

World Journal of *Orthopedics*

World J Orthop 2017 October 18; 8(10): 747-814



**DIAGNOSTIC ADVANCES**

- 747 Nuclear medicine imaging in osteonecrosis of hip: Old and current concepts
Agrawal K, Tripathy SK, Sen RK, Santhosh S, Bhattacharya A

ORIGINAL ARTICLE**Basic Study**

- 754 Bone regeneration with osteogenic matrix cell sheet and tricalcium phosphate: An experimental study in sheep
Kira T, Akahane M, Omokawa S, Shimizu T, Kawate K, Onishi T, Tanaka Y

Case Control Study

- 761 Neuropathic pain-like symptoms and pre-surgery radiographic severity contribute to patient satisfaction 4.8 years post-total joint replacement
Warner SC, Richardson H, Jenkins W, Kurien T, Doherty M, Valdes AM

Retrospective Cohort Study

- 770 Soft tissue swelling incidence using demineralized bone matrix in the outpatient setting
Chin KR, Pencle FJR, Seale JA, Valdivia JM

Retrospective Study

- 777 Total joint replacement in inhibitor-positive haemophilia: Long-term outcome analysis in fifteen patients
Danielson H, Lassila R, Ylinen P, Yrjönen T

Observational Study

- 785 Digital blinding of radiographs to mask allocation in a randomized control trial
Slobogean GP, Soswa L, Rotunno G, O'Brien PJ, Lefavivre KA
- 790 Restoration of the joint geometry and outcome after stemless TESS shoulder arthroplasty
von Engelhardt LV, Manzke M, Breil-Wirth A, Filler TJ, Jerosch J

SYSTEMATIC REVIEWS

- 798 Scaffolds based therapy for osteochondral lesions of the talus: A systematic review
Shimozono Y, Yasui Y, Ross AW, Miyamoto W, Kennedy JG

CASE REPORT

- 809 Managing extremely distal periprosthetic femoral supracondylar fractures of total knee replacements - a new PHILOS-ophy
Donnelly KJ, Tucker A, Ruiz A, Thompson NW

RETRACTION NOTE

- 814** Retraction note to: Strategy for prevention of hip fractures in patients with Parkinson's disease
Cui Q

ABOUT COVER

Editorial Board Member of *World Journal of Orthopedics*, Matthew Robert Schmitz, MD, Assistant Professor, Attending Doctor, Department of Orthopedics and Rehab, San Antonio Military Medical Center, Fort Sam, Houston, TX 78231, United States

AIM AND SCOPE

World Journal of Orthopedics (*World J Orthop*, *WJO*, online ISSN 2218-5836, DOI: 10.5312) is a peer-reviewed open access academic journal that aims to guide clinical practice and improve diagnostic and therapeutic skills of clinicians.

WJO covers topics concerning arthroscopy, evidence-based medicine, epidemiology, nursing, sports medicine, therapy of bone and spinal diseases, bone trauma, osteoarthropathy, bone tumors and osteoporosis, minimally invasive therapy, diagnostic imaging. Priority publication will be given to articles concerning diagnosis and treatment of orthopedic diseases. The following aspects are covered: Clinical diagnosis, laboratory diagnosis, differential diagnosis, imaging tests, pathological diagnosis, molecular biological diagnosis, immunological diagnosis, genetic diagnosis, functional diagnostics, and physical diagnosis; and comprehensive therapy, drug therapy, surgical therapy, interventional treatment, minimally invasive therapy, and robot-assisted therapy.

We encourage authors to submit their manuscripts to *WJO*. We will give priority to manuscripts that are supported by major national and international foundations and those that are of great basic and clinical significance.

INDEXING/ABSTRACTING

World Journal of Orthopedics is now indexed in Emerging Sources Citation Index (Web of Science), PubMed, PubMed Central and Scopus.

FLYLEAF

I-III Editorial Board

EDITORS FOR THIS ISSUE

Responsible Assistant Editor: *Xiang Li*
Responsible Electronic Editor: *Ya-Jing Lu*
Proofing Editor-in-Chief: *Lian-Sheng Ma*

Responsible Science Editor: *Jin-Xin Kong*
Proofing Editorial Office Director: *Xiu-Xia Song*

NAME OF JOURNAL
World Journal of Orthopedics

ISSN
 ISSN 2218-5836 (online)

LAUNCH DATE
 November 18, 2010

FREQUENCY
 Monthly

EDITORS-IN-CHIEF
Quanjun (Trey) Cui, MD, Professor, Department of Orthopaedic Surgery, School of Medicine, University of Virginia, Charlottesville, VA 22908, United States

Bao-Gan Peng, MD, PhD, Professor, Department of Spinal Surgery, General Hospital of Armed Police Force, Beijing 100039, China

EDITORIAL BOARD MEMBERS
 All editorial board members resources online at <http://www.wjgnet.com>

www.wjgnet.com/2218-5836/editorialboard.htm

EDITORIAL OFFICE
 Xiu-Xia Song, Director
World Journal of Orthopedics
 Baishideng Publishing Group Inc
 7901 Stoneridge Drive, Suite 501, Pleasanton, CA 94588, USA
 Telephone: +1-925-2238242
 Fax: +1-925-2238243
 E-mail: editorialoffice@wjgnet.com
 Help Desk: <http://www.fjpublishing.com/helpdesk>
<http://www.wjgnet.com>

PUBLISHER
 Baishideng Publishing Group Inc
 7901 Stoneridge Drive,
 Suite 501, Pleasanton, CA 94588, USA
 Telephone: +1-925-2238242
 Fax: +1-925-2238243
 E-mail: bpgoffice@wjgnet.com
 Help Desk: <http://www.fjpublishing.com/helpdesk>
<http://www.wjgnet.com>

PUBLICATION DATE
 October 18, 2017

COPYRIGHT
 © 2017 Baishideng Publishing Group Inc. Articles published by this Open-Access journal are distributed under the terms of the Creative Commons Attribution Non-commercial License, which permits use, distribution, and reproduction in any medium, provided the original work is properly cited, the use is non commercial and is otherwise in compliance with the license.

SPECIAL STATEMENT
 All articles published in journals owned by the Baishideng Publishing Group (BPG) represent the views and opinions of their authors, and not the views, opinions or policies of the BPG, except where otherwise explicitly indicated.

INSTRUCTIONS TO AUTHORS
<http://www.wjgnet.com/bpg/gerinfo/204>

ONLINE SUBMISSION
<http://www.fjpublishing.com>

Case Control Study

Neuropathic pain-like symptoms and pre-surgery radiographic severity contribute to patient satisfaction 4.8 years post-total joint replacement

Sophie C Warner, Helen Richardson, Wendy Jenkins, Thomas Kurien, Michael Doherty, Ana M Valdes

Sophie C Warner, Helen Richardson, Wendy Jenkins, Michael Doherty, Ana M Valdes, Academic Rheumatology, University of Nottingham, Clinical Sciences Building, Nottingham City Hospital, Nottingham NG5 1PB, United Kingdom

Sophie C Warner, Department of Cardiovascular Sciences, University of Leicester and National Institute for Health Research, Leicester Cardiovascular Biomedical Research Unit, Leicester LE3 9QP, United Kingdom

Thomas Kurien, Michael Doherty, Ana M Valdes, Arthritis Research UK Pain Centre, Nottingham NG5 1PB, United Kingdom

Thomas Kurien, Academic Division of Trauma and Orthopaedics, Queens Medical Centre, Nottingham NG7 2UH, United Kingdom

Author contributions: All authors have contributed to the conception and design of the study, the acquisition of data and/or the analysis and interpretation of the data; all authors read, provided critical feedback on intellectual content and approved the final manuscript.

Supported by PhD studentship awarded by the University of Nottingham (to Warner SC); EULAR project grant to AMV, No. 108239; ARUK Pain Centre, No. 18769.

Institutional review board statement: The North Nottinghamshire Research Ethics Committee approved the study protocol (REC number: 07/Q2501/22).

Informed consent statement: All participants gave written, informed consent prior to study inclusion.

Conflict-of-interest statement: The authors declare no relevant conflicts of interest.

Open-Access: This article is an open-access article which was selected by an in-house editor and fully peer-reviewed by external reviewers. It is distributed in accordance with the Creative Commons Attribution Non Commercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this

work non-commercially, and license their derivative works on different terms, provided the original work is properly cited and the use is non-commercial. See: <http://creativecommons.org/licenses/by-nc/4.0/>

Manuscript source: Invited manuscript

Correspondence to: Sophie C Warner, PhD, Department of Cardiovascular Sciences, University of Leicester and National Institute for Health Research, Leicester Cardiovascular Biomedical Research Unit, Glenfield Hospital Groby Road, Leicester LE3 9QP, United Kingdom. scw27@le.ac.uk
Telephone: +44-116-2044778

Received: November 29, 2016

Peer-review started: December 2, 2016

First decision: February 17, 2017

Revised: June 20, 2017

Accepted: August 15, 2017

Article in press: August 16, 2017

Published online: October 18, 2017

Abstract

AIM

To investigate a comprehensive range of factors that contribute to long-term patient satisfaction post-total joint replacement (TJR) in people who had undergone knee or hip replacement for osteoarthritis.

METHODS

Participants ($n = 1151$) were recruited from Nottinghamshire post-total hip or knee replacement. Questionnaire assessment included medication use, the pain-DETECT questionnaire (PDQ) to assess neuropathic pain-like symptoms (NP) and TJR satisfaction measured on average 4.8 years post-TJR. Individual factors were tested for an association with post-TJR satisfaction, before incorporating all factors into a full model. Data reduction was carried out using LASSO and receiver

operator characteristic (ROC) curve analysis was used to quantify the contribution of variables to post-TJR satisfaction.

RESULTS

After data reduction, the best fitting model for post-TJR satisfaction included various measures of pain, history of revision surgery, smoking, pre-surgical X-ray severity, WOMAC function scores and various comorbidities. ROC analysis of this model gave AUC = 0.83 (95%CI: 0.80-0.85). PDQ scores were found to capture much of the variation in post-TJR satisfaction outcomes: AUC = 0.79 (0.75-0.82). Pre-surgical radiographic severity was associated with higher post-TJR satisfaction: OR_{satisfied} = 2.06 (95%CI: 1.15-3.69), $P = 0.015$.

CONCLUSION

These results highlight the importance of pre-surgical radiographic severity, post-TJR function, analgesic medication use and NP in terms of post-TJR satisfaction. The PDQ appears to be a useful tool in capturing factors that contribute to post-TJR satisfaction.

Key words: Osteoarthritis; Patient satisfaction; Total joint arthroplasty; Neuropathic pain; Surgery outcomes

© **The Author(s) 2017.** Published by Baishideng Publishing Group Inc. All rights reserved.

Core tip: The growing number of total joint replacement (TJR) surgeries performed worldwide every year means that research in this area has the potential to impact millions of people. These results highlight the importance of a number of factors with regards to post-TJR satisfaction. The pain-DETECT questionnaire for neuropathic pain-like symptoms (NP) appears to be a useful tool in capturing factors that contribute to post-TJR satisfaction. Individuals with NP pre- or post-TJR could be indicated using this short questionnaire and referred for further testing and treatment to improve outcomes at every stage of their osteoarthritis treatment process.

Warner SC, Richardson H, Jenkins W, Kurien T, Doherty M, Valdes AM. Neuropathic pain-like symptoms and pre-surgery radiographic severity contribute to patient satisfaction 4.8 years post-total joint replacement. *World J Orthop* 2017; 8(10): 761-769 Available from: URL: <http://www.wjgnet.com/2218-5836/full/v8/i10/761.htm> DOI: <http://dx.doi.org/10.5312/wjo.v8.i10.761>

INTRODUCTION

A total joint replacement (TJR) is the only treatment for clinically severe osteoarthritis (OA). A TJR should be considered in individuals with marked symptoms of OA which significantly limit activity and participation and reduce quality of life if conservative treatments (*e.g.*, exercise, weight loss if overweight, analgesic medication) are insufficient^[1]. In the United Kingdom alone 160000 TJR are performed every year^[2]. Generally very good

outcomes are reported post-TJR^[3] but pain can remain a concern for some individuals. According to one study, 27% of people who had undergone total hip replacement (THR) and 44% of people who had undergone a total knee replacement (TKR) had joint pain 3-4 years after surgery^[4]. This pain can be inflammatory, nociceptive or neuropathic in nature^[5].

Patient satisfaction post-TJR has been the subject of some studies^[6-10] which have focused only on pain and function post-TJR^[11]. Pre-operative radiographic severity, co-occurrence of painful conditions, a history of revision surgery, other comorbidities, and pain catastrophizing have also been linked to post-TJR outcomes in the literature but not all in the same cohort^[7,12-15].

Neuropathic pain-like symptoms (NP) are caused by changes or damage to the nervous system, which can result from chronic nociceptive input (as seen in chronic pain states) and nerve damage during surgery^[5,16,17]. NP has been reported in people with OA and post-TJR^[4,18]. However, to our knowledge, currently no studies have investigated the role of NP on patient satisfaction post-TJR.

As TJR is currently the only long-term treatment for OA, if its effectiveness can be improved with better understanding of the individual differences in post-operative outcomes, this must be addressed. Due to the high number of TJR carried out in the United Kingdom and worldwide, research in this area has the potential to impact many individuals.

The aim of the present study was to investigate a comprehensive range of factors that contribute to long-term patient satisfaction post-TJR in people who had undergone knee or hip replacement for OA.

MATERIALS AND METHODS

Participants

The North Nottinghamshire Research Ethics Committee approved the study protocol (REC number: 07/Q2501/22). Participants who had undergone a TJR for OA were recruited from secondary care in Nottinghamshire ($n = 1151$) and gave written, informed consent. All participants had symptomatic and radiographic OA prior to TJR surgery. Between 2008 and 2011, nurse-administered questionnaires were completed by participants ($n = 1219$) on average 18 mo after surgery. These questionnaires included information on demographic variables, pain scores, TJR satisfaction and medication use. A subsequent follow-up postal questionnaire was sent to those who consented to further involvement in the study. This questionnaire was very similar in design to the baseline questionnaire. There was an average of 3.3 years between the first and second questionnaires. When the baseline and follow-up responses of participants who completed both questionnaires were compared there were no significant differences in age ($P < 0.38$), sex ($P < 0.89$), BMI ($P < 0.07$) or WOMAC pain scores ($P < 0.51$). There was not a significant difference in satisfaction levels ($P =$

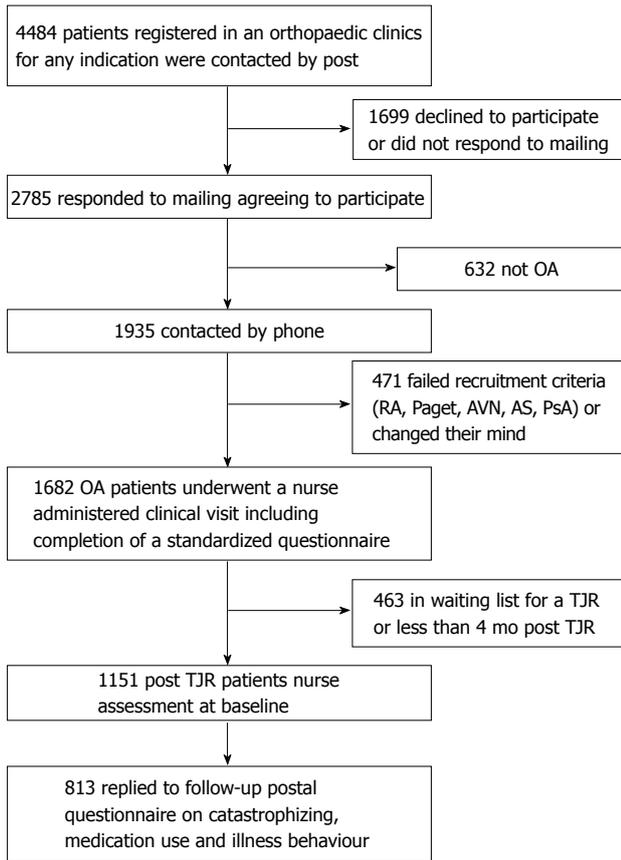


Figure 1 Flow-chart of participant recruitment for the current study.

0.22). Individuals had not been phenotyped for pain pre-surgery but pre-operative radiographic severity grade has been linked previously to TJR outcomes^[12]. The study design is presented in Figure 1.

Statistical analysis

Statistical review of this study was performed by a biomedical statistician. The statistics package R (version 3.0.2) was used to run logistic regression analyses to measure associations between TJR satisfaction and potential risk factors.

To select the risk factors contributing to post-TJR satisfaction, the least absolute shrinkage and selection operator method (LASSO) was used. LASSO is a feature selection method which, given a set of input measurements and an outcome measurement (in this case post-TJR satisfaction), fits a linear model^[19]. We employed a LASSO-regularised regression model as implemented by the R package "glmnet"^[20] (<http://cran.r-project.org/web/packages/glmnet/index.html>) using a logistic link function and the fitted LASSO coefficients derived were used.

Receiver operating characteristic (ROC) analysis was used to quantify the contribution of variables in the above models. The discrimination ability of the models was examined using the "PredictABEL" package for R (<http://cran.r-project.org/web/packages/PredictABEL/index.html>).

Trait definitions for statistical analysis

Neuropathic pain-like symptoms at the operated joints (knees or hips): The PDQ is a validated instrument for assessing NP. Scores range from 0-35, with > 12 indicating possible NP and ≥ 19 indicating likely NP^[21].

Pain severity: A visual analogue scale (VAS) was used to categorise individuals with high or low pain intensity at the operated joint (knee or hip). Scores range from 0-10, with ≥ 6 used to categorise high pain intensity.

TJR satisfaction: Individuals were asked to state how satisfied they felt with their TJR using an ordinal scale of "very satisfied", "not very satisfied" and "dissatisfied". For logistic regression analysis, individuals were dichotomised between: (1) "very satisfied"; and (2) "not very satisfied" and "dissatisfied".

Radiographic severity: The extent of joint damage evident by X-rays was categorised by assessment of pre-surgery knee and hip radiographs by a single observer. For knees, the Kellgren-Lawrence (K/L) grading system was used. Scores range from 0-4, with ≥ 3 classified as severe and 2 classified as not severe (K/L < 2 no OA)^[22]. An association with minimum joint space width (JSW) pre-surgery and pain post-surgery has been reported in a separate cohort^[12]. The minimum JSW was therefore used to classify hip OA, with minimum JSW ≤ 2.5 mm (which is a standard cut-off)^[23] being classified as radiographically severe. For bilateral surgery the joint with the most severe radiographic score was used.

Pain catastrophizing: The 13-item Pain Catastrophizing Scale (PCS) is a measure of the tendency to exaggerate the threat of a perceived harmful stimulus^[24]. Scores range from 0-52 and the highest tertile was used as a cut-off point to classify individuals as high catastrophizers, as previously described^[25].

WOMAC pain, stiffness and function: The OA-specific Western Ontario and McMaster Universities Arthritis Index (WOMAC) questionnaire includes questions about pain (scored from 0-20), stiffness (scored from 0-8) and function (scored from 0-76)^[26]. WOMAC function scores were categorised according to an OMERACT-defined PASS score of "acceptable function" (a score of ≤ 22) to allow a clinical guideline to be used in this study to put the importance of post-TJR satisfaction into a clinical context^[27,28].

Medication use

Questionnaire responses were used to classify participants as taking over-the-counter analgesics (OTC), opioids, non-steroidal anti-inflammatory drugs (NSAIDs) or other prescription medications which can be used to treat pain, as previously described^[25].

Table 1 Descriptive statistics categorised by total joint replacement satisfaction status and their contribution to the risk of dissatisfaction post-total joint replacement

Trait		Very satisfied (n = 861)	Not very satisfied (n = 227)	Dissatisfied (n = 63)	OR not very satisfied/ dissatisfied (95%CI) ¹
Demographic and morphometric	Age ± SD (yr)	73.2 ± 8.6	73.0 ± 8.8	72.2 ± 9.1	1.00 (0.98-1.01)
	% female	57.4	56.8	47.6	0.85 (0.65-1.12)
	BMI ± SD (kg/m ²)	29 ± 5.2	29.4 ± 5.1	30.7 ± 5.9	1.03 (1.00-1.06) ^a
Type of surgery	THR (n = 494)	407 (82.4%)	74 (15.0%)	13 (2.6%)	0.58 (0.44-0.77) ^e
	TKR (n = 591)	410 (69.4%)	136 (23.0%)	45 (7.6%)	2.02 (1.50-2.71) ^e
	THR + TKR (n = 66)	44 (66.7%)	17 (25.8%)	5 (7.6%)	1.63 (0.95-2.78)
Pre-operative X-ray	Years since most recent surgery	4.26	3.96	4.58	0.99 (0.96-1.03)
	Radiographically severe OA	92.10%	93.40%	96.50%	0.49 (0.27-0.87) ^a
History of surgery	Previous arthroscopic knee surgery ³	20.00%	26.40%	34.90%	1.65 (1.21-2.25) ^e
	Revision surgery	5.70%	9.30%	14.30%	2.36 (1.44-3.86) ^e
Psychological	% depression	15.9	22.0	28.6	1.64 (1.17-2.30) ^e
	PCS score (0-52)	8.2	12.8	19.7	1.06 (1.05-1.08) ^e
	Top tertile of PCS ²	20.80%	37.00%	55.60%	3.40 (2.52-4.59) ^e
Use of medication	% opioid	21.7	39.5	41.3	2.37 (1.77-3.18) ^e
	% OTC	49.0	64.5	61.9	1.33 (0.84-2.12)
	% NSAIDs	7.8	12.3	3.2	1.83 (1.38-2.42) ^e
	% other prescription analgesics	12.2	20.0	23.8	1.85 (1.29-2.66) ^e
Measures of pain	PDQ score (0-35)	4.8	10.1	14.3	1.15 (1.13-1.18) ^e
	Possible Neuropathic Pain (PDQ > 12)	10.00%	33.90%	57.10%	5.91 (4.22-8.29) ^e
	Likely Neuropathic Pain (PDQ ≥ 19)	6.50%	18.10%	34.90%	7.66 (4.80-12.22) ^e
	VAS (0-10)	3.1	5.8	7.0	1.35 (1.29-1.41) ^e
	HighVAS (> 5)	30.80%	61.20%	76.20%	6.47 (4.80-8.73) ^e
	WOMAC pain (0-20)	5.2	8.5	10.9	1.28 (1.23-1.33) ^e
	WOMAC stiffness (0-8)	2.9	4.1	4.4	1.62 (1.49-1.76) ^e
Comorbidities	WOMAC function (0-76)	25.7	38.0	47.8	1.07 (1.06-1.08) ^e
	% heart disease/angina	16.7	19.4	27.0	1.34 (0.95-1.89)
	% stroke	5.1	9.3	12.7	2.09 (1.26-3.44) ^e
	% hypertension	52.3	50.2	57.1	0.95 (0.72-1.25)
	% asthma/COPD	13.8	15.4	14.3	1.07 (0.73-1.57)
	% irritable bowel syndrome	10.2	14.5	11.1	1.44 (0.96-2.17)
	% diabetes	11.8	15.0	19.0	1.28 (0.86-1.90)
	% gout	7.5	11.9	11.1	1.50 (0.95-2.37)
	% osteoporosis	11.0	10.1	19.0	1.23 (0.80-1.88)
	% cancer	15.9	19.8	17.5	1.29 (0.91-1.83)
% current smoker	6.5	11.0	14.3	1.93 (1.22-3.07) ^e	

¹All ORs are adjusted for age, sex and BMI; ²Individuals in the top tertile of scores for the PCS questionnaire were classified as high catastrophizing; ³This classification includes any previous arthroscopic knee surgery. ^a*P* < 0.05, ^e*P* < 0.01, ^e*P* < 0.001. PCS: Pain Catastrophizing Scale; BMI: Body mass index.

Comorbidities

Comorbid conditions are commonly seen in people with OA, and people with OA are more likely to develop comorbid conditions such as cardiovascular disease and diabetes^[29]. A list of comorbidities was included in the questionnaire. Participants were asked to indicate which of these conditions they had been previously diagnosed with by a doctor.

RESULTS

The descriptive characteristics, stratified by TJR satisfaction status, are shown in Table 1. One fourth of study participants (290) were dissatisfied or not very satisfied with the outcome of their surgery. The study was thus powered (80%, *P* < 0.05) to detect associations with odds ratios of 1.75 or higher for binary traits with a prevalence of 10% or higher in the satisfied group, such as neuropathic pain (Table 1).

On univariate analysis, the majority of the variables

tested were found to be significantly associated with satisfaction post-TJR. This includes a higher BMI, various measures of pain (such as PDQ scores, high pain intensity and WOMAC pain scores), WOMAC function scores and pain catastrophizing (Table 1). Additionally, THR participants reported higher levels of being very satisfied (82.4%) than TKR patients (69.4%) (Table 1). Some factors were highly correlated with each other, such as PDQ scores and high pain intensity, PDQ scores and WOMAC pain scores and WOMAC pain scores and high pain intensity (*P* < 0.001 for all).

Given the large number of factors associated, many of them correlated with each other, we performed data reduction, using LASSO to identify which factors remain important contributors to post-TJR, post-THR and post-TKR satisfaction. After data reduction, the factors that remained in all three groups were: BMI, WOMAC function scores, PDQ scores, high pain intensity, severe pre-surgery radiographic OA and a past stroke. The full results of these analyses are shown in Table 2. Some

Table 2 The best fitting and pain-DETECT questionnaire models showing the contribution of factors to post-total joint replacement, post-total hip replacement and post-total knee replacement satisfaction

	Full/best fitting model	pain-DETECT questionnaire scores
Total joint replacement satisfaction	2.207 + (-0.013-PCS) + (0.189-sex) + (-0.398-TKR) + (0.016-BMI) + (-0.027-WOMAC function) + (-0.042-WOMAC stiffness) + (-0.012-past knee surgery) + (-0.056-PDQ) + (-0.483-highVAS) + (-0.380-revision surgery) + (0.352-severe pre-surgical radiographic OA) + (-0.066-OTC) + (-0.113-opioid) + (-0.010-current smoker) + (-0.218-stroke) + (0.051-hypertension) + (0.062-IBS) + (-0.055-gout) + (-0.123-depression)	2.796 + (-0.142-PDQ) + (-0.015-age) + (0.382-sex) + (0.004-BMI)
Total hip replacement satisfaction	2.985 + (0.014-years since surgery) + (-0.018-age) + (0.036-BMI) + (-0.036-WOMAC function) + (0.599-past knee surgery) + (-0.018-PDQ) + (-0.725-high VAS) + (-0.723-revision surgery) + (0.779-severe pre-surgical radiographic OA) + (-0.248-OTC) + (-0.119-opioid) + (-0.429-other medications for pain relief) + (-0.442-current smoker) + (-0.014-stroke) + (-0.514-depression) + (-0.022-cancer)	4.788 + (-0.121-PDQ) + (-0.038-age) + (0.036-sex) + (0.008-BMI)
Total knee replacement satisfaction	2.425 + (-0.018-PCS) + (0.190-sex) + (0.00045-BMI) + (-0.015-WOMAC function) + (-0.130-WOMAC stiffness) + (-0.076-PDQ) + (-0.313-high VAS) + (0.009-severe pre-surgical radiographic OA) + (-0.143-stroke) + (0.072-hypertension) + (0.334-IBS) + (-0.120-gout)	1.072 + (-0.146-PDQ) + (0.003-age) + (0.467-sex) + (0.011-BMI)

differences were observed in the factors that contribute to satisfaction post-THR and post-TKR, most notably a history of a revision surgery for THR and the WOMAC stiffness score for TKR. In both cases PDQ and VAS scores contribute significantly after adjustment for all other factors.

Higher pre-operative radiographic severity was also significantly associated with increased odds of TJR satisfaction: $OR_{\text{satisfied}} = 2.06$ (1.15-3.69), $P = 0.015$.

It was investigated whether patient satisfaction was related to measures of healthcare usage, specifically the use of analgesic medication. Strong associations were found between dissatisfaction and an increased likelihood of the use of some prescription analgesics (opioids and other prescription medications which can be used to treat pain) and OTC analgesics, but not prescription NSAIDs (Table 1). After adjustment for possible NP, high pain catastrophizing and high pain intensity (VAS) these associations become non-significant except in the case of opioids and OTC analgesics [$OR_{\text{dissatisfied}} = 1.68$ (1.21-2.34), $P = 0.002$; $OR_{\text{dissatisfied}} = 1.44$ (1.06-1.97), $P = 0.020$, respectively].

Post-TJR satisfaction is strongly associated with a measure of acceptable function post-TJR, according to OMERACT-defined PASS scores in the literature^[27,28]. This definition of acceptable function, according to a clinical guideline, was a very strong contributor to post-TJR satisfaction in this study after adjusting for age, sex and BMI: $OR_{\text{satisfied}} = 9.88$ (95%CI: 6.58-14.85), $P < 0.001$. This association remained significant after further adjustment for possible NP, high pain catastrophizing and high pain intensity: $OR_{\text{satisfied}} = 4.82$ (95%CI: 3.08-7.55), $P < 0.001$.

With regards to comorbidities, a history of stroke was associated with an increased risk of dissatisfaction post-TJR, as was being a current smoker (vs ex-smokers and people who have never smoked); $P < 0.01$ for both, see Table 1.

It was quantified how much these models contribute to satisfaction. The results of ROC analysis of the best-fitting model for each surgery group are shown in

Table 2 and Figure 2. The results show that the list of identified factors explains an AUC of 0.83 of patient satisfaction for post-TJR, 0.84 for post-THR and 0.83 for post-TKR. This, however, includes a large number of factors and it was investigated whether one of the factors may capture the effects of most of the other factors.

Possible NP, classified using PDQ scores, was seen in 17.3% of participants in this study. However, in the dissatisfied group the prevalence of possible NP was 3.8 times higher than in the very satisfied group $OR_{\text{possNP}} = 5.91$ (4.22-8.29), $P < 0.001$ and the prevalence of likely NP was 5.4 times higher: $OR_{\text{likNP}} = 7.66$ (4.80-12.22), $P < 0.001$ (see Table 1). Possible NP was less common in THR than TKR participants (11.9% and 22.3%, respectively) (Table 1). Likely NP has been reported previously to be present only in a small proportion of individuals post-TJR^[4] using as a definition a PDQ > 19. However strong differences exist in satisfaction at various lower cut-offs which explains why pain-DETECT scores capture such a large proportion of patient satisfaction in these data (Figure 3).

Given this strong effect we hypothesised that PDQ scores, being strongly correlated with pre-surgery X-ray scores (Spearman's rho = -0.13, $P < 0.001$) and associated with post-TJR pain intensity [$OR_{\text{highpainintensity}} = 1.35$ (1.30-1.40), $P < 0.001$], may capture much of the variation in post-TJR satisfaction outcomes, and indeed we find that this is the case. According to ROC analysis of PDQ scores (adjusted for age, sex and BMI), there is a significant contribution to post-TJR, post-THR and post-TKR satisfaction when this model is used (Table 2 and Figure 2). AUC values of 0.75 and over were reached in all three groups, even without the inclusion of any of the other available measures.

DISCUSSION

This study incorporated a comprehensive range of factors and shows that a number of factors including pain, comorbidities, smoking, history of revision surgery

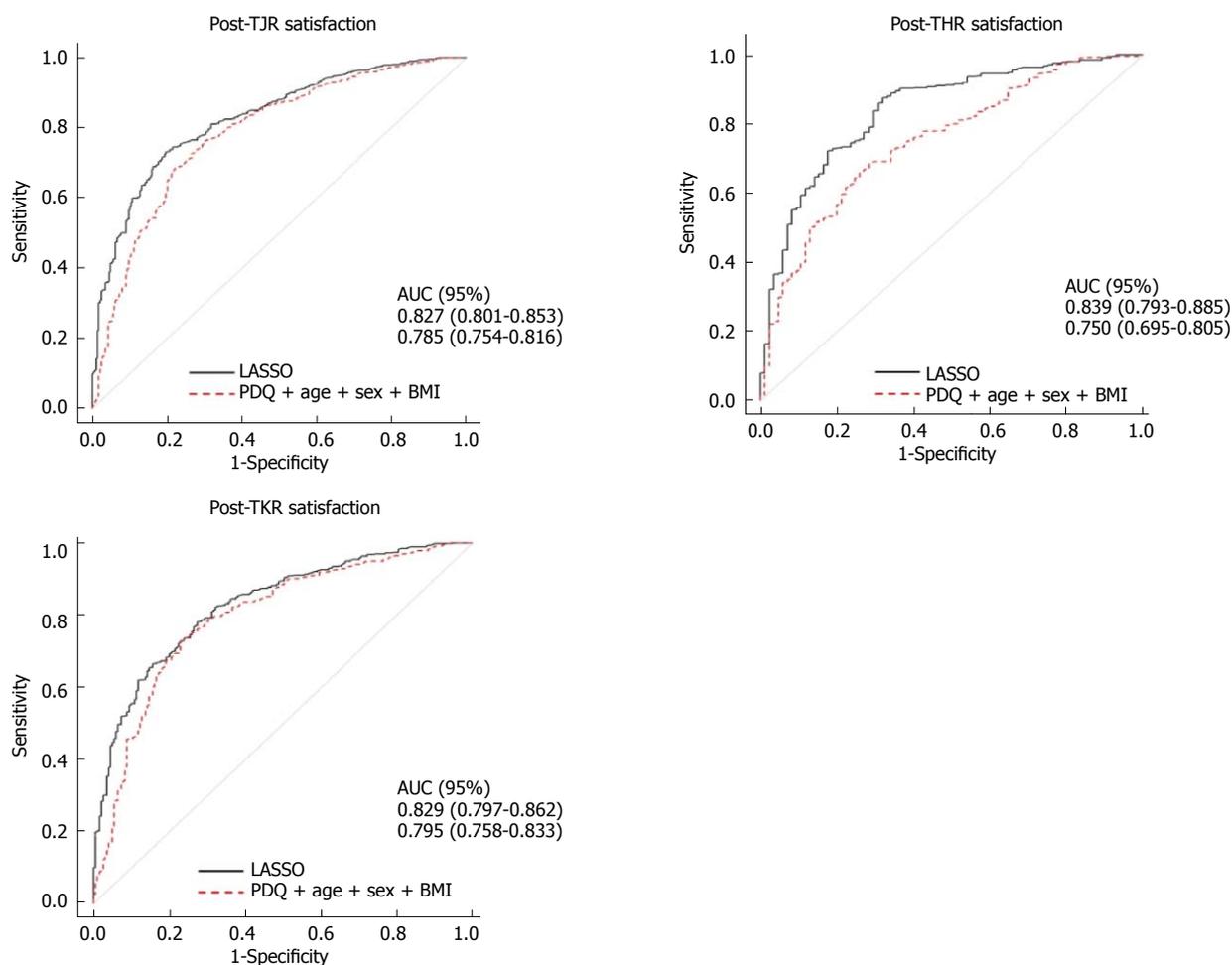


Figure 2 Receiver operator characteristic curves adjusted for age, sex and body mass index to show the amount of post-surgery satisfaction predicted by preoperative radiographic severity, pain-DETECT questionnaire scores and the best fit model. A: Post-TJR (THR and TKR combined); B: Post-TKR; C: Post-THR. TJR: Total joint replacement; THR: Total hip replacement; TKR: Total knee replacement.

and pre-surgical radiographic severity contribute to post-TJR satisfaction 4.8 years after surgery. Scores measuring the presence of NP appear to capture a large proportion of the variation seen.

Patient satisfaction is an outcome measure which is simple to use and accounts for the complex aspects of TJR^[30]. It has been recommended that patient satisfaction should be incorporated into assessments of post-TJR outcomes^[30]. Our results suggest that although post-TJR satisfaction is influenced by a large number of factors, it is well summarised by one single instrument, namely PDQ scores.

The proportion of possible NP identified in this study falls within the range reported by previous studies on NP post-TJR (reviewed in^[31]) particularly when differences in methodology and sample composition are taken into consideration^[4,15,32-37]. At first sight the importance of NP post-TJR detailed here appears to contrast with the report by Wylde *et al*^[4] who suggested that NP is a minor component of post-TJR pain.

The current data indicate that people who undergo TJR with only modest radiographic structural damage are more likely to report NP post-surgery. Although

this might suggest that the NP was also present pre-surgery, we lack the pre-operative pain assessments necessary to confirm if that is the case. In addition, pain may derive from other sources, such as bone marrow lesions, that are not evident on radiographs and may still be present post-surgery^[38]. Central nervous system involvement in OA, such as seen in NP, seems likely when the inconsistent correlation between pain and radiographic severity and the non-linear relationship between nociception and pain experienced are considered^[17] supporting the findings in this study.

In this study we fitted prediction models for patient dissatisfaction using all the contributing risk factors selected by LASSO. These models are fairly complicated in terms of the number of variables and therefore may not be applicable in clinical practice. However, we also show that PDQ scores have almost as much predictive value as the best fitting models. Therefore, in terms of clinical application our data suggest that assessing NP symptoms using the PDQ will help identify patients at highest risk of surgery dissatisfaction.

One key limitation to this study is the lack of pre-surgical pain data. However, Phillips *et al*^[39] found

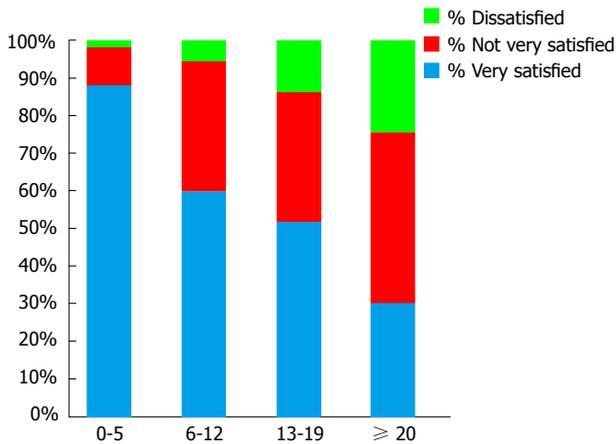


Figure 3 Proportion of post-total joint replacement patients reporting to be dissatisfied, not very satisfied or very satisfied depending on their pain-DETECT score.

that it was not possible to reliably predict post-TKR outcomes from pre-operative pain intensity and PDQ scores^[39], whereas Dualé *et al.*^[16] have reported a higher risk of NP post-surgery if peripheral NP is present pre-surgery. Although the self-administered PDQ allows data collection from a large number of individuals it does not provide definitive evidence of NP^[40]. Nonetheless, one study showed a correlation between PDQ scores and periaqueductal grey matter activation (which is involved in central sensitisation) in people with OA in areas of referred pain in response to punctate stimuli^[41].

Although in this study we did not use the widely accepted National Joint Registry agreed Patient Reported Outcomes (PROMS) data^[42], 92% of the questions in the Oxford hip and knee score (OXHS and OXKS, respectively) questionnaires are accounted for by the questionnaire used in this study, as was 83% of the content in the EQ-5D questionnaire. The questionnaire measured used in this study therefore reflects a large majority of the material covered in the PROMS. On the other hand we have examined other factors that are not usually included as part of post TJR PROMS, such as comorbidities, use of analgesic medication and pre-surgery X-ray severity all of which contribute to patient self-reported satisfaction in our data. To our knowledge this is one of the few studies to date which has looked at pain assessment integrated with comorbidities and use of medication.

Some of the factors identified as contributing to satisfaction could be addressed pre-surgery or considered when assessing outcomes post-surgery. The presence of comorbid conditions appears also to have a considerable effect on patient satisfaction, and this information may be used to manage patient expectations pre-surgery.

In conclusion, the PDQ appears to be particularly useful in capturing factors that contribute to post-TJR outcomes and may be considered as an important post-surgical assessment. These results also highlight the importance of understanding the mechanisms behind NP symptoms post-TJR, as it is a significant factor

contributing to post-TJR satisfaction and, importantly, affects a considerable proportion of individuals post-TJR.

COMMENTS

Background

Total joint replacement is a very common type of surgery. Understanding the determinants of patient satisfaction is necessary to address the increasing need for this type of surgery with population aging.

Research frontiers

The authors investigated for the first time the relationship between pre-operative radiographic severity and neuropathic pain symptoms and satisfaction post total joint replacement.

Innovations and breakthroughs

The authors show that neuropathic pain symptoms are the most important contributor to post-total joint replacement satisfaction. Other contributors are smoking and low pre-operative radiographic severity.

Applications

The prediction models used in this work can be applied to patients undergoing total joint replacement surgery for osteoarthritis.

Terminology

Neuropathic pain symptoms, caused by changes or damage to the nervous system. Pre-operative radiographic severity, refers to the extent of large joint (hip or knee) damage detected in X-rays prior to surgery.

Peer-review

This manuscript aims to evaluate factors predict satisfaction post total joint replacement. It is a vary serious paper dealing with the results of joints replacement. A well written paper and well organized study.

ACKNOWLEDGMENTS

The authors gratefully acknowledge the contributions of Sally Doherty and Maggie Wheeler to patient assessments at baseline, data collection and entry.

REFERENCES

- 1 National Institute for Health and Care Excellence. NICE Pathways: Management of osteoarthritis. [accessed 2016 May 1]. Available from: URL: <http://pathways.nice.org.uk/pathways/osteoarthritis#path=h=view%3A/pathways/osteoarthritis/management-of-osteoarthritis.xml&content=view-index>
- 2 National Joint Registry for England and Wales. 7th Annual Report. Hemel Hempstead: National Joint Registry for England and Wales; 2010. [accessed 2016 Jun 20]. Available from: URL: <http://www.njrcentre.org.uk/njrcentre/Portals/0/NJR%207th%20Annual%20Report%202010.pdf>
- 3 Losina E, Walensky RP, Kessler CL, Emrani PS, Reichmann WM, Wright EA, Holt HL, Solomon DH, Yelin E, Paltiel AD, Katz JN. Cost-effectiveness of total knee arthroplasty in the United States: patient risk and hospital volume. *Arch Intern Med* 2009; **169**: 1113-1121; discussion 1121-1122 [PMID: 19546411 DOI: 10.1001/archinternmed.2009.136]
- 4 Wylde V, Hewlett S, Learmonth ID, Dieppe P. Persistent pain after joint replacement: prevalence, sensory qualities, and postoperative determinants. *Pain* 2011; **152**: 566-572 [PMID: 21239114 DOI: 10.1016/j.pain.2010.11.023]
- 5 Kehlet H, Jensen TS, Woolf CJ. Persistent postsurgical pain: risk factors and prevention. *Lancet* 2006; **367**: 1618-1625 [PMID: 16698416 DOI: 10.1016/S0140-6736(06)68700-X]

- 6 **Dunbar MJ**, Haddad FS. Patient satisfaction after total knee replacement: new inroads. *Bone Joint J* 2014; **96-B**: 1285-1286 [PMID: 25274909 DOI: 10.1302/0301-620X.96B10.34981]
- 7 **Dunbar MJ**, Richardson G, Robertsson O. I can't get no satisfaction after my total knee replacement: rhymes and reasons. *Bone Joint J* 2013; **95-B**: 148-152 [PMID: 24187375 DOI: 10.1302/0301-620X.95B11.32767]
- 8 **Robertsson O**, Dunbar M, Pehrsson T, Knutson K, Lidgren L. Patient satisfaction after knee arthroplasty: a report on 27,372 knees operated on between 1981 and 1995 in Sweden. *Acta Orthop Scand* 2000; **71**: 262-267 [PMID: 10912927 DOI: 10.1080/000164700317411852]
- 9 **Ross CK**, Steward CA, Sinacore JM. A comparative study of seven measures of patient satisfaction. *Med Care* 1995; **33**: 392-406 [PMID: 7731280 DOI: 10.1097/00005650-199504000-00006]
- 10 **Carr AJ**, Robertsson O, Graves S, Price AJ, Arden NK, Judge A, Beard DJ. Knee replacement. *Lancet* 2012; **379**: 1331-1340 [PMID: 22398175 DOI: 10.1016/S0140-6736(11)60752-6]
- 11 **Clement N**. Patient factors that influence the outcome of total knee replacement: a critical review of the literature. *OA Orthopaedics* 2013; **1**: 11 [DOI: 10.13172/2052-9627-1-2-697]
- 12 **Valdes AM**, Doherty SA, Zhang W, Muir KR, Maciewicz RA, Doherty M. Inverse relationship between preoperative radiographic severity and postoperative pain in patients with osteoarthritis who have undergone total joint arthroplasty. *Semin Arthritis Rheum* 2012; **41**: 568-575 [PMID: 21868060 DOI: 10.1016/j.semarthrit.2011.07.002]
- 13 **Peter WF**, Dekker J, Nelissen RG, Fiocco M, van der Linden-van der Zwaag HM, Vermeulen EM, Tilbury C, Vliet Vlieland TP. Comorbidity in osteoarthritis patients following hip and knee joint replacement surgery. *Osteoarthr Cartil* 2014; **22**: S185-S6 [DOI: 10.1016/j.joca.2014.02.351]
- 14 **Edwards RR**, Haythornthwaite JA, Smith MT, Klick B, Katz JN. Catastrophizing and depressive symptoms as prospective predictors of outcomes following total knee replacement. *Pain Res Manag* 2009; **14**: 307-311 [PMID: 19714271 DOI: 10.1155/2009/273783]
- 15 **Nikolajsen L**, Brandsborg B, Lucht U, Jensen TS, Kehlet H. Chronic pain following total hip arthroplasty: a nationwide questionnaire study. *Acta Anaesthesiol Scand* 2006; **50**: 495-500 [PMID: 16548863 DOI: 10.1111/j.1399-6576.2006.00976.x]
- 16 **Dualé C**, Ouchchane L, Schoeffler P, EDONIS Investigating Group, Dubray C. Neuropathic aspects of persistent postsurgical pain: a French multicenter survey with a 6-month prospective follow-up. *J Pain* 2014; **15**: 24.e1-24.e20 [PMID: 24373573 DOI: 10.1016/j.jpain.2013.08.014]
- 17 **Graven-Nielsen T**, Wodehouse T, Langford RM, Arendt-Nielsen L, Kidd BL. Normalization of widespread hyperesthesia and facilitated spatial summation of deep-tissue pain in knee osteoarthritis patients after knee replacement. *Arthritis Rheum* 2012; **64**: 2907-2916 [PMID: 22421811 DOI: 10.1002/art.34466]
- 18 **Hochman JR**, Gagliese L, Davis AM, Hawker GA. Neuropathic pain symptoms in a community knee OA cohort. *Osteoarthritis Cartilage* 2011; **19**: 647-654 [PMID: 21440077 DOI: 10.1016/j.joca.2011.03.007]
- 19 **Tibshirani R**. Regression shrinkage and selection via the lasso: a retrospective. *J R Stat Soc: Series B (Stat Methodol)* 2011; **73**: 273-282 [DOI: 10.1111/j.1467-9868.2011.00771.x]
- 20 **Friedman J**, Hastie T, Tibshirani R. Regularization Paths for Generalized Linear Models via Coordinate Descent. *J Stat Softw* 2010; **33**: 1-22 [PMID: 20808728 DOI: 10.18637/jss.v033.i01]
- 21 **Freyenhagen R**, Baron R, Gockel U, Tölle TR. painDETECT: a new screening questionnaire to identify neuropathic components in patients with back pain. *Curr Med Res Opin* 2006; **22**: 1911-1920 [PMID: 17022849 DOI: 10.1185/030079906X132488]
- 22 **Kellgren JH**, Lawrence JS. Radiological assessment of osteoarthritis. *Ann Rheum Dis* 1957; **16**: 494-502 [PMID: 13498604]
- 23 **Croft P**, Cooper C, Wickham C, Coggon D. Defining osteoarthritis of the hip for epidemiologic studies. *Am J Epidemiol* 1990; **132**: 514-522 [PMID: 2389755 DOI: 10.1093/oxfordjournals.aje.a115687]
- 24 **Sullivan MJL**, Bishop SR, Pivik J. The Pain Catastrophizing Scale: Development and validation. *Psychol Assessment* 1995; **7**: 524-532 [DOI: 10.1037/1040-3590.7.4.524]
- 25 **Valdes AM**, Warner SC, Harvey HL, Fernandes GS, Doherty S, Jenkins W, Wheeler M, Doherty M. Use of prescription analgesic medication and pain catastrophizing after total joint replacement surgery. *Semin Arthritis Rheum* 2015; **45**: 150-155 [PMID: 26092331 DOI: 10.1016/j.semarthrit.2015.05.004]
- 26 **Bellamy N**, Buchanan WW, Goldsmith CH, Campbell J, Stitt LW. Validation study of WOMAC: a health status instrument for measuring clinically important patient relevant outcomes to antirheumatic drug therapy in patients with osteoarthritis of the hip or knee. *J Rheumatol* 1988; **15**: 1833-1840 [PMID: 3068365]
- 27 **Escobar A**, Gonzalez M, Quintana JM, Vrotsou K, Bilbao A, Herrera-Espiñeira C, Garcia-Perez L, Aizpuru F, Sarasqueta C. Patient acceptable symptom state and OMERACT-OARSI set of responder criteria in joint replacement. Identification of cut-off values. *Osteoarthritis Cartilage* 2012; **20**: 87-92 [PMID: 22155074 DOI: 10.1016/j.joca.2011.11.007]
- 28 **Maxwell JL**, Felson DT, Niu J, Wise B, Nevitt MC, Singh JA, Frey-Law L, Neogi T. Does clinically important change in function after knee replacement guarantee good absolute function? The multicenter osteoarthritis study. *J Rheumatol* 2014; **41**: 60-64 [PMID: 24293582 DOI: 10.3899/jrheum.130313]
- 29 **Stang PE**, Brandenburg NA, Lane MC, Merikangas KR, Von Korff MR, Kessler RC. Mental and physical comorbid conditions and days in role among persons with arthritis. *Psychosom Med* 2006; **68**: 152-158 [PMID: 16449426 DOI: 10.1097/01.psy.0000195821.25811.b4]
- 30 **Halawi MJ**. Outcome Measures in Total Joint Arthroplasty: Current Status, Challenges, and Future Directions. *Orthopedics* 2015; **38**: e685-e689 [PMID: 26270754 DOI: 10.3928/01477447-20150804-55]
- 31 **Haroutiunian S**, Nikolajsen L, Finnerup NB, Jensen TS. The neuropathic component in persistent postsurgical pain: a systematic literature review. *Pain* 2013; **154**: 95-102 [PMID: 23273105 DOI: 10.1016/j.pain.2012.09.010]
- 32 **Baker PN**, van der Meulen JH, Lewsey J, Gregg PJ; National Joint Registry for England and Wales. The role of pain and function in determining patient satisfaction after total knee replacement. Data from the National Joint Registry for England and Wales. *J Bone Joint Surg Br* 2007; **89**: 893-900 [PMID: 17673581 DOI: 10.1302/0301-620X.89B7.19091]
- 33 **Clarke H**, Kay J, Mitsakakis N, Katz J. Acute pain after total hip arthroplasty does not predict the development of chronic postsurgical pain 6 months later. *J Anesth* 2010; **24**: 537-543 [PMID: 20490573 DOI: 10.1007/s00540-010-0960-z]
- 34 **Martinez V**, Fletcher D, Bouhassira D, Sessler DI, Chauvin M. The evolution of primary hyperalgesia in orthopedic surgery: quantitative sensory testing and clinical evaluation before and after total knee arthroplasty. *Anesth Analg* 2007; **105**: 815-821 [PMID: 17717244 DOI: 10.1213/01.ane.0000278091.29062.63]
- 35 **Remérand F**, Le Tendre C, Baud A, Couvret C, Pourrat X, Favard L, Laffon M, Fusciardi J. The early and delayed analgesic effects of ketamine after total hip arthroplasty: a prospective, randomized, controlled, double-blind study. *Anesth Analg* 2009; **109**: 1963-1971 [PMID: 19923527 DOI: 10.1213/ANE.0b013e318181bdc8a0]
- 36 **Sanders JC**, Gerstein N, Torgeson E, Abram S. Intrathecal baclofen for postoperative analgesia after total knee arthroplasty. *J Clin Anesth* 2009; **21**: 486-492 [PMID: 20006256 DOI: 10.1016/j.jclinane.2008.12.019]
- 37 **Singh JA**, Lewallen D. Predictors of pain and use of pain medications following primary Total Hip Arthroplasty (THA): 5,707 THAs at 2-years and 3,289 THAs at 5-years. *BMC Musculoskelet Disord* 2010; **11**: 90 [PMID: 20462458 DOI: 10.1186/1471-2474-11-90]
- 38 **Kurien T**, Kerslake R, Haywood B, Pearson RG, Scammell BE. Resection and resorption of bone marrow lesions is associated with improvement of pain after knee replacement surgery. Liverpool, UK: British Orthopaedic Research Society, 2015
- 39 **Phillips JR**, Hopwood B, Arthur C, Stroud R, Toms AD. The natural history of pain and neuropathic pain after knee replacement: a prospective cohort study of the point prevalence of pain and neuropathic pain to a minimum three-year follow-up. *Bone Joint J* 2014; **96-B**: 1227-1233 [PMID: 25183595 DOI: 10.1302/0301-620X.

- 96B9.33756]
- 40 **Bouhassira D**, Attal N. Diagnosis and assessment of neuropathic pain: the saga of clinical tools. *Pain* 2011; **152**: S74-S83 [PMID: 21185120 DOI: 10.1016/j.pain.2010.11.027]
- 41 **Gwilym SE**, Keltner JR, Warnaby CE, Carr AJ, Chizh B, Chessell I, Tracey I. Psychophysical and functional imaging evidence supporting the presence of central sensitization in a cohort of osteoarthritis patients. *Arthritis Rheum* 2009; **61**: 1226-1234 [PMID: 19714588 DOI: 10.1002/art.24837]
- 42 **Health and Social Care Information Centre**. Monthly Patient Reported Outcome Measures (PROMs) in England. [accessed 2016 May 1]. Available from: URL: <http://www.hscic.gov.uk/proms>

P- Reviewer: Fisher DA, Fenichel I, Tangtrakulwanich B

S- Editor: Kong JX **L- Editor:** A **E- Editor:** Lu YJ





Published by **Baishideng Publishing Group Inc**
7901 Stoneridge Drive, Suite 501, Pleasanton, CA 94588, USA
Telephone: +1-925-223-8242
Fax: +1-925-223-8243
E-mail: bpgoffice@wjgnet.com
Help Desk: <http://www.f6publishing.com/helpdesk>
<http://www.wjgnet.com>

