

ANNEX I

SUMMARY OF PRODUCT CHARACTERISTICS

1. NAME OF THE MEDICINAL PRODUCT

Fluenz nasal spray suspension
Influenza vaccine (live, nasal)

2. QUALITATIVE AND QUANTITATIVE COMPOSITION

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

A/Victoria/4897/2022 (H1N1)pdm09-like strain
(A/Norway/31694/2022, MEDI 369815) $10^{7.0 \pm 0.5}$ FFU***

A/Thailand/8/2022 (H3N2)-like strain
(A/Thailand/8/2022, MEDI 370626) $10^{7.0 \pm 0.5}$ FFU***

B/Austria/1359417/2021-like strain
(B/Austria/1359417/2021, MEDI 355292) $10^{7.0 \pm 0.5}$ 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 2024/2025 season.

The vaccine may contain traces of the following substances: egg proteins (e.g. ovalbumin) and gentamicin. The maximum amount of ovalbumin is less than 0.024 micrograms per 0.2 ml dose (0.12 micrograms per ml), see section 4.3.

For the full list of excipients, see section 6.1.

3. PHARMACEUTICAL FORM

Nasal spray suspension

Colourless to pale yellow, clear to opalescent suspension with a pH of approximately 7.2. Small white particles may be present.

4. CLINICAL PARTICULARS

4.1 Therapeutic indications

Fluenz is indicated for active immunisation for the prevention of influenza disease in children and adolescents from 2 years to less than 18 years of age.

Fluenz should be used in accordance with official recommendations.

4.2 Posology and method of administration

Posology

Children and adolescents from 2 years to <18 years of age

The recommended dose is 0.2 ml, administered as 0.1 ml in each nostril.

For children 2 to 8 years of age who have not previously been vaccinated against seasonal influenza, a second dose should be given after an interval of at least 4 weeks.

Infants and toddlers <2 years of age

Fluenz should not be used in infants and toddlers below 2 years of age because of safety concerns regarding increased rates of hospitalisation and wheezing in this population (see section 4.8).

Method of administration

For nasal use only.

Fluenz should under no circumstances be injected.

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.

For detailed instructions on preparation and administration, see section 6.6.

4.3 Contraindications

- Hypersensitivity to the active substances or, to any of the excipients listed in section 6.1 or to gentamicin (a possible trace residue).
- Severe allergic reaction (e.g. anaphylaxis) to eggs or to egg proteins (e.g. ovalbumin).
- Children and adolescents with clinical immunodeficiency 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 (see section 4.5).

4.4 Special warnings and precautions for use

Traceability

In order to improve the traceability of biological medicinal products, the name and the batch number of the administered product should be clearly recorded.

Hypersensitivity and anaphylaxis

As with most vaccines, appropriate medical treatment and supervision should always be readily available to manage an anaphylactic event or serious hypersensitivity event following the administration of Fluenz.

Concurrent illness and chronic conditions.

Fluenz should not be administered to children and adolescents with severe asthma or who are currently wheezing, because these individuals have not been adequately studied in clinical studies (see section 4.8).

Administration of Fluenz should be postponed in individuals suffering from severe acute febrile illness or acute infection. The presence of a minor infection and/or low-grade fever should not result in the deferral of vaccination.

Vaccination should also be postponed in individuals with nasal blockage, due to the potential for reduced vaccine uptake and the absence of efficacy data in this population. The presence of mild symptoms of a common cold without nasal blockage should not result in the deferral of vaccination.

Immunocompromised individuals

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. The effectiveness of Fluenz preventing influenza disease in immunocompromised individuals has not been evaluated (see section 4.8).

General

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

Fluenz must not be administered to children and adolescents receiving salicylate therapy (see section 4.3). Salicylates should not be used in children and adolescents 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.

Fluenz can be administered concomitantly with the live attenuated vaccines: measles, mumps, rubella, varicella, and orally-administered poliovirus.

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 based on clinical judgement.

4.6 Fertility, pregnancy and lactation

Pregnancy

There is a moderate amount of data from the use of Fluenz in pregnant women. There was no evidence of significant maternal adverse outcomes in 138 pregnant women who had a record of receiving Fluenz in a US-based health insurance claims database.

In more than 300 case reports in the AstraZeneca safety database of vaccine administration to pregnant women, no unusual patterns of pregnancy complications or foetal outcomes were observed.

While animal studies do not indicate direct or indirect harmful effects with respect to reproductive toxicity, and post-marketing data offer some reassurance in the event of inadvertent administration of the vaccine, Fluenz is not recommended during pregnancy.

Breast-feeding

Limited available evidence suggests that Fluenz is not excreted in breastmilk. However, because there are limited data to assess the effects on the breast-fed infant and as some viruses are excreted in human milk, Fluenz should not be used during breast-feeding.

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

Fluenz has no or negligible influence on the ability to drive and use machines.

4.8 Undesirable effects

Summary of the safety profile

Safety data regarding use of Fluenz are based on data from Fluenz clinical studies in over 29 000 children and adolescents 2 to 17 years of age, Fluenz post-authorisation safety studies in over 84 000 children and adolescents 2 to 17 years of age, and data from influenza vaccine (live attenuated, nasal) tetravalent clinical studies in over 2 000 children and adolescents 2 to 17 years of age. Additional experience has occurred with marketed use of Fluenz and/or influenza vaccine (live attenuated, nasal) tetravalent.

In clinical studies, the safety profile of Fluenz and influenza vaccine (live attenuated, nasal) tetravalent were similar.

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

Tabulated list of adverse reactions

Adverse reaction frequencies are organised by MedDRA System Organ Class (SOC). Within each SOC, adverse reactions are arranged by decreasing frequency and then by decreasing seriousness. Frequencies of occurrence of adverse reactions are defined as: very common ($\geq 1/10$); common ($\geq 1/100$ to $< 1/10$); uncommon ($\geq 1/1\ 000$ to $< 1/100$); rare ($\geq 1/10\ 000$ to $< 1/1\ 000$); very rare ($< 1/10\ 000$) and not known (cannot be estimated from available data).

Table 1 Adverse reactions

| MedDRA SOC | Adverse reaction | Frequency |
|--|--|------------------------|
| Immune system disorders | Hypersensitivity reactions (including facial oedema, urticaria) | Uncommon |
| | Anaphylactic reactions | Very rare |
| Metabolism and nutrition disorders | Decreased appetite | Very common |
| Nervous system disorders | Headache | Common |
| | Guillain-Barré syndrome | Not known ^a |
| Respiratory, thoracic and mediastinal disorders | Nasal congestion/rhinorrhoea | Very common |
| | Epistaxis | Uncommon |
| Skin and subcutaneous tissue disorders | Rash | Uncommon |
| Musculoskeletal and connective tissue disorders | Myalgia | Common |
| General disorders and administration site conditions | Malaise | Very common |
| | Pyrexia | Common |
| Congenital, familial, and genetic disorders | Exacerbation of symptoms of Leigh syndrome (mitochondrial encephalomyopathy) | Not known ^a |

^a Post-marketing experience

Paediatric population

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). Most hospitalisations were due to gastrointestinal and respiratory tract infections and occurred more than 6 weeks post vaccination. 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 2 years and older. Fluenz is not indicated for use in infants and toddlers younger than 2 years (see section 4.2).

Concurrent illness and chronic conditions

In a study (D153-P515) of children 6 to 17 years of age with asthma (Fluenz: n=1 114, trivalent injectable influenza vaccine: n=1 115), there were no significant differences between treatment groups in the incidence of asthma exacerbations, mean peak expiratory flow rate, asthma symptom scores, or night-time awakening scores. The incidence of wheezing within 15 days after vaccination was lower in Fluenz recipients relative to inactivated vaccine recipients (19.5% vs. 23.8%, P=0.02).

In a study of children and adolescents 9 to 17 years of age with moderate to severe asthma (Fluenz: n=24, placebo: n=24), the primary safety criterion, change in percent predicted forced expiratory volume in 1 second (FEV₁) measured before and after vaccination, did not differ between treatment arms.

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, or other underlying chronic medical conditions are limited. In a study of adults 60 years of age and older (n=3 242), a high percentage of individuals had underlying chronic medical conditions (87%), including cardiac disorders (15%), respiratory, thoracic and mediastinal disorders (13%), and diabetes mellitus (9.6%). The safety profile of Fluenz in these individuals was comparable to the safety profile observed in individuals without these conditions.

Immunocompromised individuals

In HIV-infected children (n=24) and HIV-negative children (n=25) 1 through 7 years of age, and in HIV-infected children and adolescents 5 through 17 years of age receiving stable anti-retroviral therapy (Fluenz: n=122, trivalent injectable vaccine: n=121), the frequency and duration of vaccine virus shedding were comparable to that seen in healthy individuals. No adverse effects on HIV viral load or CD4 counts were identified following Fluenz administration.

Twenty mild to moderately immunocompromised children and adolescents 5 through 17 years of age (receiving chemotherapy and/or radiation therapy or who had recently received chemotherapy) were randomised 1:1 to Fluenz or placebo. Frequency and duration of vaccine virus shedding in these immunocompromised children and adolescents were comparable to that seen in healthy children and adolescents.

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

Overdose with Fluenz is unlikely due to its presentation as a single pre-filled nasal applicator. Administration of a higher than recommended dose of Fluenz was reported rarely and the adverse reaction profile was comparable to that observed with the recommended dose of Fluenz.

5. PHARMACOLOGICAL PROPERTIES

5.1 Pharmacodynamic properties

Pharmacotherapeutic group: Vaccines, influenza vaccines, ATC code: J07BB03

Mechanism of action

Fluenz is a trivalent vaccine that contains three *cold-adapted (ca)*; *temperature-sensitive (ts)*; and *attenuated (att)* influenza virus strains, an A/(H1N1) strain, an A/(H3N2) strain, and a B strain from the Victoria lineage. After intranasal administration, Fluenz replicates in the nasopharynx and induces immune responses against the three influenza strains contained in the vaccine.

Clinical efficacy

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 2 and 3 for a summary of efficacy results in the paediatric population.

Table 2 Fluenz efficacy after 2 doses against culture-confirmed influenza illness in placebo controlled paediatric studies

| Study number ^a | Region | Age range ^b | Number of study participants | Influenza season | Efficacy (95% CI) matched strains | Efficacy (95% CI) 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) |
| 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 |
| D153-P513 | Asia/ Oceania | 6 to 35 M | 1 041 | 2002 | 62.2% (43.6, 75.2) | 48.6% (28.8, 63.3) |
| D153-P522 | Europe, Asia/ Oceania, Latin America | 11 to 23 M | 1 150 | 2002-2003 | 78.4% (50.9, 91.3) | 63.8% (36.2, 79.8) |
| D153-P501 | Asia/ Oceania | 12 to 35 M | 2 764 | 2000-2001 | 72.9% (62.8, 80.5) | 70.1% (60.9, 77.3) |
| AV006 | USA | 15 to 71 M | 1 259 ^c | 1996-1997 | 93.4% (87.5, 96.5) | N/A |

^a Per protocol population, except where noted otherwise; none of the participants were previously vaccinated against influenza

^b Age range as described in the study protocol. M=months.

^c Data presented are for study participants who received two doses (ITT population).

In clinical studies AV006 and D153-P504, the efficacy of a single dose of Fluenz was evaluated in previously unvaccinated children aged 15-71 months (n=288) and aged 6-35 months (n=1 877). The efficacy against culture-confirmed influenza caused by any matched strain was 88.8% (95% CI: 64.5, 96.5) and 57.7% (95% CI: 44.7, 67.9), respectively.

Fluenz efficacy estimates against all matched strains in children who received 2 doses in year 1 and revaccination in year 2 was 100% (95% CI: 38.2, 100; n=1 110) in AV006, 84.3% (95% CI: 70.1, 92.4; n=1 265) in D153-P501, 88.7% (95% CI: 82.0, 93.2; n=1 090) in D153-P502 and 73.6% (95% CI: 33.3, 91.2; n=680) in D153-P504.

Fluenz efficacy estimates against all matched strains in children who received 2 doses in year 1 and placebo in year 2 were lower: 56.2% (95% CI: 30.5, 72.7; n=1 253) in D153-P501 and 57.0% (95% CI: 6.1, 81.7; n=718) in D153-P504.

Table 3 Fluenz relative efficacy against culture-confirmed influenza illness in active-controlled paediatric studies with injectable influenza vaccine

| Study number ^a | Region | Age range ^b | Number of study participants | Influenza season | Improved efficacy (95% CI) matched strains | Improved efficacy (95% CI) all strains regardless of match |
|---------------------------|---------------------------|------------------------|------------------------------|------------------|--|--|
| MI-CP111 | USA, Europe, Asia/Oceania | 6 to 59 M | 7 852 ^c | 2004-2005 | 44.5% (22.4, 60.6) | 54.9% (45.4, 62.9) ^d |
| D153-P514 | Europe | 6 to 71 M | 2 085 ^e | 2002-2003 | 52.7% (21.6, 72.2) | 52.4% (24.6, 70.5) ^f |
| D153-P515 | Europe | 6 to 17 Y | 2 211 ^g | 2002-2003 | 34.7% (3.9, 56.0) | 31.9% (1.1, 53.5) |

^a Per protocol population

^b Age range as described in the study protocol. M=months. Y=years.

^c Data presented are for study participants who received two doses if unvaccinated or vaccination history was unknown, and one dose for those previously vaccinated.

^d Fluenz demonstrated 55.7% (39.9, 67.6) fewer cases than injectable influenza vaccine in 3 686 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.

^e Data presented are for study participants with a history of recurrent respiratory tract infections who received two doses of intranasal influenza vaccine compared to those who received two doses of injectable influenza vaccine.

^f 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 1 609 children 24-71 months of age.

^g Data presented are for study participants clinically diagnosed with asthma who received one dose of intranasal influenza vaccine compared to those who received one dose of injectable influenza vaccine.

5.2 Pharmacokinetic properties

Not applicable.

5.3 Preclinical safety data

Non-clinical data 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
Dipotassium phosphate
Potassium dihydrogen phosphate
Gelatin
Arginine hydrochloride
Monosodium glutamate monohydrate
Water for injections

6.2 Incompatibilities

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

6.3 Shelf life

15 weeks

6.4 Special precautions for storage

Store in a refrigerator (2°C – 8°C).

Do not freeze.

Keep the nasal applicator in the outer carton in order to protect from light.

Before use, the vaccine may be taken out of the refrigerator once for a maximum period of 12 hours at a temperature not above 25°C. Stability data indicate that the vaccine components are stable for 12 hours when stored at temperatures from 8°C to 25°C. At the end of this period, Fluenz should be used immediately or discarded.

6.5 Nature and contents of container

Fluenz is supplied as a nasal spray suspension (0.2 ml) 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 sizes of 1 or 10 nasal applicators.

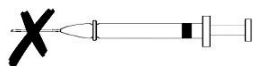
Not all pack sizes may be marketed.

6.6 Special precautions for disposal and other handling

Administration

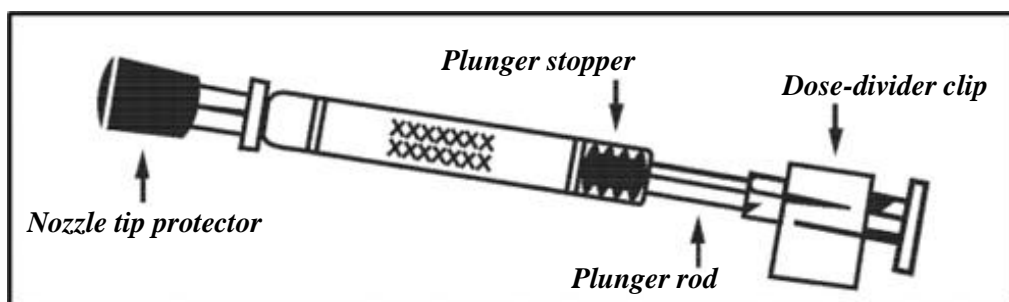
Fluenz is for single nasal use only.

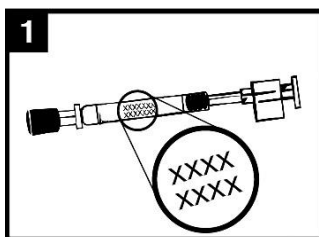
- Do not use with a needle. Do not inject.



- Do not use Fluenz if the expiry date has passed or the sprayer appears damaged, for example, if the plunger is loose or displaced from the sprayer or if there are any signs of leakage.
- Check the appearance of the vaccine before administration. The suspension should be colourless to pale yellow, clear to opalescent. Small white particles may be present.
- 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.

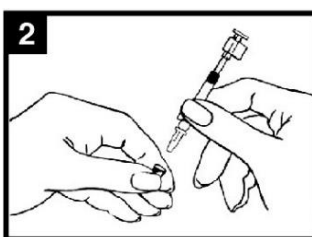
Figure 1: Fluenz Administration





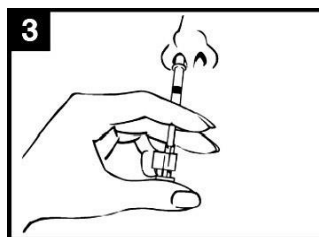
1 Check expiry date

Product must not be used after date on applicator label.



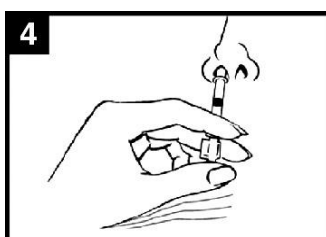
2 Prepare the applicator

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



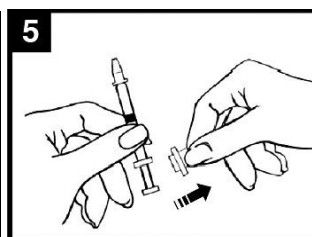
3 Position the applicator

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



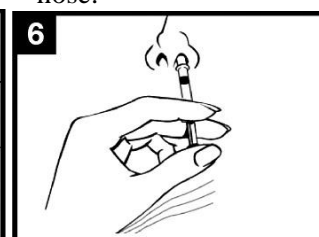
4 Depress the plunger

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



5 Remove dose-divider clip

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



6 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

AstraZeneca AB
SE-151 85 Södertälje
Sweden

8. MARKETING AUTHORISATION NUMBER(S)

EU/1/24/1816/001
EU/1/24/1816/002

9. DATE OF FIRST AUTHORISATION/RENEWAL OF THE AUTHORISATION

Date of first authorisation: 03 June 2024

10. DATE OF REVISION OF THE TEXT

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

ANNEX II

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

Name and address of the manufacturer of the biological active substances

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

Name and address of the manufacturers responsible for batch release

AstraZeneca Nijmegen B.V.,
Lagelandseweg 78
Nijmegen, 6545CG
Netherlands

B. CONDITIONS OR RESTRICTIONS REGARDING SUPPLY AND USE

Medicinal products 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 (PSURs)**

The requirements for submission of PSURs for this medicinal product are set out in the list of Union reference dates (EURD list) provided for under Article 107c(7) of Directive 2001/83/EC and any subsequent updates 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 marketing authorisation holder (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 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

TRI-FOLD CARTON

1. NAME OF THE MEDICINAL PRODUCT

Fluenz nasal spray suspension
Influenza vaccine (live, nasal)
2024/2025 season

2. STATEMENT OF ACTIVE SUBSTANCE(S)

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

A/Victoria/4897/2022 (H1N1)pdm09-like strain
(A/Norway/31694/2022, MEDI 369815) $10^{7.0 \pm 0.5}$ FFU***

A/Thailand/8/2022 (H3N2)-like strain
(A/Thailand/8/2022, MEDI 370626) $10^{7.0 \pm 0.5}$ FFU***

B/Austria/1359417/2021-like strain
(B/Austria/1359417/2021, MEDI 355292) $10^{7.0 \pm 0.5}$ 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 2024/2025 season.

3. LIST OF EXCIPIENTS

Contains also: sucrose, dipotassium phosphate, potassium dihydrogen phosphate, gelatin, arginine hydrochloride, monosodium glutamate monohydrate, water for injections. See leaflet for further information.

4. PHARMACEUTICAL FORM AND CONTENTS

Nasal spray suspension

1 single-use nasal applicator (0.2 ml)

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.

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| 6. SPECIAL WARNING THAT THE MEDICINAL PRODUCT MUST BE STORED OUT OF THE SIGHT AND REACH OF CHILDREN |
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Keep out of the sight and reach of children.

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| 7. OTHER SPECIAL WARNING(S), IF NECESSARY |
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|-----------------------|
| 8. EXPIRY DATE |
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EXP

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|--------------------------------------|
| 9. SPECIAL STORAGE CONDITIONS |
|--------------------------------------|

Store in a refrigerator.

Do not freeze.

Keep the nasal applicator in the outer carton to protect from light.

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| 10. SPECIAL PRECAUTIONS FOR DISPOSAL OF UNUSED MEDICINAL PRODUCTS OR WASTE MATERIALS DERIVED FROM SUCH MEDICINAL PRODUCTS, IF APPROPRIATE |
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| 11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER |
|---|

AstraZeneca AB
SE-151 85 Södertälje
Sweden

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| 12. MARKETING AUTHORISATION NUMBER(S) |
|--|

EU/1/24/1816/001 1 nasal applicator.
EU/1/24/1816/002 10 nasal applicators.

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|-------------------------|
| 13. BATCH NUMBER |
|-------------------------|

Lot

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| 14. GENERAL CLASSIFICATION FOR SUPPLY |
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|--------------------------------|
| 15. INSTRUCTIONS ON USE |
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|-----------------------------------|
| 16. INFORMATION IN BRAILLE |
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Justification for not including Braille accepted.

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|---|
| 17. UNIQUE IDENTIFIER - 2D BARCODE |
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2D barcode carrying the unique identifier included.

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| 18. UNIQUE IDENTIFIER - HUMAN READABLE DATA |
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PC
SN
NN

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|---|
| MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS SINGLE-USE NASAL APPLICATOR |
|---|

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

Fluenz nasal spray suspension
Influenza vaccine
2024/2025 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, nasal)

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 you get any side effects, 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
2. What you need to know before you are given Fluenz
3. How Fluenz is given
4. Possible side effects
5. How to store Fluenz
6. Contents of the pack and other information

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 2 years to less than 18 years of age. Fluenz will help to protect against the virus strains contained in the vaccine, and other strains closely related to them.

How Fluenz works

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. Each year the vaccine targets three strains of influenza, 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 the active ingredients, gentamicin (trace residue from the manufacturing process), or any of the other ingredients of this vaccine listed in section 6.
- if you have ever had a severe allergic reaction to eggs or egg proteins. For signs of allergic reactions, see section 4.
- 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 (such as acute and chronic leukaemias, lymphoma, symptomatic HIV infection, cellular immune deficiencies, and high-dose corticosteroids).
- **if you are 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

Your doctor or nurse will make sure that appropriate medical treatment and supervision is readily available in case of a rare anaphylactic reaction (a very severe allergic reaction with symptoms such as difficulty in breathing, dizziness, a weak and rapid pulse and skin rash) following the administration.

Talk to your doctor, nurse or pharmacist before vaccination:

- if you have **severe asthma** or are currently wheezing.
- if you have a severe acute illness associated with fever or infection, or if you have nasal blockage. Your doctor may decide to delay your vaccination until your fever or nasal blockage is gone.
- if you are in **close contact with someone with a severely weakened immune system** (for example, a bone marrow transplant patient needing isolation).
- if you have any birth defects that affect the bones and tissues of the skull and face, which have not undergone surgical correction.

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

Children less than 2 years of age

Children less than 2 years of age should not receive this vaccine because of the risk of side effects.

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, including medicines that do not require a prescription.

- **Do not give acetylsalicylic acid** (a substance present in many medicines used to relieve pain and lower fever) **to children** 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**, such as oseltamivir and zanamivir. 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.

Driving and using machines

- Fluenz has no or negligible influence on the ability to drive and use machines.

3. How Fluenz is given

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.

Dose

The recommended dose for children and adolescents is 0.2 ml Fluenz, administered as 0.1 ml in each nostril. **Children 2 to 8 years of age 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 if and when your child should return for the second dose.

If you have any further questions on this vaccine, ask your doctor, nurse or pharmacist.

4. Possible side effects

Like all medicines, this vaccine 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 10 000 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
- feeling generally unwell

Common (may affect up to 1 in 10 people)

- fever
- muscle aches
- headache

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 this vaccine after the expiry date which is stated on the applicator label and the carton after the letters EXP.

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

Keep the nasal applicator in the outer carton in order to protect from light.

Before use, the vaccine may be taken out of the refrigerator once 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 discarded.

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 three strains**:

| | |
|--|------------------------------|
| A/Victoria/4897/2022 (H1N1)pdm09-like strain (A/Norway/31694/2022, MEDI 369815) | 10 ^{7.0±0.5} FFU*** |
|--|------------------------------|

| | |
|--|------------------------------|
| A/Thailand/8/2022 (H3N2)-like strain (A/Thailand/8/2022, MEDI 370626) | 10 ^{7.0±0.5} FFU*** |
|--|------------------------------|

| | |
|---|------------------------------|
| B/Austria/1359417/2021-like strain (B/Austria/1359417/2021, MEDI 355292) | 10 ^{7.0±0.5} 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 2024/2025 season.

The vaccine may contain traces of the following substances: egg proteins (e.g. ovalbumin) and gentamicin. The other ingredients are sucrose, dipotassium phosphate, potassium dihydrogen phosphate, gelatin, 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 (0.2 ml) in a single-use nasal applicator (Type 1 glass). Pack size of 1 and 10 nasal applicators. Not all pack sizes may be available in your country.

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

Marketing Authorisation Holder

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Södertälje
Sweden

Manufacturer

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This leaflet was last revised in.

Other sources of information

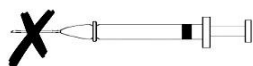
Detailed information on this medicine is available on the European Medicines Agency web site:
<https://www.ema.europa.eu>.

Instructions for healthcare professionals

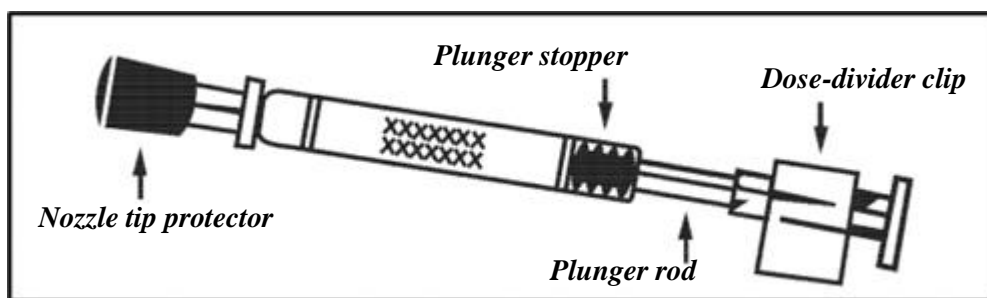
The following information is intended for healthcare professionals only:

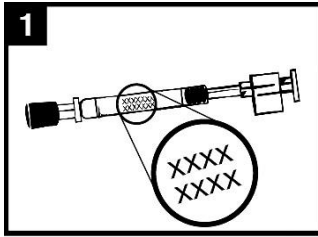
Fluenz is for single nasal use only.

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



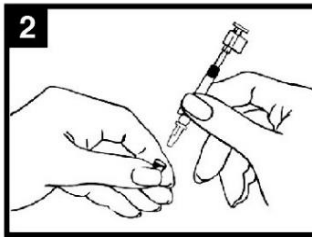
- Do not use Fluenz if the expiry date has passed or the sprayer appears damaged, for example, if the plunger is loose or displaced from the nasal applicator or if there are any signs of leakage.
- Check the appearance of the vaccine before administration. The suspension should be colourless to pale yellow, clear to opalescent. Small white particles may be present.
- Fluenz is administered as a divided dose in both nostrils as described below (see 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.





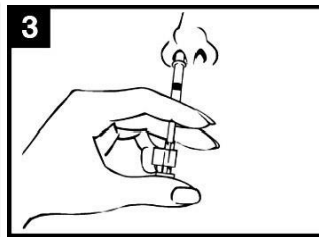
Check expiry date

Product must not be used after date on applicator label.



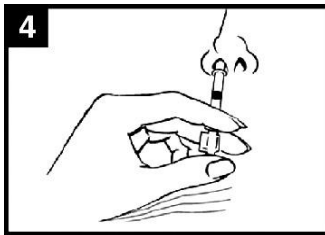
Prepare the applicator

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



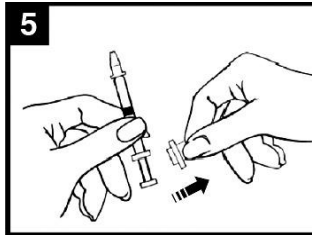
Position the applicator

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



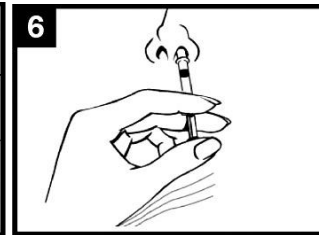
Depress the plunger

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.