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(CVMP)**

FINAL

**GUIDELINE ON DECLARATION OF HERBAL SUBSTANCES AND
HERBAL PREPARATIONS¹ IN HERBAL MEDICINAL
PRODUCTS²/TRADITIONAL HERBAL MEDICINAL PRODUCTS IN
THE SPC³**

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¹ The term "herbal substance" should be considered as equivalent to the term "herbal drug" as defined in the European Pharmacopoeia and the term "herbal preparation" should be considered as equivalent to the term "herbal drug preparation" as defined in the European Pharmacopoeia

² Throughout the guideline and unless otherwise specified, the term "herbal medicinal product" includes "traditional herbal medicinal product"

³ SPC: Summary of Product Characteristics.

**GUIDELINE ON DECLARATION OF
HERBAL SUBSTANCES AND HERBAL PREPARATIONS IN HERBAL
MEDICINAL PRODUCTS/TRADITIONAL HERBAL MEDICINAL
PRODUCTS IN THE SPC**

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EXECUTIVE SUMMARY

This guideline outlines the principles for uniform declaration in the SPC of herbal substances/preparations in herbal medicinal products as well as in traditional herbal medicinal products. It focuses on the different types of herbal substances/preparations in relation to the quality documentation given. Examples of declaration of such active substances are provided.

The guideline should be read in conjunction with current EU/(V)ICH guidelines.

1. INTRODUCTION

Common criteria for the declaration shall ensure clear differentiation between different types of herbal substances/preparations and proper description of their qualitative and quantitative particulars. As a result, a precise and consistent description of active substances of herbal medicinal products will be guaranteed within the Community.

The complex composition of herbal substances/preparations, which is essentially determined by various factors like the production process, the extraction solvent, the genuine drug extract ratio (DER genuine), and the type/physical state of the herbal substances/preparations, needs to be stated to guarantee identification and facilitate comparison of herbal substances/preparations. However, it is not feasible to provide full characterisation in the declaration as the declaration should be kept as short and precise as possible.

The declaration is primarily intended to describe the identity and quantity of the herbal substance/preparation, being the active substance of the herbal medicinal product and should focus on those characteristics found to be useful in ensuring the safety and efficacy of the herbal substance/preparation and herbal medicinal product.

Therefore, a declaration system has been established which reflects the main characteristics of herbal substances/preparations as defined in the respective specifications. For this purpose, general guidance as given in the European Pharmacopoeia (particularly the monographs “Extracts”, “Herbal Drugs”, “Herbal Drug Preparations”, and “Herbal Teas”) as well as in the guidelines listed under *References*, should be followed.

2. SCOPE

This guideline addresses the declaration of herbal substances/preparations when being the active substance of a herbal medicinal product. Examples of standardised, quantified, and other herbal substances/preparations are given.

The guideline addresses only the declaration of herbal substances/preparations in herbal medicinal products (including fixed combinations) in the SPC; it shall apply to herbal medicinal products both for human and veterinary use and to traditional herbal medicinal products for human use. Traditional herbal medicinal products may additionally contain vitamins and/or minerals. Declaration of these chemically defined substances is not covered by this guideline. Reference is given to the INN-system and to “A guideline on summary of product characteristics”.

This guideline reflects the current state of the art at the time it has been written. If necessary, the national regulatory authority/HMPC should be asked for additional guidance, especially in those cases not covered by examples in the guideline.

3. LEGAL BASIS

This guideline supports applications for marketing authorisations or registrations according to Directive 2001/83/EC and Directive 2001/82/EC as amended.

A simplified registration procedure was established for traditional herbal medicinal products for human use under Directive 2001/83/EC as amended by Directive 2004/24/EC. The principals for declaration of herbal substances/preparations in herbal medicinal products apply equally to such traditional herbal medicinal products for human use.

4. MAIN GUIDELINE TEXT

This guideline presents a brief definition of each concept and gives examples of the declaration. Generally, if the classification of a herbal preparation is not unambiguous, alternative proposals should be justified by the applicant and approved by the regulatory authority before being put into effect.

In the different chapters the characteristics, which are generally needed for the declaration of the different kinds of herbal substances/preparations, are given, followed by examples. The examples in the guideline are for illustration purposes only and not intended to reflect binding specifications. Within each example it is shown, what information is needed to form the specific declaration of the active substance of the herbal medicinal product in section 2 of the SPC. In this context it is pointed out that both excipients for adjustment (valid for standardised herbal preparations only) and/or other excipients (e.g. carrier substances) must be declared in section 6.1 of the SPC, preferably listed with a subtitle “excipients of the herbal substance/preparation”. Extraction solvents are to be declared in section 2 of the SPC only. Furthermore, section 5.3 of the (human) SPC provides for the possibility to inform on limits of unwanted (i. e. toxicologically relevant) constituents of herbal substances/preparations for safety reasons, provided that these limits are laid down in the specification as part of the quality documentation.

5. DECLARATION OF HERBAL SUBSTANCES IN THE SPC

It should be noted that this section does not apply to a herbal substance being the starting material of a herbal preparation.

The declaration of a herbal substance should cover the name and the quantity of the herbal substance. The name of the herbal substance is the scientific Latin name of the plant species according to the binomial system (genus, species, variety and author) with the Latin term of the plant part, followed by the [translated] common name of the monograph of the European Pharmacopoeia if available, or else of the Pharmacopoeia of a Member State, if available, or else the common name of the herbal substance (in brackets). In those special cases, where many different Latin plant species apply to the same herbal substance, the list of Latin names could be shortened to the genus name followed by the word “species”, e.g. “*Crataegus* species”. This option is only applicable in cases, where no restrictions concerning the species used are known from the quality documentation. In special cases, where necessary, only the scientific Latin name of the plant species may be used together with the [translated] common term for the plant part. For specific types of herbal substances (e.g. standardised, quantified) additional information may be necessary.

The following characteristics have to be stated in the declaration:

1. Name of the herbal substance.
2. Quantity of the genuine herbal substance.
3. Name and quantity of the constituent(s) with known therapeutic activity (standardised herbal substances), if applicable.
4. Name and quantity (given as a range) of the active markers (quantified herbal substances), if applicable.

5.1 Standardised herbal substances

Standardised herbal substances are adjusted within an acceptable tolerance to a given content of constituents with known therapeutic activity; standardisation is achieved by adding excipients for adjustment to the herbal substance or by blending batches of the herbal substance.

For such herbal substances the name and content of the constituent(s) with known therapeutic activity should be stated. The equivalent quantity of the genuine herbal substance should be given (as a range, if applicable).

Example:

Where a herbal medicinal product contains:

Senna leaf, cut.

Constituents with known therapeutic activity: 2.55 % hydroxyanthracene glycosides, calculated as

sennoside B.

Quantity of the genuine herbal substance as a range: 85 - 96 %.

Excipients for adjustment: 4 - 15 %.

Quantity of the standardised herbal substance (herbal substance and excipients for adjustment) in the herbal medicinal product: 1.3 g/sachet.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 tea sachet contains 1.10 g - 1.25 g *Cassia senna* L. (*C. acutifolia* Delile) and/or *Cassia angustifolia* Vahl, folium (Senna leaf), corresponding to 33 mg hydroxyanthracene glycosides, calculated as sennoside B.

5.2. Quantified herbal substances

Quantified herbal substances are adjusted to a defined range of constituents; adjustments are made by blending batches of herbal substances used in the manufacturing process.

For quantified herbal substances the name of the active markers should be stated and their content should be given in a range. The equivalent quantity of the genuine herbal substance should be given.

Example:

Where a herbal medicinal product contains:

Willow bark, cut.

Quantification: 1.5 - 1.7 % of total salicylic derivatives calculated as salicin.

Quantity of the herbal substance in the herbal medicinal product: 3.0 g/sachet.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 tea sachet contains 3.0 g of various species of genus *Salix* including *S. purpurea* L., *S. daphnoides* Vill. and *S. fragilis* L., cortex (Willow bark), corresponding to 45 mg to 51 mg of total salicylic derivatives, calculated as salicin.

5.3 Other herbal substances

For other herbal substances neither constituents with known therapeutic activity nor active markers are generally known. Therefore these herbal substances are essentially defined by their production process and their specifications.

For other herbal substances the name and content of the analytical marker(s) should not be stated. The quantity of the genuine herbal substance should be given.

Example:

Where a herbal medicinal product contains:

100 g Linseed.

Quantity of the herbal substance in the herbal medicinal product: 1 g/g.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 g herbal medicinal product contains 1 g *Linum usitatissimum* L., semen (Linseed).

6. DECLARATION OF HERBAL PREPARATIONS IN THE SPC

Herbal preparations are diverse in character ranging from simply processed, comminuted plant material to complex processed preparations such as refined extracts. The declaration of a herbal preparation should cover the name of the herbal substance and the definition of the herbal preparation including the physical state, ratio of herbal substance to genuine herbal preparation (DER genuine, also named native DER), and extraction solvent(s) if appropriate. The name of the herbal substance is the scientific Latin name of the plant species according to the binomial system (genus, species, variety and author) with the Latin term of the plant part, followed by the [translated] common name of the monograph of the European Pharmacopoeia if available, or or else of the Pharmacopoeia of a Member State, if available, or else the common name of the herbal substance (in brackets). In those special cases, where many different Latin plant species apply to the same herbal substance, the list of Latin names could be shortened to the genus name followed by the word “species”, e.g. “*Crataegus* species”. This option is only applicable in cases, where no restrictions concerning the species used are known from the quality documentation. In special cases, where necessary, only the scientific Latin name of the plant species may be used together with the [translated] common term for the plant part.

In addition, the declaration of herbal preparations needs to reflect the different extract type (type of herbal preparation) as described in the European Pharmacopoeia.

(i) **‘Standardised herbal preparations’**: are adjusted within an acceptable tolerance to a given content of constituents with known therapeutic activity; standardisation is achieved by adding excipients for adjustment to the herbal preparations or by blending batches of herbal preparations/herbal substances used in the manufacturing process.

For such preparations the name and content of the constituent(s) with known therapeutic activity should be stated. The equivalent quantity of the genuine herbal preparation should be given (as a range, if applicable).

(ii) **‘Quantified herbal preparations’**: are adjusted to a defined range of constituents (active markers); adjustments are made by blending batches of herbal preparations/herbal substances used in the manufacturing process.

For such preparations the name and content of the active markers should be stated in a range. The equivalent quantity of the genuine herbal preparation should be stated, quoting either the corresponding amount of herbal substance (given as a range) or the DER genuine.

(iii) **‘Other herbal preparations’**: are essentially defined by their production process and their specifications.

For such preparations the name and content of the analytical marker(s) should not be stated. The quantity of the genuine herbal preparation should be stated, quoting either the corresponding amount of herbal substance (given as a range) or the DER genuine.

When solvent(s) are used in the manufacture of a herbal preparation (extraction solvent(s)), the name and composition of the solvent(s) used in the first extraction step should be included in the declaration of the herbal medicinal product. If purification procedures are used in the manufacture of a herbal preparation, the word “refined” should be added to the name of the herbal preparation, where applicable.

In the SPC the following wording can be used, as appropriate: “Extraction solvent: <NAME> <COMPOSITION> % V/V” (or % m/m, as applicable).

If a fresh herbal substance is used as a starting material for manufacture of the herbal preparation, this should be added to the name of the herbal preparation, as appropriate.

6.1 Herbal preparations consisting of comminuted or powdered herbal substances

The following characteristics have to be stated in the declaration:

1. Name of the herbal substance used.
2. Physical state of the herbal preparation, if relevant.
3. Quantity of the genuine herbal preparation.

4. Name and quantity of the constituent(s) with known therapeutic activity (standardised herbal preparations), if applicable.
5. Name and quantity (given as a range) of the active markers (quantified herbal preparations), if applicable.

6.1.1 Standardised herbal preparations

Example a): Standardisation by adding excipients for adjustment

Where a herbal medicinal product contains:

Senna leaf, powdered.

Tinnevelly Senna pods, powdered.

Constituents with known therapeutic activity: 3.5 % hydroxyanthracene glycosides, calculated as sennoside B.

Quantity of the genuine herbal preparation as a range: 70 - 95 % (mixture of both senna preparations).

Excipients for adjustment: 5 - 30 %.

Other excipients: 0 %.

Quantity of the standardised herbal preparation (genuine herbal preparation + excipients for adjustment) in the herbal medicinal product: 1 g/g (200 mg Senna leaf, 500 mg - 750 mg Tinnevelly Senna pods and 50 mg - 300 mg excipients for adjustment).

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 g powder contains 200 mg *Cassia senna* L. (*C. acutifolia* Delile) and/or *Cassia angustifolia* Vahl, folium (Senna leaf) and 500 mg - 750 mg *Cassia angustifolia* Vahl, fructus (Tinnevelly Senna pods), corresponding to 35 mg hydroxyanthracene glycosides, calculated as sennoside B .

Example b): Standardisation by mixing herbal preparations

Where a herbal medicinal product contains:

Senna leaf, powdered.

Tinnevelly Senna pods, powdered.

Alexandrian Senna pods, powdered.

Constituent(s) with known therapeutic activity: 2.7 % hydroxyanthracene glycosides, calculated as sennoside B.

Quantity of the genuine herbal preparation (as a range): 100 % genuine herbal preparation (mixture of all three senna preparations).

Other excipients: 0 %.

Quantity of the genuine standardised herbal preparation in the herbal medicinal product: 1 g/g (200 mg Senna leaf, 500 mg - 750 mg Tinnevelly Senna pods, and 50 mg - 300 mg Alexandrian Senna pods).

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 g powder contains 200 mg *Cassia senna* L. (*C. acutifolia* Delile) and/or *Cassia angustifolia* Vahl, folium (Senna leaf), 500 mg - 750 mg *Cassia angustifolia* Vahl, fructus (Tinnevelly Senna pods) and 50 mg - 300 mg *Cassia senna* L. (*C. acutifolia* Delile), fructus (Alexandrian Senna pods), corresponding to 27 mg hydroxyanthracene glycosides, calculated as sennoside B.

6.1.2 Quantified herbal preparations

Example:

Where a herbal medicinal product contains:

Willow bark, powdered.

Quantification: 1.5 - 1.7 % of total salicylic derivatives, calculated as salicin.

Other excipients: 0 %.

Quantity of the genuine herbal preparation in the herbal medicinal product: 2.5 g/sachet.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 tea sachet contains 2.5 g of various species of genus *Salix* including *S. purpurea* L., *S. daphnoides* Vill. and *S. fragilis* L, cortex (Willow bark), corresponding to 37.5 mg to 42.5 mg of total salicylic derivatives, calculated as salicin.

6.1.3 Other herbal preparations

Example:

Where a herbal medicinal product contains:

Valerian root, powdered.

Other excipients: 0 %.

Quantity of the genuine herbal preparation in the herbal medicinal product: 300 mg/capsule.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 capsule contains 300 mg of *Valeriana officinalis* L. s.l., radix (Valerian root).

6.2 Herbal preparations produced by steps which exceed comminution/powdering (e.g. extracts)

The following characteristics have to be stated in the declaration:

1. Name of the herbal substance used.
2. Type/physical state of the herbal preparation.
3. Quantity of the genuine herbal preparation.
4. Name and quantity of the constituent(s) with known therapeutic activity (standardised herbal preparations), if applicable.
5. Name and quantity (given as a range) of the active markers (quantified herbal preparations), if applicable.
6. Drug extract ratio (DER genuine) or equivalence in the quantity of the herbal substance (as a range) (quantified and other herbal preparations).
7. Name and composition of extraction solvent(s).

6.2.1 Standardised extracts

Example:

Where a herbal medicinal product contains:

Dry extract from Horse chestnut seed

Constituent(s) with known therapeutic activity: 19 % triterpene glycosides, calculated as anhydrous β -aescin.

Quantity of the genuine extract (as a range): 70 - 95 % genuine extract.

DER genuine: 5 – 8 : 1.

Excipients for adjustment: 30 - 5 %.

Other excipients: 0 %.

Extraction solvent: Methanol 80 % V/V.

Quantity of the standardised extract (genuine herbal preparation and excipients for adjustment) in the herbal medicinal product: 200 mg/capsule.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 capsule contains 140 mg - 190 mg of extract (as dry extract) from *Aesculus hippocastanum* L., semen (Horse chestnut seed) corresponding to 38 mg triterpene glycosides, calculated as anhydrous β -aescin.

Extraction solvent: Methanol 80 % V/V.

6.2.2 Quantified extracts

Example:

Where a herbal medicinal product contains:

Dry extract from Ginkgo leaf, refined.

Quantity of the genuine extract: 100 % genuine extract.

DER genuine: 35 – 67 : 1.

Quantification: 22.0 to 27.0 % of flavonoids expressed as flavone glycosides.

2.8 to 3.4 % of ginkgolides A, B and C.

2.6 to 3.2 % of bilobalide.

Other excipients: 0 %.

First extraction solvent: Acetone 60 % m/m.

Quantity of the genuine quantified extract in the herbal medicinal product: 60 mg/capsule.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 capsule contains 60 mg of extract (as dry extract, refined) from *Ginkgo biloba* L., folium (Ginkgo leaf) (35 – 67 : 1), corresponding to:

13.2 mg to 16.2 mg of flavonoids expressed as flavone glycosides

1.68 mg to 2.04 mg of ginkgolides A, B and C

1.56 mg to 1.92 mg of bilobalide.

First extraction solvent: Acetone 60 % m/m.

or

1 capsule contains 60 mg of extract (as dry extract, refined) from *Ginkgo biloba* L., folium (equivalent to 2.1 g – 4.0 g of Ginkgo leaf), corresponding to:

13.2 mg to 16.2 mg of flavonoids expressed as flavone glycosides

1.68 mg to 2.04 mg of ginkgolides A, B and C

1.56 mg to 1.92 mg of bilobalide.

First extraction solvent: Acetone 60% m/m.

6.2.3 Other extracts

Example a): Other extracts such as dry extracts

Where a herbal medicinal product contains:

Dry extract from Valerian root.

Quantity of the genuine extract: 80 % genuine extract.

DER genuine: 3 – 6 : 1

Other excipients: 20 %.

Extraction solvent: Ethanol 70 % V/V.

Quantity of the dry extract (genuine herbal preparation and other excipients) in the herbal medicinal product: 200 mg/capsule.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 capsule contains 160 mg of extract (as dry extract) from *Valeriana officinalis* L. s.l., radix (Valerian root) (3 – 6 : 1).

Extraction solvent: Ethanol 70 % V/V.

or

1 capsule contains 160 mg of extract (as dry extract) from *Valeriana officinalis* L. s.l., radix (equivalent to 480 mg – 960 mg of Valerian root).

Extraction solvent: Ethanol 70 % V/V.

Example b): Other extracts such as liquid extracts

Where a herbal medicinal product contains:

Liquid extract from Matricaria flower..

Quantity of the genuine extract: 100 % genuine extract.

DER genuine: 1 : 1.

Other excipients: 0 %.

Extraction solvent: 2.5 parts ammonia solution 10 % m/m
50 parts of ethanol 96 % V/V
47.5 parts of water.

Quantity of the genuine liquid extract in the herbal medicinal product: 1 ml/ml.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 ml [≅ ... g] of oral liquid contains 1 ml of liquid extract from *Matricaria recutita* L. (*Chamomilla recutita* (L.) Rauschert), flos (Matricaria flower) (1 : 1).

Extraction solvent: Ammonia solution 10 % m/m / ethanol 96 % V/V / water (2.5/50/47.5).

or

1 ml [≅ ... g] of oral liquid contains 1 ml of liquid extract from *Matricaria recutita* L. (*Chamomilla recutita* (L.) Rauschert), flos (equivalent to 1 g Matricaria flower).

Extraction solvent: Ammonia solution 10 % m/m / ethanol 96 % V/V / water (2.5/50/47.5).

Example c): Other extracts such as tinctures

Where a herbal medicinal product contains:

Tincture from Valerian root.

Quantity of the genuine extract: 100 % genuine extract.

DER genuine: 1 : 4.0 - 4.5.

Other excipients: 0 %.

Extraction solvent: Ethanol 70 % V/V.

Quantity of the tincture in the herbal medicinal product: 1 ml/ml.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 ml [≅ ... g] of oral liquid contains 1 ml of tincture from *Valeriana officinalis* L. s.l., radix (Valerian root) (1 : 4.0 - 4.5).

Extraction solvent: Ethanol 70 % V/V.

or

1 ml [\cong ... g] of oral liquid contains 1 ml of tincture from *Valeriana officinalis* L. s.l., radix (equivalent to 220 mg - 250 mg Valerian root).

Extraction solvent: Ethanol 70 % V/V.

Example d): Other extracts such as dry extracts from a mixture

Where a herbal medicinal product contains:

Dry extract from 3 parts Valerian root
 2 parts Hop strobile
 2 parts Melissa leaf.

Quantity of the genuine extract: 80 % genuine extract.

DER genuine: 4 – 7 : 1.

Other excipients: 20 %.

Extraction solvent: Ethanol 70 % V/V.

Quantity of the dry extract (genuine herbal preparation and other excipients) in the herbal medicinal product: 200 mg/capsule.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 capsule contains 160 mg of extract (as dry extract) (4 – 7 : 1) from *Valeriana officinalis* L. s.l., radix (Valerian root) / *Humulus lupulus* L., flos (Hop strobile) / *Melissa officinalis* L., folium (Melissa leaf) (3/2/2).

Extraction solvent: Ethanol 70 % V/V.

or

1 capsule contains 160 mg of extract (as dry extract) (equivalent to 0.64 g – 1.1 g mixture of the herbal substances) from *Valeriana officinalis* L. s.l., radix (Valerian root) / *Humulus lupulus* L., flos (Hop strobile) / *Melissa officinalis* L., folium (Melissa leaf) (3/2/2).

Extraction solvent: Ethanol 70 % V/V.

6.3 Herbal preparations not covered by 6.1 or 6.2

The following characteristics have to be stated in the declaration:

1. Name of the herbal substance used.
2. Type of the herbal preparation.
3. Quantity of the genuine herbal preparation.
4. Drug extract ratio (DER genuine) or equivalent quantity of the herbal substance (as a range), if applicable.
5. Name and composition of extraction solvent(s), if applicable.

6.3.1 Other herbal preparations such as essential oils

Example:

Where a herbal medicinal product contains:

Peppermint oil.

Quantity of the essential oil: 100 % essential oil.

Other excipients: 0 %.

Quantity of the essential oil in the herbal medicinal product: 81 mg/ml oral liquid.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 ml [\cong ... g] of oral liquid contains 81 mg of *Mentha × piperita* L., aetheroleum (peppermint oil).

6.3.2 Other herbal preparations such as expressed juices

Example:

Where a herbal medicinal product contains:

Expressed juice from fresh purple coneflower herb.

Quantity of the genuine expressed juice: 100% genuine expressed juice.

DER genuine: 1.2 - 1.5 : 1.

Other excipients: 0 %.

Quantity of the genuine expressed juice in the herbal medicinal product: 1 ml/ml oral liquid.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 ml [≅ ... g] of oral liquid contains 1 ml of expressed juice from *Echinacea purpurea* (L.) Moench, herba (fresh purple coneflower herb) (1.2 - 1.5 : 1).

or

Each 1 ml [≅ ... g] of oral liquid contains 1 ml of expressed juice from *Echinacea purpurea* (L.) Moench, herba (equivalent to 1.2 g – 1.5 g fresh purple coneflower herb).

6.3.3 Other herbal preparations such as processed exudates

Example:

Where a herbal medicinal product contains:

Tincture from Myrrh.

Quantity of the processed exudate: 100 % processed exudate.

DER genuine: 1 : 4.0 - 4.5.

Other excipients: 0 %.

Extraction solvent: Ethanol 90 % V/V.

Quantity of the tincture in the herbal medicinal product: 25 mg/ml oral liquid.

The declaration in section 2 of the SPC of the herbal medicinal product is:

1 ml [≅ ... g] of oral liquid contains 25 mg of tincture from *Commiphora molmol* Engler and/or other species of *Commiphora* (Myrrh) (1 : 4.0 - 4.5).

Extraction solvent: Ethanol 90 % V/V.

or

Each ml [≅ ... g] of oral liquid contains 25 mg of tincture from *Commiphora molmol* Engler and/or other species of *Commiphora* (equivalent to 5.5 mg - 6.3 mg Myrrh).

Extraction solvent: Ethanol 90 % V/V.

DEFINITIONS

Constituents with known therapeutic activity: are chemically defined substances or groups of substances which are generally accepted to contribute substantially to the therapeutic activity of a herbal substance, a herbal preparation or a herbal medicinal product.

Drug extract ratio (DER): means the ratio between the quantity of herbal substance used in the manufacture of a herbal preparation and the quantity of herbal preparation obtained. The number (given as the actual range) written before the colon is the relative quantity of the herbal substance; the number written after the colon is the relative quantity of the herbal preparation obtained.

Excipients: In general, excipients may be defined as constituents of the medicinal product other than the active substance(s). However, in the context of this guideline only two categories of excipients are addressed:

Excipients for adjustment are used for standardisation of herbal substances/preparations.

Other excipients are technological excipients (e.g. carrier substances) which may be part of herbal preparations.

Extraction solvents: are solvents which are used for the extraction process.

Genuine (Native) herbal preparation: refers to the preparation without excipients, even if for technological reasons the genuine herbal preparation is not available. However, for soft and liquid herbal preparations the genuine herbal preparation may contain variable amounts of (extraction) solvent.

Ratio of herbal substance to genuine herbal preparation (DER genuine): is the ratio of the quantity of the herbal substance to the quantity of the resulting genuine herbal preparation. The number (given as the actual range) written before the colon is the relative quantity of the herbal substance; the number written after the colon is the relative quantity of the genuine herbal preparation obtained.

Herbal medicinal products: any medicinal product, exclusively containing as active substances one or more herbal substances or one or more herbal preparations, or one or more such herbal substances in combination with one or more such herbal preparations.

Herbal preparations: are obtained by subjecting herbal substances to treatments such as extraction, distillation, expression, fractionation, purification, concentration or fermentation. These include comminuted or powdered herbal substances, tinctures, extracts, essential oils, expressed juices and processed exudates.

Herbal substances: all mainly whole, fragmented or cut plants, plant parts, algae, fungi, lichen in an unprocessed, usually dried form but sometimes fresh. Certain exudates that have not been subjected to a specific treatment are also considered to be herbal substances. Herbal substances are precisely defined by the plant part used and the botanical name according to the binomial system (genus, species, variety and author).

Herbal teas: consist exclusively of one or more herbal substance(s) intended for oral aqueous preparations by means of decoction, infusion or maceration. The preparation is prepared immediately before use. Herbal teas are usually supplied in bulk form or in sachets.

Markers: are chemically defined constituents or groups of constituents of a herbal substance, a herbal preparation or a herbal medicinal product which are of interest for control purposes independent of whether they have any therapeutic activity. Markers serve to calculate the quantity of herbal substance(s) or herbal preparation(s) in the herbal medicinal product if the marker has been quantitatively determined in the herbal substance or herbal preparation.

There are two categories of markers:

Active markers are constituents or groups of constituents which are generally accepted to contribute to the therapeutic activity.

Analytical markers are constituents or groups of constituents that serve for analytical purposes.

Quantification: means adjusting the herbal substance or herbal preparation to a defined range of constituents (active markers) exclusively achieved by blending different batches of herbal substances

and/or herbal preparations (e.g. quantified extract).

Specification: A list of tests, references to analytical procedures, and appropriate acceptance criteria which are numerical limits, ranges, or other criteria for the tests described. It establishes the set of criteria to which a herbal substance/preparation or herbal medicinal product should conform to be considered acceptable for its intended use. "Conformance to specifications" means that the herbal substance/preparation and/or herbal medicinal product, when tested according to the listed analytical procedures, will meet the listed acceptance criteria. Specifications are binding quality standards that are agreed to between the appropriate governmental regulatory agency and the applicant.

Standardisation: means adjusting the herbal substance/preparation to a defined content of a constituent or a group of constituents with known therapeutic activity respectively either by adding excipients or by blending batches of the herbal substance and/or herbal preparation (e.g. standardised extracts).

Traditional herbal medicinal products: are medicinal products for human use that fulfil the conditions laid down in article 16a (1) of Directive 2001/83/EC, as amended.

REFERENCES

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