ANNEX I

SUMMARY OF PRODUCT CHARACTERISTICS

SUMMARY OF PRODUCT CHARACTERISTICS

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

AQUACEN OXITETRACICLINA HIDROCLORURO 1000 mg/g PREMEZCLA MEDICAMENTOSA (ES)
AQUACEN OXYTETRACYCLINE HYDROCHLORIDE 1000 mg/g MEDICATED PREMIX (EL, PT)
OSSITETRACICLINA CENAVISA 1000 mg/g premix for medicated feeding stuff (IT)

2. QUALITATIVE AND QUANTITATIVE COMPOSITION

Each g contains:

Active substance:
Oxytetracycline hydrochloride .....1000 mg

Excipients:
For a full list of excipients, see section 6.1.

3. PHARMACEUTICAL FORM

Premix for medicated feeding stuff
Yellow crystalline powder

4. CLINICAL PARTICULARS

4.1 Target species

Salmonids (*Salmo* sp, *Oncorhynchus* sp), seabass (*Dicentrarchus labrax*), sea bream (*Sparus aurata*), turbot (*Psetta maxima*), european eel (*Anguilla anguilla*), european carp (*Cyprinus carpio*)

4.2 Indications for use, specifying the target species

For the treatment of infections due to *Lactococcus garvieae* (lactococcosis), *Aeromonas hydrophila* (aeromoniosis) *and Vibrio anguillarum* (vibriosis) sensitive to oxytetracycline.

4.3 Contraindications

Do not use in case of known allergy to oxytetracycline or other substance belonging to the tetracycline group.
Do not use in cases of hypersensitivity to oxytetracycline or other substance belonging to the tetracycline groups.
Do not use in case of known resistance to tetracyclines.

4.4 Special warnings for each target species

None.

4.5 Special precautions for use

Special precautions for use in animals
Use of the product should be based on susceptibility testing of the bacteria isolated from the animal. If this is not possible, therapy should be based on local (regional, farm level) epidemiological information about susceptibility of the target bacteria.

Special precautions to be taken by the person administering the veterinary medicinal product to animals
- People with known hypersensitivity to oxytetracycline or other substance belonging to the tetracycline group should avoid any contact with the veterinary medicinal product.
- Avoid contact with skin and eyes. In case of contact, wash immediately with plenty of water.
- Avoid inhalation of veterinary medicine.
- Use personal protective equipment consisting of safety glasses, gloves and mask filtering (standard CEN FFP1).
- Wash hands after use of the veterinary medicinal product.
- If symptoms appear following exposure such as a skin rash, consult a doctor showing these warnings. The swelling of the face, lips or eyes or difficulty on breathing are more serious signs that need urgent medical attention.

4.6 Adverse reactions (frequency and seriousness)

None known.

4.7 Use during pregnancy, lactation or lay

Not applicable.

4.8 Interaction with other medicinal products and other forms of interaction

Absorption of oxytetracycline can be reduced in the presence of high amounts of Ca^{2+}, Fe^{2+}, Mg^{2+} or Al^{3+} in the diet.
It is not recommended to mix this product with feeding stuff containing other antibiotics.

4.9 Amounts to be administered and administration route

Administration in the food.

The recommended daily dose rate is 55 mg of oxytetracycline per kg of fish bodyweight, for 7-10 days, orally.

The premix should be incorporated in pelleted feed. Incorporation rates vary depending on the water temperature and the size of the fish. Corresponding to a daily intake of 1.5% body weight, the dosage of oxytetracycline hydrochloride in the feed would be about 4 kg of premix per ton of feed.

Due to the mode of administration, since feed intake depends on the clinical condition of the animal and its physiological state (age) to ensure proper dosing, the

concentration of the antimicrobial will be adjusted taking into account the daily feed intake.

Instructions for proper administration:
Medicated premix is added to pelleted feed and then fish or vegetable oil is added for adhere well the premix on the surface of the pellets.

4.10 Overdose (symptoms, emergency procedures, antidotes), if necessary

Do not exceed the stated dose.

4.11 Withdrawal period

Meat: 300 degree days.

5. PHARMACOLOGICAL PROPERTIES

Pharmacotherapeutic group: Antibacterials for systemic use.
ATC Vet Code: QJ01AA06

5.1 Pharmacodynamic properties

Oxytetracycline is a bacteriostatic antibacterial that acts by interfering bacterial protein synthesis of susceptible species. Acts on the 30S ribosomal subunit of the bacterial ribosome, which binds reversibly, preventing the binding of aminoacyl-tRNA (transfer RNA) to the complex formed by the ribosome and mRNA, thereby preventing protein synthesis.

This medicinal product is indicated in the treatment of infections due to *Lactococcus garvieae* (Gram-positive bacteria), *Aeromonas hydrophila and Vibrio anguillarum* (Gram-negative bacteria).

There are at least two mechanisms of resistance to tetracyclines. The most important mechanism is caused by a reduction of the intracellular accumulation of the drug. It is due to the establishment of an elimination route by pumping the antibacterial agent or an alteration in the transport system, resulting in a limited tetracycline energy-dependent capture to the exterior of the cell. The alteration in the transport system is produced by inducible proteins coded by plasmids and transposons. The other mechanism is evidenced by a decrease in affinity of the ribosome by tetracycline-Mg^{2+} complex due to mutation in the chromosome.

5.2 Pharmacokinetic particulars

In most species, oxytetracycline is rapidly absorbed (2-4 hours) after the oral administration in the fasted state, with a bioavailability ranging from 60% and 80%. The degree of absorption may be decreased by the presence, in the food, of soluble salts of divalent and trivalent metals, Ca^{2+}, Fe^{2+}, Mg^{2+} or Al^{3+}, with which tetracyclines form stable compounds.

In fish (rainbow trout and Chinook salmon) in water at 11°C, the bioavailability of oxytetracycline is 25-30% and the elimination half-life is 88 to 94 hours.

6. PHARMACEUTICAL PARTICULARS

6.1 List of excipients

Not applicable.

6.2 Major incompatibilities

In the absence of compatibility studies, this veterinary medicinal product must not be mixed with other veterinary medicinal products.

6.3 Shelf life

Shelf-life of the veterinary medicinal product as packaged for sale: 2 years.
Shelf-life after first opening the immediate packaging: 3 months.
Shelf life after incorporation into pelleted feed: 3 months.

6.4. Special precautions for storage

Do not store above 30°C

6.5 Nature and composition of immediate packaging

Multi-layer heat-sealed bags containing 20 kg and 1 kg.
The outermost layer of the 20 kg bag is polyester, and the innermost layer is linear polyethylene coextrusion of low-medium density.
The outermost layer of the 1 kg bag is oriented polypropylene, and the innermost layer is low-density polyethylene.

Package sizes:
Bag containing 1 kg
Bag containing 20 kg.

Not all pack sizes may be marketed.

6.6 Special precautions for the disposal of unused veterinary medicinal product or waste materials derived from the use of such products

Any unused veterinary medicinal product or waste materials derived from such veterinary medicinal product should be disposed of in accordance with local requirements.

7. MARKETING AUTHORISATION HOLDER

CENAVISA, S.L.
Camí Pedra Estela s/n
43205 Reus (SPAIN)

8. MARKETING AUTHORISATION NUMBER(S)

9. DATE OF FIRST AUTHORISATION/RENEWAL OF THE AUTHORISATION

10 DATE OF REVISION OF THE TEXT

PROHIBITION OF SALE, SUPPLY AND/OR USE

Official guidance on the incorporation of medicated premixes in feeds should be given into account.

Veterinary medicinal product subject to prescription.
To be administered by a veterinary surgeon or under their direct responsibility.