An internet portal for the development of clinical practice guidelines

W.J. Höhne¹; T. Karge¹; B. Siegmund¹; J. Preiss¹; J.C. Hoffmann²; M. Zeitz¹; U.R. Fölsch³

¹ Charité, Campus Benjamin Franklin, Medical Department I, Berlin, Germany; ² Medizinische Klinik I, St. Marienkrankenhaus, Ludwigshafen, Germany; ³ Department of General Internal Medicine, University Hospital Schleswig-Holstein, Kiel, Germany

Keywords

Clinical practice guideline, internet portal, authoring, voting, quality management

Summary

Background: The complexity and quality requirements for the development of clinical practice guidelines steadily increase. Internet technologies support this process by optimizing the development process.

Objective: The aim of this internet based solution was to facilitate the development of clinical practice guidelines.

Methods: An internet portal was developed allowing for a shared workplace to support clinical practice guideline authoring. It is based on a Content Management System and combines different tools for document handling and editing, communication as well as process and team steering.

Results: Until now, the internet portal has been successfully implicated in the development of six evidence- and consensus-based clinical practice guidelines. Additional German and European clinical practice guidelines are currently generated with support of the internet portal. The available tools allow for a flexible design of the scheduled workflow, depending on the requirements of the respective group. An additional strength of the platform is the advantage to transfer all data from a previous version of a guideline into the next 'life-cycle'.

Conclusion: The application of the portal results in a considerable reduction of costs and development time of the resulting clinical practice guidelines.

Correspondence to:

Wolfgang Höhne
Charité – Campus Benjamin Franklin
Medical Department I
Hindenburgdamm 30
D-12200 Berlin, Germany
Tel: ++49-30-8445-3950
Fax: ++49-30-8441-1308
E-mail: wolfgangj.hoehne@charite.de

Appl Clin Inf 2010; 1: 430–441

doi: 10.4338/ACI-2010-04-RA-0027

received: May 17, 2010

accepted: October 31, 2010

published: November 24, 2010

Citation: W.J. Höhne, T. Karge, B. Siegmund, J. Preiss, J.C. Hoffmann, M. Zeitz, U.R. Fölsch. An internet portal for the development of clinical practice guidelines

Appl Clin Inf 2010; 1: 430–441

<http://dx.doi.org/10.4338/ACI-2010-04-RA-0027>

1. Background

Clinical practice guidelines (CPGs) play an important role as treatment guidelines in daily clinical practice. A prerequisite of this trend was the development of rigorous quality criteria for such guidelines, which has been promoted by different national and international organisations. As a result, the requirements for the development process of clinical practice guidelines have been sharpened steadily. A modern evidence- and consensus-based guideline [1, 2], in Germany known as a S3 CPG [3], requires both the evaluation of highest possible evidence and the establishment of a consensus between all participants, from the specialized medical professional to patient organizations. To obtain acceptance, the finished guideline has to pass through an intensive reviewing process and assessment by instruments such as AGREE [4] or its German counterpart, DELBI [5]. Prerequisite for these steps is a comprehensive method report. Because of its limited lifetime moreover, the CPG also requires a cyclic update.

As a consequence, typical development time-periods for such guidelines quickly reach up to three years. At the same time, the development is expensive. There have been reported costs per guideline of € 150.000 in the Netherlands or of £ 400.000 for British NICE guidelines [6]. Consequently, reduction of expenses represents one important objective in CPG methodology [7] as well as simplification of time and cost intensive steps in the development of CPGs should be considered.

Modern Information Technology (IT) techniques such as internet portals, content management systems or communication platforms have the potential to contribute to the solution of these problems. More remarkable, no specialized IT solutions have been described for accompanying support of the CPG development so far.

2. Objective

Based on the schedule, defined by the German Association of the Scientific Medical Associations (AWMF), our objective was to develop an IT solution for the support of as many as possible steps within the CPG lifecycle (►Figure 1). In order to substantially reduce time and expenses, especially such activities as voting or collaborative work with documents had to be facilitated. A critical point was the management and surveillance of the development process by steering committees as well as workgroup leaders.

For the users the resource requirements should be as low as possible and limited to standard tools such as browsers. Platform independence (Windows, Mac OS/X, Linux/Unix) of the clients had to be guaranteed, while application of proprietary standards had to be avoided. As a consequence and to prevent licence conflicts and incalculable operational costs, the solution had to be limited to open-source and freeware components. At the same time, it was not the aim to postulate a rigid workflow, but the users should have the possibility to freely combine available tools and to scale them to their defined demands.

The main components, activities and resources, involved in CPG development are shown in ►Figure 2. The following active parts can be identified:

1. *Clinical Practice Guideline*: It is the main object of development, which has to be processed, stored / archived and communicated between the developers and later to the public. CPG self is accompanied by other documents, e.g. older versions and the CPG-specific literature.
2. *Guideline Developers*: They build the component, which drives all activities within the development process and represent the "central processing unit". Main activities are adapted according to evidence assessment, consensus finding and formulation of the CPG and its recommendations. In parallel, the developers coordinate and organize all activities. These are highly interactive tasks, which require powerful instruments for communication and data exchange.
3. *Development process*: It comprises all activities required for the development during the guideline generation. This includes in particular all processes to achieve evidence and consensus such as literature evaluation, voting or consensus conferences. The documentation of this process is one of the main parts of quality management. In addition, there is the need for tools to supervise and manage the process by authorized developers.

On the basis of this model one can identify the tasks which the IT solution should fulfill:

- management, backup and retrieval of documents;
- collaborative processing of documents;
- efficient tools for communication between members of development teams;
- instruments for evidence assessment of documents, recommendations etc.;
- instruments supporting consensus generation, e.g. different forms of voting;
- efficient management of participants, e.g. of their roles and contact data, and secure authentication and access;
- management instruments for monitoring the development process;
- simple, but effective instruments for documentation of the development process, generation of audit trails and of method reports.

None of these components is limited to one step of the development process or is restricted to a defined workflow. Accordingly, the fundamental design principle was not the development of an integrated tool, possibly limited to a single workflow, but rather a universal toolbox for the support of frequently appearing tasks.

3. Methods

To fulfill these requirements, a content management system (CMS) with powerful web interface [8] and a large degree of configurability and extensibility has been chosen. As an optimal framework, the Plone CMS has been selected [9]. Plone, as a portal solution, already includes a full-developed subsystem for accounting and access control, which is relatively simple to extend to a full-fledged, LDAP-based participant management.

A decisive advantage is the application of the universal scripting language Python as an implementation language. This offers a high flexibility for the programming of server-side business processes, compared with solutions based on web-only languages such as PHP or on more complex Java environments. Accordingly, it is an excellent platform for rapid reaction on emerging user requirements, and it comes with a multitude of extensions.

One key component of the portal is a tool for the preparation, execution and evaluation of voting via Internet [10]. It applies a formalized description of questionnaires, based on XML Schema. Various scripts for automated generation of HTML implementations of questionnaires, for monitoring of voting and finally, the statistical analysis of results simplify online voting at a considerable rate. Some features, e.g. the possibility to comment answers, are specially tailored to the demands of consent establishment.

All data, describing a CPG and generated during its development are handled by an external database, using the relational database management system MySQL, with the exception of text documents, which are managed within Plone's internal repository. To ensure a high degree of availability, the portal is hosted on a virtualized Linux server on a VMware ESX cluster.

The client-side part can be kept very simple. As software configuration, it only requires a browser, possibly supplemented by some support for the handling of PDF and office documents. But the latter are optional, since the text editing mostly takes place with internal editors, supplied by the portal. To simplify the access to the Internet, all client-server communication exclusively uses HTTP/HTTPS. Furthermore, all tools are server-side applications, and no download of any executable code is required. Thus, even in secure environments like hospitals the access does usually not raise problems.

4. Results

4.1 General usage of the portal

A view of a typical start page, as seen by a portal user, is shown in ►Figure 3. The middle part of screen represents the working area. At the left side, the main menu is located. There, the user finds all CPGs listed, for which she/he is registered. Thus the user is flexible in switching in between these CPGs. At the right column, the developer finds news and appointment information. The upper part contains general functions such as language selection (currently English / German). With the 'Preferences' function, each user can maintain the personal contact information.

The available functionality depends on the development process, defined by coordinators of the concrete CPG. Default functionality is organization including internal news and appointments, communication tools and a shared workspace for document editing and exchange. Normally, tools for consensus finding, online voting as well as for literature work and evidence assessment are available.

4.2 User management

For the management of members of development groups, a detailed, CPG-specific system of roles has been implemented. A role defines the rights of a user and also his personal view of the portal. From a total of ten guideline specific roles, the most important are:

- *coordinator*: responsible for organization, monitoring, quality management, final decisions; no access to workgroup level data; no right to vote
- *workgroup leaders*: responsible for management of their workgroup (WG); WG-local decision making; full access to WG local data; right/obligation to vote
- *workgroup member*: full access to WG local data; right/obligation to vote
- *patient*: WG member; right, but no obligation to vote
- *expert*: WG member; no right to vote

Each participant can represent different roles within a CPG development. Users, participating in different CPG developments, authenticate themselves with one single account, allowing them after login to access to all guidelines, for which they are registered. With registration of a participant, usually performed by a coordinator, the user database will be automatically updated with contact data, mailing lists etc.

4.3 Communication and collaboration

The user database builds the basis of a widespread communication subsystem. The most elementary function is the allocation of mailing lists: They can map the role scheme, but also the workgroup structure and are supplemented by efficient searching mechanisms for contact data.

For bi- and multidirectional communication discussion forums are the preferred form of communication. Each user can create a discussion thread and invite other members to participate. Compared with e-mail, these forums have some essential advantages. Firstly, they are public, either for a whole CPG team or for limited groups, e.g. coordinators or individual working groups. Thus, each member interested in a specific problem can track the discussion and contribute to the topic. Secondly, the centralized recording of a discussion thread is an effective documentation of decision processes. To perform short-time communication, chat-rooms and telephone conferences are available.

For collaborative document processing, the portal supplies some more complex tools. Within dedicated workspaces, working groups can create and edit complex documents. With FCKEditor a powerful text processor is available for this purpose (►Figure 4). In combination with version-control and history functions, including a fallback option, this tool is the core of document processing.

The portal also supplies a wiki tool, whose main scope is the development of dictionaries and ontologies. But it can be applied to compile definitions of any kind. Each developer can request the

definition of some item, whose meaning appears unclear. In this and all other cases, a wiki process will be initiated, which leads to a harmonized definition. The definition then can be declared mandatory by coordinators and adopted into the final CPG text.

4.4 Evidence assessment

The portal supplies an interface to myNCBI keyword search. CPG developer groups have access to private workspaces to perform a preselection of references on the basis of abstracts and metadata. Full texts of preselected sources can be uploaded into the portal. Because of German copyright regulations the portal guarantees an exclusive access to these documents only by authorized group members. This is part of user and session management of the portal. The documentation of evidence assessments occurs by standardized evidence tables. The assessments and also all full text documents, reported in the CPG, are archived for documentation purposes.

4.5 Consensus finding

Activities in consensus gathering are very complex, time-consuming and expensive tasks. This lies in the philosophy of the Nominal Group Process with its numerous meetings, voting and other harmonizing steps. Although this is one prerequisite for high-quality CPGs, its practical implementation often ignores modern technical possibilities, especially those of Internet.

From there, the portal provides a powerful tool to support those tasks. Principal part is a comprehensive system to generate, answer and analyze guideline-specific questionnaires. These questionnaires (►Figure 5) support all essential types of answers, e.g. Likert scales, numerical scales with freely definable endpoints, single- and multiple-choice answers or selection of percent value. In each case, the respondent has the possibility, in some situations also the responsibility, to comment on the decision. In well-founded situations a respondent can declare himself as 'not qualified'. Then she/he can decline the answering of a question.

The decision process can be supported by considerable contextual information, embedded into the questionnaire. The concrete approach depends on the policy, forced by coordinators.

A typical voting period spans over up to two months. Within this time period, each participant has the opportunity to answer the questions or to adjust them, whenever it is required.

The status of the voting can be supervised by coordinators at any time. Moreover, at any time they can request preliminary reports. For all participants, the final report of a vote (►Figure 6) is available immediately after its termination. Of special value are the comments submitted. They contain significant information for the commentary part of the CPG.

Another useful instrument, supported by the portal, is the televoting system in the context of consensus conferences and meetings. The application of these systems considerably reduces the duration and error rate of ballot counts. The results are immediately available for further processing like those of questionnaire surveys. Another important aspect for both procedures, questionnaire based voting and televoting during a meeting, is the fact, that the results are immediately available for the later generation of quality documentation, e.g. method report.

4.6 Usage results

So far, the portal has been applied to finish six evidence- and consensus-based CPGs, four new developments and two updates, both German and European. Two are already published [11, 12], the others are in editing or with the reviewing process. Currently, five additional CPG developments are hosted.

The experience from these applications indicates a considerable reduction of development time and cost. Without the portal, the duration from the designation of participants to the moment when a complete consensus is reached has been on average 2 years for new developments and one year for updates. Taking advantage of the portal, this period could be reduced to less than one year for all new developments and three respectively four months for the updates. Although there are no statistically significant data about usage characteristics up to now, at least a halving of development time seems to be representative.

Remarkable is the increasing shift from meetings toward questionnaire-based surveys for gathering consensus. In doing so, one trend is to pre-vote the key statements of the CPG in the run-up to the consensus conference. The participants of the consensus conference can then focus on really sophisticated questions, which considerably enhances the efficiency of the meeting. Questions, not answered during the meeting, can even be passed back to working groups for revision. Later, the revised version can be submitted to a short, new online voting, which further increases the flexibility.

5. Discussion

The experience, gained from different CPG developments hosted so far, indicates a high flexibility in adapting to individual methodology. Some early expectations, e.g. to fully substitute Nominal Group Processes by pure online votings, have been revised. On the contrary, skillful combination of online votings about CPG recommendations with their subsequent discussion and finalization during a consensus conference has remarkably increased the quality (accuracy, completeness) of their formulation. To optimize the interplay between those two steps, the voting tool has been completely redesigned during the test phase to improve the possibilities to comment answers.

So far, the full integration of tools, required for evidence appraisal, into the portal is not solved satisfactory. Thus, it is planned to extend the portal by a powerful Document Management System, which will support all aspects of literature work without the need of external tools.

The portal is available for further users at anytime. Detailed information can be found on the start page of the portal [8] and in an information brochure [13] available for download from the portal. Terms of portal use and prices can be inquired. The latter depend on the extent of the required tools. Referring to a CPG with up to 10 workgroups and a total of 50 participants, the cost of portal usage normally is below 10.000 €, which matches the minimum expense of a single one-day meeting of such a team.

6. Conclusions

The portal has shown its value in the development of CPGs. For developer teams it offers a high degree of flexibility in the organization. Especially, considerable parts of consensus finding, as yet often connected with numerous meetings, which are difficult to organize and very expensive, can be replaced by platform usage. In addition, the management and control of the development process can be simplified significantly. This includes measures for quality management, which are profoundly supported.

Abbreviations

AGREE	Appraisal of Guidelines Research and Evaluation
AWMF	German Association of the Scientific Medical Association
CMS	Content Management System
CPG	Clinical Practice Guideline
DELBI	German Instrument for Methodological Guideline Appraisal
HTML	Hypertext Markup Language
HTTP	Hypertext Transfer Protocol
HTTPS	Hypertext Transfer Protocol Secure
IT	Information technology
LDAP	Lightweight Directory Access Protocol
NCBI	National Center for Biotechnology Information
NICE	British National Institute for Health and Clinical Excellence
PDF	Portable Document Format
PHP	PHP Hypertext Processor

TMF	German Technology, Methods, and Infrastructure for Networked Medical Research Association
WG	Workgroup
XML	Extensible Markup Language

Conflict of Interest Statement

The authors have not to report any Conflicts of Interest with respect to the content of this manuscript.

Protection of Human Subjects and Animals in Research

The present manuscript does not report results from research including experiments on human subjects or animals.

Acknowledgements

The development and the current operation of the portal have been and are supported by the German Technology, Methods, and Infrastructure for Networked Medical Research Association TMF e.V. The project has also been supported by German Federal Ministry of Education and Research within competence network "Inflammatory bowel diseases" (Grant 01 GI 0486).

Also, our thanks are directed to the colleagues of the Charité Berlin, supporting the daily operation of the portal, especially to Mrs. Schönberg. We also thank the developers of the ESPGHAN Celiac disease guideline for permission to use some snapshots for illustration purposes.

Clinical Relevance Statement

As evidence-based recommendations for health professionals, clinical practice guidelines play an important role in their daily work. The aim of the portal, presented here, is to optimize the quality of these recommendations, by supporting their development. This increased quality also results in increased confidence in CPGs, connected with improved implementation into clinical practice.

Fig. 1 Lifecycle of a clinical practice guideline according to German AWMF schedule

Fig. 2 Model of guideline development

The screenshot shows the TMF Clinical Practice Guidelines Development Portal for Coeliac Disease. The page has a blue header with the TMF logo and navigation links (Home, Contact, Imprint). Below the header, a breadcrumb trail reads 'You are here: Home → Guidelines → Coeliac Disease'. The main content area is titled 'Coeliac Disease' and includes a 'Content' section with sub-sections: Organization, Communication, Survey, and Literature. Each sub-section lists relevant documents and their content counts. For example, under 'Organization', there are links to 'Members and workgroup structure', 'List of group members and email addresses (excel file)', 'Munich (Herrsching) Meeting' (2 elements), and 'London Meeting' (5 elements). Under 'Survey', there are links to 'Personal survey progress', 'Pre Istanbul voting - evaluations' (5 elements), and 'Recent surveys' (3 elements). Under 'Literature', there are links to '15 studies sens spec ppv npv prev 20 08 2009.xls', 'Abstraction of 20 studies 20 08 2009.doc', 'Exclusion criteria 20 08 2009.doc', 'Marsh criteria 20 08 2009.xls', 'Tests studied 20 08 2009.doc', and 'Guidelines' (6 elements). The right sidebar contains 'News' (NICE response, Survey evaluations) and 'Upcoming Appointments' (Consensus Conference). At the bottom right, there is a calendar for September 2010.

Coeliac Disease

ESPGHAN guidelines for the diagnosis of coeliac disease in children

Content

Organization

- Members and workgroup structure
- List of group members and email addresses (excel file)
- Munich (Herrsching) Meeting Contents: 2 element(s)
- Documents related to the meeting in Munich (Herrsching) august 31st 2009
- London Meeting Contents: 5 element(s)

Communication

- Mailing lists
- Discussion forum Contents: 3 element(s)

Survey

- Personal survey progress
- Pre Istanbul voting - evaluations Contents: 5 element(s)
- Recent surveys Contents: 3 element(s)

Literature

- 15 studies sens spec ppv npv prev 20 08 2009.xls
- Abstraction of 20 studies 20 08 2009.doc
- Exclusion criteria 20 08 2009.doc
- Marsh criteria 20 08 2009.xls
- Tests studied 20 08 2009.doc
- Guidelines Contents: 6 element(s)

News

- NICE response (Oct 5) on our LEVEL 3 study selection (Guideline Coeliac Disease) 2009-10-05
- Survey evaluations now available (Guideline Demo-LL) 2008-02-18

Upcoming Appointments

- Consensus Conference Frankfurt/Main, 2011-02-27 (Guideline Demo-LL)

September 2010

Su	Mo	Tu	We	Th	Fr	Sa
			1	2	3	4
5	6	7	8	9	10	11
12	13	14	15	16	17	18
19	20	21	22	23	24	25
26	27	28	29	30		

Fig. 3 Start page with index of contents of a typical CPG workspace

Edit Document

by Troncone — last modified 2009-10-28 09:13

[History](#)
[Recent Versions](#)

A page in the site. Can contain rich text.

Title ■

Biopsy (old document)

Description

A short summary of the content

no longer relevant, refer to New Biopsy paper to be voted on in section 'Survey'

Body Text ■

Text Format HTML

Biopsy

Who should undergo diagnostic biopsies?

Small bowel biopsy has so far been considered as the gold or reference standard for the diagnosis of CD. According to the ESPGHAN 1990 definition, CD is in fact a "gluten-dependent enteropathy". Since then a wider spectrum of histological lesions (not only villous atrophy) has been related to CD, HLA typing is increasingly been used mainly to rule out CD and, finally, evidence has accumulated on the diagnostic value of CD-related antibodies. It could then be possible to switch from a situation where the biopsy is the only criterion to a new one where clinical presentation, serology, histology, family history and genetics, as well improvement on strict gluten free diet (GFD), all aid in establishing a diagnosis.

The pooled specificity for IgA anti-tissue transglutaminase antibodies (anti-tTG) is 93% (92-95); it is even higher for

Fig. 4 Handling text with embedded FCKEditor

VI) How to perform a biopsy?

Biopsy can be retrieved by upper endoscopy or in special circumstances by Crosby capsule. Although jejunal biopsies obtained by means of Crosby capsule are usually of a better quality, the general consensus is that biopsies should be taken during upper endoscopy. Such a policy has several advantages (shorter procedure time, absence of radiation, possibility of inspecting the mucosa, more fragments obtained to overcome the possibility of focal lesions). Biopsies (at least four) should be taken from the first (the bulb) and second or third portion of duodenum. Capsule endoscopy performance is not close enough to duodenal biopsies and could be considered only in patients unable or unwilling to undergo upper endoscopy

Please make your decision!

☐ agree
☐ disagree
☐ abstention from voting

Comment:

Editing status:

Remark: The question is completely answered, if one of the answer options is selected.

Fig. 5 Typical question layout

VI) How to perform a biopsy?

Biopsy can be retrieved by upper endoscopy or in special circumstances by Crosby capsule. Although jejunal biopsies obtained by means of Crosby capsule are usually of a better quality, the general consensus is that biopsies should be taken during upper endoscopy. Such a policy has several advantages (shorter procedure time, absence of radiation, possibility of inspecting the mucosa, more fragments obtained to overcome the possibility of focal lesions). Biopsies (at least four) should be taken from the first (the bulb) and second or third portion of duodenum. Capsule endoscopy performance is not close enough to duodenal biopsies and could be considered only in patients unable or unwilling to undergo upper endoscopy

Please make your decision!

Total number of votes:

13

Distribution of answers:

agree:	9 of 13	69.23 %
disagree:	4 of 13	30.77 %
abstention from voting:	0 of 13	0.00 %

Comments:

- for answer 'agree'
 - The last sentence is not clear to me. What does it mean "capsule endoscopy performance"?
 - Watson or Crosby capsule. According to my knowledge Crosby capsules are not any more manufactured, but Watson capsule is still available.
 - Biopsies (at least four) should be taken from the first (the bulb) and second or third portion of duodenum. Histological lesion in at least one specimen is sufficient for Dx, but normal histology in all 4 biopsy specimens is needed to exclude the Dx .
 - The wording may be changed: capsule endoscopy does not have the diagnostic quality...
- for answer 'disagree'
 - Videocapsule should be separated or in the comment only
 - Biopsies should be taken from both the first (at least one) and the third/fourth portion of the duodenum (at least two). I agree with the other statements.
 - I do not agree with "Capsule endoscopy could be considered only in patients unable or unwilling to undergo upper endoscopy"
 - I think that "the special circumstances" should be specified, for example as "or if not possible, by Crosby capsule". Now the text seems a little obscure to me.

Fig. 6 Evaluation report for question from ► Figure 5

References

1. Council of Europe. Developing a methodology for drawing up guidelines on best medical practices. Recommendation (2001) 13. 2002.
2. Harbour RT, editor. SIGN 50 – a guideline developer's handbook. Scottish Intercollegiate Guidelines Network. 2008.
3. German Association of the Scientific Medical Associations. Classification of S3 guidelines. German. [cited 2010 September 27]. Available from: <http://www.uni-duesseldorf.de/AWMF/II/klass-s3.htm>.
4. AGREE Collaboration. Appraisal of Guidelines Research and Evaluation (AGREE) instrument [cited 2010 September 27]. Available from: <http://www.agreecollaboration.org/instrument/>.
5. Agency for Quality in Medicine. German Instrument for Methodological Guideline Appraisal (DELBI) [cited 2010 September 27]. Available from: <http://www.leitlinien.de/leitlinienmethodik/leitlinienbewertung/delbi/delbi-english-version>.
6. Grol R. Has guideline development gone astray? BMJ 2010; 340: c306.
7. Eccles M, Mason J: How to develop cost-conscious guidelines. Health Tech Assess 2001;5(16):1-69.
8. cpg-development.com [Internet]. Berlin: Charité; c2008- [cited 2010 September 27]. Available from: <http://www.cpg-development.com/>.
9. Plone.org [Internet]. Plone foundation c2000- [cited 2010 September 27]. Available from: <http://plone.org/>.
10. Hohne W, Eckhold R, Hoffmann JC. [Guideline development and XML-based rapid prototyping]. German. In: Steyer G, Tolxdorff T, editors. Telemed 2005: bit for bit - Halbzeit auf dem Weg zur Telematik-infrastruktur. Berlin: Akademische Verlagsgesellschaft AKA; 2005. p. 284-93.
11. Hoffmann JC, Preiß JC, Autschbach F, Buhr HJ, Hauser W, Herrling K, et al. S3-Leitlinie „Diagnostik und Therapie des Morbus Crohn“ - Ergebnisse einer Evidenz-basierten Konsensuskonferenz der Deutschen Gesellschaft für Verdauungs- und Stoffwechselkrankheiten zusammen mit dem Kompetenz-netz Chronisch entzündliche Darmerkrankungen. German. Z Gastroenterol. 2008; 46(9): 1094-1146.
12. Fischbach W, Malfertheiner P, Hoffmann JC, Bolten W, Bornschein J, Gotze O, et al. S3-Guideline "Helicobacter pylori and gastroduodenal ulcer disease" of the German Society for Digestive and Metabolic Diseases (DGVS) in cooperation with the German Society for Hygiene and Microbiology, Society for Pediatric Gastroenterology and Nutrition e. V., German Society for Rheumatology, AWMF-Registration-no. 021 / 001. Z Gastroenterol. 2009; 47(12): 1230-1263.
13. Hohne W, Karge T, Schutt A. CPG development – Clinical Guidelines: TMF Portal for Web Based Development [Internet]. Berlin: TMF e.V.; 2010 [cited 2010 September 27]. Available from: <http://www.cpg-development.com/hilfe/information-brochure-about-the-cpg-development-portal>.