The mediating role of psychological symptoms on falls risk among older adults with osteoarthritis

Abstract: The purpose of this study was to investigate the role of fear of falling (FoF) and psychological symptoms in explaining the relationship between osteoarthritis (OA) symptom severity and falls. Individuals aged ≥65 years with ≥2 falls or ≥1 injurious fall over the past 12 months were included in the falls group, while volunteers aged ≥65 years with no history of falls over 12 months were recruited as controls. The presence of lower extremity OA was determined radiologically and clinically. Severity of symptoms was assessed using the Western Ontario and McMaster Universities Arthritis Index (WOMAC) questionnaire. FoF and psychological status were measured with the shortened version of the Falls Efficacy Scale-International and the 21-item Depression, Anxiety and Stress Scale (DASS-21), respectively. Of 389 (229 fallers, 160 non-fallers) potential participants, mean (SD) age: 73.74 (6.60) years, 141 had clinical OA and 171 had radiological OA. Fallers with both radiological OA and clinical OA had significantly higher FoF and DASS-21 scores than non-fallers. FoF was positively correlated with symptom severity in fallers and non-fallers with radiological and clinical OA. Depression, anxiety, and stress scores were only significantly correlated with symptom severity among fallers but not non-fallers with radiological and clinical OA. The relationship between mild symptoms and reduced risk of falls compared to no symptoms in those with radiological OA was attenuated by increased anxiety. The increased falls risk associated with severe symptoms compared to mild symptoms in clinical OA was attenuated by FoF. FoF may, therefore, be a potentially modifiable risk factor for OA-associated falls which could be considered in future intervention studies.

Keywords: falls, fear of falling, osteoarthritis, psychological status, aged

Introduction

Osteoarthritis (OA) is the most prevalent chronic degenerative joint disease world-wide.}\(^1\) Aging is a major risk factor for OA with one in three individuals aged ≥75 years reported to have severe OA changes on their knee radiographs.}\(^2\) Its prevalence, alongside that of many chronic medical conditions, is expected to rise with the rapidly aging global population.

OA is associated with deficits in balance control.}\(^3\) Among individuals with severe OA, 50% have reported at least one fall in the preceding year.}\(^2\) However, one in three older adults aged ≥65 years also experience at least one fall annually regardless of OA status.}\(^4,5\) Existing published evidence on the relationship between OA and OA have used a variety of different methods for the identification of their OA subjects, including presence of knee pain, self-reported physical diagnosed OA, clinical OA based on agreed criteria, and radiological evidence of OA. These studies also often reported the relationship between OA and balance measures or falls risk rather than actual fall events.}\(^5-9\) Furthermore, a recent study revealed a biphasic relationship...
between OA and falls with older individuals with radiological evidence of OA with mild OA symptoms less likely to experience falls compared to those with radiological OA with no symptoms, while individuals with clinical OA and severe symptoms are more likely to experience falls than those with mild symptoms.10

Both falls and OA adversely affect the older individuals’ psychological well-being and quality of life (QoL).11,12 Strong negative associations were observed between depressive mood and fear of falling (FoF) with QoL in older fallers,13 while increased pain and functional limitations were found to decrease QoL in patients with OA.14

Psychological issues among fallers with OA have not been examined previously. We therefore conducted a case–control study comparing older fallers and non-fallers with knee OA, determined using both radiological and clinical criteria, to identify potential differences in FoF and psychological symptoms among fallers and non-fallers with OA.

**Methods**

**Study population**

This was a cross-sectional case–control study. Subjects aged ≥65 years with either a history of two falls or one injurious fall over the past 12 months were recruited from the Departments of Emergency Medicine, Primary Care, and Geriatric Medicine at a teaching hospital in greater Kuala Lumpur. Control participants with no falls (non-fallers) over the previous 12 months were recruited through media and word-of-mouth advertising. Participants were excluded if they had clinically diagnosed dementia (ICD-10 definition), severe physical disabilities, or major psychiatric illnesses. Demographic details were obtained from all participants. This study was approved by the University of Malaya Medical Centre Medical Ethics Committee (925.4) and complied with the Helsinki Declaration of 1975, revised in 1983. Written informed consent was obtained from all study participants.

**Diagnosis of OA**

All participants were assessed for the presence of knee OA using two different definitions of OA, radiological OA and clinical diagnosis of OA, based on predefined criteria. The influence of OA according to each definition on falls was evaluated separately.

**Radiographic diagnosis of OA**

Standardized knee radiographs were obtained in the standing antero-posterior view in all consenting participants. These images were assessed by a radiologist blinded of clinical data. The severity of OA was determined by the Kellgren–Lawrence (KL) grading scale,14 with a score of 0 indicating no knee OA and the maximal score of 4 indicating severe OA. Subjects with KL grades of 2–4 were considered subjects with radiological OA. Participants without radiological OA were excluded in subsequent analysis involving those with radiological OA only.

**Clinical diagnosis of OA**

Clinical diagnosis of OA was made by a rheumatologist, blinded from clinical data, based on the American College of Rheumatology criteria. Clinical diagnosis of knee OA required the presence of knee arthralgia, plus any 3 of the following: 1) age >50 years, 2) morning stiffness, 3) crepitus on active motion in at least one side, 4) bony enlargement in at least one side, and 5) no palpable warmth of synovia in both knees.15 Participants without clinical OA were excluded from analyses involving those with clinical OA only.

**Pain, stiffness, and functional ability**

Pain, stiffness, and functional ability were assessed by the Western Ontario and McMaster Universities Arthritis Index (WOMAC) questionnaire visual analog scale version, in English, Malay, Chinese, or Tamil. The scores for each item were obtained by measuring the distance along the line from the marker at the far left of each analog scale to the point at which the patient’s mark intersects the horizontal line on the analog scale. The score of each subscale as well as the grand total score was calculated. The maximum total length or score for pain, stiffness, and physical function subscales were 500 mm, 200 mm, and 1,700 mm, respectively.16 Individuals with no pain, no stiffness, and no impairment in functional ability were those with a score of zero for pain, stiffness, and physical function. The sum of the individual component scores for pain, stiffness, and physical function was calculated to obtain the total WOMAC score. Using 25th, 50th, and 75th percentile values as ordinal cutoffs, the non-normally distributed total WOMAC score was further categorized into 4 groups: no symptoms (0 mm), mild symptoms (1–200 mm), moderate symptoms (201–465 mm), and severe symptoms (≥466 mm).

**Falls efficacy and psychological status**

FoF

The shortened version of the Falls Efficacy Scale-International (short FES-I) was used. The short FES-I consists of seven items on a 4-point Likert scale, with 1 indicating no concern...
and 4 indicating severe concern. The minimum and maximum scores for the short FES-I are therefore 7 and 28, respectively. To allow for adjustment for potential confounders, cutoffs were developed using the median values; subjects with a score of ≥11 were considered to have greater FoF.

Depression, anxiety, and stress
Depression, anxiety, and stress were measured with the 21-item Depression, Anxiety, and Stress Scale (DASS-21). This is a self-reported measure in which subjects rate the frequency and severity of the negative emotions of depression, anxiety, and stress over the previous week. Frequency or severity ratings are made on a series of 4-point scales, with 0 indicating “did not apply to me at all” and 3 indicating “applied to me very much, or most of the time.” The scores were calculated individually for the three components: depression, anxiety, and stress. Because the DASS-21 is the short version of DASS (which has 42 items), the final score for each item were multiplied by 2. The total score for each component was dichotomized using median values as the cutoffs, depression ≥4, anxiety ≥4, and stress ≥8, respectively.

Statistical analysis
The SPSS 20.0 (IBM Corporation, Armonk, NY, USA) statistical package was used for statistical analysis. Normality was assessed using the Kolmogorov–Smirnov test. Mean values with SD or median with interquartile range were expressed for normally and non-normally distributed continuous data accordingly. The independent t-test and Mann–Whitney U test were employed for parametric and non-parametric data, respectively. Categorical variables were expressed as frequencies with percentages in parentheses and compared with the chi-squared test. Spearman’s rank correlation was used to evaluate the correlation between WOMAC subscales with FoF and DASS-21 domain scores. Using cutoffs described earlier, continuous data were grouped into dichotomous variables. Subsequently, logistic regression analysis using fall as a dependent variable was performed to determine potentially mediating effects of FoF and psychological status on the relationship between OA severity and falls. Dummy variables were created for comparisons between multiple categories, with the lowest category (no arthritis or mild arthritis) as the reference category. The exponential value of the parameter estimate, B, for each variable, which would represent the adjusted odds ratio (OR) as a measure of the strength of association was presented along with its 95% CI.

Results
Demographics
A total of 389 subjects, 229 fallers and 160 non-fallers, with a mean age of 72.2±6.1 years, were evaluated and screened for the presence of OA. One hundred and forty-one (36.2%) individuals had clinically diagnosed OA based on the ACR criteria. Knee X-rays were available for 205 participants, and 171 (83%) had radiological evidence of OA. Characteristics of fallers and non-fallers with radiological and clinical OA are summarized in Table 1. Fallers were significantly older than non-fallers in both the groups. There were ethnic differences between fallers and non-fallers with clinical OA. A greater proportion of fallers compared to non-fallers had severe OA symptoms in clinical OA. A higher percentage of non-fallers were seen with “mild” OA symptoms in radiological OA, compared to asymptomatic OA (Table 1).

FoF and psychological status among fallers and non-fallers with different types of OA
In Table 2, fallers with either radiological or clinical OA had significantly greater FoF and reported significantly more severe depressive symptoms than non-fallers. Due to the non-normally distributed data, the individual domain scores were then dichotomized using median cutoffs to allow for adjustment for potential confounders using logistic regression analyses. After adjustment for the potential confounders of age, gender, ethnicity, and comorbidities, the association between greater FoF and falls remained significant in both the types of OA. However, the association between depression and falls in radiological OA group was no longer statistically significant after the same adjustment (Table 2).

FoF and psychological domains correlated with OA symptom severity in fallers and non-fallers
Table 3 documents the correlation coefficients (Spearman’s rho) between FoF, psychological status, and grand total of WOMAC score in fallers and non-fallers who had OA, diagnosed radiologically or clinically. Among fallers with radiological OA or clinical OA, symptom severity was significantly and positively correlated with FoF and all three psychological domains (p<0.01). In contrast, among the non-fallers, symptom severity was only significantly correlated with FoF regardless of the method of OA diagnosis. Symptom severity correlated with depression only in those with radiological OA (Table 3).
Table 1 Characteristics of individuals with radiological OA and clinical OA according to fall status

<table>
<thead>
<tr>
<th></th>
<th>Radiological OA (n=171)</th>
<th>Clinical OA (n=141)</th>
<th>p-value</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Fallers (n=106)</td>
<td>Non-fallers (n=65)</td>
<td></td>
</tr>
<tr>
<td>Age (years), mean (SD)</td>
<td>75.28 (6.92)</td>
<td>71.31 (4.65)</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Female sex, n (%)</td>
<td>75 (70.8)</td>
<td>50 (76.9)</td>
<td>0.377</td>
</tr>
<tr>
<td>BMI (kg/m²), mean (SD)</td>
<td>24.73 (4.22)</td>
<td>24.42 (3.34)</td>
<td>0.623</td>
</tr>
<tr>
<td>Ethnicity*, n (%)</td>
<td>Malay 14 (13.2)</td>
<td>12 (18.5)</td>
<td>Reference</td>
</tr>
<tr>
<td></td>
<td>Chinese 71 (67.0)</td>
<td>46 (70.8)</td>
<td>0.521</td>
</tr>
<tr>
<td></td>
<td>Indian 17 (16.0)</td>
<td>4 (6.2)</td>
<td>0.058</td>
</tr>
<tr>
<td></td>
<td>Others 4 (3.8)</td>
<td>3 (4.6)</td>
<td>0.876</td>
</tr>
<tr>
<td>Comorbidities, n (%)</td>
<td>Heart diseases 9 (8.5)</td>
<td>3 (4.6)</td>
<td>0.336</td>
</tr>
<tr>
<td></td>
<td>Hypertension 57 (53.8)</td>
<td>34 (52.3)</td>
<td>0.852</td>
</tr>
<tr>
<td></td>
<td>Diabetes mellitus 32 (30.2)</td>
<td>13 (20.0)</td>
<td>0.142</td>
</tr>
<tr>
<td></td>
<td>Stroke 7 (6.6)</td>
<td>3 (4.6)</td>
<td>0.591</td>
</tr>
<tr>
<td></td>
<td>Atrial fibrillation 2 (1.9)</td>
<td>1 (1.5)</td>
<td>1.000</td>
</tr>
<tr>
<td></td>
<td>Visual impairment 26 (24.5)</td>
<td>11 (16.9)</td>
<td>0.241</td>
</tr>
<tr>
<td>WOMAC grand total score*, n (%)</td>
<td>No symptoms 27 (25.5)</td>
<td>16 (24.6)</td>
<td>Reference</td>
</tr>
<tr>
<td></td>
<td>Mild 19 (17.9)</td>
<td>28 (43.1)</td>
<td>0.036</td>
</tr>
<tr>
<td></td>
<td>Moderate 25 (23.6)</td>
<td>13 (20.0)</td>
<td>0.779</td>
</tr>
<tr>
<td></td>
<td>Severe 35 (33.0)</td>
<td>8 (12.3)</td>
<td>0.058</td>
</tr>
</tbody>
</table>

Notes: *Logistic regression with dummy variables. Categorized using percentile cutoffs from total WOMAC score. "No symptoms": 0 mm. "Mild": 1–200 mm. "Moderate": 201–465 mm. "Severe": ≥466 mm. Mean age and BMI were compared using Student’s independent t-test. The bold font indicates statistical significance.

Abbreviations: BMI, body mass index; OA, osteoarthritis; IQR, interquartile range; FES-I, Falls Efficacy Scale-International; DASS-21, 21-item Depression, Anxiety, and Stress Scale; WOMAC, Western Ontario and McMaster Universities questionnaire; NA, not applicable.

Multivariate logistic regression evaluating the relationship between OA severity and falls

Table 4 explores the mediating roles of FoF and psychological domains on the association between symptom severity and falls according to OA diagnosis. Models 1–5 assessed the relationship between symptom severity and falls in those with radiological OA, while models 6–10 explored the relationship between symptom severity and falls in those with clinical OA.

Symptom severity and falls in radiological OA

Comparisons were made with falls as the dependent variable and categorical severity of OA as independent variables using no OA symptoms as the reference group. In Model 1, following adjustments for the potential confounders of age, gender, ethnicity, and comorbidities, mild OA symptoms in radiological OA was associated with reduced risk of falls (OR =0.38, 95% CI =0.15 to 0.98) compared to those with radiological OA and no symptoms. No significant difference in falls risk was observed in those with moderate OA symptoms or severe OA symptoms compared to those with no symptoms in this model. In Model 2, the protective relationship between mild OA symptoms remained after adjustments were made for short FES-I scores (OR =0.33, 95% CI =0.12 to 0.87). This relationship also remained statistically significant after adjustment for DASS depression (Model 3: OR =0.38, 95% CI =0.15 to 0.97) and DASS stress scores (Model 4: OR =0.39, 95% CI =0.15 to 0.97).

However, adjustment for DASS anxiety scores led to attenuation of the protective effect of mild OA symptoms against falls compared to no OA symptoms in those with radiological OA (Model 4: OR =0.40, 95% CI =0.16 to 1.01).

Symptom severity and falls in clinical OA

Comparisons were made using falls as the dependent variable and OA symptom severity category. As the presence of symptom is requisite for the diagnosis of clinical OA, OA symptom severity as determined by the WOMAC scores was categorized only according to the mild, moderate, and severe categories. Model 6 suggests that individuals with clinical OA and severe OA symptoms were significantly more likely to experience falls compared to those with mild OA symptoms (OR =3.92, 95% CI =1.51 to 10.20). This relationship was attenuated after adjustment for short FES-I scores (OR =2.56, 95% CI =0.90 to 7.28). However, DASS depression, anxiety, and stress scores did
Table 2  Association between falls and psychological scores for individuals with radiological OA and clinical OA

<table>
<thead>
<tr>
<th></th>
<th>Radiological OA (n=171)</th>
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<tbody>
<tr>
<td></td>
<td>Fallers (n=106)</td>
<td>Non-fallers (n=65)</td>
</tr>
<tr>
<td>Short FES-I, median (IQR)</td>
<td>14 (10–20)</td>
<td>9 (7–13)</td>
</tr>
<tr>
<td>DASS-21, median (IQR)</td>
<td>Depression score</td>
<td>12 (1–23)</td>
</tr>
<tr>
<td></td>
<td>Anxiety score</td>
<td>4 (0–8)</td>
</tr>
<tr>
<td></td>
<td>Stress score</td>
<td>8 (4–24)</td>
</tr>
<tr>
<td>Short FES-I ≥11, n (%)</td>
<td>70 (66.7)</td>
<td>19 (31.1)</td>
</tr>
<tr>
<td>DASS-21, n (%)</td>
<td>Depression ≥4</td>
<td>78 (75.0)</td>
</tr>
<tr>
<td></td>
<td>Anxiety ≥4</td>
<td>68 (66.0)</td>
</tr>
<tr>
<td></td>
<td>Stress ≥8</td>
<td>30 (47.6)</td>
</tr>
</tbody>
</table>

Notes: The bold font indicates statistical significance. *Mann–Whitney U test was used for the non-parametric continuous data. Dichotomized data were categorized using median values. **Adjusted for age, gender, ethnicity, and comorbidities.

Abbreviations: OA, osteoarthritis; OR, odds ratio; IQR, interquartile range; FES-I, Falls Efficacy Scale-International; DASS-21, 21-item Depression, Anxiety, and Stress Scale.

Discussion

Among individuals with OA, fallers were significantly more likely to experience FoF and depression regardless of whether the presence of OA was determined using radiological or clinical criteria. When the relationship imaging or clinical assessment. When the relationship between OA symptoms and psychological status was assessed independently among fallers and non-fallers, however, there was a significant correlation between severity of OA and depression, anxiety, and stress with OA severity scores in falls risk in individuals with OA. Our findings, therefore, highlight a potential role of psychological status and FoF in determining the risk of falls in those individuals with OA.

Further multivariate analyses revealed that the increased risk of falls among women with OA was mediated by both depression and FoF. Regardless of OA severity, fallers had significantly higher levels of depression and anxiety compared to non-fallers.

Among individuals with moderate OA symptoms, the relationship between falls and psychological status was not significant. Among individuals with severe OA symptoms, however, there was a significant correlation between falls risk and depression.

Among fallers, the relationship between falls and psychological status was significant, with higher levels of depression and anxiety associated with an increased risk of falls. This relationship was stronger among non-fallers, indicating that psychological status may play a role in falls risk even among those without a history of falls.

We also examined the relationship between falls and total WOMAC scores in those with severe OA symptoms. We found a significant correlation between falls risk and total WOMAC scores in those with severe OA symptoms. This relationship was stronger among non-fallers, indicating that falls risk is associated with worse physical function in those with severe OA symptoms.

These findings suggest that psychological factors may play a role in falls risk among individuals with OA, and that interventions targeting psychological well-being may be beneficial in reducing falls risk.
physical performance measure of function. As highlighted earlier, these previous studies had used varying definitions of OA and may not necessarily employ actual fall events as their primary outcome. Our study has therefore emphasized that psychological factors do play a role in falls risk regardless of OA definitions. In addition, our study had compared actual fallers with non-fallers rather than employing surrogate measures such as balance performance, as in previous studies.

The etiology of psychological symptoms in OA and the increased psychological burden in fallers with OA has not been well elucidated. Pain and joint stiffness associated with OA are chronic, though often branded less severe than other inflammatory joint conditions. The long-term effects of joint pain and stiffness could have a direct consequence on mood. In addition, a previous study has demonstrated a significant association between FoF and pain. The presence of OA, however, also influences joint stability, which occurs as a result of ligamentous laxity from deconditioning occurring as a consequence of activity avoidance due to joint symptoms. Knee buckling, which occurs as a result of joint instability, is in turn, associated with FoF. Furthermore, joint pain and stiffness, instability, FoF, and reduced physical activity all lead to reduced functional ability, which will also adversely affect psychological well-being. The increased risk of FoF and depression among fallers with OA compared to non-fallers may imply that fallers were either more likely to experience the adverse psychological consequences of OA or that fallers with OA experience additional psychological burden over non-fallers with OA. However, when further within group analyses were employed, while the dose–response relationship is observed between FoF and OA symptom severity in both fallers and non-fallers with OA, the dose–response relationship between depression, stress, and anxiety and OA symptom severity only among fallers. The cross-sectional design of our study does not allow for the determination of the temporal relationships between FoF and depression with falls.

We further explored the effects of FoF, depression, anxiety, and stress on the differential relationship of severity of OA symptoms with falls using multivariate logistic regression. The reduced risk of falls observed among those with radiological OA and mild symptoms had previously been reported. Our mediation analysis, however, suggests that symptoms of anxiety may account for the reduced falls risk in those with radiological OA and mild symptoms over those with radiological OA and no symptoms. This suggests that those with increased anxiety may exhibit increased vigilance or avoidance of challenging activities which then reduces their risk of falling. The exposure of these individuals to extrinsic risk factors for falls such as physical environment, assistive devices, or footwear may also be reduce.

Conversely, while individuals with clinical OA and severe symptoms were significantly more likely to experience falls than those with mild symptoms, this relationship was not influenced by psychological status. Our findings were consistent with that of Quach and Burr who also found no interaction between clinically significant depressive symptoms with falls in their subjects with OA. FoF, on the other hand, seemed to account for the excess falls risk associated with severe OA compared to mild OA among individuals with
clinical OA. Our cross-sectional findings, as before, does not allow us to determine any causal relationship between FoF and falls in individuals with clinical OA, but does, however, highlight FoF as a potentially modifiable risk factor for falls among those with severe clinical OA. A recent systematic review suggested that exercise interventions may reduce FoF in community-dwelling general older populations.29 Now, the use of actual psychological interventions in the management of FoF is also being evaluated.30 The determination of the value of exercise and psychological interventions on FoF and falls may therefore be worthwhile in individuals with clinical OA.

As mentioned earlier, the cross-sectional design of our study precludes the establishment of temporality or causation. Fallers in our study also consisted of those with recurrent or injury falls, who will be considered high-risk fallers. The risk of injury and healthcare burden are particularly high among this group of fallers, which may limit the applicability of our findings to those who sustain one accidental fall with no injury, but may suffer adverse psychosocial consequences as a result. In addition, the presence of falls in the previous 12 months was determined through self-report, which may be subject to recall bias.31 A larger, more robust, prospective study to help establish the temporal relationship between FoF and other psychological symptoms and falls in individuals with OA should now be conducted to aid further understanding of the complex relationships between psychological status, OA, and falls.32,33

Conclusion
Older fallers with radiological and clinical OA have an increased FoF and depressive symptoms compared to non-fallers. A dose–response relationship exists between FoF, depression, anxiety, and stress with OA symptom severity among fallers with OA. Further exploratory analyses suggest mediating effects of anxiety on the protective effect of mild symptoms among those with radiological OA and FoF on the increased risk of falls among those with severe clinical OA. Our findings highlight the important role of psychological status in falls among older individuals with OA, which will help determine the direction of future studies on strategies to reduce the burden on falls in those with OA.

Acknowledgments
We thank Dr Karen Morgan for her valuable input during the preparation of this manuscript. This study has been funded by a University of Malaya Grand Challenge Research Fund (GC002-14HTM) and a University of Malaya Postgraduate Research Fund (PPP) grant (PG013-2014B).

Disclosure
The study sponsors had no role in the collection, analysis and interpretation of data; in the writing of the manuscript; or in the decision to submit the manuscript for publication. The authors report no conflicts of interest in this work.

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