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Committee on Herbal Medicinal Products (HMPC)

European Union herbal monograph on *Species digestivae* Draft

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BG (bulgarski): Чай при стомашно-чревни оплаквания, комбинации	LT (lietuvių kalba): Virškinimą skatinantis ir gerinantis mišinys
CS (čeština): žaludeční čajová směs	LV (latviešu valoda): Tēja gremošanas veicināšanai/gremošanas sistēmai
DA (dansk): Fordøjelseste	MT (Malti): speci digestivi
DE (Deutsch): Verdauungstees	NL (Nederlands): Maag- en darmkruiden
EL (elliniká): Μίγματα φυτικών ειδών για εγχύματα/αφεψήματα με ευστόμαχη δράση	PL (polski):
EN (English): Herbal tea combinations for use in gastrointestinal complaints	PT (português): associações de substâncias vegetais para utilização em tisanas com acção digestiva / estomáquica
ES (español): Combinaciones de sustancias vegetales digestivas para uso en tisana	RO (română): ceai gastric
ET (eesti keel):	SK (slovenčina): žalúdková čajovina
FI (suomi): ruoansulatusvaivoissa perinteisesti käytetyt yhdistelmäteevalmisteet	SL (slovenščina): zdravlilni čaji za prebavo/želodec
FR (français): espèces digestives /stomachiques	SV (svenska): örtteer vid matsmältningsbesvär/mag-tarmbesvär
HR (hrvatski): čajevi za probavu/želudac	IS (íslenska):
HU (magyar): Emésztést elősegítő teakeverék	NO (norsk):
IT (italiano): Specie composte per tisane digestive/stomachiche	

European Union herbal monograph on *Species digestivae*

1. Name of the medicinal product

To be specified for the individual finished product.

2. Qualitative and quantitative composition^{1, 2}

Well-established use	Traditional use
	<p>With regard to the registration application of Article 16d(1) of Directive 2001/83/EC</p> <p>Herbal tea combinations containing several herbal substances out of the following list:</p> <p><i>Achillea millefolium</i> L., herba, flos</p> <p><i>Aloysia citrodora</i> Paláu, folium (Verbenae citriodora folium)</p> <p><i>Althaea officinalis</i> L., radix</p> <p><i>Artemisia absinthium</i> L., herba</p> <p><i>Carum carvi</i> L., aetheroleum, fructus</p> <p><i>Centaurium erythraea</i> Rafn. S.l., herba</p> <p><i>Cinnamomum verum</i> J. S. Presl, (<i>Cinnamomum zeylanicum</i> Nees), cortex</p> <p><i>Curcuma xanthorrhiza</i> Roxb. (<i>C. xanthorrhiza</i> D. Dietrich), rhizoma</p> <p><i>Foeniculum vulgare</i> Miller subsp. <i>vulgare</i> var. <i>dulce</i> (Miller) Thellung., fructus</p> <p><i>Foeniculum vulgare</i> Miller subsp. <i>vulgare</i> var. <i>vulgare</i>, fructus</p> <p><i>Fumaria officinalis</i> L., herba</p> <p><i>Gentiana lutea</i> L., radix</p> <p><i>Glycyrrhiza glabra</i> L. and/or <i>Glycyrrhiza inflata</i> Bat. and/or <i>Glycyrrhiza uralensis</i> Fisch., radix</p> <p><i>Hypericum perforatum</i> L., herba</p> <p><i>Marrubium vulgare</i> L., herba</p>

¹ The declaration of the active substance(s) for an individual finished product should be in accordance with relevant herbal quality guidance.

² The material complies with the respective monographs in the Ph. Eur.

Well-established use	Traditional use
	<p><i>Matricaria recutita</i> L., flos</p> <p><i>Melissa officinalis</i> L., folium</p> <p><i>Mentha x piperita</i> L., folium</p> <p><i>Menyanthes trifoliata</i> L., folium</p> <p><i>Peumus boldus</i> Molina, folium</p> <p><i>Pimpinella anisum</i> L., fructus</p> <p><i>Rosmarinus officinalis</i> L. folium</p> <p><i>Salvia officinalis</i> L., folium</p> <p><i>Taraxacum officinale</i> Weber ex Wigg., radix cum herba</p> <p>i) Herbal substance</p> <p>Not applicable.</p> <p>ii) Herbal preparations</p> <p>Combinations of the comminuted herbal substances (see combinations table below)</p>

	Absinthii herba	Althaeae radix	Anisi fructus	Boldi folium	Carvi fructus	Centaurii herba	Cinnamomi cortex	Curcumae xanthorrhizae rhizoma	Foeniculi amari fructus	Foeniculi dulcis fructus	Fumariae herba	Gentianae radix	Hyperici herba	Liquiritiae radix	Marrubii herba	Matricariae flos	Melissae folium	Menthae piperitae herba	Menyanthis folium	Millefolii herba/flos	Rosmarini folium	Salviae officinalis folium	Taraxaci radix cum herba	Verbenae citriodora folium	Range in combination (excl. excipients)
Absinthii herba						+	+					+					+	+		+					10-50%
Althaeae radix			+						+					+		+		+						+	10%
Anisi fructus		+			+	+			+							+	+			+	+			+	10-40%
Boldi folium											+				+	+		+	+		+		+		10-32%
Carvi fructus			+					+	+							+		+		+			+		10-40%
Centaurii herba	+						+		+			+	+			+		+		+	+		+		10-35%
Cinnamomi cortex	+					+						+													10%
Curcumae xanthorrhizae rhizoma					+													+		+			+		15-20%
Foeniculi amari fructus			+		+	+										+		+	+	+					14-40%
Foeniculi dulcis fructus		+				+							+	+		+		+							10-22%
Fumariae herba				+														+	+		+		+		13%
Gentianae radix	+					+	+			+				+		+				+			+		10-30%
Hyperici herba						+			+							+		+							10%
Liquiritiae radix		+							+							+		+		+		+			10-35%
Marrubii herba				+												+		+					+		25%
Matricariae flos		+			+	+			+								+	+	+	+	+			+	10-40%
Melissae folium	+		+						+							+				+					10-35%
Menthae piperitae herba	+	+	+	+	+	+		+	+		+		+	+	+	+	+		+	+	+	+	+	+	10-40%
Menyanthis folium				+							+							+			+		+		11%
Millefolii herba/flos		+	+	+	+	+			+					+		+	+	+							10-50%
Rosmarini folium			+	+							+					+		+	+				+	+	10-16%
Salviae officinalis folium													+		+		+								10%
Taraxaci radix cum herba				+		+					+							+	+		+				19-25%
Verbenae citriodora folium		+	+													+		+			+				10%

A + indicates the evidence of traditional medicinal use of a certain combination.

Out of these documented combinations the combination partners for an application for a traditional herbal medicinal product may be chosen.

The number of active substances in an herbal tea combination is recommended to be limited to a **maximum of 4**. Further herbal substances may be added as excipients. Their number, amount and function should be justified.

The very right row indicates the traditional amount of the herbal substance in the combinations. The calculation is based on those combination partners only, which contribute to the plausibility of the traditional medicinal use. The amount in a traditional herbal medicinal product should be within this range. The sum of active ingredients should end up with 100%. Excipients are not considered in this range.

3. Pharmaceutical form

Well-established use	Traditional use
	<p>Comminuted herbal substances as herbal tea for oral use.</p> <p>The pharmaceutical form should be described by the European Pharmacopoeia full standard term.</p>

4. Clinical particulars

4.1. Therapeutic indications

Well-established use	Traditional use
	<p>Traditional herbal medicinal product used for the symptomatic relief of mild dyspeptic/gastrointestinal disorders including bloating and flatulence.</p> <p>The product is a traditional herbal medicinal product for use in the specified indication exclusively based upon long-standing use.</p>

4.2. Posology and method of administration ³

Well-established use	Traditional use
	<p>Posology</p> <p><i>Adults and Elderly</i></p> <p><i>Combinations containing (among other active ingredients) Absinthii herba, Anisi fructus, Boldi folium, Carvi fructus, Centaurii herba, Cinnamomi cortex, Curcumae xanthorrhizae rhizoma, Fumariae herba, Gentianae radix, Hyperici herba, Liquiritiae radix, Melissa folium, Menyanthidis trifoliatae folium, Millefolii herba, Rosmarini herba, Salviae officinalis folium or Taraxaci radix cum herba:</i></p> <p>Herbal tea: 1.5-2 g of the herbal tea combination in 150 ml of boiling water as an herbal infusion 3 times daily.</p> <p><i>Adolescents, Adults and Elderly</i></p>

³ For guidance on herbal substance/herbal preparation administered as herbal tea or as infusion/decoction/macerate preparation, please refer to the HMPC 'Glossary on herbal teas' (EMA/HMPC/5829/2010 Rev.1).

Well-established use	Traditional use
	<p><i>Combinations containing Verbenae citriodoraefolium, Althaeae radix, Foeniculi amari fructus, Marrubii herba, Matricariae flos or Menthae piperitae herba:</i></p> <p>Herbal tea: 1.5-2 g of the herbal tea combination in 150 ml of boiling water as an herbal infusion 1-2 times daily.</p> <p><i>Depending on the herbal substances included:</i></p> <p>The use in children and adolescents under 18 years of age is not recommended (see section 4.4 'Special warnings and precautions for use').</p> <p><i>or</i></p> <p>The use in children under 12 years of age is not recommended (see section 4.4 'Special warnings and precautions for use').</p> <p>Duration of use</p> <p>If the symptoms persist longer than 2 weeks during the use of the medicinal product, a doctor or a qualified health care practitioner should be consulted.</p> <p><i>Combinations containing Foeniculi fructus and/or Anisi fructus:</i></p> <p>Not to be used for more than 2 weeks.</p> <p>Method of administration</p> <p>Oral use.</p>

4.3. Contraindications

Well-established use	Traditional use
	<p><i>All combinations:</i></p> <p>Hypersensitivity to the active substances.</p> <p><i>Combinations containing Absinthii herba, Matricariae flos, Millefolii herba or Taraxaci radix cum herba:</i></p> <p><i>Additionally:</i> Hypersensitivity to other plants of the Asteraceae family.</p> <p><i>Combinations containing Anisi fructus, Carvi fructus or Foeniculi fructus:</i></p> <p><i>Additionally:</i> Hypersensitivity to other plants of the Apiaceae family (fennel, anise, celery,</p>

Well-established use	Traditional use
	<p>coriander and dill).</p> <p><i>Combinations containing Verbenae citriodorae folium:</i></p> <p><i>Additionally:</i> Hypersensitivity to other plants of the Verbenaceae family.</p> <p><i>Combinations containing Marrubii herba:</i></p> <p><i>Additionally:</i> Hypersensitivity to other plants of the Lamiaceae family.</p> <p><i>Combinations containing Menyanthidis trifoliatae folium:</i></p> <p><i>Additionally:</i> Active gastric or duodenal ulcer.</p> <p><i>Combinations containing Absinthii herba, Boldi folium, Fumariae herba, Marrubii herba, Rosmarini herba or Taraxaci radix cum herba:</i></p> <p><i>Additionally:</i> Obstruction of the bile duct, cholangitis, liver disease, gallstones and any other biliary disorders.</p> <p><i>Combinations containing Centaurii herba; Gentianae radix or Taraxaci radix cum herba:</i></p> <p><i>Additionally:</i> Peptic ulcer.</p> <p><i>Combinations containing Anisi fructus or Foeniculi fructus:</i></p> <p><i>Additionally:</i> Hypersensitivity to anethole.</p> <p><i>Combinations containing Carvi fructus:</i></p> <p><i>Additionally:</i> Hypersensitivity to mugwort or to birch.</p> <p><i>Combinations containing Cinnamomi cortex:</i></p> <p><i>Additionally:</i> Hypersensitivity to Peru balsam.</p> <p><i>Combinations containing Menthae piperitae herba:</i></p> <p><i>Additionally:</i> Hypersensitivity to menthol.</p>

4.4. Special warnings and precautions for use

Well-established use	Traditional use
	<p><i>Depending on the herbal substances included:</i></p> <p>The use in children and adolescents under 18 years of age has not been established due to lack of adequate data.</p> <p><i>or</i></p>

Well-established use	Traditional use
	<p>The use in children under 12 years of age has not been established due to lack of adequate data.</p> <p><i>Combinations containing Absinthii herba, Carvi fructus, Curcumae xanthorrhizae rhizoma, Marrubii herba or Menthae piperitae herba:</i></p> <p>Gallstones, liver disease, cholangitis and any other biliary disorders.</p> <p><i>Combinations containing Althaeae radix:</i></p> <p>Not to be taken ½ to 1h before or after intake of other medicinal products.</p> <p><i>Combinations containing Liquiritiae radix:</i></p> <p>Hypertension, cardiac rhythm disorders, kidney diseases, liver or cardiovascular disorders or hypokalaemia. Concomitant use with diuretics, cardiac glycosides, corticosteroids, stimulant laxatives or other medications which may aggravate electrolyte imbalance is not recommended.</p> <p><i>Combinations containing Marrubii herba:</i></p> <p>Active peptic ulcer.</p> <p><i>Combinations containing Menthae piperitae herba:</i></p> <p>Patients with gastro-oesophageal reflux (heartburn).</p> <p><i>Combinations containing Taraxaci radix cum herba:</i></p> <p>Patients with renal failure and/or diabetes, and/or heart failure because of possible risks due to hyperkalaemia.</p> <p>If the symptoms worsen during the use of the medicinal product, a doctor or a qualified health care practitioner should be consulted.</p>

4.5. Interactions with other medicinal products and other forms of interaction

Well-established use	Traditional use
	<p><i>All combinations except for those containing Liquiritiae radix and/or Matricariae flos:</i></p>

Well-established use	Traditional use
	<p>None reported.</p> <p><i>Combinations containing Liquiritiae radix:</i></p> <p>Not to use concomitantly with diuretics, cardiac glycosides, corticosteroids, stimulant laxatives or other medications which may aggravate electrolyte imbalance.</p> <p><i>Combinations containing Matricariae flos:</i></p> <p>For patients after renal transplantation taking high doses for longer periods (about two months) interactions based on effects on CYP450 have been reported.</p>

4.6. Fertility, pregnancy and lactation

Well-established use	Traditional use
	<p><i>All combinations:</i></p> <p>Safety during pregnancy and lactation has not been established. In the absence of sufficient data, the use during pregnancy and lactation is not recommended.</p> <p>No fertility data available.</p>

4.7. Effects on ability to drive and use machines

Well-established use	Traditional use
	<p><i>All combinations except for those containing Melissa folium:</i></p> <p>No studies on the effect on the ability to drive and use machines have been performed.</p> <p><i>Combinations containing Melissa folium:</i></p> <p>May impair ability to drive and use machines. Affected patients should not drive or operate machinery.</p>

4.8. Undesirable effects

Well-established use	Traditional use
	<p><i>Combinations containing Absinthii herba, Althaeae radix, Carvi fructus, Centaurii herba, Cinnamomi cortex, Fumariae herba, Liquiritiae radix, Melissa</i></p>

Well-established use	Traditional use
	<p><i>folium, Marrubii herba or Salviae officinalis folium:</i></p> <p>None known.</p> <p>If adverse reactions occur, a doctor or a qualified health care practitioner should be consulted.</p> <p><i>Combinations containing Anisi fructus:</i></p> <p>Allergic reactions to aniseed affecting the skin or the respiratory system have been reported. The frequency is not known.</p> <p><i>Combinations containing Boldi folium:</i></p> <p>Hypersensitivity (anaphylaxis) has been reported.</p> <p>The frequency is not known</p> <p><i>Combinations containing Foeniculi fructus or Rosmarini herba:</i></p> <p>Allergic reactions affecting the skin, or the respiratory system may occur. The frequency is not known.</p> <p><i>Combinations containing Curcumae xanthorrhizae rhizoma:</i></p> <p>Mild gastrointestinal symptoms such as dry mouth, flatulence and gastric irritation may occur. The frequency is not known.</p> <p><i>Combinations containing Gentianae radix:</i></p> <p>Gastrointestinal disorders have been observed. The frequency was uncommon.</p> <p>In rare cases, tachycardia and pruritus have been reported.</p> <p>Headache may occur. The frequency is not known.</p> <p><i>Combinations containing Hyperici herba:</i></p> <p>Gastrointestinal disorders, allergic skin reactions, fatigue and restlessness may occur. The frequency is not known.</p> <p>Fair-skinned individuals may react with intensified sunburn-like symptoms under intense sunlight.</p> <p><i>Combinations containing Matricariae flos:</i></p> <p>Hypersensitivity reactions including severe allergic reaction (dyspnoea, Quincke's disease, vascular collapse, anaphylactic shock) following mucosal contact with liquid chamomile preparations have been reported. The frequency</p>

Well-established use	Traditional use
	<p>is not known.</p> <p><i>Combinations containing Menthae piperitae herba:</i></p> <p>The gastro-oesophageal reflux may worsen, and heartburn may increase. The frequency is not known.</p> <p><i>Combinations containing Millefolii herba:</i></p> <p>Hypersensitivity reactions of the skin have been reported. The frequency is not known.</p> <p><i>Combinations containing Taraxaci radix cum herba:</i></p> <p>Epigastric pain and hyperacidity may occur. The frequency is not known.</p> <p>Allergic reactions may occur. The frequency is not known.</p> <p>If other adverse reactions not mentioned above occur, a doctor or a qualified health care practitioner should be consulted.</p>

4.9. Overdose

Well-established use	Traditional use
	<p><i>All combinations except for those containing Centaurii herba and/or Liquiritiae radix:</i></p> <p>No case of overdose has been reported.</p> <p><i>Combinations containing Centaurii herba:</i></p> <p>Stomach disturbances and nausea have been reported.</p> <p><i>Combinations containing Liquiritiae radix:</i></p> <p>Water retention, hypokalaemia, hypertension, cardiac rhythm disorders, hypertensive encephalopathy.</p>

5. Pharmacological properties

5.1. Pharmacodynamic properties

Well-established use	Traditional use
	<p>Not required as per Article 16c(1)(a)(iii) of Directive 2001/83/EC.</p>

5.2. Pharmacokinetic properties

Well-established use	Traditional use
	Not required as per Article 16c(1)(a)(iii) of Directive 2001/83/EC.

5.3. Preclinical safety data

Well-established use	Traditional use
	<p>Not required as per Article 16c(1)(a)(iii) of Directive 2001/83/EC, unless necessary for the safe use of the product.</p> <p><i>Combinations containing Absinthii herba:</i></p> <p>Thujone is reported to be neurotoxic and chemotypes with low content of thujone should be preferred.</p> <p>Tests on reproductive toxicity have been performed with a dry ethanolic extract of Absinthii herba administered orally to pregnant rats. Results showed reduced sites of implantations and a reduced rate of born pups. Thujone is known for its uterus stimulating activity.</p> <p><i>Combinations containing Anisi fructus:</i></p> <p>Results from non-clinical studies showed a weak mutagenic potential of anethole. Anethole and its metabolites are unlikely to be genotoxic <i>in vivo</i>.</p> <p>The genotoxic risk related to estragole is not considered to be relevant in the specified conditions of use (short term use in adolescents, adults and elderly) due to the small amount present in herbal infusions prepared from aniseed.</p> <p><i>Combinations containing Boldi folium:</i></p> <p>Tests on genotoxicity and carcinogenicity have not been performed.</p> <p><i>Combinations containing Foeniculi fructus:</i></p> <p>A fennel aqueous extract was tested in an Ames test on <i>Salmonella typhimurium</i> strains TA98, TA100 and turned out as negative. Results from studies carried out in laboratory animals showed a weak mutagenic activity of anethole.</p>

Well-established use	Traditional use
	<p>The genotoxic risk related to estragole is not considered to be relevant in the specified conditions of use due to the small amount present in herbal infusions prepared from fennel.</p> <p><i>Combinations containing Gentianae radix:</i></p> <p>For some xanthenes which are among the constituents of <i>Gentiana lutea</i>, positive results were found in the AMES test (pre-incubation method).</p> <p>Assessment of preclinical safety requires further studies towards these effects.</p> <p><i>Combinations containing Hyperici herba:</i></p> <p>Studies on acute toxicity and repeated dose toxicity did not show signs of toxic effects.</p> <p>Tests on reproductive toxicity revealed equivocal results.</p> <p>Tests on genotoxicity and carcinogenicity have not been performed.</p>

6. Pharmaceutical particulars

Well-established use	Traditional use
	<p><i>Combinations containing Absinthii herba or Salviae officinalis folium:</i></p> <p>The amount of thujone has to be specified in the given product. The daily exposure has to be below 6.0 mg.</p> <p><i>Combinations containing Boldi folium:</i></p> <p>The levels of ascaridole in herbal preparations and herbal medicinal products should be determined.</p> <p><i>Combinations containing Hyperici herba:</i></p> <p>The amount of hyperforin has to be specified in herbal preparation for oral use. The daily intake of hyperforin has to be below 1 mg.</p>

7. Date of compilation/last revision

07 July 2021