European Medicines Agency
*Evaluation of Medicines for Human Use*

London, 30th July 2009 corr 1
EMEA/CHMP/473955/2009

**COMMITTEE FOR MEDICINAL PRODUCTS FOR HUMAN USE**
**JULY 2009 PLENARY MEETING**
**MONTHLY REPORT**

The Committee for Medicinal Products for Human Use (CHMP) held its July plenary meeting on 20-23 July 2009.

## CENTRALISED PROCEDURE

### Initial applications for marketing authorisation

The CHMP adopted ten positive opinions by consensus on initial marketing authorisation applications.

*New medicinal products*

- **Arcalyst** (rilonacept), from Regeneron UK Limited, indicated for the treatment of adults and children aged 12 years and older with severe symptoms of Cryopyrin-Associated Periodic Syndromes (CAPS), including Familial Cold Autoinflammatory Syndrome (FCAS) and Muckle-Wells Syndrome (MWS). CAPS is a rare, inherited condition causing recurrent fever, rash and joint pain. Arcalyst is the **57th orphan medicine** to receive a positive opinion from the CHMP. The review began on 23 July 2008, with an active review time of 197 days**.**
- **Exforge HCT** (amlodipine besylate/valsartan/hydrochlorothiazide), **Copalia HCT** (amlodipine besylate/valsartan/hydrochlorothiazide), **Imprida HCT** (amlodipine besylate/valsartan /hydrochlorothiazide) and **Dafiro HCT** (amlodipine besylate/valsartan/hydrochlorothiazide), all from Novartis Europharm Limited, indicated for the treatment of essential hypertension, as substitution therapy in adult patients whose blood pressure is adequately controlled on the combination of amlodipine, valsartan and hydrochlorothiazide. The review for Exforge HCT began on 24 September 2008, with an active review time of 206 days. The review for the three other products began on 29 March 2009, with an active review time of 89 days and was aligned with that of Exforge HCT**.**
- **Ilaris** (canakinumab), from Novartis Europharm Ltd., indicated for the treatment of adults, adolescents and children aged 4 years and older with Cryopyrin-Associated Periodic Syndromes (CAPS), including Muckle-Wells Syndrome (MWS), Neonatal-Onset Multisystem Inflammatory Disease (NOMID) / Chronic Infantile Neurological, Cutaneous, Articular Syndrome (CINCA), and severe forms of Familial Cold Autoinflammatory Syndrome (FCAS). CAPS is a rare, inherited condition causing recurrent fever, rash and joint pain. Ilaris is the **58th orphan medicine** to receive a positive opinion from the CHMP. The review began on 24 December 2008, with an active review time of 176 days.
- **Ratioepo (**epoetin theta), **Eporatio** (epoetin theta), both from Ratiopharm GmbH, and **Biopoin** (epoetin theta), from CT Arzneimittel GmbH, indicated for the treatment of symptomatic anaemia associated with chronic renal failure in adult patients and for the treatment of symptomatic anaemia in adult cancer patients with non-myeloid malignancies receiving chemotherapy. Reviews began on 25 June 2008, with an active review time of 205 days**.**
- **Resolor** (prucalopride), from Movetis NV, indicated for the symptomatic treatment of chronic constipation in women in whom laxatives fail to provide adequate relief. The review began on 28 May 2008, with an active review time of 206 days**.**

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*Positive opinions for 'informed consent' duplicate applications*

The Committee adopted three positive opinions by consensus for duplicate applications for pandemic influenza mock-up vaccines.

- **Pandemic Influenza Vaccine H5N1 Baxter AG** (whole virion, Vero cell derived, inactivated), from Baxter AG, a duplicate of **Celvapan**, from Baxter AG.
- **Foclivia** (H5N1 virus surface antigens), from Novartis Vaccines and Diagnostics S.r.L., a duplicate of **Focetria**, from Novartis Vaccines and Diagnostics S.r.L.
- **Pandemic influenza vaccine (H5N1) (split virion, inactivated, adjuvanted) GlaxoSmithKline Biologicals** (H5N1 split virion, inactivated, adjuvanted with AS03), from GSK Biologicals, a duplicate of **Pandemrix**, from GSK Biologicals.

A separate press release and a question-and-answer document with an update on the Agency's pandemic vaccines activities are available.

The Committee also adopted a positive opinion by consensus for **Alendronate sodium and colecalciferol MSD**, from Merck Sharp & Dohme Ltd, a duplicate application to Fosavance, which is already authorised in the EU for the treatment of postmenopausal osteoporosis in patients at risk of vitamin D insufficiency.

When submitting 'informed consent' duplicate applications, applicants can make reference to a medicine that is already authorised in the EU, provided they have obtained consent from the marketing authorisation holder of the reference medicine to do so.

*Generic medicinal products*

The Committee adopted five positive opinions by consensus for the following generic medicines, for which a reference medicine is already authorised in the European Union. The medicines concerned are:

- **Enyglid (**repaglinide), and **Repaglinide Krka** (repaglinide), both from Krka, d.d., Novo mesto, generics of Novonorm, intended to treat patients with type-2 diabetes.
- **Irbesartan Teva (**irbesartan), from Teva Pharma B.V., a generic of Aprovel, intended for the treatment of essential hypertension.
- **Lamivudine Teva** (lamivudine), from Teva Pharma B.V., a generic of Zeffix, intended for the treatment of chronic hepatitis B in adults.
- **Clopidogrel MYLAN Pharma** (clopidogrel, as besilate), from Mylan S.A.S., a generic of Plavix, intended for the prevention of atherothrombotic events in patients suffering from myocardial infarction or established peripheral arterial disease.

Summaries of opinion for these medicinal products are available here. Further information will be included in the European Public Assessment Reports (EPARs) once the European Commission has granted final approval.

*Negative opinions for new medicines*

The Committee adopted three negative opinions by consensus, recommending that marketing authorisations for the following two medicines be refused:

- **Gemesis** (bercaplermin), from Biomimetic Therapeutics Ltd, intended for the treatment of periodontally related defects.
- **Milnacipran Pierre Fabre Médicament** (milnacipran) and **Impulsor** (milnacipran), both from Pierre Fabre Médicament, intended for the treatment of fibromyalgia syndrome.

Question and-answer documents with more information about the reasons for the refusals can be found here.

## Post-authorisation procedures

Extensions of indication and other recommendations

The Committee gave four positive opinions by consensus and one by majority (Abilify) for applications for the extension of indication, adding new treatment options, for the following medicines:

- **Isentress** (raltegravir), from Merck Sharp & Dohme Ltd, to extend the indication to include antiretroviral-treatment naïve adult patients. Isentress is currently authorised as combination-therapy with other anti-retroviral medicines for the treatment of human immunodeficiency virus (HIV-1) infection in treatment-experienced adult patients.
- **Keppra** (levetiracetam), from UCB S.A., to extend the age range for use of the medicine to children and infants from 1 month of age in the indication of adjunctive treatment of partial seizures (fits) with or without secondary generalisation. Keppra is currently authorised in this indication in patients from 4 years of age. It is also currently authorised as monotherapy to treat partial onset seizures with or without secondary generalisation in patients from 16 years of age with newly diagnosed epilepsy. Keppra is also authorised as combination therapy to treat myoclonic seizures in patients from 12 years of age with juvenile myoclonic epilepsy and primary generalised tonic-clonic seizures in patients from 12 years of age with idiopathic generalised epilepsy.
- **Mabthera** (rituximab), from Roche Registration Ltd, to extend the indication to include the treatment of relapsed/refractory chronic lymphocytic leukaemia. Mabthera is currently authorised for the treatment of Non-Hodgkin's Lymphoma, first-line treatment of patients with chronic lymphocytic leukaemia (CLL) and second-line treatment of adult patients with severe active rheumatoid arthritis.
- **Torisel** (temsirolismus), from Wyeth Europa Ltd., to extend the indication to include the treatment of adult patients with relapsed and/or refractory mantle cell lymphoma. Torisel is currently authorised for the first-line treatment of patients with advanced renal cell carcinoma.
- **Abilify** (aripiprazole), from Otsuka Pharmaceutical Europe Ltd., to extend the indication to include treatment of schizophrenia in adolescents 15 years and older. Abilify is currently authorised for the treatment of schizophrenia and for the treatment and prevention of moderate to severe manic episodes in Bipolar I Disorder.

Summaries of opinion for these extensions of indication are available here. Further information will be included in the EPARs once the European Commission has granted final approval.

Negative opinion for extension of indication

The Committee adopted a negative opinion by majority, recommending the refusal of an extension of indication for **Erbitux** (cetuximab), from Merck KGaA. The indication applied for was first-line treatment of patients with epidermal growth factor receptor (EGFR)-expressing advanced or metastatic non-small cell lung cancer in combination with platinum-based chemotherapy. Erbitux is currently authorised for single and combination treatment of patients with EGFR-expressing metastatic colorectal cancer and as combination therapy for the treatment of patients with squamous cell cancer of the head and neck. A question and answer document with more information about the reasons for the refusal is available.

Re-examination procedure under Article 6(9) of Regulation (EC) No. 1085/2003 concluded

Following re-examination of its negative opinion adopted in April 2009, the CHMP adopted a final negative opinion by consensus, for **Lyrica** (pregabalin) from Pfizer Ltd, for an extension of the indication to the treatment of fibromyalgia in adults experiencing moderate to severe pain. Lyrica is currently authorised for the treatment of neuropathic pain, epilepsy and generalised anxiety disorder in adults. A question and answer document with more information about the re-examination is available.

Withdrawal

The European Medicines Agency has been formally notified by Gendux Molecular Limited of its decision to withdraw its application for a centralised marketing authorisation for the medicine **Contusugene ladenovec Gendux** (contusugene ladenovec), suspension for injection. Contusugene ladenovec Gendux was expected to be used for the treatment of squamous cell carcinoma in head and neck cancer. A question-and-answer document with more information is available.

Other information

The CHMP adopted an amendment to sections 4.4. and 4.8 of the Summary of Product Characteristics (SPC) of **Sutent** (sunitinib) from Pfizer Limited, regarding the events "hypersensitivity/angioedema" and "fistula" reported during post marketing following a cumulative safety review conducted by the MAH. In almost all the cases of Adverse Drug Reactions (ADRs) for angioedema and fistula, the treatment with Sutent was completely interrupted in order to recover from the symptoms. Therefore, information regarding the management of these ADRs has been added to SPC in the above mentioned sections. The Package Leaflet has been updated accordingly.

Following a safety review of serious unlabelled ADRs of **Luminity** from Bristol-Myers Squibb Pharma EEIG, the CHMP requested the MAH to update sections 4.2 and 4.4 of the SPC. Consequently, the MAH submitted a variation to update the above mentioned sections to include additional safety information on cardiopulmonary reactions. Section 4.8 was also updated to add ventricular arrhythmias (primary ventricular tachycardia and premature ventricular contractions but also ventricular fibrillation asystole and severe respiratory distress). The Package Leaflet has been updated accordingly.

Update on safety of insulin glargine

Following review of all available information on a possible relationship between insulin analogues, in particular insulin glargine, and the risk of cancer, the Committee concluded that the available data does not provide a cause for concern and that changes to the prescribing advice are therefore not necessary. Insulin glargine is a long-acting insulin analogue, authorised in the European Union (EU) as Lantus and Optisulin, for the treatment of adults, adolescents and children aged six years or above with diabetes, when treatment with insulin is required. More information is available in a separate press release.

Opinion on paediatric extension for Arimidex adopted

The CHMP adopted an opinion for **Arimidex** (anastrozole), from AstraZeneca AB, to reflect new clinical data on the use of anastrozole in the treatment of short stature in pubertal boys with growth hormone deficiency in association with exogenous growth hormone, and in the treatment of testotoxicosis. These data were generated in accordance with an agreed paediatric investigation plan (PIP).
AstraZeneca AB had applied for an extension of indication to include treatment of short stature in pubertal boys with growth hormone deficiency in association with exogenous growth hormone. The Committee's opinion was that the data do not support the authorisation of the medicine for children. However the information on the paediatric studies will be included in the product information. This opinion will be transformed into a decision by the European Commission, after which the Member States must implement the changes adopted.
The opinion was adopted in accordance with Article 29 of the Paediatric Regulation (1901/2006).

## OTHER INFORMATION ON THE CENTRALISED PROCEDURE

### Lists of Questions

The Committee adopted eight Lists of Questions on initial applications (including two under the mandatory scope and six under the optional scope).

## Detailed information on the centralised procedure

An overview of centralised procedures since 1995 is given in **Annex 1**. The post-authorisation centralised procedures finalised during this meeting are summarised in **Annex 2**. The list of medicinal products for which marketing authorisations have been granted by the European Commission since the CHMP plenary meeting in June 2009 is provided in **Annex 3.**

## Name Review Group (NRG)

Following the declaration of an influenza pandemic by the World Health Organization (WHO) on 11 June 2009 and the initiation by the EMEA of its pandemic crisis-management plan, the NRG in collaboration with the Vaccines Working Party and the Biologics Working Party has initiated an accelerated name review procedure for the proposed names of (pandemic) influenza vaccines. The CHMP endorsed the conclusions on 16th July 2009.

Statistical information on the outcome of the checking of acceptability of proposed invented names for (pandemic) influenza vaccines through the centralised procedure is provided in **Annex 4**.

# REFERRAL PROCEDURES

## Referral procedures concluded

The CHMP concluded a referral procedure under Article 30 of Directive 2001/83/EC, as amended, for **Meronem** and associated names (meropenem), from AstraZeneca group of companies and associated companies, used as an anti-infective. The CHMP adopted a harmonised product information.

This type of procedure is initiated with a view to harmonising product information for medicinal products authorised at Member State level. The CHMP recommended the amendment of the Summary of Product Characteristics, labelling and package leaflet.

A question-and-answer document with more information about these referrals can be found here

## Referral procedures started

The CHMP started a referral procedure under Article 30 of Directive 2001/83/EC as amended for **Atacand** and associated names (candesartan), from AstraZeneca group of companies and associated companies, indicated in the treatment of essential hypertension and the treatment of patients with heart failure and impaired left ventricle systolic function.

The CHMP also initiated three referrals under Article 6(12) of Commission Regulation EC No 1084/2003 for the following authorised medicinal products:

- **Yaz 24+4** (ethinylestradiol/drospirenone) and associated names from Bayer B.V. indicated for female contraception.
- **Ethinylestradiol/drospirenon 24+4** (ethinylestradiol/drospirenone) and associated names from Bayer B.V. indicated for female contraception.
- **Extraneal** (icodextrin, sodium chloride, sodium lactate, calcium chloride dehydrate, magnesium chloride hexahydrate) from Baxter Healthcare Limited indicated for use as a once daily replacement for a single glucose exchange as part of a continuous ambulatory peritoneal dialysis (CAPD) or automated peritoneal dialysis (APD) regimen for the treatment of chronic renal failure, particularly for patients who have lost ultrafiltration on glucose solutions.

### Re-examination procedure concluded

The Committee finalised a re-examination procedure for a referral under Article 29 of Directive 2001/83/EC, as amended, for **Ciclosporin IDL** and associated names (ciclosporin), 25 mg, 50 mg and 100 mg, capsules from International Drug Licensing (IDL), indicated in transplantation and auto-immune diseases. The Committee concluded that Ciclosporin IDL is not bioequivalent to the reference product

and recommended the refusal of the Marketing Authorisation in the Concerned Member States and the suspension of the Marketing Authorisation for Ciclosporin IDL in the Member States where the product is currently authorised.

Referrals under Article 29 of Directive 2001/83/EC, as amended are initiated by one or more Member States in cases where an agreement cannot be reached in the context of the mutual recognition procedure or the decentralised procedure.

A question-and-answer document with more information about this procedure can be found here.

Re-examination procedures started under Article 32(4) of the Directive 2001/83/EC, as amended

The EMEA has been formally requested by Hospira UK Limited to re-examine the negative opinion adopted during the CHMP meeting on 22-25 June 2009 for **Teicoplanin Hospira and associated names** (teicoplanin), 200 mg and 400 mg powder and solvent for injection or infusion. The CHMP had concluded during a referral procedure under Article 29 of Directive 2001/83/EC, as amended, that sufficient evidence was not presented to demonstrate that Teicoplanin Hospira was a generic of the reference product. The CHMP recommended by consensus the refusal of the granting of the Marketing Authorisation in the Concerned Member States for Teicoplanin Hospira and associated names.

The EMEA has been formally requested by a group of Marketing Authorisation Holders to re-examine the opinion, adopted during the CHMP meeting on 22–25 June 2009 on a referral procedure under Article 31 of Directive 2001/83/EC[1], as amended, for **dextropropoxyphene containing medicinal products**, recommending the withdrawal of the Marketing Authorisations. Dextropropoxyphene-containing medicinal products are used in the treatment of acute and chronic pain.

Withdrawal

The EMEA has been formally notified by Cangene Europe Ltd. of its decision to withdraw all its applications and all marketing authorizations for **WinRho SDF** (Human anti-D immunoglobulin), 1500U and 5000U powder for solution for injection/infusion used to increase patients' platelet counts. In April 2009 the CHMP had started a referral procedure under Article 29 of Directive 2001/83/EC, as amended because of disagreements over bioequivalence.

## MUTUAL RECOGNITION AND DECENTRALISED PROCEDURES - HUMAN

The CHMP noted the report from the 42nd CMD(h) (Co-ordination Group for Mutual Recognition and Decentralised procedures-Human) held on 20-21 July 2009. For further details, please see the relevant press release on the CMD(h) website under the heading 'Press Releases': http://www.hma.eu/

## CHMP WORKING PARTIES

The CHMP was informed of the outcome of the discussions of the Scientific Advice Working Party (SAWP) meeting, which was held on 6-8 July 2009. For further details, please see **Annex 5**.

Documents prepared by the CHMP Working Parties adopted during the July 2009 CHMP meeting are listed in **Annex 6**.

## UPCOMING MEETINGS FOLLOWING THE JULY 2009 CHMP PLENARY MEETING

- The 58th meeting of the CHMP will be held at the Agency on 21-24 September 2009.
- The next Name Review Group meeting will be held at the Agency on 15 September 2009.
- The 43rd CMD(h) (Co-ordination Group for Mutual Recognition and Decentralised Procedures) will be held at the Agency on 21-23 September 2009.

[1] This referral was made under Article 31 of Directive 2001/83/EC and not under Article 31 of Regulation (EC) 83/2001 as mentioned in the monthly report previously published.

**ORGANISATIONAL MATTERS**

The main topics addressed during the July 2009 CHMP meeting related to:

- The agreement to create a new EWP biostatistics drafting group.
- Follow-on discussion of a report from the SAG drafting group regarding better optimization of Specialised Experts and SAGs' consultation.
- An update regarding the mid-year report on Working Parties Work Programme.
- A general presentation by Dr. Dal Pan on FDA initiatives in Pharmacovigilance and Drug Safety.
- The adoption of the revision of Volume 9A of the Rules governing medicinal products in the EU following the public consultation together with the overview of comments received.
- The adoption of the joint Technical Report from ECDC and EMEA on "The bacterial challenge - time to react. A call to narrow the gap between multidrug-resistant bacteria in the EU and development of new antibacterial agents" which will be published later in September 2009.
- Preliminary discussion regarding the appointment of Rapporteurs and Co-Rapporteurs in the context of Work Sharing variations. Follow-on discussion will be held in September 2009.
- A revised European Pharmacovigilance Strategy for Antivirals (EMEA/429123/2009).
- Follow-on discussion on the latest developments regarding EMEA activities concerning the novel influenza virus (H1N1). A separate press release and a question-and-answer document with an update on the Agency's pandemic vaccines activities are available.

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This CHMP Monthly Report and other documents are available on the Internet at the following address: http://www.emea.europa.eu

## PRE-AUTHORISATION: MARKETING AUTHORISATION APPLICATIONS

| Activity | 2009 | | | | | | | | 1995 onwards |
|---|---|---|---|---|---|---|---|---|---|
| | Optional Scope | | | | Mandatory scope | | | Total | Overall total |
| | NAS | Significant innovation | Interest of Patients | Generics | Biotech | Indications | Orphans | | |
| Applications for MA submitted | 12 | 3 | 0 | 34 | 7 | 2 | 4 | 62 | 860 |
| Positive opinions | 14 | 9 | 0 | 32 | 11 | 3 | 5 | 74 | 567 |
| Negative opinions[2] | 0 | 2 | 0 | 0 | 2 | 0 | 1 | 5 | 26 |
| Withdrawals prior to opinion | 2 | 0 | 0 | 1 | 3 | 2 | 2 | 10 | 149 |
| Marketing authorisation granted by the Commission | 12 | 10 | 0 | 5 | 8 | 2 | 5 | 42 | 527 |

## PRE-AUTHORISATION: SCIENTIFIC SERVICES

| Activity (submissions) | 2009 | 1995 onwards |
|---|---|---|
| Compassionate use applications | 0 | 0 |
| Art. 58 applications | 0 | 4 |
| Consultation for medical devices[3] | 1 | 6 |
| PMF (Click here for a list of PMF certifications) | 1 | 14 |
| VAMF | 0 | 0 |

[2] In case of Re-examination under Art. 9(2) of Regulation (EC) No. 726/2004, the opinion will not be counted twice.
[3] Consultation in accordance with Council Directive 93/42/EEC concerning medical devices as amended by Directive 2000/70/EC as regards medical devices incorporating stable derivates of human blood or plasma and Directive 2001/104/EC

## OUTCOME OF THE JULY 2009 CHMP MEETING IN RELATION TO ACCELERATED ASSESSMENT PROCEDURES

| Substance | Intended indications(s) | Accelerated Assessment Requests | |
|---|---|---|---|
| | | Accepted | Rejected |
| Chemical | N/A | N/A | N/A |
| Biological | N/A | N/A | N/A |

**ANNEX 2 TO CHMP MONTHLY REPORT JULY 2009**

## POST-AUTHORISATION: TYPE I AND II VARIATIONS, ANNEX II, RENEWALS AND ANNUAL RE-ASSESSMENT APPLICATIONS

| Activity | 2009 | Overall total 1995 onwards |
|---|---|---|
| Type I Variations (positive notifications) | 679 | 7048 |
| Type II Variations (positive opinions) | 681 | 5224 |
| Type II Variations (negative opinions) | 2 | 18 |
| Annex II Applications (positive opinions) | 40 | 223 |
| Annual Re-assessments (positive opinions) | 11 | - |
| Opinions for renewals of conditional MA's (positive opinions) | 2 | 8 |
| 5-year Renewals (positive opinions) | 44 | - |

| Opinions for Type II Variation applications | |
|---|---|
| **Number of Opinions** | **Outcome** |
| 9 Extension of indication | 8 Positive opinions<br>1 Negative Opinion |
| 79 SPC changes | 79 Positive opinions |
| 30 Quality changes | 30 Positive opinions |

| Opinions for Annual Re-Assessment applications | | |
|---|---|---|
| **Name of Medicinal Product (INN) MAH** | **Outcome** | **Comments** |
| N/A | N/A | N/A |

| Opinion for renewals of conditional MA's | | |
|---|---|---|
| **Name of Medicinal Product (INN) MAH** | **Outcome** | **Comments** |
| N/A | N/A | N/A |

| Opinions for 5-Year Renewal applications | | |
|---|---|---|
| **Name of Medicinal Product (INN) MAH** | **Outcome** | **Comments** |
| **Ammonap** (sodium phenylbutyrate), Swedish Orphan International AB | Positive Opinion adopted | Unlimited validity |
| **Wilzin** (zinc acetate dihydrate), Orphan Europe S.A.R.L | Positive Opinion adopted | Unlimited validity |
| **Xagrid** (anagrelide), Shire Pharmaceutical Contracts Limited | Positive Opinion adopted | Unlimited validity |
| **Protelos** (strontium ranelate), Les Laboratoires Servier | Positive Opinion adopted | Recommending additional renewal |
| **Osseor** (strontium ranelate), Les Laboratoires Servier | Positive Opinion adopted | Recommending additional renewal |

**ANNEX 3 TO CHMP MONTHLY REPORT JULY 2009**

**MEDICINAL PRODUCTS GRANTED A COMMUNITY MARKETING AUTHORISATION UNDER THE CENTRALISED PROCEDURE SINCE THE JUNE 2009 CHMP MONTHLY REPORT**

| | |
|---|---|
| **Invented Name** | Vedrop |
| **INN** | tocofersolan |
| **Marketing Authorisation Holder** | Orphan Europe S.A.R.L. |
| **Proposed ATC code** | A11HA08 |
| **Indication** | Vedrop is indicated in vitamin E deficiency due to digestive malabsorption in paediatric patients suffering from congenital chronic cholestasis or hereditary chronic cholestasis, from birth (in term newborns) to 16 or 18 years of age, depending on the region. |
| **CHMP Opinion date** | 29.05.2009 |
| **Marketing Authorisation Date** | 24.07.2009 |

| | |
|---|---|
| **Invented Name** | Nymusa |
| **INN** | caffeine citrate |
| **Marketing Authorisation Holder** | Chiesi Farmaceutici SpA |
| **Proposed ATC code** | N06BC01 |
| **Indication** | Treatment of primary apnoea of premature newborns |
| **CHMP Opinion date** | 23.04.2009 |
| **Marketing Authorisation Date** | 02.07.2009 |

| | |
|---|---|
| **Invented Name** | Iressa |
| **INN** | gefitinib |
| **Marketing Authorisation Holder** | AstraZeneca AB |
| **Proposed ATC code** | L01XE02 |
| **Indication** | IRESSA is indicated for the treatment of adult patients with locally advanced or metastatic non-small cell lung cancer (NSCLC) with activating mutations of EGFR-TK. |
| **CHMP Opinion date** | 23.04.2009 |
| **Marketing Authorisation Date** | 24.06.2009 |

| | |
|---|---|
| **Invented Name** | Victoza |
| **INN** | liraglutide |
| **Marketing Authorisation Holder** | Novo Nordisk A/S |
| **Proposed ATC code** | A10BX07 |
| **Indication** | Victoza is indicated for treatment of adults with type 2 diabetes mellitus to achieve glycaemic control:<br>In combination with: Metformin or a sulphonylurea, in patients with insufficient glycaemic control despite maximal tolerated dose of monotherapy with metformin or sulphonylurea.<br>In combination with: Metformin and a sulphonylurea or metformin and a thiazolidinedione in patients with insufficient glycaemic control despite dual therapy. |
| **CHMP Opinion date** | 23.04.2009 |
| **Marketing Authorisation Date** | 30.06.2009 |

| | |
|---|---|
| **Invented Name** | Ribavirin Teva B.V |
| **INN** | ribavirin |
| **Marketing Authorisation Holder** | Teva Pharma B.V |
| **Proposed ATC code** | J05AB04 |
| **Indication** | Ribavirin Teva Pharma B.V. is indicated for the treatment of chronic hepatitis C and must only be used as part of a combination regimen with peginterferon alfa-2b (adults) or interferon alfa-2b (adults, children (3-years of age or older), and adolescents). Ribavirin monotherapy must not be used. |
| **CHMP Opinion date** | 23.04.2009 |
| **Marketing Authorisation Date** | 01.07.2009 |

| | |
|---|---|
| **Invented Name** | Repaglinide Teva |
| **INN** | repaglinide |
| **Marketing Authorisation Holder** | Teva Pharma B.V |
| **Proposed ATC code** | A10B X02 |
| **Indication** | Repaglinide is indicated in patients with type 2 diabetes (Non Insulin-Dependent Diabetes Mellitus (NIDDM)) whose hyperglycaemia can no longer be controlled satisfactorily by diet, weight reduction and exercise. Repaglinide is also indicated in combination with metformin in type 2 diabetes patients who are not satisfactorily controlled on metformin alone. Treatment should be initiated as an adjunct to diet and exercise to lower the blood glucose in relation to meals. |
| **CHMP Opinion date** | 23.04.2009 |
| **Marketing Authorisation Date** | 29.06.2009 |

**ANNEX 4 TO CHMP MONTHLY REPORT JULY 2009**

**INVENTED NAME REVIEW GROUP (NRG)**

| | **NRG meeting 27 Jan 2009** | | **NRG meeting 17 Mar 2009** | | **NRG meeting 12 May 2009** | | **NRG accelerated name review for (pandemic) influenza vaccines 16 Jul 2009** | | **2009** | |
|---|---|---|---|---|---|---|---|---|---|---|
| | Accepted | Rejected | Accepted | Rejected | Accepted | Rejected | Accepted | Rejected | Accepted | Rejected |
| Proposed invented names | 47 | 52 | 30 | 36 | 27 | 28 | 17 | 6 | 121 | 122 |
| Justification for retention of invented name * | 5 | 1 | 3 | 1 | 2 | 1 | 0 | 0 | 10 | 3 |

*In case of objections to the proposed invented name(s), the applicant may justify the retention of the proposed invented name using the relevant justification form available on the EMEA website.

**ANNEX 5 TO CHMP MONTHLY REPORT JULY 2009**

## PRE-AUTHORISATION: SCIENTIFIC ADVICE AND PROTOCOL ASSISTANCE EMEA CENTRALISED PROCEDURES

| | 1995 - 2008 | 2009 | Overall Total |
|---|---|---|---|
| Scientific Advice | 887 | **149** | **1036** |
| Follow-up to Scientific Advice | 171 | **26** | **197** |
| Protocol Assistance | 198 | **25** | **223** |
| Follow-up to Protocol Assistance | 90 | **11** | **101** |
| | **1346** | **211** | **1557** |

## OUTCOME OF THE JULY 2009 CHMP MEETING IN RELATION TO SCIENTIFIC ADVICE PROCEDURES

### Final Scientific Advice Procedures

| Substance | Intended indications(s) | Type of Request | | | | Topic | | | |
|---|---|---|---|---|---|---|---|---|---|
| | | New | | Follow-up | | Pharmaceutical | Pre-clinical | Clinical | Significant Benefit |
| | | SA | PA | SA | PA | | | | |
| Biological | Enzyme replacement therapy to improve growth and development | X | | | | | | X | |
| Chemical | Treatment of Chronic Lymphocytic Leukaemia | X | | | | X | X | X | |
| Chemical | Treatment of Acute Myeloid Leukaemia | | X | - | | | X | X | X |
| Chemical | Generic of Xeloda | X | | | | | | X | |
| Biological | Treatment of relapsed or refractory Hairy Cell Leukaemia | | X | | | | X | X | |

| Substance | Intended indications(s) | Type of Request | | | | Topic | | | |
|---|---|---|---|---|---|---|---|---|---|
| | | New | | Follow-up | | Pharmaceutical | Pre-clinical | Clinical | Significant Benefit |
| | | SA | PA | SA | PA | | | | |
| Chemical | Treatment of Stage IV Melanoma | X | | | | | X | X | |
| Biological | Radiotherapy treatment for patients with stage I and II breast cancer who have undergone breast surgery | X | | | | | | X | |
| Biological | Treatment of Diffuse Large B Cell Lymphoma | X | | | | | | X | |
| Chemical | Treatment of Multiple Myeloma | | X | | | | | X | |
| Biological | Treatment of Hodgkin Lymphoma | | X | | | X | X | X | |
| Chemical | Treatment of Multiple Myeloma | | | | X | | | X | |
| Biological | Treatment of Rheumatoid Arthritis | X | | | | X | X | X | |
| Chemical | Treatment of Glioblastoma Multiforme | X | | | | | | X | |
| Biological | Prevention and treatment of haemorrhage or surgical bleeding in von Willebrand disease | X | | | | X | | X | |
| Biological | Treatment and prophylaxis of bleeding in Hemophilia A | X | | | | X | X | X | |
| Chemical | Treatment of Pulmonary Arterial Hypertension | X | | | | | X | X | |

| Substance | Intended indications(s) | Type of Request | | | | Topic | | | |
|---|---|---|---|---|---|---|---|---|---|
| | | New | | Follow-up | | Pharma ceutical | Pre-clinical | Clinical | Significant Benefit |
| | | SA | PA | SA | PA | | | | |
| Biological | Treatment of Rheumatoid Arthritis, Psoriatic Arthritis, Severe Ankylosing Spondylitis, Psoriasis, Polyarticular Juvenile Idiopathic Arthritis | X | | | | X | | | |
| Chemical | Treatment of Hyperlipidemias | | | X | | | | X | |
| Chemical | Treatment of Acute Coronary Syndrome, chronic protection in patients with a history of Myocardial Infraction (MI), Thrombotic Stroke or Peripheral Arterial Disease | X | | | | X | | | |
| Advanced Therapy | Treatment of Epodermolysis Bullosa | | X | | | | X | | |
| Chemical | Treatment of Solar Urticaria | | X | | | | X | | |
| Biological | Prevention of Influenza infection | | | X | | X | | X | |
| Biological | Prevention of Influenza infection | X | | | | | X | | |
| Chemical | Treatment of mild to moderate and prodromal Alzheimer's Disease | X | | | | | X | X | |
| Chemical | Generic to Keppra | X | | | | X | | X | |

| Substance | Intended indications(s) | Type of Request | | | | Topic | | | |
|---|---|---|---|---|---|---|---|---|---|
| | | New | | Follow-up | | Pharma ceutical | Pre-clinical | Clinical | Significant Benefit |
| | | SA | PA | SA | PA | | | | |
| Chemical | Generic to olanzapine | X | | | | X | | X | |
| Chemical | Treatment of Chronic Obstructive Pulmonary Disease | X | | | | X | X | X | |
| Chemical | Treatment of Allergic Rhinitis and Chronic Idiopathic Urticaria | X | | | | | | X | |
| Biological | Treatment of Asthma | X | | | | | X | | |
| Advanced Therapy | Treatment of total bilateral and unilateral limbal stem cell deficiency | X | | | | X | X | X | |
| Biological | Treatment of post-menopausal osteoporosis | | | X | | | | X | |
| Biological | Treatment of Immunglobulin-E mediated allergic diseases | | | X | | | | X | |
| Chemical | Agent for diagnostic Magnetic Resonance Imaging for the detection of lesions of the liver suspected to be due to metastatic disease | X | | | | | X | X | |
| Chemical | Treatment of Hyperphosphata-emia | X | | | | X | X | X | |
| Chemical | Control of Hyperphosphata-emia | X | | | | X | | | |

SA: Scientific Advice
PA: Protocol Assistance

The above-mentioned 24 Scientific Advice letters, 6 Protocol Assistance letters, 4 Follow-up Scientific Advice and 1 Follow-up Protocol Assistance letters were adopted at the 20-23 July 2009 CHMP meeting.

**New requests for Scientific Advice Procedures**

The Committee accepted 46 new Requests for which the procedure started at the SAWP meeting held on 6-8 July 2009. The new requests are divided as follows: 29 Initial Scientific Advice, 9 Follow-up Scientific Advice, 7 Initial Protocol Assistance and 1 Follow-up Protocol Assistance.

# DOCUMENTS PREPARED BY THE CHMP WORKING PARTIES ADOPTED DURING THE JULY 2009 CHMP MEETING

## BIOLOGIC WORKING PARTY (BWP)

| Reference number | Document | Status[4] |
|---|---|---|
| (EMEA/52104/2008) | Draft Report on Expert Workshop on CJD Risk and Urine-derived Medicinal Products | Adopted |
| (EMEA/CHMP/BWP/ 290688/2009 | Concept Paper on a Revision of the Guideline on Pharmaceutical Aspects of the Product Information for Human Vaccines EMEA/CPMP/BWP/2758/02 | Adopted |
| (EMEA/CPMP/BWP/ 2879/02 rev.1) | Concept Paper on the Need to update the CHMP position statement on CJD and Plasma-derived and Urine-derived Medicinal Products | Adopted |

## BLOOD PRODUCT WORKING PARTY (BPWP)

| Reference number | Document | Status[4] |
|---|---|---|
| EMEA/CHMP/BPWP/ 144533/2009 | Guideline on the Clinical Investigation of Recombinant and Human Plasma-derived Factor VIII Products | Adopted for 3-month public consultation |
| EMEA/CHMP/BPWP/ 144552/2009 | Guideline on the Clinical Investigation of Recombinant and Human Plasma-derived Factor IX Products | Adopted for 3-month public consultation |
| EMEA/CHMP/BPWP/ 161104/2009 | Concept Paper on the Need for a Guideline on the Clinical Investigation of Specific Immunoglobulins | Adopted |

## GENE THERAPY WORKING PARTY (GTWP)

| Reference number | Document | Status[4] |
|---|---|---|
| EMEA/CHMP/ICH/ 449035/2009 | Final version of the 'ICH Considerations on the General Principles to Address Virus and Vector Shedding' adopted by ICH Steering Committee in June 2009 | Adopted |

## WORKING PARTY ON BIOSIMILAR MEDICINAL PRODUCTS (BMWP)

| Reference number | Document | Status[4] |
|---|---|---|
| EMEA/CHMP/BMWP/ 233377/2008 | Overview of comments received on the Reflection Paper on Similar Medicinal Products containing Recombinant Interferon Alpha - EMEA/CHMP/BMWP/102046/2006 | Adopted |
| EMEA/CHMP/BMWP/ 301636/2008 | Draft revised Guideline on Non-Clinical and Clinical development of Similar Biological Medicinal products containing recombinant erythropoietins | Adopted for 4-month public consultation |

[4] Adopted or release for consultation documents can be found at the EMEA website (under "What's new-recent publications" or under Human Medicines-Guidance documents).

## EFFICACY WORKING PARTY (EWP)

| Reference number | Document | Status[4] |
|---|---|---|
| CPMP/EWP/1119/98 Rev.1 | Guideline on Clinical Evaluation of Diagnostic Agents | Adopted |
| (EMEA/CHMP/EWP/ 321180/2008 | Appendix 1 to the Guideline on Clinical Evaluation of Diagnostic Agents (CPMP/EWP/1119/98 Rev 1) - on Imaging Agents | Adopted |
| EMEA/CHMP/EWP/ 350495/2009 | Concept Paper on the Need to update the Note for Guidance on Clinical Investigation of Medicinal Products and Note for Guidance on the Clinical Investigation on Medicinal Products in the Treatment of Hypertension (CPMP/EWP/238/95 rev. 2) to discuss the need for outcome studies basis on safety data at the time of MAA | Adopted |

## PHARMACOVIGILANCE WORKING PARTY (PhVWP)

| Reference number | Document | Status[4] |
|---|---|---|
| EMEA/CHMP/PhVWP/ 594810/2008 | Guideline on Preparation of Assessment Report on PSURs | Adopted |
| EMEA/426669/2009<br><br>EMEA/426674/2009 | Guideline on the Conduct of Pharmacovigilance for vaccines for Pre- and Post-Exposure Prophylaxis against infectious diseases<br><br>Overview of comments | Adopted |