ANNEX I

SUMMARY OF PRODUCT CHARACTERISTICS

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

PREVEXXION RN+HVT+IBD concentrate and solvent for suspension for injection

2. QUALITATIVE AND QUANTITATIVE COMPOSITION

Each dose (0.2 ml for subcutaneous or 0.05 ml for *in ovo*) of the vaccine suspension contains:

Active substances:
Cell-associated, live recombinant Marek's disease (MD) virus, serotype 1, strain RN1250: 2.9 to 3.9 $\log_{10}$ PFU*

Cell-associated, live recombinant turkey herpesvirus (HVT), strain vHVT013-69, expressing the VP2 protein gene of infectious bursal disease (IBD) virus: 3.6 to 4.4 $\log_{10}$ PFU*

*PFU: plaque-forming units

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

3. PHARMACEUTICAL FORM

Concentrate and solvent for suspension for injection.
Concentrate: yellow to reddish pink opalescent homogeneous suspension.
Solvent: red-orange limpid solution.

4. CLINICAL PARTICULARS

4.1 Target species

Chickens.

4.2 Indications for use, specifying the target species

For active immunisation of one-day-old chicks or 18-day-old embryonated chicken eggs:
- to prevent mortality and clinical signs and reduce lesions caused by MD virus (including very virulent MD virus), and
- to prevent mortality and clinical signs and lesions caused by IBD (also known as Gumboro disease) virus.

Onset of immunity: MD: 5 days post-hatch.
IBD: 14 days post-hatch (subcutaneous) or 28 days post-hatch (*in ovo)*.

Duration of immunity: MD: A single vaccination is sufficient to provide protection for the entire risk period.
IBD: 10 weeks post-hatch.

4.3 Contraindications

None.

4.4 Special warnings for each target species

Vaccinate healthy animals only.

Chickens with maternally derived antibodies against MD when vaccinated with this veterinary medicinal product, may have a delayed onset of immunity against IBD.

4.5 Special precautions for use

Special precautions for use in animals
Apply the usual aseptic precautions to all administration procedures.
As this is a live vaccine, both vaccine strains may be excreted from vaccinated birds. The RN1250 vaccine strain has not been shown to spread in experimental conditions. The vHVT013-69 vaccine strain may be spread to unvaccinated chickens and turkeys. Appropriate veterinary and husbandry measures should be taken to avoid spread of the vaccine strains to unvaccinated chickens, turkeys and other susceptible species.

Special precautions to be taken by the person administering the veterinary medicinal product to animals
Personal protective equipment consisting of gloves, spectacles and boots should be worn when handling the veterinary medicinal product, before withdrawing from liquid nitrogen, during the ampoule thawing and opening operations. Frozen glass ampoules may explode during sudden temperature changes. Store and use liquid nitrogen only in a dry and well-ventilated place. Inhalation of the liquid nitrogen is dangerous.

4.6 Adverse reactions (frequency and seriousness)

None.

4.7 Use during pregnancy, lactation or lay

This veterinary medicinal product is designed for one-day-old chicks and 18-day-old embryonated chicken eggs, therefore the safety of the veterinary medicinal product has not been established during lay.

4.8 Interaction with other medicinal products and other forms of interaction

No information is available on the safety and efficacy of this vaccine when used with any other veterinary medicinal product. 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

Subcutaneous and *in ovo* use.
Preparation of the vaccine suspension:

- Wear protective gloves, spectacles and boots during the ampoule thawing and opening operations. The handling of liquid nitrogen should take place in a well-ventilated area.
- Preparation of the vaccine shall be planned before the ampoules are taken from the liquid nitrogen. The exact amount of vaccine ampoules and amount of solvent needed shall be calculated first according to the table below provided as example:

Solvent bag	Number of vaccine ampoules (subcutaneous use)	Number of vaccine ampoules (*in ovo* use)
1 bag of 200 ml solvent	1 ampoule (1000 doses)	4 ampoules (1000 doses) or 2 ampoules (2000 doses) or 1 ampoule (4000 doses)
1 bag of 400 ml solvent	2 ampoules (1000 doses) or 1 ampoule (2000 doses)	8 ampoules (1000 doses) or 4 ampoules (2000 doses) or 2 ampoules (4000 doses)
1 bag of 800 ml solvent	4 ampoules (1000 doses) or 2 ampoules (2000 doses) or 1 ampoule (4000 doses)	16 ampoules (1000 doses) or 8 ampoules (2000 doses) or 4 ampoules (4000 doses)

- Remove from the liquid nitrogen container only those ampoules, which are to be used immediately.
- Thaw the contents of the ampoules rapidly by gentle agitation in water at 25 °C─30 °C. The thawing process should not exceed 90 seconds. Proceed immediately to the next step.
- As soon as they are thawed, wipe the ampoules with a clean paper towel and then open them while holding them at arm's length (in order to prevent injury if any ampoule breaks).
- Select an appropriately sized sterile syringe to withdraw the vaccine from all the ampoules that are thawed, and fit it with a needle of 18 gauge or larger.
- Tear the overpouch on the solvent bag, and then gently insert the syringe needle through the septum of one of the bag-connecting tubes and withdraw 2 ml of solvent.
- Then draw up the complete contents of all the thawed ampoules into the syringe. Do this by slowly drawing up the contents from each ampoule by gently tilting the ampoule forward and inserting the needle with the bevel edge facing downwards towards the bottom of the ampoule. Continue until all the vaccine is drawn out of the ampoule.
- Transfer the syringe contents into the solvent bag (do not use the solvent if it is cloudy).
- Gently mix the vaccine in the solvent bag by moving the bag back and forth.
- It is important to rinse the ampoules and ampoule tips. To do this, draw up a small volume of the solvent containing the vaccine into the syringe. Then slowly fill the ampoule bodies and tips with it. Withdraw the content from the ampoule bodies and tips, and inject it back into the solvent bag.
- Repeat this rinsing operation once.
- Repeat the thawing, opening, transfer and rinsing operations for the appropriate number of ampoules to be diluted in the solvent bag.
- The vaccine is ready for use and should be mixed by gentle agitation and used immediately. During vaccination, gently swirl the bag frequently to ensure the vaccine remains homogenously mixed.
- The vaccine is a clear, red-orange coloured suspension for injection to be used within two hours. Do not freeze it under any circumstances. Do not re-use opened containers of vaccine.

Posology:
One single injection of 0.2 ml per one-day-old chick or 0.05 ml per 18-day-old embryonated chicken egg.

Method of administration:
The vaccine must be administered by subcutaneous injection in the neck or by *in ovo* injection.

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

A limited and transient effect on growth was observed when 10-fold maximum release dose was administered subcutaneously to White Leghorn specified pathogen free chickens.

4.11 Withdrawal period(s)

Zero days.

5. IMMUNOLOGICAL PROPERTIES

Pharmacotherapeutic group: Immunologicals for Aves, live viral vaccines.
ATCvet code QI01AD15

The vaccine contains the recombinant viruses RN1250 and vHVT013-69 within chicken embryo cells.
The RN1250 virus is an engineered MD virus composed of three serotype 1 strains. Its genome also contains long terminal repeats of reticuloendotheliosis virus.
The vHVT013-69 virus is a recombinant HVT expressing the protective antigen (VP2) of the IBD virus strain Faragher 52/70.
The vaccine induces an active immunity and a serological response against Marek's disease and IBD in chickens.

6. PHARMACEUTICAL PARTICULARS

6.1 List of excipients

Vaccine concentrate:
Dimethyl sulfoxide
199 Earle medium
Sodium hydrogen carbonate
Hydrochloric acid
Water for injections

Solvent:
Sucrose
Casein hydrolysate
Phenolsulfonphthalein (Phenol red)
Dipotassium phosphate
Potassium dihydrogen phosphate
Sodium hydroxide or hydrochloric acid
Water for injections

6.2 Major incompatibilities

Do not mix with any other veterinary medicinal product, except 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: 2 years.
Shelf life of the solvent as packaged for sale: 2 years.
Shelf life after vaccine preparation according to directions: 2 hours at a temperature below 25 °C.

6.4 Special precautions for storage

Vaccine concentrate:
Store and transport frozen in liquid nitrogen.
The liquid nitrogen containers must be checked regularly for liquid nitrogen level and must be refilled as needed.
Discard any ampoules that have been accidentally thawed.

Solvent:
Store below 30 °C. Do not freeze. Protect from light.

6.5 Nature and composition of immediate packaging

Vaccine concentrate:
- Type I glass ampoule of 1,000 doses of vaccine, 5-ampoule carrier.
- Type I glass ampoule of 2,000 doses of vaccine, 5-ampoule carrier.
- Type I glass ampoule of 4,000 doses of vaccine, 4-ampoule carrier.

The ampoule carriers are stored firstly in canisters and these canisters are then stored later in the liquid nitrogen containers.

Solvent:
- Polyvinylchloride bag containing 200 ml, 400 ml, 600 ml, 800 ml, 1,000 ml, 1,200 ml, 1,600 ml, 1,800 ml or 2,400 ml.

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

Boehringer Ingelheim Vetmedica GmbH
55216 Ingelheim/Rhein
GERMANY

8. MARKETING AUTHORISATION NUMBER(S)

EU/2/20/255/001-003

9. DATE OF FIRST AUTHORISATION/RENEWAL OF THE AUTHORISATION

Date of first authorisation: 20 July 2020

10. DATE OF REVISION OF THE TEXT

<{MM/YYYY}>
<{DD/MM/YYYY}>
<{DD month YYYY}>

Detailed information on this veterinary medicinal product is available on the website of the European Medicines Agency (http://www.ema.europa.eu/).

PROHIBITION OF SALE, SUPPLY AND/OR USE

Not applicable.

ANNEX II

A. MANUFACTURER OF THE BIOLOGICAL ACTIVE SUBSTANCE AND MANUFACTURER RESPONSIBLE FOR BATCH RELEASE

B. CONDITIONS OR RESTRICTIONS REGARDING SUPPLY AND USE

C. STATEMENT OF THE MRLs

A. MANUFACTURER OF THE BIOLOGICAL ACTIVE SUBSTANCE AND MANUFACTURER RESPONSIBLE FOR BATCH RELEASE

Name and address of the manufacturer of the biological active substance
Manufacturer of the active substance

Boehringer Ingelheim Animal Health France SCS
Laboratoire Porte des Alpes
Rue de l'Aviation
69800 Saint Priest
FRANCE

Name and address of the manufacturer(s) responsible for batch release
Vaccine:
Boehringer Ingelheim Animal Health France SCS
Laboratoire Porte des Alpes
Rue de l'Aviation
69800 Saint Priest
FRANCE

Solvent:
Boehringer Ingelheim Animal Health France SCS
Laboratoire Porte des Alpes
Rue de l'Aviation
69800 Saint Priest
FRANCE

Laboratoire Bioluz
Zone Industrielle de Jalday
64500 Saint Jean de Luz
FRANCE

The printed package leaflet of the medicinal product must state the name and address of the manufacturer responsible for the release of the concerned batch.

B. CONDITIONS OR RESTRICTIONS REGARDING SUPPLY AND USE

Veterinary medicinal product subject to prescription

According to Article 71 of Directive 2001/82/EC of the European Parliament and of the Council as amended, a Member State may, in accordance with its national legislation, prohibit the manufacture, import, possession, sale, supply and/or use of immunological veterinary medicinal products on the whole or part of its territory if it is established that:

a) the administration of the product to animals will interfere with the implementation of a national programme for the diagnosis, control or eradication of animal diseases, or will cause difficulties in certifying the absence of contamination in live animals or in foodstuffs or other products obtained from treated animals.

b) the disease to which the product is intended to confer immunity is largely absent from the territory in question.

C. STATEMENT OF THE MRLs

The active substance being a principle of biological origin intended to produce active immunity is not within the scope of Regulation (EC) No 470/2009.

The excipients listed in section 6.1 of the SPC are either allowed substances for which table 1 of the annex to Commission Regulation (EU) No 37/2010 indicates that no MRLs are required or considered as not falling within the scope of Regulation (EC) No 470/2009 when used as in this veterinary medicinal product.

ANNEX III

LABELLING AND PACKAGE LEAFLET

A. LABELLING

MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS

AMPOULE

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

PREVEXXION RN+HVT+IBD

2. QUANTITY OF THE ACTIVE SUBSTANCE(S)

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

1,000
2,000
4,000

4. ROUTE(S) OF ADMINISTRATION

s.c./SC/*in ovo*

5. WITHDRAWAL PERIOD(S)

6. BATCH NUMBER

Lot {number}

7. EXPIRY DATE

EXP {month/year}

8. THE WORDS "FOR ANIMAL TREATMENT ONLY"

For animal treatment only.

PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGING (LABEL) OF THE DILUENT
(bag)

1. NAME OF THE DILUENT

Solvent for cell associated poultry vaccines.

2. CONTENT BY WEIGHT, BY VOLUME OR BY NUMBER OF DOSES

200 ml
400 ml
600 ml
800 ml
1000 ml
1200 ml
1600 ml
1800 ml
2400 ml

3. ROUTE(S) OF ADMINISTRATION

Read the package leaflet supplied with the vaccine before use.

4. STORAGE CONDITIONS

Store below 30 °C. Do not freeze. Protect from light.

5. BATCH NUMBER

Lot {number}

6. EXPIRY DATE

EXP {month/year}

7. THE WORDS "FOR ANIMAL TREATMENT ONLY"

For animal treatment only.

Boehringer Ingelheim

B. PACKAGE LEAFLET

PACKAGE LEAFLET:
PREVEXXION RN+HVT+IBD 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
Boehringer Ingelheim Vetmedica GmbH
55216 Ingelheim/Rhein
GERMANY

Manufacturer(s) responsible for batch release
Vaccine:
Boehringer Ingelheim Animal Health France SCS
Laboratoire Porte des Alpes
Rue de l'Aviation
69800 Saint-Priest
FRANCE

Solvent:
Boehringer Ingelheim Animal Health France SCS
Laboratoire Porte des Alpes
Rue de l'Aviation
69800 Saint-Priest
FRANCE

Laboratoire Bioluz
Zone Industrielle de Jalday
64500 Saint Jean de Luz
FRANCE

2. NAME OF THE VETERINARY MEDICINAL PRODUCT

PREVEXXION RN+HVT+IBD concentrate and solvent for suspension for injection

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

Each dose (0.2 ml for subcutaneous or 0.05 ml for *in ovo*) of the vaccine suspension contains:

Active substances:
Cell-associated, live recombinant Marek's disease (MD) virus, serotype 1, strain RN1250: 2.9 to 3.9 $\log_{10}$ PFU*

Cell-associated, live recombinant turkey herpesvirus (HVT, strain vHVT013-69), expressing the VP2 protein gene of infectious bursal disease (IBD) virus: 3.6 to 4.4 $\log_{10}$ PFU*

*PFU: plaque-forming units

Concentrate and solvent for suspension for injection.
Concentrate: yellow to reddish pink opalescent homogeneous suspension
Solvent: red-orange limpid solution.

4. INDICATION(S)

For active immunisation of one-day-old chicks or 18-day-old embryonated chicken eggs:
- to prevent mortality and clinical signs and reduce lesions caused by MD virus (including very virulent MD virus), and
- to prevent mortality and clinical signs and lesions caused by IBD (also known as Gumboro disease) virus.

Onset of immunity:	MD: 5 days post-hatch. IBD: 14 days post-hatch (subcutaneous) or 28 days post-hatch (*in ovo)*.
Duration of immunity:	MD: A single vaccination is sufficient to provide protection for the entire risk period. IBD: 10 weeks post-hatch.

5. CONTRAINDICATIONS

None.

6. ADVERSE REACTIONS

None.
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.

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

One single injection of 0.2 ml per one-day-old chick or 0.05 ml per embryonated chicken egg.
The vaccine must be administered by subcutaneous injection in the neck or by *in ovo* injection.

9. ADVICE ON CORRECT ADMINISTRATION

Preparation of the vaccine suspension:
- Wear protective gloves, spectacles and boots during the ampoule thawing and opening operations. The handling of liquid nitrogen should take place in a well-ventilated area.
- Preparation of the vaccine shall be planned before the ampoules are taken from the liquid nitrogen. The exact amount of vaccine ampoules and amount of solvent needed shall be calculated first according to the table below provided as example:

Solvent bag	Number of vaccine ampoules (subcutaneous use)	Number of vaccine ampoules (*in ovo* use)
1 bag of 200 ml solvent	1 ampoule (1000 doses)	4 ampoules (1000 doses) or 2 ampoules (2000 doses) or 1 ampoule (4000 doses)
1 bag of 400 ml solvent	2 ampoules (1000 doses) or 1 ampoule (2000 doses)	8 ampoules (1000 doses) or 4 ampoules (2000 doses) or 2 ampoules (4000 doses)
1 bag of 800 ml solvent	4 ampoules (1000 doses) or 2 ampoules (2000 doses) or 1 ampoule (4000 doses)	16 ampoules (1000 doses) or 8 ampoules (2000 doses) or 4 ampoules (4000 doses)

- Remove from the liquid nitrogen container only those ampoules, which are to be used immediately.
- Thaw the contents of the ampoules rapidly by gentle agitation in water at 25 °C─30 °C. The thawing process should not exceed 90 seconds. Proceed immediately to the next step.
- As soon as they are thawed, wipe the ampoules with a clean paper towel and then open them while holding them at arm's length (in order to prevent injury if any ampoule breaks).
- Select an appropriately sized sterile syringe to withdraw the vaccine from all the ampoules that are thawed, and fit it with a needle of 18 gauge or larger.
- Tear the overpouch on the solvent bag, and then gently insert the syringe needle through the septum of one of the bag-connecting tubes and withdraw 2 ml of solvent.
- Then draw up the complete contents of all the thawed ampoules into the syringe. Do this by slowly drawing up the contents from each ampoule by gently tilting the ampoule forward and inserting the needle with the bevel edge facing downwards towards the bottom of the ampoule. Continue until all the vaccine is drawn out of the ampoule.
- Transfer the syringe contents into the solvent bag (do not use the solvent if it is cloudy).
- Gently mix the vaccine in the solvent bag by moving the bag back and forth.
- It is important to rinse the ampoules and ampoule tips. To do this, draw up a small volume of the solvent containing the vaccine into the syringe. Then slowly fill the ampoule bodies and tips with it. Withdraw the content from the ampoule bodies and tips, and inject it back into the solvent bag.
- Repeat this rinsing operation once.
- Repeat the thawing, opening, transfer and rinsing operations for the appropriate number of ampoules to be diluted in the solvent bag.
- The vaccine is ready for use and should be mixed by gentle agitation and used immediately. During vaccination, gently swirl the bag frequently to ensure the vaccine remains homogenously mixed.
- The vaccine is a clear, red-orange coloured suspension for injection to be used within two hours. Do not freeze the vaccine under any circumstances. Do not re-use opened containers of vaccine.

10. WITHDRAWAL PERIOD(S)

Zero days.

11. SPECIAL STORAGE PRECAUTIONS

Keep out of the sight and reach of children.
The vaccine concentrate must be stored and transported frozen in liquid nitrogen.
The liquid nitrogen containers must be checked regularly for liquid nitrogen level and must be refilled as needed.
Solvent must be stored at a temperature below 30 °C.
Do not freeze.

Protect from light.
Shelf life after vaccine preparation according to directions: 2 hours at a temperature below 25 °C.
Do not use the vaccine after the expiry date stated on the ampoule.

12. SPECIAL WARNING(S)

Special warnings for each target species:
Vaccinate healthy animals only.
Chickens with maternally derived antibodies against MD when vaccinated with this veterinary medicinal product, may have a delayed onset of immunity against IBD.

Special precautions for use in animals
Apply the usual aseptic precautions to all administration procedures.
As this is a live vaccine, both vaccine strains may be excreted from vaccinated birds. The RN1250 vaccine strain has not been shown to spread in experimental conditions. The vHVT013-69 vaccine strain may be spread to unvaccinated chickens and turkeys. Appropriate veterinary and husbandry measures should be taken to avoid spread of the vaccine strains to unvaccinated chickens, turkeys and other susceptible species.

Special precautions to be taken by the person administering the veterinary medicinal product to animals
Personal protective equipment consisting of gloves, spectacles and boots should be worn when handling the veterinary medicinal product, before withdrawing from liquid nitrogen, during the ampoule thawing and opening operations. Frozen glass ampoules may explode during sudden temperature changes. Store and use liquid nitrogen only in a dry and well-ventilated place. Inhalation of the liquid nitrogen is dangerous.

Lay:
This veterinary medicinal product is designed for one-day-old chicks and 18-day-old embryonated chicken eggs, therefore the safety of the veterinary medicinal product has not been established during lay.

Interaction with other medicinal products and other forms of interaction:
No information is available on the safety and efficacy of this vaccine when used with any other veterinary medicinal product. 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), if necessary:
A limited and transient effect on growth was observed when 10-fold maximum release dose was administered subcutaneously to White Leghorn specified pathogen free chickens.

Major incompatibilities:
Do not mix with any other veterinary medicinal product, except solvent supplied for use with the veterinary medicinal product.

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

Discard any ampoules that have been accidentally thawed. Do not re-freeze under any circumstances. Do not re-use opened containers of vaccine.

Medicines should not be disposed of via wastewater or household waste. Ask your veterinary surgeon how to dispose of medicines no longer required. These measures should help to protect the environment.

14. DATE ON WHICH THE PACKAGE LEAFLET WAS LAST APPROVED

Detailed information on this veterinary medicinal product is available on the website of the European Medicines Agency http://www.ema.europa.eu/.

15. OTHER INFORMATION

The vaccine contains the recombinant viruses RN1250 and vHVT013-69 within chicken embryo cells.
The RN1250 virus is an engineered MD virus composed of three serotype 1 strains. Its genome also contains long terminal repeats of reticuloendotheliosis virus.
The vHVT013-69 virus is a recombinant HVT expressing the protective antigen (VP2) of the IBD virus strain Faragher 52/70.
The vaccine induces an active immunity and a serological response against Marek's disease and IBD in chickens.

Package sizes:
Frozen vaccine concentrate:
- Type I glass ampoule of 1,000 doses of vaccine, 5-ampoule carrier.
- Type I glass ampoule of 2,000 doses of vaccine, 5-ampoule carrier.
- Type I glass ampoule of 4,000 doses of vaccine, 4-ampoule carrier.

The ampoule carriers are stored firstly in canisters, and these canisters are then stored later in the liquid nitrogen containers.

Solvent:
- polyvinylchloride bag of 200 ml, 400 ml, 600 ml, 800 ml, 1,000 ml, 1,200 ml, 1,600 ml, 1,800 ml or 2,400 ml.

Not all pack sizes may be marketed.