94245 – COLOURLESS CAPSULES 2

1. IDENTIFICATION OF THE SUBSTANCE OR PREPARATION.

1.1 Identification of the substance or preparation.
Name: Colourless capsules 2
Code: 94245
Internal code: 55200
Valid from batch: 0116447

1.2 Synonyms.
No information available.

2. DESCRIPTION

Appearance: Two pieces of hard gelatine. The gelatine used meets all the requirements of the European Pharmacopoeia and USP.
The sections of the hard gelatine capsule are freely separated before filling, and the separated edges of the lid and the body are cleanly and uniformly cut without visible roughness.

Size: 2
Cap/body colour: Colourless/Colourless.
Cap external diameter (mm): 6.37±0.03
Body external diameter (mm): 6.08±0.03
Odour: Odourless.
Flavour: Insipid.
Origin: Croatia.
Method of production: The capsules are manufactured with pharmaceutical-grade gelatine.

Approximate capsule filling content in mg

Density / Size	0.6 g/ml	0.8 g/ml	1.0 g/ml	1.2 g/ml
2	222	296	370	444

3. COMPOSITION/INFORMATION ON COMPONENTS.

Capsule composition:
Water: 14-15%
100% csp gelatine

Regulatory information under food law on the colourant used, where applicable:

Products to be consumed in the European Union use colours that comply with (EU) 231/2012, which sets forth specifications for food additives.
Where applicable, as in products to be consumed in the US, the colourants are FDA certified in accordance with 21CFR Parts 73 and 74 for food and drugs.
The amount of colour additive used in the above product is in accordance with the ADI as per the WHO/FAO as applicable.
When used, globally accepted colourants, such as iron oxides, comply with the acceptance criteria of regulated markets such as the EU and US.
The elemental iron content does not exceed 5 mg per day according to 21CFR Part 73.120 for food and Part 73.1200 for medicine.

4. PHYSICO-CHEMICAL DATA.

For more information, see the analysis report.

Hard gelatine capsules disintegrate in less than 15 minutes.

5. PROPERTIES/USES.

The capsules are FIT for human consumption.
The hard gelatine capsules are distributed worldwide and are designed for use in pharmaceuticals and foodstuffs.
Hard gelatine capsules are soluble in hot water at 98.6°F ± 1.8°F (37°C ± 1°C).
The moisture content of the hard gelatine capsules is between 13% and 16%.

6. DOSAGE.

No information available.

7. REMARKS.

SPECIFICATION:

The capsules comply with all the requirements set forth in the EP and USP.

STORAGE:

The capsules are manufactured with a carefully controlled moisture content, and this should be taken into account to obtain the best filling performance; if the conditions are not respected, maintenance problems will appear. The capsules will be too dry and brittle and

will contract due to loss of moisture, or they will become sticky and soft due to increased moisture.

The capsules preserve their characteristics as long as they are stored at temperatures between 15°C and 25°C and at a relative humidity of 35% to 65%, and provided that the following precautions are taken:

- Capsules should not be exposed directly to sunlight or stored near heat sources or moisture. They should be stored in a dry, cool place and they should be allowed to balance the conditions in the filling area before use.
- The capsules should be kept in their original packaging. The containers should be tightly closed during transport and storage to ensure their quality over time. If any number of capsules is extracted for examination or filling, it is essential to properly close the container.
- It is recommended that during capsule filling, only the required number of capsules be available to avoid exposing them to the heat that the filling machines generate. Filling areas should be between 20°C and 25°C and at a relative humidity between 45% and 55% to keep the moisture content of the capsules in the desired range of 13% to 16%.

The documentation available related to the product's regulatory compliance is included below.

BSE/TSE:

The gelatine used is a mixture of pharmaceutical-grade gelatine based on product quality requirements and regulations. All gelatine used is compliant with the pharmacopoeia and applicable food standards. Bovine gelatine, when used, is derived from the bones and skins of healthy animals that have been certified as fit for human consumption by a veterinary officer at the slaughterhouse. The tissues used to manufacture the gelatine are free of specified risk materials.
In addition, the bovine gelatine is compliant with the following BSE regulations applicable in the country of use, such as:
• Regulation (EC) No 999/2001 and its amendments.
• Regulation (EC) No 853/2004.
• USFDA, September 1997, Guidance for Industry on the Sourcing and Processing of Gelatine to Reduce Potential Risk Posed by BSE in FDA-Regulated Products.
• USFDA 21 CFR Parts 189, 589, 700 relating to livestock materials prohibited in human food, medicines and cosmetics and animal feed.
• USFDA 9 CFR Part 94.23 on import of bovine gelatine.
• Commission Directive 2003/63/EC and Ph. Eur – 1483, accredited by the certificate of suitability.

• Note for guidance on minimising the risk of transmitting animal spongiform encephalopathy agents via human and veterinary medicinal products (EMA/410/01 Rev. 03).
• In addition, no ingredients of animal origin are used in the manufacture of the above product.

Therefore, hard gelatine capsule shells do not pose any risk of BSE.

GMOs:
Hard gelatine capsule shells are not subjected to or derived from any genetic modification technique as defined in Article 2(2) of European Directive 2001/187/EC.
The ingredients used to manufacture the above product are not subjected to or derived from any genetic modification technique as defined above.
The ingredients used to manufacture the above product can be traced back to their origin.
The product does not contain GMOs (genetically modified organisms).
Accordingly, the hard gelatine capsule shells are compliant with EC 298/2008 (formerly 189/2003) and EC 1830/2003 on genetically modified food and feed.

NANOMATERIALS:
The product is not considered a nanomaterial as defined in EC Directive 1169/2011.
Furthermore, the ingredients used in the above product are not manufactured as nanomaterials nor is nanotechnology used.

RESIDUAL SOLVENTS:
The hard gelatine capsule shells do not contain any Class I and II solvents as per CPMP/ICH/283/95 and the current edition of the USP General Chapter <467> on Residual Solvents.
The following Class III solvents (glacial acetic acid, isopropyl alcohol, ethyl alcohol and n-butyl alcohol) are likely to be present in our product in amounts below 5,000 ppm. No other solvents are used in the manufacturing process.

ELEMENTAL IMPURITIES:
The product is compliant with:

- ICHQ3D Guideline for elemental impurities, final version dated 26 April 2022.
- EMA/CHMP/ICH/353369/2013 ICH Q3D Guideline for elemental impurities dated 22 March 2019.
- USP <232> Elemental Impurities – Limits.

Compliance is based on three components of the ICH Q3D: assessment of potential elemental impurities, comparison of potential levels with permitted daily exposure (PDE) for each element of toxicological concern and the development of controls designed to limit the concentration of elemental impurities in medicinal products to below the PDE.

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CONTAMINANTS:

Melamine: The capsules do not contain melamine.
Gluten: The capsules are gluten free.
Aflatoxins: The capsules do not contain aflatoxins.
Pesticides: The capsules do not contain residual pesticides and are safe for human consumption.
Dioxins: The capsules do not contain dioxins.
PAHs and PCBs: The capsules do not contain PAHs or PCBs.

ALLERGENS:

Allergen free according to VO (EU) No. 1169/2011 and the US FDA Guidance for Industry (food allergen labelling and consumer protection).
The possible presence of SO_2 comes from the gelatine used and is found in amounts lower than 50 ppm.

PRESERVATIVES:

The product does not contain preservatives.

MANUFACTURER CERTIFICATIONS:

cGMP, ISO 9001:2015, 14001:2015, Kosher, Halal