

## **SUMMARY OF PRODUCT CHARACTERISTICS**

### **1. NAME OF THE VETERINARY MEDICINAL PRODUCT**

Moxodex LA 100 mg/ml Solution for Injection for Cattle.

### **2. QUALITATIVE AND QUANTITATIVE COMPOSITION**

Each ml contains:

#### **Active substance:**

Moxidectin                              100 mg

#### **Excipients:**

<b>Qualitative composition of excipients and other constituents</b>	<b>Quantitative composition if that information is essential for proper administration of the veterinary medicinal product</b>
Benzyl Alcohol (E1519)	70 mg
Butylhydroxytoluene (BHT)	0.6 mg
Sorbitan Oleate	-
Propylene Glycol Dicaprylocaprate	-

Clear, yellow liquid, free of visible particles.

### **3. CLINICAL INFORMATION**

#### **3.1 Target species**

Cattle.

#### **3.2 Indications for use for each target species**

In cattle weighing from 100 to 500 kg body weight, treatment and prevention of mixed infestations by the following gastro-intestinal nematodes, respiratory nematodes and certain arthropod parasites:

Adult and immature gastro-intestinal nematodes:

- . *Haemonchus placei*
- . *Haemonchus contortus*
- . *Ostertagia ostertagi* (including inhibited larvae)
- . *Trichostrongylus axei*

- . *Trichostrongylus colubriformis*
- . *Nematodirus helvetianus* (adults only)
- . *Nematodirus spathiger*
- . *Cooperia surnabada*
- . *Cooperia oncophora*
- . *Cooperia pectinata*
- . *Cooperia punctata*
- . *Oesophagostomum radiatum*
- . *Bunostomum phlebotomum* (adults only)
- . *Chabertia ovina* (adults only)
- . *Trichuris spp.* (adults only)

Adult and immature respiratory tract nematode

- . *Dictyocaulus viviparus*

Warble grubs (migrating larvae)

- . *Hypoderma bovis*
- . *Hypoderma lineatum*

Lice

- . *Linognathus vituli*
- . *Haematopinus eurysternus*
- . *Solenopotes capillatus*
- . *Bovicola bovis* (aid in control)

Mange mites

- . *Sarcoptes scabiei*
- . *Psoroptes ovis*
- . *Chorioptes bovis* (aid in control)

The veterinary medicinal product has a persistent action and protects cattle for a certain duration against infection or re-infection with the following parasites for the period indicated:

<b>Species</b>	<b>Protection period (days)</b>
<i>Dictyocaulus viviparus</i>	120
<i>Ostertagia ostertagi</i>	120
<i>Haemonchus placei</i>	90
<i>Oesophagostomum radiatum</i>	150
<i>Trichostrongylus axei</i>	90
<i>Linognathus vituli</i>	133

The veterinary medicinal product is effective against *Hypoderma* larvae at the time of treatment but its persistent activity against *Hypoderma* has not been evaluated. If the product is given before the end of the fly season complimentary treatment with a product effective against *Hypoderma* may be required.

Persistent efficacy periods have not been established for parasite species other than those included in the above list. Therefore, re-infection of animals on pasture

contaminated by parasites other than these remains possible before the end of the 90 day minimum persistency period demonstrated for specific species.

### **3.3 Contraindications**

Do not use in animals less than 100 kg bodyweight or greater than 500 kg.

Do not inject the product by intravascular route. Intravascular injection may result in ataxia, paralysis, convulsions, collapse and death. To prevent any intravascular injection, carefully follow the administration procedure described in item "Amounts to be administered and administration route."

Do not use in cases of hypersensitivity to the active substance or any of the excipients.

### **3.4 Special warnings**

Unnecessary use of antiparasitics or use deviating from the instructions given in the SPC may increase the resistance selection pressure and lead to reduced efficacy. The decision to use the product should be based on confirmation of the parasitic species and burden, or of the risk of infestation based on its epidemiological features, for each individual animal and/or herd.

Repeated use for an extended period, particularly when using the same class of substances, increases the risk of resistance development. Within a herd, maintenance of susceptible refugia is essential to reduce that risk. Systematically applied interval-based treatment and treatment of a whole herd should be avoided. Instead, if feasible, only selected individual animals or subgroups should be treated (targeted selective treatment). This should be combined with appropriate husbandry and pasture management measures. Guidance for each specific herd should be sought from the responsible veterinarian.

Underdosing may result in ineffective use and may favour resistance development; therefore, underestimation of bodyweight and misadministration of the product must be avoided (see section 3.9. Administration routes and dosage).

The use of this product should take into account local information about susceptibility of the target parasites, where available.

It is recommended to further investigate cases of suspected resistance, using an appropriate diagnostic method (e.g. Faecal Egg Count Reduction Test). Confirmed resistance should be reported to the marketing authorisation holder or to the competent authority.

Where the results of tests strongly suggest resistance to a particular anthelmintic, an anthelmintic belonging to another pharmacological class and having a different mode of action should be used.

In the absence of risk of co-infection, a narrow spectrum product should be used.

### 3.5 Special precautions for use

#### Special precautions for safe use in the target species:

In order to prevent abscesses, a strict aseptic technique is recommended. The veterinary medicinal product has been formulated specifically for subcutaneous injection in dorsal surface of the ear of cattle and must not be given by any other route of administration or to any other species.

To avoid possible secondary reactions by the death of *Hypoderma* larvae in the spine or the oesophagus of animals, it is recommended to administer a veterinary medicinal product effective against *Hypoderma* larvae after the end of fly activity and before the larvae reach their resting sites. Consult your veterinary surgeon on the correct timing of this treatment.

Immunity to nematodes depends on adequate exposure to infection. Although not normally the case, circumstances could occur in which anthelmintic control measures might increase the vulnerability of cattle to re-infection. Animals may be at risk towards the end of their first grazing season, particularly if the season is long, or in the following year if they move onto heavily contaminated pasture. In such instances, further control measures may be necessary.

#### Special precautions to be taken by the person administering the veterinary medicinal product to animals:

This product can cause skin and eye irritation.

Avoid direct contact with skin and eyes.

Do not smoke, drink, or eat while handling the product.

Wash hands after use.

If accidental skin contact occurs, wash the affected area with soap and water.

If accidental eye exposure occurs, immediately rinse the eyes thoroughly with water.

If skin or eye irritation persists, seek medical attention.

Take care to avoid self-injection.

In case of accidental self-injection, seek medical advice immediately and show the package leaflet or the label to the physician.

Advice to medical practitioners: In case of accidental self-injection: Treat symptomatically.

#### Special precautions for the protection of the environment:

Moxidectin fulfils the criteria for a (very) persistent, bioaccumulative and toxic (PBT) substance; therefore, exposure of the environment to moxidectin must be limited to the extent possible. Treatments should be administered only when necessary and should be based on faecal egg counts or evaluation of the risk of infestation at the animal and/or herd level.

Like other macrocyclic lactones, moxidectin has the potential to adversely affect non-target organisms:

Faeces containing moxidectin excreted onto pasture by treated animals may temporarily reduce the abundance of dung feeding organisms. Following treatment of

cattle with the veterinary medicinal product, levels of moxidectin that are potentially toxic to dung fly species may be excreted over a period more than 4 weeks and may decrease dung fly abundance during that period. It has been established in laboratory tests that moxidectin may temporarily affect dung beetle reproduction; however, field studies indicate no long-term effects. Nevertheless, in case of repeated treatments with moxidectin (as with veterinary medicinal products of the same anthelmintic class) it is advisable not to treat animals every time on the same pasture to allow dung fauna populations to recover.

Moxidectin is inherently toxic to aquatic organisms including fish. The veterinary medicinal product should be used only according to the label instructions. Based on the excretion profile of moxidectin when administered as the injectable formulation, treated animals should not have access to watercourses during the 10 days after treatment.

### 3.6 Adverse events

Cattle:

Rare (1 to 10 animals / 10,000 animals treated):	Injection site swelling <sup>1,2</sup> Depression and ataxia
Very rare (<1 animal / 10,000 animals treated, including isolated reports):	Injection site abscess <sup>2</sup>

<sup>1</sup>immediate or delayed, may further develop into injection site abscesses, frequency tends to be higher in heavier animals.

<sup>2</sup>generally disappear without treatment within 14 days after administration, although may persist up to 5 weeks (<5% of cases) or longer (very rare occasions). In case of hypersensitivity reactions, a symptomatic treatment should be applied.

Reporting adverse events is important. It allows continuous safety monitoring of a veterinary medicinal product. Reports should be sent, preferably via a veterinarian, to either the marketing authorisation holder or the national competent authority via the national reporting system. See the package leaflet for respective contact details.

### 3.7 Use during pregnancy, lactation or lay

Pregnancy and lactation:

Can be used during pregnancy. Note section 3.3. Contraindications and section 3.12 Withdrawal periods.

### 3.8 Interaction with other medicinal products and other forms of interaction

The effects of GABA agonists are increased by moxidectin.

### 3.9 Administration routes and dosage

For subcutaneous use.

Dosage is 0.5 ml/50 kg bodyweight, equivalent to 1.0 mg moxidectin/kg bodyweight, given by a single subcutaneous injection in the ear using an 18 gauge, 25 – 40 mm hypodermic needle. The 50 ml vial stoppers must not be broached more than 20 times. Use automatic syringe equipment for the 250 ml vial, no more than five piercings should be made.

Shake well before use.

Underdosing could result in ineffective use and may favour resistance development. To ensure that the correct dosage is used, an animal's bodyweight should be determined as accurately as possible. If animals are to be treated collectively, reasonably homogeneous groups should be set up, and all animals in a group should be dosed at a rate corresponding to the heaviest one.

Accuracy of the dosing device should be thoroughly checked.

The injection should be given subcutaneously in the loose tissues on the dorsal surface of the ear, just distal to the distal edge of the auricular cartilage.

The dorsal (outer) surface of the ear should first be cleansed with antiseptic and allowed to briefly air dry. Palpate the edge of the auricular cartilage closest to the head, on the dorsal (hairy) surface of the ear. From this landmark, taking care to avoid blood vessels (artery, vein), the needle should be inserted subcutaneously starting at a point approximately 3 to 3.5 cm distal to this edge (away from the head), and directed towards the base of the ear, and the needle advanced to the hub. At this point, gently aspirate the syringe to confirm that the needle is not in a blood vessel.

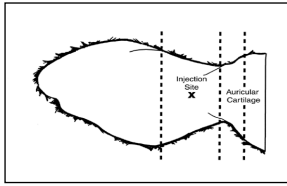
Upon injection, the resulting depot should reside just distal to the edge of the auricular cartilage.

Following administration, the needle is withdrawn from the skin as pressure is applied for several seconds with the thumb at the point of insertion.

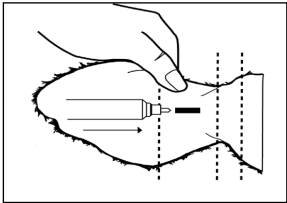
Due to the long-lasting protection against *Dictyocaulus viviparus* and the stomach worms, *Ostertagia ostertagi* and *Haemonchus placei*, a single treatment with the formulation at turn-out helps control parasitic bronchitis (lungworm) and parasitic gastro-enteritis throughout the grazing season by reducing the build-up of infective larvae on pasture associated with these parasites.

For best results the injection should be given to each calf of target weight to be grazed together immediately prior to being turned out to pasture. Animals should be set stocked throughout the grazing season or moved to a pasture which has not been grazed by other cattle earlier in the season.

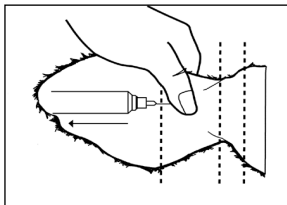
**Diagram: Ear injection procedure.**



- The injection site is approximately 3.5 cm (1.5 inches) distal to the distal edge of the auricular cartilage.



- Use one hand to grasp and steady the ear.
- Inject subcutaneously using an 18 gauge x 1 inch needle.



- Inject contents. Depot should be just distal to the distal edge of the auricular cartilage.
- Apply pressure at the point of insertion as the needle is withdrawn from the skin to help seal the opening.

**3.10 Symptoms of overdose (and where applicable, emergency procedures and antidotes)**

Reactions at the injection site have to be expected more frequently and severe depending on the injected volume. Systemic clinical signs of overdoses are consistent with the mode of action of moxidectin. These clinical signs are manifested as transient salivation, depression, drowsiness and ataxia 24 to 36 hours post-treatment. The systemic clinical signs usually disappear within 36 to 72 hours without treatment. At doses >3 times the recommended dose divided on both ears, the systemic clinical signs included recumbency, muscle tremor, ruminal tympany and dehydration, which were resolved after treatment with fluids. The systemic clinical signs can last for a few days to ten days. There is no specific antidote.

**3.11 Special restrictions for use and special conditions for use, including restrictions on the use of antimicrobial and antiparasitic veterinary medicinal products in order to limit the risk of development of resistance**

Not applicable.

### **3.12 Withdrawal periods**

Meat and offal: 108 days.

Milk: Not permitted for use in lactating animals producing milk for human consumption or industrial purposes or within 80 days of expected parturition.

The withdrawal period is based solely on a single injection at the ear site of injection.

## **4. PHARMACOLOGICAL INFORMATION**

### **4.1 ATCvet code:**

QP54AB02

### **4.2 Pharmacodynamics**

Moxidectin is an endectocide active against a wide range of internal and external parasites and is a second generation macrocyclic lactone of the milbemycin family.

Moxidectin interacts with GABA receptors and glutamate gated chloride channels. The net effect is to open the chloride channels on the postsynaptic junction to allow the inflow of chloride ions and induce an irreversible resting state. This results in flaccid paralysis and eventual death of parasites exposed to the active substance.

The exact mechanisms of parasite resistance to moxidectin have not been elucidated. A resistance mechanism involving metabolism by p-glycoproteins and efflux from the cells by ABC transporters has been proposed for ivermectin and a similar mechanism is thought to play a role in moxidectin resistance. However, parasites resistant to ivermectin are known to show some degree but not complete cross-resistance to moxidectin. It has been proposed that the reason for the incomplete cross-resistance is that there are multiple avenues of moxidectin action in target parasites that may include receptors other than the Glutamate-gated chloride channels.

### **4.3 Pharmacokinetics**

Moxidectin is absorbed following subcutaneous injection with maximum blood concentrations being achieved 24 to 48 hours post injection. The active substance is distributed throughout the body tissues but due to its lipophilicity it is concentrated mainly in the fat. The depletion half-life in fat is 26 – 32 days.

Moxidectin undergoes limited biotransformation by hydroxylation in the body. The only significant route of excretion is the faeces.

### **Environmental properties**

Moxidectin fulfils the criteria for a (very) persistent, bioaccumulative and toxic (PBT) substance. In particular, in acute and chronic toxicity studies with algae, crustaceans and fish, moxidectin showed toxicity to these organisms, yielding the following endpoints:



Organism		EC <sub>50</sub>	NOEC
Algae	<i>S. capricornutum</i>	>86.9 µg/l	86.9 µg/l
Crustaceans (Water fleas)	<i>Daphnia magna</i> (acute)	0.0302 µg/l	0.011 µg/l
	<i>Daphnia magna</i> (reproduction)	0.0031 µg/l	0.010 µg/l
Fish	<i>O. mykiss</i>	0.160 µg/l	Not determined
	<i>L. macrochirus</i>	0.620 µg/l	0.52 µg/l
	<i>P. promelas</i> (early life)	Not	0.0032 µg/l
	<i>Cyprinus carpio</i>	0.11 µg/l	Not determined

EC<sub>50</sub>: the concentration which results in 50% of the test species individuals being adversely affected, i.e. both mortality and sub-lethal effects.

NOEC: the concentration in the study at which no effects are observed.

This implies that when allowing moxidectin to enter water bodies, this may have a severe and lasting impact on aquatic life. To mitigate this risk, all precautions for use and disposal must be adhered to.

## 5. PHARMACEUTICAL PARTICULARS

### 5.1 Major incompatibilities

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

### 5.2 Shelf life

Shelf life of the veterinary medicinal product as packaged for sale: 2 years.

Shelf life after first opening the immediate packaging: 28 days.

### 5.3 Special precautions for storage

Do not store above 25°C. Keep the container in the outer carton in order to protect from light.

### 5.4 Nature and composition of immediate packaging

Nature of the primary container: HDPE vial with a chlorinated butyl rubber stopper and an Aluminium flip off seal (50 ml vial)/Aluminium seal (250 ml).

Pack sizes:

Box containing 1 vial of 50 mL size.

Box containing 1 vial of 250 mL size.

Not all pack sizes may be marketed.

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

Medicines should not be disposed of via wastewater.

The veterinary medicinal product should not enter water courses as moxidectin may be dangerous for fish and other aquatic organisms.

Use take-back schemes for the disposal of any unused veterinary medicinal product or waste materials derived thereof in accordance with local requirements and with any national collection systems applicable to the veterinary medicinal product concerned.

## **6. NAME OF THE MARKETING AUTHORISATION HOLDER**

EU Pharmaceuticals Ltd

## **7. MARKETING AUTHORISATION NUMBERS**

GB: Vm 39787/5006

NI: Vm 39787/3006

## **8. DATE OF FIRST AUTHORISATION**

11 September 2024

## **9. DATE OF THE LAST REVISION OF THE SUMMARY OF THE PRODUCT CHARACTERISTICS**

September 2024

## **10. CLASSIFICATION OF VETERINARY MEDICINAL PRODUCT**

Veterinary medicinal product subject to prescription.

Find more product information by searching for the 'Product Information Database' on [www.gov.uk](http://www.gov.uk).

*Gavin Hall*

Approved: 11 September 2024