Health Related Quality of Life in Functional Gastrointestinal Disorders in Asia

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The importance of health-related quality of life (HRQOL) continues to grow, as clinicians and clinical researchers have recognized the impact of the functional gastrointestinal disorders. Limited information is available on the performance of HRQOL questionnaires in Asia. Furthermore, the effect across different cultural settings of functional gastrointestinal disorders on HRQOL has been little studied in Eastern countries. We summarized recent studies on HRQOL in Korean patients with functional gastrointestinal disorders as well as other Asian literatures. Functional gastrointestinal disorders-related symptoms had a great effect on the HRQOL of Korean patients. These results and their considerable prevalence in Korea indicate that functional gastrointestinal disorders have a substantial social impact in this country.

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Key Words
Dyspepsia; Irritable bowel syndrome; Quality of life

Introduction

Functional gastrointestinal disorders (FGIDs) are chronic conditions that are considered to have no structural or biochemical abnormalities that account for the symptoms.¹ FGIDs are not life threatening conditions, however, they significantly could impair the quality of life (QOL). FGIDs are important to public health because they are remarkably common, can hinder daily activities and induce a major social and economic burden. Thus, the impact of FGIDs on population health is probably huge. Despite its potential health relevance, the impact of FGIDs has received relatively little attention compared to other life threatening diseases in Asia. Furthermore, the effect across different cultural settings of FGIDs on health-related quality of life (HRQOL) has been little studied in Asian countries. The majority of published studies on FGIDs are ethnocentric and mostly reported from Western, Caucasian populations. Recently, more attention was given to the health economic implication of FGIDs in Asia along with the progress of economy. Here, we summarized recent studies on HRQOL in Asian patients with FGIDs.
Definition and Assessment of Health-Related Quality of Life in Asian Literature

World Health Organization defined health as condition that has the absence of disease and infirmity but also has the presence of physical, mental, and social well-being. QOL can be defined as the functional effect of an illness and its consequent therapy upon a patient. It describes health status from the patients' perspective. In this context, when we are managing patients, we must care not only about objective findings but also about patient's perception. HRQOL is made up of subject perceptions and objective health status which comprised of physical, psychological and social domain. The assessment of HRQOL continues to grow in importance, as clinicians and clinical researchers have recognized the impact of the FGIDs.

Both generic and disease-specific QOL instruments are to assess HRQOL. Generic instruments use generalized questions that are not specific to any particular diseases. An example of commonly used generic questionnaires in Asian studies is the Short Form 36 Health Survey (SF-36). It has 36 scaled items about health. Items are clustered in sub-scores (domain), such as "vitality" and "mental health". Lower scores indicate poorer QOL, with a score of 100 indicating perfect health status. There are published normative data for the general population of several Asian countries, facilitating comparisons between the subjects with disease and healthy controls. Generic instruments are used in general populations to assess a variety of health status, conditions, and diseases. They also permit comparison among diseases and may even detect unanticipated effects. However, they are time consuming and complicated with multiple questions. Generic instruments are not specific tools to assess any particular disease status or susceptible population of patients. Thus, they are insensitive to change with time or treatment. Disease-specific questionnaires are designed to detect HRQOL changes that may not be targeted in generic instruments, but are important manifestations of a certain disorder. Disease-specific instruments may not be widely applied to other diseases as these symptoms may not appear in other disorders. The advantage over the generic instruments is that they reflect problems most important to a specific population. They can detect smaller and more specifically relevant changes with time or treatment in health status, which may be overlooked by generic instruments. Disease-specific instruments are also too complex to administer and score. They are not capable of discriminating among different medical conditions.

Examples of commonly used disease-specific questionnaires in FGIDS in Asian studies are the IBS-QOL and the Nepean Dyspepsia Index (NDI). The IBS-QOL was developed by Patrick et al., which was comprised of 34 irritable bowel syndrome (IBS) specific QOL items with 8 Domains: dysphoria, interference with activity, body image, health worry, food avoidance, social relations, sexual relations and relationships. Each item was rated using a 5-point Likert response scale. Those scales are added up and transformed to 0-100 scale, indicating 0 as the poorest QOL and 100 as the maximum QOL. A patient responding “not at all” for all 34 items would score 100 in IBS-QOL. If responded as “slightly”, for all items, the overall IBS-QOL score would be 80. NDI is a dyspepsia-specific QOL instrument developed by Talley et al. This is a measuring instrument of symptom status and QOL in functional dyspepsia. A 15-item symptom checklist measures the frequency, severity, and bothersomeness of each upper gastrointestinal symptom on Likert scales. A total score for each symptom is created by adding 3 scales. A total dyspepsia HRQOL score is derived from the 25-item NDI, which is comprised of 5 sub-domains; interference, knowledge/control, eat/drink, sleep disturbance and work/study.

Humans interpret health events in a manner that is partly dependent on their social and cultural background. Thus, sociocultural validation of a questionnaire is needed to see its relevance to sociocultural influence. In most Asian studies, translation of the original questionnaires was done appropriately according to the accepted linguistic validation guidelines, including forward translation, backward translation and patient testing before their use. In addition to the linguistic validation, cross-cultural validation was performed in some studies.

We performed a systematic review of the literature to compare the HRQOL of patients with FGIDs with either healthy controls or those with other diseases. Full-length published papers from Asia during 1980-2010 were included if (1) patients had FGIDs and (2) HRQOL was measured using validated generic or dyspepsia-specific instruments, but not global assessment alone.

Chronic Gastrointestinal Symptoms and Their Effect on Health-Related Quality of Life in Community

Only a few studies were conducted on HRQOL in people in the community who did not visit the clinic (Table 1). One study from Korea has conducted a cross-sectional survey of gastro-
Table 1. Summary of the 8 Studies on Quality of Life in Korean Patients With Irritable Bowel Syndrome

| Author          | Population                | Subjects N, (female %, age) | Definition of IBS | QOL and other instrument | Results regarding QOL                                                                 |
|-----------------|---------------------------|----------------------------|-------------------|--------------------------|--------------------------------------------------------------------------------------|
| Lee and Chang24 | Clinic patients and public advertisement | 47 consultants, 32 non-consulters, (59.6%, 37.8) | Rome II | WHO-QOL BREF, SCL90R | Lower scores in all domains. Lower somatization and anxiety in IBS subjects and lower interpersonal sensitivity in consulters in SCL90R. |
| Park and Shin23  | 1,456 college students    | 374 (100%, 20.5)           | Rome II | IBS-QOL, GARS, Diet questionnaire | The lowest score in physical function, and the diarrhea-type ranked energy-related QOL the lowest. IBS subjects scored higher stress. No differences in eating habits.  |
| Shin and Ha22    | 973 nurses                | 254 (100%, 25.6)           | Rome II | SF-36          | Poorer QOL in all domains except physical functioning. IBS subjects visit doctors, take diagnostic tests and medication more frequently. |
| Hong25           | 587 nurses                | 208 (99%, 28.4)            | Rome II | SF-36, SCL90R  | Poorer QOL in all domains except physical functioning.                               |
| Lim26            | 1,000 community people    | 157 (61.5%, 36.7)          | Rome III | SF-36, SCL90, GARS | Poorer QOL in all domains. No difference among subtypes. IBS subjects visit doctors, take diagnostic tests and medication more frequently. Stress more common in IBS subjects. |
| Lee27            | 557 college students      | 157 (61.1%, 23.1)          | Rome II | WHO-QOL BREF health related questionnaire | The score of overall QOL and physical health domain were lower in IBS. IBS may be related to reported food intolerance, history of food poisoning, regular use of medication, extra-colonic symptoms, dietary habits, alterations in bowel habits during menses. |
| Jeong et al16    | Random sample of 1,427 residents in Asan-si | 31 (55.4%) | Rome II | The SF-36 | Poorer QOL on all domains vs. subject not having chronic gastrointestinal symptoms. |
| Park et al7      | Patients at 12 research institutions | 932 (55.3%, 43.0) | Rome II | The F-36, the IBS-QOL | Poorer QOL in all domains. Lower QOL scores in female, in severe symptoms, and in high level of education. |

Developed by Korean Dietetic Association; Health related questionnaire recorded a subject's dietary habits, medical history extra-colonic symptoms. IBS, irritable bowel syndrome; QOL, quality of life; D, diarrhea; C, constipation; A, alternating; U, unspecified; M, mixed; WHO-QOL BREF, Korean version of the World Health Organization Quality of Life Measure; SCL90R, symptom checklist-revised; GARS, Korean version of the Global Assessment of Recent Stress Scale; SF-36, Short Form 36 Health Survey.
Functional Dyspepsia

Functional dyspepsia is believed to be common in Asia as well as in Western countries. In Korea, it has been estimated that approximately 12% of adult population have dyspepsia, and a large number of patients with organic causes of dyspepsia were found in the survey. The management of dyspepsia is a major issue in clinical practice. Functional dyspepsia is the most common FGID in both primary and secondary care in both male and female population. The study also found that the prevalence of functional dyspepsia was higher in women than in men.

Table 2. Summary of the 6 Studies on Quality of Life in Asian Patients With Irritable Bowel Syndrome

| Author          | Population                                      | Subjects (n [female %, age]) | Definition of IBS | QOL and other instrument | Results regarding QOL                                                                 |
|-----------------|-------------------------------------------------|------------------------------|-------------------|--------------------------|--------------------------------------------------------------------------------------|
| Gwee et al28    | Random sample of 3,000 households in Singapore | 196 (54.6%, 40.5)           | Rome II           | SF-20, Spielberger       | IBS had poor HRQOL compared to healthy controls; IBS subjects had higher anxiety scores, and prone to abdominal surgery and to bladder symptoms. |
| Kanazawa et al29| Annual health check-up and 56 patients          | 46 (54.3%, 29), 56 (51.8%, 33) | SF-36, GSRS, STAI, SDS, PSS | Poorer QOL on all domains except physical functioning; Non-consulters had better QOL than consulters in the areas of role physical, body pain, general health, and social functioning. |
| Si et al30      | Hospital patients                               | 662 (52.9%, 44.8)           | Rome II           | SF-36                    | HRQOL was significantly impaired among the subjects with dyspepsia compared to healthy controls; Lower scores on all domains except physical functioning. No difference among subtypes. |
| Tung et al31    | Hospital patients                               | 99 (50.5%, 46.8)            | Rome III          | WHOQOL-BREF, HADS        | Psychiatric morbidity was frequent in IBS subjects and also linked to poorer QOL; The HADS depression and anxiety scores have negative correlations with QOL scores. |
| Lau et al32     | Face-to-face interviews on 1,288 residents in Shatin | 41 (65.8%)                  | Rome II           | SF-36                    | IBS had lower vitality scores on the SF-36 scale; Female IBS had lower mental health scores. 5% of men and 10% of women consulted a doctor last 12 months. 7.7% had > 1 days off work due to IBS last 12 months. |
| Xiong et al33   | Face-to-face interview on 4,178 random sample in South China | 239 (59.8%), Subtype D:C:M (74:15:11) | Rome II           | SF-36                    | IBS was associated with a decrement in health-related QOL score. Poorer QOL on all domains. |

IBS, irritable bowel syndrome; QOL, quality of life; D, diarrhea; C, constipation; M, mixed; U, unspecified; SF-20, Short Form 20 Health Survey; STAI, State-Trait Anxiety Inventory; SF-36, Short Form 36 Health Survey; GSRS, Gastrointestinal Symptoms Rating Scale; SDS, Self-Rating Depression Scale; PSS, Perceived Stress Scale; WHOQOL-BREF, Hong Kong Chinese version of the World Health Organization Quality of Life Measure; HADS, Hospital Anxiety and Depression Scale.
pepsia questionnaire and the SF-36. Endoscopic examination revealed the organic causes of dyspepsia in 28% of dyspeptic patients; gastric ulcer in 5.3%, duodenal ulcer in 9.3%, gastric cancer in 1.4% and reflux esophagitis in 2.6%. Patients with functional dyspepsia by Rome II criteria had lower HRQOL score on all 8 domains of the SF-36 compared to general population. Both mental and physical scores were equally lower than healthy subjects. Patients with functional dyspepsia reported poorer QOL in 4 domains including general health, vitality, role emotional and mental health than the patients with peptic ulcer. Dyspeptic patients who are seen in a referral setting had worse HRQOL scores than primary care patients and non-consulters who were detected in the general population.

Cho et al.6 evaluated QOL using NDI. Validation study of Korean version of NDI revealed an excellent test-retest reproducibility and internal consistency which was expressed by Cronbach’s alpha, 0.92. Construct was validated by comparison with SF-36, global rating of QOL, and anxiety and intrusiveness. The scores of all subscales were significantly low in dyspeptic patients demonstrating that the NDI is able to discriminate between the 2 groups. The greatest impairment was seen in the knowledge/control domain of all subscales, and the lowest impairment was seen in the sleep disturbance subscore. Responsiveness of the NDI instrument was performed as part of the cross-cultural validation, which took place in 3 centers in an open trial of mosapride.6 One hundred and 29 patients with functional dyspepsia by Rome II participated in this study and completed the SF-36 and the NDI at baseline and the NDI after mosapride treatment. All dyspeptic symptoms were significantly decreased after mosapride treatment. Whereas dyspepsia score decreased after medication, NDI score increased after treatment. In other words, patient’s symptoms improved as well as their QOL. An increase of NDI score after mosapride treatment, which meant a better QOL, was observed in summary scale as well as in all 5 subscales. The greatest improvement was observed in the knowledge/control domain among 5 subscales which was mostly impaired prior to treatment. This could explain the nature of improvement after drug treatment. The subjects were compared as responders and non-responders to mosapride treatment for evaluating the degree to which the NDI questionnaires and their subscales improved with treatment. All NDI subscales showed 154%-302% greater improvement for responders over non-responders (P < 0.001 for all subscales). This finding attests to the fact that the NDI is sensitive enough to detect changes with treatment. In this study the NDI was shown to be highly responsive to treatment effects thereby leading to its recommendation to be used in treatment trials. However, further studies are needed in the large placebo controlled trials.

Irritable Bowel Syndrome

The reported prevalence of IBS in Asian countries varies from 5 to 10% depending on the study population and diagnostic criteria (Table 2).19,20 Asian doctors and medical professionals are beginning to notice the burden of IBS along with increasing prevalence of the organic colonic diseases in Asian countries. A Taiwanese survey assessed the social and sleep quality impact of IBS subjects receiving physical check-up.21 36% of IBS subjects visited physicians due to gastrointestinal symptoms and the mean number of visits was 2.3 during the previous year. The absenteeism from work or school was marked in the IBS subjects, whether due to any illness or any particular gastrointestinal symptoms. An almost 2-fold increase in the frequency of sleep disturbance was noted in subjects with IBS symptoms.21 Studies regarding QOL in Korean and Asian patients with IBS were summarized in Tables 17,16,22-27 and 2.28-33 Most studies consistently showed poorer QOL in IBS subjects in both mental and physical summary of the SF 36, while some showed no difference in physical functioning domain. Poorer QOL in mental component can be explained by psychiatric comorbidity in IBS subjects. Even though the social and cultural environment in which study was carried out differ from that in the West, the results are in agreement with those studies conducted in the West. The QOL of IBS patients was significantly impaired in all dimensions of generic HRQOL compared with the general population or for healthy subjects. Interestingly, the score for social functioning in the SF-36 and that for the sexual and relationship domain of the IBS-QOL were somewhat higher than those of Western studies.24,36 suggesting that IBS has little impact on social function in Korean and Chinese patients.7,37 These responses may be related to the Eastern cultural milieu. Such aspects should be elaborated further by studies across the cultures. Data from nationwide cross-sectional survey in Korea7 showed functional limitations to be noted in all 8 subscales of the IBS-QOL with the mean overall score at 74.2. The lowest scores were noted on the health concern and food avoidance domains which is consistent with the West. Therefore, health was the greatest concern among IBS patients. It indicates that patients experience emotional dysphoria, ie, feeling helpless, out of control, depressed and irritable. It is noteworthy that the scores for the sexual relationship domains were
greater than 80, which meant that most responses were “not at all” or “slightly”, indicating sexual activity was not interfered by bowel symptoms. It is contrary to the findings of the West. The high scores in the Sexual domain may mean that IBS marginally affects the sexual life in some Korean patients. Asians are raised and educated in a Confucian society in which both males and females avoid discussing sexual concerns.

Demographic and social factors had an effect on the HRQOL of patients with IBS. The IBS-QOL was significantly associated with female gender, symptom severity, and low level of education in Korean study. Female patients with IBS had a lower HRQOL than males, although the difference was negligible to discriminate the gender difference in HRQOL. Significant impairment of HRQOL was only observed in patients with severe symptoms, whereas patients with mild and moderate symptoms showed only mild impairment. Patients who experienced mild symptoms scored over 80 for all domains except health worry. By contrast, the overall score for IBS-QOL was less than 65 in the severe symptom group. These findings support the concept that symptom severity is an important determinant of QOL and IBS-specific QOL questionnaire has strong power to discriminate. Thus, the measurement of a patient’s HRQOL could be an important element of clinical research and patient care. In a retrospective review of HRQOL in chronic disease, patients who were older, female, less educated, not living with a partner, and at least with 1 co-morbid condition reported the poorer level of QOL. Subjects with psychiatric morbidity also had a poorer QOL, thereby screening for psychiatric disorders in gastroenterology clinics is worthwhile.

There were no differences in HRQOL between subsets of IBS, which is consistent with previous studies in which the HRQOL of subgroups of IBS was evaluated using standardized Rome criteria. Responsiveness of the IBS-QOL was validated in an open trial of tegaserod in Korean patients. All subscales of the IBS-QOL showed greater improvement for responders over non-responders with the greatest changes in the dysphoria, interference with activities, body image, health worry and food avoidance subscales. These findings suggest that disease-specific HRQOL could more accurately reflect the disease status of IBS and the response to treatment. The IBS-QOL questionnaire could be employed as an important outcome variable in the clinical trials of FGIDs. When looking at the subscale responses to treatments, it is clear that the psychosocial subscale scores are more responsive in explaining the nature of the improvement. For the IBS-QOL, the subscales of dysphoria, interference with activity and food avoidance improved more. It is perhaps not surprising that these were the scales that were more impaired prior to treatment.

**Conclusion**

We extensively reviewed the Asian studies that have measured HRQOL in FGIDs. We have confirmed that Asians with FGIDs showed decreased HRQOL. However, we do not have sufficient Asian data at the moment and most of them are on cross-sectional epidemiologic study. We need to know what factors would lead to patients feeling poorer QOL in Asian region. Further studies are needed to evaluate the factors influencing HRQOL, for instance, non-intestinal symptoms or psychological comorbidity. This will help not only to obtain specific patient’s needs but also to provide information for making guidelines on FGIDs management.

FGIDs are common in Asia and healthcare utilization is very high. Despite this, FGIDs which are defined under Rome criteria are not being widely applied in clinical practice in Asia. Many primary care doctors do not fully comprehend the concept of functional diseases. In addition, lack of understanding FGIDs by patients might affect mental components of QOL. This could be solved through educating general practitioners as well as providing adequate information about FGIDs to general public. We know that the impact of FGIDs on population health is probably immense in Asia, however, it is still under-recognized. In light of this reality, we should place extra focus on clinical research on FGIDs.

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