The impact of sarcopenia on low back pain and quality of life in patients with osteoporosis

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Abstract

Purpose: Osteoporosis combined with sarcopenia contributes to a high risk of falling, fracture, and even mortality. However, sarcopenia’s impact on low back pain and quality of life (QOL) in patients with osteoporosis is still unknown. The purpose of this study is to investigate low back pain and QOL in osteoporosis patients with sarcopenia.

Methods: We assessed 100 ambulatory patients who came to our hospital for osteoporosis treatment. Low back pain was evaluated using the Visual Analogue Scale (VAS) with 100 being an extreme amount of pain and 0 no pain. The Japanese Orthopaedic Association Back Pain Evaluation Questionnaire (JOABPEQ) score was used to assess QOL after adjustment for age, history of vertebral fracture, and adult spinal deformity. Differences in low back pain intensity assessed by VAS between groups were evaluated by the Wilcoxon rank-sum test. Covariance analysis was used to assess QOL. All data are expressed as either median, interquartile range, or average, standard error.

Results: Patients were classified into the sarcopenia group (n = 32) and the non-sarcopenia group (n = 68). Low back pain intensity assessed by VAS was significantly higher in the sarcopenia group than in the non-sarcopenia group (33.0 [0-46.6] vs. 8.5 [0-40.0]; p < 0.05). The subscales of the JOABPEQ for low back pain were significantly lower in the sarcopenia group than in the non-sarcopenia group (65.0 ± 4.63 vs. 84.0 ± 3.1; p < 0.05).

Conclusion: In this cross-sectional study, sarcopenia affected low back pain and QOL in ambulatory patients with osteoporosis. Sarcopenia may exacerbate low back pain and QOL.

Keywords: Osteoporosis, Adults spinal deformity, Sarcopenia, Quality of life, Sagittal alignments

Introduction

Muscle mass has been observed to decrease with age at a rate of approximately 1% annually after age 40 [1]. Sarcopenia is defined as the loss of skeletal muscle mass and strength that occurs with advancing age [2]. Osteoporosis is defined as a skeletal disorder characterized by compromised bone strength leading to an increased risk of fracture [3]. Both sarcopenia and osteoporosis are geriatric diseases that decrease activity in daily living, and are interconnected physically and chemically. Recently, combined osteoporosis and sarcopenia are called “osteosarcopenia” [4]. Osteosarcopenia contributes to a high risk of falling, fracture, and even mortality [5]. Since osteosarcopenia is a new concept, studies of osteosarcopenia are insufficient. Thus, it remains unclear how sarcopenia affects low back pain and quality of life (QOL) in patients with osteoporosis.

Aging and decline of physical activity lead to decreased QOL. Spinal misalignment induced by loss of skeletal muscle mass and vertebral fracture also leads to decreased QOL [6, 7]. In particular, increased thoracic kyphosis leads to decreased QOL in patients with
osteoporosis [8]. The SRS-Schwab classification system is an assessment scale for spinal deformity using radiographic parameters including the sagittal vertical axis, pelvic tilt (PT), and pelvic incidence minus lumbar lordosis (PI-LL) [9]. Among them, PT is one of the most important parameters for assessing spinal deformity, PT is significantly correlated with pain, disability, and QOL [10].

In this cross-sectional study, we hypothesized that osteoporosis patients with sarcopenia would have more low back pain and poorer QOL than osteoporosis patients without sarcopenia. The purpose of this study was to investigate the effect of sarcopenia on low back pain and QOL in ambulatory patients with osteoporosis.

Methods

Ethics

The study protocol conformed to the ethics guidelines of the Declaration of Helsinki, as reflected in prior approval given by the institutional review board of Munakata Suikokai General Hospital (approval ID: 19,002). Informed consent from patients was obtained using an opt-out approach.

Patients

We did a cross-sectional study and assessed 100 ambulatory patients when they came to our hospital (Munakata Suikoukai General Hospital, Japan) for osteoporosis treatment between January 2018 and December 2018. We summarized inclusion and exclusion criteria in Table 1. According to the Japanese Society for Bone and Mineral Research, we diagnosed osteoporosis and osteopenia [11].

Diagnosis of sarcopenia

We assessed sarcopenia according to the Asian Working Group for Sarcopenia guidelines [12]. (1) Cutoff values for handgrip strength are <26 kg for men and <18 kg for women. The cutoff value for walking speed is the usual gait speed of <0.8 m/s. Patients with either grip strength or walking speed less than the cutoff value are suspected to have sarcopenia. (2) In cases of suspected sarcopenia, the next-step is to evaluate muscle mass. Cutoff values for skeletal muscle mass index (SMI) are 7.0 kg/m² for men and 5.7 kg/m² for women measured by bioimpedance analysis. (3) Patients with less than the muscle mass cutoff are diagnosed with sarcopenia.

Outcome measures

Bone mineral density

Areal BMD of the posterior-anterior lumbar spine was measured by dual-energy x-ray absorptiometry using a Hologic QDR 4500 A densitometer (Hologic, Waltham, MA). All scans of an individual subject were performed using the same densitometer. Quality control measurements were performed daily with a Hologic anthropomorphic spine phantom.

Isometric knee extension strength

Isometric knee extension strength on the dominant side was measured at discharge using a manual muscle strength monitor (Mobie; Sakai Medical Co., Ltd.) [13]. Patients sat on the edge of a bed with their feet not touching the floor and with their arms crossed in front of their body. The highest value of three assessments was used in the analysis.

Evaluation of pain

Low back pain, pain from the buttocks to lower limbs, and numbness of the buttocks to lower limbs were evaluated using a Visual Analogue Scale (VAS) with 100 being an extreme amount of pain and 0 no pain.

Spinal alignment

The patient was radiographed at the clavicle position [Poosiripinyo T (2017) J Orthop Sci 22 (1):34-37.]. LL, PI, PT, and sacral slope (SS) were measured from various lumbar spine lateral views (Fig. 1). We defined adult spinal deformity as PT>30 degrees, based on the SRS-Schwab classification [10].

Table 1 Inclusion and exclusion criteria

| Inclusion Criteria | Exclusion criteria |
|--------------------|--------------------|
| Patients 60 years of age or over, with osteoporosis or osteopenia | Unable to stand |
| Osteoporosis: History of fragility fracture and bone mineral density (BMD) <80% of the young adult mean, or BMD<70% | Unable to walk 10 meters |
| Osteopenia: BMD was 70% to 80% of the young adult mean | Unable to be measured in various physical tests |
| Pacemaker | Long-term steroid use |
| Fresh vertebral fracture | Hemiplegia |
Disease-specific QOL measure for patients with low back pain
The Japanese Orthopaedic Association Back Pain Evaluation Questionnaire (JOABPEQ) was designed for the assessment of disease-specific QOL and low back pain and lumbar spinal disease. It is a disease-specific tool that contains 25 items tapping into five subscales: low back pain (four items), lumbar function (six items), walking ability (five items), social function (four items), and mental health (seven items). The score for each subscale ranges from 0 to 100, with higher scores indicating better conditions. The JOABPEQ is used in Japan and other countries, and its reliability has been proven [14]. Recently, the JOABPEQ is used to assess low back pain patients and can be used to assess other lumbar spine disorders such as lumbar canal stenosis, lumbar disc herniation, and neuropathic pain in a low back pain patient [14].

Statistical analysis
All the statistical analysis were performed by JMP Version 14.0 statistical software (SAS Institute Inc., Cary, NC, USA), with $P < 0.05$ considered statistically significant in all cases. Differences in patient characteristics at the time of visit and JOABPEQ results between the sarcopenia and non-sarcopenia groups were evaluated by the Wilcoxon rank-sum test. QOL is affected by age, history of vertebral fracture and adult spinal deformity, so we used analysis of covariance to adjust these factors. Pearson correlation coefficient was used to determine the correlation between SMI and spinal alignment parameters or the JOABPEQ. The Pearson correlation coefficient measures the strength of the linear relationship [15, 16]. It ranges from -1 to 1. The correlation coefficient value may be expressed from very weak to very strong in increments of 0.2. 0 to 0.19 is considered very weak, either positive or negative, and 0.8 to 1.0 is considered very strong. A multivariate Cox regression analysis with stepwise variable selection was used to identify independent variables associated with low back pain in the JOABPEQ. All data are expressed as either median, (interquartile range [IQR]), range, or average, standard error.
Results

Patient characteristics at the time of visit

Patients were classified into sarcopenia (n = 32) and non-sarcopenia groups (n = 68), and their clinical characteristics at the time of visit are shown in Table 2. Patients in the sarcopenia group were significantly older than those in the non-sarcopenia group (p < 0.05). Body mass index (BMI) and SMI were significantly lower in the sarcopenia group than in the non-sarcopenia group (p < 0.05). The ratio of history of vertebral fracture and adult spinal deformity was significantly higher in the sarcopenia group than in the non-sarcopenia group (p < 0.05).

Bone mineral density, Isometric knee extension strength and Evaluation of Pain

BMD, and isometric knee extension torque were significantly lower in the sarcopenia group than in the non-sarcopenia group ([BMD (g/cm²): 0.50 [0.42-0.56] vs. 0.56 [0.50-0.64]; p = 0.0079, isometric knee extension strength (kgf/kg): 11.7 [6.4-15.8] vs. 17.1 [13.1-21.5]; p < 0.0001]). Low back pain intensity assessed by VAS was significantly higher in the sarcopenia group than in the non-sarcopenia group (33.0 [0-46.6] vs. 8.5 [0-40.0]; 0.0312). There was no significant difference in VAS for pain from buttocks to lower limbs and numbness from buttocks to lower limbs (Pain from buttocks to lower limbs assessed by VAS (mm): 0 [0-0] vs. 0 [0-0]; 0.5409, Numbness of buttocks to lower limbs assessed by VAS (mm): 0 [0-0] vs. 0 [0-0]; p = 0.8124).

Spinal Alignment

PT in the sarcopenia group was significantly higher than in the non-sarcopenia group (p < 0.05). LL and SS were significantly lower in the sarcopenia group than in the non-sarcopenia group (p < 0.05). There was no significant difference in PI (Table 2).

JOABPEQ (Disease-specific QOL) before adjusting

The results for QOL evaluated with the JOABPEQ are shown in Table 3. The subscales for low back pain, lumbar function, walking ability, and social life function were significantly lower in the sarcopenia group than in the non-sarcopenia group (p < 0.05). No significant differences were observed in the other subscales.

Table 2

|                      | Sarcopenia (N = 32) | Non-sarcopenia (N = 68) | p       |
|----------------------|---------------------|-------------------------|---------|
|                      | Median (IQR) | Range (min-max) | Median (IQR) | Range (min-max) |         |
| Age                  | 81.5 (74.5-87.8) | 63-97 | 71.5 (68.3-80.0) | 60-91 | 0.0001 |
| Sex (female/male)    | 28/4      |       | 60/8       |       | 0.9159 |
| BMI (kg/m²)          | 20.7 (18.8-23.1) | 14.3-26.3 | 22.5 (20.7-24.7) | 17.5-32.2 | 0.0072 |
| SMI (kg/m²)          | 5.1 (4.7-5.5) | 4.2-6.8 | 6.1 (5.7-6.7) | 4.9-7.9 | <.0001 |
| BMD (femoral neck)   | 0.50 (0.42-0.56) | 0.38-0.83 | 0.56 (0.50-0.64) | 0.38-0.90 | 0.0079 |
| Knee extension torque (kgf/kg) | 11.7 (6.4-15.8) | 3.4-20.3 | 17.1 (13.1-21.5) | 3.9-46.3 | <.0001 |
| History of vertebral fracture (N) | 19 (55.88%) |       | 14 (20.89%) |       | 0.0004 |
| Adult spinal deformity (N) | 14 (43.75%) |       | 12 (17.91%) |       | 0.0063 |
| Pelvic Tilt (°)      | 29.0 (21.2-37.8) | 8.7-57.0 | 22.0 (16.0-28.0) | 4.0-39.7 | 0.005  |
| Lumbar Lordosis (°)  | 36.0 (26.3-49.8) | 12.0-68.0 | 43.0 (36.0-52.9) | 3.0-79.0 | 0.0445 |
| Pelvic Incidence (°) | 54.0 (49.0-68.5) | 38.0-91.0 | 55.0 (50.0-61.0) | 34.0-85.0 | 0.6614 |
| Sacral Slope (°)     | 28.0 (23.3-36.8) | 13.0-56.2 | 32.2 (27.0-40.0) | 2.0-54.0 | 0.0441 |

Abbreviations: BMI body mass index, BMD bone mineral density, SMI skeletal muscle mass, VAS Visual Analogue Scale

Table 3

|                      | Sarcopenia (N = 32) | Non-sarcopenia (N = 68) | p       |
|----------------------|---------------------|-------------------------|---------|
|                      | Median (IQR) | Range (min-max) | Median (IQR) | Range (min-max) |         |
| Low back pain        | 50 (43-100)  | 14-100 | 100 (71-100)  | 29-100 | 0.0003 |
| Lumbar function      | 79 (17-83)   | 0-100 | 83 (75-100)   | 8-100 | 0.0031 |
| Walking ability      | 64 (29-91)   | 0-100 | 93 (66-100)   | 0-100 | <.0001 |
| Social life function | 65 (30-78)   | 0-100 | 78 (57-100)   | 14-100 | 0.0014 |
| Mental health        | 54 (49-74)   | 21-94 | 63 (51-78)    | 39-96 | 0.1041 |
difference in mental health was found between the two groups.

**JOABPEQ (Disease-specific QOL) after adjusted by age, history of vertebral fracture, and adult spinal deformity**
The subscale score for low back pain was significantly lower in the sarcopenia group than in the non-sarcopenia group ($p < 0.05$). No significant difference between the two groups was found for lumbar function, walking ability, social life function, and mental health (Table 4).

**Correlation between variables**
Correlations done for spinal alignment revealed a significant negative correlation between PT and both SMI and BMD (SMI; $r = -0.232; p = 0.0207$, BMD; $r = -0.253; p = 0.0125$). PT had a significant negative correlation with all categories in the JOABPEQ except for mental health (low back pain; $r = -0.298; p = 0.003$, lumbar function; $r = -0.249; p = 0.0126$, walking ability; $r = -0.348; p = 0.0004$, social function; $r = -0.294, p = 0.0031$, mental health; $r = -0.0436, p = 0.6684$). PT had a significant positive correlation with low back pain intensity assessed by VAS ($r = 0.2357; p = 0.0188$) (Fig. 2).

**Independent factors associated with low back pain in JOABPEQ**
A multivariate Cox regression analysis was done with variables related to low back pain; age, BMI, gender, PT, LL, SS, BMD, SMI, gait-speed, grip, and sarcopenia. Sarcopenia and walking speed were the only significant factors for low back pain in the JOABPEQ (sarcopenia: estimate -9.694; 95% confidence interval -15.378 — -4.011; $p = 0.010$, walking speed: estimate 15.186; 95% confidence interval 3.027 — 27.345; $p = 0.0149$).

**Discussion**
We assessed QOL and low back pain in ambulatory patients with osteoporosis. The ratio of sarcopenia was 32% in this study. Low back pain intensity assessed by VAS was significantly higher in the sarcopenia group than in the non-sarcopenia group. A multivariate Cox regression analysis extracted sarcopenia as the independent factor associated with back pain in the JOABPEQ. Sarcopenia might affect QOL and low back pain in patients with osteoporosis.

**Ratio of sarcopenia**
In this study, we investigated the ratio of sarcopenia in osteoporosis patients. The proportion of osteoporosis patients with sarcopenia was 32% (average age 76.4 years). Yoshimura et al. investigated 1,099 Japanese patients with osteoporosis (average age 72.1 years) and reported a sarcopenia ratio of 19.1% [17]. They also showed a yearly increase with age of about 2.0% (male 2.2%, female 1.9%) in the ratio of sarcopenia in patients with osteoporosis. Morley et al. reported that the ratio of sarcopenia varies by age, with the ratio in a 60- to 70-year-old group of 5-13% and a ratio of 11-50% for those over 80 [12]. Taken together, the higher average age of our patients may have influenced the ratio of sarcopenia.

**Low back pain and sarcopenia**
Low back pain intensity assessed by VAS in the sarcopenia group was significantly higher than in the non-sarcopenia group in this study. Sarcopenia was a significant factor for low back pain in JOABPEQ using multivariate Cox regression analysis. Sakai et al. investigated the relationship between low back pain and sarcopenia. The ratio of sarcopenia was 40.0% in the low back pain group and 26.6% in the non-low back pain group. Low back pain is correlated with SMI rather than with BMD. They concluded that sarcopenia was associated with low back pain [15]. Moreover, Spinal alignment including PT is one of the causes of low back pain. Schwab found PT is significantly correlated with an Oswestry Disability Index Score which suggests low back pain (cut off of PT is 22) [10]. Eguchi et al.

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**Table 4** Japanese Orthopaedic Association Back Pain Evaluation Questionnaire (JOABPEQ : Disease-specific QOL) adjusted by age, history of vertebral fracture, and adult spinal deformity

| Sarcopenia (N = 32) | Non-sarcopenia (N = 68) | p |
|---------------------|-------------------------|---|
| Low back pain       | 64.44 ± 4.42            | 84.25 ± 3.10 | 0.0005 |
| Lumbar function     | 62.98 ± 4.84            | 73.31 ± 4.07 | 0.0995 |
| Walking ability     | 67.55 ± 4.78            | 75.26 ± 3.42 | 0.1777 |
| Social life function| 63.08 ± 4.19            | 71.22 ± 3.53 | 0.13298 |
| Mental health       | 61.94 ± 3.04            | 65.66 ± 2.18 | 0.3057 |

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(See figure on next page.)

**Fig. 2** Correlations with pelvic tilt (PT). A statistically significant negative correlation was observed between PT and both skeletal muscle mass (A) and bone mineral density (B). A statistically significant negative correlation was observed between PT and the subscale scores of JOABPEQ for low back pain (C), lumbar function (D), walking ability (E), and social life function (F). A statistically significant positive correlation was observed between PT and Low back pain intensity assessed by VAS (G). Abbreviations: JOABPEQ, Japanese Orthopaedic Association Back Pain Evaluation Questionnaire; PT, pelvic tilt; VAS, Visual Analogue Scale
Fig. 2 (See legend on previous page.)

A  $r=-0.232, *p=0.0207$

B  $r=-0.253, *p=0.0125$

C  $r=-0.298, *p=0.003$

D  $r=-0.249, *p=0.0126$

E  $r=-0.348, *p=0.0004$

F  $r=-0.294, *p=0.0031$

G  $r=0.2357, *p=0.0188$
reported that low back pain had a negative correlation with SMI and a positive correlation with PT, suggesting that sarcopenia may be associated with low back pain as a result of posterior pelvic tilt [16]. PT was significantly higher in the sarcopenia group than the non-sarcopenia group in this study (29.0 vs. 22.0). PT was also significantly correlated with low back pain intensity assessed by VAS. Thus, our results were similar to the previous reports. Sarcopenia may be related to posterior pelvic tilt and low back pain.

QOL and sarcopenia
Sarcopenia affects not only physical function but also fracture and mortality, although it was previously unclear that sarcopenia affects QOL in patients with osteoporosis. We first showed that sarcopenia affects QOL in patients with osteoporosis in this study. We assessed QOL using the JOABPEQ in this study. Subscale scores for low back pain, lumbar function, walking ability, and social life function were significantly lower in the sarcopenia group than in the non-sarcopenia group. In particular, the subscale score for low back pain was significantly lower in the sarcopenia group than in the non-sarcopenia group after adjusting for age, history of vertebral fracture, and adult spinal deformity. We also investigated the relationship between spinal alignment and both SMI and QOL by using the Pearson correlation coefficient. PT was significantly correlated with each item except for mental health in the QOL assessment by JOABPEQ and also significantly correlated with SMI. Appendicular and trunk muscle mass form the pelvic/lumbar stabilization structure. Eguchi et al. reported that sarcopenia was one of the causes for spinal deformity affecting QOL. They showed that PT is an important factor that affects QOL in patients with scoliosis which is consistent with our findings [16]. They discussed that sarcopenia worsens QOL through spinal deformity caused by decreasing muscle mass. Jung et al. also showed that PT is correlated with QOL in osteoporosis patients [18]. Moreover, Miyakoshi et al. concluded that the lower QOL of osteoporosis patients may be associated with deformity of spinal alignment related to generalized muscle weakness [6]. Thus, sarcopenia may worsen QOL through spinal deformity caused by decreasing muscle mass in patients with osteoporosis.

Pathological mechanisms underlying the correlation between sarcopenia, a low back pain, and reduction of QOL
In this study, sarcopenia affected quality of life and low back pain intensity. A possible explanation is explained by the following pathological mechanisms. It is crucial to maintain an upright position against gravity for humans walking upright on two legs. Anti-gravity muscles such as the multifidus, erector spinae, and vastus lateralis are involved in holding the upright position [19]. These anti-gravity muscles include type I and type II muscle fibers. However, they contain largely type II muscle fibers. Sarcopenia is caused by inflammatory cytokines that cause fatty infiltration of skeletal muscle and induce apoptosis, especially of type II muscle fibers [20]. Fatty infiltration of antigravity muscles such as the multifidus muscle is associated with low back pain [21]. Fat infiltration is also associated with reduced lumbar function [22]. Patients in the sarcopenia group was significantly older than those in the non-sarcopenia group in this study. Age-related type II fiber atrophy and fat infiltration may have affected low back pain and QOL. The ratio of history of vertebral fracture was significantly higher in the sarcopenia group than in the non-sarcopenia group. Dysmobility syndrome is correlating sarcopenia and osteoporosis together with mobility disturbances, obesity, fractures, and falls [23]. Dysmobility syndrome is a useful concept to identify fragility fracture and to do an early intervention, rather than focusing on a single disease, osteoporosis, sarcopenia, or obesity [24]. Dysmobility syndrome is a risk factor of fragility fracture in patients with osteoporosis [25, 26]. Langella et al. investigated the effect of vertebral fractures on spinal deformity. They reported that thoracic kyphosis caused by vertebral body fracture increased the burden on skeletal muscles to maintain alignment in the sagittal plane, resulting in pain [7]. Thus, dysmobility syndrome, combined with sarcopenia and osteoporosis may have caused the fracture to be caused in this study, and the vertebral fracture may have exacerbated the pain. Taken together with our results, the sarcopenia group seems to have exacerbated pain and QOL due to age-related decrease in antigravity muscles and vertebral body fractures resulting from sarcopenia.

Limitation
This study has several limitations. First, it is a single-center cross-sectional study, there were a small number of cases, and the gender ratio was symmetric. Second, we were unable to assess disease affecting low back pain including lumbar vertebrae herniated disk, ossification of posterior longitudinal ligament, and spinal canal stenosis. These diseases may affect a patient’s low back pain. Third, we showed that sarcopenia exacerbates low back pain and QOL, however, it remains unclear how sarcopenia affects treatments for osteoporosis. It is necessary to perform a multicenter prospective cohort study to investigate the impact of sarcopenia on treatments for osteoporosis.
Conclusions
In this cross-sectional study, we investigated the effect of sarcopenia on low back pain and QOL in ambulatory patients with osteoporosis. Sarcopenia may worsen low back pain and QOL through spinal deformity caused by decreasing muscle mass in patients with osteoporosis.

Abbreviations
QOL: Quality of life; SMI: Skeletal muscle mass index; JOABPEQ: Japanese Orthopaedic Association Back Pain Evaluation Questionnaire; VAS: Visual Analogue Scale; LL: Lumbar lordosis; PI: Pelvic incidence; SS: Sacral slope; PT: Pelvic tilt; BMI: Body mass index; BMD: Bone mineral density.

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Authors' contributions
RH, SI, HM participated in study conception and design, interpretation of data, and drafting the manuscript. MB, SI, KH, TH, EH, YH, AK, HI, YM, KM performed data acquisition. KY, KS, TY, KY, NS, participated in interpretation of data and critical revision. The authors read and approved the final manuscript.

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Availability of data and materials
The datasets generated and/or analysed during the current study are available in the University Hospital Medical Information Network Clinical Trial Registry (UMIN000042021). https://center6.umin.ac.jp/cgi-open-bin/ctr_e/ctr_view.cgi?recptno=R000047958.

Declarations
Ethics approval and consent to participate
The study protocol conformed to the ethics guidelines of the Declaration of Helsinki, as reflected in prior approval given by the institutional review board of Munakata Suikokai General Hospital. Informed consent from patients was obtained using an opt-out approach.

Consent for publication
Not applicable.

Competing interests
All authors declare no competing interests.

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