Medicines & Healthcare products Regulatory Agency



Influenza Reagent Influenza virus infectious BVR-11 NIBSC code: 19/172 Instructions for use (Version 5.0, Dated 19/11/2019)

1. INTENDED USE

Reagent 19/172 is prepared from BVR-11 (B/Victoria/705/2018 x B/Brisbane/46/2015) which was processed for freeze drying in 250µl volumes as described by Campbell, PJ, Journal of Biological Standardisation, 1974, 2,249-267. The derivation and known passage history of BVR-11 is attached.

2. CAUTION

This preparation is not for administration to humans or animals in the human food chain

The material is not of human or bovine origin. As with all materials of biological origin, this preparation should be regarded as potentially hazardous to health. It should be used and discarded according to your own laboratory's safety procedures. Such safety procedures should include the wearing of protective gloves and avoiding the generation of aerosols. Care should be exercised in opening ampoules or vials, to avoid cuts.

3. UNITAGE

No unitage is assigned to this material

4. CONTENTS

Country of origin of biological material: United Kingdom. Each ampoule contains 250µl (nominal) of infectious influenza virus as allantoic fluid from SPF embryonated hen's eggs.

5. STORAGE

Store in the dark at -20°C or below

Please note: because of the inherent stability of lyophilized material, NIBSC may ship these materials at ambient temperature.

6. DIRECTIONS FOR OPENING

Vials have a 'flip-up' circular cap. Either on the cap or the collar of the vial, there is an indication of the point at which to lever off the cap. This exposes an area of the stopper through which reconstitution and withdrawal of the preparation can be made using a hypodermic needle and syringe. If use of a pipette is preferred, then fully remove the metal collar using, for example, forceps, taking care to avoid cuts by wearing appropriate gloves. Remove the stopper for access. Care should be taken to prevent loss of the contents.

7. USE OF MATERIAL

Reconstitute the contents of one ampoule of reagent with 250µl of sterile distilled water. Leave for a minimum of 5 minutes before use to allow for complete solution of freeze-dried material. A range of dilutions (e.g. 10^{-3} to 10^{-5}) should be made in a suitable medium for initial cultivation.

8. STABILITY

Reference Materials should be stored on receipt as indicated on the label.

NIBSC follows the policy of WHO with respect to its reference materials.

9. REFERENCES

NA



Potters Bar, Hertfordshire, EN6 3QG. T +44 (0)1707 641000, nibsc.org WHO International Laboratory for Biological Standards, UK Official Medicines Control Laboratory

10. ACKNOWLEDGEMENTS

NA

11. FURTHER INFORMATION

Further information can be obtained as follows; This material: enquiries@nibsc.org WHO Biological Standards: http://www.who.int/biologicals/en/ JCTLM Higher order reference materials: http://www.bipm.org/en/committees/jc/jctlm/ Derivation of International Units: http://www.nibsc.org/standardisation/international_standards.aspx Ordering standards from NIBSC: http://www.nibsc.org/products/ordering.aspx NIBSC Terms & Conditions: http://www.nibsc.org/terms_and_conditions.aspx

12. CUSTOMER FEEDBACK

Customers are encouraged to provide feedback on the suitability or use of the material provided or other aspects of our service. Please send any comments to enquiries@nibsc.org

13. CITATION

In all publications, including data sheets, in which this material is referenced, it is important that the preparation's title, its status, the NIBSC code number, and the name and address of NIBSC are cited and cited correctly.

14. MATERIAL SAFETY SHEET

Classification in accordance with Directive 2000/54/EC, Regulation (EC) No 1272/2008: Not applicable or not classified

Physical and Chemical properties					
Physical appeara white powder	nce:	Corrosive:	No		
Stable:	Yes	Oxidising:	No		
Hygroscopic:	No	Irritant:	No		
Flammable:	No	Handling:See	caution, Section 2		
Other (specify):	Live influenza virus				
Toxicological properties					

Effects of inhalation:	Likelihood of influenza virus infection	
Effects of ingestion:	Not established, avoid ingestion	
Effects of skin absorption:	Not established, avoid contact with skin	

Suggested First Aid

Inhalation:	Seek medical advice
Ingestion:	Seek medical advice
Contact with eyes:	Wash with copious amounts of water. Seek medical advice
Contact with skin:	Wash thoroughly with water.

Action on Spillage and Method of Disposal

Spillage of contents should be taken up with absorbent material wetted with an appropriate virucidal agent. Rinse area with an appropriate virucidal agent followed by water. Absorbent materials used to treat spillage should be treated as biologically hazardous waste.

15. LIABILITY AND LOSS

In the event that this document is translated into another language, the English language version shall prevail in the event of any inconsistencies between the documents.

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Unless expressly stated otherwise by NIBSC, NIBSC's Standard Terms and Conditions for the Supply of Materials (available at http://www.nibsc.org/About_Us/Terms_and_Conditions.aspx or upon request by the Recipient) ("Conditions") apply to the exclusion of all other terms and are hereby incorporated into this document by reference. The Recipient's attention is drawn in particular to the provisions of clause 11 of the Conditions.

16. INFORMATION FOR CUSTOMS USE ONLY

Country of origin for customs purposes*: United Kingdom * Defined as the country where the goods have been produced and/or sufficiently processed to be classed as originating from the country of supply, for example a change of state such as freeze-drying. Net weight: 0.25g per ampoule Toxicity Statement: Non-toxic Veterinary certificate or other statement if applicable.

Attached: No







Derivation of BVR-11 B/Victoria/705/2018 – High Growth Reassortant

B/Victoria/705/2018 (BVR-11) is a B high growth reassortant influenza virus.

PREPARATION

The preparation of B/Victoria/705/2018 (BVR-11) high growth reassortant influenza virus was conducted in R&D Influenza Operations Department at Sequrus.

The high yielding parent strain used was B/Brisbane/46/2015.

MATERIALS

The following materials of biological origin were used during the preparation of high growth reassortant BVR-11:

Virus Isolate:

The virus isolate was obtained from the WHO Collaborating Centre for Reference & Research on Influenza, Melbourne.

Supply details are:	
B/Victoria/705/2018	
WHO-CC Storage Lot number:	SL10015015
Passages prior to receipt at WHO-CC:	0
Passages undertaken in WHO-CC:	3

Eggs:

Specific Pathogen Free (SPF) eggs were used for all passages at Sequrus.

Antiserum:

Monoclonal Antibody BBR3.7E8.1B9 (15/03/19B) raised against influenza virus B/Brisbane/60/2008.

The MAb was prepared from murine hybridoma cells derived from mice immunized with inactivated B/Brisbane/60/2008 influenza virus.

Note on Transmissible Spongiform Encephalopathies (TSEs):

Australia and New Zealand have been declared TSE free in accordance with OIE guidelines. Detailed information on Australia's animal health status can be obtained from the following Animal Health Australia website link: http://www.animalhealthaustralia.com.au/programs/biosecurity

The MAb was gamma irradiated at 35kGrays prior to use.





REPORT

PASSAGE HISTORY:

Mixed infection passage:	B/Victoria/705/2018 wild type virus	HA titre $= 57$
	@10-3 x B/Brisbane/46/2015 @10-3	
	.↓	
1 st Antiserum Passage**	Inoculum @ 10-3 with MAb to	HA titre $= 320$
4	B/Brisbane/60/2008	
	\downarrow	
2 nd Antiserum Passage**	Inoculum @ 10-5 with Mab to	HA titre = 640
	B/Brisbane/60/2008)	
	\downarrow	
1 st Limit Dilution Passage	Inoculum @ 10-9	HA titre = 57
	\downarrow	
2 ^{ud} Limit Dilution Passage	Inoculum @ 10-8	HA titre $= 49$
	\downarrow	
3 rd Limit Dilution Passage	Inoculum @ 10-9	HA titre = 98
	\downarrow	
7 th Passage	Inoculum @ 10-5	Mean HA = 197

Total number of passages post mixed infection = 6Total number of passages since this virus was received from an approved laboratory = 7HA titres were determined using fowl red blood cells at room temperature.

** Virus sample diluted to 10-3 dilution was mixed with Mab to B/Brisbane/60/2008 and incubated for 1 hour at room temperature. Incubated virus/MAb sample was serially diluted and inoculated into eggs at indicated dilution.



Passage history of BVR-11 (post-mixed infection)

Passage	Lot	Laboratory
E4-E10		Seqirus, Australia
E11	VW10028796	Seqirus, Australia
E12	44830	NIBSC, Hertfordshire, UK

Sterility: No visible contamination was detected in a variety of media (tryptose soya broth, thioglycolate broth, Sabouraud's broth and blood agar plates) after 14 days incubation.

The HA and NA sequence of this virus is available at GISAID with the accession number EPI_ISL_397113.