

**Improved method for preparing ginkgo extracts having a low
content of 4'-O-methyl pyridoxine and/or biflavones**

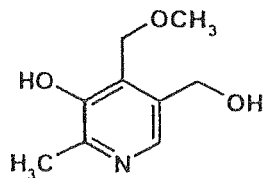
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The present invention relates to an improved multi-step method for preparing an extract from ginkgo biloba having a reduced content of 4'-O-methyl pyridoxine and/or biflavones, wherein the depletion is effected by filtration over an adsorber resin and/or an ion exchanger and the substances to be removed are retained on
10 the resin. The invention further relates to an extract from ginkgo biloba having a reduced content of 4'-O-methyl pyridoxine and/or biflavones which is obtainable by the method according to the present invention, as well as to its use.

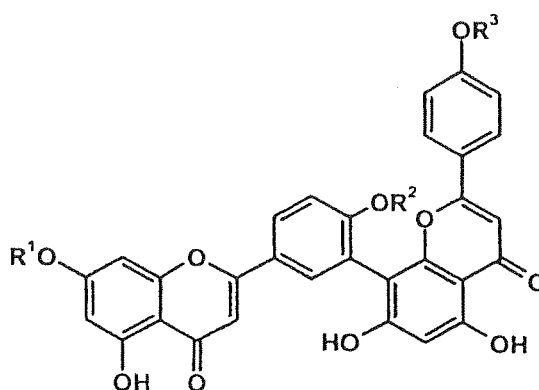
Since decades, extracts from the leaves of ginkgo biloba are used as a medica-
15 ment. They are currently used for the treatment of different kinds of dementia and symptoms thereof as well as cerebral and peripheral blood circulation disorders. Ingredients, the efficacy is associated with, are terpene lactones (ginkgolides A, B, C and bilobalide) as well as glycosides of flavones (quercetin, kaempferol and isorhamnetin). The leaves of ginkgo biloba also contain components which do not
20 contribute to the desired efficacy, but which may be responsible for risks and side effects. In addition to unpolar plant ingredients such as ginkgolic acids, those components are 4'-O-methyl pyridoxine and biflavones. In a ginkgo extract, which is efficacious and at the same time as safe as possible and as low in side effects as possible, these compounds should thus not be present to the largest possible
25 extent.

4'-O-methyl pyridoxine may cause symptoms of poisoning such as convulsive sei-
zures and unconsciousness. Thus, this compound is also referred to as ginkgo-
toxin. The biflavones contained in ginkgo exhibit an immunotoxic potential and
30 may elicit contact allergies. These biflavones contained in ginkgo are predomi-
nantly the compounds amentoflavone, bilobetin, ginkgetin, isoginkgetin and
sciadopitysin.

4'-O-Methyl pyridoxine:



5 Biflavones:



Amentoflavone:	$R^1 = R^2 = R^3 = H$
Bilobetin:	$R^1 = R^3 = H, R^2 = OCH_3$
Ginkgetin:	$R^3 = H, R^1 = R^2 = OCH_3$
Isoginkgetin:	$R^1 = H, R^2 = R^3 = OCH_3$
Sciadopitysin:	$R^1 = R^2 = R^3 = OCH_3$

Methods for depleting 4'-O-methyl pyridoxine and simultaneously biflavones as well as the extracts obtained are already described in EP 1 037 646 B1. Therein, 4'-O-methyl pyridoxine is retained using an acidic cation exchanger and the biflavones are adsorbed on activated carbon. Preferably these depletion steps are carried out after step f) of the method according to EP 1 037 646 B1 (page 3), i.e., after "treating the solution with a lead compound or an insoluble polyamide". The method of EP 1 037 646 B1 is based on the method originally described in DE 39 40 091 C2.

In EP 1 037 646 B1 the depletion of the biflavones and 4'-O-methyl pyridoxine is carried out at a stage of the method (step f)), wherein relatively high contents of lead salts and ammonium salts are present such that an increased amount of ion exchanger has to be employed.

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Furthermore, the depletions of biflavones and 4'-O-methyl pyridoxine are described and claimed in combination only. However, depending on the limits to be possibly established in the future and in view of the requirement of as low a modification of the extract composition as possible, it may be desirable to remove the
10 biflavones or 4'-O-methyl pyridoxine only.

Furthermore, the contents of less than 50 ppm of 4'-O-methyl pyridoxine and less than 100 ppm of biflavones which are aspired in EP 1 037 646 B1, are still comparatively high. Since EP 1 037 646 B1 does not describe a comparative example,
15 also the actual degree of depletion achieved in Examples 2 and 3 cannot be seen from this reference.

Thus, it is the object underlying the present invention to provide a method for preparing ginkgo extracts having a low content of 4'-O-methyl pyridoxine and/or a low
20 content of bioflavones which does not have the disadvantages indicated above and, in particular,

- wherein well recyclable and/or very inexpensive adsorbents are employed,
- which leads to contents of 4'-O-methyl pyridoxine of less than 20 ppm, preferably less than 10 ppm and particularly preferred less than 2 ppm and/or
25 contents of biflavones of less than 20 ppm, preferably less than 10 ppm and particularly preferred less than 5 ppm and
- which leads to extracts having preferred contents of flavonoides of 20 to 30% by weight (particularly preferred 22.0 to 27.0%), of terpene lactones of 4.5 to 8.5% by weight (particularly preferred 5.0 to 7.0%, preferably 2.8 to 3.4%
30 ginkgolides A, B and C and 2.6 to 3.2% bilobalid) and of ginkgolic acids (5 ppm at the most), wherein said contents of flavonoids of 22.0 to 27.0%, of terpene lactones of 5.0 to 7.0% (2.8 to 3.4% thereof ginkgolides A, B and C

and 2.6 to 3.2% thereof bilobalid) and of ginkgolic acids (5 ppm at the most) fulfil the requirements of the German Pharmacopoeia 2003 ("Deutsches Arzneibuch"; DAB).

- 5 According to the DAB the flavonoids are determined in the form of quercetin, kaempferol and isorhamnetin after acidic hydrolysis and calculated as flavonoid glycosides.

This object could be solved by separating the biflavones using adsorber resins
10 and/or separating 4'-O-methyl pyridoxine via acidic ion exchangers.

In order to produce the extracts according to the present invention, preferably the following method is carried out, wherein steps a) to i) as well as m) are essentially described already in DE 39 40 091 C2, the total disclosure content of which, in
15 particular with respect to the details of the method, shall be incorporated into the present application by reference:

- a) extracting the leaves of ginkgo biloba (drug) at a temperature of 20-100°C, preferably 40-60°C, using water-containing acetone having a content of 20-90% by weight, a water-containing alkanol having 1 to 3 carbon atoms (methanol, ethanol,
20 n-propanol and isopropanol) and a content of 20-90% by weight or anhydrous methanol, wherein the ratio drug : solvent amounts to 1:4 to 1:20, preferably 1:5 to 1:10,
- b) separating the organic solvent by evaporating the solvent to a content of 10% by weight at the most (preferably 5% by weight at the most), wherein water may
25 be added during the final distillation steps,
- c) diluting the remaining aqueous solution with water to a solids content of 5 to 25% by weight (preferably 15 to 20% by weight), cooling the solution to a temperature below 25°C (preferably about 10 to 12°C) and allowing the remaining aqueous solution to stand until a precipitation is formed,
- 30 d) treating the remaining aqueous solution with ammonium sulfate (preferably to a content of up to 40% by weight, particularly preferred 25 to 35% by weight) and subsequently extracting the solution at least once using methyl ethyl ketone or a

mixture of methyl ethyl ketone and acetone (preferably in a ratio of 9:1 to 4:6, in particular 6:4),

e) concentrating the extract obtained (preferably to a solids content of 50 to 70% by weight) and diluting the extract with a mixture of ethanol and water to obtain a
5 solution containing 50% by weight water and 50% by weight ethanol and having a solids content of 10% by weight,

f) treating the solution with a lead compound (preferably with lead acetate, lead hydroxide acetate, lead nitrate or an aqueous suspension of lead hydroxide) until a change in colour from brown to umber occurs, or an insoluble polyamide (preferably polyamide 6, polyamide 6,6 or crosslinked polyvinylpyrrolidone (polyvi-
10 done)),

g) extracting the filtered solution using an aliphatic or cycloaliphatic solvent having a boiling point of 60 to 100°C,

h) concentrating the remaining aqueous-alcoholic solution, subsequently treating
15 the solution with ammonium sulfate and extracting the solution with methyl ethyl ketone and ethanol, (the solution is preferably concentrated under reduced pressure to an ethanol content of about 5% at the most, added with ammonium sulfate to a content of 20% by weight and extracted with the solvent in a ratio of 8:2 to 5:5, preferably 6:4),

i) concentrating the organic phase obtained to a solids content of 50 to 70% by
20 weight,

k) dissolving the concentrate in an aqueous alkanol having 1 to 3 carbon atoms and optionally filtering the product,

l) filtration over an adsorber resin and/or an ion exchanger, wherein the sub-
25 stances to be removed are retained on the resin and

m) drying the concentrate (preferably at a maximum temperature of 60 to 80°C) under reduced pressure to obtain a dry extract having a water content of less than 5%.

30 It is the advantage of the method according to the present invention that the depletion step is carried out at the end of the method. Thus, a smaller amount of the

ion exchanger has to be employed and particularly low contents of 4'-O-methyl pyridoxine and/or biflavones are obtained.

Preferred adsorber resins in step l) are resins based on optionally substituted styrenes/divinylbenzenes such as Diaion HP-20, HP-21 or Sepabeads SP-207 and SP-850. Preferred ion exchangers are strongly acidic ion exchangers such as Merck I or Amberlite IR-120. The filtration optionally includes the usual rinsing using further solvent. The aqueous alkanol in step k) is aqueous methanol, ethanol, n-propanol or isopropanol having a content of 20-90% by weight, respectively, preferably aqueous ethanol, a concentration of 40 to 60% by weight being particularly preferred.

A subject of the present invention are extracts, in particular dry extracts, which are obtainable by the method according to the present invention and which exhibit the following contents:

- less than 20 ppm, preferably less than 10 ppm and in particular less than 2 ppm 4'-O-methyl pyridoxine and/or
- less than 20 ppm, preferably less than 10 ppm and in particular less than 5 ppm biflavones.

Furthermore, subject of the present invention are extracts, in particular dry extracts, which are obtainable by the method according to the present invention and which exhibit the following further contents:

- 20 to 30% by weight, preferably 22.0 to 27.0% by weight flavonoids,
- 4.5 to 8.5% by weight, preferably 5.0 to 7.0% by weight terpene lactones and
- less than 10 ppm, preferably not more than 5 ppm ginkgolic acids.

A further subject of the present invention are extracts, in particular dry extracts, which are obtainable by the method according to the present invention and which exhibit the following contents:

- 20 to 30% by weight, preferably 22.0 to 27.0% by weight flavonoids,
- 2.5 to 4.5% by weight ginkgolides A, B and C in total, preferably 2.8 to 3.4% by weight ginkgolides A, B and C in total,
- 5 • 2.0 to 4.0% by weight, preferably 2.6 to 3.2% by weight bilobalide,
- less than 10 ppm, preferably not more than 5 ppm ginkgolic acids and
- less than 20 ppm, preferably less than 10 ppm, particularly preferred less than 2 ppm of 4'-O-methyl pyridoxine and/or
- less than 20 ppm, preferably less than 10 ppm and particularly preferred less
- 10 than 5 ppm of biflavones.

According to the European Pharmacopoeia, dry extracts generally have a dry residue of at least 95% by weight.

- 15 The extracts according to the present invention can be administered in the form of powders, granules, tablets, dragées (coated tablets) or capsules, preferably orally. In order to prepare tablets, the extract is mixed with suitable pharmaceutically acceptable adjuvants such as lactose, cellulose, silicon dioxide, croscarmellose and magnesium stearate, and pressed into tablets which are optionally provided with a
- 20 suitable coating, for example made of hydroxymethylcellulose, polyethyleneglycol, pigments (such as titanium dioxide, iron oxide) and talcum. The extract according to the present invention can also be filled into capsules, optionally under the addition of adjuvants such as stabilizers, fillers and the like. The dosage is such that 10 to 2000 mg, preferably 50 to 1000 mg and particularly preferred 100 to 500 mg
- 25 extract are administered per day.

- A further subject of the present invention are medicaments, food products and other preparations containing these extracts, optionally in combination with other substances such as active ingredients and/or pharmaceutically acceptable adju-
- 30 vants. The term "food product" as used herein particularly refers to dietetic food products, dietary supplement products as well as medical food and dietary supplements.

Examples

Preparation of the extract solution used in Examples 1 and 2 as well as in 5 Comparative Examples 1 and 2

Dried leaves of ginkgo biloba (drug) were extracted at a temperature of about 50°C using seven times their weight (w/w) (i.e. drug-solvent ratio is 1:7) made up of acetone/water 60/40 (w/w) (step a)).

10

The organic solvent was largely separated from the extract (step b)) and the remaining concentrated aqueous solution was diluted with water to a solids content of about 10% by weight. The solution was cooled to a temperature of about 12°C under agitation and the resulting precipitate was removed (step c)).

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About 30% by weight ammonium sulfate was added to the remaining aqueous solution and the solution formed was extracted using a mixture of methyl ethyl ketone and acetone in a ratio of 6:4 (step d)).

20

The extract obtained was largely concentrated and the concentrate thus obtained was diluted with water and ethanol such that a solution containing 50% by weight water and 50% by weight ethanol at a solids content of about 10% by weight was obtained (step e)). The solution was filtered and the filtrate was added with an aqueous solution of lead hydroxide acetate (step f)).

25

After a further filtration, the remaining aqueous alcoholic solution was extracted with heptane (step g)).

30

Subsequently, the remaining aqueous alcoholic solution was concentrated under reduced pressure to an ethanol content of about 5% and ammonium sulfate was added such that a content of about 20% by weight based on the water content was achieved.

The solution obtained was extracted using a mixture of methyl ethyl ketone and ethanol in a ratio of 6:4 (step h)).

- 5 The resulting organic phase was largely concentrated (solids content of about 55% by weight) (step i)). Then the product was diluted with ethanol and water such that a solution having an ethanol content of about 75% and a solids content of about 15% was obtained (step k)).

10 **Example 1: Ginkgo extract having a reduced content of 4'-O-methyl pyridoxine**

1249 g of the extract solution obtained according to the preparation method described above was adjusted with water to an ethanol content of about 40% (w/w)
15 and applied to 50 ml (corresponding to 0.28 ml/g of the dry extract) Amberlite IR-120 (strongly acidic ion exchanger) (column: about 2 x 16.5 cm; flow: about 8 ml/min) (step l)), followed by rinsing with 200 ml of 50% ethanol (w/w). The resulting solution was concentrated on a rotary evaporator and dried in vacuum at 50°C (step m)): 179.4 g.

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As can be seen from the following table, a content of 4'-O-methyl pyridoxine of 0.4 ppm was achieved without essentially modifying the contents of ingredients which are relevant with respect to the efficacy.

25 **Comparative Example 1: Ginkgo extract without depletion according to step l) of the present invention**

98.4 g of the extract solution employed in Example 1 were concentrated by evaporation and dried in vacuum at 50°C: 14.7 g.

30

Example 2a): Ginkgo extract having a reduced content of biflavones

500 g of the extract solution obtained according to the preparation method described above were adjusted with water to an ethanol content of about 40% (w/w) and applied to 400 ml (corresponding to 5.7 ml/g of the dry extract) Sepabeads SP-850 (adsorber resin) (column: about 4 x 32 cm; flow: about 8 ml/min) (step l)), followed by rinsing with about 2 l of 40% ethanol (w/w) in total. 10% of the resulting solution (259.3 g) were concentrated on a rotary evaporator and freeze-dried (step m)): 6.92 g. The remaining 90% of the solution were employed in Example 2b).

As can be seen from the following table, biflavone contents of less than 1 ppm were achieved, respectively, without essentially modifying the contents of ingredients which are relevant with respect to the efficacy.

Example 2b): Ginkgo extract having a reduced content of 4'-O-methyl pyridoxine and biflavones

90% of the solution obtained in Example 2a) (2334.1 g) were applied to 25 ml (corresponding to 0.40 ml/g of the dry extract) Amberlite IR-120 (strongly acidic ion exchanger) (column: about 1.5 x 11 cm; flow: about 10 ml/min) (step l)), followed by rinsing with 100 ml of 40% ethanol (w/w). The resulting solution was concentrated on a rotary evaporator and freeze-dried (step m)): 61.67 g.

As can be seen from the following table, a content of 4'-O-methyl pyridoxine of 0.4 ppm and contents of biflavones of less than 1 ppm were achieved, respectively, without essentially modifying the contents of ingredients that are relevant with respect to the efficacy.

Comparative Example 2: Ginkgo extract without depletion according to step l) of the present invention

5 50 g of the extract solution employed in Example 2a) were concentrated by evaporation and freeze-dried: 7.05 g.

Comparative Example 3: Ginkgo extract with biflavone depletion and 4'-O-methyl pyridoxine depletion after treatment with a lead compound (step f)) according to EP 1 037 646 B1

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Dried leaves of ginkgo biloba (drug) were extracted at a temperature of about 50°C using seven times their weight (w/w) (i.e. drug-solvent ratio is 1:7) made up of acetone/water 60/40 (w/w).

15 The organic solvent was largely separated from the extract and the remaining concentrated aqueous solution was diluted with water to a solids content of about 10% by weight, cooled to a temperature of about 12°C under agitation and the resulting precipitate was removed.

20 About 30% by weight ammonium sulfate was added to the remaining aqueous solution and the solution formed was extracted using a mixture of methyl ethyl ketone and acetone in a ratio of 6:4.

25 The extract obtained was largely concentrated and the concentrate thus obtained was diluted with water and ethanol such that a solution containing 50% by weight water and 50% by weight ethanol at a solids content of about 10% by weight was obtained. The solution was filtered, the filtrate was added with an aqueous solution of lead hydroxide acetate and filtered again (step f)).

30 150 ml of the solution thus obtained (portion of dry extract: 7.1%; ethanol content: about 50%) were stirred with 2.5 g (corresponding to 0.17 g/g of the final product) active carbon for 30 minutes at room temperature. The filtrate was applied to a

column with 4.2 ml (corresponding to 0.28 ml/g of the final product) strongly acidic cation exchanger (Merck I). The ion exchanger column was washed with 4 x 5 ml EtOH (50%).

- 5 The combined eluates were extracted by shaking using 3 x 70 ml n-hexane. The ethanol/water phase was concentrated on a rotary evaporator and diluted with water to 100 g. The extract solution thus obtained was extracted with 1 x 40 ml, 2 x 30 ml and 1 x 20 ml of a mixture of methyl ethyl ketone:ethanol in a ratio of 3:2, wherein 15 g ammonium sulfate was added to achieve phase separation.

10

The organic phase was stirred for 1 h at room temperature after adding 25 g ammonium sulfate. The undissolved ammonium sulfate as well as the aqueous phase formed were separated. The extract solution was concentrated on a rotary evaporator at 50°C and dried in vacuum at 60°C: 14.8 g.

15

As can be seen from the following table, a 4'-O-methyl pyridoxine content of 52 ppm and a biflavone content of 31.3 ppm and, thus, significantly higher contents and inferior depletion results are achieved compared to Example 2b) according to the present invention.

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The determination of the contents of flavonoids, terpene lactones and ginkgolic acids listed in Table 1 was carried in accordance with the DAB.

- 25 The quantitative determination of 4'-O-methyl pyridoxine was performed by means of HPLC and fluorescence detection after enrichment via Sep Pak Plus C18 cartridges, followed by enrichment via Bond Elut LRC-SCX cartridges, wherein a Phenomenex Prodigy 5 μ ODS-3 HPLC column and a gradient of trifluoroacetic acid (pH 2) : methanol = 9:1 (v/v) $\rightarrow$ methanol, were employed.

- 30 The determination of the contents of the biflavones listed in Table 1 was performed by means of HPLC and UV detection at 340 nm via a Waters Nova Pak

C18 column using a gradient of water: acetonitrile of 9:1 (v/v) + 0.3% by volume phosphoric acid (85%) → acetonitrile + 0.3% by volume phosphoric acid.

In each case, calibration was performed using reference substances having a
5 high, defined purity grade.

Table 1: Composition of the extracts according to examples given above

Extract according to	Example 1	Comparative Example 1	Example 2a)	Example 2b)	Comparative Example 2	Comparative Example 3
Flavonoids [%]	24.86	24.51	27.17	25.32	25.89	22.78
Ginkgolides A, B, C [%]	3.28	2.94	2.97	3.04	2.97	3.40
Bilobalide [%]	3.17	3.03	3.18	3.14	3.07	3.07
Terpene lactones [%]	6.45	5.97	6.15	6.18	6.04	6.47
Ginkgolic acids [ppm]	< 5	< 5	< 5	< 5	< 5	< 5
4'-O-Methyl pyridoxine [ppm]	0.4	104.5	164	0.8	89	52
Amentoflavone [ppm]	68.5	78.7	n. d. (< 1)	n. d. (< 1)	81.6	12.6
Bilobetin [ppm]	123.3	120.4	n. d. (< 1)	n. d. (< 1)	136.4	10.4
Ginkgetin [ppm]	86.3	82.1	n. d. (< 1)	n. d. (< 1)	86.1	2.7
Isoginkgetin [ppm]	103.3	104.4	n. d. (< 1)	n. d. (< 1)	111.5	4.8
Sciadopitysin [ppm]	44.0	43.0	n. d. (< 1)	n. d. (< 1)	44.0	0.8
Total amount of biflavones [ppm]	425.4	428.6	n. d. (< 5)	n. d. (< 5)	459.6	31.3

n. d. = not detectable

Claims

1. Method for producing an extract from ginkgo biloba having a reduced content of 4'-O-methyl pyridoxine and/or biflavones, characterized by the following steps of the method:
- a) extracting leaves of ginkgo biloba using water-containing acetone, a water-containing alkanol having 1 to 3 carbon atoms or anhydrous methanol,
 - b) separating the organic solvent by concentrating the solvent to a content of 10% by weight at the most, wherein water may be added during the final distillation steps,
 - c) diluting the remaining aqueous solution with water to a solids content of 5 to 25% by weight, cooling to a temperature below 25°C and allowing to stand until a precipitate is formed,
 - d) treating the remaining aqueous solution with ammonium sulfate and subsequently extracting at least once using methyl ethyl ketone or a mixture of methyl ethyl ketone and acetone,
 - e) concentrating the extract obtained and diluting with an ethanol-water mixture to obtain a solution containing 50% by weight water and 50% by weight ethanol at a solids content of 10% by weight,
 - f) treating the solution with a lead compound or an insoluble polyamide,
 - g) extracting the filtered solution using an aliphatic or cycloaliphatic solvent having a boiling point of 60 to 100°C,
 - h) concentrating the remaining aqueous-alcoholic solution, followed by treating with ammonium sulfate and extracting with methyl ethyl ketone and ethanol,
 - i) concentrating the organic phase obtained to a solids content of 50 to 70% by weight,
 - k) dissolving the concentrate in an aqueous alkanol having 1 to 3 carbon atoms and optionally filtering,
 - l) filtration over an adsorber resin to reduce the content of biflavones and/or an acidic ion exchanger to reduce the content of 4'-O-methyl pyridoxine, wherein the substances to be removed are retained on the resin and/or the acidic ion exchanger and

m) drying the concentrate at reduced pressure to obtain a dry extract having a water content of less than 5%.

- 5 2. Method according to claim 1, wherein the aqueous alkanol in step k) is aqueous ethanol.
3. Method according to claim 2, wherein the aqueous ethanol is ethanol having a content of 40 to 60% by weight.
- 10 4. Method according to any one of claims 1 to 3, wherein the adsorber resin in step l) is a resin based on optionally substituted styrenes/divinylbenzenes.
5. Method according to any one of claims 1 to 3, wherein the acidic ion exchanger in step l) is a strongly acidic ion exchanger.
- 15 6. Extract from leaves of ginkgo biloba, characterized by having the following contents:
 - less than 10 ppm 4'-O-methyl pyridoxine and/or
 - less than 20 ppm biflavones selected from the group comprising
- 20 amentoflavone, bilobetin, ginkgetin, isoginkgetin and sciadopitysin.
7. Extract according to claim 6, characterized by having a content of less than 2 ppm 4'-O-methyl pyridoxine.
- 25 8. Extract according to claim 6 or 7, characterized by having a content of less than 10 ppm biflavones.
9. Extract according to claim 6 or 7, characterized by having a content of less than 5 ppm biflavones.

10. Extract according to any of claims 6 to 9, characterized by having the following contents:
- 20 to 30% by weight flavonoids, selected from the group comprising quercetin, kaempferol and isorhamnetin glycosides,
 - 5 • 4.5 to 8.5% by weight terpene lactones and
 - less than 10 ppm ginkgolic acids.
11. Extract according to any of claims 6 to 9, characterized by having the following contents:
- 10 • 22.0 to 27.0 % by weight flavonoids, selected from the group comprising quercetin, kaempferol and isorhamnetin glycosides,
 - 5.0 to 7.0% by weight terpene lactones and
 - 5 ppm ginkgolic acids at the most.
- 15 12. Extract according to any of claims 6 to 9, characterized by having the following contents:
- 22.0 to 27.0 % by weight flavonoids, selected from the group comprising quercetin, kaempferol and isorhamnetin glycosides,
 - 2.8 to 3.4% by weight ginkgolides A, B and C in total,
 - 20 • 2.6 to 3.2% by weight bilobalid and
 - 5 ppm ginkgolic acids at the most.
13. Use of an extract according to any one of claims 6 to 12 for the manufacture of a medicament, food product or other preparation for the treatment of dementia and symptoms thereof and/or cerebral or peripheral blood circulation disorders.
- 25 14. Medicament, food product or other preparation, characterized by a content of a ginkgo extract according to any one of claims 6 to 12.