

# Method

**Searching for evidence.** Based on the previous edition of GEMA<sup>2</sup>, published in 2009, and following the recommendations for Updating Clinical Practice Guidelines in the National Health System<sup>3</sup>, the members of the Executive Committee undertook a systematic search to select and evaluate articles on asthma published between 2009 and 2014 (Pro-GEMA Project). After reviewing high impact factor journals of Pneumology, Allergology, Pediatrics, Primary Care, Internal Medicine and Otorhinolaryngology, which were also classified between the two first quartiles of their specialty field, a total of **184** documents were selected (abstracts available at <http://www.progema-gemasthma.com/foco.html>) that were considered of interest for updating this guideline. All these documents were provided to the authors for evaluation. Furthermore, authors were encouraged to perform their own literature searches for specific topics. To this end, the procedure normally used to develop clinical practice guidelines was followed<sup>4</sup>. Also, the reference lists of the main international practice guidelines<sup>5,6</sup> were reviewed in order to identify the most relevant systematic reviews and clinical trials. These guidelines were searched in specialized databases (*National Guideline Clearinghouse*, *National Library of Guidelines*) and the TRIP medical literature meta-search engine database. Databases from the *Centre for Reviews and Dissemination (DARE y HTA database)* and *The Cochrane Library* were consulted with a view to identifying systematic reviews and evaluation of additional technologies. The search was completed with an update of the systematic reviews from the date of search and relevant studies included in the main electronic databases of original studies (*MEDLINE*, *CENTRAL* and *EMBASE*).

**Classification of the evidence.** To assess the quality of evidence, an alphabetic classification was used (table 0.1) that classifies the information into four categories (A, B, C, D) reflecting the grade of confidence in the results obtained in the available studies. Category A would correspond to a high quality evidence and D to a very low quality. For category A, confidence in the results is high and the potential modification of available findings by further studies is unlikely. In contrast, for lower categories, C or D, the confidence level will be low or very low, and there is a high probability that further studies will modify the results, or even the direction of the effect. However, it must be remember that this system is very useful to categorize the evidence regarding therapeutic efficacy of drugs or other treatment, but the effect of other interventions may be

underestimated. This can explain why evidence from studies aimed at determining the appropriateness of some diagnostic procedures has often been assigned a level of evidence C.

Taking into account the recent emergence of new approaches used to classify the quality of evidence based on aspects other than the study design<sup>7,8</sup>, some of the characteristics of the GRADE framework were used (<http://www.gradeworkinggroup.org/>), although the GRADE system was not applied in full.

**Classification of recommendations.** To classify the relevance and consistency of clinical recommendations, the same method used in the previous edition of GEMA<sup>2</sup> was followed, in which recommendations were categorized in two levels: robust recommendations (R1), that is, those associated with more benefits than risks according to the opinion of the group of authors, and weak recommendations (R2), that is, those in which some uncertainty exists as to whether its application might entail more benefits than risks. To carry out this distribution in R1 or R2, the quality of information was weighed (based on the above-mentioned classification), along with the balance between risks and benefits of interventions, the costs (according to the available specialized literature), and the patients' values and preferences (through the participation of FENAER members).

Table 0.1. Classification of the quality of evidence

Categories of evidence	
A	SR of RCTs with or without MA; and RCTs with low risk of bias. Evidence based on a substantial number of well-designed studies with consistent results.
B	SR of RCTs with or without MA; and RCTs with moderate risk of bias. Evidence obtained from a limited number of studies and/or inconsistent results.
C	Evidence obtained from non-randomized, observational or uncontrolled studies.
D	Clinical experience or scientific literature that cannot be included in category C.

Abbreviations: MA, Meta-analysis; RCTs, randomized controlled clinical trials; SR, Systematic reviews.

The categorization of the recommendation level was established by consensus, first of the authors (see below for the working method used) and finally by the agreement of reviewers (through the Delphi method), whose opinions were binding for the final version of all recommendations.

**Text and recommendations: drafting and consensus building.** The writing process was based on a pyramidal consensus system going from a multidisciplinary thematic mini-consensus by chapter to a large final consensus among all authors and reviewers. Based on the document of the previous edition and the new references on asthma published between 2009 and 2015, a group of authors and coordinators made up by experts from the participating scientific societies drew up the new chapter sections they were assigned (including the classification of evidence and recommendations). The authors submitted their texts to each chapter coordinators who were members of the GEMA Executive Committee. After unifying

and reviewing the texts, the chapter coordinator submitted the draft to the authors of each chapter in order to reach the first partial consensus. After implementation of changes, all chapters were brought together in one single document which, in turn, was sent to all authors and coordinators for telematics discussion (and for face-to-face discussion, when necessary) and approval. The resulting document was submitted to experts in the methodology of clinical practice guidelines from the INPECS (Instituto para la Excelencia Clínica y Sanitaria [Institute for Clinical and Healthcare Excellence]), who made a critical review of the methodology and writing of both the text and the recommendations. Finally, after these modifications and improvements, recommendations were revised and agreed on (through the Delphi method) by a group of experts in asthma from the participating societies. Recommendations not achieving a certain consensus level were removed from the final document.