WATER FOR INJECTIONS BP

PL 02848/0228

UKPAR

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WATER FOR INJECTIONS BP

PL 02848/0228

LAY SUMMARY

The MHRA has granted Antigen International Limited a Marketing Authorisation (licence) for the medicinal product Water for Injections BP (PL 02848/0228).

This is a prescription-only medicine (POM) mainly used to dissolve and reconstitute medicinal products acting as a carrier solution for injectable drugs.

No new or unexpected safety concerns arose from this application and it was therefore judged that the benefits of using Water for Injections BP outweigh the risks, hence a Marketing Authorisation has been granted.

WATER FOR INJECTIONS BP

PL 02848/0228

SCIENTIFIC DISCUSSION

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INTRODUCTION

Based on the review of the data on quality, safety and efficacy, the UK granted a marketing authorisation for the medicinal product Water for Injections BP to Antigen International Limited on 31st of August 2007. The product is a prescription-only medicine.

This application was submitted as an abridged application according to Article 10(1) of Directive 2001/83/EC, claiming essential similarity to the original product Water for Injections BP, PL 02848/0152 licensed to Antigen International Limited in 1991 and so the 10-year period of data exclusivity has expired.

The product contains the active ingredient water for injections and is indicated to serve as a vehicle for dilution and reconstitution of suitable medicinal products for parenteral administration.

PHARMACEUTICAL ASSESSMENT

DRUG SUBSTANCE

No materials of animal or human origin are used in the production of the active substance.

An appropriate specification based on the European Pharmacopoeia has been provided.

Analytical methods have been appropriately validated and are satisfactory for ensuring compliance with the relevant specifications.

Active water for injections is stored in appropriate packaging. The specifications and typical analytical test reports are provided and are satisfactory.

Batch analysis data are provided and comply with the proposed specification.

No stability data are provided and deemed necessary for this widely used active ingredient.

DRUG PRODUCT

Other ingredients

Water for Injections is the only ingredient in the product. No materials of animal or human origin are contained in or used in the manufacture of this product. No genetically modified organisms are included in this product.

Manufacture

A description and flow-chart of the manufacturing method have been provided.

In-process controls are appropriate considering the nature of the product and the method of manufacture. Process validation has been carried out on all batches. The results are satisfactory.

Finished product specification

The finished product specification is satisfactory. Acceptance limits have been justified with respect to conventional pharmaceutical requirements and, where appropriate, safety. Test methods have been described and have been adequately validated, as appropriate. Batch data have been provided and comply with the release specification. Certificates of analysis have been provided for any working standards used.

Container Closure System

The product is packaged in 5ml and 10ml ampoules of Polyethylene. This polyethylene does not contain any additives and is in compliance with the Ph Eur monograph . Specifications and Certificates of Analysis for packaging used have been provided. This is satisfactory. All primary product packaging complies with EU legislation regarding contact with solutions for parenteral and ophthalmic use Directive 2002/72/EC (as amended).

Stability

Finished product stability studies have been conducted in accordance with current guidelines. Based on the results, a shelf-life of 2 years when stored above 25ºC has been set, which is satisfactory.

Conclusion

It is recommended that Marketing Authorisation is granted for this application.

PRECLINICAL ASSESSMENT

This application for a generic product claims essential similarity to Water for Injections, PL 02848/0152 licensed to Antigen International Limited in 1991, which has been licensed within the EEA for over 10 years.

No new preclinical data have been supplied with this application and none is required for an application of this type.

CLINICAL ASSESSMENT

1. INTRODUCTION

This is a national standard application for Water for Injections BP, PL 02848/0228. This application has been submitted under article 10.1 first paragraph of EC Directive 2001/83, as an essentially similar product to water for Injections BP (PL 02848/0152), licensed to Antigen International Limited.

2. BACKGROUND

Water for injections is a well-known and widely used solvent. It is compatible with an extensive number of substances and can be used as a vehicle/diluent for the parenteral administration of other medicinal products.

3. INDICATIONS

Water for Injections is indicated to serve as a vehicle for dilution and reconstitution of suitable medicinal products for parenteral administration.

4. DOSE & DOSE SCHEDULE

The dosage administered will be dictated by the nature of the additive used. The administration rate will be dependent upon the dose regimen of the prescribed drug.

Administration:
For parenteral use. The directions for use related to the added medicinal product will dictate the appropriate volumes as well as the administration route.

5. TOXICOLOGY

No new toxicology data has been submitted and none are required.

6. CLINICAL PHARMACOLOGY

6.1 PHARMACOKINETICS

Water for Injections being only the vehicle for the administration of the added medicinal product, the pharmacokinetics will depend on the nature of the drug added.

6.2 PHARMACODYNAMICS

Water for Injections being only the vehicle for the administration of the added medicinal product, the pharmacodynamics will depend on the nature of the drug added.

7. EFFICACY

No formal efficacy data derived from studies of patients have been provided for this application and none are required.

8. SAFETY

No formal safety data have been provided for this application and none are required.

9. EXPERT REPORT

No new clinical data has been submitted and none are required.

10. SUMMARY OF PRODUCT CHARACTERISTICS

The SPC is fully in line with that for the reference product.

11. PATIENT INFORMATION LEAFLET

The PIL is satisfactory.

12. LABELLING

The labelling is satisfactory.

13. MARKETING AUTHORISATION FORM

This is satisfactory.

14. DISCUSSION

This is a national standard application for Water for Injections BP.

Water for Injection is well established as a vehicle/diluent for the parenteral administration of other medicinal products. It has been used in different countries for decades with a satisfactory efficacy and safety profile.

15. CONCLUSIONS

The efficacy and safety of Water for Injections BP are satisfactory for the grant of a product licence.

OVERALL CONCLUSION AND RISK BENEFIT ASSESSMENT

QUALITY

The important quality characteristics of Water for Injections BP are well defined and controlled. The specifications and batch analytical results indicate consistency from batch to batch. There are no outstanding quality issues that would have a negative impact on the benefit/risk balance.

PRECLINICAL

No new preclinical data were submitted and none are required for application of this type.

EFFICACY

No new data were submitted and none are required for application of this type.

The SPC, PIL and labelling are satisfactory and consistent with that for reference product.

RISK BENEFIT ASSESSMENT

The quality of the product is acceptable and no new preclinical or clinical safety concerns have been identified. Extensive clinical experience with Water for Injections is considered to have demonstrated the therapeutic value of the compound. The benefit/risk balance is considered to be positive.

WATER FOR INJECTIONS BP

PL 02848/0228

STEPS TAKEN FOR ASSESSMENT

1	The MHRA received the marketing authorisation application on 6th January 2004
2	Following standard checks and communication with the applicant the MHRA considered the application valid on 10th February 2004
3	Following assessment of the application the MHRA requested further information relating to the clinical dossier on 15th June 2004, and on quality dossiers on 9th July 2004 and 9th November 2006
4	The applicant responded to the MHRA's requests, providing further information on clinical dossier on 22nd August 2006 and quality dossier on 10th November 2006
5	The application was determined on 31st August 2007

SUMMARY OF PRODUCT CHARACTERISTICS

1. NAME OF THE MEDICINAL PRODUCT

Water for Injections BP.

2. QUALITATIVE AND QUANTITATIVE COMPOSITION

Each 1ml of solution contains 1ml of Water for Injections.

3. PHARMACEUTICAL FORM

Solution for Injection.
Clear, colourless, odourless, sterile solution.

4. CLINICAL PARTICULARS

4.1 Therapeutic indications

Water for Injections is indicated to serve as a vehicle for dilution and reconstitution of suitable medicinal products for parenteral administration.

4.2 Posology and method of administration

Dosage:
The dosage administered will be dictated by the nature of the additive used. The administration rate will be dependent upon the dose regimen of the prescribed drug.

Administration:
For parenteral use. The directions for use related to the added medicinal product will dictate the appropriate volumes as well as the administration route.

4.3 Contraindications

None known.
The contraindications related to the added medicinal product should be considered.

4.4 Special warnings and precautions for use

Water for Injections is hypotonic and it should not be administered alone because it may cause haemolysis.

4.5 Interaction with other medicinal products and other forms of interaction

None known.

4.6 Pregnancy and lactation

May be used during pregnancy and lactation.
The risks during use in pregnancy and in lactating women are determined by the nature of the added medicinal products.

4.7 Effects on ability to drive and use machines

None.

4.8 Undesirable effects

None known.

The nature of the additive will determine the likelihood of any undesirable effects.

4.9. Overdose

No effects anticipated with the proposed use.
Haemolysis may occur following infusion of large volumes of hypotonic solutions using sterile Water for Injections as diluent.
The signs and symptoms of overdose will also be related to the nature of the medicinal product being added. In the event of accidental overdose, the treatment should be discontinued and the patient should be observed for the appropriate signs and symptoms related to the medicinal product administered.

5. PHARMACOLOGICAL PROPERTIES

5.1. Pharmacodynamic properties

Not applicable.

Water for Injections being only the vehicle for the administration of the added medicinal product, the pharmacodynamics will depend on the nature of the drug added.

5.2. Pharmacokinetic properties

Not applicable.

Water for Injections being only the vehicle for the administration of the added medicinal product, the pharmacokinetics will depend on the nature of the drug added.

5.3. Preclinical safety data

No relevant information other than that which is included in other sections of the Summary of Product Characteristics.
Water for Injections being only the vehicle for the administration of the added medicinal product, the preclinical safety data will depend on the nature of the drug added.

6. PHARMACEUTICAL PARTICULARS

6.1. List of excipients

Not applicable.

6.2. Incompatibilities

Water for Injections BP should not be mixed with any other agents unless their compatibility has been established.

6.3. Shelf life

2 years.
Use immediately after first opening of the ampoule. Discard unused contents.

6.4. Special precautions for storage

Do not store above 25°C.

6.5. Nature and contents of container

5 ml or 10 ml hermetically sealed translucent plastic ampoules, polyethylene Ph. Eur. packed in cardboard cartons to contain 10 or100 ampoules.

6.6 SPECIAL PRECAUTIONS FOR DISPOSAL

For parenteral use.
Use as directed by a medical practitioner.
If only part of the ampoule is used, discard the remaining solution.
Keep out of the reach and sight of children.

7. MARKETING AUTHORISATION HOLDER

Antigen International Ltd
Roscrea
County Tipperary
Ireland.

8. MARKETING AUTHORISATION NUMBER

PL 02848/0228

9 DATE OF FIRST AUTHORISATION/RENEWAL OF THE AUTHORISATION

31/08/2007

10 DATE OF REVISION OF THE TEXT

31/08/2007

PATIENT INFORMATION LEAFLET 0000000-00

Water for Injection BP

Please read this leaflet carefully. It provides a summary of information about the product you may be about to use or be given by your doctor. Keep this leaflet you may need to read it again.

1. WHAT IS WATER FOR INJECTION BP?

Water for Injection BP is sterile (clean) water and is presented in either 5ml or 10ml polyethylene ampoules (Mini- Plasco®), in packs of 10 or 100 ampoules.

2. WHAT IS WATER FOR INJECTION BP USED FOR?

Water for Injection BP is mainly used to dissolve and reconstitute medicinal products acting as a carrier solution for injectable drugs.

3. BEFORE YOU USE WATER FOR INJECTION BP

Check the solution in the ampoule is clear and colourless. Do not use the solution if it is not clear or colourless.

Check the ampoule is intact. Do not use the solution if the ampoule is not intact.

4. HOW TO USE WATER FOR INJECTION BP

Water for Injection BP is for parenteral use (delivery of a drug into the body without using the oral tract). It is used to reconstitute or dissolve a drug, before it is injected into the body.

Water for Injections can be used during pregnancy or if you are breastfeeding. You should discuss it with your doctor if you are pregnant, become pregnant or are breastfeeding as its use will depend on the medicine given with Water for Injection BP.

Always use Water for Injection BP exactly as your doctor has told you.

- If you miss a dose: follow the advice given to you by your doctor.
- If you take too much Water for Injection BP see your pharmacist or doctor for advice.

5. SIDE EFFECTS

Along with desirable effects, a medicine may cause unwanted effects. It is very rare to experience side effects from Water for Injection BP. If you experience any side effects inform your doctor or pharmacist.

The main cause of side effects would be the medication prescribed by your doctor for use with Water for Injection BP. Please read the information leaflet for the medicine prescribed to you by your doctor to check for possible side effects.

6. STORING WATER FOR INJECTION BP

Keep Water for Injection BP and all medicines out of the reach and sight of children.

Do not store above 25°C. Do not use Water for Injection BP after the expiry date shown on the label and carton.

- Use immediately after opening the ampoule.
- Discard any unused contents.

7. FURTHER INFORMATION

This leaflet does not contain all the information about your medicine. If you have any questions or are not sure about anything, ask your doctor or pharmacist.

The Marketing Authorisation holder is Antigen International Limited, Roscrea, Co. Tipperary, Ireland.

PL No. : 02848/0228

The company responsible for manufacture is Aguettant, Parc Scientifique Tony Garnier-Rhône, 1 rue Alexander Fleming, BP1744, 69353 Lyon Cedex 07, France.

Antigen Pharmaceuticals

Antigen International Ltd.
Roscrea, Co. Tipperary, Ireland.

Date of last revision : May, 2007.

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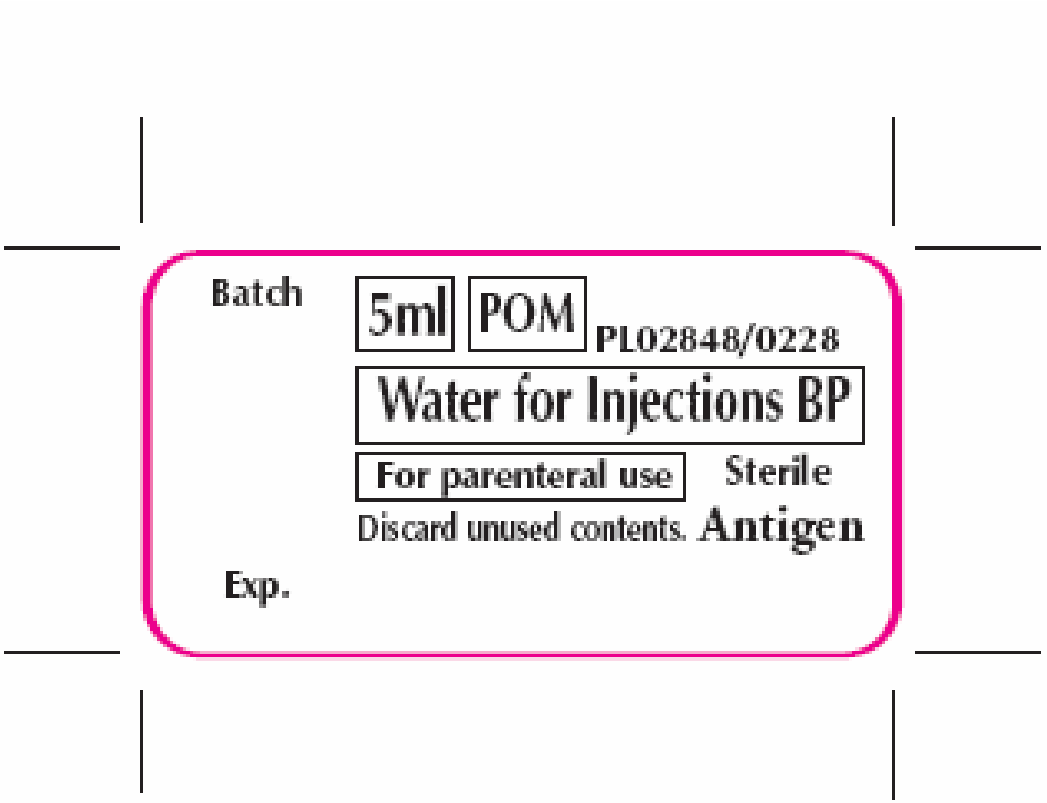
IWA015410ELF/0406/A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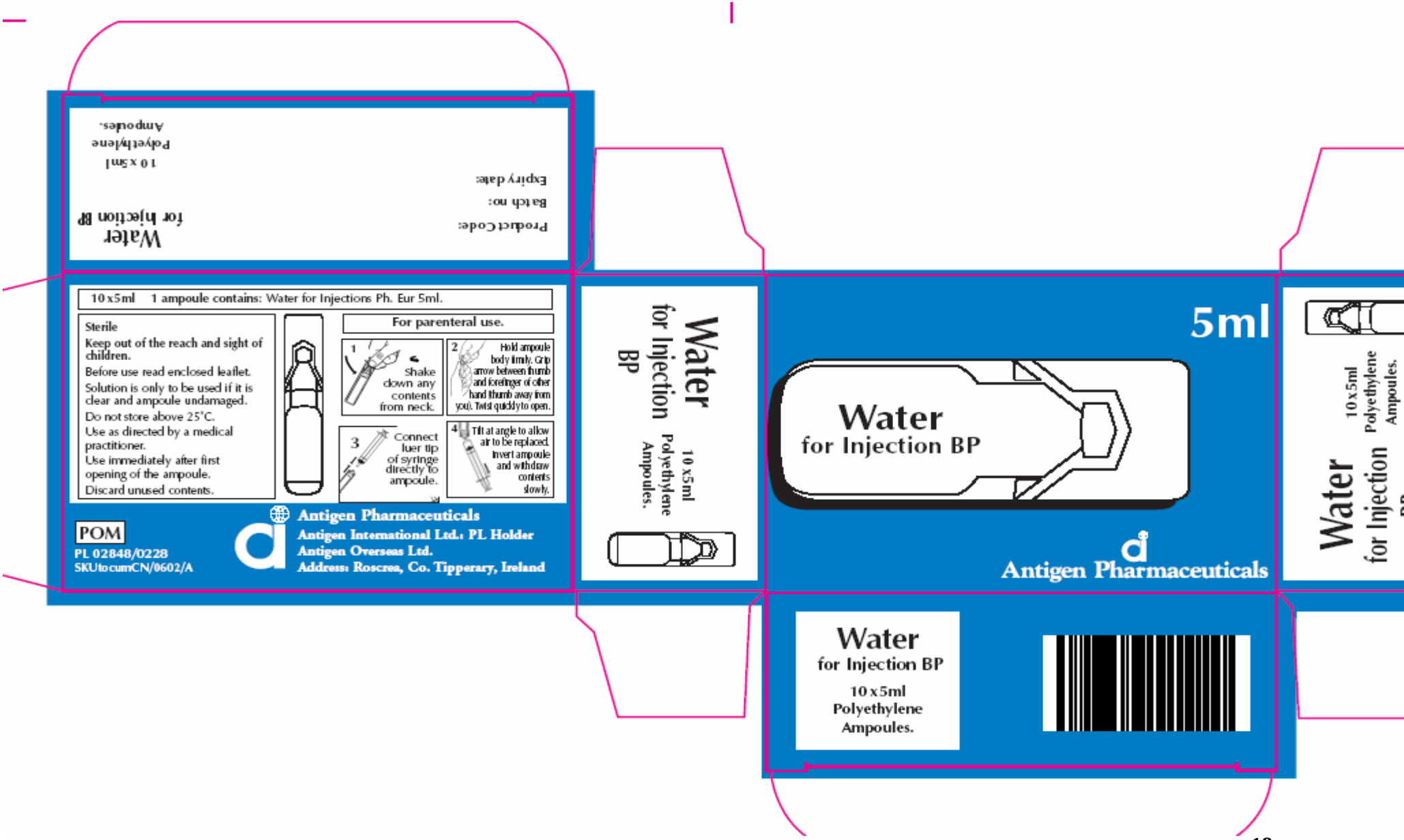
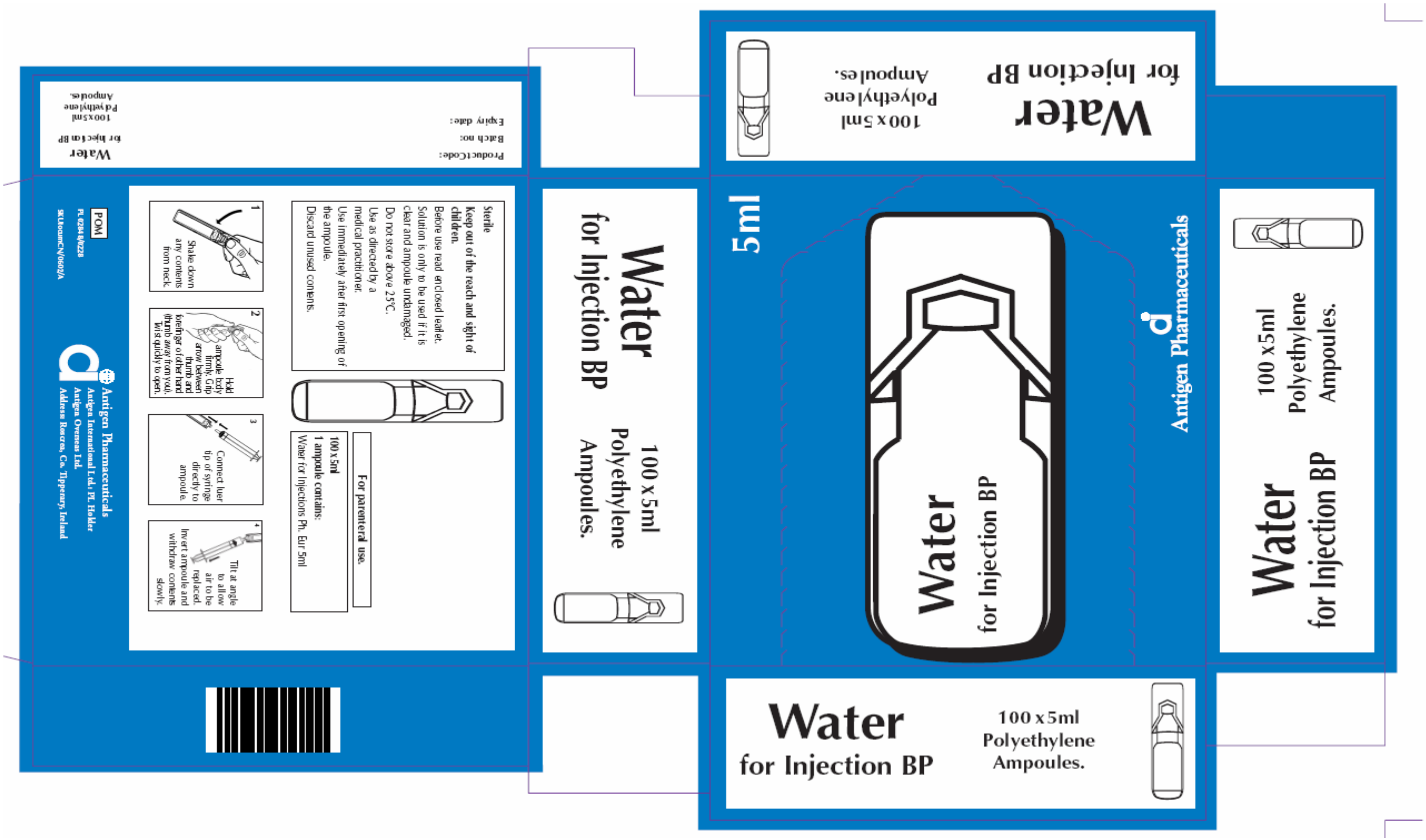
5ml
POM
PL02848/0228
Water for Injections BP
For parenteral use
Sterile
Discard unused contents.
Antigen
Exp.


Water
for Injection BP
Expiry date:
Batch no:
ProductCode:
10 x 5ml
Polyethylene
Ampoules.
10 x5ml 1 ampoule contains: Water for Injections Ph. Eur 5ml.
Sterile
Keep out of the reach and sight of children.
Before use read enclosed leaflet.
Solution is only to be used if it is clear and ampoule undamaged.
Do not store above 25°C.
Use as directed by a medical practitioner.
Use immediately after first opening of the ampoule.
Discard unused contents.
For parenteral use.
1 Shake down any contents from neck.
2 Hold ampoule body firmly. Grip arrow between thumb and forefinger of other hand thumb away from you). Twist quickly to open.
3 Connect luer tip of syringe directly to ampoule.
4 Tilt at angle to allow air to be replaced. Invert ampoule and withdraw contents slowly.
POM
PL 02848/0228
SKUtocumCN/0602/A
Antigen Pharmaceuticals
Antigen International Ltd.: PL Holder
Antigen Overseas Ltd.
Address: Roscrea, Co. Tipperary, Ireland
Water
for Injection
BP
10 x5ml
Polyethylene
Ampoules.
5ml
Water
for Injection BP
Antigen Pharmaceuticals
Water
for Injection BP
10 x5ml
Polyethylene
Ampoules.
Water
for Injection
10 x5ml
Polyethylene
Ampoules.


Water
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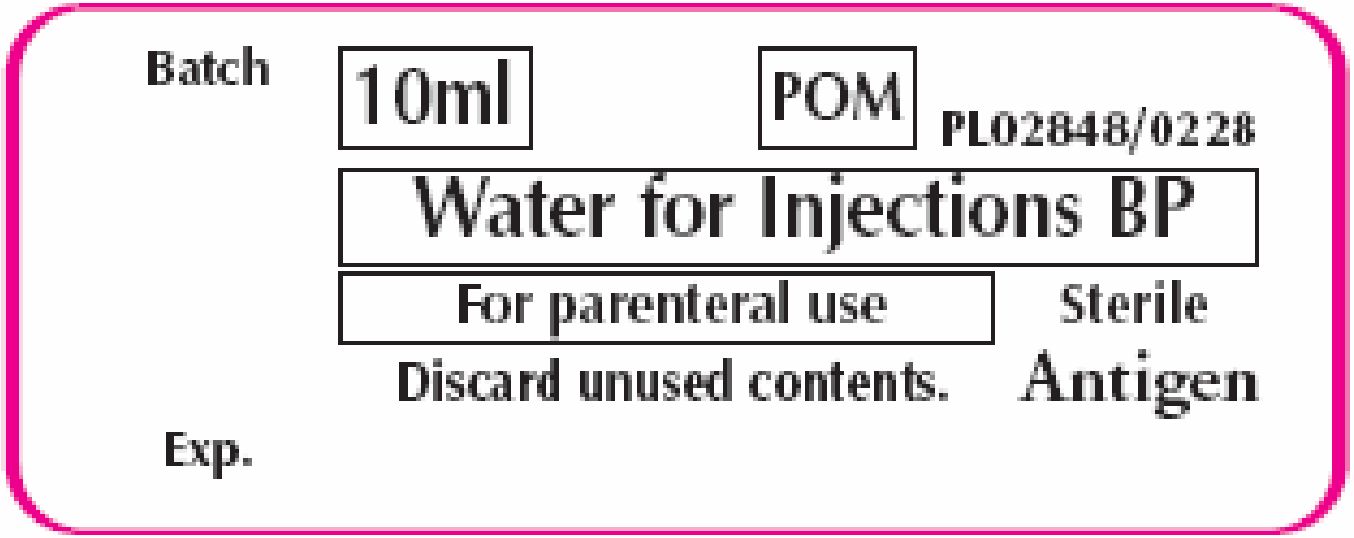
Antigen Pharmaceuticals
Antigen International Ltd.: PL Holder
Antigen Overseas Ltd.
Address Roscrea, Co. Tipperary, Ireland
5ml
Antigen Pharmaceuticals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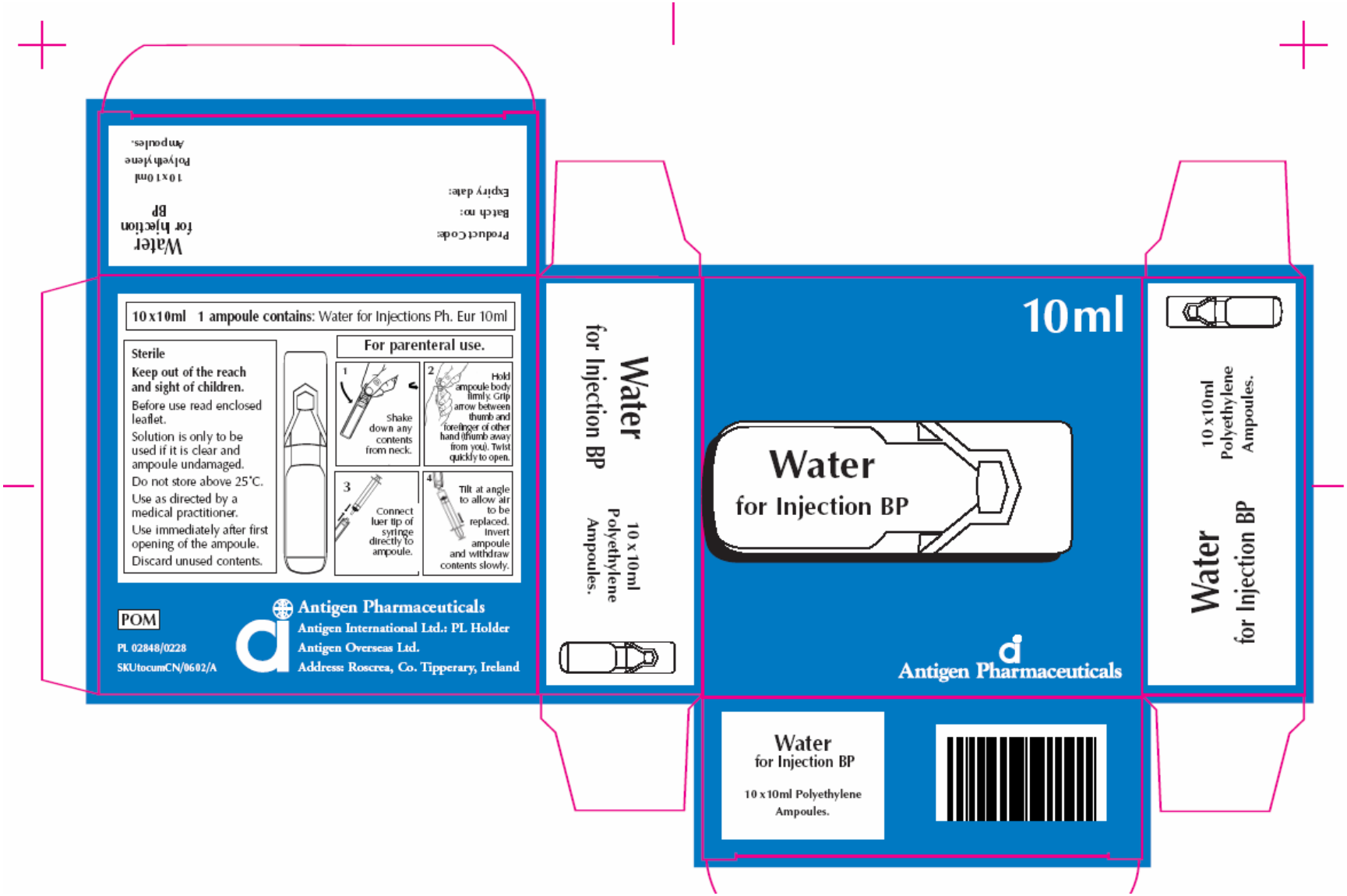
10ml
POM
PL02848/0228
Water for Injections BP
For parenteral use
Sterile
Discard unused contents.
Antigen
Exp.


Water
for Injection
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Polyethylene
Ampoules.
Product Code:
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Expiry date:
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Water
for Injection BP
10 x 10ml
Polyethylene
Ampoules.
10ml
Water
for Injection BP
Antigen Pharmaceuticals
Water
for Injection BP
10 x 10ml Polyethylene
Ampoules.
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Polyethylene
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Water
for Injection BP

Water
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Ampoules.
Antigen Pharmaceuticals
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Discard unused contents.
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100 x 10ml
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Water for Injections Ph. Eur 10ml
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Shake down any contents from neck.
2
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3
Connect luer tip of syringe directly to ampoule.
4
Tilt at angle to allow air to be replaced. Invert ampoule and withdraw contents slowly.
Product Code:
Batch no:
Expiry date:
Water
for Injection BP
100 x 10ml
Polyethylene
Ampoules.
POM
PL 02848/0228
SKUtxxmmCVI0603/A
Antigen Pharmaceuticals
Antigen International Ltd.: PL Holder
Antigen Overseas Ltd.
Address Rosrea, Co. Tipperary, Ireland