MACROGOL COMPOUND ORAL POWDER
(sodium chloride, sodium hydrogen carbonate, potassium chloride, macrogol 3350)

PL 33579/0001

UKPAR

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LAY SUMMARY

Macrogol Compound Oral Powder
(sodium chloride, sodium hydrogen carbonate, potassium chloride, macrogol 3350)

This is a summary of the public assessment report (PAR) for Macrogol Compound Oral Powder (PL 33579/0001). It explains how Macrogol Compound Oral Powder was assessed and its authorisation recommended, as well as the conditions of use. It is not intended to provide practical advice on how to use Macrogol Compound Oral Powder.

For practical information about using Macrogol Compound Oral Powder, patients should read the package leaflet or contact their doctor or pharmacist.

What is Macrogol Compound Oral Powder and what is it used for?

Macrogol Compound Oral Powder contains the active substances sodium chloride, sodium hydrogen carbonate, potassium chloride and macrogol 3350. Macrogol Compound Oral Powder is a laxative prescribed for the treatment of long-term constipation.

This medicine is identical to Macroherm Powder for Oral Solution (PL 17740/0008), which was authorised in the UK to Hermes Arzneimittel GmbH on 12 October 2010. Hermes Arzneimittel GmbH has agreed that scientific data presented for Macroherm Powder for Oral Solution (PL 17740/0008) can be used for this application for Macrogol Compound Oral Powder.

How is Macrogol Compound Oral Powder used?

Macrogol Compound Oral Powder is for oral use. This medicine can be obtained without a prescription from a pharmacy. The recommended dose for constipation in adults, children (aged 12 years and above) and the elderly is one sachet taken one to three times daily. The contents of each sachet must be poured into a glass and dissolved in about 125 ml or a quarter pint of water. As for all laxatives, prolonged use is not usually recommended.

How does Macrogol Compound Oral Powder work?

Macrogol Compound Oral Powder contains the active ingredients sodium chloride, sodium hydrogen carbonate, potassium chloride and macrogol 3350. Macrogol Compound Oral Powder is a laxative used for the treatment of long-term constipation. It works by making the faeces softer and easier to pass, therefore, providing relief from constipation.

How has Macrogol Compound Oral Powder been studied?

This application is identical to the previously granted application for Macroherm Powder for Oral Solution (PL 17740/0008; Hermes Arzneimittel GmbH). The applicant (Neopharma Limited) referred to data provided by Hermes Arzneimittel GmbH for the grant of Macroherm Powder for Oral Solution (PL 17740/0008) as the

basis for the grant of an identical licence for Macrogol Compound Oral Powder (PL 33579/0001).

What are the benefits and risks of Macrogol Compound Oral Powder?

Macrogol Compound Oral Powder is considered identical to Macroherm Powder for Oral Solution, with benefits and risks taken as being the same as those for Macroherm Powder for Oral Solution.

Why is Macrogol Compound Oral Powder approved?

No new or unexpected safety concerns arose from this application. It was, therefore, considered that the benefits of Macrogol Compound Oral Powder outweigh their risks; and the grant of a Marketing Authorisation was recommended.

What measures are being taken to ensure the safe and effective use of Macrogol Compound Oral Powder?

A risk management plan has been developed to ensure that Macrogol Compound Oral Powder is used as safely as possible. Based on this plan, safety information has been included in the Summary of Product Characteristics and the package leaflet for Macrogol Compound Oral Powder, including the appropriate precautions to be followed by healthcare professionals and patients.

Other information about Macrogol Compound Oral Powder

A Marketing Authorisation was granted in the UK on 09 May 2014. For more information about treatment with Macrogol Compound Oral Powder, read the package leaflet, or contact your doctor or pharmacist.

This summary was last updated in July 2014.

The full PAR for Macrogol Compound Oral Powder follows this summary.

MACROGOL COMPOUND ORAL POWDER (PL 33579/0001)

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 Macrogol Compound Oral Powder (PL 33579/0001) on 09 May 2014 to Neopharma Limited.

This application for Macrogol Compound Oral Powder was submitted as an abridged simple application according to Article 10c of Directive 2001/83/EC, as amended, cross-referring to Macroherm Powder for Oral Solution, which was granted a licence to Hermes Arzneimittel GmbH on 12 October 2010 (PL 17740/0008).

This medicine can be obtained without a prescription from a pharmacy (legal status P) and is indicated for the treatment of chronic constipation.

The product contains the active ingredients sodium chloride, sodium hydrogen carbonate, potassium chloride and Macrogol 3350. Macrogol Compound Oral Powder is an osmotically acting laxative. Macrogol 3350 induces a laxative effect through its osmotic action in the gut. Macrogol 3350 increases the stool volume, which in turn causes colon motility via neuromuscular pathways. The physiological consequence is an improved propulsive colonic transportation of the softened stools and a facilitation of the defaecation. Electrolytes combined with macrogol 3350 are exchanged across the intestinal barrier (mucosa) with serum electrolytes, and excreted in faecal water without net gain or loss of sodium, potassium and water.

PHARMACEUTICAL ASSESSMENT

LICENCE NO: PL 33579/0001
PROPRIETARY NAME: Macrogol Compound Oral Powder
ACTIVE(S): sodium chloride, sodium hydrogen carbonate, potassium chloride, macrogol 3350
COMPANY NAME: Neopharma Limited
E.C. ARTICLE: Article 10c of Directive 2001/83/EC, as amended
LEGAL STATUS: P

1. INTRODUCTION

This is a simple, piggyback application for Macrogol Compound Oral Powder submitted under Article 10c of Directive 2001/83/EC, as amended. The proposed Marketing Authorisation Holder is Neopharma Limited, 57 High Street, Odiham, Hampshire, RG29 1LF.

This application cross-refers to Macroherm Powder for Oral Solution (PL 17740/0008), which was granted a licence to Hermes Arzneimittel GmbH on 12 October 2010.

The current application is considered valid.

2. MARKETING AUTHORISATION APPLICATION FORM

2.1 NAME(S)

The proposed name of the product is Macrogol Compound Oral Powder. The product has been named in line with current requirements.

2.2 Strength, pharmaceutical form, route of administration, container and pack sizes

Each sachet contains the following quantitative composition of active ingredients:

Sodium chloride	0.3507g
Sodium hydrogen carbonate	0.1785g
Potassium chloride	0.0466g
Macrogol 3350	13.125g

The content of electrolyte ions per sachet following reconstitution in 125 ml of water is equivalent to:

Sodium	65 mmol/l
Chloride	53 mmol/l
Hydrogen carbonate (bicarbonate)	17 mmol/l
Potassium	5 mmol/l

Each sachet is composed of paper, ethylene-methacrylic acid co-polymer and aluminium. The sachets are packed in cartons of 2, 6, 8, 10, 20, 30, 50, 60 (2x30) and 100 (2x50).

No mock-ups have been assessed as part of this Marketing Authorisation. Therefore the Marketing Authorisation Holder has committed to providing the MHRA with the mock-ups for any pack size before placing the product on the market.

The proposed shelf-life for the sachet (36 months) and the reconstituted solution (24 hours), and the storage conditions for the sachet (Do not store above 25 °C) and the reconstituted solution (Store covered in a refrigerator [2°C to 8°C]) are consistent with the details registered for the cross-reference product.

2.3 Legal status

On approval, the product will be available without prescription from a pharmacy (legal status P).

2.4 Marketing authorisation holder/Contact Persons/Company

Neopharma Limited, 57 High Street, Odiham, Hampshire, RG29 1LF.

The QP responsible for pharmacovigilance is stated and his CV is included.

2.5 Manufacturers

The proposed manufacturing sites are consistent with those registered for the cross-reference product and evidence of Good Manufacturing Practice (GMP) compliance has been provided.

2.6 Qualitative and quantitative composition

The proposed composition is consistent with the details registered for the cross-reference product.

2.7 Manufacturing process

The proposed manufacturing process is consistent with the details registered for the cross-reference product and the maximum batch sizes are stated.

2.8 Finished product/shelf-life specification

The proposed finished product specification is in line with the details registered for the cross-reference product.

2.9 Drug substance specification

The proposed drug substance specification is consistent with the details registered for the cross-reference product.

2.10 TSE Compliance

No materials of animal or human origin are included in this product. This is consistent with the cross-reference product.

3. EXPERT REPORTS

The applicant has included detailed expert reports in Module 2 of the application. Signed declarations and copies of the experts' CVs are enclosed in Module 1.4 for the quality, non-clinical and clinical experts. All are considered to have sufficient experience for their responsibilities.

4. PRODUCT NAME & APPEARANCE

See 2.1 for details of the proposed product names. The appearance of the product is identical to the cross-reference product.

5. SUMMARY OF PRODUCT CHARACTERISTICS (SmPC)

The proposed SmPC is consistent with the details registered for the cross-reference product.

6. PATIENT INFORMATION LEAFLET (PIL)/CARTON

PIL

The patient information leaflet has been prepared in line with the details registered for the cross-reference product.

Carton and blister text

The proposed text is comparable to the artwork registered for the cross-reference product and complies with statutory requirements. In line with current legislation, the applicant has also specified that the name of the product in Braille will be included on the outer packaging.

7. CONCLUSIONS

The data submitted with this application are acceptable. From a quality perspective, a Marketing Authorisation should be granted.

NON-CLINICAL ASSESSMENT

No new non-clinical data have been supplied with this application and none are required for applications of this type.

CLINICAL ASSESSMENT

No new clinical data have been supplied with this application and none are required for applications of this type.

OVERALL CONCLUSION AND BENEFIT/RISK ASSESSMENT

QUALITY

The important quality characteristics of Macrogol Compound Oral Powder are identical to those of the already granted reference product. There are no outstanding quality issues that would have a negative impact on the benefit/risk balance.

NON-CLINICAL

No new non-clinical data were submitted and none are required for applications of this type.

CLINICAL PHARMACOLOGY/EFFICACY

No new clinical pharmacology/efficacy data have been submitted with this application and none are required for applications of this type.

SAFETY

No new safety data have been submitted with this application and none are required for applications of this type.

No new or unexpected safety concerns arise from this application.

The SmPC, PIL and labelling are satisfactory.

Pharmacovigilance System and Risk Management Plan

The Pharmacovigilance System, as described by the applicant, fulfils the requirements and provides adequate evidence that the applicant has the services of a qualified person responsible for pharmacovigilance, and has the necessary means for the notification of any adverse reaction suspected of occurring either in the Community or in a third country.

A risk management plan has been developed to ensure that Macrogol Compound Oral Powder is used as safely as possible. Based on this plan, safety information has been included in the Summary of Product Characteristics and the package leaflet for Macrogol Compound Oral Powder, including the appropriate precautions to be followed by healthcare professionals and patients.

BENEFIT-RISK ASSESSMENT

The quality of the product is acceptable, and no new non-clinical or clinical safety concerns have been identified. Extensive clinical experience with Macrogol Compound Oral Powder is considered to have demonstrated the therapeutic value of the product. The benefit/risk is, therefore, considered to be positive.

MACROGOL COMPOUND ORAL POWDER (PL 33579/0001)

STEPS TAKEN FOR ASSESMENT

1	The MHRA received the marketing authorisation application on 20 December 2012
2	Following standard checks and communication with the applicant, the MHRA considered the application valid on 23 May 2013
3	Following assessment of the application, the MHRA requested further information relating to the dossiers on 15 August 2013, 05 November 2013 and 13 January 2014
4	The applicant responded to the MHRA's requests, providing further information on 07 October 2013, 19 November 2013 and 24 January 2014
5	The application was determined on 09 May 2014

MACROGOL COMPOUND ORAL POWDER
(PL 33579/0001)
STEPS TAKEN AFTER AUTHORISATION - SUMMARY

Date submitted	Application type	Scope	Outcome

Summary of Product Characteristics and Patient Information Leaflet

The current approved UK versions of the Summary of Product Characteristics (SmPC) and Patient Information Leaflet (PILs) for these products are available on the MHRA website.

Labelling

PARTICULARS TO APPEAR ON THE OUTER PACKAGING

CARTON

1. NAME OF THE MEDICINAL PRODUCT

Macrogol Compound Oral Powder

2. STATEMENT OF ACTIVE SUBSTANCE(S)

Each sachet contains:

Macrogol 3350 (Polyethylene glycol 3350)	13.125 g
Sodium chloride	0.3507 g
Sodium hydrogen carbonate	0.1785 g
Potassium chloride	0.0466 g

On reconstitution in 125 ml of water, each sachet provides:

Sodium	65 mmol/l
Chloride	53 mmol/l
Hydrogen carbonate	17 mmol/l
Potassium	5 mmol/l

3. LIST OF EXCIPIENTS

Contains sorbitol (E420).
See leaflet for further information.

4. PHARMACEUTICAL FORM AND CONTENTS

Powder for oral solution

2 sachets
6 sachets
8 sachets
10 sachets
20 sachets
30 sachets
50 sachets
60 (2x30) sachets
100 (2x50) sachets

5. METHOD AND ROUTE(S) OF ADMINISTRATION

Oral use

Dissolve the contents of one sachet in 125 ml of water then drink it.

Read the package leaflet before use.

6. SPECIAL WARNING THAT THE MEDICINAL PRODUCT MUST BE STORED OUT OF THE REACH AND SIGHT OF CHILDREN

Keep out of the sight and reach of children.

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP

9. SPECIAL STORAGE CONDITIONS

Do not store above 25 °C.

Reconstituted solution can be covered and stored in the refrigerator (2°C to 8°C), and should be used within 24 hours.

10. SPECIAL PRECAUTIONS FOR DISPOSAL OF UNUSED MEDICINAL PRODUCTS OR WASTE MATERIALS DERIVED FROM SUCH MEDICINAL PRODUCTS, IF APPROPRIATE

After 24 hours, any unused solution should be discarded.

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Neopharma Limited
57 High Street
Odiham
Hampshire
RG29 1LF

12. MARKETING AUTHORISATION NUMBER(S)

PL 33579/0001

13. BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

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