

# TABLE OF EXTRANEOUS AGENTS TO BE TESTED FOR IN RELATION TO THE GENERAL AND SPECIES SPECIFIC GUIDELINES ON PRODUCTION AND CONTROL OF MAMMALIAN VETERINARY VACCINES

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| Guideline Title          | Table of Extraneous Agents to be Tested for in relation to the General and Species Specific Guidelines on Production and Control of Mammalian Veterinary Vaccines  |
| Legislative Basis        | Directive 81/852/EEC as amended  |
| Date of First Adoption   | prior to September 1994  |
| Date of Entry into Force | prior to September 1994  |
| Status                   | Last revised September 1994  |
| Previous Titles          | None   |
| Other References         |  |
| Additional Notes         | Testing for extraneous agents is required in a number of the general and specific documents detailing requirements for vaccines. This note takes the form of a table containing a list of agents which must be taken into account when considering what extraneous agents testing is appropriate. The presence of an agent on the list does not mean that a specific test for that agent must be carried out. However, the applicant must provide in the dossier, a justification for not carrying out a specific test for an agent. |

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# TABLE OF EXTRANEEOUS AGENTS TO BE TESTED FOR IN RELATION TO THE GENERAL AND SPECIES-SPECIFIC GUIDELINES ON PRODUCTION AND CONTROL OF MAMMALIAN VETERINARY VACCINES

## 1. EXPLANATORY NOTE

There are a number of paragraphs in the GRIMV and GRLMV documents and the species - specific guidelines that require testing for extraneous agents. The following table contains a list of agents which must be taken into account when considering what extraneous agents testing is appropriate. The presence of an agent on the list does not mean that a specific test for that agent must be carried out. However, the applicant must provide in the dossier, a justification for not carrying out a specific test for an agent.

The types of justification that can be given include:

- a) Disease does not occur in country origin (and material could not have been contaminated by this agent, subsequently). Supporting documentary evidence must be provided.
- b) Substance in question cannot be contaminated with this agent, e.g. agent does not cross placenta.
- c) Where the applicant has shown that his general test can be shown to detect the specified agent.

For points a and b, these justifications may be given by referring to published literature.

The table is divided into sections by species. As indicated in the guidelines, consideration has to be given to the species of origin of the test material and the target species for the product. In addition to consideration of relevance of the agents listed in the table, the applicant must also take into account changes in the disease situation in the country of origin of materials and, particularly, take note of new emerging disease.

## 2. TABLE

Explanation of the following table

- SPF herd: include serological tests, clinical examination and, if necessary, detection tests for microbiological agents (e.g. GRIMV 1.2.3, GRLMV 1.2.3, Bovine vaccines guideline 1.2.3).
- General tests: Are those where it is specified that the appropriate cell cultures are examined for cytopathic effect and for haemadsorption as described in general guidelines (e.g. GRIMV and GRMLV 1.2.2.1.4 – 2.2.1.4.2 and 1.2.2.1.4.4, 1.2.3.1.4 – 1.2.3.1.4.2 and 1.2.3.1.4.4, 1.3.5).
- Specified tests: Are carried out by immunofluorescence or other test methods as indicated in the guidelines (e.g. GRIMV and GRLMV 1.1.4.1, 1.2.2.1.4.3 and 1.2.2.1.4.4, 1.2.3.1.4.3 and 1.2.3.1.4.4, 1.3.5, Bovine vaccines guideline 1.2.2, 1.2.3).

| SPECIES                   | AGENT  | SPF HERD | VIRUS SEED    |                |
|---------------------------|--|----------|---------------|----------------|
|                           |  |          | GENERAL TESTS | SPECIFIC TESTS |
| BOVINE, OVINE AND CAPRINE | ADENOVIRUSES, SUBGROUPS 1 AND 2                                      | YES      | YES           |                |
| BOVINE, OVINE AND CAPRINE | AKABANE VIRUS  | YES      |               | YES            |
| BOVINE, OVINE AND CAPRINE | AUESZKY'S DISEASE VIRUS  |          | YES           |                |
| BOVINE, OVINE AND CAPRINE | BLUETONGUE VIRUS and EPIZOOTIC HEMORRAGIC DISEASE VIRUS              | YES      |               | YES            |
| BOVINE                    | BOVINE CORONAVIRUS   | YES      | YES           |                |
| BOVINE                    | BOVINE EPHEMERAL FEVER VIRUS   | YES      |               |                |
| BOVINE, OVINE AND CAPRINE | BOVINE HERPESVIRUSES 1, 2, 4   | YES      | YES           |                |
| BOVINE, OVINE             | BOVINE LEUKEMIA VIRUS (ENZOOTIC BOVINE LEUKOSIS)                     | YES      |               | YES            |
| BOVINE, OVINE             | BOVINE PAPILLOMAVIRUSES  | YES      |               |                |
| BOVINE                    | BOVINE PARVOVIRUS  | YES      | YES           |                |
| BOVINE                    | BOVINE PAPULAR STOMATITIS VIRUS, PSEUDOCOWPOXVIRUS                   | YES      | YES           |                |
| BOVINE                    | BOVINE RESPIRATORY SYNCYTIAL VIRUS                                   | YES      |               | YES            |
| BOVINE                    | BOVINE ROTAVIRUS   | YES      |               |                |
| BOVINE AND OVINE          | BOVINE VIRAL DIARRHOEA VIRUS   | YES      |               | YES            |
| BOVINE                    | BRUCELLA ABORTUS   | YES      |               | YES            |
| BOVINE, OVINE AND CAPRINE | COWPOXVIRUS, VACCINIA VIRUS  | YES      | YES           |                |
| BOVINE, OVINE AND CAPRINE | FOOT AND MOUTH DISEASE VIRUS, A, O, C, ASIAI, SATI, SAT1, SAT2, SAT3 | YES      |               | YES            |
| BOVINE, OVINE AND CAPRINE | LEPTOSPIRA SPP.  | YES      |               |                |
| BOVINE                    | LUMPY SKIN DISEASE VIRUS   | YES      | YES           |                |
| BOVINE                    | MALIGNANT CATARRHAL FEVER (AFRICAN FORM) VIRUS                       | YES      |               |                |
| BOVINE                    | MALIGNANT CATARRHAL FEVER (EUROPEAN FORM)                            | YES      |               |                |
| BOVINE, OVINE AND CAPRINE | MYCOBACTERIUM TUBERCULOSIS AND PARATUBERCULOSIS                      | YES      |               | YES            |
| BOVINE, OVINE AND CAPRINE | MYCOPLASMA SP.   | YES      |               | YES            |
| BOVINE, OVINE AND CAPRINE | PARAINFLUENZA 3 VIRUS  | YES      | YES           |                |
| BOVINE                    | COXIELLA BURNETTI (Q-FEVER)  | YES      | YES           |                |
| BOVINE                    | RABIES VIRUS   | YES      |               | YES            |
| BOVINE, OVINE AND CAPRINE | RIFT VALLEY FEVER VIRUS  | YES      |               | YES            |
| BOVINE                    | RINDERPEST VIRUS   | YES      | YES           |                |
| BOVINE, OVINE AND CAPRINE | SALMONELLA SP.   | YES      |               |                |
| BOVINE                    | VESICULAR STOMATITIS VIRUS, INDIANA AND NEW JERSEY                   | YES      | YES           |                |

| SPECIES           | AGENT  | SPF HERD | SUBSTANCES OF ANIMAL ORIGIN/CELL SUBSTRATES/VIRUS SEED |                |
|-------------------|--|----------|--|----------------|
|                   |  |          | GENERAL TESTS  | SPECIFIC TESTS |
| CAPRINE           | CAPRINE ARTHRITIS ENCEPHALITIS VIRUS             | YES      |  | YES            |
| CAPRINE           | CAPRINE HERPESVIRUS                              | YES      | YES  |                |
| OVINE             | BORDER DISEASE VIRUS                             | YES      |  | YES            |
| OVINE             | BORNA DISEASE VIRUS                              | YES      |  | YES            |
| OVINE             | BRUCELLA OVIS                                    | YES      |  | YES            |
| OVINE             | CHLAMYDIA OVIS                                   | YES      |  | YES            |
| OVINE             | JAAAGSIEKTE (OVINE PULMONARY ADENOMATOSIS VIRUS) | YES      |  |                |
| OVINE             | LOUPING ILL VIRUS                                | YES      |  | YES            |
| OVINE             | NAIROBI SHEEP DISEASE VIRUS                      | YES      |  | YES            |
| OVINE             | ROSS RIVER VIRUS                                 | YES      | YES  |                |
| OVINE             | SCRAPIE  | YES(1)   |  |                |
| OVINE AND CAPRINE | BRUCELLA MELITENSIS                              | YES      |  | YES            |
| OVINE AND CAPRINE | ECTHYMA CONTAGIOSUM VIRUS (ORF VIRUS)            | YES      | YES  |                |
| OVINE AND CAPRINE | MAEDI VISNA VIRUS                                | YES      |  |                |
| OVINE AND CAPRINE | PESTE DES PETITS RUMINANTS VIRUS                 | YES      | YES  |                |

(1) SCRAPIE: Demonstration of freedom requires monitoring for absence of disease for at least two years and examination of brain material from cull animals by histology and/or examination for scrapie associated fibrils.

| SPECIES | AGENT   | SPF HERD | ORIGIN/CELL SUBSTRATES/VIRUS SEED |                |
|---------|---|----------|-----------------------------------|----------------|
|         |   |          | GENERAL TESTS                     | SPECIFIC TESTS |
| PORCINE | AFRICAN SWINE FEVER VIRUS   | YES      |                                   | YES            |
|         | AUESZKY'S DISEASE VIRUS   | YES      | YES                               |                |
|         | ACTINOBACILLUS PLEUROPNEUMONIAE   | YES      |                                   |                |
|         | BORDETELLA BRONCHISEPTICA   | YES      |                                   | YES            |
|         | BOVINE VIRAL DIARRHOEA VIRUS  | YES      |                                   | YES            |
|         | BRUCELLA SUIIS  | YES      |                                   | YES            |
|         | CLASSICAL SWINE FEVER VIRUS   | YES      |                                   | YES            |
|         | ENCEPHALOMYOCARDITIS VIRUS  | YES      | YES                               |                |
|         | ERYSPELOTHRIX RHUSIOPATHIAE   | YES      |                                   |                |
|         | FOOT AND MOUTH DISEASE VIRUS A, O, C, ASIAT, SAT1, SAT2, SAT3           | YES      |                                   | YES            |
|         | HAEMAGGLUTININATING ENCEPHALOMYELITIS VIRUS                             | YES      | YES                               |                |
|         | LEPTOSPIRA SPP  | YES      |                                   |                |
|         | MYCOPLASMA HYOPNEUMONIAE  | YES      |                                   | YES            |
|         | MYCOPLASMA HYORHINIS  | YES      |                                   | YES            |
|         | PASTEURELLA MULTOCIDA   | YES      |                                   |                |
|         | TRANSMISSIBLE GASTROENTERITIS VIRUS and PORCINE RESPIRATORY CORONAVIRUS | YES      | YES                               |                |
|         | PORCINE ADENOVIRUSES  | YES      | YES                               |                |
|         | PORCINE CYTOMEGALOVIRUS   | YES      |                                   | YES            |
|         | PORCINE EPIDEMIC DIARRHOEA VIRUS  | YES      |                                   |                |
|         | PORCINE ENTEROVIRUSES (including TESCHENT-ALFEN)                        | YES      | YES                               |                |
|         | PORCINE INFLUENZA VIRUS   | YES      |                                   | YES            |
|         | PORCINE PARVOVIRUS  | YES      |                                   | YES            |
|         | PORCINE RESPIRATORY AND REPRODUCTIVE SYNDROME VIRUS                     | YES      |                                   | YES            |
|         | PORCINE ROTAVIRUS   | YES      |                                   |                |
|         | PORCINE VESICULAR EXANTHEMA VIRUS                                       | YES      | YES                               |                |
|         | RABIES VIRUS  | YES      |                                   | YES            |
|         | SWINE POXVIRUS  | YES      | YES                               |                |
|         | SWINE VESICULAR DISEASE VIRUS   | YES      | YES                               |                |
|         | VESICULAR STOMATITIS VIRUS  | YES      |                                   | YES            |

| SPECIES | AGENT   | SPF HERD | SUBSTANCES OF ANIMAL ORIGIN/CELL SUBSTRATES/VIRUS SEED |                |
|---------|---|----------|--|----------------|
|         |   |          | GENERAL TEST   | SPECIFIC TESTS |
| EQUINE  | AFRICAN HORSE SICKNESS VIRUS                                    | YES      |  | YES            |
|         | BORNA DISEASE VIRUS   | YES      |  |                |
|         | EQUINE ARTERITIS VIRUS  | YES      | YES  |                |
|         | EQUINE ENCEPHALOMYELITIS VIRUSES EASTERN, WESTERN, VENEZUELLIAN | YES      |  | YES            |
|         | EQUINE HERPES VIRUSES 1, 2, 3, 4,                               | YES      | YES  |                |
|         | EQUINE INFECTIOUS ANEMIA VIRUS                                  | YES      |  | YES            |
|         | EQUINE INFLUENZA VIRUS  | YES      |  | YES            |
|         | JAPANESE B ENCEPHALITIS VIRUS                                   | YES      |  | YES            |
|         | MYCOBACTERIUM PSEUDOMALLEI                                      | YES      |  |                |
|         | RABIES VIRUS  | YES      |  | YES            |
|         | STREPTOCOCCUS EQUI  | YES      |  |                |
|         | TRYPANOSOMA SP.   | YES      |  |                |
|         | VESICULAR STOMATITIS VIRUS                                      | YES      |  | YES            |
| FELINE  | AUESZKY'S DISEASE VIRUS   |          | YES  |                |
|         | CHLAMYDIA PSITTACI  | YES      |  | YES            |
|         | COWPOXVIRUS   | YES      | YES  |                |
|         | FELINE CALICIVIRUS  | YES      | YES  |                |
|         | FELINE HERPESVIRUS 1  | YES      | YES  |                |
|         | FELINE IMMUNODEFICIENCY VIRUS                                   | YES      |  | YES            |
|         | FELINE LEUKEMIA VIRUS/FELINE SARCOMA VIRUS                      | YES      |  | YES            |
|         | FELINE INFECTIOUS PERITONITIS VIRUS, FELINE ENTERIC CORONAVIRUS | YES      |  |                |
|         | FELINE PANLEUKOPENIA VIRUS                                      | YES      | YES  |                |
|         | FELINE SYNCYTTIA FORMING VIRUS                                  | YES      |  | YES            |
|         | RABIES VIRUS  | YES      |  | YES            |

| SPECIES           | AGENT   | SPF HERD | ORIGIN/CELL SUBSTRATES/VIRUS SEED |                |
|-------------------|---|----------|-----------------------------------|----------------|
|                   |   |          | GENERAL TESTS                     | SPECIFIC TESTS |
| <b>CANINE</b>     | AUESKY'S DISEASE VIRUS                          |          | YES                               |                |
|                   | BRUCELLA CANIS                                  | YES      |                                   | YES            |
|                   | CANINE ADENOVIRUS 1 AND 2                       | YES      | YES                               |                |
|                   | CANINE CORONAVIRUS                              | YES      | YES                               |                |
|                   | CANINE DISTEMPER VIRUS                          | YES      | YES                               |                |
|                   | CANINE HERPESVIRUS                              | YES      | YES                               |                |
|                   | CANINE PARAVOVIRUS                              | YES      | YES                               |                |
|                   | LEPTOSPIRA SPP.                                 | YES      |                                   |                |
|                   | PARAINFLUENZA 2 VIRUS (SV5)                     | YES      | YES                               |                |
| RABIES VIRUS      | YES   |          | YES                               |                |
| <b>RABBIT</b>     | ARENAVIRUS                                      | YES      |                                   | YES            |
|                   | AUESKY'S DISEASE VIRUS                          |          | YES                               |                |
|                   | ENCEPHALOMYOCARDITIS VIRUS                      | YES      | YES                               |                |
|                   | MYXOMA VIRUS, SHOPE FIBROMA VIRUS               | YES      |                                   | YES            |
|                   | RABBIT HEMORRAGIC DISEASE VIRUS                 | YES      | YES                               |                |
|                   | RABIES VIRUS                                    | YES      |                                   | YES            |
| <b>RODENT</b>     | ARENAVIRUS (LYMPHOCYTIC CHORIOMENINGITIS VIRUS) | YES      |                                   | YES            |
|                   | ENCEPHALOMYOCARDITIS VIRUS                      |          |                                   | YES            |
|                   | RABIES VIRUS                                    | YES      |                                   | YES            |
| <b>VERO CELLS</b> | BOVINE VIRAL DIARRHOEA VIRUS                    |          |                                   | YES            |
|                   | ENDOGENOUS RETROVIRUSES (C, D-TYPES PARTICLES)  |          | YES                               | YES            |
|                   | HERPESVIRUSES                                   |          |                                   | YES            |
|                   | MYCOPLASMA SPP.                                 |          | YES                               |                |
|                   | REOVIRUSES                                      |          | YES                               |                |
|                   | SV 40 VIRUS                                     |          | YES                               |                |
|                   | SV 5 VIRUS                                      |          | YES                               |                |