ANNEX I

SUMMARY OF PRODUCT CHARACTERISTICS

▼This medicinal product is subject to additional monitoring. This will allow quick identification of new safety information. Healthcare professionals are asked to report any suspected adverse reactions. See section 4.8 for how to report adverse reactions.

1. NAME OF THE MEDICINAL PRODUCT

FLUENZ nasal spray suspension
Influenza vaccine (live attenuated, nasal)

2. QUALITATIVE AND QUANTITATIVE COMPOSITION

Reassortant influenza virus* (live attenuated) of the following strains**:

A/California/7/2009 (H1N1)pdm09-like strain (A/California/7/2009, MEDI 228029)	$10^{7.0\pm0.5}$ FFU***
A/Victoria/361/2011 (H3N2)-like strain (A/Texas/50/2012, MEDI 237514)	$10^{7.0\pm0.5}$ FFU***
B/Massachusetts/2/2012-like strain (B/Massachusetts/2/2012, MEDI 237751)	$10^{7.0\pm0.5}$ FFU***

..... per 0.2 ml dose

* propagated in fertilised hens' eggs from healthy chicken flocks.
** produced in VERO cells by reverse genetic technology. This product contains genetically modified organisms (GMOs).
*** fluorescent focus units

This vaccine complies with the WHO recommendation (Northern Hemisphere) and EU decision for the 2013/2014 season.

The vaccine may contain residues of the following substances: egg proteins (e.g. ovalbumin) and gentamicin.

For the full list of excipients, see section 6.1.

3. PHARMACEUTICAL FORM

Nasal spray, suspension

The suspension is colourless to pale yellow, clear to opalescent. Small white particles may be present.

4. CLINICAL PARTICULARS

4.1 Therapeutic indications

Prophylaxis of influenza in individuals 24 months to less than 18 years of age.

The use of FLUENZ should be based on official recommendations.

4.2 Posology and method of administration

Posology

Children and adolescents from 24 months:
0.2 ml (administered as 0.1 ml per nostril).

For children who have not previously been vaccinated against seasonal influenza, a second dose should be given after an interval of at least 4 weeks.

FLUENZ should not be used in infants and toddlers below 24 months of age because of safety concerns (see section 4.4).

Method of administration
Immunisation must be carried out by nasal administration.

DO NOT INJECT FLUENZ.

See section 6.6 for administration instructions.

4.3 Contraindications

Hypersensitivity to the active substances or to any of the excipients listed in section 6.1 (e.g. gelatin), or to gentamicin (a possible trace residue), eggs or egg proteins (e.g. ovalbumin).

Children and adolescents who are clinically immunodeficient due to conditions or immunosuppressive therapy such as: acute and chronic leukaemias; lymphoma; symptomatic HIV infection; cellular immune deficiencies; and high-dose corticosteroids. FLUENZ is not contraindicated for use in individuals with asymptomatic HIV infection; or individuals who are receiving topical/inhaled corticosteroids or low-dose systemic corticosteroids or those receiving corticosteroids as replacement therapy, e.g. for adrenal insufficiency.

Children and adolescents younger than 18 years of age receiving salicylate therapy because of the association of Reye's syndrome with salicylates and wild-type influenza infection.

4.4 Special warnings and precautions for use

As with most vaccines, appropriate medical treatment and supervision should always be readily available in case of an anaphylactic event following the administration of FLUENZ.

FLUENZ should not be administered to children and adolescents with severe asthma or active wheezing because these individuals have not been adequately studied in clinical studies.

Do not administer FLUENZ to infants and toddlers younger than 12 months. In a clinical study, an increase in hospitalisations was observed in infants and toddlers younger than 12 months after vaccination (see section 4.8). It is not recommended to administer FLUENZ to infants and toddlers 12-23 months of age. In a clinical study, an increased rate of wheezing was observed in infants and toddlers 12-23 months of age after vaccination (see section 4.8).

Vaccine recipients should be informed that FLUENZ is an attenuated live virus vaccine and has the potential for transmission to immunocompromised contacts. Vaccine recipients should attempt to avoid, whenever possible, close association with severely immunocompromised individuals (e.g. bone marrow transplant recipients requiring isolation) for 1-2 weeks following vaccination. Peak incidence of vaccine virus recovery occurred 2-3 days post-vaccination in clinical studies. In circumstances where contact with severely immunocompromised individuals is unavoidable, the potential risk of transmission of the influenza vaccine virus should be weighed against the risk of acquiring and transmitting wild-type influenza virus.

FLUENZ should under no circumstances be injected.

No data exist regarding the safety of intranasal administration of FLUENZ in children with unrepaired craniofacial malformations.

4.5 Interaction with other medicinal products and other forms of interaction

Do not administer FLUENZ to children and adolescents younger than 18 years of age receiving salicylate therapy (see section 4.3). Do not use salicylates in children and adolescents younger than 18 years of age for 4 weeks after vaccination unless medically indicated as Reye's syndrome has been reported following the use of salicylates during wild-type influenza infection.

The co-administration of FLUENZ with the live attenuated vaccines: measles, mumps, rubella, varicella, and orally-administered poliovirus has been studied. No clinically meaningful changes in immune responses to measles, mumps, varicella, orally-administered poliovirus or FLUENZ have been observed. The immune response to rubella vaccine was significantly altered. However, this alteration might not be of clinical relevance with the two dose immunisation schedule of the rubella vaccine.

The co-administration of FLUENZ with inactivated vaccines has not been studied.

The concurrent use of FLUENZ with antiviral agents that are active against influenza A and/or B viruses has not been evaluated. However, based upon the potential for influenza antiviral agents to reduce the effectiveness of FLUENZ, it is recommended not to administer the vaccine until 48 hours after the cessation of influenza antiviral therapy. Administration of influenza antiviral agents within two weeks of vaccination may affect the response of the vaccine.

If influenza antiviral agents and FLUENZ are administered concomitantly, revaccination should be considered when appropriate.

4.6 Fertility, pregnancy and lactation

Pregnancy
There are limited data from the use of FLUENZ in pregnant women.

While animal studies do not indicate direct or indirect harmful effects with respect to reproductive toxicity, FLUENZ is not recommended during pregnancy.

Breastfeeding
It is not known whether FLUENZ is excreted in human milk. Therefore, as some viruses are excreted in human milk, FLUENZ should not be used during breastfeeding.

Fertility
No data exist regarding the possible effects of FLUENZ on male and female fertility.

4.7 Effects on ability to drive and use machines

The vaccine is unlikely to have an effect on the ability to drive and use machines.

4.8 Undesirable effects

Summary of the safety profile

Safety data regarding use of FLUENZ have been compiled from over 28,500 children and adolescents 2 to 17 years of age from clinical studies and over 52,500 children and adolescents

from post-authorisation safety studies. Additional experience has occurred with marketed use of this vaccine.

Although safety in children and adolescents with mild to moderate asthma has been established, data in children with other pulmonary diseases or with chronic cardiovascular, metabolic or renal diseases are limited. In studies of adults in which a high percentage of individuals had underlying chronic medical conditions, the safety profile of FLUENZ was comparable to the safety profile observed in individuals without these conditions.

Summary of adverse reactions

The most common adverse reaction observed in clinical studies was nasal congestion/rhinorrhoea.

Adverse reaction frequencies are reported as:
Very common (≥ 1/10)
Common (≥ 1/100 to < 1/10)
Uncommon (≥ 1/1,000 to < 1/100)
Very rare (< 1/10,000)

Immune system disorders
Uncommon: Hypersensitivity reactions (including facial oedema, urticaria and very rare anaphylactic reactions)

Metabolism and nutrition disorders
Very common: Decreased appetite

Nervous system disorders
Very common: Headache

Respiratory, thoracic, and mediastinal disorders
Very common: Nasal congestion/rhinorrhoea
Uncommon: Epistaxis

Skin and subcutaneous tissue disorders
Uncommon: Rash

Musculoskeletal and connective tissue disorders
Common: Myalgia

General disorders and administration site conditions
Very common: Malaise
Common: Pyrexia

In an active-controlled clinical study (MI-CP111), an increased rate of hospitalisations (for any cause) through 180 days after final vaccination dose was observed in infants and toddlers 6-11 months of age (6.1% FLUENZ versus 2.6% injectable influenza vaccine). The rate of hospitalisations was not increased in FLUENZ recipients 12 months and older. In the same study, an increased rate of wheezing through 42 days was observed in infants and toddlers 6-23 months of age (5.9% FLUENZ versus 3.8% injectable influenza vaccine). The rate of wheezing was not increased in FLUENZ recipients 24 months and older. FLUENZ is not indicated for use in infants and toddlers younger than 24 months (see section 4.4).

Very rare reports of Guillain-Barré syndrome and exacerbation of symptoms of Leigh syndrome (mitochondrial encephalomyopathy) have also been observed in the post-marketing setting.

Reporting of suspected adverse reactions
Reporting suspected adverse reactions after authorisation of the medicinal product is important. It allows continued monitoring of the benefit/risk balance of the medicinal product. Healthcare professionals are asked to report any suspected adverse reactions via the national reporting system listed in Appendix V.

4.9 Overdose

There have been occasional reports of administration of twice the recommended dose of FLUENZ in the post-marketing setting. The adverse reactions reported were similar to those seen with the recommended single dose of FLUENZ.

5. PHARMACOLOGICAL PROPERTIES

5.1 Pharmacodynamic properties

Pharmacotherapeutic group: Influenza vaccines, influenza live attenuated; ATC Code: J07BB03

The influenza virus strains in FLUENZ are (a) *cold-adapted (ca)*; (b) *temperature-sensitive (ts)*; and (c) *attenuated (att)*. As a result, they replicate in the nasopharynx and induce protective immunity.

Efficacy
FLUENZ has been administered to over 30,000 individuals in controlled clinical studies over multiple years, in various regions and using different vaccine strains.

Paediatric studies

FLUENZ's efficacy data in the paediatric population consist of 9 controlled studies comprising over 20,000 infants and toddlers, children and adolescents, conducted during 7 influenza seasons. Four placebo-controlled studies included second season revaccination. FLUENZ has demonstrated superiority in 3 active-controlled studies with injectable influenza vaccine. See Table 1 and 2 for a summary of efficacy results in the paediatric population.

Table 1 FLUENZ Efficacy in Placebo Controlled Paediatric Studies

Study Number	Region	Age Range[a]	Number of Study Participants	Influenza Season	Efficacy (95% CI)[b] Matched strains	Efficacy (95% CI)[b] All strains regardless of match
D153-P502	Europe	6 to 35 M	1,616	2000-2001	85.4% (74.3, 92.2)	85.9% (76.3, 92.0)
				2001-2002	88.7% (82.0, 93.2)	85.8% (78.6, 90.9)
D153-P504	Africa, Latin America	6 to 35 M	1,886	2001	73.5% (63.6, 81.0)[c]	72.0% (61.9, 79.8)[c]
				2002	73.6% (33.3, 91.2)	46.6% (14.9, 67.2)
D153-P513	Asia/ Oceania	6 to 35 M	2,107	2002	62.2% (43.6, 75.2)	48.6% (28.8, 63.3)
D153-P522	Europe, Asia/ Oceania, Latin America	11 to 24 M	1,150	2002-2003	78.4% (50.9, 91.3)	63.8% (36.2, 79.8)

Study Number	Region	Age Range[a]	Number of Study Participants	Influenza Season	Efficacy (95% CI)[b] Matched strains	Efficacy (95% CI)[b] All strains regardless of match
D153-P501	Asia/Oceania	12 to 35 M	2,764	2000-2001	72.9% (62.8, 80.5)	70.1% (60.9, 77.3)
				2001-2002	84.3% (70.1, 92.4)[d]	64.2% (44.2, 77.3)[d]
AV006	USA	15 to 71 M	1,259	1996-1997	93.4% (87.5, 96.5)	93.4% (87.5, 96.5)
				1997-1998	100% (63.1, 100)	87.1% (77.7, 92.6)[e]

[a] M = months
[b] Reduction in culture-confirmed influenza illness relative to placebo.
[c] Data presented for clinical trial D153-P504 are for study participants who received two doses of study vaccine. In previously unvaccinated study participants who received one dose in year 1, efficacy was 57.7% (95% CI: 44.7, 67.9) and 56.3% (95% CI: 43.1, 66.7), respectively, thus supporting the need for two doses of vaccine in previously unvaccinated children.
[d] In study participants who received 2 doses in year 1 and placebo in year 2, efficacy in year 2 was 56.2% (95% CI: 30.5, 72.7) and 44.8% (95% CI: 18.2, 62.9), respectively, in D153-P501, thus supporting the need for second-season revaccination.
[e] The primary circulating strain was antigenically dissimilar from the H3N2 strain represented in the vaccine; efficacy against the mismatched A/H3N2 strain was 85.9% (95% CI: 75.3, 91.9).

Table 2 FLUENZ Relative Efficacy in Active-controlled Paediatric Studies with Injectable Influenza Vaccine

Study Number	Region	Age Range[a]	Number of Study Participants	Influenza Season	Improved Efficacy (95% CI)[b] Matched strains	Improved Efficacy (95% CI)[b] All strains regardless of match
MI-CP111	USA, Europe, Asia/Oceania	6 to 59 M	7,852	2004-2005	44.5% (22.4, 60.0) fewer cases than injectable	54.9% (45.4, 62.9)[c] fewer cases than injectable
D153-P514	Europe	6 to 71 M	2,085	2002-2003	52.7% (21.6, 72.2) fewer cases than injectable	52.4% (24.6, 70.5)[d] fewer cases than injectable
D153-P515	Europe	6 to 17 Y	2,211	2002-2003	34.7% (3.9, 56.0) fewer cases than injectable	31.9% (1.1, 53.5) fewer cases than injectable

[a] M = months. Y = years. Age range as described in the protocol for the study.
[b] Reduction in culture-confirmed influenza illness relative to injectable influenza vaccine.
[c] FLUENZ demonstrated 55.7% (39.9, 67.6) fewer cases than injectable influenza vaccine in 3,659 infants and toddlers 6-23 months of age and 54.4% (41.8, 64.5) fewer cases in 4,166 children 24-59 months of age.
[d] FLUENZ demonstrated 64.4% (1.4, 88.8) fewer cases than injectable influenza vaccine in 476 infants and toddlers 6-23 months of age and 48.2% (12.7, 70.0) fewer cases in 1,579 children 24-71 months of age.

Adult studies
Several studies against placebo have shown that FLUENZ may have some efficacy in adults. However, a conclusion on clinical benefit of this vaccine in adults could not be made given that results observed in some studies versus injectable influenza vaccines were suggestive of a lower efficacy of FLUENZ.

5.2 Pharmacokinetic properties

Not applicable.

5.3 Preclinical safety data

Non-clinical data with FLUENZ reveal no special hazard for humans based on conventional non-clinical studies of repeated dose toxicity, reproduction and developmental toxicity, local tolerance, and neurovirulence.

6. PHARMACEUTICAL PARTICULARS

6.1 List of excipients

Sucrose
Dibasic potassium phosphate
Monobasic potassium phosphate
Gelatin (porcine, Type A)
Arginine hydrochloride
Monosodium glutamate monohydrate
Water for injections

6.2 Incompatibilities

In the absence of compatibility studies, this medicinal product must not be mixed with other medicinal products.

6.3 Shelf life

18 weeks.

6.4 Special precautions for storage

Store in a refrigerator (2°C – 8°C).

Do not freeze.

Protect from light.

Before use, the vaccine may be taken out of the refrigerator, without being replaced, for a maximum period of 12 hours at a temperature not above 25°C. If the vaccine has not been used after this 12-hour period, it should be disposed of.

6.5 Nature and contents of container

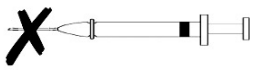
FLUENZ is supplied as a 0.2 ml suspension in a single-use nasal applicator (Type 1 glass), with nozzle (polypropylene with polyethylene transfer valve), nozzle tip-protector cap (synthetic rubber), plunger rod, plunger-stopper (butyl rubber), and a dose-divider clip.

Pack size of 10.

6.6 Special precautions for disposal and other handling

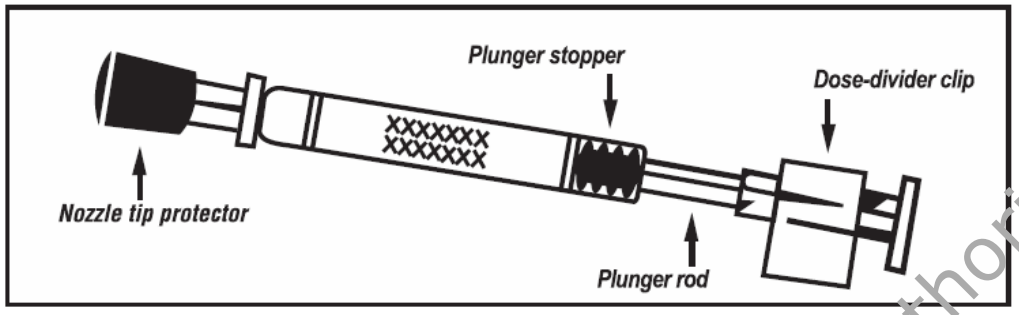
Administration
FLUENZ IS FOR NASAL USE ONLY.

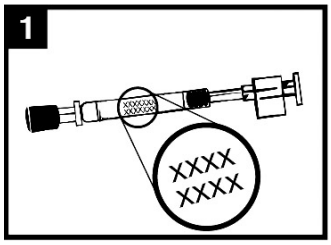
- DO NOT USE WITH A NEEDLE. Do not inject.



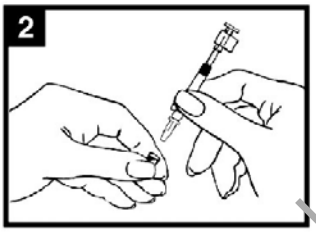
- FLUENZ is administered as a divided dose in both nostrils.
- After administering half of the dose in one nostril, administer the other half of the dose in the other nostril immediately or shortly thereafter.
- The patient can breathe normally while the vaccine is being administered – there is no need to actively inhale or sniff.
- Refer to the FLUENZ administration diagram (Figure 1) for step-by-step administration instructions.

Figure 1 FLUENZ Administration

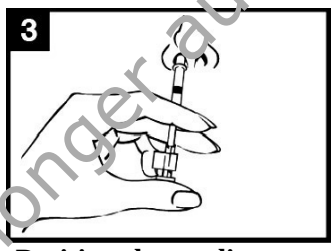




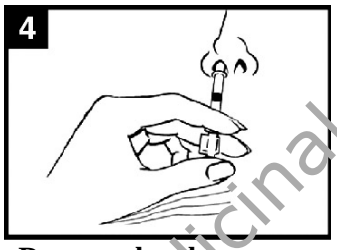
Check expiry date
Product must be used before date on applicator label.



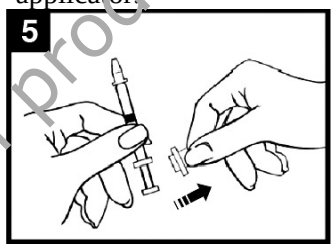
Prepare the applicator
Remove rubber tip protector. Do not remove dose-divider clip at the other end of the applicator.



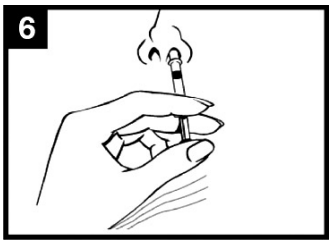
Position the applicator
With the patient in an upright position, place the tip just inside the nostril to ensure FLUENZ is delivered into the nose.



Depress the plunger
With a single motion, depress plunger **as rapidly as possible** until the dose-divider clip prevents you from going further.



Remove dose-divider clip
For administration in the other nostril, pinch and remove the dose-divider clip from plunger.



Spray in other nostril
Place the tip just **inside the other nostril** and with a single motion, depress plunger **as rapidly as possible** to deliver remaining vaccine.

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

7. MARKETING AUTHORISATION HOLDER

MedImmune, LLC
Lagelandseweg 78
6545 CG Nijmegen
Netherlands
(Tel) +31 24 371 7310

8. MARKETING AUTHORISATION NUMBER(S)

EU/1/10/661/002

9. DATE OF FIRST AUTHORISATION/RENEWAL OF THE AUTHORISATION

Date of first authorisation: 27 January 2011

10. DATE OF REVISION OF THE TEXT

Detailed information on this medicinal product is available on the website of the European Medicines Agency http://www.ema.europa.eu/.

ANNEX II

A. MANUFACTURERS OF THE BIOLOGICAL ACTIVE SUBSTANCE(S) AND MANUFACTURER RESPONSIBLE FOR BATCH RELEASE

B. CONDITIONS OR RESTRICTIONS REGARDING SUPPLY AND USE

C. OTHER CONDITIONS AND REQUIREMENTS OF THE MARKETING AUTHORISATION

D. CONDITIONS OR RESTRICTIONS WITH REGARD TO THE SAFE AND EFFECTIVE USE OF THE MEDICINAL PRODUCT

A. MANUFACTURERS OF THE BIOLOGICAL ACTIVE SUBSTANCE(S) AND MANUFACTURER RESPONSIBLE FOR BATCH RELEASE

Name and address of the manufacturers of the biological active substance(s)

MedImmune, LLC
297 North Bernardo Avenue,
Moutain View
California, 94043
USA

MedImmune, LLC
3055 Patrick Henry Drive
Santa Clara
California, 95054
USA

MedImmune, UK Limited
Plot 6, Renaissance Way, Boulevard Industry Park, Speke
Liverpool
L24 9JW
UK

Name and address of the manufacturer responsible for batch release

MedImmune, UK Limited
Plot 6, Renaissance Way, Boulevard Industry Park, Speke
Liverpool
L24 9JW
UK

The printed package leaflet of the medicinal product must state the name and address of the manufacturer responsible for the release of the concerned batch.

B. CONDITIONS OR RESTRICTIONS REGARDING SUPPLY AND USE

Medicinal product subject to medical prescription.

- **Official batch release**

In accordance with Article 114 of Directive 2001/83/EC, the official batch release will be undertaken by a state laboratory or a laboratory designated for that purpose.

C. OTHER CONDITIONS AND REQUIREMENTS OF THE MARKETING AUTHORISATION

- **Periodic Safety Update Reports**

The marketing authorisation holder shall submit periodic safety update reports for this product in accordance with the requirements set out in the list of Union reference dates (EURD list) provided for under Article 107c(7) of Directive 2001/83/EC and published on the European medicines web-portal.

D. CONDITIONS OR RESTRICTIONS WITH REGARD TO THE SAFE AND EFFECTIVE USE OF THE MEDICINAL PRODUCT

- **Risk Management Plan (RMP)**

The MAH shall perform the required pharmacovigilance activities and interventions detailed in the agreed RMP presented in Module 1.8.2 of the Marketing Authorisation and any agreed subsequent updates of the RMP.

An updated RMP shall be submitted annually until renewal.

When the submission of a PSUR and the update of a RMP coincide, they should be submitted at the same time.

In addition, an updated RMP should be submitted:

- At the request of the European Medicines Agency;
- Whenever the risk management system is modified, especially as the result of new information being received that may lead to a significant change to the benefit/risk profile or as the result of an important (pharmacovigilance or risk minimisation) milestone being reached.

ANNEX III

LABELLING AND PACKAGE LEAFLET

A. LABELLING

PARTICULARS TO APPEAR ON THE OUTER PACKAGING

PACK SIZE OF 10 SINGLE-USE NASAL APPLICATORS (2 X 5 NASAL APPLICATORS)

1. NAME OF THE MEDICINAL PRODUCT

FLUENZ nasal spray suspension
Influenza vaccine (live attenuated, nasal)
2013/2014 season

2. STATEMENT OF ACTIVE SUBSTANCE(S)

Reassortant influenza virus* (live attenuated) of the following strains**:

A/California/7/2009 (H1N1)pdm09-like strain
(A/California/7/2009, MEDI 228029) $10^{7.0\pm0.5}$ FFU***

A/Victoria/361/2011 (H3N2)-like strain
(A/Texas/50/2012, MEDI 237514) $10^{7.0\pm0.5}$ FFU***

B/Massachusetts/2/2012-like strain
(B/Massachusetts/2/2012, MEDI 237751) $10^{7.0\pm0.5}$ FFU***

.....per 0.2 ml dose

* propagated in fertilised hens' eggs from healthy chicken flocks.
** produced in VERO cells by reverse genetic technology.
*** fluorescent focus units.

This vaccine complies with the WHO recommendations (Northern Hemisphere) and EU decision for the 2013/2014 season.

3. LIST OF EXCIPIENTS

Contains also: sucrose, dibasic potassium phosphate, monobasic potassium phosphate, gelatin (porcine, Type A), arginine hydrochloride, monosodium glutamate monohydrate, water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

Nasal spray, suspension
10 single-use nasal applicators (0.2 ml each)

5. METHOD AND ROUTE(S) OF ADMINISTRATION

For nasal use only. Do not inject.
Read the package leaflet before use.

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

Store in a refrigerator.

Do not freeze.

Protect from light.

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

Please read the package leaflet for disposal of medicines no longer required.

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

MedImmune, LLC
Lagelandseweg 78
6545 CG Nijmegen
Netherlands

12. MARKETING AUTHORISATION NUMBER(S)

EU/1/10/661/002 < – 10 sprayers>

13. BATCH NUMBER

Lot

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription.

15. INSTRUCTIONS ON USE

16. INFORMATION IN BRAILLE

Justification for not including Braille accepted

MINIMUM PARTICULARS TO APPEAR ON BLISTERS OR STRIPS

PACK SIZE OF 5 SINGLE-USE NASAL APPLICATORS

1. NAME OF THE MEDICINAL PRODUCT

FLUENZ nasal spray suspension
Influenza vaccine (live attenuated, nasal)
2013/2014 season

2. NAME OF THE MARKETING AUTHORISATION HOLDER

MedImmune, LLC

3. EXPIRY DATE

EXP

4. BATCH NUMBER

Lot

5. OTHER

For nasal use only. Do not inject.

5 single-use nasal applicators (0.2 ml each)

Store in a refrigerator. Do not freeze.

MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS

SINGLE-USE NASAL APPLICATOR

1. NAME OF THE MEDICINAL PRODUCT AND ROUTE(S) OF ADMINISTRATION

FLUENZ
Influenza vaccine
2013/2014 season

2. METHOD OF ADMINISTRATION

For nasal use only.

3. EXPIRY DATE

EXP

4. BATCH NUMBER

Lot

5. CONTENTS BY WEIGHT, BY VOLUME OR BY UNIT

0.2 ml

6. OTHER

B. PACKAGE LEAFLET

Package Leaflet: Information for the user

Fluenz nasal spray suspension
Influenza vaccine (live attenuated, nasal)

▼This medicine is subject to additional monitoring. This will allow quick identification of new safety information. You can help by reporting any side effects you may get. See the end of section 4 for how to report side effects.

Read all of this leaflet carefully before the vaccine is given because it contains important information for you or your child.

- Keep this leaflet. You may need to read it again.
- If you have any further questions, ask your doctor, nurse or pharmacist.
- This vaccine has been prescribed for you or your child only. Do not pass it on to others.
- If any of the side effects gets serious, talk to your doctor, nurse or pharmacist. This includes any possible side effects not listed in this leaflet. See section 4.

What is in this leaflet:

1. What Fluenz is and what it is used for

Fluenz is a vaccine to prevent influenza (flu). It is used in children and adolescents 24 months to less than 18 years of age.

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

Fluenz vaccine viruses are grown in chicken eggs. The vaccine targets three strains of influenza virus each year, following the annual recommendations by the World Health Organisation.

2. What you need to know before you are given Fluenz

You will not be given Fluenz

- **if you are allergic** to eggs, egg proteins, gentamicin, or gelatin or any of the other ingredients of Fluenz (listed in section 6 "Contents of the pack and other information"). For signs of allergic reactions, see section 4 "Possible side effects".
- if you have a **blood disorder** or a **cancer** that **affects the immune system**.
- if you have been **told by your doctor** that you have **a weakened immune system** as a result of a disease, medicine, or other treatment.
- **if you are under 18 years** of age and **already taking acetylsalicylic acid** *(a substance present in many medicines used to relieve pain and lower fever)*. This is because of the risk of a very rare but serious disease *(Reye's syndrome)*.

If any of these apply, **tell your doctor, nurse or pharmacist**.

Warnings and precautions

Talk to your doctor, nurse or pharmacist before vaccination:
- if the **child is less than 24 months of age**. Children less than 24 months of age should not receive this vaccine because of the risk of side effects.
- if you have **severe asthma** or are currently wheezing.
- if you are in **close contact with someone with a severely weakened immune system** (for example, a bone marrow transplant patient needing isolation).

If any of these apply, **tell your doctor, nurse or pharmacist before vaccination**. He or she will decide if Fluenz is suitable for you.

Other medicines, other vaccines and Fluenz

Tell your doctor, nurse or pharmacist if the person being vaccinated is taking, has recently taken or might take any other medicines.
- **Do not give acetylsalicylic acid to children aged less than 18 years** for 4 weeks after vaccination with Fluenz unless your doctor, nurse or pharmacist tells you otherwise. This is because of the risk of Reye's syndrome, a very rare but serious disease that can affect the brain and liver.
- **It is recommended that Fluenz is not given** at the same time as influenza-specific **antiviral medicines**. This is because the vaccine may work less effectively.

Your doctor, nurse or pharmacist will decide if Fluenz can be given at the same time as other vaccines.

Pregnancy and breast-feeding
- If you are **pregnant**, think you may be pregnant, plan to become pregnant soon or are breast feeding, **tell your doctor, nurse or pharmacist before receiving this vaccine**. Fluenz is **not recommended** for women who are pregnant or are breast-feeding.

3. How Fluenz is given

Fluenz will be administered under the supervision of a doctor, nurse or pharmacist.

Fluenz must only be used as a nasal spray.

Fluenz must not be injected.

Fluenz will be given as a spray in each nostril. You can breathe normally while you are given Fluenz. You do not need to actively inhale or sniff.

Dosage
- **The recommended dose** for children and adolescents is 0.2 ml Fluenz, administered as 0.1 ml in each nostril.
- **Children who have not previously had an influenza vaccine** will receive a second, follow-up dose after an interval of at least 4 weeks. Follow your doctor, nurse or pharmacist's instructions about when your child should return for the second dose.

4. Possible side effects

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

Ask your doctor, nurse or pharmacist if you want more information about possible side effects from Fluenz.

Some side effects may be serious

Very rare
(may affect up to 1 in 1,000,000 people):
- severe allergic reaction: signs of a severe allergic reaction may include shortness of breath and swelling of the face or tongue.

Tell your doctor straight away or seek urgent medical care if you experience any of the effects above.

Other possible side effects of Fluenz

Very common
(may affect more than 1 in 10 people):
- runny or stuffy nose
- reduced appetite
- weakness
- headache

Common
(may affect up to 1 in 10 people):
- fever
- muscle aches

Uncommon
(may affect up to 1 in 100 people):
- rash
- nose bleed
- allergic reactions

Reporting of side effects
If you get any side effects, talk to your doctor, nurse or pharmacist. This includes any possible side effects not listed in this leaflet. You can also report side effects directly via the national reporting system listed in Appendix V. By reporting side effects you can help provide more information on the safety of this medicine.

5. How to store Fluenz

Keep this vaccine out of the sight and reach of children.

Do not use Fluenz after the expiry date which is stated on the applicator label after the letters EXP.

Store in a refrigerator (2°C to 8°C). Do not freeze.

Protect from light.

Before use, the vaccine may be taken out of the refrigerator, without being replaced, for a maximum period of 12 hours at a temperature not above 25°C. If the vaccine has not been used after this 12-hour period, it should be disposed of.

Any unused product or waste material should be disposed of in accordance with local requirements for medical waste. Do not throw away any medicines via wastewater or household waste. Ask your pharmacist how to throw away medicines you no longer use. These measures will help protect the environment.

6. Contents of the pack and other information

What Fluenz contains

The active substances are:
Reassortant influenza virus* (live attenuated) of the following strains**:

A/California/7/2009 (H1N1)pdm09-like strain (A/California/7/2009, MEDI 228029)	$10^{7.0\pm0.5}$ FFU***
A/Victoria/361/2011 (H3N2)-like strain (A/Texas/50/2012, MEDI 237514)	$10^{7.0\pm0.5}$ FFU***
B/Massachusetts/2/2012-like strain (B/Massachusetts/2/2012, MEDI 237751)	$10^{7.0\pm0.5}$ FFU***

……..……per 0.2 ml dose

* propagated in fertilised hens' eggs from healthy chicken flocks.
** produced in VERO cells by reverse genetic technology. This product contains genetically modified organisms (GMOs).
*** fluorescent focus units

This vaccine complies with the WHO (World Health Organisation) recommendations (Northern Hemisphere) and EU decision for the 2013/2014 season.

The other ingredients are sucrose, dibasic potassium phosphate, monobasic potassium phosphate, gelatin (porcine, Type A), arginine hydrochloride, monosodium glutamate monohydrate and water for injections.

What Fluenz looks like and contents of the pack

This vaccine is presented as a nasal spray suspension in a single-use nasal applicator (0.2 ml) in a pack size of 10.

The suspension is a colourless to pale yellow liquid that is clear to slightly cloudy. Small white particles may be present.

Marketing Authorisation Holder and Manufacturer

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Manufacturer: MedImmune, UK Limited, Plot 6, Renaissance Way, Boulevard Industry Park, Speke, Liverpool, L24 9JW, UK

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This leaflet was last revised in.

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

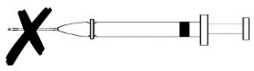
Fluenz is a trademark of MedImmune, LLC.

Instructions for health professionals

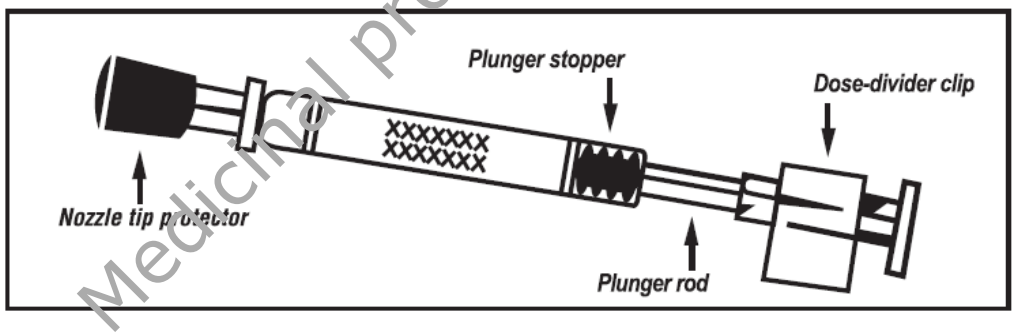
The following information is intended for medical or healthcare professionals only:

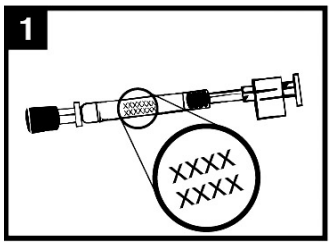
Fluenz is for nasal use only.

- **Do not use with a needle.** Do not inject.

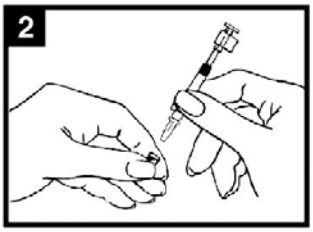


- Fluenz is administered as a divided dose in both nostrils as described below. (See also, *How Fluenz is given*, in section 3).
- After administering half of the dose in one nostril, administer the other half of the dose in the other nostril immediately or shortly thereafter.
- The patient can breathe normally while the vaccine is being administered – there is no need to actively inhale or sniff.

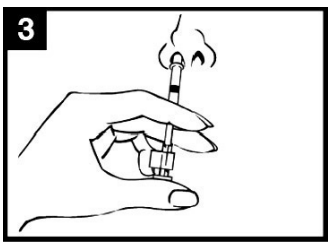




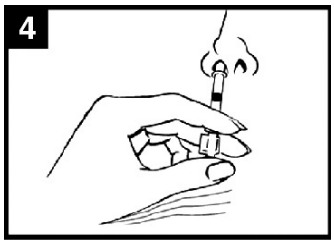
Check expiry date
Product must be used before date on applicator label.



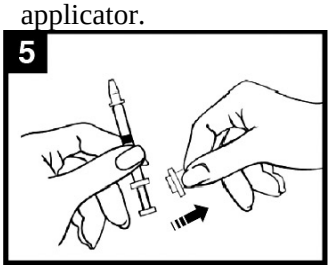
Prepare the applicator
Remove rubber tip protector. Do not remove dose-divider clip at the other end of the applicator.



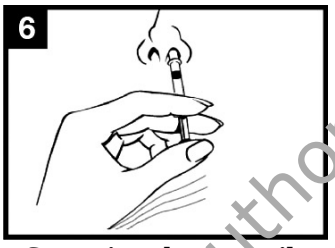
Position the applicator
With the patient in an upright position, place the tip just inside the nostril to ensure Fluenz is delivered into the nose.



Depress the plunger
With a single motion, depress plunger **as rapidly as possible** until the dose-divider clip prevents you from going further.



Remove dose-divider clip
For administration in the other nostril, pinch and remove the dose-divider clip from plunger.



Spray in other nostril
Place the tip just **inside the other nostril** and with a single motion, depress plunger **as rapidly as possible** to deliver remaining vaccine.

See section 5 for advice on storage and disposal.