**Questionnaire for Risk Assessment of the Presence of Nitrosamine Precursors on Packaging Material Used in the Manufacture of Medicines**

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| **Material Commercial name** | Click here to enter text. |
| **Material General Name / Type** | Click here to enter text. |
| **Manufacturer Name** | Click here to enter text. |
| **Manufacturing Site Address** | Click here to enter text. |
|  |
| **Nitrosamines are substances classified as probable cancer-causing agents in humans. In 2018, the presence of these substances was detected in several blood pressure medications known as "sartans" and, in 2019, impurities of the same type were detected in batches of the active pharmaceutical ingredient ranitidine.****Therefore, in alignment with health authorities in several countries, ANVISA recommends a reassessment of the synthesis routes of the raw materials used in the manufacture of medicines to ensure that the amounts of nitrosamines in pharmaceutical products do not present risks to patients.****Still in 2019, the presence of nitrosamines was detected in medicines packaged in blisters, in which the occurrence of these substances was not expected, due to their formulations and manufacturing processes.****Subsequent investigations pointed out that the presence of nitrosamines could have occurred due to a reaction between the nitrocellulose of the primer layer that covers some types of materials used in the primary packaging of these medicines and amines present on inks used to print out information on it.****This questionnaire must be answered by the qualified person of the packaging materials manufacture and aims to identify possible sources of contamination by nitrosamines in these materials.****Answer the following questions according to the manufacturing process for the packaging material identified above:** |

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| 1) Is Sodium Nitrite (NaNO2) or any other nitrite or nitrosating agent: |  |  |  |
| - Used in any of the material manufacturing process? | [ ] **YES** | [ ] **NO** |  |
| - Known to be used in the preparation of raw materials or intermediates used in the manufacturing process? | [ ] **YES** | [ ] **NO** | [ ] **Unavailable information** |
| - Known to be used in the preparation of reagents or processing aids used in the manufacturing process? | [ ] **YES** | [ ] **NO** | [ ] **Unavailable information** |
| - Known to be generated as impurities during the manufacturing process? | [ ] **YES** | [ ] **NO** | [ ] **Unavailable information** |

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| 2) Is Nitrocellulose or any other substance containing nitrogen used in the composition of the material? | [ ] **YES** | [ ] **NO** |  |
| If **YES**, is it possible to replace these substances keeping the material properties and reducing the possibilities of generating nitrosamines? | [ ] **YES** | [ ] **NO** |  |
| If **YES**, please inform the substance that could be alternatively used:**Click here to enter text.** | ☐ **Not applicable** |

**Issued by:**

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| --- | --- |
| Name | **Click here to enter text.** |
| Job Title | **Click here to enter text.** |
| Signature |  |