Quality of Life and Treatment Satisfaction with Pharmacological Interventions in Chinese Adults with Chronic Pain Due to Osteoarthritis

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Research article

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Abstract

**Background:** Aim of this multicenter, observational, cross-sectional study was to evaluate health-related quality of life (HRQoL) and treatment satisfaction of current medications in Chinese knee OA patients.

**Methods:** Brief Pain Inventory (BPI), Treatment Satisfaction Questionnaire (TSQM-1.4), and HRQoL (EQ-5D-5L) were assessed in total of 601 OA of knee patients.

**Results:** Mean score of EQ-5D-5L of patients with BPI-Severity $\geq 4$ was significantly lower than those with BPI-Severity $<4$. All the scores of TSQM in 4 dimensions were lower in patients with BPI-Severity $\geq 4$ than in those with BPI-Severity $<4$. Both HRQoL scores and TSQM scores showed a statistically significant decreasing trend with increasing BPI-Severity pain score.

**Conclusion:** Chronic knee OA pain has a significant impact on patients’ HRQoL. More severe patients with OA were less satisfied with current treatments.

**Trial registration (Clinical Trials identifier):** Not applicable

**Background**

Osteoarthritis (OA) is one of the most prevalent chronic musculoskeletal disorders and a leading cause of disability worldwide, especially among the elderly [1]. Globally, the prevalence of knee OA in men is lower compared to women, with 9.6% of men and 18% of women aged over 60 years affected [1, 2]. In China, the prevalence of radiographic OA was 42.8% in women and 21.5% in men; whereas, symptomatic OA occurred in 15% of women and 5.6% of men. The prevalence of radiographic and symptomatic OA in Chinese men was similar to that in white men in the United States (US). However, Chinese women had a higher prevalence of radiographic and symptomatic OA than women in the US [2, 3].

Chronic pain is one of the most common health issues that exerts a significant social and financial burden on the individual and society. Patients with inadequate pain relief are more likely to have worse quality of life (QoL), greater function loss, and greater pain interference with daily activities [4]. OA is a leading cause of deteriorated QoL due to chronic pain [5, 6]. Compared with the radiographic OA without pain, painful OA has been associated with higher cardiovascular risk and mortality [7]. Pain is recognized as one of the hallmark symptoms in OA and is a common reason patients seek medical attention. Mechanisms underlying chronic pain include a complex interaction of physiological, emotional, cognitive, social, and environmental factors [8]. When considering the complex nature of chronic pain, treatment often necessitates the use of a blend of different approaches. In terms of nonsurgical standard interventions for OA, multimodal pain management is a comprehensive treatment of complex chronic pain syndromes that includes 4 core disciplines of multimodal pain management: pain medicine, psychotherapy, exercise therapy (including physiotherapy), and assistant medical professions including nurses. Multimodal pain management protocols aim to address pain control, facilitate functional recovery, and maintain patient satisfaction [9, 10].
Patient-reported outcome is an important consideration in the treatment of patients with OA. All aspects of QoL are compromised when pain is inadequately treated, and effective pain relief has been shown to improve health-related quality of life (HRQoL) [11, 12]. When patients with OA were asked to rank aspects of QoL impacted by their condition, they highlighted enjoyment of life, emotional well-being, fatigue, weakness, and sleep-related problems as the most important areas they would consider when evaluating the success of their pain treatment [13]. The pain caused by OA can have a substantial impact on patients’ QoL [11–13]. In a 2012 online survey of patients with OA in the United Kingdom [14], 52% of the 2001 respondents reported that OA had a large impact on their life, 71% reported having persistent pain even after taking their prescribed pain medication, and 12% said their pain was often unbearable. In a cross-sectional study conducted in 2014 by Kantar Health, only 14% of patients in Japan with diagnosed pain who suffered from joint pain were highly satisfied with their pain medications [4]. Furthermore, a multinational longitudinal survey showed that patients with inadequate pain relief were more likely to have a worse QoL, greater function loss, and greater pain interference [4].

Patient satisfaction is an important indicator of the quality of care provided to patients with OA [15]. Patient-reported outcomes, such as HRQoL and patient satisfaction, were used to capture patients’ experience of chronic disease and can support the physician in clinical practice to facilitate patient-centered care [16]. Thus, QoL and treatment satisfaction assessments are crucial to evaluating the clinical effectiveness of treatment in OA.

Little is known about the impact of chronic knee OA pain on HRQoL and treatment satisfaction in a real-world setting in China. Therefore, the cross-sectional survey presented in this article has been designed to understand the impact of chronic knee OA pain on HRQoL and to evaluate treatment satisfaction of current medications among Chinese patients with knee OA.

**Methods**

**Study design and subjects**

This site-based, multicenter, observational, cross-sectional study in China enrolled 601 outpatients with knee OA from 2 orthopedics, 2 rheumatology, and 1 pain department in 5 tertiary hospitals from March to October 2018. Written informed consent was obtained from each patient before they participated in any study-related procedures.

Chinese adult patients (aged ≥ 40 years) with diagnosed knee OA experiencing chronic pain for at least 3 months and receiving oral medications during the past 12 months were eligible for the study. Patients with rheumatoid arthritis or other inflammatory arthritis; knee pain caused by other diseases (eg, traumatic fracture history or tumor); mental illness, including cognitive disorders such as Alzheimer’s disease, schizophrenia; and bedridden patients who were undergoing knee replacement surgery were excluded. Patients with pain level higher than knee pain due to cancer or other reasons such, as gout and chondrocalcinosis, were also excluded. Socio-demographics, disease characteristics, Brief Pain Inventory (BPI), treatment information, and patient responses to HRQoL (5-level of Chinese Quality of Life-5
Dimensions version [EQ-5D-5L)] and Treatment Satisfaction Questionnaire for Medication (TSQM-1.4) interviews were also assessed.

**Measures**

**Patient characteristics**

The characteristics measured were age, sex, body mass index, ethnicity, employment status, education status, insurance status, and comorbidity. The following OA characteristics were measured for each enrolled patient: age and location at first diagnosis, current department of visits, number and location of painful sites, and severity of pain. The average number of weekly days of paid work or housework lost due to OA was also recorded. In addition, information related to the current treatment for OA pain management (including non-pharmacotherapy) was collected from each enrolled patient.

**Outcome measures**

The BPI is a validated self-reported questionnaire that assesses pain severity using the Numerical Rating Scale for Pain Intensity (NRS-PI, 0 to 10 scale, where 0 = no pain and 10 = worst possible pain) for the conditions of worst, least, and average pain, as well as “pain right now”. The 5-level Chinese Quality of Life-5 Dimensions version (EQ-5D-5L) [17] comprises 5 dimensions: mobility, self-care, usual activities, pain/discomfort, and anxiety/depression. Each dimension has 5 levels: no problems, slight problems, moderate problems, severe problems, and extreme problems. Self-health care assessment was performed using the EuroQol (EQ) visual analogue scale (EQ VAS). The EQ VAS self-rating records the respondent’s own assessment of their health status on a 20-cm vertical VAS with endpoints labelled ‘the best health you can imagine’ and ‘the worst health you can imagine.’ [18] The TSQM was designed to assess treatment satisfaction for patients with chronic diseases. The TSQM 1.4 is a 14-item psychometrically robust and validated instrument consisting of 4 scales: effectiveness, side effects, convenience, and global satisfaction, each on a scale of 0–100 with higher scores indicating a higher level of satisfaction.

**Statistical analyses**

Demographic and clinical characteristics were assessed using frequencies and percentages for categorical variables and mean values and standard deviations (SDs) for continuous variables (descriptive analysis) in the whole patient population. Impact on QoL (EQ-5D-5L) and treatment satisfaction (TSQM-1.4) by BPI-Severity score (< 4 and \(\geq 4\)) were presented using mean (SD) and were compared using a t-test. For each of self-assessed health, EQ-5D-5L, and TSQM, a linear regression model was used to estimate the regression coefficient along with corresponding 95% confidence interval (CI) for BPI-Severity, adjusting for age (continuous), number of pain sites (continuous), and comorbidity (yes or no). Missing data were not analyzed. Statistical analyses were conducted using SAS 9.4 (SAS Institute, Cary, NC), and a 2-sided \(P\) value of 0.05 was considered statistically significant.

**Results**
A total of 601 patients met the eligibility criteria and completed this survey (Fig. 1). The mean (SD) age of enrolled patients was 61.77 (9.53) years and the majority of patients were female. More than 50% of patients had at least 1 comorbidity of gastrointestinal or cardiovascular disease (Table 1). The most commonly used current treatments for knee OA were oral medication, a patch or ointment, or intra-articular hyaluronic acid injection (Table 2). More than half of patients were rated with BPI-Severity ≥ 4. Pain interfered with work productivity, with 37.1% of patients self-reporting that more than 4 days/week of work or housework were lost due to OA pain.
Table 1
Patient demographics and baseline characteristics

| Characteristics                          | N = 601                       |
|-----------------------------------------|-------------------------------|
| **Age, mean (SD)**                      | 61.77 (9.53)                 |
| **Body mass index, mean (SD)**          | 24.66 (3.16)                 |
| **Gender, n (%)**                       |                               |
| Male                                    | 149 (24.79)                  |
| Female                                  | 452 (75.21)                  |
| **Nationalities, n (%)**                |                               |
| Han                                     | 587 (97.67)                  |
| Others                                  | 14 (2.33)                    |
| **Working status, n (%)**               |                               |
| Unemployed                              | 37 (6.17)                    |
| Part-time                               | 7 (1.17)                     |
| Full-time                               | 144 (24)                     |
| Retired                                 | 412 (68.67)                  |
| **Educational status, n (%)**           |                               |
| Below senior high school                | 293 (48.75)                  |
| Senior high school                      | 137 (22.8)                   |
| Junior college                          | 88 (14.64)                   |
| Undergraduate                           | 76 (12.65)                   |
| Postgraduate or above                   | 7 (1.16)                     |
| **Insurance types, n (%)**              |                               |
| Urban resident basic medical insurance  | 177 (29.45)                  |
| Urban employee basic medical insurance  | 242 (40.27)                  |
| New rural cooperative medical system    | 158 (26.29)                  |
| Commercial health insurance             | 5 (0.83)                     |
| Uninsured                               | 19 (3.16)                    |
| **Comorbidity, n (%)**                  |                               |

n: number of subjects; SD: standard deviation.
| Characteristics                  | N = 601 |
|---------------------------------|---------|
| Any comorbidities               | 331 (55.07) |
| Hypertension                    | 239 (39.77) |
| Coronary heart disease          | 75 (12.48) |
| Myocardial infarction           | 2 (0.33) |
| Stroke                          | 12 (2) |
| Cerebral hemorrhage             | 1 (0.17) |
| Gastritis                       | 86 (14.31) |
| Nephropathy                     | 15 (2.5) |
| Diabetes                        | 92 (15.31) |
| Stomach or duodenal ulcers      | 20 (3.33) |

n: number of subjects; SD: standard deviation.
Table 2  
Clinical characteristics of knee osteoarthritis

| Characteristics                                      | N = 601                  |
|------------------------------------------------------|--------------------------|
| **Age at first diagnosis, mean (SD)**                 | 58.13 (9.62)             |
| **Location at first diagnosis, n (%)**                |                          |
| Unilateral knee                                      | 290 (48.33)              |
| Bilateral knee                                       | 278 (46.33)              |
| Others (shoulders, elbows, hips, etc.)               | 32 (5.33)                |
| **Current department, n (%)**                        |                          |
| Rheumatology                                         | 155 (25.79)              |
| Orthopedics                                          | 326 (54.24)              |
| Pain                                                 | 120 (19.97)              |
| **Brief Pain Inventory score, mean (SD)**             |                          |
| Pain Severity (full score: 10)                       | 3.78 (1.62)              |
| Pain Interference (full score: 10)                   | 2.97 (1.70)              |
| **Treatment pattern, n (%)a**                        |                          |
| Oral medication (Western/traditional Chinese medicine)| 469 (78.04)              |
| Patch/ointment                                       | 271 (45.09)              |
| Intra-articular hyaluronic acid injection            | 189 (31.45)              |
| Intra-articular steroid injection                    | 125 (20.8)               |
| Physiotherapy (electrotherapy/hyperthermia)          | 88 (14.64)               |
| Kinesitherapy (rehabilitation treatment)             | 22 (3.66)                |
| Orthoses (cane, etc.)                                | 4 (0.67)                 |
| Others                                               | 30 (4.99)                |
| **Average weekly days of paid work or housework loss due to osteoarthritis, mean (SD) in the past month** |               |
| 0 day                                                | 235 (39.30)              |
| 1 day                                                | 44 (7.36)                |
| 2–3 days                                             | 97 (16.22)               |

n: number of subjects; SD: standard deviation.
The mean score of EQ-5D-5L of patients with BPI-Severity ≥ 4 was significantly lower than those with BPI-Severity < 4 (0.62 vs 0.84, \( P < 0.0001 \)) (Table 3). A similar trend was observed for self-assessed health outcomes, where the mean self-assessed health score of patients with BPI-Severity ≥ 4 was significantly lower than those with BPI-Severity < 4 (66.88 vs 73.8, \( P < 0.0001 \)). There were statistically significant differences in all 4 domains of TSQM-1.4 between both patient subgroups (BPI-Severity ≥ 4 and < 4) (Table 4). The mean score of TSQM for the patients with BPI-Severity ≥ 4 was significantly lower than those with BPI-Severity < 4 for Effectiveness (51.0 vs 57.8, \( P < 0.0001 \)), Side Effects (94.9 vs 97.2, \( P = 0.0099 \)), Convenience (60.2 vs 64.7, \( P < 0.0001 \)), and Global Satisfaction (57.7 vs 60.4, \( P = 0.0402 \)). As shown by the TSQM score, treatment satisfaction was significantly lower in patients with BPI-Severity ≥ 4 than in those with BPI-Severity < 4.

The BPI-Pain Severity scores were inversely associated with the self-assessed health, EQ-5D-5L, and TSQM scores. In linear regression models adjusted for age, number of pain sites, and comorbidity, HRQoL scores (self-assessed health [-3.20; \( P < 0.0001 \]) and EQ-5D-5L [-0.08; \( P < 0.0001 \]) showed a significant decreasing trend with each unit increase in BPI-Severity pain score, indicating that reduction in knee pain was statistically significantly associated with improvements in HRQoL scores (Table 5). The score of TSQM also showed a significant decreasing trend in Effectiveness (-2.69, \( P < 0.0001 \)), Side Effects (-0.56, \( P = 0.0483 \)), Convenience (-1.40, \( P < 0.0001 \)), and Global Satisfaction (-1.16, \( P = 0.0038 \)) with each unit increase in BPI-Severity pain score (Table 6), indicating that lower knee pain was significantly associated with higher TSQM Effectiveness, Side Effects, Convenience, and Global Satisfaction scores.

### Table 5
The association between BPI and HRQoL

| EQ-5D-5L and self-assessed health per BPI | Parameter estimatea | 95% CI          | P value | Parameter estimateb | 95% CI          | P value |
|------------------------------------------|---------------------|-----------------|---------|---------------------|-----------------|---------|
| EQ-5D-5L                                 | -0.08               | (-0.09,-0.08)   | < 0.0001| -0.08               | (-0.09,-0.07)   | < 0.0001|
| Self-assessed health                     | -3.31               | (-4.03,-2.59)   | 0.0523  | -3.20               | (-3.92,-2.48)   | < 0.0001|

aAdjusted for age. bAdjusted for age, number of pain site, comorbidity. BPI: Brief Pain Inventory; CI: confidence interval; EQ-5D-5L: EQ 5 dimension-5-level; HRQoL: Health-Related Quality of Life.
Table 6
The association between BPI and TSQM

| TSQM per BPI          | Parameter estimatea | 95% CI       | P value | Parameter estimateb | 95% CI       | P value |
|-----------------------|---------------------|--------------|---------|---------------------|--------------|---------|
| TSQM- Effectiveness   | -2.64               | (-3.33,-1.94)| < 0.0001| -2.69               | (-3.38,-1.99)| < 0.0001|
| TSQM- Side effect     | -0.59               | (-1.14,-0.03)| 0.0381  | -0.56               | (-1.12,0.00)| 0.0483  |
| TSQM- Convenience     | -1.42               | (-1.94,-0.90)| < 0.0001| -1.40               | (-1.93,-0.88)| < 0.0001|
| TSQM- Global satisfaction | -1.08             | (-1.86,-0.29)| 0.0073  | -1.16               | (-1.94,-0.38)| 0.0038  |

aAdjusted for age. bAdjusted for age, number of pain site, comorbidity. BPI: Brief Pain Inventory; CI: confidence interval; TSQM: Treatment Satisfaction Questionnaire for Medication.

Discussion

The cross-sectional survey presented in this article is the first large-scale, multicenter real-world study to explore the impact of OA pain on HRQoL and treatment satisfaction among Chinese patients with OA. The results show that chronic pain has not been well managed since 78% of Chinese patients with OA who were treated with pharmacological therapy combined with other therapies still experienced moderate-to-severe pain (BPI $\geq 4$) and significantly lower HRQoL and treatment satisfaction. Moreover, more than 35% of patients self-reported that they lost more than 4 days/week of work due to OA pain. These observations indicate that the patients with OA were not satisfied with current treatments. The cross-sectional survey results suggest that patients with moderate-to-severe OA pain had significantly lower HRQoL and treatment satisfaction scores as compared to patients with mild OA pain. Overall, pain severity plays an important role in predicting HRQoL and treatment satisfaction in Chinese patients with knee OA. Also, the study results suggest that increased pain severity is associated with a decrease in the levels of HRQoL and treatment satisfaction among Chinese patients with OA. Reduction in knee pain was statistically significantly associated with improvements in HRQoL and treatment satisfaction among Chinese patients with OA.

The analysis results suggest that pain severity plays an important role in predicting HRQoL, and our findings are consistent with the previous studies [19–21]. A published study demonstrated that patients experiencing OA pain in both knees have poorer HRQoL compared to patients with unilateral knee pain or no knee pain [22]. A population-based study in Japan revealed that patients with severe knee OA had significantly lower physical HRQoL than those with mild and moderate knee OA [23]. A large population-based cohort study from southern Sweden also confirmed that participants with knee OA (defined either clinically or radiographically) reported lower HRQoL scores than those with no knee OA [24]. The results of another study showed that patients with radiographic knee OA had considerably lower scores in all
subgroups of SF-36 compared with healthy controls [25]. The results obtained from a cross-sectional study revealed that the lower HRQoL scores were associated with increased pain severity in patients with knee OA [26].

Patient satisfaction with treatment is essential in OA and is a measure of therapeutic effectiveness [10, 27]. In this study, TSQM scores in 4 dimensions were significantly lower in patients with OA with moderate-to-severe pain intensity (BPI-Severity ≥ 4) than in those with mild OA pain intensity (BPI-Severity < 4). This indicates that treatment satisfaction was found to be higher in OA patients with lower pain, which is consistent with an earlier study showing that decreased pain was associated with increased treatment efficacy and, thereby, patient satisfaction [27]. Thus, switching treatments to achieve lower pain levels might enhance treatment satisfaction among patients with knee OA. Stahmer et al [28] reported that patient satisfaction with pain management is associated with the amount of pain relief achieved. Moreover, the findings regarding pain as an important factor in predicting treatment satisfaction may be extrapolated to patients with knee OA globally. In summary, pain severity has a great impact on HRQoL and treatment satisfaction in Chinese patients with knee OA. Pain relief may help improve patients’ HRQoL and treatment satisfaction. This real-world study provided the evidence that relieving pain should be the first choice of therapy for knee OA.

Our study does have some limitations. Since data were derived from a cross-sectional survey, the association between knee OA pain severity and HRQoL and treatment satisfaction cannot be viewed as causal. Longitudinal studies are needed to examine the relationship between knee OA pain severity and HRQoL and treatment satisfaction. Moreover, the study was conducted at 5 tertiary hospitals in China, and no randomization mechanism was used in their selection; hence, it is difficult to generalize the findings.

**Conclusions**

Chronic pain due to OA, especially in those patients with moderate-to-severe pain, has a significant impact on patients’ HRQoL. In our study, patients with more severe OA were less satisfied with current treatments. Appropriate pain management in China is important in improving HRQoL and the treatment satisfaction for medication.

**List Of Abbreviations**
Declarations

Ethics approval and consent to participate

The authors state that they have obtained appropriate institutional review board approval or have followed the principles outlined in the Declaration of Helsinki for all human investigations. In addition, for all subjects participating in this study, informed consent was obtained prior to any study procedures being performed.

The following is the summary of institutional review board approval obtained for this study project:

| Institute name                              | Ethics committee name                                      | Committee's reference number                        |
|---------------------------------------------|-----------------------------------------------------------|-----------------------------------------------------|
| Peking University People’s Hospital         | The Medical Ethics Committee of Peking University People’s Hospital | 2018PHB015-01; 2018PHB015-02                          |
| Beijing Hospital                            | The Ethics Committee of Beijing Hospital                   | 2017BJYYEC-158-01; 2017BJYYEC-158-02                 |
| First Affiliated Hospital of Harbin Medical University | The Ethics Committee of First Affiliated Hospital of Harbin Medical University | 201804; 2018XS09-02;                                 |
| Shanghai Sixth People’s Hospital            | The Ethics Committee of Shanghai Sixth People’s Hospital  | 2018-KY-002; 2018-KY-002- (K)-(1)                     |
| The Second Xiangya Hospital of Central South University | The Ethics Committee of the Second Xiangya Hospital of Central South University | 2018-01004; 2018K048;                                 |
Consent for publication

Not applicable.

Author contributions

QX, HL, JL, DD, JZ, JC, SL, YZ, YC, XM, ZZ: made substantial contributions to the conception or design of the work; or the acquisition, analysis, or interpretation of data for the work; AND drafted the work or revised it critically for important intellectual content; AND gave final approval of the version to be published; AND agreed to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

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Financial & competing interests disclosure

The authors declare that they have no competing interests. YZ, YC, and XM are employees of Eli Lilly and Company.

Availability of data and materials/Data sharing statement

Data will be shared upon request.

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**Figures**
Figure 1

Flow chart of survey sampling. n: number of subjects; OA: osteoarthritis.

Supplementary Files

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