
<th>No.</th>
<th>Test Compound Structure</th>
<th>52</th>
<th>53</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
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<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td></td>
<td><img src="image" alt="Structure 52" /></td>
<td><img src="image" alt="Structure 53" /></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Potassium lactate foodstuff</th>
<th>15/26</th>
<th>30 ppm</th>
<th>18/32</th>
<th>30 ppm</th>
</tr>
</thead>
<tbody>
<tr>
<td>KCl foodstuff</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>18/26</td>
<td>10 ppm</td>
<td>n=10</td>
<td>11/16</td>
<td>30 ppm</td>
</tr>
<tr>
<td>KCl solution</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>10 ppm</td>
<td>n=10</td>
<td>25 ppm</td>
<td>n=9</td>
</tr>
</tbody>
</table>

**Figure 5C**
<table>
<thead>
<tr>
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<th>Test Compound Structure</th>
</tr>
</thead>
<tbody>
<tr>
<td>54</td>
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</tr>
<tr>
<td>55</td>
<td><img src="image2" alt="Chemical Structure" /></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Potassium lactate foodstuff</th>
<th>Potassium lactate solution</th>
<th>KCl foodstuff</th>
<th>KCl solution</th>
</tr>
</thead>
<tbody>
<tr>
<td>19/22 10 ppm c</td>
<td>1 ppm n=10 d</td>
<td>16/30 1 ppm d</td>
<td>25/36 1 ppm b</td>
</tr>
<tr>
<td><img src="image3" alt="Diagram" /></td>
<td><img src="image4" alt="Diagram" /></td>
<td><img src="image5" alt="Diagram" /></td>
<td><img src="image6" alt="Diagram" /></td>
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</tbody>
</table>

**FIGURE 5D**
<table>
<thead>
<tr>
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<th>Test Compound Structure</th>
</tr>
</thead>
<tbody>
<tr>
<td>56</td>
<td><img src="image" alt="Chemical Structure" /></td>
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</tbody>
</table>

<table>
<thead>
<tr>
<th>Potassium lactate foodstuff</th>
<th>18.26 ppm</th>
</tr>
</thead>
<tbody>
<tr>
<td>Potassium lactate solution</td>
<td><img src="image" alt="Graph" /> 10 ppm</td>
</tr>
<tr>
<td>KCl foodstuff</td>
<td>11.6 ppm</td>
</tr>
<tr>
<td>KCl solution</td>
<td>30 ppm</td>
</tr>
<tr>
<td></td>
<td><img src="image" alt="Graph" /> 1 ppm</td>
</tr>
</tbody>
</table>

**Figure 5E**