WHO Reference Reagent
The 1st International Reference Preparation for Anti-Yellow Fever Serum, Monkey
NIBSC code: YF
Instructions for use
(Version 5.0, Dated 24/08/2011)

1. INTENDED USE
This material has been prepared and characterised by the Statens Serum Institut (SSI), Copenhagen, Denmark. With effect from 1st July 1997, the National Institute for Biological Standards and Control (NIBSC), Potters Bar, UK is the custodian and distributor of this material.

For details of this International Reference Preparation, please refer to the enclosed package insert from the Statens Serum Institut.

The preparation is labelled ‘Anti-yellow fever serum’.

The package insert from SSI 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
143 International units (IU) of Anti Yellow Fever Serum per ampoule.

4. CONTENTS
Country of origin of biological material: Denmark.

5. STORAGE
Store at -20° C.

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
No attempt should be made to weigh out any portion of the freeze-dried material prior to reconstitution.

8. STABILITY
Reference materials are held at NIBSC within assured, temperature-controlled storage facilities. 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
See attached package insert from SSI

10. ACKNOWLEDGEMENTS
Not applicable

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 1272/2006: Not applicable or not classified

| Physical appearance: Freeze dried powder | Corrosive: No |
| Stable: Yes | Oxidising: No |
| Hygroscopic: No | Irritant: No |
| Flammable: No | Handling: See caution, Section 2 |
| Other (specify): Contains material of monkey origin |

Toxicological properties
Effects of inhalation: Not established, avoid inhalation
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 ampoule contents should be taken up with absorbent material wetted with an appropriate disinfectant. Rinse area with an appropriate disinfectant followed by water. Absorbent materials used to treat spillage should be treated as biological 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.

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*: Denmark
| 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: 1.0g  |
| Toxicity Statement: Non-toxic  |
| Veterinary certificate or other statement if applicable.  |
| Attached: No  |

17. CERTIFICATE OF ANALYSIS
NIBSC does not provide a Certificate of Analysis for WHO Biological Reference Materials because they are internationally recognised primary reference materials fully described in the instructions for use. The reference materials are established according to the WHO Recommendations for the preparation, characterization and establishment of international and other biological reference standards http://www.who.int/bloodproducts/publications/TRS932Annex2_Inter_biolstandardsrev2004.pdf (revised 2004). They are officially endorsed by the WHO Expert Committee on Biological Standardization (ECBS) based on the report of the international collaborative study which established their suitability for the intended use.
THE INTERNATIONAL REFERENCE PREPARATION
of
ANTI-YELLOW FEVER SERUM, MONKEY
(1st international reference preparation)

1. THE REFERENCE PREPARATION
The reference preparation was established in 1962\textsuperscript{1,2}. It is prepared from a
pool of sera from three monkeys (Cercopithecus aethiops tanta) immunized
against the pantropic Asibi strain of yellow fever virus. The serum pool was
prepared in the laboratory of the West African Council for Medical Research,
Lagos, Nigeria.

2. AMPOULE CONTENTS
The pool was distributed into ampoules (1 ml serum/ampoule) and freeze-dried.
By definition\textsuperscript{4} the total contents of each ampoule contains 143 International
Units (IU) of Anti-Yellow Fever Serum.

3. USE OF THE STANDARD
The primary purpose for this preparation was to use it as a positive control
serum in the mouse protection test used in the control of yellow fever
vaccine.

The preparation was examined in an international collaborative assay in 11
laboratories in 10 countries\textsuperscript{a}.

An IU was defined because it was found, that the reference preparation was
also useful in potency assays of anti-yellow-fever sera.

4. GENERAL REMARKS ABOUT INTERNATIONAL REFERENCE MATERIALS
International biological standards and international biological reference
reagents provide a means of ensuring uniformity throughout the world in the
designation of the potency or activity of preparations used in the prophylax-
ics, therapy, or diagnosis of disease, where this cannot be expressed in terms
of physical or chemical quantities. The International Units are units of
quantities of "effective constituent"\textsuperscript{3}.

The standard is the material as it exists in the ampoules; the "material" thus
includes the effective constituents together with all the other constituents
that may be present (moisture, carrier, buffer, salt etc., according to the
form in which the standard is available).

International biological reference materials are intended for use in the
calibration of the contents of "effective constituent" in national or working
standard preparations and for the expression of these contents in the
respective International Units. For the routine use in the laboratory the
national or working standards should be used in order to save as much as
possible the international reference materials. These are only sent to
individual laboratories in very limited amounts. The preparations are sent
free of charges but sometimes a small charge might be claimed for the air-
freighting.

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YF

Statens Seruminstitut · S. Artillerivej · DK-2300 Copenhagen S · Denmark
Phone +45 3268 3268 · Telex 31316 serum.dk · Telefax +45 3268 3868

National Institute for Biological Standards and Control, Potters Bar, Hertfordshire, EN6 3QG. Tel 44 (0)1707 641000, nibsc.org
WHO International Laboratory for Biological Standards, UK Official Medicines Control Laboratory
5. REFERENCES

