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| <b>Doc Number:</b><br><b>GLF-XX-ZZA</b> | <b>ANNEXURE G</b><br><b>CONSOLIDATED VARIATIONS GUIDELINE</b><br><b>(VETERINARY)</b> |  |
| Revision: 1.0                           |  | Effective date: <b>30 September 2025</b>  |

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## SCOPE

This annexure contains details on the specific categories and codes for variations specific to the safety and efficacy of registered medicines. This guideline is relevant only to submissions made for variations affecting the safety and efficacy of orthodox human medicines. It does not include veterinary, biological, and complementary medicines.

### Classification of clinical variations

The below classifications have been adapted from the EMA variation classification, article 5, xxxxxxxxxxxxxxxxxxxx.

The variations classifications for Type I and Type II changes are detailed below.

Type IA<sub>AN</sub> variations are implemented by the application and notification sent to SAHPRA within 12 month of implementation.

Type IA<sub>IN</sub> variations are immediately implementable; however, the authority should be notified within 30 working days.

Type IB variations become implementable after 67 working days, if no communication is received from SAHPRA.

If one or more conditions are not met, the applicant must classify the change using a higher code e.g. If a condition of type IA is not met, the variation is upgraded to a Type IB.

#### 4.2. Veterinary (C.II)

##### a) Summary of SAHPRA codes and variation classifications

All relevant C.I codes will apply to veterinary medicines as they do for human medicines – see section 4.1 of this addendum for the details of these exceptions. Note that any reference to a PIL for veterinary medicines is not applicable to SAHPRA – relevant codes should be interpreted with respect to the Label and PI only.

The adoption of C.II codes specific to veterinary medicines is explicitly covered below.

*Table 4.2.1 – summary of codes adopted/excluded for Veterinary*

| SAHPRA/EMA code | Code description | Exception type | Exception number |
|-----------------|------------------|----------------|------------------|
|-----------------|------------------|----------------|------------------|

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|-------------------|--|---|----------------|
| All C.I codes     | C.I codes relevant to veterinary medicines are adopted with the same exceptions detailed in section 4.1 of this addendum document  | Various                                     | 4.1.1 – 4.1.13 |
| C.II.1            | Variations concerning a change to or addition of a non-food producing target species   | None <sup>[1]</sup>                         | N/A            |
| C.II.2a – C.II.2b | Deletion of a food producing or non-food producing target species<br>b) Deletion as a result of a safety and/or efficacy issue<br>b) Deletion not resulting from a safety and/or efficacy issue                                    | Adopted with clarification                  | 4.2.1          |
| C.II.3            | Changes to the withdrawal period for a veterinary medicine   | None <sup>4</sup>                           | N/A            |
| C.II.4            | Variations concerning the replacement or addition of a serotype, strain, antigen or combination of serotypes, strains or antigens for veterinary vaccine relevant diseases listed on the “Controlled and Notifiable diseases list” | Alteration                                  | 4.2.2          |
| C.II.5            | Variations concerning the replacement of a strain for a veterinary vaccine against equine influenza  | Exclusion – combined with code C.II.4 above | 4.2.2          |
| C.II.6            | Changes to the label which are not connected with the PI   | None <sup>4</sup>                           | N/A            |
| C.II.7a – C.II.7b | Introduction of a new Pharmacovigilance system   | Exclusion                                   | 4.2.3          |
| C.II.8            | Change in the frequency and/or date of submission of period safety update reports (PSUR)   | None <sup>4</sup>                           | N/A            |

b) Code-related exceptions

|                         |   |                                  |                          |
|-------------------------|---|----------------------------------|--------------------------|
| <b>4.2.1</b>            | <b>Clarification</b>  |                                  |                          |
| <b>EMA/SAHPRA code</b>  | C.II.2a – C.II.2b   | <b>EMA/SAHPRA classification</b> | Type II (a); Type IB (b) |
| <b>Code description</b> | Deletion of a food producing or non-food producing target species<br>b) Deletion as a result of a safety and/or efficacy issue<br>b) Deletion not resulting from a safety and/or efficacy issue |                                  |                          |
| <b>Details</b>          | Code clarified to include deletions related to safety and efficacy issues.  |                                  |                          |

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| <b>4.2.2</b> | <b>Exception type</b>     | Alteration  | <b>EMA code</b>              | C.II.4  |
|              | <b>EMA classification</b> | Type II   | <b>SAHPRA classification</b> | Type II |
|              | <b>Code description</b>   | Variations concerning the replacement or addition of a serotype, strain, antigen or combination of serotypes, strains or antigens for veterinary vaccine relevant diseases listed on the “Controlled and Notifiable diseases list”                  |                              |         |
|              | <b>Details</b>            | SAHPRA has expanded the scope of diseases relevant to this code. Note that code C.II.5 falls away with an exclusion, as the content is now incorporated into code C.II.4 (i.e. there is no longer a need for a separate code for equine influenza). |                              |         |

|              |                           |   |                              |                   |
|--------------|---------------------------|---|------------------------------|-------------------|
| <b>4.2.3</b> | <b>Exception type</b>     | Exclusion   | <b>EMA code</b>              | C.II.7a – C.II.7b |
|              | <b>EMA classification</b> | Type II (a); Type IB (b)  | <b>SAHPRA classification</b> | N/A               |
|              | <b>Code description</b>   | Introduction of a new Pharmacovigilance system  |                              |                   |
|              | <b>Details</b>            | SAHPRA will only adopt this code once the PV function for Veterinary medicines has been formalised. |                              |                   |

[1] Code adopted as-is, without exceptions.