

Regulations on the Foundation of Japanese Center for the Validation of Alternative Methods

(The foundation of JaCVAM)

Article 1:

Japanese Center for the Validation of Alternative Methods (JaCVAM) is founded in the Biological Safety Research Center (BSRC), the National Institute of Health Sciences (NIHS).

(The aim and duties of JaCVAM)

Article 2:

1. The aim of JaCVAM is to contribute to the introduction of the alternative methods, which assist the enhancement of 3Rs (Reduction, Refinement and Replacement) concerning animal studies, into regulatory studies as much as possible, while securing the safety of citizens, in the evaluation of the safety of the materials related to the duties of NIHS. This is also helpful for the preparation of the documents applied for the approval of the manufacture and the sale of domestic quasi-drugs.

2. For this aim, JaCVAM evaluates the utility of studies on the safety of materials related to the duties of NIHS and its limitations, and conducts validations necessary for the evaluation. In addition, JaCVAM is concerned with the domestic and international cooperation and deals with international matters in related fields.

(The foundation of the organizations for the activity of JaCVAM)

Article 3:

1. For the proper management of JaCVAM, the Advisory Council, the Steering Committee and the Regulatory Acceptance Board are founded. Upon the commission by the Steering Committee, the Peer Review Panel and the Validation Management Team are organized.

2. The Director General of NIHS commissions the members of these organizations. The term of office of the members is 2 years, and reappointment is possible.

3. The Section for the Evaluation of the Novel Methods in the Division of Pharmacology, BSRC, NIHS serves as the secretariats of these organizations.

4. The meetings of these organizations are effective when 2 thirds or more of the members attend. The decisions are made by the consent of all the attendance. If such consent is not achieved, the judgments agreed by 2 thirds or more of the attendants are regarded as decisions. In these cases, the points not obtaining the full consent are described in the reports.

(The Advisory Council)

Article 4:

1. The Advisory Council receives the reports from the Steering Committee once or more a year, and discusses about and provides advice on them.

2. The Advisory Council consists of about 10 members including the Director General of NIHS, the Head of BSRC, the administrator of the agency in charge, the expert of animal welfare, the representatives of related societies, the representatives of the industry and other persons judged necessary by the chairperson. The Director General of NIHS serves as the chairperson.

(The Steering Committee)

Article 5:

1. The Steering Committee discusses about and decides the scientific validity and the budget and manpower necessary for the execution of the evaluation, on the plans for the selection and the evaluation of novel and modified methods to be examined by JaCVAM. In addition, the Steering Committee discusses about the reports from the Regulatory Acceptance Board, adds opinions to the documents of those from JaCVAM about the methods judged proper as administrative ones, and transmits the documents to the agencies of the Minister of Health, Labour and Welfare (MHLW) in charge as well as announces them officially. In addition, the Steering Committee appoints the chairpersons of the Peer Review Panel and the Validation Management Team.

2. The Steering Committee consists of the Director General of NIHS, the

members of the Steering Committee of BSRC (the Head of BSRC, the Directors of the Divisions of Cellular and Molecular Toxicology, Pathology, Pharmacology, Genetics and Mutagenesis, Risk Assessment, and the Head of the Animal Management Section in the Division of Cellular and Molecular Toxicology), the administrator of MHLW in charge, the administrator of the Independent Administrative Cooperation, Pharmaceuticals and Medical Devices Agency in charge and the Head of the Section of the Evaluation of Novel Methods in the Division of Pharmacology, BSRC. The Head of BSRC serves as the chairperson.

(The Regulatory Acceptance Board)

Article 6:

- 1) The Regulatory Acceptance Board examines the opinions from the reports of the Peer Review Panel, the background information, and the public comments, discusses about the submitted methods on the basis of their scientific validity, administrative utility, and social acceptability, and prepares the final reports.
- 2) The Regulatory Acceptance Board consists of the Head of BSRC and the experts of the safety and the statistical analysis and the persons necessary considered by the chairperson. The Head of BSRC serves as the chairperson.

(The Peer Review Panel)

Article 7:

- 1) The Peer Review Panel evaluates the submitted methods as a third party, prepares the proposal of the evaluation reports, and serves it to public comments. If necessary, the Peer Review Panel proposes the execution of the validation and the points to be examined in the validation. Considering the results obtained, the Peer Review Panel discusses, prepares the reports as the Peer Review Panel, and presents them to the Steering Committee.
- 2) The Peer Review Panel is organized on individual submitted methods by the Steering Committee, and consists of the experts of the safety of chemicals and the statistical analysis who do not contribute to the development or the

validation of the method. The chairperson of the Peer Review Panel appointed by the Steering Committee selects the members of the Peer Review Panel with discussion with the Secretariat.

(The Validation Management Team)

Article 8:

1) The Validation Management Team plans the validations and executes them. In addition, considering the results of the validations, the Validation Management Team prepares the reports of the validations including recommended protocols, and submits them to the Steering Committee. The Validation Management Team is organized when the Peer Review Panel admits the validation is necessary.

2) The chairperson of the Validation Management Team is appointed by the Steering Committee, and selects the members with discussion with the Secretariat.

(The Secretariat)

Article 9:

The Secretariat conducts official works related to the management of JaCVAM, supports the organizations prescribed from the Articles 4 to 8, and cooperates and copes with domestic and international societies and organizations related to the evaluation of alternative methods for the safety evaluation of substances including chemicals. In addition, the Secretariat collects, classifies and presents the information related to the evaluation of studies. When necessary, the Secretariat recommends candidates for the chairpersons of the Peer Review Panel and the Validation Management Team to the Steering Committee. Further, the Secretariat advises the chairpersons on the selection of the members of each committee.

(Detailed regulations)

Article 10:

Other than these regulations, particles necessary for the activity of JaCVAM are established separately as detailed regulations by the Steering

Committee.

(Revision of the regulations)

Article 11:

These regulations are revised by the discussion in the Steering Committee and the approval of the Director General of NIHS.

(Additional rules)

Article 1:

These regulations are in force from May 1st, 2007.

Article 2:

The revision of the part of these regulations is in force from July 31st, 2009.

Article 3:

The revision of the part of these regulations is in force from April 20th, 2011.