Requirements
to the Registration of
Medicinal products
in the Republic of Armenia

Yerevan

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Requirements to the Registration of Medicinal products in the Republic of Armenia

Current requirements to registration are based on the below mentioned legislative acts: The Laws of the Republic of Armenia "On Medicinal products", "On State Taxes", and the Decree of the Government of the Republic of Armenia No 347 of 25 April, 2001 "On adopting the Rule of Registration of Medicinal products and Expertise Fees for Registration of Medicinal products in the Republic of Armenia", amended by Government Decrees No 148-N of 3 February, 2005 and No 1000-N of 3 September, 2009, the Order No 123-N of the Ministry of Health of the RA dated 7 February, 2006 on approval of ''The Procedure of Expertise for Registration of Medicinal products in the Republic of Armenia, Form and Description of the Registration Certificate and the List of variations of medicinal products registered in the Republic of Armenia that do not require new registration.''

1. General provisions

1.1. It is allowed to import, produce, store, distribute, sell and use only those medicinal products on the territory of the Republic of Armenia, which are registered in the Republic of Armenia.

1.2. Registration of medicinal products, rejection and withdrawal of registration is carried out by the Ministry of Health of the Republic of Armenia, and of veterinary vaccines, serums and diagnostics - by the Ministry of Agriculture of the Republic of Armenia.

1.3. Registration of medicinal products is conducted based on the results of the scientifically justified criteria and expertise of safety, efficacy and quality of medicinal products. Expertise of medicinal products for registration is carried out by the Scientific Center of Drug and Medical Technology Expertise (hereinafter referred to as 'Scientific Center'[1]).

1.4. Every registration of medicinal products is carried out according to each manufacturer (firm), and also each country of origin, if the production of the same medicinal product is carried out in different countries by the same manufacturer.

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1.5. The quality of medicinal products registered in the Republic of Armenia shall comply with the requirements of currently used official Pharmacopoeias in the Republic of Armenia: the XI State Pharmacopoeia of the former USSR, the European Pharmacopoeia(Ph Eur), the International Pharmacopoeia (Ph Int), the American Pharmacopoeia (USP), the British Pharmacopoeia (BP), the German Pharmacopoeia (DAP), the German Homeopathic Pharmacopoeia (HAB), the French Pharmacopoeia (PhF) and in some cases - temporary Pharmacopoeial monographs approved by the Ministry of Health of the Republic of Armenia.

1.6. The following is subject to registration in the Republic of Armenia:

- new original and generic medicinal products (including immunological, veterinary, homeopathic),
- additional dosage strengths, pharmaceutical forms and new indications of registered medicinal products,
- new combination of medicinal products.

1.7. Registration is not required for medicinal products, which are made in Pharmacies in accordance with Prescriptions and in the cases defined by the Government of the Republic of Armenia.

1.8. The period of validity of registration of medicinal products in the Republic of Armenia is five years. At the expiry date of the term of the registration of the medicinal product it is subject to new registration.

1.9. In case of changes in the composition, manufacturing technology, international nonproprietary names of registered medicinal products, as well as in case of new therapeutic indications, medicinal products are subject to new registration. The Ministry of Health defines the List of variations of medicinal products registered in the Republic of Armenia that do not require new registration (Appendix 2).

1.10. In accordance with the order of the Ministry of Health during 30 days after issuing the Registration certificate the information on registered medicinal product is included in the State Register of Medicinal products of the Republic of Armenia which is published according to the regulation.

2. Submission of Application for registration

2.1. For the purpose of registration of medicinal products, the manufacturer or its authorized representative (hereinafter referred to as 'applicant') submits a required documentation according to the approved lists (Appendix 1.1-1.9), samples of medicinal products and reference standards to the Scientific Center. Documents are submitted in Armenian, Russian or English and also on CD, if available.

2.2. The applicant is responsible for the authenticity of documents and correctness of information.

2.3. Applicant shall submit samples of medicinal products in Armenian, or in Russian, or in English (for prescription medicinal products) packaging and labeling: two consumer packages (checking-identification and laboratory-arbitrage) and in necessary quantities (in consumer packages) required for laboratory expertise complied with the specifications and methods of analyses (pharmacopoeial monographs, etc.)

3. Registration procedure

3.1. Preliminary examination of submitted documentation and samples is carried out by the Scientific Center within maximum 10 days about which the applicant receives a written notification with indication of the expertise fee (Appendix 3).

3.2. Expertise for registration is started after payment of expertise fee as an advanced payment. The date of payment is considered as the start point of the expertise. Maximal duration of expertise is 180 days.

3.3 The applicant may request for withdrawing the application at any time before the end of the expertise. In this case the submitted documentation, samples and reference standard as well as the expertise fee are not returned to the applicant.

3.4 In case of failing to pay the expertise fee within 6 months upon receiving the written notification on payment, the applicant has to submit a new application.

3.5 For medicinal product of major therapeutic and public health interest intended for the treatment of serious or life threatening diseases or condition the expertise for registration may be carried out within the framework of state budget by the Government order. The list of this low demand but vital medicines is adopted by the Ministry of Health (available in the website of Scientific center).

3.6 The expertise of pharmacological, toxicological, clinical and pre-clinical studies, technological procedures, specifications and methods of analyses, methods of manufacturing and quality control is carried out in terms of assuring the compliance of a medicinal product with the approved requirements of quality, safety, efficacy and manufacturing standards.

3.7 If the results of the laboratory expertise of the medicinal product are negative, the applicant may submit new samples of the medicinal product of two series different from the previous one, in a sufficient quantity to carry out two laboratory testing.

3.8 If the information provided for the purpose of evaluating the quality, safety and efficacy of medicinal products is inadequate, the Scientific Center may request additional documentations, samples and data. The period for providing of the required supplementary documents, samples of medicinal products and information is not included in the expertise period. In case the applicant fails to submit required documentation and/or samples and/or reference standards within 6 months, the expertise is suspended and the application is annulated.

3.9 After the expertise the Scientific Center submits the assessment report to the Pharmacological Council of the Ministry of Health within 5 days.

3.10 Receiving the results of the expertise, the Pharmacological Council of the Ministry of Health provides recommendation about registration or refusal of registration of the medicinal product in the Republic of Armenia, as well as conclusion about including the medicinal products in the lists (Controlled medicinal product, Non-prescription medicinal product, Essential medicinal products) adopted in the Republic of Armenia within 15 days. The notice about the conclusion of the Pharmacological Council of the Ministry of Health should be sent to the applicant within 5 days.

3.11 The decision about registration of medicinal product is made by the Ministry of Health within 10 days on the base of the expertise results, conclusion of the Pharmacological Council and payment of state tax in accordance with established procedure and amount (Appendix 4) to the appropriate account of the State Treasury of the Republic of Armenia (in case the payment is made in foreign currency – in accordance with the actual at date exchange rate established by the Central Bank of RA).

3.12 The registration procedure will be suspended if the state tax is not paid in accordance with the established procedure and amount by the applicant within 30

days after notification about positive conclusion of the Pharmacological Council of the Ministry of Health. In future the expertise for registration of medicinal product will be conducted due to the established procedure by applying new application.

3.13 The registration certificate should be issued to the applicant within 10 calendar days according to the order of the Ministry of Health about registration of the medicinal product.

3.14 The manufacturer should inform the Scientific Center about any changes of the registered medicinal product by submitting relevant documentation on changes. The submitted documentation (sample) is evaluated within 30 days, and after the approval by the Ministry of Health of the Republic of Armenia is included into the registration documentation. Variations not requiring new registration (Appendix 2) are taken into consideration. In case of changes in the name of the medicinal product, the name of manufacturer or the marketing authorization holder, additional presentation and packaging - the registration certificate is re-formulated by adding number of variation to the number of registration certificate.

3.15 Both approved documentation and sample are used as a base for identification, quality control and official information regulation in all stages of regulation in the Republic of Armenia. The sample of a medicinal product includes, immediate and/or outer packaging, labeling, instruction for use as well as color mock-ups.

3.16 The results of expertise for registration of medicinal products can be appealed according to the legislation of the Republic of Armenia.

4. Rejecting the registration of medicinal products and withdrawal.

4.1. The registration of medicinal products is rejected if the following is available:

- a negative conclusion of the expertise.
- alerts on the medicinal product received from international specialized sources
- the medicinal product contains chlorofluorocarbons (CFC), except those medicinal products, which list is approved by the Ministry of Health

4.2. The applicant is informed on the rejection of the medicinal product registration within 10 days.

4.3. The registration of a medicinal product may be withdrawn and the circulation of the medicinal product may be suspended, if the following is available:

- a notification from the manufacturer,
- non-compliance with the adopted criteria of quality, safety and efficacy of the medicinal products, including new serious adverse reactions,
- alerts on the medicinal product received from international specialized sources.

4.4. The information on withdrawal of the medicinal product registration is provided by the Ministry of Health in the specialized and official publications within 10 days.

4.5. The manufacturer covers the expenses of withdrawing the medicinal product from regional pharmaceutical market.

4.6. Decisions on registration withdrawal of the medicinal product may be appealed according to the Legislation of the Republic of Armenia.

Appendix 1.1

List

Of the documents required for the registration of generic medicinal products in the Republic of Armenia

1. Application form (Appendix 1.8).
2. Registration certificate of the medicinal product issued by the country of origin (either original or verified copy).
3. Certificate of Good Manufacturing Practice (GMP) issued by the authorized body of the country of origin (for manufacturers in the Republic of Armenia and CIS countries- manufacturing license if the GMP certificate is not available-either original or verified copy).
4. Registration status in other countries.
5. Summary of Product Characteristics (Appendix 1.9).
6. Instruction for use for specialists and patients.
7. Qualitative and quantitative composition of the medicinal product (including excipients).
8. Pharmacopoeial monograph(s) and/or control method(s) or specification(s) of the finished medicinal product and its ingredients, packaging specification(s) (2 copies).
9. Quality certificates of the active substances and excipients of the medicinal product.
10. Summary lot protocol of vaccines and serums and the Lot Release certificate issued by the NRA of the country of origin.
11. Data on stability study and shelf life of medicinal product.
12. Brief description of the technological process, chemical, technological and equipment schemes of the production, including controls of critical steps.
13. Data on pharmacokinetic and/or bio-equivalence and/or limited clinical trials of the medicinal product. If they are not available, for manufacturers of the Republic of Armenia and CIS countries- data on acute toxicity study.
14. Information on pharmacological, toxicological and clinical trials (literature references or own data).
15. For veterinary medicinal products - information on maximum residue limits in the foodstuff (meat, milk, egg, etc.). The time limitation of foodstuff use.
16. The label and packaging of the medicinal product and/or its color mock-ups and specimens (also electronic version) for all presentations mentioned in the application.

17. Certificate (verified copy) or verified extract from appropriate register about legal protection of trademark issued by the Intellectual Property Agency of the Ministry of Economy of the Republic of Armenia
18. Periodic Safety Update Report.
19. TSE-Certificate of Suitability for the material of animal origin.

Appendix 1.2

List

of the documents required for the expertise of medicinal products containing new active substances for registration in the Republic of Armenia

1. Application form (Appendix 1.8).
2. Registration certificate of the medicinal product issued by the country of origin (either original or verified copy).
3. Certificate of Good Manufacturing Practice (GMP) issued by the authorized body of the country of origin (for manufacturers in the Republic of Armenia and CIS countries-manufacturing license if the GMP certificate is not available-either original or verified copy).
4. Registration status in other countries.
5. Summary of Product Characteristics (Appendix 1.9).
6. Instruction for use for specialists and patients .
7. Qualitative and quantitative composition of the medicinal product (including excipients).
8. Pharmacopoeial monograph(s) and/or control method(s) or specification(s) of the finished medicinal product and its ingredients, packaging specification(s) (2 copies).
9. Quality certificates of the active substances and excipients of the medicinal product.
10. Data on stability study and shelf life of medicinal product.
11. Brief description of the technological process, chemical, technological and equipment schemes of production, including control of critical steps.
12. Reports on the pre-clinical studies of pharmacological activities, pharmacodynamic, pharmacokinetic and adverse reactions of the medicinal product.
13. Reports on the pre-clinical studies of the safety (acute, sub-chronic and chronic toxicity, genotoxicity, carcinogenicity, reproductive and developmental toxicity, local tolerance, antigenicity, Immono-toxicity and other toxicity studies).
14. Reports on the clinical trials on the specific activity, pharmacodynamic, pharmacokinetic and adverse reactions of the medicinal product.
15. For veterinary medicinal products - information on maximum residue limits in the foodstuff (meat, milk, egg, etc.). The time limitation of use of the foodstuff use.
16. The label and packaging of the medicinal product and/or its color mock-ups and specimens (also electronic version) for all presentations mentioned in the application.

17. Certificate (verified copy) or verified extract from appropriate register about legal protection of trademark and/or patent issued by the Intellectual Property Agency of the Ministry of Economy of the Republic of Armenia
18. Periodic Safety Update Report.
19. TSE-Certificate of Suitability for the material of animal origin.

Appendix 1.3

List

of the documents required for expertise of homeopathic medicinal products for registration

in the Republic of Armenia

1. Application form (Appendix 1.8).
2. Registration certificate of the medicinal product issued by the country of origin (either original or verified copy).
3. Certificate of Good Manufacturing Practice (GMP) issued by the authorized body of the country of origin (for manufacturers in the Republic of Armenia and CIS countries-manufacturing license if the GMP certificate is not available) (either original or verified copy)
4. Registration status in other countries.
5. Summary of Product Characteristics (Appendix 1.9).
6. Instruction for use of the combined homeopathic medicinal products.
7. Qualitative and quantitative composition of the medicinal product (including excipients).
8. Pharmacopoeial monograph(s) and/or control method(s) or specification(s) of the finished medicinal product and its ingredients, packaging specification(s) (2 copies).
9. Quality certificate of the medicinal product.
10. Data on stability study and shelf life of medicinal product.
11. Data on the safety and efficacy of the medicinal product.
12. The label and packaging of the medicinal product or its color mock-ups and specimens (also electronic version) for all presentations mentioned in the application.

Appendix 1.4

List

of the documents required for registration expertise of the medicinal product registered in the Republic of Armenia and produced in other countries by the same manufacturer

1. Application form (Appendix 1.8).
2. Registration certificate of the medicinal product issued by the country of origin (either original or verified copy).
3. Summary of Product Characteristics (Appendix 1.9).
4. Instruction for use for specialists and patients.
5. Certificate of Good Manufacturing Practice (GMP) issued by the authorized body of the country of origin (for manufacturers in the Republic of Armenia and CIS countries-manufacturing license if the GMP certificate is not available) (either original or verified copy)
6. Brief description of the technological process, chemical, technological and equipment schemes of production, including control of critical steps
7. Data on pharmacokinetic and/or bioequivalence and acute toxicity studies of the medicinal product.
8. Pharmacopoeial monograph(s) and/or control method(s) or specification(s) of the finished medicinal product and its ingredients, packaging specification(s) (2 copies).
9. Reference stating that above the mentioned documents of the medicinal product, it has not been changed since last registration.
10. The label and packaging of the medicinal product or its color mock-ups and specimens (also electronic version) for all presentations mentioned in the application .

Appendix 1.5

List

of the documents required for registration expertise of the additional dosages of the medicinal product registered in the Republic of Armenia

1. Application form (Appendix 1.8).
2. Registration certificate of the medicinal product issued by the country of origin (either original or verified copy).
3. Summary of Product Characteristics (original or verified copy).
4. Instruction for use for specialists and patients..
5. Qualitative and quantitative composition of the medicinal product (including excipients).
6. Pharmacopoeial monograph(s) and/or control method(s) or specification(s) of the finished medicinal product and its ingredients, packaging specification(s) (2 copies).
7. Quality certificates of the active substances and excipients of the medicinal product.
8. Data on stability study and shelf life of medicinal product.
9. For veterinary medicinal products-information on maximum residue limits in the foodstuff (meat, milk, egg, etc.). Time limitation of foodstuff use.
10. The label and packaging of the medicinal product or its color mock-ups (also electronic version) for all presentations mentioned in the application.
11. TSE-Certificate of Suitability for the material of animal origin.

Appendix 1.6

List

of the documents required for registration expertise of the additional pharmaceutical forms of the medicinal product registered in the Republic of Armenia

1. Application form (Appendix 1.8).
2. Registration certificate of the medicinal product issued by the country of origin (either original or verified copy).
3. Summary of Product Characteristics (Appendix 1.9).
4. Instruction for use for specialists and patients..
5. Qualitative and quantitative composition of the medicinal product (including excipients).
6. Pharmacopoeial monograph(s) and/or control method(s) or specification(s) of the finished medicinal product and its ingredients, packaging specification(s) (2 copies).
7. Quality certificates of the active substances and excipients of the medicinal product.
8. Data on stability study and shelf life of medicinal product.
9. Data on pharmacokinetic and/or bio-equivalence and/or limited clinical studies of the medicinal product.
10. Data on toxicity studies of the medicinal product.
11. Data on clinical trials.
12. For veterinary medicinal products-information on maximum residue limits in the foodstuff (meat, milk, egg, etc.). Time limitation of foodstuff use.
13. The label and packaging of the medicinal product or its color mock-ups and specimens (also electronic version) for all presentations mentioned in the application.
14. TSE-Certificate of Suitability for the material of animal origin.

Appendix 1.7

List

of the documents required for registration expertise of the new indications of the medicinal product registered in the Republic of Armenia.

1. Application form (Appendix 1.8).
2. Registration certificate of the medicinal product issued by the country of origin (either original or verified copy).
3. Summary of Product Characteristics (Appendix 1.9).
4. Instruction for use for specialists and patients.
5. Data on clinical efficacy

Data should be included in the application form

1. Name of the medicinal product
2. International Nonproprietary Name
3. Composition

* active substances
* excipients

4. Dosage strength
5. Pharmaceutical form and route of administration
6. Anatomical-clinical- chemical code (ATC)
7. Presentation and packaging
8. Indications
9. Shelf-life
10. Storage conditions
11. Legal status for supply to the patient in the country of origin
12. Manufacturer (name, address, country)
13. Marketing authorization holder (name, address)
13. Number and expiry date of the registration certificate of the patent and trade mark
14. Applicant (manufacturer or its authorized representative), address, phone number(s), fax, signature, stamp or seal, date of signing.

Appendix 1.9

The summary of the product characteristics

1. **Name of the medicinal product**

2. **Qualitative and quantitative composition** with indication of active substances (international nonproprietary name or chemical name), indication of other excipients knowledge of which is essential for safe and proper administration of the medicinal product.

3. **Pharmaceutical form**

4. **Clinical particulars**
   - 4.1 Therapeutic indications
   - 4.2 Posology and method of administration (where appropriate dosage adjustments in specific patient group should be stated)
   - 4.3 Contraindications
   - 4.4 Special warnings and precautions for use
   - 4.5 Interactions
   - 4.6 Pregnancy and lactation
   - 4.7 Effects on ability to drive and use machines
   - 4.8 Undesirable effects
   - 4.9 Overdose

5. **Pharmacological properties**
   - 5.1. Pharmacodynamic properties
   - 5.2 Pharmacokinetic properties
   - 5.3 Preclinical safety data

6. Pharmaceutical particulars
   - 6.1 Excipients
   - 6.2 Incompatibilities
   - 6.3 Shelf-life

6.4 Storage conditions

6.5 Nature and contents of container

6.6. Special precautions for disposal

7. **Manufacturer** (name, address, country)

8. **Marketing Authorization holder** (name, address, country)

9. **Date of final revision of the text**.

Appendix 2

List

of the minor changes of medicinal products registered in the Republic of Armenia that do not require new registration

1. Changes in the content of GMP certificate or manufacturing license adopted by the relevant bodies of the country of origin that do not include the name, address or country of the manufacturer.
2. Change in the name of a medicinal product (trade name and/or international nonproprietary name) in case the composition and indication of the finished product remain unchanged.
3. Change in the name of manufacturer and/or the name of the marketing authorization holder, in case the country of origin remains unchanged.
4. Replacement of an excipient with a comparable excipient, except for the adjuvant for vaccines or a biological excipient.
5. Change or replacement of coloring agent currently used in the finished product.
6. Addition, deletion or replacement of neutral flavoring agent currently used in the finished product.
7. Change in coating weight of tablets and change in weight of capsule shells.
8. Changes in qualitative composition of immediate packaging except for sterile products.
9. Deletion of one of the therapeutic indications of the finished product (in case safety characteristics remain unchanged).
10. Deletion of one of the routes of administration.
11. Change in the manufacturer of the active substance.
12. Change in batch size of an active substance, in case quality control data of active substance indicate that consistency of manufacturing process remained unaffected and physical properties of active substance remain unchanged
13. Changes in the specification of an active substance due to improvement of test procedure, addition of new methods and tightening of specification limits.
14. Changes in manufacturing process that do not involve a change of specification of finished product, in case the registration expertise of medicinal product for new

manufacturing process proves that safety, quality and efficiency characteristics are unchanged.

15. Change in batch size of the finished product, in case the consistency of manufacture remains unaffected.
16. Changes in the specification of the finished product due to improvement of test procedure, addition of new methods and tightening of specification limits.
17. Changes in synthesis of excipients of the finished product, in case specifications, composition and quantitative impurity profile remain unchanged.
18. Changes in specifications of excipients due to improvement test procedure, addition of new methods and tightening of specification limits.
19. Change in the shelf life of the finished product in case of its prolongation, not exceeding five years.
20. Change in the shelf life of the finished product after first opening.
21. Change in the shelf life of the finished product after dilution or reconstitution
22. Changes in the storage conditions of the finished product
23. Changes in the methods of quality control of active substances of the finished product
24. Changes in the methods of quality control of the finished product
25. Change to comply with an update of the relevant monograph of the Pharmacopoeias
26. Changes in testing methods of non-pharmacopoeial excipients
27. Change to a test procedure of the immediate packaging of the finished product
28. Changes in testing methods of the supplier or devices
29. Changes in pack shape, size, design and number of units (e.g. tablets, ampoules, etc.) in a pack
30. Change or addition of imprints, bossing or other markings on tablets or printing on capsules.
31. Change of dimensions of tablets, capsules, suppositories or pessaries without a change in the quantitative composition and mean mass.

Appendix 3

FEES
payable for medicinal product registration expertise in the Republic of Armenia

№	**Type of application for registration**	**Expertise Fee (including VAT)** (thousand Armenian dram)
1	**The first dosage form and dosage strength of generic medicinal products**	**900**
	• **each additional pharmaceutical form**	**450**
	• **each additional dosage strength**	**240**
	• **each new indication**	**450**
2	**New combination of known medicinal products**	**1200**
3	**The first pharmaceutical form and dosage strength of the medicinal products containing new active substances**	**2250**
	• **each additional pharmaceutical form and dosage strength**	**1200**
4	**The first pharmaceutical form and dosage strength of Homeopathic medicinal products**	**240**
	• **each additional pharmaceutical form, dosage strength and new indication**	**60**
5	**Herbal preparations and other preparations of natural substances and Dietary supplements**	**240**
6	**Re-formulating of the Registration Certificates, in cases of the following changes, connected with the name of the product and/or manufacturing company, packaging and other minor changes which do not affect the approved quality, safety and efficacy of the product.**	**24**

If the same manufacturer's medicinal product, already registered in the Republic of Armenia, is also manufacturing in other countries, the payment amount for every additional country-manufacturer's medicinal product registration will be half of the approved amount indicated in the table above.

Rate of the State Tax for Registration in the Republic of Armenia

№	Type of application for registration	Rate of the State Tax (Armenian dram)
1.	First and additional pharmaceutical form and dosage strength of medicinal products containing new active substances	70000
2.	New combination of known medicinal products	40000
3.	First and additional pharmaceutical form and dosage strength of generic medicinal products	40000
4.	New indications	10000
5.	Herbal preparations and other preparations of natural substance	10000
6.	Homeopathic medicinal products	2000
7.	Dietary supplements	20000
8.	Re-formulating of the Registration Certificates, in cases of the following changes, connected with name of product and/or manufacturing company, packaging and other minor changes which do not affect the approved safety, quality and efficacy of the product.	5000