

**INFLUENZA VACCINE (SPLIT VIRION,
INACTIVATED) MONOVALENT POOLED
HARVEST**

ORIGINAL STRAIN B/Victoria lineage

Working virus seed lot used :
PRODUCTION

Dates of inoculation, harvest, disruption and inactivation

Inoculation	Harvest	Disruption	Inactivation

Disruption agent : Octoxynol 9
Treatment \geq 1 hour

Storage conditions before inactivation : $+ 5^{\circ} \pm 3^{\circ}C$

Inactivation : By formaldehyde action

Concentration of inactivating agent : 0.2 %

Time of inactivation : 25-26 hours at $+20^{\circ}C \pm 5^{\circ}C$ followed by a stage at $+5^{\circ}C \pm 3^{\circ}C$ for 24-96 hours

Storage conditions after inactivation : $+ 5^{\circ} \pm 3^{\circ}C$

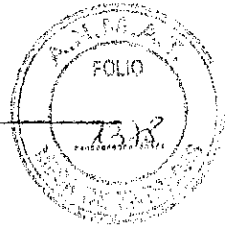
Concentration and purification procedure : Isopycnic ultracentrifugation on a saccharose gradient

Antibiotics used during preparation : Neomycine sulfate in the inoculum

Filtration : On 0.22 μm pore size membrane

Destinataire : A. Patention de l'ANSM

Export officiel non contrôlé
Modèle : Batch Release protocol Q1Y



Certificate of Analysis

Product Name : INFLUENZA VACCINE (SPLIT VIRION, INACTIVATED)

Process Stage : WORKING SEED LOT

Volume / Quantity :

Storage Condition : ≤ -60°C

Date of Manufacture :

Working Seed Lot

Test(s)	Acceptance Criteria	Result	Approval Date
TEST FOR MYCOPLASMA BY MICROBIOLOGICAL CULTURE <i>By Inoculation to suitable culture media</i>	Negative for the presence of mycoplasma when cultured on agar and broth		
BACTERIAL AND FUNGAL STERILITY <i>By Membrane filtration</i>	No microbial growth		
TEST FOR MYCOPLASMA BY EPIFLUORESCENCE IN CELLULAR CULTURE <i>By Inoculation to indicator cells and specific staining</i>	Negative for the presence of mycoplasma when cultured in cells and treated with a fluorescent dye		
NEURAMINIDASE ANTIGEN IDENTIFICATION <i>By RT-PCR</i>	POSITIVE		
HAEMAGGLUTININ ANTIGEN IDENTIFICATION <i>By Haemagglutination inhibition</i>	POSITIVE		
INFECTIOUS TITER <i>By EID50</i>	≥ 6 log10 EID50/mL	log10 EID50/mL	

Destinataire : A l'attention de l'ANSM

Export officiel non contrôlé
Motif : Batch Release PROCECOQV

Information Confidentielle
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ROXANA MONTEMILONE
DIRECTORA TÉCNICA
APODERADA
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Número : Q_0565202
Version : 2.0
Categoría : I

Par : BOSSUT Isabelle
Le : 24 oct. 2016 13:56:56

Statut : Applicable



**INFLUENZA VACCINE (SPLIT VIRION,
INACTIVATED)
WORKING VIRUS SEED LOT
B/Victoria lineage -**

A/ PRODUCTION

Lot number :

Inoculation

Date of inoculation :

Inoculum : Primary seed lot

Passage number from primary seed lot :

Inoculated dilution (log10) :

Eggs used :

Inoculated volume per egg : mL

Number of eggs inoculated :

Temperature of incubation : **Optimal incubation conditions**

Added antibiotics : None

Harvest and purification

Date :

Storage conditions : ≤ - 60°C

Release date :

Destinataire : A l'attention de FANSM

Export officiel non contrôlé
Motif : Batch Release Protocole

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Catégorie : 1





B – CONTROLS

Test for mycoplasma

- by culture method

Method : Agar culture

Volume tested : ml

Media used : Hayflick – Frey

Temperature of incubation :

Positive controls :
Mycoplasma pneumoniae, Mycoplasma
orale, Mycoplasma gallisepticum,
Mycoplasma synoviae, Acholeplasma
laidlawii

. Test on :

. Test off :

Subculture at the 3rd day

. Test on :

. Test off :

Subculture at the 7th day

. Test on :

. Test off :

Subculture at the 14th day

. Test on :

. Test off :

Subculture at the 21th day

. Test on :

. Test off :

Result :

Positive control :

Mycoplasma pneumoniae :

Mycoplasma orale :

Mycoplasma gallisepticum :

Mycoplasma synoviae :

Acholeplasma laidlawii :

Result :

Destinataire : A l'attention de l'ANSM

Export officiel non contrôlé
Motif : Batch release procedure

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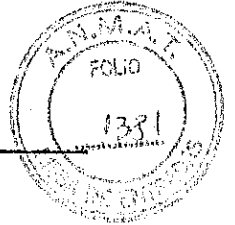
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Test for Mycoplasma
- by epifluorescence in cell culture

Method : DNA fluorescence staining
Cell used : ,th passage
Volume tested : ml
Temperature of incubation : °C
Positive control : Mycoplasma hyorhinis and Mycoplasma orale
Test on :
Test off :
Result :

Bacterial and fungal sterility

Method : Membrane filtration

Media	Thioglycollate medium	Soybean Casein Digest medium
Temperatures :	30-35°C	20-25°C
Volume tested :	ml	ml
Inoculum per Steritest :	ml	ml
Volume of medium :	100 ml	100 ml
Number of inoculated Steritest :	1	1
Test on :		
Test off :		
Result :		

Destinataire : A l'attention de PANSM

Export officiel non contrôlé
Motif : Batch release procedure

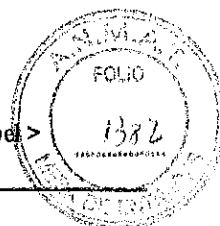
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Categoría : 1





Haemagglutinin Antigen Identification

Method : Haemagglutination inhibition test
Date :

Antisera used : = B/Yamagata lineage
= A/H1N1
= A/H3N2
= B/Yamagata lineage

Antigens used : = B/Victoria lineage
= A/H1N1
= A/H3N2
= B/Yamagata lineage

Result:

ANTIGEN	ANTISERA			
	ANTI B/Victoria /	ANTI H1 A/	ANTI H3 A/	ANTI B/Yamagata
B/Victoria				
A/H1N1				
A/H3N2				
B/Yamagata				
WORKING VIRUS SEED LOT				

Conclusion

Desimulair - A. Falcantoni de FANSM

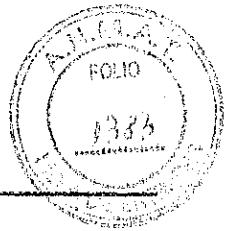
Export officiel non contrôlé
Mont F. Batch release 16/06/2016

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Infectious titer

Method : Haemagglutination (HA) System
used : Embryonated hen eggs Number
of tests :
Test on : Test
off :

Virus concentration per ml
expressed in log 10 : and Infectious Dose 50/ml

Mean of the titers : Infectious Dose 50/ml

Destinataire : A l'attention de FANSM

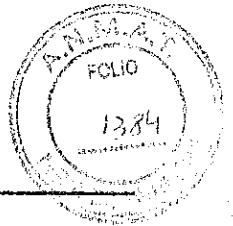
Export officiel non contrôlé
Motif : Batch release procédure

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Certificate of Analysis

Product Name : INFLUENZA VACCINE (SPLIT VIRION. INACTIVATED)

Process Stage : PRIMARY SEED LOT

Volume / Quantity :

Storage Condition : ≤ -60°C

Date of Manufacture :

Master Seed Lot

Test(s)	Acceptance Criteria	Result	Approval Date
HAEMAGGLUTININ ANTIGEN IDENTIFICATION <i>By Haemagglutination inhibition</i>	POSITIVE		

Export officiel non contrôlé
Motif : Batch Release protocol QIV
Destinataire : A l'attention de l'ANSM

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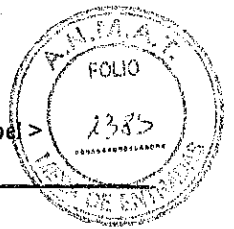
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Catégorie : 1





**INFLUENZA VACCINE (SPLIT VIRION, INACTIVATED)
MASTER VIRUS SEED LOT**

ORIGINAL STRAIN

Name : B/Victoria lineage
Origin :
Lot number :
Received :
Passage history on receipt :

**MASTER VIRUS SEED LOT
A - PRODUCTION**

Lot number :

Inoculation

Date of inoculation :
Passage number from reassortant received :
Dilutions of inoculum (log10) :
Eggs used :
Number of eggs per dilution :
Inoculated volume per egg . ml
Temperature of incubation : °C

Harvest of allantoic fluids

Date of harvest :
Pool of allantoic fluids at the dilution (log10) :
Filling and storage at $\leq - 60^{\circ}\text{C}$

Release date :

Des-antaire : A l'attention de l'ANSM
Export officiel non contrôlé
Motif : Batch Release protocol QIV

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Catégorie : 1

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B -- CONTROLS

Haemagglutinin Antigen Identification

Method Haemagglutination inhibition test

Date

Antisera used : = B/Victoria lineage
= A/H1N1
= A/H3N2
= B/Yamagata lineage

Antigens used : = B/Victoria lineage
= A/H1N1
= A/H3N2
= B/Yamagata lineage

Result:

ANTIGEN	ANTISERA			
	ANTI B/Victoria	ANTI H1 A/	ANTI H3 A/	ANTI B/Yamagata
B/Victoria				
A/H1N1				
A/H3N2				
B/Yamagata				
MASTER VIRUS SEED LOT				

Conclusion

Destinataire : A l'attention de l'ANSM

Export officiel non contrôlé
Motif : Batch Release protocol QV

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Número : Q_0565202
Version : 2.0
Categoría : 1



Certificate of Analysis

Product Name : INFLUENZA VACCINE (SPLIT VIRION, INACTIVATED)

Process Stage : BULK

Volume / Quantity :

Storage Condition : +5°C+/-3°C

Date of Manufacture :

BULK

Expiry Date :

Test(s)	Acceptance Criteria	Result	Approval Date
OCTOXYNOL-9 CONTENT <i>By HPLC</i>	≤ 800 µg/mL	µg/mL	
HAEMAGGLUTININ ANTIGEN IDENTIFICATION <i>By Haemagglutination Inhibition</i>	Positive		
HAEMAGGLUTININ ANTIGEN CONTENT <i>By SRID</i>	For formulation Mean : No specification		
	Lower confidence limit (P=0.95)		
	Higher confidence limit (P=0.95)		
BACTERIAL AND FUNGAL STERILITY <i>By Membrane filtration</i>	No microbial growth		
RESIDUAL INFECTIOUS VIRUS <i>By Inoculation of eggs</i>	No residual infectious virus		
NEURAMINIDASE ANTIGEN IDENTIFICATION <i>By RT-PCR</i>	Positive (RT-PCR method performed on the first three monovalent bulk batches manufactured from any new working seed lot)		
NEURAMINIDASE ENZYMATIC ACTIVITY <i>By Enzymatic method</i>	Presence of neuraminidase activity (Enzymatic method performed on the three first batches manufactured from any new working seed lot)		

Export officiel non contrôlé - Destinataire : A. Patistion de FANSM
Motif : Batch Release protocol QIV

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