

**ANNEX I**

**SUMMARY OF PRODUCT CHARACTERISTICS**

## 1 NAME OF THE VETERINARY MEDICINAL PRODUCT

CRYOMAREX RISPENS

Concentrate and solvent for suspension for injection.

## 2 QUALITATIVE AND QUANTITATIVE COMPOSITION

Each 0,2 ml dose of diluted vaccine contains:

### Active substance:

Attenuated live Marek's disease virus, serotype 1, Rispens strain .....  $10^3 - 10^4$  PFU\*

PFU\* Plaque Forming Unit

### Excipient:

Phenol red .....2  $\mu$ l

For the full list of excipients, see section 6.1.

## 3 PHARMACEUTICAL FORM

Concentrate and solvent for suspension for injection.

Frozen concentrate: homogeneous yellow to red-orange suspension

Solvent: red-orange limpid solution.

## 4 CLINICAL PARTICULARS

### 4.1 Target Species

Chickens (Broiler and replacement chick i.e. chicken for reproduction or layer hen).

### 4.2 Indications for Use, specifying the target species

For active immunization of broilers and replacement chicks i.e. chicken for reproduction or layer hen, to reduce mortality, clinical signs and lesions of Marek's disease.

Onset of immunity: 9 days following vaccination.

Duration of immunity: a single vaccination is sufficient to provide protection during the risk period.

### 4.3 Contra-indications

None.

### 4.4 Special Warnings for each Target Species

Vaccinate healthy animals only.

#### **4.5 Special Precautions for use**

##### Special Precautions for use in animals

Vaccine strain spreads from vaccinated to unvaccinated birds.

##### Special Precautions to be taken by the Person Administering the veterinary medicinal product to animals

The thawing of the ampoules removed from liquid nitrogen may cause an ampoule to break. Personal protective equipment consisting of protecting gloves and goggles should be worn when handling the veterinary medicinal product

Warnings to be taken are indicated in the section “**Amounts to be administered and administration route**”.

#### **4.6 Adverse reactions (frequency and seriousness)**

None known.

#### **4.7 Use during pregnancy, lactation or lay**

Do not use in birds in lay.

#### **4.8 Interaction with other medicinal products and other forms of Interaction**

Safety and efficacy data are available which demonstrate that this vaccine can be mixed with Vaxxitek HVT+IBD, a live HVT vector vaccine expressing viral protein 2 (VP2) of Infectious Bursal Disease (IBD).

No information is available on the safety and efficacy of this vaccine when used with any other veterinary medicinal product except the products mentioned above. A decision to use this vaccine before or after any other veterinary medicinal product therefore needs to be made on a case by case basis.

#### **4.9 Amounts to be administered and administration route**

##### Posology:

The dilution possibilities of the different presentations are the following:

- 1 ampoule of 1,000 doses of frozen concentrate in 1 bag of 200 ml of solvent
- 2 ampoules of 1,000 doses of frozen concentrate in 1 bag of 400 ml of solvent
- 3 ampoules of 1,000 doses of frozen concentrate in 1 bag of 600 ml of solvent
- 6 ampoules of 1,000 doses of frozen concentrate in 1 bag of 1200 ml of solvent
- 7 ampoules of 1,000 doses of frozen concentrate in 1 bag of 1400 ml of solvent
- 9 ampoules of 1,000 doses of frozen concentrate in 1 bag of 1800 ml of solvent
- 1 ampoule of 2,000 doses of frozen concentrate in 1 bag of 400 ml of solvent
- 2 ampoules of 2,000 doses of frozen concentrate in 1 bag of 800 ml of solvent
- 6 ampoules of 2,000 doses of frozen concentrate in 1 bag of 2400 ml of solvent

##### Method of administration:

Inoculate to chicks aged 1 day (at the hatchery) by intramuscular route in the leg or by subcutaneous route in the neck, 0.2 ml (1 dose) of diluted vaccine.

##### Dilution of vaccine:

- Remove from liquid nitrogen container only those ampoules carried by the green cane, which are to be used.
- Thaw rapidly the content of the ampoules by stirring in water at 25-30°C.
- As soon as they are completely thawed, open the ampoules holding them at arm's length in order to prevent any risk of injury should the ampoule breaks.
- Draw up the content of an ampoule into a 5-ml sterile syringe immediately after opening of the ampoule.
- Transfer the concentrate into a bag of sterile solvent.
- Draw up 2ml of content of the bag of solvent into the syringe.
- Rinse the ampoule with these 2ml, and then transfer the rinsing liquid into the bag of solvent.

- Repeat the rinsing operation once or twice.
- Repeat the thawing, opening, transfer and rinsing operations for the appropriate number of ampoules to be diluted in the solvent; either 1 ampoule of 1,000 doses of vaccine per 200 ml of solvent (or 1 ampoule of 2,000 doses of vaccine per 400 ml of solvent).
- The diluted vaccine prepared as described is mixed by gentle stirring so as to be ready for use.
- It should be used immediately after preparation.
- Gently agitate the solvent bag with the diluted vaccine from time to time so as to mix the vaccine suspension during the vaccination process.
- Regulate syringe and maintain a volume of 0.2 ml during the operation.

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

No side effects have been reported after administration of a 10-fold overdose of the vaccine.

#### **4.11 Withdrawal Period (s)**

Zero days.

### **5. IMMUNOLOGICAL PROPERTIES**

ATC vet-Code: QI01AD03

Pharmacotherapeutic group: Avian attenuated vaccines: Marek's disease virus.

The vaccine virus is in form of infected chicken embryo cells.

The vaccine stimulates active immunity against Marek's disease.

### **6. PHARMACEUTICAL PARTICULARS**

#### **6.1 List of excipients**

Frozen concentrate:

Dimethyl sulfoxide

Dilution medium (BME vitamin, sodium hydrogen carbonate, phenol red, bovine serum, hydrochloric acid, water for injections)

Solvent:

Sucrose

Casein hydrolysate

Dipotassium phosphate

Potassium dihydrogen phosphate

Sodium hydroxide

Phenol red

Hydrochloric acid

Water for injections

#### **6.2 Major Incompatibilities**

Do not mix with any other veterinary medicinal product except those mentioned in section 4.8 and the solvent supplied for use with the veterinary medicinal product.

### **6.3 Shelf Life**

Shelf-life of the veterinary medicinal product as packaged for sale:

Frozen concentrate: 3 years.

Solvent: 3 years.

Shelf-life after dilution according to directions; 2 hours.

### **6.4 Special Precautions for Storage**

Concentrate:

Store and transport frozen in liquid nitrogen at -196°C.

Solvent:

Store below 30°C. Do not freeze

Storage after dilution: below 25°C.

### **6.5 Nature and composition of immediate packaging**

Frozen concentrate: 2-ml glass (type I) ampoule of 1,000 or 2,000 doses.

Solvent: PVC (polyvinylchloride) bag of 200ml, 400ml, 600ml, 800ml, 1200ml, 1400ml, 1800ml or 2400ml.

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**

*[countries specific]*

## **8. MARKETING AUTHORISATION NUMBER**

*[countries specific]*

## **9. DATE OF FIRST AUTHORISATION/RENEWAL OF THE AUTHORISATION**

*[countries specific]*

## **10. DATE OF REVISION OF THE TEXT**

*[countries specific]*

## **PROHIBITION OF SALE, SUPPLY AND/OR USE**

Not applicable

**ANNEX III**  
**LABELLING AND PACKAGE LEAFLET**

## **A. LABELLING**

**MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS**

1000 doses ampoules  
2000 doses ampoules

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

CRYOMAREX Rispens

**2. QUANTITY OF THE ACTIVE SUBSTANCE(S)**

MDV - Rispens

**3. CONTENTS BY WEIGHT, BY VOLUME OR BY NUMBER OF DOSES**

1000   
2000

**4. ROUTE(S) OF ADMINISTRATION**

IM  
SC

**5. WITHDRAWAL PERIOD**

**6. BATCH NUMBER**

Lot {number}

**7. EXPIRY DATE**

EXP {month/year}

**8. THE WORDS “FOR ANIMAL TREATMENT ONLY”**

Ad us vet

**PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGE**

Bag of solvent of 200 ml , 400 ml, 600 ml, 800 ml, 1200 ml, 1400 ml, 1800 ml or 2400 ml.

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

CRYOMAREX Rispens  
Sterile solvent

**2. STATEMENT OF ACTIVE SUBSTANCES**

**3. PHARMACEUTICAL FORM**

Solvent

**4. PACKAGE SIZE**

200 ml  
400 ml  
600 ml  
800 ml  
1200 ml  
1400 ml  
1800 ml  
2400 ml

**5. TARGET SPECIES**

Chickens (Broilers and replacement chicks)

**6. INDICATION(S)**

**7. METHOD AND ROUTE(S) OF ADMINISTRATION**

Read the package leaflet before use.

**8. WITHDRAWAL PERIOD(S)**

Withdrawal period(s): Zero days

**9. SPECIAL WARNING(S), IF NECESSARY**

**10. EXPIRY DATE**

EXP {month/year}  
Once diluted: 2 hours

**11. SPECIAL STORAGE CONDITIONS**

Store and transport below 30°C. Do not freeze

**12. SPECIAL PRECAUTIONS FOR THE DISPOSAL OF UNUSED PRODUCTS OR WASTE MATERIALS, IF ANY**

Dispose of waste material in accordance with local requirements

**13. THE WORDS “FOR ANIMAL TREATMENT ONLY” AND CONDITIONS OR RESTRICTIONS REGARDING SUPPLY AND USE, IF APPLICABLE**

For animal treatment only. To be supplied only on veterinary prescription.

**14. THE WORDS “KEEP OUT OF THE SIGHT AND REACH OF CHILDREN”**

Keep out of the sight and reach of children

**15. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER**

*To be completed nationally.]*

**16. MARKETING AUTHORISATION NUMBER(S)**

*To be completed nationally.]*

**17. MANUFACTURER’S BATCH NUMBER**

Lot {number}

**B. PACKAGE LEAFLET**

**PACKAGE LEAFLET FOR:**  
CRYOMAREX RISPENS concentrate and solvent for suspension for injection

**1. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER AND OF THE MANUFACTURING AUTHORISATION HOLDER RESPONSIBLE FOR BATCH RELEASE, IF DIFFERENT**

Marketing authorisation holder:

*To be completed nationally.]*

Manufacturer responsible for batch release:

Merial Laboratoire Porte des Alpes  
99 rue de l'Aviation, 69800 Saint Priest  
France

**2. NAME OF THE VETERINARY MEDICINAL PRODUCT**

CRYOMAREX RISPENS  
Concentrate and solvent for suspension for injection

**3. STATEMENT OF THE ACTIVE SUBSTANCE(S) AND OTHER INGREDIENT(S)**

Per dose (0,2 ml):

**Active substance:**

Attenuated live Marek's disease virus, serotype 1, Rispens strain 10<sup>3</sup>-10<sup>4</sup> PFU\*

\*PFU: Plaque forming unit

**Excipient :**

Phenol red 2 µl

Concentrate and solvent for suspension for injection.

Frozen concentrate: homogeneous yellow to red-orange suspension

Solvent: red-orange limpid solution.

**4. INDICATION(S)**

For active immunisation of broilers and replacement chicks i.e. reproductive animals or layers, to reduce mortality, clinical signs and lesions of Marek's disease.

Onset of immunity: 9 days following vaccination.

Duration of immunity: a single vaccination is sufficient to provide protection during the risky period.

**5. CONTRAINDICATIONS**

None

**6. ADVERSE REACTIONS**

None known.

If you notice any side effects, even those not already listed in this package leaflet or you think that the medicine has not worked, please inform your veterinary surgeon.

## **7. TARGET SPECIES**

Chickens (broiler and replacement chick i.e. chicken for reproduction or layer hen).

## **8. DOSAGE FOR EACH SPECIES, ROUTE(S) AND METHOD OF ADMINISTRATION**

The dilution possibilities of the different presentations are the following:

- 1 ampoule of 1,000 doses of frozen concentrate in 1 bag of 200 ml of solvent
- 2 ampoules of 1,000 doses of frozen concentrate in 1 bag of 400 ml of solvent
- 3 ampoules of 1,000 doses of frozen concentrate in 1 bag of 600 ml of solvent
- 6 ampoules of 1,000 doses of frozen concentrate in 1 bag of 1200 ml of solvent
- 7 ampoules of 1,000 doses of frozen concentrate in 1 bag of 1400 ml of solvent
- 9 ampoules of 1,000 doses of frozen concentrate in 1 bag of 1800 ml of solvent
- 1 ampoule of 2,000 doses of frozen concentrate in 1 bag of 400 ml of solvent
- 2 ampoules of 2,000 doses of frozen concentrate in 1 bag of 800 ml of solvent
- 6 ampoules of 2,000 doses of frozen concentrate in 1 bag of 2400 ml of solvent

Inoculate to chicks aged 1 day (at the hatchery), by intramuscular route in the leg or by subcutaneous route in the neck, 0.2 ml (1 dose) of diluted vaccine.

## **9. ADVICE ON CORRECT ADMINISTRATION**

Remove from the liquid nitrogen container only the ampoules carried by the green cane which are to be used.

Thaw rapidly the contents of the ampoules by agitation in water at 25-30°C.

As soon as they are completely thawed, open the ampoules holding them at arm's length in order to prevent any risk of injury should the ampoule breaks. Draw up the vaccine contents into the 5-ml sterile syringe, once the ampoule is opened.

Transfer the concentrate into a bag of sterile solvent.

Draw up to 2 ml of the contents of the bag of solvent into the syringe.

Rinse the ampoule with these 2 ml, then transfer the rinsing liquid into the bag of solvent.

Repeat this rinsing operation once or twice.

The diluted vaccine prepared as described is homogenized by gentle agitation so as to be ready for use.

Repeat the operations of thawing, transfer and rinsing for each ampoule to be diluted with solvent, considering that an ampoule of 1000 doses should be diluted with 200 ml of solvent (and an ampoule of 2000 doses with 400 ml of solvent).

The diluted vaccine prepared as described is homogenized by gentle agitation so as to be ready for use.

It should be used immediately after preparation.

Gently agitate the solvent bag with the diluted vaccine from time to time so as to mix the vaccine suspension during the vaccination process.

Regulate syringe and maintain a volume of 0.2 ml during the operation.

## **10. WITHDRAWAL PERIOD(S)**

Zero days.

## **11. SPECIAL STORAGE PRECAUTIONS**

Keep out of the sight and reach of children.

### Concentrate:

Store and transport frozen in liquid nitrogen at -196°C.

### Solvent:

Store below 30°C. Do not freeze

Storage after dilution: 2 hours at a temperature below 25°C.

Do not use this veterinary medicinal product after the expiry date which is stated on the label after EXP.

## **12. SPECIAL WARNING(S)**

### Special warnings for each target species:

Vaccinate healthy animals only.

### Special precautions for use in animals:

Vaccine strain spreads from vaccinated to unvaccinated birds.

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

The thawing of the ampoules removed from liquid nitrogen may cause an ampoule to break.

Personal protective equipment consisting of protecting gloves and goggles should be worn when handling the veterinary medicinal product. Warnings to be taken are indicated in the section "Advice on correct administration".

### Lay:

Do not use in birds in lay.

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

Safety and efficacy data are available which demonstrate that this vaccine can be mixed with Vaxxitek HVT+IBD, a live HVT vector vaccine expressing the viral protein 2 (VP2) of Infectious Bursal disease (IBD).

No information is available on the safety and efficacy of this vaccine when used with any other veterinary medicinal product, except the product mentioned above. A decision to use this vaccine before or after any other veterinary medicinal product therefore needs to be made on a case by case basis.

### Overdose (symptoms, emergency procedures, antidotes):

No adverse effects have been observed after the administration of an overdose (10 doses).

### Incompatibilities:

Do not mix with another veterinary medicinal product except those mentioned in section SPECIAL WARNING and the solvent supplied for use with the veterinary medicinal product

## **13. SPECIAL PRECAUTIONS FOR THE DISPOSAL OF UNUSED PRODUCT OR WASTE MATERIALS, IF ANY**

Medicinal products should not be poured into sewage or through household glass.

Ask your veterinarian or pharmacist how to get rid of useless medicinal products. Those precautions aim at protecting the environment.

**14. DATE ON WHICH THE PACKAGE LEAFLET WAS LAST APPROVED**

*To be completed nationally.]*

**15. OTHER INFORMATION**

Pack sizes:

Frozen concentrate: 2-ml glass (type I) ampoule of 1,000 or 2,000 doses.

Solvent: PVC (polyvinylchloride) bag of 200ml, 400ml, 600ml, 800ml, 1200ml, 1400ml, 1800ml or 2400ml.

All pack sizes may not be marketed.

Veterinary medicinal product subject to prescription.

Administer under the control of a veterinarian.



is used to replace the wording “doses”