

ANNEX III
LABELLING AND PACKAGE LEAFLET

A. LABELLING

PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGE

5 LITRE PACK

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Pyceze 500 mg/mL concentrate for solution for fish treatment

Bronopol

2. STATEMENT OF ACTIVE SUBSTANCES

Bronopol 500 mg/mL

3. PHARMACEUTICAL FORM

Concentrate for solution for fish treatment.

4. PACKAGE SIZE

5 litre

5. TARGET SPECIES

Farmed Atlantic salmon and rainbow trout eggs, Atlantic salmon and rainbow trout.

6. INDICATION(S)

7. METHOD AND ROUTE(S) OF ADMINISTRATION

For administration to holding water.

EGGS:

Treat once daily at 50 mg bronopol/litre (1 mL product /10 litres incubator water) for 30 minutes daily commencing 24 hours after fertilisation and until hatch.

If appropriate to the farm management system, dead eggs should be removed on a regular basis to avoid this becoming a source of further fungal infection.

The correct amount of product per volume of incubator should be measured and mixed with at least 10 litres of water prior to addition to the incubator water. Undiluted product in direct contact may damage the surface of plastic troughs.

Use an appropriate measuring device to measure out the correct amount of product. After use, rinse the measuring device carefully and add the rinsings back into the incubator water.

Flow rates to the incubator must be such that a complete exchange of the incubator water is achieved in 30 minutes or less following the end of the treatment period.

It is essential that the fish eggs be exposed to a homogeneous concentration of bronopol throughout the treatment period. This can be achieved by ensuring there is continual water movement in the incubator during the treatment period by following the recommendations below for either re-circulation or continuous addition depending on your choice of application. It is important to maintain the flow rate in the incubator during the treatment period at that normally used on the farm to prevent any physical damage to the sensitive eggs.

Use the equations below to calculate the amount of product required, depending on your choice of application. *It is recommended that you do not change the configuration of your system.* The treatment time may appear longer than 30 minutes – this is to allow time for the product to disperse so that each egg receives the correct dose for the correct time.

RE-CIRCULATION: (DOSE = 1 ML PER 10 LITRES WATER = 50 MG BRONOPOL/LITRE)

The total treatment dose is added to the system over the time it takes for the water to circulate once through the system (Administration Time) to achieve proper mixing. Circulation of the medicated water is then continued for the treatment time of 30 minutes.

$$\text{Administration Time (min)} = \frac{\text{Incubator and pipework volume (litres)}}{\text{Flow rate (litres/min)}}$$

$$\text{Dose of product (mL)} = \frac{\text{Incubator and pipework volume* (litres)}}{10}$$

* = including volume of water the product is diluted in for administration to incubator (Administration Volume)

Add the dose of product during the calculated Administration Time, and then circulate for 30 minutes. The flow of unmedicated water should then resume.

CONTINUOUS ADDITION: (DOSE = 1 ML PER 10 LITRES WATER = 50 MG BRONOPOL/LITRE)

Treatment is added to the system over the time it takes for the water already in the system to achieve the correct dose level. Addition of medicated water is then continued for the treatment time of 30 minutes.

$$\text{Administration Time (min)} = \left(\frac{\text{Incubator and pipework volume (litres)}}{\text{Flow rate (litres/min)}} \right) + 30$$

$$\text{Dose of product (mL)} = \left[\frac{\text{Flow rate (litres/min)} \times \text{Administration Time (mins)}}{10} \right] + \frac{\text{Volume used to dilute the product (Litres)}^*}{10}$$

* = Volume of water product is diluted in for administration to inflow of incubator (Administration Volume)

Add the dose of product during the calculated Administration Time, leaving the in-flow water unchanged.

See package leaflet to illustrate these calculations.

FISH:

Treat once daily at 20 mg bronopol/litre (1 mL product/25 litres water) for 30 minutes daily, for up to 14 consecutive days. Use the equation below to calculate the amount of product required.

$$\text{Dose of product (mL)} = \frac{\text{Treatment Volume (litres)}}{25}$$

After 30 minutes treatment time resume the unmedicated water flow to the treatment unit.

Using an appropriate measuring device add the required volume of product to a bucket containing at least 10 litres of water from the treatment bath. Mix the bucket contents thoroughly and, after stopping the water flow, add to the treatment bath at several positions, spreading the treatment around the treatment bath as much as possible to aid mixing. After use, rinse the measure and the bucket carefully and add the rinsings back into the treatment bath. Undiluted product in direct contact may damage the surface of plastic tanks.

FLOW RATES TO THE TREATMENT UNIT MUST BE SUCH THAT A COMPLETE EXCHANGE OF THE WATER IS ACHIEVED IN 60 MINUTES OR LESS FOLLOWING THE END OF THE TREATMENT PERIOD.

It is recommended that oxygen be diffused into the treatment unit throughout the treatment period to maintain an oxygen level above 7 mg/litre and to ensure adequate mixing of the product.

Movement of the fish and oxygen diffusion during treatment will ensure mixing of the product through the treatment water. The treatment unit volume may be reduced to limit the amount of product used. Care should be taken when doing this not to unduly stress the fish by lack of space or oxygen availability.

For treatment of freshwater cages, the net should be raised to a depth of 1 to 2 metres depending on the biomass of the fish. The cage to be treated should then be surrounded by an impervious tarpaulin to completely isolate it. Oxygen should be

supplied and treatment carried out as above. Once the 30 minutes treatment time is completed, the tarpaulin should be removed and the nets lowered to the original depth.

Treat daily for up to 14 consecutive days according to veterinary assessment.

Read the package leaflet before use.

8. WITHDRAWAL PERIOD(S)

Do not use in salmonid eggs intended for human consumption. Meat – zero degree days.

9. SPECIAL WARNING(S), IF NECESSARY

CONTRAINDICATIONS/WARNINGS: Do not use in smolting Atlantic salmon or rainbow trout alevins as studies have indicated increased toxicity at this stage. Do not conduct bath treatments with other products at the same time as bath treatments with bronopol. Use in broodstock should only be in accordance with a benefit/risk assessment by the responsible veterinarian because reproductive toxicity has not been established in the target species. Treatment with Pyceze is most effective if the infection is treated early and before there is a chance for the fungus to invade the underlying muscle or before there is a build-up of a dense mat of dead eggs.

Data appears to demonstrate that Pyceze, once diluted in a fixed volume of water, may be degraded by prolonged or repeated exposure to high intensity ultraviolet light. Degradation of Pyceze may alter its toxicity profile. It is therefore recommended that Pyceze treated water is not passed repeatedly through ultraviolet sterilising filters.

Accurately estimate the volume of water to be treated.

Use at water temperatures of more than 13 °C only in accordance with a benefit/risk assessment by the responsible veterinarian, as efficacy and safety at temperatures above 13 °C has not been demonstrated.

OPERATOR WARNINGS:

Irritating to eyes, lungs and skin.

Wear protective clothing, impervious gloves, face mask and suitable eye protection when mixing and handling the product. (For instance 0.3 mm nitrile rubber gloves, disposable half mask respirator conforming to European Standard EN149 together with suitable eye protection or a non-disposable respirator conforming to European Standard EN140 with a filter to EN143).

If in contact with the skin, wash immediately and – in case of contact with the eyes – rinse immediately with plenty of water and seek medical advice.

Do not smoke, drink or eat when using the product and wash hands carefully after use. For full warnings and precautions refer to package leaflet.

10. EXPIRY DATE

EXP {month/year}

11. SPECIAL STORAGE CONDITIONS

Store in the original container. Keep the container tightly closed. Do not store above 25 °C. Shelf-life after first opening the immediate packaging: 6 months
Once opened, use by
Once diluted, the product must be used immediately.

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

Dispose of waste material in accordance with local requirements.
Disposal: read package leaflet.

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

CZ Vaccines S.A.U.
36410 O Porriño – Spain

16. MARKETING AUTHORISATION NUMBER(S)

Vm 30824/4004

17. MANUFACTURER’S BATCH NUMBER

Lot {number}

PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGE

1 LITRE PACK

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Pyceze 500 mg/mL concentrate for solution for fish treatment

Bronopol

2. STATEMENT OF ACTIVE SUBSTANCES

Bronopol 500 mg/mL

3. PHARMACEUTICAL FORM

Concentrate for solution for fish treatment.

4. PACKAGE SIZE

1 litre

5. TARGET SPECIES

Farmed Atlantic salmon and rainbow trout eggs, Atlantic salmon and rainbow trout.

6. INDICATION(S)

7. METHOD AND ROUTE(S) OF ADMINISTRATION

For administration to holding water.

EGGS:

Treat once daily at 50 mg bronopol/litre (1 mL product /10 litres incubator water) for 30 minutes daily commencing 24 hours after fertilisation and until hatch.

If appropriate to the farm management system, dead eggs should be removed on a regular basis to avoid this becoming a source of further fungal infection.

The correct amount of product per volume of incubator should be measured and mixed with at least 10 litres of water prior to addition to the incubator water. Undiluted product in direct contact may damage the surface of plastic troughs.

Use an appropriate measuring device to measure out the correct amount of product. After use, rinse the measuring device carefully and add the rinsings back into the incubator water.

Flow rates to the incubator must be such that a complete exchange of the incubator water is achieved in 30 minutes or less following the end of the treatment period.

It is essential that the fish eggs be exposed to a homogeneous concentration of bronopol throughout the treatment period. This can be achieved by ensuring there is continual water movement in the incubator during the treatment period by following the recommendations below for either re-circulation or continuous addition depending on your choice of application. It is important to maintain the flow rate in the incubator during the treatment period at that normally used on the farm to prevent any physical damage to the sensitive eggs.

Use the equations below to calculate the amount of product required, depending on your choice of application. *It is recommended that you do not change the configuration of your system.* The treatment time may appear longer than 30 minutes – this is to allow time for the product to disperse so that each egg receives the correct dose for the correct time.

RE-CIRCULATION: (DOSE = 1 ML PER 10 LITRES WATER = 50 MG BRONOPOL/LITRE)

The total treatment dose is added to the system over the time it takes for the water to circulate once through the system (Administration Time) to achieve proper mixing. Circulation of the medicated water is then continued for the treatment time of 30 minutes.

$$\text{Administration Time (min)} = \frac{\text{Incubator and pipework volume (litres)}}{\text{Flow rate (litres/min)}}$$

$$\text{Dose of product (mL)} = \frac{\text{Incubator and pipework volume* (litres)}}{10}$$

* = including volume of water the product is diluted in for administration to incubator (Administration Volume)

Add the dose of product during the calculated Administration Time, and then circulate for 30 minutes. The flow of unmedicated water should then resume.

CONTINUOUS ADDITION: (DOSE = 1 ML PER 10 LITRES WATER = 50 MG BRONOPOL/LITRE)

Treatment is added to the system over the time it takes for the water already in the system to achieve the correct dose level. Addition of medicated water is then continued for the treatment time of 30 minutes.

$$\text{Administration Time (min)} = \left(\frac{\text{Incubator and pipework volume (litres)}}{\text{Flow rate (litres/min)}} \right) + 30$$

$$\text{Dose of product (mL)} = \left[\frac{\text{Flow rate (litres/min)} \times \text{Administration Time (mins)}}{10} \right] + \frac{\text{Volume used to dilute the product (Litres)}^*}{10}$$

* = Volume of water product is diluted in for administration to inflow of incubator (Administration Volume)

Add the dose of product during the calculated Administration Time, leaving the in-flow water unchanged.

See package leaflet to illustrate these calculations.

FISH:

Treat once daily at 20 mg bronopol/litre (1 mL product/25 litres water) for 30 minutes daily, for up to 14 consecutive days. Use the equation below to calculate the amount of product required.

$$\text{Dose of product (mL)} = \frac{\text{Treatment Volume (litres)}}{25}$$

After 30 minutes treatment time resume the unmedicated water flow to the treatment unit.

Using an appropriate measuring device add the required volume of product to a bucket containing at least 10 litres of water from the treatment bath. Mix the bucket contents thoroughly and, after stopping the water flow, add to the treatment bath at several positions, spreading the treatment around the treatment bath as much as possible to aid mixing. After use, rinse the measure and the bucket carefully and add the rinsings back into the treatment bath. Undiluted product in direct contact may damage the surface of plastic tanks.

FLOW RATES TO THE TREATMENT UNIT MUST BE SUCH THAT A COMPLETE EXCHANGE OF THE WATER IS ACHIEVED IN 60 MINUTES OR LESS FOLLOWING THE END OF THE TREATMENT PERIOD.

It is recommended that oxygen be diffused into the treatment unit throughout the treatment period to maintain an oxygen level above 7 mg/litre and to ensure adequate mixing of the product.

Movement of the fish and oxygen diffusion during treatment will ensure mixing of the product through the treatment water. The treatment unit volume may be reduced to limit the amount of product used. Care should be taken when doing this not to unduly stress the fish by lack of space or oxygen availability.

For treatment of freshwater cages, the net should be raised to a depth of 1 to 2 metres depending on the biomass of the fish. The cage to be treated should then be surrounded by an impervious tarpaulin to completely isolate it. Oxygen should be

supplied and treatment carried out as above. Once the 30 minutes treatment time is completed, the tarpaulin should be removed and the nets lowered to the original depth.

Treat daily for up to 14 consecutive days according to veterinary assessment.

Read the package leaflet before use.

8. WITHDRAWAL PERIOD(S)

Do not use in salmonid eggs intended for human consumption. Meat – zero degree days.

9. SPECIAL WARNING(S), IF NECESSARY

CONTRAINDICATIONS/WARNINGS: Do not use in smolting Atlantic salmon or rainbow trout alevins as studies have indicated increased toxicity at this stage. Do not conduct bath treatments with other products at the same time as bath treatments with bronopol. Use in broodstock should only be in accordance with a benefit/risk assessment by the responsible veterinarian because reproductive toxicity has not been established in the target species. Treatment with Pyceze is most effective if the infection is treated early and before there is a chance for the fungus to invade the underlying muscle or before there is a build-up of a dense mat of dead eggs.

Data appears to demonstrate that Pyceze, once diluted in a fixed volume of water, may be degraded by prolonged or repeated exposure to high intensity ultraviolet light. Degradation of Pyceze may alter its toxicity profile. It is therefore recommended that Pyceze treated water is not passed repeatedly through ultraviolet sterilising filters.

Accurately estimate the volume of water to be treated.

Use at water temperatures of more than 13 °C only in accordance with a benefit/risk assessment by the responsible veterinarian, as efficacy and safety at temperatures above 13 °C has not been demonstrated.

OPERATOR WARNINGS:

Irritating to eyes, lungs and skin.

Wear protective clothing, impervious gloves, face mask and suitable eye protection when mixing and handling the product. (For instance 0.3 mm nitrile rubber gloves, disposable half mask respirator conforming to European Standard EN149 together with suitable eye protection or a non-disposable respirator conforming to European Standard EN140 with a filter to EN143).

If in contact with the skin, wash immediately and – in case of contact with the eyes – rinse immediately with plenty of water and seek medical advice.

Do not smoke, drink or eat when using the product and wash hands carefully after use. For full warnings and precautions refer to package leaflet.

10. EXPIRY DATE

EXP {month/year}

11. SPECIAL STORAGE CONDITIONS

Store in the original container. Keep the container tightly closed. Do not store above 25 °C. Shelf-life after first opening the immediate packaging: 6 months
Once opened, use by
Once diluted, the product must be used immediately.

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

This product is designed for use following dilution and waterborne administration. Without any dilution the product is harmful to other aquatic life.

A discharge consent may be required for release of this product into the aquatic environment. This must be obtained, prior to use of the product, from the relevant authority. Otherwise the product should only be used if the flow rate of untreated waters allows for a minimum dilution of the volume of treated water. For fish treatment the product should only be used if the flow rate of untreated water allows a dilution of 1:2000 times the volume of treated water. For egg treatment the product should only be used if the flow rate of untreated water allows a dilution of 1:5000 times the volume of treated water. Where the appropriate dilution of treated water cannot be achieved, the farm must have a discharge process to limit the release of product into the environment to within the parameters described. This can be achieved by the use of holding tanks and ponds, discharge lagoons and biofilters to clean treated water. Where this applies, the user must monitor the discharge concentration to ensure the parameters are not exceeded.

Do not contaminate surface water or ditches with product or used container.

Any unused veterinary medicinal product or waste materials derived from such veterinary medicinal products should be disposed of 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

CZ Vaccines S.A.U.
36410 O Porriño – Spain

16. MARKETING AUTHORISATION NUMBER(S)

Vm 30824/4004

17. MANUFACTURER'S BATCH NUMBER

Lot {number}

PARTICULARS TO APPEAR ON THE OUTER PACKAGE

5 LITRE CARTON

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Pyceze 500 mg/mL concentrate for solution for fish treatment

Bronopol

2. STATEMENT OF ACTIVE SUBSTANCES

Bronopol 500 mg/mL

3. PHARMACEUTICAL FORM

Concentrate for solution for fish treatment.

4. PACKAGE SIZE

5 litre

5. TARGET SPECIES

Farmed Atlantic salmon and rainbow trout eggs, Atlantic salmon and rainbow trout.

6. INDICATION(S)

7. METHOD AND ROUTE(S) OF ADMINISTRATION

For administration to holding water.

EGGS:

Treat once daily at 50 mg bronopol/litre (1 mL product /10 litres incubator water) for 30 minutes daily commencing 24 hours after fertilisation and until hatch.

If appropriate to the farm management system, dead eggs should be removed on a regular basis to avoid this becoming a source of further fungal infection.

The correct amount of product per volume of incubator should be measured and mixed with at least 10 litres of water prior to addition to the incubator water. Undiluted product in direct contact may damage the surface of plastic troughs.

Use an appropriate measuring device to measure out the correct amount of product. After use, rinse the measuring device carefully and add the rinsings back into the incubator water.

Flow rates to the incubator must be such that a complete exchange of the incubator water is achieved in 30 minutes or less following the end of the treatment period.

It is essential that the fish eggs be exposed to a homogeneous concentration of bronopol throughout the treatment period. This can be achieved by ensuring there is continual water movement in the incubator during the treatment period by following the recommendations below for either re-circulation or continuous addition depending on your choice of application. It is important to maintain the flow rate in the incubator during the treatment period at that normally used on the farm to prevent any physical damage to the sensitive eggs.

Use the equations below to calculate the amount of product required, depending on your choice of application. *It is recommended that you do not change the configuration of your system.* The treatment time may appear longer than 30 minutes – this is to allow time for the product to disperse so that each egg receives the correct dose for the correct time.

RE-CIRCULATION: (DOSE = 1 ML PER 10 LITRES WATER = 50 MG BRONOPOL/LITRE)

The total treatment dose is added to the system over the time it takes for the water to circulate once through the system (Administration Time) to achieve proper mixing. Circulation of the medicated water is then continued for the treatment time of 30 minutes.

$$\text{Administration Time (min)} = \frac{\text{Incubator and pipework volume (litres)}}{\text{Flow rate (litres/min)}}$$

$$\text{Dose of product (mL)} = \frac{\text{Incubator and pipework volume* (litres)}}{10}$$

* = including volume of water the product is diluted in for administration to incubator (Administration Volume)

Add the dose of product during the calculated Administration Time, and then circulate for 30 minutes. The flow of unmedicated water should then resume.

CONTINUOUS ADDITION: (DOSE = 1 ML PER 10 LITRES WATER = 50 MG BRONOPOL/LITRE)

Treatment is added to the system over the time it takes for the water already in the system to achieve the correct dose level. Addition of medicated water is then continued for the treatment time of 30 minutes.

$$\text{Administration Time (min)} = \left(\frac{\text{Incubator and pipework volume (litres)}}{\text{Flow rate (litres/min)}} \right) + 30$$

$$\text{Dose of product (mL)} = \left[\frac{\text{Flow rate (litres/min)} \times \text{Administration Time (mins)}}{10} \right] + \frac{\text{Volume used to dilute the product (Litres)}^*}{10}$$

* = Volume of water product is diluted in for administration to inflow of incubator (Administration Volume)

Add the dose of product during the calculated Administration Time, leaving the in-flow water unchanged.

See package leaflet to illustrate these calculations.

FISH:

Treat once daily at 20 mg bronopol/litre (1 mL product/25 litres water) for 30 minutes daily, for up to 14 consecutive days. Use the equation below to calculate the amount of product required.

$$\text{Dose of product (mL)} = \frac{\text{Treatment Volume (litres)}}{25}$$

After 30 minutes treatment time resume the unmedicated water flow to the treatment unit.

Using an appropriate measuring device add the required volume of product to a bucket containing at least 10 litres of water from the treatment bath. Mix the bucket contents thoroughly and, after stopping the water flow, add to the treatment bath at several positions, spreading the treatment around the treatment bath as much as possible to aid mixing. After use, rinse the measure and the bucket carefully and add the rinsings back into the treatment bath. Undiluted product in direct contact may damage the surface of plastic tanks.

FLOW RATES TO THE TREATMENT UNIT MUST BE SUCH THAT A COMPLETE EXCHANGE OF THE WATER IS ACHIEVED IN 60 MINUTES OR LESS FOLLOWING THE END OF THE TREATMENT PERIOD.

It is recommended that oxygen be diffused into the treatment unit throughout the treatment period to maintain an oxygen level above 7 mg/litre and to ensure adequate mixing of the product.

Movement of the fish and oxygen diffusion during treatment will ensure mixing of the product through the treatment water. The treatment unit volume may be reduced to limit the amount of product used. Care should be taken when doing this not to unduly stress the fish by lack of space or oxygen availability.

For treatment of freshwater cages, the net should be raised to a depth of 1 to 2 metres depending on the biomass of the fish. The cage to be treated should then be surrounded by an impervious tarpaulin to completely isolate it. Oxygen should be

supplied and treatment carried out as above. Once the 30 minutes treatment time is completed, the tarpaulin should be removed and the nets lowered to the original depth.

Treat daily for up to 14 consecutive days according to veterinary assessment.

Read the package leaflet before use.

8. WITHDRAWAL PERIOD(S)

Do not use in salmonid eggs intended for human consumption. Meat – zero degree days.

9. SPECIAL WARNING(S), IF NECESSARY

CONTRAINDICATIONS/WARNINGS: Do not use in smolting Atlantic salmon or rainbow trout alevins as studies have indicated increased toxicity at this stage. Do not conduct bath treatments with other products at the same time as bath treatments with bronopol. Use in broodstock should only be in accordance with a benefit/risk assessment by the responsible veterinarian because reproductive toxicity has not been established in the target species. Treatment with Pyceze is most effective if the infection is treated early and before there is a chance for the fungus to invade the underlying muscle or before there is a build-up of a dense mat of dead eggs.

Data appears to demonstrate that Pyceze, once diluted in a fixed volume of water, may be degraded by prolonged or repeated exposure to high intensity ultraviolet light. Degradation of Pyceze may alter its toxicity profile. It is therefore recommended that Pyceze treated water is not passed repeatedly through ultraviolet sterilising filters.

Accurately estimate the volume of water to be treated.

Use at water temperatures of more than 13 °C only in accordance with a benefit/risk assessment by the responsible veterinarian, as efficacy and safety at temperatures above 13 °C has not been demonstrated.

OPERATOR WARNINGS:

Irritating to eyes, lungs and skin.

Wear protective clothing, impervious gloves, face mask and suitable eye protection when mixing and handling the product. (For instance 0.3 mm nitrile rubber gloves, disposable half mask respirator conforming to European Standard EN149 together with suitable eye protection or a non-disposable respirator conforming to European Standard EN140 with a filter to EN143).

If in contact with the skin, wash immediately and – in case of contact with the eyes – rinse immediately with plenty of water and seek medical advice.

Do not smoke, drink or eat when using the product and wash hands carefully after use. For full warnings and precautions refer to package leaflet.

10. EXPIRY DATE

EXP {month/year}

11. SPECIAL STORAGE CONDITIONS

Store in the original container. Keep the container tightly closed. Do not store above 25 °C. Shelf-life after first opening the immediate packaging: 6 months
Once opened, use by
Once diluted, the product must be used immediately.

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

Dispose of waste material in accordance with local requirements.
Disposal: read package leaflet.

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

CZ Vaccines S.A.U.
36410 O Porriño – Spain

16. MARKETING AUTHORISATION NUMBER(S)

Vm 30824/4004

17. MANUFACTURER’S BATCH NUMBER

Lot {number}

PARTICULARS TO APPEAR ON THE OUTER PACKAGE

4 X 5 LITRE CARTONS

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Pyceze 500 mg/mL concentrate for solution for fish treatment

Bronopol

2. STATEMENT OF ACTIVE SUBSTANCES

Bronopol 500 mg/mL

3. PHARMACEUTICAL FORM

Concentrate for solution for fish treatment.

4. PACKAGE SIZE

4 x 5 litre bottles

5. TARGET SPECIES

Farmed Atlantic salmon and rainbow trout eggs, Atlantic salmon and rainbow trout.

6. INDICATION(S)

7. METHOD AND ROUTE(S) OF ADMINISTRATION

For administration in holding water.

Dosage instructions appear on the container label and package leaflet. You should receive a measure with this bottle.

8. WITHDRAWAL PERIOD(S)

Do not use in salmonid eggs intended for human consumption. Meat – zero degree days.

9. SPECIAL WARNING(S), IF NECESSARY

OPERATOR WARNINGS:

Irritating to eyes, lungs and skin.

Wear protective clothing, impervious gloves, face mask and suitable eye protection when mixing and handling the product. (For instance 0.3 mm nitrile rubber gloves, disposable half mask respirator conforming to European Standard EN149 together with suitable eye protection or a non-disposable respirator conforming to European Standard EN140 with a filter to EN143).

If in contact with the skin, wash immediately and – in case of contact with the eyes – rinse immediately with plenty of water and seek medical advice.

Do not smoke, drink or eat when using the product and wash hands carefully after use. For full warnings and precautions refer to package leaflet.

10. EXPIRY DATE

EXP {month/year}

11. SPECIAL STORAGE CONDITIONS

Store in the original container. Keep the container tightly closed. Do not store above 25 °C. Shelf-life after first opening the immediate packaging: 6 months
Once diluted, the product must be used immediately.

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

Dispose of waste material in accordance with local requirements.
Disposal: read package leaflet.

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

CZ Vaccines S.A.U.
36410 O Porriño – Spain

16. MARKETING AUTHORISATION NUMBER(S)

Vm 30824/4004

17. MANUFACTURER'S BATCH NUMBER

Lot {number}

B. PACKAGE LEAFLET

PACKAGE LEAFLET:
Pyceze 500 mg/mL concentrate for solution for fish treatment
Bronopol

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

Marketing authorisation holder and manufacturer responsible for batch release:

CZ Vaccines S.A.U.
A Relva s/n – Torneiros
36410 O Porriño
Pontevedra
Spain

2. NAME OF THE VETERINARY MEDICINAL PRODUCT

Pyceze 500 mg/mL concentrate for solution for fish treatment
Bronopol

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

Each 1 mL contains: Bronopol 500 mg

4. INDICATION(S)

The product is indicated for:

- Prevention of growth of fungal infections (*Saprolegnia* spp) in the face of suspected or known challenge in farmed Atlantic salmon eggs and rainbow trout eggs.
- Prevention or reduction of fungal infections (*Saprolegnia* spp) in farmed Atlantic salmon and rainbow trout kept in fresh water.

The product is most effective if used at the first signs of fungal infection.

5. CONTRAINDICATIONS

Do not use in smolting Atlantic salmon or rainbow trout alevins as studies have indicated increased toxicity at this stage.

Do not conduct bath treatments with other products at the same time as bath treatments with bronopol.

6. ADVERSE REACTIONS

Occasionally some restlessness has been noted among fish during addition of the product to treatment water.

If you notice any serious effects or other effects not mentioned in this leaflet, please inform your veterinarian.

7. TARGET SPECIES

Farmed Atlantic salmon and rainbow trout eggs, Atlantic salmon and rainbow trout.

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

For administration to holding water.

Tear off the internal aluminium seal carefully, and avoid the falling of small pieces into the bottle.

EGGS:

Treat once daily at 50 mg bronopol/litre (1 mL product /10 litres incubator water) for 30 minutes daily commencing 24 hours after fertilisation and until hatch.

If appropriate to the farm management system, dead eggs should be removed on a regular basis to avoid this becoming a source of further fungal infection.

The correct amount of product per volume of incubator should be measured and mixed with at least 10 litres of water prior to addition to the incubator water. Undiluted product in direct contact may damage the surface of plastic troughs.

Use an appropriate measuring device to measure out the correct amount of product. After use, rinse the measuring device carefully and add the rinsings back into the incubator water. **Flow rates to the incubator must be such that a complete exchange of the incubator water is achieved in 30 minutes or less following the end of the treatment period.**

It is essential that the fish eggs be exposed to a homogeneous concentration of bronopol throughout the treatment period. This can be achieved by ensuring there is continual water movement in the incubator during the treatment period by following the recommendations below for either re-circulation or continuous addition depending on your choice of application. It is important to maintain the flow rate in the incubator during the treatment period at that normally used on the farm to prevent any physical damage to the sensitive eggs.

Use the equations below to calculate the amount of product required, depending on your choice of application. *It is recommended that you do not change the configuration of your system.* The treatment time may appear longer than 30 minutes – this is to allow time for the product to disperse so that each egg receives the correct dose for the correct time.

RE-CIRCULATION: (DOSE = 1 ML PER 10 LITRES WATER = 50 MG BRONOPOL/LITRE)

The total treatment dose is added to the system over the time it takes for the water to circulate once through the system (Administration Time) to achieve proper mixing. Circulation of the medicated water is then continued for the treatment time of 30 minutes.

$$\text{Administration Time (min)} = \frac{\text{Incubator and pipework volume (litres)}}{\text{Flow rate (litres/min)}}$$

$$\text{Dose of product (mL)} = \frac{\text{Incubator and pipework volume}^* \text{ (litres)}}{10}$$

* = including volume of water the product is diluted in for administration to incubator (Administration Volume)

Add the dose of product during the calculated Administration Time, and then circulate for 30 minutes. The flow of unmedicated water should then resume.

CONTINUOUS ADDITION: (DOSE = 1 ML PER 10 LITRES WATER = 50 MG BRONOPOL/LITRE)

Treatment is added to the system over the time it takes for the water already in the system to achieve the correct dose level. Addition of medicated water is then continued for the treatment time of 30 minutes.

$$\text{Administration Time (min)} \left\{ = \frac{\text{Incubator and pipework volume (litres)}}{\text{Flow rate (litres/min)}} \right\} + 30$$

$$\text{Dose of product (mL)} = \frac{[\text{Flow rate (litres/min)} \times \text{Administration Time (mins)}]}{10} + \frac{\text{Volume used to dilute the product (Litres)}^*}{10}$$

* = Volume of water product is diluted in for administration to inflow of incubator (Administration Volume)

Add the dose of product during the calculated Administration Time, leaving the in-flow water unchanged.

See under Dose Calculation for examples of these calculations.

FISH:

Treat once daily at 20 mg bronopol/litre (1 mL product/25 litres water) for 30 minutes daily, for up to 14 consecutive days. Use the equation below to calculate the amount of product required.

$$\text{Dose of product (mL)} = \frac{\text{Treatment Volume (litres)}}{25}$$

After 30 minutes treatment time resume the unmedicated water flow to the treatment unit.

Using an appropriate measuring device add the required volume of product to a bucket containing at least 10 litres of water from the treatment bath. Mix the bucket contents thoroughly and, after stopping the water flow, add to the treatment bath at several positions, spreading the treatment around the treatment bath as much as

possible to aid mixing. After use, rinse the measure and the bucket carefully and add the rinsings back into the treatment bath. Undiluted product in direct contact may damage the surface of plastic tanks.

FLOW RATES TO THE TREATMENT UNIT MUST BE SUCH THAT A COMPLETE EXCHANGE OF THE WATER IS ACHIEVED IN 60 MINUTES OR LESS FOLLOWING THE END OF THE TREATMENT PERIOD.

It is recommended that oxygen be diffused into the treatment unit throughout the treatment period to maintain an oxygen level above 7 mg/litre and to ensure adequate mixing of the product.

Movement of the fish and oxygen diffusion during treatment will ensure mixing of the product through the treatment water. The treatment unit volume may be reduced to limit the amount of product used. Care should be taken when doing this not to unduly stress the fish by lack of space or oxygen availability.

For treatment of freshwater cages, the net should be raised to a depth of 1 to 2 metres depending on the biomass of the fish. The cage to be treated should then be surrounded by an impervious tarpaulin to completely isolate it. Oxygen should be supplied and treatment carried out as above. Once the 30 minutes treatment time is completed, the tarpaulin should be removed and the nets lowered to the original depth.

Treat daily for up to 14 consecutive days according to veterinary assessment.

9. ADVICE ON CORRECT ADMINISTRATION

GUIDELINES FOR PYCEZE TREATMENT OF ATLANTIC SALMON AND RAINBOW TROUT

Graduated 500 mL measuring device is available with the 5 litre pack size.

Treatment duration is 30 minutes.

DOSE CALCULATION:

$$\text{Dose of Pyceze (mL)} = \frac{\text{Treatment Volume (litres)}}{25}$$

PYCEZE TREATMENT OF FISH IS BY CONVENTIONAL BATH TREATMENT

- Where fish are in tanks, raceways or earth ponds the water supply should be turned off. The treatment unit volume may be reduced to limit the amount of product used. Care should be taken when doing this not to unduly stress the fish by lack of space or oxygen availability.
- For treatment of freshwater cages, the net should be raised to a depth of 1 m to 2 m depending on the biomass of the fish and a tarpaulin should be used to completely enclose the fish in a known volume of water. It is important to accurately determine the volume of water to be medicated. Once the 30 minute treatment time is completed, the tarpaulin should be removed and the

- nets lowered to the original depth.
- In all cases it is recommended that oxygen be diffused into the treatment unit throughout the treatment period to maintain an oxygen level above 7 mg/L and to ensure adequate mixing of the product.
- Flow rates to the treatment unit must be such that a complete exchange of water is achieved in 60 minutes or less following the end of the treatment period.

TREATMENT:

1. Using an appropriate measuring device to add the required volume of Pyceze to a bucket containing at least 10 litres of water from the treatment bath.
2. Mix the bucket contents thoroughly and, after stopping the water flow, add to the treatment bath at several positions, spreading the treatment around the bath as much as possible to aid mixing.
3. After use, rinse the measure and bucket used to mix the Pyceze carefully and add the rinsings to the treatment bath to ensure all product is used.
4. Undiluted product in direct contact may damage the surface of plastic tanks.
5. Once all the product is added, check the time and keep the fish in the treatment bath for 30 minutes.
6. If fish become stressed or agitated abandon treatment, turn on the water supply and contact your veterinarian surgeon.
7. At the end of the treatment period, the water flow should be turned back on and the tank/raceway/pond allowed to fill and treatment flushed out. Only now should the additional oxygen supply be removed. For cages, remove tarpaulin.
8. Consecutive daily treatments for up to 14 days are recommended.

salmonid rearing system		Treatment bath dimensions	Treatment volume	Dose of Pyceze
Tank	e.g. 1	2 m diameter x 1 m depth	3,142 litres	126 mL
	e.g. 2	5 m diameter x 2 m depth	39,270 litres	1.57 litres
Raceway/pond	e.g. 1	5 m long x 2 m wide x 1 m depth	10 m ³ (10,000 litres)	400 mL
	e.g. 2	10 m long x 5 m wide x 1.5 m depth	75 m ³ (75,000 litres)	3.0 litres
Freshwater cage (square)	e.g. 1	5 m x 5 m x 1 m	25 m ³ (25,000 litres)	1.0 litre
	e.g. 2	12 m x 12 m x 1 m ^o	144 m ³ (144,000 litres)	5.76 litres

GUIDELINES FOR PYCEZE TREATMENT OF SALMONID EGGS

It is recommended that you do NOT change from your normal incubation system whilst medicating with Pyceze.

Treat once daily at 50 mg bronopol/litre (1 mL Pyceze/10 litres incubator water) for 30 minutes commencing 24 hours after fertilisation until hatch.

The correct amount of product per volume of incubator should be measured and mixed with at least 10 litres of incubator water prior to addition to the incubator. Undiluted product in direct contact may damage the surface of plastic troughs.

Use an appropriate measuring device to measure out the correct amount of product. After use, rinse the measuring device carefully and add the rinsings back into the incubator water. Flow rates to the incubator must be such that a complete exchange of the incubator water is achieved in 30 minutes or less following the end of the treatment period.

It is essential that the fish eggs be exposed to a homogeneous concentration of Pyceze throughout the treatment period. This can be achieved by ensuring there is continual water movement in the incubator during the treatment period by following the recommendations below for either re-circulation or continuous addition depending on the choice of application. It is important to maintain the flow rate in the incubator during the treatment period at that normally used on the farm to prevent any physical damage to the sensitive eggs.

Use the equations below to calculate the amount of product required, depending on your choice of application. *It is recommended that you do not change the configuration of your system.* The treatment time may appear longer than 30 minutes – this is to allow time for the product to disperse so that each egg received the correct dose for the correct time.

DECISION:

Will you be re-circulating water during treatment? If YES then follow treatment and dose calculation 1. If NO then follow treatment and dose calculation 2.

1. RE-CIRCULATION: (DOSE = 1 ML PER 10 LITRES WATER = 50 MG BRONOPOL/LITRE)

Total treatment dose is added to the system over the time it takes for the water to circulate once through the system (Administration Time), to achieve proper mixing.

Circulation of the medicated water is then continued for the treatment time of 30 minutes.

$$\text{Administration Time (min)} = \frac{\text{Incubator and pipework volume (litres)}}{\text{Flow rate (litres/min)}}$$

$$\text{Dose of product (mL)} = \frac{\text{Incubator and pipework volume}^* \text{ (litres)}}{10}$$

* = including volume of water the product is diluted in for administration to incubator (Administration Volume)

1. Using an appropriate measuring device add the required volume of Pyceze to a bucket of water containing at least 10 litres ('Volume used to dilute Pyceze in above'*). This water is the Administration Volume.
2. Mix the bucket contents thoroughly and add to your Administration Vessel (e.g. header tank).
3. If the system does not usually function as a re-circulation unit, couple the re-circulation up and start running ensuring the same flow rate of water as pre-re-circulation. During the treatment period there should be no water leaving the treatment system.
4. Turn on the tap of the Administration Vessel allowing the Administration Volume to be added to the incubator over the calculated Administration Time. It is good practice to try this with no Pyceze present to ensure the Pyceze is added over the correct time.
5. Once the product has been fully added to the re-circulating water and the unit allowed to re-circulate for a further 30 minutes (treatment time) resume circulation of un- medicated water.

DOSE EXAMPLES:

Treatment regime	Incubator vessel type	Volume (litres)			Flow rate (Inflow) (litres/minute)	Admin. Time (minutes)	Pyceze dose (mL)
		Incubator	Pipework/re - circ. unit	Admin. Volume			
Re-circulation	Small incubator	10 litres x rack of 20	90	10	1.5 litres x 20	9.7	30
	Large incubator	80	90	10	16	10.6	18
	Egg trough	200 litres x 2 in	90	10	25	19.6	50

2. CONTINUOUS ADDITION: (DOSE = 1 ML PER 10 LITRES WATER = 50 MG BRONOPOL/LITRE)

Treatment is added to the system over the time it takes for the water already in the system to achieve the correct dose level. Addition of the medicated water is then continued for the treatment time of 30 minutes.

$$\text{Administration Time (min)} = \left(\frac{\text{Incubator and pipework volume (litres)}}{\text{Flow rate (litres/min)}} \right) + 30$$

$$\text{Dose of product (mL)} = \frac{[\text{Flow rate (litres/min)} \times \text{Administration Time (mins)}]}{10} + \frac{\text{Volume used to dilute the product (Litres)*}}{10}$$

* = Volume of water Pyceze is diluted in for administration to inflow of incubator (Administration Volume)

- a) Using an appropriate measuring device add the required volume of Pyceze to a bucket of water containing at least 10 litres ('Volume used to dilute Pyceze in above'*). This water is the Administration Volume.

- b) Mix the bucket contents thoroughly and add to your Administration Vessel (e.g. header tank).
- c) Turn on the tap of the Administration Vessel allowing the Administration Volume of water to be added to the incubator over the calculated Administration Time leaving the inflow water flow rate unchanged. It is good practice to try this with no Pyceze present to ensure the Pyceze (Administration Volume) is added over the correct time.
- d) The flow rate to the incubator needs to be such that a complete exchange of the incubator water is achieved in 30 minutes following the end of the treatment period.

DOSE EXAMPLES:

Treatment regime	Incubator vessel type	Volume (litres)			Flow rate (Inflow) (litres/minute)	Admin. Time (minutes)	Pyceze dose (mL)
		Incubator	Pipework	Admin. volume			
Continuous addition	Small incubator	10 litres x rack of 10	20	10	1.5 litres x 20	38.0	58
	Large incubator	70	0	10	16	34.4	56
	Egg trough	100 litres x 2 in series	10	10	12	47.5	58

It is recommended that you telephone your prescribing veterinarian to check the dose and treatment regime you plan to use BEFORE your first use of Pyceze. If a positive recovery is not seen after 14 days consult your veterinarian surgeon.

10. WITHDRAWAL PERIOD(S)

Do not use in salmonid eggs intended for human consumption. Meat – zero degree days.

11. SPECIAL STORAGE PRECAUTIONS

Store in the original container. Keep the container tightly closed. Do not store above 25 °C. Shelf-life after first opening the immediate packaging: 6 months

Once the container is opened for the first time, apply the in-use shelf-life which is specified on this package leaflet (section 11). Calculate the date by which any product remaining in the container should be used and write the corresponding discard date in the space provided.

Once diluted, the product must be used immediately.

12. SPECIAL WARNING(S)

CONTRAINDICATIONS/WARNINGS: Do not use in smolting Atlantic salmon or rainbow trout alevins as studies have indicated increased toxicity at this stage. Do not conduct bath treatments with other products at the same time as bath treatments with bronopol. Use in broodstock should only be in accordance with a benefit/risk

assessment by the responsible veterinarian because reproductive toxicity has not been established in the target species. Treatment with Pyceze is most effective if the infection is treated early and before there is a chance for the fungus to invade the underlying muscle or before there is a build-up of a dense mat of dead eggs.

Data appears to demonstrate that Pyceze, once diluted in a fixed volume of water, may be degraded by prolonged or repeated exposure to high intensity ultraviolet light. Degradation of Pyceze may alter its toxicity profile. It is therefore recommended that Pyceze treated water is not passed repeatedly through ultraviolet sterilising filters.

Accurately estimate the volume of water to be treated.

Use at water temperatures of more than 13 °C only in accordance with a benefit/risk assessment by the responsible veterinarian, as efficacy and safety at temperatures above 13 °C has not been demonstrated.

Treatment with the product is most effective if the infection is treated early and before there is a chance for the fungus to invade the underlying muscle or before there is a build up of a dense mat of dead eggs.

The product must be correctly administered to ensure efficacy

Use in broodstock should only be in accordance with a benefit/risk assessment by the responsible veterinarian because reproductive toxicity has not been established in the target species.

Safety studies on both eggs and fish conducted at five times the recommended dose and double the recommended duration (60 minutes) for up to 28 days indicate no adverse effects.

SPECIAL PRECAUTIONS TO BE TAKEN BY THE PERSON ADMINISTERING THE VETERINARY MEDICINAL PRODUCT TO ANIMALS

Irritating to eyes, lungs and skin.

Wear protective clothing, impervious gloves, face mask and suitable eye protection when mixing and handling the product. (For instance 0.3 mm nitrile rubber gloves, disposable half mask respirator conforming to European Standard EN149 together with suitable eye protection or a non-disposable respirator conforming to European Standard EN140 with a filter to EN143).

If in contact with the skin, wash immediately and – in case of contact with the eyes – rinse immediately with plenty of water and seek medical advice.

Do not smoke, drink or eat when using the product and wash hands carefully after use. For animal treatment only.

To be supplied only on veterinary prescription.

Keep out of the reach and sight of children.

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

This product is designed for use following dilution and waterborne administration. Without any dilution the product is harmful to other aquatic life.

A discharge consent may be required for release of this product into the aquatic environment. This must be obtained, prior to use of the product, from the relevant authority. Otherwise the product should only be used if the flow rate of untreated waters allows for a minimum dilution of the volume of treated water. For fish treatment the product should only be used if the flow rate of untreated water allows a dilution of 1:2000 times the volume of treated water. For egg treatment the product should only be used if the flow rate of untreated water allows a dilution of 1:5000 times the volume of treated water. Where the appropriate dilution of treated water cannot be achieved, the farm must have a discharge process to limit the release of product into the environment to within the parameters described. This can be achieved by the use of holding tanks and ponds, discharge lagoons and biofilters to clean treated water. Where this applies, the user must monitor the discharge concentration to ensure the parameters are not exceeded.

Do not contaminate surface water or ditches with product or used container.

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

14. DATE ON WHICH THE PACKAGE LEAFLET WAS LAST APPROVED

November 2023

15. OTHER INFORMATION

Pack sizes:

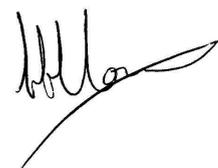
Cardboard box with 1 bottle containing 5 Litres and 1 Polypropylene graduated measuring device (500 mL).

Cardboard box with 4 bottles containing 5 Litres and 4 Polypropylene graduated measuring device (500 mL).

Bottle containing 1 Litre.

Not all pack sizes may be marketed.

For any information about this veterinary medicinal product, please contact the local representative of the marketing authorisation holder.



Approved 11 May 2024