B. PACKAGE LEAFLET

PACKAGE LEAFLET: INFORMATION FOR THE USER

Pandemrix suspension and emulsion for emulsion for injection
Pandemic influenza vaccine (H1N1) (split virion, inactivated, adjuvanted)

Read all of this leaflet carefully before you receive this vaccine .
- Keep this leaflet. You may need to read it again.
- If you have any further questions, ask your doctor or nurse.
- If any of the side effects gets serious, or if you notice any side effects not listed in this leaflet, please tell your doctor.

In this leaflet:
1. What Pandemrix is and what it is used for
2. Before you receive Pandemrix
3. How Pandemrix is given
4. Possible side effects
5. How to store Pandemrix
6. Further information

1. What Pandemrix is and what it is used for

Pandemrix is a vaccine to prevent pandemic influenza (flu).

Pandemic flu is a type of influenza that occurs every few decades and which spreads rapidly around the world. The symptoms of pandemic flu are similar to those of ordinary flu but may be more severe.

When a person is given the vaccine, the immune system (the body's natural defence system) will produce its own protection (antibodies) against the disease. None of the ingredients in the vaccine can cause flu.

2. Before you receive Pandemrix

You should not receive Pandemrix:

- if you have previously had a sudden life-threatening allergic reaction to any ingredient of Pandemrix (these are listed at the end of the leaflet) or to any of the substances that may be present in trace amounts as follows: egg and chicken protein, ovalbumin, formaldehyde, gentamicin sulphate (antibiotic) or sodium deoxycholate. Signs of an allergic reaction may include itchy skin rash, shortness of breath and swelling of the face or tongue. However, in a pandemic situation, it may be appropriate for you to have the vaccine provided that appropriate medical treatment is immediately available in case of an allergic reaction.

If you are not sure, talk to your doctor or nurse before having this vaccine.

Take special care with Pandemrix:

- if you have had any allergic reaction other than a sudden life-threatening allergic reaction to any ingredient contained in the vaccine, to thiomersal, to egg and chicken protein, ovalbumin, formaldehyde, gentamicin sulphate (antibiotic) or to sodium deoxycholate. (see section 6. Further information).
- if you have a severe infection with a high temperature (over 38°C). If this applies to you then your vaccination will usually be postponed until you are feeling better. A minor infection such

as a cold should not be a problem, but your doctor or nurse will advise whether you could still be vaccinated with Pandemrix,
- if you are having a blood test to look for evidence of infection with certain viruses. In the first few weeks after vaccination with Pandemrix the results of these tests may not be correct. Tell the doctor requesting these tests that you have recently been given Pandemrix.

In any of these cases, TELL YOUR DOCTOR OR NURSE, as vaccination may not be recommended, or may need to be delayed.

If your child receives the vaccine, you should be aware that the side effects may be more intense after the second dose, especially temperature over 38°C. Therefore monitoring of temperature and measures to lower the temperature (such as giving paracetamol or other medicines that lower fever) after each dose are recommended.

Please inform your doctor or nurse if you have a bleeding problem or bruise easily.

Taking other medicines
Please tell your doctor or nurse if you are taking or have recently taken any other medicines, including medicines obtained without a prescription or have recently been given any other vaccine.

Pandemrix can be given at the same time as seasonal influenza vaccines that do not contain an adjuvant.

Persons who have received a seasonal influenza vaccine that does not contain an adjuvant may receive Pandemrix after an interval of at least three weeks.

There is no information on administration of Pandemrix with other vaccines. However, if this cannot be avoided, the vaccines should be injected into separate limbs. In such cases, you should be aware that the side effects may be more intense.

Pregnancy and breast-feeding
Tell your doctor if you are pregnant, think you may be pregnant, plan to become pregnant. You should discuss with your doctor whether you should receive Pandemrix.
The vaccine may be used during breast-feeding.

Driving and using machines
Some effects mentioned under section 4. "Possible side effects" may affect the ability to drive or use machines.

Important information about some of the ingredients of Pandemrix
This vaccine contains thiomersal as a preservative and it is possible that you may experience an allergic reaction. Tell your doctor if you have any known allergies.

This medicinal product contains less than 1 mmol sodium (23 mg) and less than 1 mmol of potassium (39 mg) per dose, i.e. essentially sodium- and potassium-free.

3. How Pandemrix is given

Your doctor or nurse will administer the vaccine in accordance with official recommendations.

The vaccine will be injected into a muscle (usually in the upper arm).

<u>Adults, including the elderly and children from the age of 10 years onwards</u>
A dose (0.5 ml) of the vaccine will be given.
Clinical data suggest that a single dose may be sufficient.
If a second dose is administered there should be an interval of at least three weeks between the first and second dose.

Children from 6 months to 9 years of age
A dose (0.25 ml) of the vaccine will be given.
If a second dose of 0.25 ml is given this will be administered at least three weeks after the first dose.

Children aged less than 6 months of age
Vaccination is currently not recommended in this age group.

When Pandemrix is given for the first dose, it is recommended that Pandemrix (and not another vaccine against H1N1) be given for the complete vaccination course.

4. Possible side effects

Like all medicines, Pandemrix can cause side effects, although not everybody gets them.

Allergic reactions may occur following vaccination, in rare cases leading to shock. Doctors are aware of this possibility and have emergency treatment available for use in such cases.

The frequency of possible side effects listed below is defined using the following convention:

Very common (affects more than 1 user in 10)
Common (affects 1 to 10 users in 100)
Uncommon (affects 1 to 10 users in 1,000)
Rare (affects 1 to 10 users in 10,000)
Very rare (affects less than 1 user in 10,000)

The side effects listed below have occurred with Pandemrix (H5N1) in clinical studies in adults, including the elderly. In these clinical studies most side effects were mild in nature and short term. The side-effects are generally similar to those related to seasonal flu vaccines.

These side effects have also been observed with similar frequencies in clinical studies in adults including the elderly and in children aged 10 to 17 years with Pandemrix (H1N1), except for redness (uncommon in the adults and common in the elderly) and fever (uncommon in the adults and elderly). Gastro-intestinal symptoms and shivering were at a higher rate in the children 10-17 years of age. In children aged 3-9 years who received a first half adult dose of Pandemrix (H1N1), the side effects were similar compared to the side effects reported in adults, with the exception of shivering, sweating and gastro-intestinal symptoms which were reported at a higher rate in children aged 3 to 9 years. Additionally, in children aged 3 to 5 years of age, drowsiness, irritability and loss of appetite were reported very commonly.

Very common:
- Headache
- Tiredness
- Pain, redness, swelling or a hard lump at the injection site
- Fever
- Aching muscles, joint pain

Common:
- Warmth, itching or bruising at the injection site
- Increased sweating, shivering, flu-like symptoms
- Swollen glands in the neck, armpit or groin

Uncommon:
- Tingling or numbness of the hands or feet
- Sleepiness

- Dizziness
- Diarrhoea, vomiting, stomach pain, feeling sick
- Itching, rash
- Generally feeling unwell
- Sleeplessness

In children aged 6-35 months who received a half of the adult dose (0.25 ml) of Pandemrix (H1N1), fever and irritability occurred more often compared to the children 3-9 years who received a half of the adult dose (0.25 ml) of Pandemrix (H5N1).

In children aged 6-35 months who received two doses of 0.25 ml (half of the adult dose) the side effects after the second dose were more intense, especially fever (≥38°C), which occurred very commonly.

These side effects usually disappear within 1-2 days without treatment. If they persist, CONSULT YOUR DOCTOR.

The side effects listed below have occurred during post-marketing experience with Pandemrix H1N1 vaccine:

- Allergic reactions leading to a dangerous decrease of blood pressure, which, if untreated, may lead to shock. Doctors are aware of this possibility and have emergency treatment available for use in such cases.
- Generalised skin reactions including facial swelling and urticaria (hives)
- Fits due to fever

The side effects listed below have occurred in the days or weeks after vaccination with vaccines given routinely every year to prevent flu. These side effects may occur with Pandemrix.

Rare
- Severe stabbing or throbbing pain along one or more nerves
- Low blood platelet count which can result in bleeding or bruising

Very rare
- Vasculitis (inflammation of the blood vessels which can cause skin rashes, joint pain and kidney problems)
- Neurological disorders such as encephalomyelitis (inflammation of the central nervous system), neuritis (inflammation of nerves) and a type of paralysis known a Guillain-Barré Syndrome

If any of these side effects occur, please tell your doctor or nurse immediately.

If any of the side effects gets serious, or if you notice any side effects not listed in this leaflet, please tell your doctor.

5. How to store Pandemrix

Keep out of the reach and sight of children.

Before the vaccine is mixed:
Do not use the suspension and the emulsion after the expiry date which is stated on the carton. The expiry date refers to the last day of that month.

Store in a refrigerator (2°C - 8°C).
Store in the original package in order to protect from light.
Do not freeze.

After the vaccine is mixed:
After mixing, use the vaccine within 24 hours and do not store above 25°C.

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

6. Further information

What Pandemrix contains

- Active substance:
  Split influenza virus, inactivated, containing antigen* equivalent to:

  A/California/7/2009 (H1N1)v-like strain (X-179A) 3.75 micrograms** per 0.5 ml dose

  *propagated in eggs
  **expressed in microgram haemagglutinin

  This vaccine complies with the WHO recommendation and EU decision for the pandemic.

- Adjuvant:
  The vaccine contains an 'adjuvant' AS03 to stimulate a better response. This adjuvant contains squalene (10.69 milligrams), DL-α-tocopherol (11.86 milligrams) and polysorbate 80 (4.86 milligrams).

- Other ingredients:
  The other ingredients are: polysorbate 80, octoxynol 10, thiomersal, sodium chloride, disodium hydrogen phosphate, potassium dihydrogen phosphate, potassium chloride, magnesium chloride, water for injections

What Pandemrix looks like and contents of the pack

Suspension and emulsion for emulsion for injection.
The suspension is a colourless light opalescent liquid.
The emulsion is a whitish homogeneous liquid.

Prior to administration, the two components should be mixed. The mixed vaccine is a whitish emulsion.

One pack of Pandemrix consists of:
- one pack containing 50 vials of 2.5 ml suspension (antigen)
- two packs containing 25 vials of 2.5 ml emulsion (adjuvant)

Marketing Authorisation Holder and Manufacturer

GlaxoSmithKline Biologicals s.a.
Rue de l'Institut 89
B-1330 Rixensart
Belgium

For any information about this medicine, please contact the local representative of the Marketing Authorisation Holder:

België/Belgique/Belgien **Luxembourg/Luxemburg**

GlaxoSmithKline s.a./n.v.
Tél/Tel: + 32 2 656 21 11

България
ГлаксоСмитКлайн ЕООД
Тел.: + 359 2 953 10 34

Česká republika
GlaxoSmithKline s.r.o.
Tel: + 420 2 22 00 11 11
gsk.czmail@gsk.com

Danmark
GlaxoSmithKline Pharma A/S
Tlf: + 45 36 35 91 00
dk-info@gsk.com

Deutschland
GlaxoSmithKline GmbH & Co. KG
Tel: + 49 (0)89 360448701
produkt.info@gsk.com

Eesti
GlaxoSmithKline Eesti OÜ
Tel: +372 667 6900
estonia@gsk.com

Ελλάδα
GlaxoSmithKline A.E.B.E
Τηλ: + 30 210 68 82 100

España
GlaxoSmithKline, S.A.
Tel: + 34 902 202 700
es-ci@gsk.com

France
Laboratoire GlaxoSmithKline
Tél: + 33 (0) 800 00 12 12
grippeA@gsk.com

Ireland
GlaxoSmithKline (Ireland) Ltd
Tel: + 353 (0)1 4955000

Ísland
GlaxoSmithKline ehf.
Sími: +354-530 3700

Italia
GlaxoSmithKline S.p.A.
Tel:+ 39 04 59 21 81 11

Κύπρος
GlaxoSmithKline s.a./n.v.
Tél/Tel: + 32 2 656 21 11

Magyarország
GlaxoSmithKline Kft.
Tel.: + 36-1-2255300

Malta
GlaxoSmithKline Malta
Tel: + 356 21 238131

Nederland
GlaxoSmithKline BV
Tel: + 31 (0)30 69 38 100
nlinfo@gsk.com

Norge
GlaxoSmithKline AS
Tlf: + 47 22 70 20 00
firmapost@gsk.no

Österreich
GlaxoSmithKline Pharma GmbH.
Tel: + 43 1 970 75-0
at.info@gsk.com

Polska
GSK Commercial Sp. z o.o.
Tel.: + 48 (22) 576 9000

Portugal
GlaxoSmithKline, Produtos Farmacêuticos, Lda.
Tel: + 351 21 412 95 00
FI.PT@gsk.com

România
GlaxoSmithKline (GSK) SRL
Tel: + 40 (0)21 3028 208

Slovenija
GlaxoSmithKline d.o.o.
Tel: + 386 (0) 1 280 25 00
medical.x.si@gsk.com

Slovenská republika
GlaxoSmithKline Slovakia s.r.o.
Tel: + 421 (0)2 48 26 11 11
recepcia.sk@gsk.com

Suomi/Finland
GlaxoSmithKline Oy
Puh/Tel: + 358 10 30 30 30
Finland.tuoteinfo@gsk.com

Sverige

GlaxoSmithKline Cyprus Ltd
Τηλ: + 357 22 39 70 00

Latvija
GlaxoSmithKline Latvia SIA
Tel: + 371 67312687
lv-epasts@gsk.com

Lietuva
GlaxoSmithKline Lietuva UAB
Tel. +370 5 264 90 00
info.lt@gsk.com

GlaxoSmithKline AB
Tel: + 46 (0)8 638 93 00
info.produkt@gsk.com

United Kingdom
GlaxoSmithKline UK
Tel: + 44 (0)808 100 9997
customercontactuk@gsk.com

This leaflet was last approved in {MM/YYYY}.

Pandemrix has been authorised under "Exceptional Circumstances".
The European Medicines Agency (EMEA) will regularly review any new information on the medicine and this package leaflet will be updated as necessary.

Detailed information on this medicine is available on the European Medicines Agency (EMEA) web site: http://www.emea.europa.eu/

The following information is intended for medical or healthcare professionals only:

Pandemrix consists of two containers:
Suspension: multidose vial containing the antigen,
Emulsion: multidose vial containing the adjuvant.

Prior to administration, the two components should be mixed.

Instructions for mixing and administration of the vaccine:

1. Before mixing the two components, the emulsion (adjuvant) and suspension (antigen) should be allowed to reach room temperature; each vial should be shaken and inspected visually for any foreign particulate matter and/or abnormal physical appearance. In the event of either being observed (including rubber particles from the stopper), discard the vaccine.
2. The vaccine is mixed by withdrawing the entire contents of the vial containing the adjuvant by means of a syringe and by adding it to the vial containing the antigen.
3. After the addition of the adjuvant to the antigen, the mixture should be well shaken. The mixed vaccine is a whitish emulsion. In the event of other variation being observed, discard the vaccine.
4. The volume of the Pandemrix vial after mixing is at least 5 ml. The vaccine should be administered in accordance with the recommended posology (see section 3 "How Pandemrix is given").
5. The vial should be shaken prior to each administration and inspected visually for any foreign particulate matter and/or abnormal physical appearance. In the event of either being observed (including rubber particles from the stopper), discard the vaccine.
6. Each vaccine dose of 0.5 ml (full dose) or 0.25 ml (half dose) is withdrawn into a syringe for injection and administered intramuscularly.
7. After mixing, use the vaccine within 24 hours. The mixed vaccine can either be stored in a refrigerator (2°C - 8°C) or at room temperature not exceeding 25°C. If the mixed vaccine is stored in a refrigerator, it should be allowed to reach room temperature before each withdrawal.

The vaccine should not be administered intravascularly.

Any unused product or waste material should be disposed of in accordance with local requirements.