



U.S. Food and Drug Administration
Protecting and Promoting Public Health

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Vaccine Substances

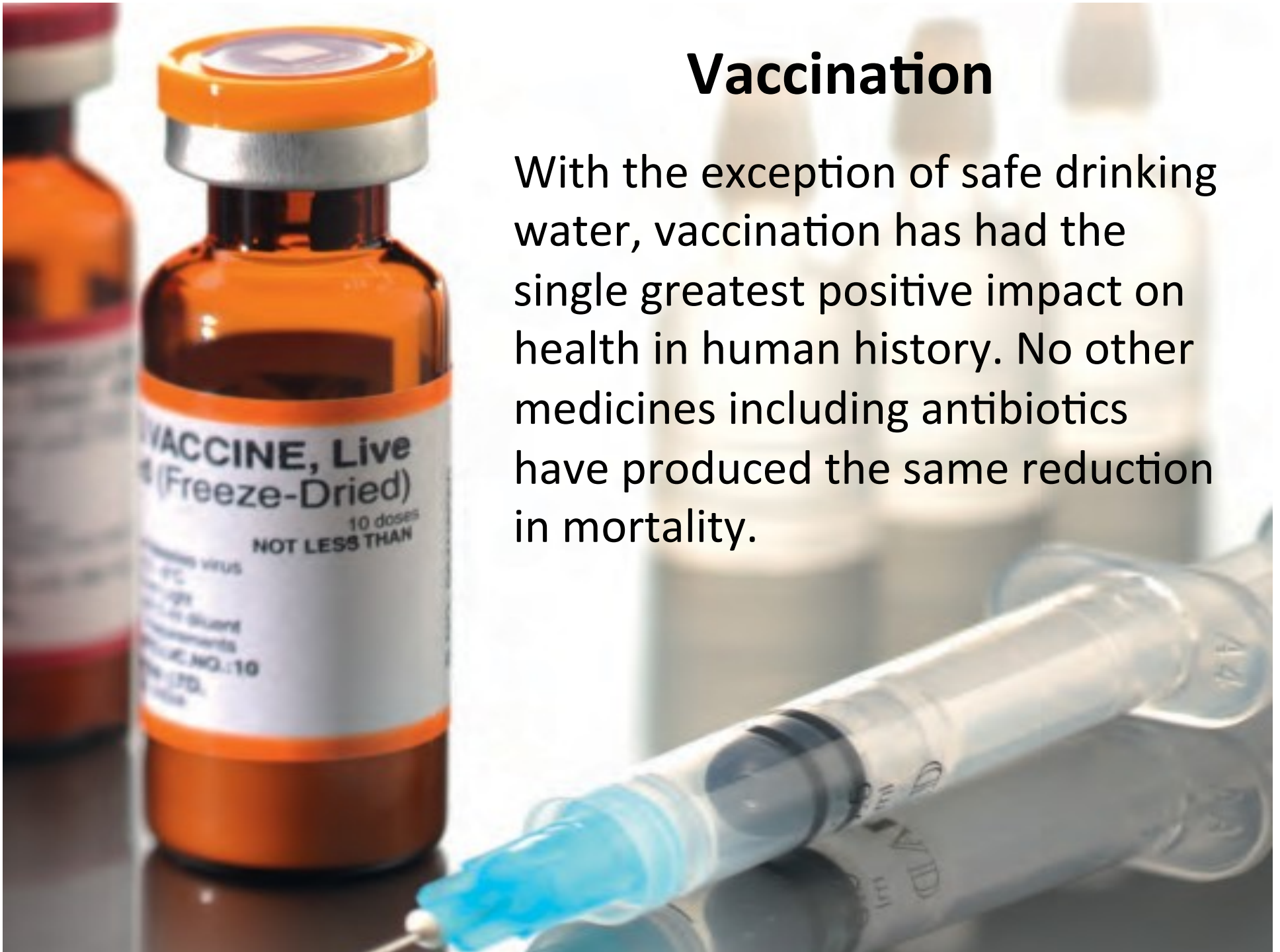
Frank Switzer, PhD
US Food & Drug Administration
frank,switzer@fda.hhs.gov



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Vaccination

With the exception of safe drinking water, vaccination has had the single greatest positive impact on health in human history. No other medicines including antibiotics have produced the same reduction in mortality.





What is a Vaccine?

- A biological preparation that provides active acquired immunity to a particular disease
- Presents itself to the immune system as a disease-causing microorganism
- Often made from weakened or killed forms of the microorganism or a specific cellular component (usually on the organism's surface)



What is a Vaccine?

It stimulates the body's immune system to:

- Recognize the agent as a threat
- Destroy it
- Keep a record of it so that the immune system can more easily recognize and destroy any of these microorganisms that it later encounters



Vaccine Cell Components

- Lysates
- Envelope Glycoprotein Mixtures
- Pure Cellular Proteins (Protein)
- Capsule Polysaccharides (Polymer)

Polysaccharide Types B & C

β -methylglycosides) and on supportive chemical evidence. The data indicate that the serogroup B polysaccharide is a 2 \rightarrow 8- α -linked homopolymer of sialic acid, identical in structure with colominic acid from *Escherichia coli*, whereas the de-*O*-acetylated serogroup C polysaccharide is a 2 \rightarrow 9- α -linked homopolymer. The native serogroup C polysaccharide is *O*-acetylated (1.16 mol of *O*-acetyl per sialic acid residue), all the *O*-acetyl substituents being located only at C-7 and C-8 of the sialic acid residues, and in addition contains unacetylated residues (24%). The polysaccharide contains di-*O*-acetylated residues (*O*-acetyl on C-7 and C-8), and at least one of the possible monoacetylated residues at C-7 or C-8.

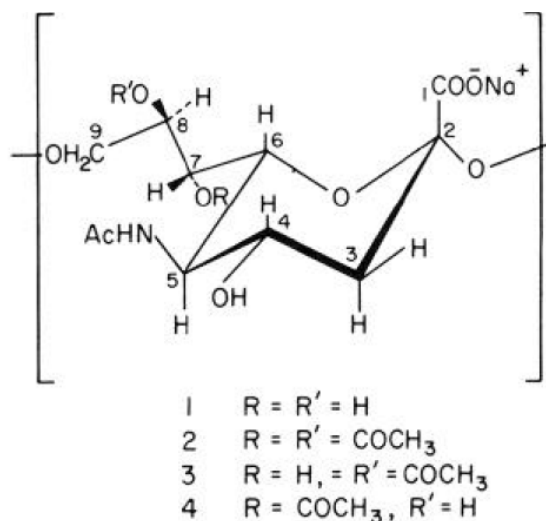
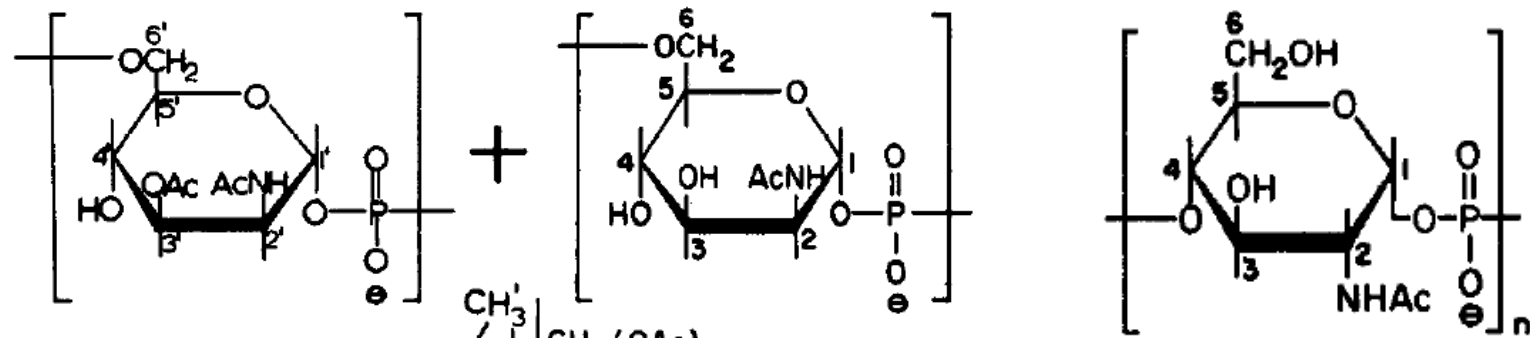


FIG. 3. The four different *N*-acetylneuraminic acid residues possible in the serogroup C polysaccharide.

Polysaccharide Types A and X



phorylated polysaccharides. The serogroup A polysaccharide has been shown to be essentially a 1→6-linked homopolymer of partially *O*-acetylated 2-acetamido-2-deoxymannosyl phosphate (8) and that of serogroup X to be a homopolymer of 2-acetamido-2-deoxyglucosyl phosphate linked either 1→3 or 1→4 (9). Due

Polysaccharide Type 11A

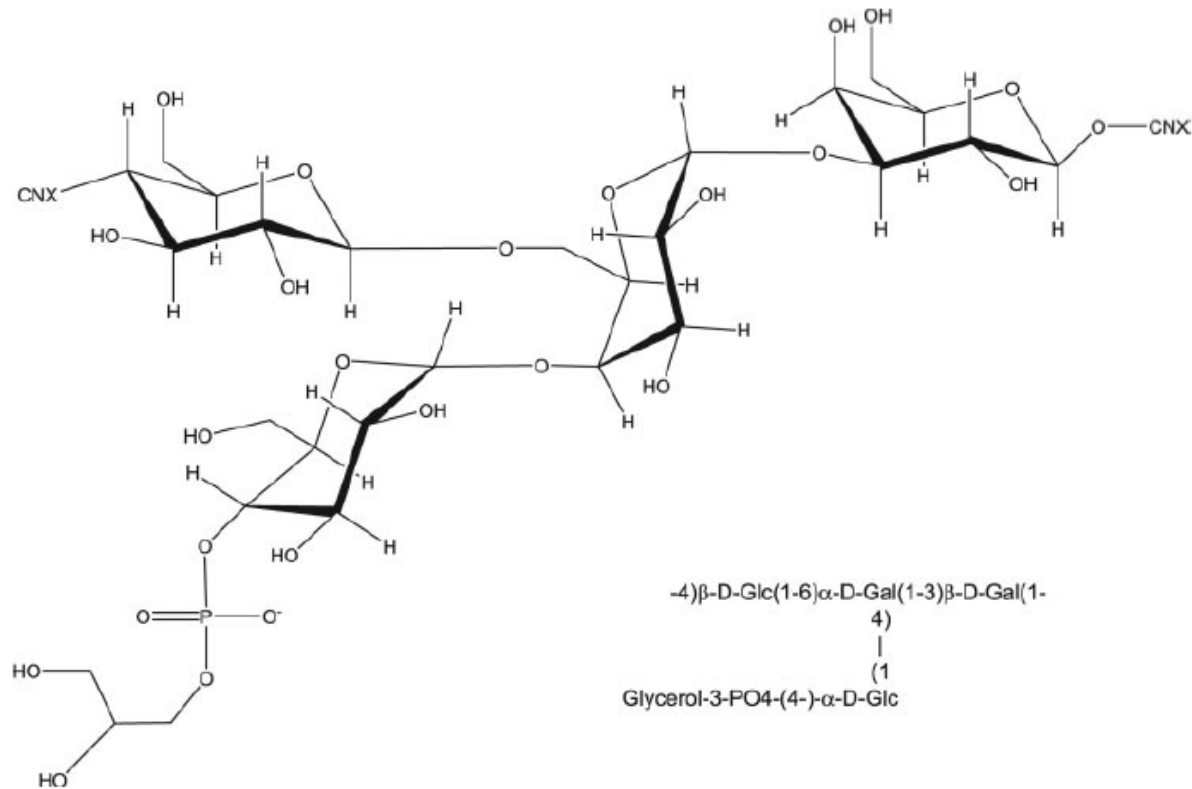


FIGURE 4. Revised structure of de-O-acetylated *S. pneumoniae* 11A PS.



Antigen Immunogenicity Modification

- Attenuated Strains
- Inactivation
- Conjugation



Live Attenuated Strains

- Produced by modifying a disease-producing (“wild”) virus or bacterium
- Resulting vaccine organism retains the ability to replicate and produce immunity but usually does not cause illness



Vaccine Microorganism Taxonomy

- Bacteria
 - ITIS/NCBI
 - Antigens usually described at Serotype/group level
 - Attenuated strains
- Viruses
 - ICTV (In Catalogue of Life)/NCBI
 - Antigens described at Strain level



Inactivated Antigens

- Agent/Physical modifications of (protein or structurally-diverse protein mixtures) antigens
- Crosslinking agents
 - Formaldehyde
 - Glutaraldehyde
 - Propiolactone
 - Ultraviolet light
 - Heat



Conjugated Antigens

- Used if the cellular component is not sufficiently antigenic
- Conjugate is typically a toxoid protein
 - Inactivated toxin (inactivation agent)
 - Mutated toxin (CRM197)
- May or may not be able to describe the details of how the conjugating toxoid is bonded to the antigen

Subunits 2



Subunit 1

10	20	30	40	50	60
SLTDLGGEL	IKIKNEDLTF	IAEKNSFSEE	PFQDEIVSYN	TKNKPLNFNY	SLDKIIVDYN
70	80	90	100	110	120
LQSKITLPND	RTPPVTKGIP	YAPEYKSNAA	STIEIHNIDD	NTIYQYLYAQ	KSP TTLQRIT
130	140	150	160	170	180
MTNSVDDALI	NSTKIYSYFP	SVISKVNQGA	QGILFLQWVR	DIIDDFTNES	SQKTTIDKIS
190	200	210	220	230	240
DVSTIVPYIG	PALNIVKQGY	EGNFIGALET	TGVVLLLEYI	PEITLPVIAA	LSIAESSTQK
250	260	270	280	290	300
EKIIKTIDNF	LEKRYEKWIE	VYKLVKAKWL	GTVNTQFQKR	SYQMYRSLEY	QVDAIKKIID
310	320	330	340	350	360
YEYKIYSGPD	KEQIADENN	LKNKLEEKAN	KAMININIFM	RESSRSFLVN	QMINEAKKQL
370	380	390	400	410	420
LEFDTQSKNI	LMQYIKANSK	FIGITELKKL	ESKINKVFST	PIPFYSYKNL	DCWVDNEEDI
430	440	450	460	470	480
DVILKKSTIL	NLDINNDIIS	DISGFNSSVI	TYPDAQLVPG	INGKAIHLVN	NESSEVIVHK
490	500	510	520	530	540
AMDIEYNDMF	NNFTVSFWLR	VPKVSASHLE	QYGTNEYSII	SSMKKHSLSI	SGGWSVSLKG
550	560	570	580	590	600
NNLIWTLKDS	AGEVRQITFR	DLPDKFNAYL	ANKWVFITIT	NDRLSSANLY	INGVLMGSAE
610	620	630	640	650	660
ITGLGAIED	NNITLKLDR	NNNNQYVSID	KFRIFKALN	PKEIEKLYTS	YLSITFLRDF
670	680	690	700	710	720
WGNPLRYDTE	YYLIPVASSS	KDVQLKNITD	YMYLTNAPSY	TNGKLNIIYYR	RLYNGLKFI
730	740	750	760	770	780
KRYTPNNEID	SFVKSGDFIK	LYVSYNNNEH	IVGYPKDGNA	FNNLDRIILRV	GYNAPGIPLY
790	800	810	820	830	840
KKMEAVKL RD	LKTYSVQLKL	YDDKNASLGL	VGTHNGQIGN	DPNRDILIAS	NWYFNHLKDK
850	860				
ILGCDWYFVP	TDEGWTND				

Subunit 2

10	20	30	40	50	60
PITINNFRYS	DPVNNDTIIM	MEPPYCKGLD	IYYKAFKITD	RIWIVPERYE	FGTKPEDFNP
70	80	90	100	110	120
PSSLIEGASE	YYDPNYLR TD	SDKDRFLQTM	VKLFNRIKNN	VAGEALLDKI	INAIPLYGNS
130	140	150	160	170	180
YSLLDKFD TN	SNSVSFNLE	QDPSGATTKS	AMLTNLIIFG	PGPVLNKNEV	RGIVLRVDNK
190	200	210	220	230	240
NYFPCRDGFG	SIMQMAFCPE	YVPTFDNVIE	NITSLTIGKS	KYFQDPALLL	MHELIVLHG
250	260	270	280	290	300
LYGMQVSSHE	IIPSKQEIYM	QHTYPISAEE	LFTFGGQDAN	LISIDIKNDL	YEKTLNDYKA
310	320	330	340	350	360
IANKLSQVTS	CNDPNIDIDS	YKQIYQKKYQ	FDKDSNGQYI	VNEDKFQILY	NSIMYGFT EI
370	380	390	400	410	420
ELGKKFN IKT	RLSYFSMNH D	PVKIPNLLDD	TIYNDTEGFN	IESKDLKSEY	KGQNM RVNTN
430	440	450	460		
AFRNVDGSL	VSKLIGLKK	IIPPTNIREN	LYNR TA		

Disulfide Links 2

Residue Index

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1_ 620

2_ 438

1_ 10

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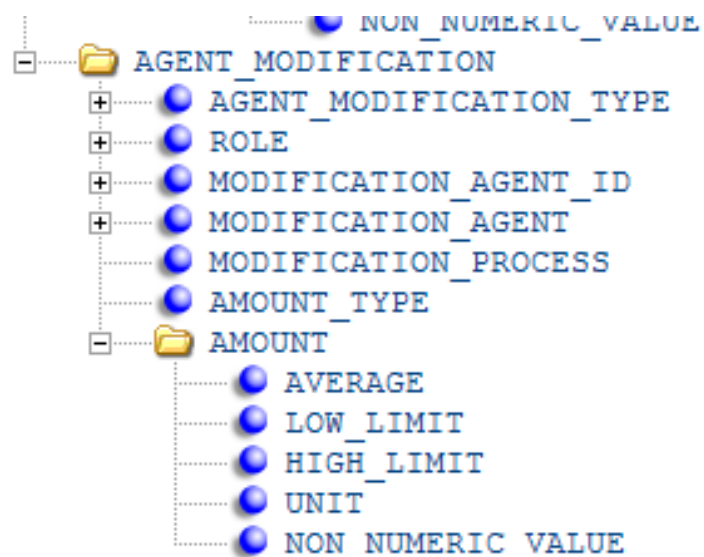


Vaccine Products

- Most are polyvalent - multiple antigens
 - Antigens focused on different serotypes of the same organism
 - Antigens to vaccinate against different diseases
 - All are distinct active ingredients
- Many contain also adjuvating agents that increase antigen immunogenicity

ACTIVE INGREDIENT/ACTIVE MOIETY		
Ingredient Name	Basis of Strength	Strength
INFLUENZA A VIRUS A/CHRISTCHURCH/16/2010 NIB-74XP (H1N1) ANTIGEN (FORMALDEHYDE INACTIVATED) (UNII: F6VKT8IL26) (INFLUENZA A VIRUS A/CHRISTCHURCH/16/2010 NIB-74XP (H1N1) HEMAGGLUTININ ANTIGEN (FORMALDEHYDE INACTIVATED) - UNII:9PM6202D07)	INFLUENZA A VIRUS A/CHRISTCHURCH/16/2010 NIB-74XP (H1N1) ANTIGEN (FORMALDEHYDE INACTIVATED)	15 ug in 0.5 mL
INFLUENZA A VIRUS A/TEXAS/50/2012 X-223A (H3N2) ANTIGEN (FORMALDEHYDE INACTIVATED) (UNII: 4127F11825) (INFLUENZA A VIRUS A/TEXAS/50/2012 X-223A (H3N2) HEMAGGLUTININ ANTIGEN (FORMALDEHYDE INACTIVATED) - UNII:S477VIE5TZ)	INFLUENZA A VIRUS A/TEXAS/50/2012 X-223A (H3N2) ANTIGEN (FORMALDEHYDE INACTIVATED)	15 ug in 0.5 mL
INFLUENZA B VIRUS B/MASSACHUSETTS/2/2012 BX-51B ANTIGEN (FORMALDEHYDE INACTIVATED) (UNII: 3W4UKF8P6Z) (INFLUENZA B VIRUS B/MASSACHUSETTS/2/2012 BX-51B HEMAGGLUTININ ANTIGEN (FORMALDEHYDE INACTIVATED) - UNII:N784VJP7ZF)	INFLUENZA B VIRUS B/MASSACHUSETTS/2/2012 BX-51B ANTIGEN (FORMALDEHYDE INACTIVATED)	15 ug in 0.5 mL
INFLUENZA B VIRUS B/BRISBANE/60/2008 ANTIGEN (FORMALDEHYDE INACTIVATED) (UNII: W45Z4CJE2J) (INFLUENZA B VIRUS B/BRISBANE/60/2008 HEMAGGLUTININ ANTIGEN (FORMALDEHYDE INACTIVATED) - UNII:166D8OWM7B)	INFLUENZA B VIRUS B/BRISBANE/60/2008 ANTIGEN (FORMALDEHYDE INACTIVATED)	15 ug in 0.5 mL

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SOURCE_MATERIAL	
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+ SOURCE_MATERIAL_TYPE	VIRUS
SOURCE_MATERIAL_STATE	
+ PARENT_SUBSTANCE_ID	9306C9VO2C
+ PARENT_SUBSTANCE_NAME	INFLUENZA A VIRUS H1N1
DEVELOPMENTAL_STAGE	
PART_GROUP	
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FRACTION	
MATERIAL_TYPE	
+ FRACTION	AVIAN TYPE GLYCOPROTEIN ANTIGENS
ORGANISM	
KINGDOM	
PHYLUM	
CLASS	
ORDER	
FAMILY	
GENUS	
SPECIES	
HYBRID_SPECIES_MATERNAL_ORGANISM	
HYBRID_SPECIES_PATERNAL_ORGANISM	
HYBRID_TYPE	
+ INFRASPECIFIC_TYPE	STRAIN
+ INFRASPECIFIC_DESCRIPTION	A/CHRISTCHURCH/16/2010 NIB-74XP (H1N1)



CHEMICAL
INACTIVATION
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FORMALDEHYDE

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Accepted scientific name: *Influenzavirus a: Influenza A virus* ICTV (accepted name)

Synonyms: -


Common names: -

Classification:	Kingdom	Viruses	ICTV_MSL	LSID ▶
	Phylum	Not assigned	ICTV_MSL	LSID ▶
	Class	Not assigned	ICTV_MSL	LSID ▶
	Order	Not assigned	ICTV_MSL	LSID ▶
	Family	Orthomyxoviridae	ICTV_MSL	LSID ▶
	Genus	<i>Influenzavirus a:</i>	ICTV_MSL	LSID ▶

Distribution: -

Lifezones: -

Additional data: -

Source database: [ICTV_MSL](#), 1, Mar 2014  100% ★★★★★

Latest taxonomic scrutiny: A.M.Q.King, M.J.Adams, E.B. Carstens & E.J. Lefkowitz (Eds), Mar-2014

Online resource: <http://ictvonline.org/virusTaxonomy.asp?version=2013>