# Annex 4

## Collaborative procedure between the World Health Organization Prequalification of Medicines Programme and national medicines regulatory authorities in the assessment and accelerated national registration of WHO-prequalified pharmaceutical products

# 1. Definitions

## Collaborative procedure (Procedure)

Procedure for collaboration between the WHO Prequalification of Medicines Programme (WHO/PQP) and interested national medicines regulatory authorities (NMRAs) in the assessment and accelerated national registration of WHO-prequalified pharmaceutical products.

## Participating authorities or participating NMRAs

NMRAs that voluntarily agree to implement this collaborative procedure and accept the task of processing applications for registration of WHO-prequalified pharmaceutical products in accordance with the terms of the Procedure. A list of participating authorities is posted on the WHO/PQP web site (http://www.who.int/prequal/).

# 2. Background information

National assessment of applications for registration of pharmaceutical products (marketing authorization) is the key regulatory process that enables NMRAs to evaluate and monitor the quality, safety and efficacy of pharmaceutical products. For most countries the approach to registration of pharmaceutical products is a combination of two components:

- the NMRA's own assessment of application documentation combined with verification of compliance with relevant good practices by inspections (mostly focusing on good manufacturing practices (GMP) and inspections of manufacturing sites);
- consideration by the NMRA of decisions and outcomes of assessments and inspections made by NMRAs in other countries.

Consideration of the outcomes of assessments and inspections by trusted authorities substantially contributes to savings in regulatory resources and improvements in the quality of regulatory decisions, while retaining the prerogative of NMRAs to conclude their assessment by sovereign decisions, which reflect their own judgement of the benefit–risk balance as it relates to their specific country situation and the legislation in place.

Taking into consideration the regulatory decisions of other NMRAs requires setting up a system that will permit:

- identification of reference authorities whose regulatory decisions are based on acceptable standards and identification of documents

associated with such regulatory decisions, which are relevant to the regulatory environment in the country wishing to rely on such decisions;

- assurance that the product for which the decision has been taken by the reference NMRA is identical to the product being assessed, or, if it is not identical, that a clear understanding exists of the differences between the products subjected to assessment in the two regulatory environments;
- efficient use of available scientific expertise and human and financial resources to decide, with reasonable certainty, on the benefit–risk profile of an evaluated pharmaceutical product when used in a given country;
- the choice by each NMRA of the approaches that will make best use of the resources, workload and competence of individual NMRAs. Approaches could range from completely independent data reviews and inspections to adoption of regulatory decisions of trusted authorities without any further scientific review. A pragmatic approach is to assess only those areas which relate to use of the product in the country concerned and where failure to comply with regulatory standards could pose health risks. In the other areas, the outcomes of trusted authorities may be adopted.

This Procedure is based on the above-mentioned considerations. In line with the *Procedure for prequalification of pharmaceutical products*,[1] it aims at providing a convenient tool for NMRAs wishing to enhance their pre-marketing evaluation and registration system by taking advantage of the scientific assessment work conducted by WHO/PQP. The present procedure is complementary to the WHO/PQP collaboration procedure with NMRAs in inspection activities (http://www.who.int/prequal, "Inspections").

It is expected that enhanced collaboration and information exchange between NMRAs and WHO/PQP will benefit both partners. Subject to the agreement of the WHO prequalification holders concerned, NMRAs will gain access to assessment outcomes that are not in the public domain and that have been prepared in conformity with the WHO recommended standards on which the *Procedure for prequalification of pharmaceutical products* is based. Such reports will help NMRAs to make their decisions and may also assist in educating national regulatory staff. At the same time, feedback from NMRAs on the information and documentation received from WHO/PQP under the Procedure

[1] Procedure for prequalification of pharmaceutical products. In: *WHO Expert Committee on Specifications for Pharmaceutical Preparations. Forty-fifth report*. Geneva, World Health Organization, 2011 (WHO Technical Report Series, No. 961), Annex 10.

will allow WHO/PQP to improve its work and to ensure that the outcomes of prequalification assessments are relevant to NMRAs. As a consequence patients will benefit from this collaboration by gaining faster access to medicines which have been found acceptable in principle for procurement by United Nations agencies. Depending on available resources, participating authorities may be given the opportunity to participate in the assessment process and in inspections organized by WHO/PQP.

This collaborative procedure also benefits manufacturers of prequalified medicines through faster and better harmonized regulatory approvals in participating countries. This Procedure, when combined with the *Collaboration Procedure with NMRAs in inspection activities*, may also alleviate the burden of national inspections on manufacturers.

## 3. Principles of collaboration

3.1 This collaborative procedure is limited to those pharmaceutical products that have been assessed and inspected by WHO/PQP in line with the procedures and standards available at www.who.int/prequal ("Information for applicants") and have been found to be acceptable in principle for procurement by United Nations agencies as listed in the *List of WHO prequalified medicines* available at www.who.int/prequal. It is not, however, applicable to medicines which have been listed as prequalified on the basis of approval by stringent regulatory authorities.[2] Although it is expected that the Procedure will mostly serve to accelerate the assessment and registration of multisource (generic) pharmaceutical products,[3] it is also applicable to any pharmaceutical product for which the safety and efficacy has been documented to WHO/PQP by the submission of preclinical and clinical data. The Procedure has three major stakeholders: WHO/PQP, interested NMRAs and those WHO prequalification holders/applicant[4] who agree that this procedure is used for applications for national registration of their WHO-prequalified product submitted to an NMRA.

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[2] Products listed as prequalified according to the procedures described in the *Guidelines on submission of documentation for prequalification of innovator finished pharmaceutical products approved by stringent regulatory authorities*.

[3] *Guideline on submission of documentation for prequalification of multisource (generic) finished pharmaceutical products (FPPs) approved by stringent regulatory authorities* (http://www.who.int/prequal/); and in: *WHO Expert Committee on Specifications for Pharmaceutical Preparations. Forty-fifth report*. Geneva, World Health Organization, 2011 (WHO Technical Report Series, No. 961), Annex 11.

[4] If the applicant for national registration is not the same as the WHO prequalification holder, the WHO prequalification holder must confirm to the NMRA and WHO/PQP by an authorization letter (as per the template annexed to Appendix 3, Part A) that the applicant is acting for, or pursuant to rights derived from, the WHO prequalification holder and that the prequalification holder agrees with the application of the procedure in the country concerned.

3.2 WHO/PQP and participating authorities receive applications for the same pharmaceutical product. Within the context of this Procedure, the same pharmaceutical product is characterized by:

- the same product dossier;[5]
- the same manufacturing chain, processes and control of materials;
- the same active pharmaceutical ingredient (API) and finished pharmaceutical product (FPP) specifications;
- the same essential elements of product information.[6]

3.3 WHO/PQP, with the agreement of the WHO prequalification holder, shares the full outcome, of prequalification assessments and inspections, including final assessment and inspection reports, with participating authorities, under appropriate obligations of confidentiality and restrictions on use (see below). As regards sharing the outcomes of assessments and inspections, only data owned by the WHO prequalification holder are shared. Sharing of any other data is subject to additional agreement of the data owners concerned.

3.4 For the purpose of this collaborative procedure, participating authorities accept the product documentation and reports, in the format in which they are routinely prepared by WHO in accordance with the *Procedure for prequalification of pharmaceutical products* published on WHO/PQP's web site at www.who.int/prequal, and as Annex 10 in WHO Technical Report Series, No. 961. It should be noted, however, that participating authorities may require applicants to comply with specific requirements for local regulatory review. Each participating authority should make such specific requirements public.

3.5 Fees to be paid by the applicants to participating authorities will continue to follow standard national procedures. Similarly, the submission by manufacturers of samples for laboratory testing – if required – will continue to follow standard procedures as defined in national legislation and/or as defined by national regulatory authorities.

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[5] Only the technical data included in the dossier must be the same. There may be country-specific differences in administrative data, or if required by NMRAs under exceptional circumstances, additional technical data can be provided (e.g. bioequivalence with a country-specific comparator).

[6] The essential elements of product information include in particular the indications, contraindications, posology (dosing), special warnings and precautions for use, adverse reactions, storage conditions, primary packaging and shelf-life. Differences in brand name, the name of applicant or prequalification holder, language, format and degree of detail of the product information, labelling of internal and external packaging, among others, are not considered essential for the purposes of this procedure. The language of the product information may be different as long as the information content is the same as that approved by WHO/PQP.

3.6 Consistent with the terms of Appendix 1A and Appendix 3, Part B, each participating authority commits itself:

- to treat any information and documentation provided to it by WHO/PQP pursuant to this Procedure as confidential in accordance with the terms of Appendix 1A, and to allow access to such information and documentation only to persons:[7]
  - who have a need to know for the purpose of the assessment and accelerated registration of the product in question in the country and any post-registration processes that may be required;
  - who are bound by confidentiality undertakings in respect of such information and documentation which are no less stringent than those reproduced in Appendix 1A;
  - to issue its national regulatory decision on a given prequalified pharmaceutical product (whether positive or negative) within 90 calendar days after being given access to the confidential information and documentation concerning each product.[8]

These commitments are provided by each participating authority to WHO/PQP in writing by entering into the agreement for participation in this Procedure as reproduced in Appendix 1A and are reconfirmed for each pharmaceutical product for which collaboration is sought (see Appendix 3, Part B).

Each participating NMRA nominates a maximum of two focal points who will access the restricted-access web site, through which WHO/PQP will communicate all confidential information and documentation. Focal points designated by the NMRA must sign the undertaking reproduced in Appendix 1B before they will be granted access to the restricted-access web site. Any change in designated focal points must be communicated to WHO/PQP in writing without delay and must be accompanied by an undertaking (Appendix 1B) signed by the new focal point(s).

[7] This includes the focal point(s) and all other persons in the NMRA who have access to any information and documentation provided by WHO/PQP.

[8] Participating authorities should issue their national regulatory decisions at the earliest opportunity after being given access to the confidential information and documentation on a given prequalified product. Although a time limit of 90 days is defined in the Procedure, the decision should normally be taken within 60 days. This deadline can be extended to a maximum of 90 days if predefined dates of technical or decision-making meetings do not allow a participating authority to issue its decision within 60 days. If a participating authority does not issue its decision within 90 days and does not communicate valid reasons for the delay to WHO/PQP, WHO/PQP will follow up with the head of the NMRA to clarify the situation.

3.7 The decision whether or not to register a given product in a particular country remains the prerogative and responsibility of each participating authority. Accordingly, a participating authority may come to a different conclusion from that reached by WHO/PQP. Within 30 calendar days of having taken its decision, the participating authority reports this decision, together with the dates of submission and registration and, if applicable, any deviations from the WHO/PQP's decision on prequalification and the reasons for such deviations,[9] to WHO/PQP. It does so through the restricted-access web site by completing the form in Part C of Appendix 3. The NMRA provides a copy of the completed form to the applicant.

3.8 Participation by WHO prequalification holders/applicants is voluntary, through the submission to a participating NMRA of the expression of interest reproduced in Part A of Appendix 3. For each product, such participation will be subject to the WHO prequalification holder/applicant accepting the terms of this Procedure, including the confidential exchange of information and documentation between WHO/PQP and the NMRA (see Appendix 2). The WHO prequalification holder/applicant can cease participation in this procedure at any time provided that he or she informs WHO/PQP and the participating NMRAs in writing of his or her decision. In such a case the NMRA shall cease all use of the information disclosed to it for the respective product(s) as per the terms of the participation agreement (see Appendix 1).

3.9 The requirements and procedures in case of a variation (as defined in the *WHO guidelines on variations to a prequalified product*[10]) may differ between NMRAs and WHO/PQP. The present collaborative procedure includes a variation procedure (see below under "Post-registration processes") which is aimed at promoting consistency between variations accepted by WHO/PQP and variations accepted by participating authorities. There could be situations in which a manufacturer of a WHO-prequalified pharmaceutical product submits a variation application to a participating authority and not to WHO/PQP, or vice versa. In such a case, the conditions of the national registration, which were initially "harmonized" with the WHO prequalification decision,

[9] This refers to a decision not to approve the marketing authorization of a WHO-prequalified product and to a decision to approve the marketing authorization, but with deviations in indications, contraindications, posology (dosing), special warnings and precautions for use, adverse drug reactions, storage conditions and shelf-life. Differences in brand name, name of applicant or prequalification holder, format of product information, level of detail of product information, labelling of internal and external packaging and language of product information are not considered to be deviations from the prequalification conclusions.

[10] WHO guidelines on variations to a prequalified product. In: *WHO Expert Committee on Specifications for Pharmaceutical Preparations. Forty-seventh report*. Geneva, World Health Organization, 2013 (WHO Technical Report Series, No. 981), Annex 3 (and any updates thereto).

may become essentially different through the product life-cycle. In such a case a pharmaceutical product registered and procured in a participating country would no longer be the same as the "WHO-prequalified" product because the specifications and/or other essential parameters would no longer be the ones accepted by WHO/PQP. As a result, applicants are required to submit the variations which are submitted to WHO/PQP without delay to participating authorities, and participating authorities are encouraged to follow the outcomes of the WHO variation procedures for nationally-approved WHO-prequalified products. WHO/PQP will inform the NMRA which registered individual prequalified products, through the restricted-access web site, about variations to the prequalification status of such products, if and when regulatory action is deemed to be justified. If a national variation procedure results in the nationally registered product being no longer the same[11] as the WHO-prequalified product, or in the event that a variation of a WHO-prequalified product is not followed by the same variation of the nationally registered product, the participating authority informs WHO/PQP of the situation by submitting the form in Appendix 4, clearly specifying the deviations. Other participating NMRAs, which have registered the WHO-prequalified product in question pursuant to this Procedure, will be made aware of such deviations through the restricted-access web site. In addition, if the fact that a WHO-prequalified product has been registered in a particular country pursuant to this Procedure has been made public, any subsequent deviations should also be made public.

3.10 If a prequalified product is withdrawn by the WHO prequalification holder, or is suspended or delisted by WHO/PQP, WHO/PQP will inform each participating authority which has approved, or is in the process of reviewing, the product pursuant to this collaborative procedure, of the withdrawal, suspension or delisting and the reasons for taking this action, through the restricted-access web site and subject to the obligations of confidentiality contained in Appendix 1A. Similarly, when an NMRA deregisters or suspends the registration of a prequalified pharmaceutical product for any reason, it will inform WHO/PQP of this decision and of its reasons through the restricted-access web site. Other participating NMRAs which have registered the WHO-prequalified product in question pursuant to this Procedure will be made aware of such national deregistration or suspension through the restricted-access web site. In addition, if the fact

[11] Within the context of this Procedure, the same pharmaceutical product is characterized by the same product dossier, the same manufacturing chain, processes and control of materials, the same API and FPP specifications and the same essential elements of product information, as further described in paragraph 3.2 above.

that a WHO-prequalified product has been registered in a country pursuant to this Procedure has been made public, any subsequent deregistration or suspension should also be made public.

3.11 Participation in this Procedure does not exempt applicants for national registration and holders of national registration from the respective national regulatory requirements. Participating authorities retain the right to assess submitted data and organize site inspections to the extent they deem appropriate.

## 4. Steps in the collaboration for national registration of a pharmaceutical product[12]

4.1 The applicant submits the product dossier for a WHO-prequalified pharmaceutical product to a participating NMRA. The technical part of the dossier is updated to reflect the data submitted to WHO/PQP during the initial prequalification procedure, and consecutive variation procedures and requalification (where applicable). The applicant must provide the participating authority with:

- an application dossier complying with established national requirements, including the same technical information as that submitted to WHO/PQP. To the extent that national regulatory requirements allow, the technical part of the dossier will be identical to the current version of the WHO/PQP dossier;
- an expression of interest reproduced in Part A of Appendix 3;
- country-specific data;
- any fees that may be payable to the NMRA pursuant to national requirements.

Wherever possible, to minimize the workload of the NMRA and facilitate the process, applicants should ensure that they express their interest to use the Procedure (Appendix 3, Part A) to the NMRA and to WHO/PQP before submitting a national application for registration. If acceptable to NMRAs, not only should the technical content of the dossiers be the same, but also the format in which data are presented should closely follow the common technical document (CTD) format in which dossiers are submitted to WHO/PQP.

In situations where the applicant wishes to apply the Procedure to an application which is already pending within the NMRA, the applicant should

[12] In addition, to complement the steps of this collaborative procedure, joint inspections may be arranged under the collaborative procedure for joint inspections posted on the WHO/PQP web site (www.who.int/prequal, "Inspections").

first update the dossier to ensure that the technical part of the information is the same as that submitted to WHO/PQP. It is the decision of individual NMRAs whether to apply the Procedure in such cases.

4.2 For each application under this Procedure, WHO/PQP is informed by the WHO prequalification holder/applicant about the submission to the participating NMRA by providing a copy of completed Appendix 3, Part A. The WHO prequalification holder provides WHO at this time with its written consent for WHO/PQP to provide the product-related information in compliance with the applicable confidentiality requirements to the NMRA of the country concerned (see Appendix 2).

4.3 The participating NMRA informs WHO/PQP and the respective applicant of each application which it accepts or declines to include in this Procedure, and requests WHO/PQP to provide it with the necessary information and documentation (Appendix 3, Part B). The Procedure applies only to applications that the NMRA has accepted as complete.

4.4 Within 30 calendar days of receipt of the above-mentioned request WHO/PQP shares the most recent product-related information and assessment and inspection outcomes through the restricted-access web site with the participating authority. This information is subject to the obligations of confidentiality and restrictions on use and may include assessment report(s), variation assessment report(s) if applicable, full inspection report(s) of the most recent inspection(s) and the letter of prequalification or requalification. At the request of the participating authority, WHO/PQP provides explanations and/or more detailed information.

4.5 After receiving the information and documentation from WHO/PQP, the participating authority undertakes an accelerated assessment of the product in question. For each application, the participating authority is required to issue the relevant national decision within 90 calendar days from the day it received access to the complete prequalification documentation. Within 30 days of having taken its decision, the participating authority reports this decision, together with an indication of the dates of submission and registration, and, if applicable, any deviations from the WHO prequalification conclusion and the reasons for such deviations, to WHO/PQP through the restricted-access web site. This report is provided to WHO/PQP using Part C of Appendix 3 and is copied to the applicant. WHO/PQP lists pharmaceutical products registered according to this Procedure by participating NMRAs on its public web site. The steps in the collaboration for national registrations of a pharmaceutical product are summarized in Figure 1.

Figure 1
**Flowchart showing the principal steps of the collaborative procedure**

The national medicines regulatory authority (NMRA) confirms to the WHO Prequalification of Medicines Programme (WHO/PQP) its interest in participating in the collaborative procedure and nominates focal point(s) for access to the restricted-access web site. The NMRA completes, signs, and submits to WHO/PQP, the agreement reproduced in Appendix 1A. The focal person(s) who are nominated to access the restricted-access web site complete and submit the undertaking reproduced in Appendix 1B, to WHO/PQP.

Appendix 1, Part A and Appendix 1, Part B

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WHO/PQP lists the participating NMRAs on its public web site.

**Registration process**

The applicant submits the application for national registration of the WHO-prequalified pharmaceutical product to the participating authority, and informs the authority of its interest in following the collaborative procedure by completing the expression of interest reproduced in Appendix 3, Part A. If the applicant for national registration is not the same as the WHO prequalification holder, the WHO prequalification holder confirms to the NMRA and WHO/PQP by an authorization letter (as per the form annexed to Appendix 3, Part A) that the applicant is acting for, or pursuant to rights derived from, the WHO prequalification holder and that the prequalification holder agrees with the application of the procedure in the country concerned.

Appendix 3

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The WHO prequalification holder/applicant informs WHO/PQP about the submission of its application to the NMRA(s) (by providing a copy of completed Appendix 3, Part A) and, for each product and country, provides WHO/PQP with its written consent to share the product-related information and documentation, under confidential cover, with the participating authority. The WHO prequalification holder completes and signs the consent form reproduced in Appendix 2 and submits it to WHO.

Appendix 2

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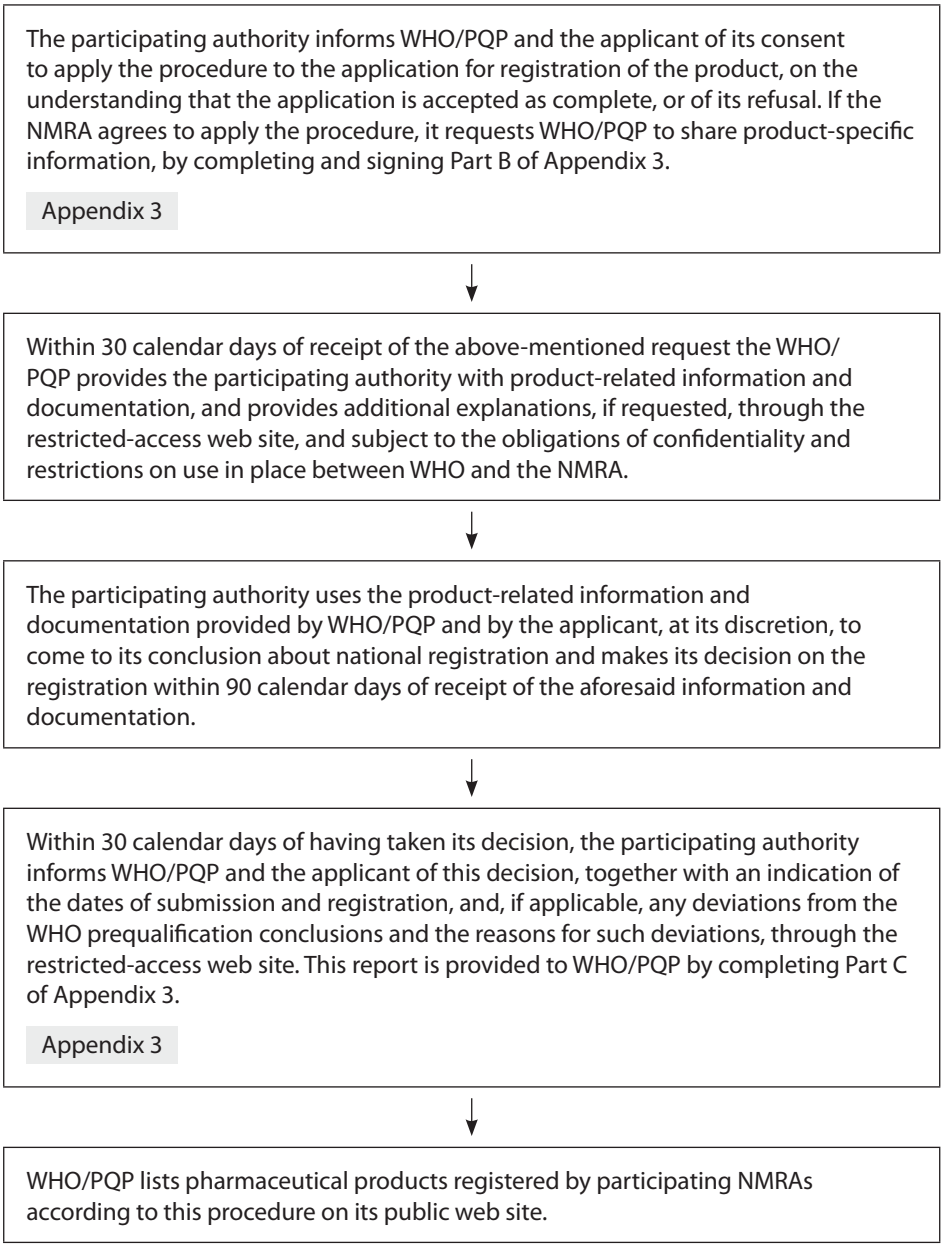
*continues*

Figure 1 *continued*




*continues*

Figure 1 *continued*

**Post-registration processes**

The WHO prequalification holder/applicant submits variations simultaneously to WHO/PQP and relevant participating authorities. If regulatory action is deemed to be justified, WHO/PQP promptly provides the participating authorities concerned, through the restricted-access web site, and subject to the above-mentioned obligations of confidentiality and restrictions on use, with variation assessment reports and post-prequalification inspection reports, and any related information it considers relevant. If a national variation procedure results in the nationally-registered product being no longer the same[13] as the WHO-prequalified product, or in the event that a variation of a WHO-prequalified product is not followed by the same variation of the nationally-registered product, the participating authority informs WHO of the situation within 30 calendar days of obtaining access to the information and documentation provided by WHO/PQP, by submitting the form reproduced in Appendix 4, clearly specifying the deviations. Other participating NMRAs which have registered the WHO-prequalified product in question pursuant to this procedure will be made aware of such deviations through the restricted-access web site.

Appendix 4

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WHO/PQP informs the participating authority, through the restricted-access web site, and subject to the above-mentioned obligations of confidentiality and restrictions on use, about withdrawals, suspensions or delistings of prequalified pharmaceutical products. The participating authority informs WHO/PQP, through the restricted-access web site, of national deregistration or suspension (for any reason) of a prequalified pharmaceutical product and the reasons for doing so.

Other participating NMRAs which have registered the WHO-prequalified product in question pursuant to this procedure will be made aware of such national deregistration or suspension, through the restricted-access web site.

Appendix 4

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WHO/PQP removes a product from the list published in line with this procedure:

- if the nationally-registered product is no longer the same[14] as the WHO-prequalified product, or
- if the NMRA deregisters a WHO-prequalified product, or
- if WHO/PQP delists a WHO-prequalified product.

WHO/PQP will also publish the reasons for the removal from the list.

[13] See footnote 11.

[14] See footnote 11.

## 5. Collaboration mechanisms for post-registration variations

5.1 Post-prequalification variations submitted to WHO/PQP are expected to be submitted simultaneously to any relevant participating authorities, and vice versa. Submission of variations to NMRAs should respect national regulatory requirements.

5.2 WHO/PQP promptly shares the variation assessment reports and post-prequalification inspection reports, through the restricted-access web site, and subject to the above-mentioned obligations of confidentiality and restrictions on use, with the relevant participating authorities, in all cases in which variation (including "notification" according to WHO/PQP's variation procedure[15]) requires regulatory action (e.g. where product safety, efficacy or patient information materials are concerned). Within 30 days of obtaining access to the information and documentation from WHO/PQP, each participating authority informs WHO/PQP through the restricted-access web site if and to what extent a variation of a WHO-prequalified product is not followed by the same variation of the nationally-registered product and, as a consequence, the nationally-registered product is no longer the same[16] as the WHO-prequalified product.

5.3 If a national variation procedure results in the nationally-registered product being no longer the same[17] as the WHO-prequalified product, the participating authority informs WHO/PQP within 30 days about the subject and outcome of this national variation procedure.

5.4 Deviations under 5.2 and 5.3 above may include change of source of active ingredients and/or manufacturing sites, product specifications, testing methods, storage conditions, shelf-life, packaging material, indications, contraindications, posology (dosing), special warnings and precautions for use, and adverse reactions. Differences in brand name, name of applicant or WHO prequalification holder, format of product information, level of detail of product information, labelling of internal and external packaging and language of product information are not considered to be deviations from the prequalification conclusions.

[15] Guidance on variations to prequalified dossiers is available at: http://www.who.int/prequal/info_applicants/info_for_applicants_guidelines.htm

[16] See footnote 11.

[17] See footnote 11.

5.5 WHO/PQP removes a product from the list published in line with this procedure if the nationally-registered product is no longer the same[18] as the WHO-prequalified product.

## 6. Withdrawals, suspensions or delistings of prequalified pharmaceutical products and national deregistrations

6.1 If a WHO-prequalified product is withdrawn from prequalification by the WHO prequalification holder, or if a product is suspended or delisted by WHO/PQP, WHO/PQP will promptly, through the restricted-access web site, and subject to the above-mentioned obligations of confidentiality and restrictions on use, inform relevant participating authorities accordingly, providing the reasons whenever needed.

6.2 In the case that a participating NMRA deregisters or suspends the registration of a prequalified pharmaceutical product for any reason, the participating authority informs WHO/PQP of the decision (together with an indication of the reasons), through the restricted-access web site. The information should be provided promptly whenever product quality, safety or efficacy are concerned and in all other cases within 30 working days. A participating authority is encouraged to consult WHO/PQP before adopting a decision about deregistration or suspension of registration of a WHO-prequalified product.

6.3 In case a WHO-prequalified product is deregistered at the national level, or in case WHO/PQP delists a prequalified product, WHO/PQP adjusts the information about this product on its web site accordingly.

[18] See footnote 11.

# Appendix 1

## NMRA participation agreement and undertaking for NMRA focal point(s)

### Appendix 1, Part A

*Agreement to participate in the collaborative procedure between the World Health Organization Prequalification of Medicines Programme and national medicines regulatory authorities (NMRAs) in the assessment and accelerated national registration of WHO-prequalified pharmaceutical products*

#### Details of national medicines regulatory authority (NMRA)

Name of NMRA _____ ("the NMRA")
Postal address: _____

_____

Country: _____ ("the Country")
Telephone number (please include codes): _____
E-mail: _____

#### Scope of agreement

Applicants for national registration of a WHO-prequalified pharmaceutical product (hereafter referred to as "Applicants") may express their interest to the NMRA for the assessment and accelerated registration of this product ("the Product") in the Country under the "collaborative procedure between the World Health Organization Prequalification of Medicines Programme (WHO/PQP) and national medicines regulatory authorities in the assessment and accelerated national registration of WHO-prequalified pharmaceutical products" (hereafter referred to as "the Procedure").[1]

Subject to the NMRA agreeing to conduct such assessment and consider such accelerated registration of the Product under the Procedure (by submitting the form reproduced in Part B of Appendix 3 attached to the Procedure to WHO/PQP through the restricted-access web site), the NMRA hereby confirms for each

[1] If the applicant for national registration is not the same as the WHO prequalification holder, the WHO prequalification holder must confirm to the NMRA and to WHO/PQP by an authorization letter (as per the template annexed to Appendix 3, Part A) that the applicant is acting for, or pursuant to rights derived from, the WHO prequalification holder, and that the WHO prequalification holder agrees with the application of the Procedure in the country concerned.

such Product that it will adhere to, and collaborate with the WHO/PQP and the applicant of the Product in accordance with, the terms of the Procedure.

### Confidentiality of information

Any information and documentation relating to the Product and provided by WHO/PQP to the NMRA under the Procedure may include but shall not necessarily be limited to:

- the full WHO/PQP assessment and inspection outcomes (reports);
- information and documentation on variations (as defined in the *WHO guidelines on variations to a prequalified product*, WHO Technical Report Series, No. 981, and any updates thereto), as well as information and documentation on any actions taken by WHO/PQP or NMRAs post-prequalification of the Product;
- all such data, reports, information and documentation being hereinafter referred to as "the Information".

As regards sharing the outcomes of assessments and inspections, only data owned by the WHO prequalification holder are shared. Sharing of any other data is subject to additional agreement of the data owners concerned.

WHO/PQP agrees to make such Information available to the NMRA through a restricted-access web site exclusively for the purpose of the assessment and accelerated registration of the Product in the Country and any post-registration processes that may be required, in accordance with and subject to the terms of the Procedure ("the Purpose"). The NMRA agrees to treat any Information provided by WHO/PQP as aforesaid as strictly confidential and proprietary to WHO/PQP, the WHO prequalification holder/applicant and/or parties collaborating with WHO/PQP and/or the WHO prequalification holder/applicant. In this regard, the NMRA agrees to use such Information only for the Purpose and to make no other use thereof. Thus, the NMRA undertakes to maintain the Information received from WHO/PQP in strict confidence, and to take all reasonable measures to ensure that:

- the Information received from WHO/PQP shall not be used for any purpose other than the Purpose;
- the Information shall only be disclosed to persons who have a need to know for the aforesaid Purpose and are bound by confidentiality undertakings in respect of such information and documentation which are no less stringent than those contained herein.

The NMRA warrants and represents that it has adequate procedures in place to ensure compliance with its aforesaid obligations.

The obligations of confidentiality and restrictions on use contained herein shall not cease on completion of the Purpose.

The obligations of confidentiality and restrictions on use contained herein shall not apply to any part of the Information which the NMRA is clearly able to demonstrate:

- was in the public domain or the subject of public knowledge at the time of disclosure by WHO/PQP to the NMRA under the Procedure; or
- becomes part of the public domain or the subject of public knowledge through no fault of the NMRA; or
- is required to be disclosed by law, provided that the NMRA shall in such event immediately notify WHO/PQP and the applicant in writing of such obligation and shall provide adequate opportunity to WHO/PQP and/or the applicant to object to such disclosure or request confidential treatment thereof (provided always, however, that nothing contained herein shall be construed as a waiver of the privileges and immunities enjoyed by WHO/PQP and/or as submitting WHO/PQP to any national court jurisdiction).

Upon completion of the Purpose, the NMRA shall cease all use and make no further use of the Information disclosed to it under the Procedure, and shall promptly destroy all of the Information received from WHO/PQP which is in tangible or other form, except that the NMRA may retain copies of the Information in accordance with its established archival procedures, subject always, however, to the above-mentioned obligations of confidentiality and restrictions on use.

The Purpose for each product shall be deemed completed as soon as:

- the WHO prequalification holder/Applicant discontinues participation in the Procedure for the particular product;
- the Product is deregistered by the NMRA and/or delisted by WHO/PQP.

The access right of the NMRA's focal person(s) to the restricted-access web site will cease automatically upon the NMRA ceasing to participate in the Procedure. If and as soon as an NMRA focal point is replaced by a new focal point or ceases to be an employee of the NMRA, such focal point's access to the restricted-access web site shall automatically terminate.

The NMRA agrees that it has no right in or to the Information and that nothing contained herein shall be construed, by implication or otherwise, as the grant of a licence to the NMRA to use the Information other than for the Purpose.

## Timelines

In respect of each Product which the NMRA accepts to assess and consider for accelerated registration under the Procedure, the NMRA undertakes to abide by the terms of the Procedure, including but not limited to the following timelines for processing each application:

- within 90 calendar days of obtaining access (through the restricted-access web site) to:
  - the data submitted to WHO/PQP for prequalification of the Product and owned by the WHO prequalification holder,
  - the full WHO/PQP assessment and inspection outcomes (reports), the NMRA undertakes to take a decision on the national registration of the Product;
- within 30 working days of the NMRA's decision on national registration of the Product, the NMRA undertakes to inform WHO/PQP of this decision and of any deviations from the WHO prequalification conclusions (with an indication of the reasons for such deviations) by completing and submitting the form attached as Appendix 3, Part C to the Procedure to WHO/PQP through the restricted-access web site;
- if a national variation procedure results in the nationally registered product being no longer the same[2] as the WHO-prequalified product, or if and to the extent a variation of a WHO-prequalified product is not followed by a variation of the nationally-registered product and as a consequence, the nationally-registered product is no longer the same[2] as the WHO-prequalified product, the NMRA undertakes to inform WHO/PQP thereof (together with an indication of the reasons for such deviations) within 30 days of the conclusion of the national variation procedure or within 30 days of having received access to the information and documentation provided by WHO/PQP, as the case may be (i.e. by completing and submitting the form attached to the Procedure as Appendix 4 to WHO/PQP through the restricted-access web site);[3]

[2] Within the context of this Procedure, the same pharmaceutical product is characterized by the same product dossier, the same manufacturing chain, processes and control of materials, the same API and FPP specifications and the same essential elements of product information, as further described in paragraph 3.2 of the Procedure.

[3] If the fact that a WHO-prequalified product has been registered in a country pursuant to this Procedure has been made public any subsequent deviations should be made public also.

- the NMRA undertakes to inform WHO/PQP in case the NMRA deregisters or suspends the registration of the Product in the Country, by completing and submitting the form attached to the Procedure as an Appendix 4 to WHO/PQP through the restricted-access web site, and to do so promptly if this decision is based on quality, safety of efficacy concerns, and within 30 working days if this decision is based on other reasons.

### Focal points for access to restricted-access web site

The NMRA has designated the person(s) listed below to act as focal point(s) for access to WHO/PQP's restricted-access web site. The undertaking(s) completed and signed by the focal point(s) is(are) attached hereto as an Appendix to this agreement.

Any change in designated focal points must be communicated to WHO/PQP without delay in writing and will be subject to the new focal point having signed and submitted to WHO the undertaking reproduced in Appendix 1B to the Procedure. The NMRA also undertakes to inform WHO/PQP if and as soon as a designated focal point ceases to be an employee of the NMRA.

### Focal point for inspections

*If applicable, this should be the same focal point as for the "WHO/PQP collaborative procedure between WHO/PQP and selected NMRAs in inspection activities" (http://who.int/prequal).*

Mr/Ms/Dr:
First name (and initials): _____
Surname/family name: _____
Title in NMRA: _____
E-mail: _____
Phone: _____

☐ A signed undertaking is attached

### Focal point for dossier assessment

*The same person as above may be nominated. If a different person is nominated, please complete details below.*

Mr/Ms/Dr:
First name (and initials): _____
Surname/family name: _____

Title in NMRA: _____
E-mail: _____
Phone: _____

☐ A signed undertaking is attached

## Miscellaneous

The NMRA agrees that WHO/PQP may list its name on the WHO/PQP web site as a participant in the Procedure. Except as provided hereinbefore, neither party shall, without the prior written consent of the other party, refer to the relationship of the parties under this Agreement and/or to the relationship of the other party to the Product, the Information and/or the Purpose, in any statement or material of an advertising or promotional nature.

This Agreement shall not be modified except by mutual agreement of WHO and the NMRA in writing. The NMRA furthermore undertakes to promptly inform WHO/PQP of any circumstances or change in circumstances that may affect the implementation of this Agreement.

The parties shall use their best efforts to settle amicably any dispute relating to the interpretation or execution of this Agreement. In the event of failure of the latter, the dispute shall be settled by arbitration. The arbitration shall be conducted in accordance with the modalities to be agreed upon by the parties or in the absence of agreement, with the UNCITRAL Arbitration Rules in effect on the date of this Agreement. The parties shall accept the arbitral award as final.

It is agreed furthermore that nothing contained in this Agreement shall be construed as a waiver of any of the privileges and immunities enjoyed by WHO under national and international law, and/or as submitting WHO to any national court jurisdiction.

Agreed and accepted
For the NMRA

Signature: _____
Name: _____
Title: _____
Place and date: _____

Attachments:
1. Signed undertakings of NMRA focal point(s) (Appendix 1, Part B).

## Appendix 1, Part B

### Undertaking for NMRA focal point(s)

**The undersigned:**

Mr/Ms/Dr:
First name (and initials): _____
Surname/family name: _____
Title in NMRA: _____
Name of NMRA _____ ("the NMRA")
Country: _____ ("the Country")
E-mail: _____
Phone: _____

Applicants for national registration of WHO-prequalified pharmaceutical products (hereafter referred to as "Applicants") may express their interest to the NMRA for the assessment and accelerated national registration of such products under the "collaborative procedure between the World Health Organization Prequalification of Medicines Programme (WHO/PQP) and national medicines regulatory authorities (NMRAs) in the assessment and accelerated national registration of WHO-prequalified pharmaceutical products" (hereafter referred to as "the Procedure").[1]

Subject to the NMRA agreeing to conduct such assessment and consider such accelerated registration of a WHO-prequalified product under the Procedure, WHO/PQP will communicate confidential Information (as hereinafter defined) relating to each such product to the NMRA, and the NMRA will communicate outcomes of the national registration procedure and post-registration actions in respect of such products to WHO/PQP, through a restricted-access web site, which can be accessed only by the focal points designated by the NMRA, including the undersigned. For the purpose of accessing the restricted-access web site and downloading Information and uploading reports in accordance with and subject to the terms of the Procedure, WHO/PQP will provide the undersigned with a secret access code. The undersigned undertakes to treat this access code as strictly confidential and not to disclose it to any other person whatsoever. The undersigned furthermore undertakes to take all precautionary measures that may be needed to prevent any other person whatsoever from obtaining the aforesaid

[1] If the applicant for national registration is not the same as the WHO prequalification holder, the WHO prequalification holder must confirm to the NMRA and to WHO/PQP by an authorization letter (as per the template annexed to Appendix 3, Part A) that the applicant is acting for, or pursuant to rights derived from, the WHO prequalification holder, and that the prequalification holder agrees with the application of the Procedure in the country concerned.

secret access code and from accessing the restricted-access web site (i.e. except for the other designated focal points who have signed this undertaking).

"Information" as aforesaid means any information and documentation relating to a WHO-prequalified product to be provided by WHO/PQP to the NMRA under the Procedure, including but not necessarily limited to:

- the full WHO/PQP assessment and inspection outcomes (reports);
- information and documentation on subsequent variations (as defined in the *WHO guidelines on variations to a prequalified product*, WHO Technical Report Series, No. 981, and any updates thereto), as well as information and documentation on any actions taken by WHO/PQP or NMRAs post-prequalification of the Product.

As regards sharing the outcomes of assessments and inspections, only data owned by the WHO prequalification holder are shared. Sharing of any other data is subject to additional agreement of the data owners concerned.

The undersigned confirms that:

1. the NMRA has bound him or her to obligations of confidentiality and restrictions on use no less stringent than those contained in Appendix 1A to the Procedure; and that
2. the aforesaid obligations of confidentiality and restrictions on use shall not cease on completion of the assessment and accelerated registration of any product in the Country, nor on completion of any post-registration processes that may be required, nor on the undersigned ceasing to be an employee of (or ceasing to have another relationship with) the NMRA.

The undersigned shall automatically cease having the right to access the restricted-access web site when the NMRA designates a new focal point to replace the undersigned or when the undersigned ceases to be an employee of the NMRA.

This Undertaking shall not be modified except by mutual agreement of WHO and the undersigned in writing. The undersigned furthermore undertakes to promptly inform WHO/PQP of any circumstances or change in circumstances that may affect the implementation of this Undertaking.

The parties shall use their best efforts to settle amicably any dispute relating to the interpretation or execution of this Undertaking. In the event of failure of the latter, the dispute shall be settled by arbitration. The arbitration shall be conducted in accordance with the modalities to be agreed upon by the parties or in the absence of agreement, with the UNCITRAL Arbitration Rules in effect on the date of this Undertaking. The parties shall accept the arbitral award as final.

It is agreed furthermore that nothing contained in this Undertaking shall be construed as a waiver of any of the privileges and immunities enjoyed by WHO under national and international law, and/or as submitting WHO to any national court jurisdiction.

Agreed and accepted by the Undersigned:

Signature: _____
Name: _____
Title in NMRA: _____
Place and date: _____

# Appendix 2

## Consent of WHO prequalification holder for WHO to share information with NMRA confidentially under the Procedure

Reference is made to the attached expression of interest for the assessment and accelerated national registration under the Procedure of the following WHO-prequalified pharmaceutical product (hereafter referred to as "the Product") in _____ [**country**].[1]

### WHO prequalification details:

WHO prequalification reference number: _____
Date of prequalification (dd/mm/yyyy): _____
Date of requalification (if applicable): _____
WHO prequalification holder:[2] _____

### Application details:

Name of entity: _____ ("the Applicant")
Street: _____
City and country: _____
E-mail: _____
Phone: _____

The WHO prequalification holder hereby consents to WHO/PQP providing the following information and documentation to the NMRA of _____ [**country**] ("the NMRA") for the assessment and accelerated registration of the Product in the country under the Procedure and to freely discuss the same with the aforesaid NMRA for this purpose:

- the full WHO/PQP assessment and inspection outcomes (reports);

[1] Please complete a separate form of this Annex for each country.

[2] If the applicant for national registration is not the same as the WHO prequalification holder, the WHO prequalification holder must confirm to the NMRA and to WHO/PQP by an authorization letter (as per the template annexed to Appendix 3, Part A) that the applicant is acting for, or pursuant to rights derived from, the WHO prequalification holder, and that the prequalification holder agrees with the application of the Procedure in the country concerned.

- information and documentation on subsequent variations (as defined in the *WHO guidelines on variations to a prequalified product*, WHO Technical Report Series, No. 981, and any updates thereto), as well as information and documentation on any actions taken by WHO/PQP post-prequalification of the Product.
- all such data, reports, information and documentation being hereinafter referred to as "the Information".

As regards sharing the outcomes of assessments and inspections, only data owned by the WHO prequalification holder are shared. Sharing of any other data is subject to additional agreement of the data owners concerned.[3]

Such consent is subject to the NMRA having entered into an agreement with WHO/PQP as per Appendix 1A to the Procedure and having agreed to conduct the assessment and consider the accelerated registration of the Product under the Procedure, by having submitted the form reproduced in Part B of Appendix 3 to the Procedure to WHO.

If a national variation procedure results in the nationally-registered product being no longer the same[4] as the WHO-prequalified Product, or if a variation of the WHO-prequalified Product is not followed by a variation of the nationally-registered product and, as a consequence, the nationally-registered product is no longer the same, the WHO prequalification holder/Applicant will inform WHO/PQP of the differences and their reasons.

### For the WHO prequalification holder

Signature: _____

Name: _____

Title: _____

Place: _____

Date (dd/mm/yyyy): _____

[3] In case that certain data submitted to WHO/PQP by the WHO prequalification holder in relation to prequalification of the Product are not in his/her ownership, the WHO prequalification holder specifies such data in an annex to this declaration of consent.

[4] Within the context of this Procedure, the same pharmaceutical product is characterized by the same product dossier, the same manufacturing chain, processes and control of materials, the same API and FPP specifications and the same essential elements of product information, as further described in paragraph 3.2 of the Procedure.

# Appendix 3

## Expression of interest to NMRA for the assessment and accelerated national registration, acceptance by NMRA and notification of Procedure outcomes

### Appendix 3, Part A

*Expression of interest to the national medicines regulatory authority (NMRAs) for the assessment and accelerated national registration of a WHO-prequalified pharmaceutical product*

In line with the Procedure, the undersigned Applicant[1] expresses its interest in the application of the above-mentioned Procedure by the NMRA of _____ [country] ("the NMRA") in respect of the following submission for national registration:

**Application details:**

Name of entity: _____ ("the Applicant")
Street: _____
City and country: _____
E-mail): _____
Phone: _____
Date of application: _____
(dd/mm/yyyy, e.g. 31/07/2011): _____
Product name in national system (if known): _____
National reference number (if known): _____

**Product details:**

API(s) (INN): _____
_____
Dosage form and strength: _____
Packaging: _____
Manufacturing site(s), including block(s)/unit(s) if appropriate: _____

[1] If the applicant for national registration is not the same as the WHO prequalification holder, the WHO prequalification holder must confirm to the NMRA and to WHO/PQP by an authorization letter (as per the template annexed to Appendix 3, Part A) that the applicant is acting for, or pursuant to rights derived from, the WHO prequalification holder, and that the prequalification holder agrees with the application of the Procedure in the country concerned.

**WHO prequalification details:**

WHO prequalification reference number: _____
Date of prequalification (dd/mm/yyyy): _____
WHO prequalification holder: _____

The Applicant confirms that the information and documentation provided in support of the above-mentioned submission for national registration is true and correct, that the pharmaceutical product submitted for national registration is the same[2] as the WHO-prequalified product and that the technical part of the information is the same[3] as that submitted to the WHO Prequalification of Medicines Programme (WHO/PQP). Non-essential differences[4] from the information submitted to WHO/PQP, are the following: _____

_____

_____

Subject to the NMRA agreeing to conduct the assessment and consider the accelerated registration of the Product under the Procedure, the Applicant:

1. undertakes to adhere to, and collaborate with the NMRA and WHO/PQP in accordance with the terms of the Procedure; and
2. will authorize WHO/PQP[5] to provide the NMRA confidential access to the following information and documentation and to freely discuss the same with the aforesaid NMRA for the above-mentioned Purpose:

   – the full WHO/PQP assessment and inspection outcomes (reports);

[2] Within the context of this Procedure, the same pharmaceutical product is characterized by the same product dossier, the same manufacturing chain, processes and control of materials, the same API and FPP specifications and the same essential elements of product information, as further described in paragraph 3.2 of the Procedure.

[3] Only the technical data included in the dossier must be the same. There may be country-specific differences in administrative data, or if required by NMRAs under exceptional circumstances, additional technical data can be provided (e.g. bioequivalence with a country-specific comparator).

[4] As defined in Section 3.2 of the Procedure, differences in administrative information, brand name, name of applicant/prequalification holder (provided that the applicant is acting for, and has the authority to represent the WHO prequalification holder), format of product information, level of detail of product information, labelling of internal and external packaging and language of product information are not considered to be essential differences.

[5] If the applicant for national registration is not the same as the WHO prequalification holder, then the authorization to WHO/PQP must be provided by the WHO prequalification holder or their legal representative.

– information and documentation on subsequent variations (as defined in the *WHO guidelines on variations to a prequalified product*, WHO Technical Report Series, No. 981, and any updates thereto), as well as information and documentation on any actions taken by WHO/PQP post-prequalification of the Product.

As regards sharing the outcomes of assessments and inspections, only data owned by the WHO prequalification holder are shared. Sharing of any other data is subject to additional agreement of the data owners concerned.

3. authorizes the NMRA to freely share and discuss all registration and the Product related information provided by the Applicant to the NMRA, with WHO/PQP, subject to the obligations of confidentiality and restrictions on use as contained in the NMRA's participation agreement and focal points' undertakings.

☐ The application for national registration was submitted before the Applicant decided to apply the Procedure to the Product and therefore at the time of submission the registration dossier did not respect conditions of the Procedure. Steps taken to update the submission to the NMRA to make the dossier "the same" as required by the Procedure, are listed and referenced in the attached letter.

☐ The applicant is not the WHO prequalification holder. An authorization letter from the WHO prequalification holder is attached.

### For the Applicant

Signature: _____
Name: _____
Title: _____
Place: _____
Date (dd/mm/yyyy): _____

### Template for authorization letter

(To be provided if the applicant is not the WHO prequalification holder. Please provide a separate letter for each NMRA concerned, with a copy to WHO/PQP).

This is to confirm that _____ (*name of applicant*) _____ seeking registration for prequalified product number _____ (*WHO/PQ number*) _____ in _____ (*name country*) _____ under the WHO collaborative procedure for accelerated registration of WHO prequalified products, is acting for, or pursuant to rights derived from _____ (*name of WHO prequalification holder*)

_____ and that _____ (*name of WHO prequalification holder*) _____ agrees with the application of the procedure in the country concerned.

For _____ (*name of WHO Prequalification holder*) _____ :

Signature _____
Name _____
Title _____
Date _____

## Appendix 3, Part B

*Acceptance by the NMRA to apply the Procedure to a specified WHO-prequalified pharmaceutical product and request for access to product-specific information and documentation*

If there have been changes to the details as completed in Part A, please complete the relevant fields below. Where fields below are left blank, the data in Part A are considered to be valid.

### Application details:

Name of entity: _____ ("the Applicant")
Street: _____
City and country: _____
E-mail: _____
Phone: _____
Date of application: _____
(dd/mm/yyyy, e.g. 31/07/2011): _____
Product name in national system (if known): _____
National reference number (if known): _____

### Product details:

API(s) (INN): _____
Dosage form and strength: _____
Packaging: _____
Manufacturing site(s), including block(s)/unit(s) if appropriate: _____

### WHO prequalification details:

WHO prequalification reference number: _____

Date of prequalification (dd/mm/yyyy): _____
WHO prequalification holder _____

**Please complete either section A or section B below:**

☐ **Section A**

The NMRA agrees to conduct the assessment and the accelerated registration of the above-mentioned product ("the Product") under the Procedure and requests access to product-specific information, in accordance with and subject to the terms of the Procedure and the Agreement between WHO/PQP and the NMRA dated ____ /____ /____ (dd/mm/yyyy).

☐ **Section B**

The NMRA has decided not to apply the Procedure to the above-mentioned Product for the following reasons: _____

**For the NMRA**

Signature: _____
Name: _____
Title: _____
Place: _____
Date (dd/mm/yyyy): _____

## Appendix 3, Part C

*Notification of outcomes of national registration procedure by the NMRA*

Product and application details as completed in Parts A and B above apply.

**Please complete either Section A or B below:**

☐ **Section A**

Registration has been granted, and the above-mentioned product ("the Product") is identified as follows in the national medicines register:
**Name of the Product** _____
**National registration number** _____
**Date of registration** _____ (dd/mm/yyyy)
**Product details** (if different from those specified in Parts A and B): _____
API(s) (INN) _____
Dosage form and strength _____
Packaging _____

Manufacturing site(s), including block(s)/unit(s) if appropriate _____

Registration **holder** (if different from the Applicant as specified in Parts A and B: _____

Name of entity _____

Street _____

City and country _____

E-mail _____

Phone _____

**Are the national registration conclusions different from prequalification outcomes?**[6]

_____ **(Yes/No)**

### If you answered Yes to the above question:

| Deviation | Reason |
|---|---|
| | |
| | |
| | |

Please specify whether registration is subject to specific commitments, the registration is provisional or conditional, use of the Product is limited by specific prescribing restrictions, or additional clinical trials or additional data are required:

_____

☐ **Section B**

The application for registration of the Product was rejected for the following reasons: _____

### For the NMRA

Signature: _____

Name: _____

Title: _____

Place: _____

Date: (dd/mm/yyyy) _____

[6] This refers to deviations in indications, contraindications, posology (dosing), special warnings and precautions for use, adverse drug reactions, storage conditions and shelf-life. Differences in brand name, name of applicant/prequalification holder, format of a product information, level of detail of product information, labelling of internal and external packaging and language of product information are not considered to be a deviation from the prequalification conclusions.

# Appendix 4

## Report on post-registration actions in respect of a product registered under the Procedure

- Variation of the national registration resulting in the national registration conditions being inconsistent with the WHO/PQP prequalification conclusions
- Deregistration or suspension of the registration of the product

### Product details:

Product name in national system: _____ ("the Product")
National registration number: _____
Date of registration (dd/mm/yyyy): _____

### WHO prequalification details:

WHO prequalification reference number: _____
Date of prequalification (dd/mm/yyyy): _____
WHO prequalification holder: _____

☐ The national variation procedure has resulted in the nationally-registered Product being no longer the same[1] as the WHO-prequalified product.

| Deviations[2] | Reasons |
|---|---|
| | |
| | |
| | |
| | |

[1] Within the context of this Procedure, the same pharmaceutical product is characterized by the same product dossier, the same manufacturing chain, processes and control of materials, the same API and FPP specifications and the same essential elements of product information, as further described in paragraph 3.2 of the Procedure.

[2] This refers to deviations in indications, contraindications, posology (dosing), special warnings and precautions for use, adverse drug reactions, storage conditions and shelf-life. Differences in brand name, name of applicant/prequalification holder, format of product information, level of detail of product information, labelling of internal and external packaging and language of product information are not considered to be a deviation from the prequalification conclusions.

☐ The variation notified to NMRA by WHO/PQP has not been followed by a variation of the nationally-registered Product and, as a consequence, the nationally-registered product is no longer the same[3] as the WHO-prequalified product.

| Deviations[4] | Reasons |
|---|---|
| | |
| | |
| | |

☐ The Product has been deregistered or the registration of the Product has been suspended.

Deregistration **(Yes/No)**: _____
Suspension of registration: **(Yes/No)**: _____
Effective date (dd/mm/yyyy): ____ / ____ / ____
Reasons: _____

### For the NMRA

Signature: _____
Name: _____
Title: _____
Place: _____
Date: (dd/mm/yyyy) _____

[3] See footnote 1.
[4] See footnote 2.