ANNEX I

SUMMARY OF PRODUCT CHARACTERISTICS

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. QUALITATIVE AND QUANTITATIVE COMPOSITION

Each 0.5 ml adjuvanted dose contains:

Active substances:
Purified diphtheria toxoid ..... equal to or greater than 20 IU* (30 Lf)
Purified tetanus toxoid ..... equal to or greater than 40 IU* (10 Lf)
Purified pertussis toxoid ..... 25 micrograms
Purified pertussis filamentous haemagglutinin ..... 25 micrograms
Hepatitis B surface antigen ** ..... 5.0 micrograms
Inactivated type 1 poliovirus (Mahoney) ..... D antigen^: 40 units†
Inactivated type 2 poliovirus (MEF 1) ..... D antigen^: 8 units†
Inactivated type 3 poliovirus (Saukett) ..... D antigen^: 32 units†
Haemophilus influenzae type b polysaccharide (polyribosylribitol phosphate) 12 micrograms conjugated to tetanus toxoid (24 micrograms)

Adjuvanted on aluminium hydroxide (0.3 mg)

* As lower confidence limit (p = 0.95).
** Surface antigen of hepatitis B virus produced from recombinant strain 2150-2-3 of the yeast *Saccharomyces cerevisiae*.
^ Quantity of antigen in the final bulk product, according to W.H.O. (TRS 673, 1992)
† Or equivalent antigenic quantity determined by a suitable immunochemical method

For excipients, see 6.1

3. PHARMACEUTICAL FORM

Suspension for injection in pre-filled syringe
HEXAVAC is a slightly opaque white suspension

4. CLINICAL PARTICULARS

4.1 Therapeutic indications

This combined vaccine is indicated for primary and booster vaccination of children against diphtheria, tetanus, pertussis, hepatitis B caused by all known subtypes of viruses, poliomyelitis and invasive infections caused by *Haemophilus influenzae* type b.

4.2 Posology and method of administration

Primary vaccination :
The primary vaccination schedule consists of two or three doses of 0.5 ml administered within the first year of life according to official recommendations. There should be an interval of at least 1 month between doses : such as 2 , 3, 4 months; 2, 4, 6 months; 3, 5 months.

Booster :
After a primary vaccination with 2 doses of HEXAVAC (i.e. 3, 5 months), a booster dose must be given between 11 and 13 months of age; after a primary vaccination with 3 doses of Hexavac (e.g. 2, 3, 4 months; 2, 4, 6 months), a booster dose must be given between 12 and 18 months of age, according to official recommendations.
HEXAVAC can be used for the booster dose provided the toddler has received a full primary vaccination course of each of the antigens contained in HEXAVAC regardless of whether they were administered as monovalent or combination vaccines produced by Sanofi Pasteur MSD.

Method of administration
HEXAVAC should be administered intramuscularly into the quadriceps or deltoid preferably at alternating sites for subsequent injections.

This vaccine should not be used in newborns, adolescents or adults.

4.3 Contraindications

Known hypersensitivity to any component of the vaccine or severe reaction after previous administration of the vaccine.

Encephalopathy within 7 days of administration of a previous dose of any vaccine containing pertussis antigens (whole cell or acellular pertussis vaccines).

In these circumstances the vaccination course should be continued with vaccine not containing a pertussis component.
Vaccination should be postponed in the case of fever or acute disease.

4.4 Special warnings and special precautions for use

This vaccine should not be used in newborns, adolescents or adults.

Infants born of hepatitis B virus surface antigen (HBsAg)-positive mothers should receive Hepatitis B Immune Globulin (HBIG) and Hepatitis B Vaccine (Recombinant) at birth and should complete the hepatitis B vaccination series. The subsequent administration of HEXAVAC for completion of the hepatitis B vaccination series in infants who were born of HBsAg-positive mothers and received HBIG or infants born of mothers of unknown status has not been studied. HEXAVAC should not be used as the birth dose or subsequent doses during the first year of life for children born to HbsAg-positive mothers.

HEXAVAC should be administered with caution to subjects with thrombocytopenia or a bleeding disorder since bleeding may occur following an intramuscular administration to these subjects.

HEXAVAC should under no circumstances be administered intravascularly. The intradermal or subcutaneous routes must not be used either.

If any of the following events are known to have occurred in temporal relation to receipt of the vaccine, the decision to give further doses of pertussis-containing vaccines should be carefully considered:

Temperature of ≥ 40.0 °C within 48 hours, not due to another identifiable cause.
Collapse or shock-like state (hypotonic-hyporesponsiveness episode) within 48 hours of vaccination.
Persistent, inconsolable crying lasting ≥ 3 hours, occurring within 48 hours of vaccination.
Convulsions with or without fever, occurring within 3 days of vaccination.

Antipyretic treatment should be initiated according to local guidelines.

As with all injectable vaccines, appropriate medical treatment and supervision should be readily available for immediate use in case of a rare anaphylactic reaction following the administration of vaccine.

In subjects who have a history of a severe reaction within 48 hours of a previous injection with a vaccine containing similar components, the course of vaccination should be carefully considered.

Because of the long incubation period of hepatitis B, it is possible for unrecognised hepatitis B infection to be present at the time of immunisation. The vaccine may not prevent hepatitis B infection in such cases.

HEXAVAC will not prevent hepatitis infection caused by other agents such as hepatitis A, hepatitis C and hepatitis E or by other liver pathogens.

HEXAVAC does not protect against invasive diseases due to serotypes other than *Haemophilus influenzae* type b or against meningitis of other origins.

As each dose may contain undetectable traces of neomycin, streptomycin and polymyxin B, caution should be exercised when the vaccine is administered to subjects with hypersensitivity to these antibiotics.

The immunogenicity of HEXAVAC could be reduced by immunosuppressive treatment or immunodeficiency. In such cases it is recommended to postpone the vaccination until the end of the disease or treatment. Nevertheless, vaccination of subjects with chronic immunodeficiency such as HIV infection is recommended even if the antibody response might be limited.

There are currently no sufficient data available regarding immunogenicity of the concomitant administration of HEXAVAC with PREVENAR (pneumococcal polysaccharide conjugated vaccine, adsorbed). However when HEXAVAC was co-administered with PREVENAR (pneumococcal polysaccharide conjugated vaccine, adsorbed) in clinical studies, the rate of febrile reactions was higher compared to that occurring following the administration of hexavalent vaccines alone. These reactions were mostly moderate (less than or equal to 39° C) and transient.

HEXAVAC must not be mixed in the same syringe with other vaccines or other parenterally administered drugs.

4.5 Interaction with other medicinal products and other forms of interaction

Except in the case of immunosuppressive therapy (see 4.4 Special warnings and special precautions for use), no significant clinical interaction with other treatments or biological products has been documented.

There are no data in regards to the efficacy and safety of concomitant administration of HEXAVAC with Measles, Mumps and Rubella Virus Vaccine, live.

There are currently no sufficient data available regarding immunogenicity of the concomitant administration of HEXAVAC with PREVENAR (pneumococcal polysaccharide conjugated vaccine, adsorbed).

4.6 Pregnancy and lactation

Not applicable

4.7 Effects on ability to drive and use machines

Not applicable

4.8 Undesirable effects

- ***Clinical trials experience***

In clinical trials, more than 3,900 infants and 4,400 toddlers (from 12 to 20 months of age) have received HEXAVAC.

Commonly reported reactions included redness and/or induration/swelling/pain at the injection site, fever equal to or greater than 38 °C, irritability, drowsiness, loss of appetite, insomnia, diarrhoea and vomiting. Less commonly, fever equal to or greater than 40 °C, tenderness at the injection site, prolonged inconsolable crying and redness and/or induration > 7 cm at the injection site or swelling of the entire limb have been reported. Febrile convulsion and high-pitched crying have been reported rarely. A single bilateral oedematous reaction of the lower limbs and a single, hypotonic hypo-responsive episode have been reported.

These signs and symptoms usually occurred within 48 hours following the vaccination. They were mostly mild, generally lasting for up to 72 hours and resolved spontaneously.

No increase in the number of undesirable reactions was noted between first, second and third doses of the primary series except for an increase in the rate of fever equal to or greater than 38 °C after the second dose in the primary series.

The rate of fever equal to or greater than 40 °C increased after booster immunisation but remained < 1 %. Redness and/or induration > 7 cm at the injection site increased after booster immunisation but remained < 1 %. In rare instances, these cases were associated with swelling of the entire limb.

- ***Post marketing experience***

The following additional undesirable effects have been reported following the widespread use of HEXAVAC.

Common (>1/100 and <1/10)
Application site disorders (reactions at the injection site): Oedema / Pruritus / Urticaria.

Rare (>1/10,000 and <1/1,000)
Body as a whole - General disorders: Prolonged or abnormal crying.

Very rare (<1/10,000)
Body as a whole - General disorders: Allergic reaction / Chills / Fatigue / Hypotonic-hyporesponsive episode / Malaise / Oedema / Pallor / Swelling or oedema of the entire limb(s) / Transient local lymph node swelling.

Central and peripheral nervous system disorders: Convulsions (febrile and non febrile) / Encephalitis / Encephalopathy with acute brain oedema / Eyes rolling / Guillain Barré Syndrome / Hypotonia / Neuritis.

Gastro-intestinal system disorders: Abdominal pain / Meteorism / Nausea.

Platelets, bleeding & clotting disorders: Petechiae / Purpura / Purpura thrombocytopenic / Thrombocytopenia.

Psychiatric disorders: Agitation / Sleep disorder.

Respiratory system disorders: Dyspnoea or Stridor inspiratory.

Skin and appendages disorders: Angioedema / Erythema / Pruritus / Rash / **U**rticaria .

Vascular (extracardiac) disorders: Flushing.

- ***Potential undesirable effects***

In addition, other undesirable effects have been reported with the marketed use of vaccines closely related to HEXAVAC.
Undesirable effects reported in clinical trials and the marketed use to date of Sanofi Pasteur MSD diphtheria, tetanus, acellular pertussis, *Haemophilus influenzae* type b and inactivated poliomyelitis adsorbed vaccine are included in the list of undesirable effects for HEXAVAC.
Very rare reactions following the use of Sanofi Pasteur MSD hepatitis B (recombinant) vaccine include alopecia, hypotension, optic neuritis, facial paralysis, erythema multiforme, and anaphylaxis. As with other hepatitis B vaccines, in many instances, the causal relationship to the vaccine has not been established.

4.9 Overdose

Not applicable.

5. PHARMACOLOGICAL PROPERTIES

5.1 Pharmacodynamic properties

Pharmacotherapeutic group: Bacterial and viral vaccines, combined, ATC code : J07CA

The diphtheria and tetanus toxoids are prepared from the toxins of cultures of *Corynebacterium diphtheriae* and *Clostridium tetani* by formaldehyde detoxification followed by purification. The surface antigen of hepatitis B virus is produced by culture of a recombinant strain of yeast cells (*Saccharomyces cerevisiae)*.

The poliomyelitis vaccine is obtained from the propagation of poliomyelitis viruses types 1, 2 and 3 on Vero cells, purified, then inactivated by formaldehyde.

The acellular pertussis components (pertussis toxin: PT and filamentous haemagglutinin: FHA) are extracted from *Bordetella pertussis* cultures then separately purified. The pertussis toxin (PT) is detoxified separately with glutaraldehyde to create the toxoid (PTxd). The FHA is not detoxified. It has been shown that PTxd and FHA play a major role in protection against pertussis.
This vaccine contains the purified capsular polysaccharide (polyribosyl ribitol phosphate: PRP) of *Haemophilus influenzae* type b conjugated to tetanus toxoid. When administered alone PRP induces a serological response, but it is weakly immunogenic in infants. The covalent binding of PRP to tetanus toxoid makes it a T-cell dependent antigen which induces a specific IgG anti-PRP response in infants and which elicits immune memory.

This vaccine induces specific humoral antibodies against HBsAg (anti-HBs) and against diphtheria and tetanus toxoids (anti-D and anti-T). Development of anti-HBs titre equal to or greater than 10 mIU/ml and of anti-D and anti-T equal to or greater than 0.01 IU/ml measured 1-2 months after the third injection correlates with protection against hepatitis B infection and against diphtheria and tetanus respectively.

Immune response after primary vaccination
In the pivotal clinical study, all infants (100 %) developed a seroprotective antibody level (equal to or greater than 0.01 IU/ml) to both diphtheria and tetanus antigens one month after completion of the primary series. For pertussis, 91.8 % and 90.5 % of infants achieved a four-fold rise in PT and FHA antibody titres respectively. The 4-fold increase in post immunisation titres is considered a sign of seroconversion of which the clinical significance is unknown in the absence of a serological correlate of protection. Protective levels of anti-HBs (equal to or greater than 10 mIU/ml) were achieved in

96.6 % of infants ; the geometric mean titres (GMTs) were diminished compared to the control group. Anti-poliovirus titres above the threshold of 5 (reciprocal of dilution in seroneutralisation) against poliovirus types 1, 2 and 3 were developed in 100 % of infants and these were considered protected against poliomyelitis. After primary vaccination, 93.7 % of infants had an anti-PRP titre equal to or greater than 0.15 µg/ml; the GMTs were diminished compared to the control group (2.06 µg/ml versus 3.69 µg/ml).

Immune response after booster injection
In the pivotal clinical study where toddlers received HEXAVAC as a booster dose after having been primed with HEXAVAC, antibody titres equal to or greater than 0.1 IU/ml were achieved by all toddlers to tetanus and by 98.8 % to diphtheria. A mean 7.4 and 4.3-fold rise in antibody titres to PT and FHA respectively was achieved and all toddlers developed protective antibody titres against poliovirus types 1, 2 and 3. Just before the booster injection anti-PRP GMTs were 0.40 µg/ml and 0.64 µg/ml for HEXAVAC and for the control group respectively. After booster, GMTs increased to 16.7 µg/ml and 23.0 µg/ml in each group respectively, indicating a strong anamnestic response. Anti-PRP titres equal to or greater than 0.15 µg/ml and equal to or greater than 1 µg/ml were achieved in 100 % and 96.6 % of toddlers respectively. Following the booster dose, 96.6 % of toddlers developed anti-HBs titres equal to or greater than 10 mIU/ml. A mean 20.5-fold rise in antibody titres to HBs was observed after the booster dose. Other trials gave similar or higher results. Surveillance and antibody long term persistence studies are ongoing and will provide additional information in regards to the duration of protection.
After a 3, 5, 12 months schedule, immune responses were compatible with the sought clinical protection and of the same magnitude as those reported previously for HEXAVAC or other licensed combination vaccines during the second year of life

5.2 Pharmacokinetic properties

Evaluation of pharmacokinetic properties is not required for vaccines.

5.3 Preclinical safety data

Preclinical data including single-dose, repeated dose and local tolerance studies revealed no unexpected findings and no target organ toxicity.

6. PHARMACEUTICAL PARTICULARS

6.1 List of excipients

Formulation contains aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

6.2 Incompatibilities

The vaccine should not be mixed in the same syringe with other vaccines or parenterally administered substances.

6.3 Shelf life

36 months

6.4 Special precautions for storage

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

6.5 Nature and contents of container

0.5 ml of suspension in pre-filled syringe (type I glass) with a plunger stopper (chlorobromobutyl) with attached needle - pack of 1, 10, 25 and 50.
0.5 ml of suspension in pre-filled syringe (type I glass) with a plunger stopper (chlorobromobutyl) without needle - pack of 1, 10, 25 and 50.
0.5 ml of suspension in pre-filled syringe (type I glass) with a plunger stopper (chlorobromobutyl), with 1 or 2 separate needles - pack of 1 and 10.

Not all pack sizes may be marketed.

6.6 Instructions for use and handling

Before use, the vaccine should be well shaken in order to obtain a homogeneous slightly opaque white suspension.

For needle free syringes, the needle should be pushed firmly on to the end of the prefilled syringe and rotated through 90 degrees.

7. MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

8. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/001- 012

9. DATE OF FIRST AUTHORISATION/RENEWAL OF THE AUTHORISATION

October 23rd, 2000

10. DATE OF REVISION OF THE TEXT

ANNEX II

A. MANUFACTURER OF THE BIOLOGICAL ACTIVE SUBSTANCES AND MANUFACTURING AUTHORISATION HOLDER RESPONSIBLE FOR BATCH RELEASE

B. CONDITIONS OF THE MARKETING AUTHORISATION

A. MANUFACTURER OF THE BIOLOGICAL ACTIVE SUBSTANCES AND MANUFACTURING AUTHORISATION HOLDER RESPONSIBLE FOR BATCH RELEASE

Name and address of the manufacturer of the biological active substances

For Hepatitis B surface antigen:
Merck & Co. Inc.
Sumneytown Pike
West Point
Pennsylvania 19486 USA

For the other components:
Sanofi Pasteur SA
Campus Mérieux
1541 Avenue Marcel Mérieux
F-69280 Marcy L'Etoile

Name and address of the manufacturer responsible for batch release

Sanofi Pasteur SA
Campus Mérieux
1541 Avenue Marcel Mérieux
F-69280 Marcy L'Etoile

B. CONDITIONS OF THE MARKETING AUTHORISATION

- **CONDITIONS OR RESTRICTIONS REGARDING SUPPLY AND USE IMPOSED ON THE MARKETING AUTHORISATION HOLDER**

Medicinal product subject to medical prescription.

- **OTHER CONDITIONS**

The holder of this marketing authorisation must inform the European Commission about the marketing plans for the medicinal product authorised by this decision.

Official batch release: in accordance with Article 114 Directive 2001/83/EEC, the official batch release will be undertaken by a state laboratory or a laboratory designated for that purpose.

ANNEX III

LABELLING AND PACKAGE LEAFLET

A. LABELLING

PARTICULARS TO APPEAR ON THE OUTER PACKAGING OR, WHERE THERE IS NO OUTER PACKAGING, ON THE IMMEDIATE PACKAGING

HEXAVAC - single dose pre-filled syringe with attached needle - Pack of 1

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. STATEMENT OF ACTIVE SUBSTANCE(S)

1 adjuvanted dose (0.5 ml) contains:

Purified diphtheria toxoid	≥ 20 IU
Purified tetanus toxoid	≥ 40 IU
Purified pertussis toxoid	25 µg
Purified pertussis filamentous haemagglutinin	25 µg
Hepatitis B surface antigen	5.0 µg
Inactivated type 1 poliovirus	40 D units
Inactivated type 2 poliovirus	8 D units
Inactivated type 3 poliovirus	32 D units
Haemophilus influenzae type b conjugated to tetanus toxoid	12 µg

3. LIST OF EXCIPIENTS

Excipients: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

1 single dose 0.5 ml pre-filled syringe with attached needle
Suspension for injection in pre-filled syringe

5. METHOD AND, IF NECESSARY, ROUTE(S) OF ADMINISTRATION

Shake well before use.
Intramuscular use.

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

Keep out of the reach and sight of children.

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

9. SPECIAL STORAGE CONDITIONS

Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze

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

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

12. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/001

13. MANUFACTURER'S BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription

15. INSTRUCTIONS ON USE

PARTICULARS TO APPEAR ON THE OUTER PACKAGING OR, WHERE THERE IS NO OUTER PACKAGING, ON THE IMMEDIATE PACKAGING

HEXAVAC - single dose pre-filled syringe without needle - Pack of 1

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. STATEMENT OF ACTIVE SUBSTANCE(S)

1 adjuvanted dose (0.5 ml) contains:

Purified diphtheria toxoid	≥ 20 IU
Purified tetanus toxoid	≥ 40 IU
Purified pertussis toxoid	25 µg
Purified pertussis filamentous haemagglutinin	25 µg
Hepatitis B surface antigen	5.0 µg
Inactivated type 1 poliovirus	40 D units
Inactivated type 2 poliovirus	8 D units
Inactivated type 3 poliovirus	32 D units
Haemophilus influenzae type b conjugated to tetanus toxoid	12 µg

3. LIST OF EXCIPIENTS

Excipients: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

1 single dose 0.5 ml pre-filled syringe without needle
Suspension for injection in pre-filled syringe

5. METHOD AND, IF NECESSARY, ROUTE(S) OF ADMINISTRATION

Shake well before use.
Intramuscular use.

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

Keep out of the reach and sight of children

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

9. SPECIAL STORAGE CONDITIONS

Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze

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

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

12. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/005

13. MANUFACTURER'S BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription

15. INSTRUCTIONS ON USE

PARTICULARS TO APPEAR ON THE OUTER PACKAGING OR, WHERE THERE IS NO OUTER PACKAGING, ON THE IMMEDIATE PACKAGING

HEXAVAC - single dose pre-filled syringe with 1 separate needle - Pack of 1

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. STATEMENT OF ACTIVE SUBSTANCE(S)

1 adjuvanted dose (0.5 ml) contains:

Purified diphtheria toxoid	≥ 20 IU
Purified tetanus toxoid	≥ 40 IU
Purified pertussis toxoid	25 µg
Purified pertussis filamentous haemagglutinin	25 µg
Hepatitis B surface antigen	5.0 µg
Inactivated type 1 poliovirus	40 D units
Inactivated type 2 poliovirus	8 D units
Inactivated type 3 poliovirus	32 D units
Haemophilus influenzae type b conjugated to tetanus toxoid	12 µg

3. LIST OF EXCIPIENTS

Excipients: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

1 single dose 0.5 ml pre-filled syringe with 1 separate needle
Suspension for injection in pre-filled syringe

5. METHOD AND, IF NECESSARY, ROUTE(S) OF ADMINISTRATION

Shake well before use.
Intramuscular use.

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

Keep out of the reach and sight of children

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

9. SPECIAL STORAGE CONDITIONS

Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze

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

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

12. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/009

13. MANUFACTURER'S BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription

15. INSTRUCTIONS ON USE

PARTICULARS TO APPEAR ON THE OUTER PACKAGING OR, WHERE THERE IS NO OUTER PACKAGING, ON THE IMMEDIATE PACKAGING

HEXAVAC - single dose pre-filled syringe with 2 separate needles - Pack of 1

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. STATEMENT OF ACTIVE SUBSTANCE(S)

1 adjuvanted dose (0.5 ml) contains:

Purified diphtheria toxoid	≥ 20 IU
Purified tetanus toxoid	≥ 40 IU
Purified pertussis toxoid	25 µg
Purified pertussis filamentous haemagglutinin	25 µg
Hepatitis B surface antigen	5.0 µg
Inactivated type 1 poliovirus	40 D units
Inactivated type 2 poliovirus	8 D units
Inactivated type 3 poliovirus	32 D units
Haemophilus influenzae type b conjugated to tetanus toxoid	12 µg

3. LIST OF EXCIPIENTS

Excipients: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

1 single dose 0.5 ml pre-filled syringe with 2 separate needles
Suspension for injection in pre-filled syringe

5. METHOD AND, IF NECESSARY, ROUTE(S) OF ADMINISTRATION

Shake well before use.
Intramuscular use.

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

Keep out of the reach and sight of children

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

9. SPECIAL STORAGE CONDITIONS

Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze

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

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

12. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/010

13. MANUFACTURER'S BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription

15. INSTRUCTIONS ON USE

PARTICULARS TO APPEAR ON THE OUTER PACKAGING OR, WHERE THERE IS NO OUTER PACKAGING, ON THE IMMEDIATE PACKAGING

HEXAVAC - single dose pre-filled syringe with attached needle - Pack of 10

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. STATEMENT OF ACTIVE SUBSTANCE(S)

1 adjuvanted dose (0.5 ml) contains:

Purified diphtheria toxoid	≥ 20 IU
Purified tetanus toxoid	≥ 40 IU
Purified pertussis toxoid	25 µg
Purified pertussis filamentous haemagglutinin	25 µg
Hepatitis B surface antigen	5.0 µg
Inactivated type 1 poliovirus	40 D units
Inactivated type 2 poliovirus	8 D units
Inactivated type 3 poliovirus	32 D units
Haemophilus influenzae type b conjugated to tetanus toxoid	12 µg

3. LIST OF EXCIPIENTS

Excipients: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

10 single doses 0.5 ml pre-filled syringes with attached needle.
Suspension for injection in pre-filled syringe

5. METHOD AND, IF NECESSARY, ROUTE(S) OF ADMINISTRATION

Shake well before use.
Intramuscular use.

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

Keep out of the reach and sight of children.

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

9. SPECIAL STORAGE CONDITIONS

Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze

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

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

12. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/002

13. MANUFACTURER'S BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription

15. INSTRUCTIONS ON USE

PARTICULARS TO APPEAR ON THE OUTER PACKAGING OR, WHERE THERE IS NO OUTER PACKAGING, ON THE IMMEDIATE PACKAGING

HEXAVAC - single dose pre-filled syringe without needle - Pack of 10

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. STATEMENT OF ACTIVE SUBSTANCE(S)

1 adjuvanted dose (0.5 ml) contains:

Purified diphtheria toxoid	≥ 20 IU
Purified tetanus toxoid	≥ 40 IU
Purified pertussis toxoid	25 µg
Purified pertussis filamentous haemagglutinin	25 µg
Hepatitis B surface antigen	5.0 µg
Inactivated type 1 poliovirus	40 D units
Inactivated type 2 poliovirus	8 D units
Inactivated type 3 poliovirus	32 D units
Haemophilus influenzae type b conjugated to tetanus toxoid	12 µg

3. LIST OF EXCIPIENTS

Excipients: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

10 single doses 0.5 ml pre-filled syringes without needle
Suspension for injection in pre-filled syringe

5. METHOD AND, IF NECESSARY, ROUTE(S) OF ADMINISTRATION

Shake well before use.
Intramuscular use.

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

Keep out of the reach and sight of children

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

9. SPECIAL STORAGE CONDITIONS

Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze

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

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

12. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/006

13. MANUFACTURER'S BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription

15. INSTRUCTIONS ON USE

PARTICULARS TO APPEAR ON THE OUTER PACKAGING OR, WHERE THERE IS NO OUTER PACKAGING, ON THE IMMEDIATE PACKAGING

HEXAVAC - single dose pre-filled syringe with 1 separate needle - Pack of 10

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. STATEMENT OF ACTIVE SUBSTANCE(S)

1 adjuvanted dose (0.5 ml) contains:

Purified diphtheria toxoid	≥ 20 IU
Purified tetanus toxoid	≥ 40 IU
Purified pertussis toxoid	25 µg
Purified pertussis filamentous haemagglutinin	25 µg
Hepatitis B surface antigen	5.0 µg
Inactivated type 1 poliovirus	40 D units
Inactivated type 2 poliovirus	8 D units
Inactivated type 3 poliovirus	32 D units
Haemophilus influenzae type b conjugated to tetanus toxoid	12 µg

3. LIST OF EXCIPIENTS

Excipients: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

10 single doses 0.5 ml pre-filled syringes with 1 separate needle (for each syringe).
Suspension for injection in pre-filled syringe

5. METHOD AND, IF NECESSARY, ROUTE(S) OF ADMINISTRATION

Shake well before use.
Intramuscular use.

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

Keep out of the reach and sight of children

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

9. SPECIAL STORAGE CONDITIONS

Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze

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

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

12. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/011

13. MANUFACTURER'S BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription

15. INSTRUCTIONS ON USE

PARTICULARS TO APPEAR ON THE OUTER PACKAGING OR, WHERE THERE IS NO OUTER PACKAGING, ON THE IMMEDIATE PACKAGING

HEXAVAC - single dose pre-filled syringe with 2 separate needles - Pack of 10

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. STATEMENT OF ACTIVE SUBSTANCE(S)

1 adjuvanted dose (0.5 ml) contains:

Purified diphtheria toxoid	≥ 20 IU
Purified tetanus toxoid	≥ 40 IU
Purified pertussis toxoid	25 µg
Purified pertussis filamentous haemagglutinin	25 µg
Hepatitis B surface antigen	5.0 µg
Inactivated type 1 poliovirus	40 D units
Inactivated type 2 poliovirus	8 D units
Inactivated type 3 poliovirus	32 D units
Haemophilus influenzae type b conjugated to tetanus toxoid	12 µg

3. LIST OF EXCIPIENTS

Excipients: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

10 single doses 0.5 ml pre-filled syringes with 2 separate needles (for each syringe)
Suspension for injection in pre-filled syringe

5. METHOD AND, IF NECESSARY, ROUTE(S) OF ADMINISTRATION

Shake well before use.
Intramuscular use.

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

Keep out of the reach and sight of children

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

9. SPECIAL STORAGE CONDITIONS

Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze

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

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

12. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/012

13. MANUFACTURER'S BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription

15. INSTRUCTIONS ON USE

PARTICULARS TO APPEAR ON THE OUTER PACKAGING OR, WHERE THERE IS NO OUTER PACKAGING, ON THE IMMEDIATE PACKAGING

HEXAVAC - single dose pre-filled syringe with attached needle - Pack of 25

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. STATEMENT OF ACTIVE SUBSTANCE(S)

1 adjuvanted dose (0.5 ml) contains:

Purified diphtheria toxoid	≥ 20 IU
Purified tetanus toxoid	≥ 40 IU
Purified pertussis toxoid	25 µg
Purified pertussis filamentous haemagglutinin	25 µg
Hepatitis B surface antigen	5.0 µg
Inactivated type 1 poliovirus	40 D units
Inactivated type 2 poliovirus	8 D units
Inactivated type 3 poliovirus	32 D units
Haemophilus influenzae type b conjugated to tetanus toxoid	12 µg

3. LIST OF EXCIPIENTS

Excipients: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

25 single doses 0.5 ml pre-filled syringes with attached needle.
Suspension for injection in pre-filled syringe

5. METHOD AND, IF NECESSARY, ROUTE(S) OF ADMINISTRATION

Shake well before use.
Intramuscular use.

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

Keep out of the reach and sight of children.

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

9. SPECIAL STORAGE CONDITIONS

Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze

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

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

12. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/003

13. MANUFACTURER'S BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription

15. INSTRUCTIONS ON USE

PARTICULARS TO APPEAR ON THE OUTER PACKAGING OR, WHERE THERE IS NO OUTER PACKAGING, ON THE IMMEDIATE PACKAGING

HEXAVAC - single dose pre-filled syringe without needle - Pack of 25

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. STATEMENT OF ACTIVE SUBSTANCE(S)

1 adjuvanted dose (0.5 ml) contains:

Purified diphtheria toxoid	≥ 20 IU
Purified tetanus toxoid	≥ 40 IU
Purified pertussis toxoid	25 µg
Purified pertussis filamentous haemagglutinin	25 µg
Hepatitis B surface antigen	5.0 µg
Inactivated type 1 poliovirus	40 D units
Inactivated type 2 poliovirus	8 D units
Inactivated type 3 poliovirus	32 D units
Haemophilus influenzae type b conjugated to tetanus toxoid	12 µg

3. LIST OF EXCIPIENTS

Excipients: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

25 single doses 0.5 ml pre-filled syringes without needle
Suspension for injection in pre-filled syringe

5. METHOD AND, IF NECESSARY, ROUTE(S) OF ADMINISTRATION

Shake well before use.
Intramuscular use.

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

Keep out of the reach and sight of children

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

9. SPECIAL STORAGE CONDITIONS

Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze

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

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

12. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/007

13. MANUFACTURER'S BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription

15. INSTRUCTIONS ON USE

PARTICULARS TO APPEAR ON THE OUTER PACKAGING OR, WHERE THERE IS NO OUTER PACKAGING, ON THE IMMEDIATE PACKAGING

HEXAVAC - single dose pre-filled syringe with attached needle - Pack of 50

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. STATEMENT OF ACTIVE SUBSTANCE(S)

1 adjuvanted dose (0.5 ml) contains:

Purified diphtheria toxoid	≥ 20 IU
Purified tetanus toxoid	≥ 40 IU
Purified pertussis toxoid	25 µg
Purified pertussis filamentous haemagglutinin	25 µg
Hepatitis B surface antigen	5.0 µg
Inactivated type 1 poliovirus	40 D units
Inactivated type 2 poliovirus	8 D units
Inactivated type 3 poliovirus	32 D units
Haemophilus influenzae type b conjugated to tetanus toxoid	12 µg

3. LIST OF EXCIPIENTS

Excipients: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

50 single doses 0.5 ml pre-filled syringes with attached needle.
Suspension for injection in pre-filled syringe

5. METHOD AND, IF NECESSARY, ROUTE(S) OF ADMINISTRATION

Shake well before use.
Intramuscular use.

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

Keep out of the reach and sight of children.

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

9. SPECIAL STORAGE CONDITIONS

Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze

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

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

12. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/004

13. MANUFACTURER'S BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription

15. INSTRUCTIONS ON USE

PARTICULARS TO APPEAR ON THE OUTER PACKAGING OR, WHERE THERE IS NO OUTER PACKAGING, ON THE IMMEDIATE PACKAGING

HEXAVAC - single dose pre-filled syringe without needle - Pack of 50

1. NAME OF THE MEDICINAL PRODUCT

HEXAVAC suspension for injection in pre-filled syringe

Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

2. STATEMENT OF ACTIVE SUBSTANCE(S)

1 adjuvanted dose (0.5 ml) contains:

Purified diphtheria toxoid	≥ 20 IU
Purified tetanus toxoid	≥ 40 IU
Purified pertussis toxoid	25 µg
Purified pertussis filamentous haemagglutinin	25 µg
Hepatitis B surface antigen	5.0 µg
Inactivated type 1 poliovirus	40 D units
Inactivated type 2 poliovirus	8 D units
Inactivated type 3 poliovirus	32 D units
Haemophilus influenzae type b conjugated to tetanus toxoid	12 µg

3. LIST OF EXCIPIENTS

Excipients: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

4. PHARMACEUTICAL FORM AND CONTENTS

50 single doses 0.5 ml pre-filled syringes without needle
Suspension for injection in pre-filled syringe

5. METHOD AND, IF NECESSARY, ROUTE(S) OF ADMINISTRATION

Shake well before use.
Intramuscular use.

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

Keep out of the reach and sight of children

7. OTHER SPECIAL WARNING(S), IF NECESSARY

8. EXPIRY DATE

EXP:

9. SPECIAL STORAGE CONDITIONS

Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze

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

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

Sanofi Pasteur MSD SNC
8, rue Jonas Salk
F-69007 Lyon

12. NUMBER(S) IN THE COMMUNITY REGISTER OF MEDICINAL PRODUCTS

EU/1/00/147/008

13. MANUFACTURER'S BATCH NUMBER

Batch

14. GENERAL CLASSIFICATION FOR SUPPLY

Medicinal product subject to medical prescription

15. INSTRUCTIONS ON USE

MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS

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

HEXAVAC

Intramuscular use

2. METHOD OF ADMINISTRATION

Store at 2 °C – 8 °C
Do not freeze
Shake well before use

3. EXPIRY DATE

EXP:

4. BATCH NUMBER

Batch

5. CONTENTS BY WEIGHT, BY VOLUME OR BY UNIT

1 dose = 0.5 ml

Sanofi Pasteur MSD SNC

B. PACKAGE LEAFLET

PACKAGE LEAFLET

Read all of this leaflet carefully before your child is vaccinated.
- Keep this leaflet. You may need to read it again.
- If you have further questions, please ask your doctor or your pharmacist.
- This vaccine has been prescribed for your child and you should not pass it on to others.

In this leaflet:

HEXAVAC suspension for injection in pre-filled syringe
Diphtheria, tetanus, acellular pertussis, inactivated poliomyelitis, hepatitis B (recombinant) and *Haemophilus influenzae* type b conjugate vaccine, adjuvanted.

The active substances are:
Purified diphtheria toxoid ..... equal to or greater than 20 IU* (30 Lf)
Purified tetanus toxoid ..... equal to or greater than 40 IU* (10 Lf)
Purified pertussis toxoid ..... 25 micrograms
Purified pertussis filamentous haemagglutinin ..... 25 micrograms
Hepatitis B surface antigen ** ..... 5.0 micrograms
Inactivated type 1 poliovirus (Mahoney) ..... D antigen^: 40 units†
Inactivated type 2 poliovirus (MEF 1) ..... D antigen^: 8 units†
Inactivated type 3 poliovirus (Saukett) ..... D antigen^: 32 units†
Haemophilus influenzae type b polysaccharide (polyribosylribitol phosphate) 12 micrograms conjugated to tetanus toxoid (24 micrograms)
for one adjuvanted dose of 0.5 ml

* As lower confidence limit ($p = 0.95$).
** Surface antigen of hepatitis B virus produced from recombinant strain 2150-2-3 of the yeast *Saccharomyces cerevisiae*.
^ Quantity of antigen in the final bulk product, according to W.H.O. (TRS 673, 1992)
† Or equivalent antigenic quantity determined by a suitable immunochemical method

The other ingredients are: aluminium hydroxide and a buffer solution of disodium phosphate, monopotassium phosphate, sodium carbonate, sodium bicarbonate, trometamol, sucrose, medium 199 (complex blend of amino acids, mineral salts, vitamins and other ingredients) and water for injections.

Marketing Authorisation Holder: Sanofi Pasteur MSD SNC, 8 rue Jonas Salk, F-69007 Lyon
Manufactured by: Sanofi Pasteur SA, F-69280 Marcy l'Etoile

1. WHAT IS HEXAVAC AND WHAT IT IS USED FOR

HEXAVAC is an injectable vaccine in a 0.5 ml single dose syringe.

HEXAVAC is indicated to help protect your child against diphtheria, tetanus, pertussis, poliomyelitis, infection of the liver caused by all known subtypes of hepatitis B virus and invasive disease (infection of brain and spinal cord tissues, infection of the blood, etc.) caused by *Haemophilus influenzae* type b (Hib) bacterium in children 8 weeks to 18 months of age.

HEXAVAC is available in packs of 1, 10, 25 and 50 with or without neddles.

2. BEFORE YOU USE HEXAVAC

Do not use HEXAVAC:

- if your child is allergic to any component of the vaccine.
- in case of newborns, adolescents or adults.
- if your child has fever or other illness, particularly a cough, cold or flu (vaccination should be delayed).
- if your infant has suffered brain damage (encephalopathy) following a previous dose of a whole cell or acellular pertussis vaccine.

Take special care with HEXAVAC:

- if your child has an hypersensitivity to neomycin, streptomycin and polymyxin B, due to the use of these substances during production.
- if your child has a thrombocytopenia or a bleeding disorder since bleeding may occur following an intramuscular administration to these subjects.
- if any of the following events are known to have occurred in temporal relation to receipt of the vaccine (The decision to give further doses of pertussis-containing vaccines to your child should be carefully considered if any of the following events are known to have occurred following the administration of the vaccine) :
  Temperature of ≥ 40.0 °C within 48 hours, not due to another identifiable cause.
  Collapse or shock-like state (hypotonic-hyporesponsiveness episode) within 48 hours of vaccination.
  Persistent, inconsolable crying lasting ≥ 3 hours, occurring within 48 hours of vaccination.
  Convulsions with or without fever, occurring within 3 days of vaccination.
- if your child has any present or past medical problems or allergies, including allergic reactions after any dose of HEXAVAC.
- if you are a hepatitis B virus surface antigen (HBsAg)-positive mother, your infant should receive Hepatitis B Immune Globulin (HBIG) and hepatitis B vaccine (Recombinant) at birth and should complete the hepatitis B vaccination series. The subsequent administration of HEXAVAC for completion of the hepatitis B vaccination series in infants who were born of HBsAg-positive mothers and received HBIG or infants born of mothers of unknown status has not been studied. HEXAVAC should not be used as the birth dose or subsequent doses during the first year of life for children born to HbsAg-positive mothers .
- because as with other similar vaccines, cases of *Haemophilus* b disease may occur in the week after vaccination prior to the onset of the protective effects of the vaccine.
- because hepatitis B infection can go undetected for a long period of time, it is possible that an individual may already be infected at the time the vaccine is given. The vaccine may not prevent hepatitis B in these individuals.

Using other vaccines
There are currently not sufficient data available regarding immune response of the concomitant administration of HEXAVAC with PREVENAR (pneumococcal polysaccharide conjugated vaccine, adsorbed). However, when HEXAVAC was co-administered in clinical studies the rate of febrile reactions was higher compared to that occurring following administration of hexavalent vaccines alone. These reactions were mostly moderate and transient. If your child is due to be vaccinated with HEXAVAC and other vaccines simultaneously, please ask your doctor for further information.

3. HOW TO USE HEXAVAC

The primary vaccination schedule consists of two or three doses of 0.5 ml administered within the first year of life according to the official recommendations. There should be an interval of at least 1 month between doses : such as 2 , 3, 4 months; 2, 4, 6 months; 3, 5 months.

After a primary vaccination with 2 doses of HEXAVAC (e.g 3, 5 months), a booster dose must be given between 11 and 13 months of age; after a primary vaccination with 3 doses of Hexavac (e.g. 2, 3, 4 months; 2, 4, 6 months), a booster dose must be given between 12 and 18 months of age, according to official recommendations.
HEXAVAC can be used for the booster dose provided the toddler has received a full primary vaccination course of each of the antigens contained in Hexavac regardless of whether they were administered as monovalent or combination vaccines produced by Sanofi Pasteur MSD.

HEXAVAC should be injected into the quadriceps or deltoid muscle preferably at alternating sites for subsequent injections.
HEXAVAC should under no circumstances be administered intravascularly. The intradermal or subcutaneous routes must not be used either.

HEXAVAC must not be mixed in the same syringe with other vaccines or other parenterally administered drugs.

For needle free syringes, the needle should be pushed firmly on to the end of the prefilled syringe and rotated through 90 degrees.

If you forget to take HEXAVAC:
Your doctor will decide when to give the missed dose.

4. POSSIBLE SIDE EFFECTS

As with other vaccines your child may have some side effects. HEXAVAC has been generally well tolerated in clinical trials. Side effects included injection-site reactions such as tenderness, redness, swelling and pain. Other side effects included irritability, sleepiness, fever, sleeplessness, diarrhoea, vomiting, loss of appetite and prolonged crying where the child cannot be consoled.

Following the widespread use of HEXAVAC, additional undesirable effects have been reported:
Among reactions found at the injection site, itching and hives have been also reported.
Rarely, the patient may have prolonged or abnormal crying.
Very rarely there may be allergic reaction; chills; fatigue; malaise; oedema; swelling of the entire limb(s); Guillain Barré Syndrome; hypotonic hyporesponsive episode; pallor; convulsions (with or without fever); brain inflammation, acute brain swelling; eyes rolling; decrease in muscle tone; neuritis; nausea; abdominal gas and/or pain; low platelet count; purple or red brown spots visible through the skin; agitation; difficulty sleeping; breathing difficulty ; wheezing ; swelling of the face, lips, mouth, tongue or throat which may cause difficulty in swallowing or breathing; rash; skin redness; generalized hives; generalized itching; flushing; temporary swelling of local lymph nodes.

Tell your doctor promptly about these symptoms. If the condition persists or worsens, you may have to bring your child to the doctor.

In addition, tell your doctor if your child experiences any symptoms that suggest an allergic reaction such as rash, redness, itching, pallor or oedema after any dose in the vaccination series.

If you notice any side effects not mentioned in this leaflet, please inform your doctor or pharmacist.

5. STORING HEXAVAC

Keep out of the reach and sight of children.
Store at 2 °C – 8 °C (in a refrigerator)
Do not freeze
Do not use after the expiry date stated on the label.

6. FURTHER INFORMATION

For any information about this medicinal product, please contact the local representative of the Marketing Authorisation Holder.

België/Belgique/Belgien
Sanofi Pasteur MSD sa/nv
Tél: 32.2.726.95.84

Česká republika
AVENTIS PASTEUR OFFICE PRAHA
Tel: 420 222 522 523

Danmark
Sanofi Pasteur MSD sa/nv
Tél: 32.2.726.95.84

Deutschland
Sanofi Pasteur MSD GmbH
Tel: 49.6224.594.0

Eesti
AS Oriola – Tallinn
Tel: 370 5 273 0967

Ελλάδα
BIANEΞ A.E.
Τηλ. 30.210.8009111

España
Sanofi Pasteur MSD SA
Tel: 349.1.371 78.00

France
Sanofi Pasteur MSD SNC
Tél: 33.4.37.28.40.00

Ireland
Sanofi Pasteur MSD Ltd
Tel: 3531.295.2226

Ísland
Sanofi Pasteur MSD sa/nv
Tel: +32.2.726.95.84

Luxembourg/Luxemburg
Sanofi Pasteur MSD sa/nv
Tél: 32.2.726.95.84

Magyarország
AVENTIS PASTEUR Representative Office
Tel.: 36 13 28 39 80

Malta
CHERUBINO LTD
Tel: 356 21 343 270

Nederland
Sanofi Pasteur MSD sa/nv branch
Tel: 32.2.726.95.84

Norge
Sanofi Pasteur MSD sa/nv
Tlf: +32.2.726.95.84

Österreich
Sanofi Pasteur MSD GmbH
Tel: 43.1.866.70.22.202

Polska
AVENTIS PASTEUR Sp.Z.o.o.
Tel.: 48 22 661 55 39

Portugal
UCB PHARMA Lda
Tel: 351.21.302.53.00

Slovenija
Aventis Pasteur GmbH Representative
Tel: 386 4 33 74 14

Slovenská republika
Aventis Pasteur GmbH
Tel: 421 41 700 2711

Italia
Sanofi Pasteur MSD SpA
Tel: 390.6.664.092.11

Κύπρος
XANTOS LYSSIOTIS AND SON
Τηλ: 357 22 34 80 06

Latvija
AVENTIS PASTEUR GmbH
Tel: 370 5 273 0967

Lietuva
AVENTIS PASTEUR Representative Office
Tel. 370 5 273 0967

Suomi/Finland
Sanofi Pasteur MSD sa/nv
Tél: 32.2.726.95.84

Sverige
Sanofi Pasteur MSD SNC France Filial i Sverige
Tél: 32.2.726.95.84

United Kingdom
Sanofi Pasteur MSD Ltd
Tel: 44.1.628.785.291

This leaflet was last approved on: