

Patient-Reported Allergies, a Marker of Preoperative Pain and Disability in Elective Spine Surgery

Global Spine Journal 2020, Vol. 10(5) 592-597 © The Author(s) 2019 Article reuse guidelines: sagepub.com/journals-permissions DOI: 10.1177/2192568219865188 journals.sagepub.com/home/gsj



Keith W. Lyons, MD¹, Tracy M. Borsinger, MD¹, Adriana P. Lucas, MS¹, Kevin J. McGuire, MD, MS¹, Adam M. Pearson MD, MS¹, and William A. Abdu, MD, MS¹

Abstract

Study Design: Retrospective review.

Objective: Previous literature demonstrates mixed results regarding the relationship between patient-reported allergies and pain, function, and satisfaction scores. The objective of this study was to investigate the correlation between patient-reported allergies and preoperative Oswestry Disability Index (ODI), Neck Disability Index (NDI), and Patient-Reported Outcomes Measurement System (PROMIS) scores.

Methods: All patients undergoing elective cervical, lumbar procedures between May 2017 and October 2018 were included. Baseline demographic information was recorded, as well as all reported allergies or adverse reactions. Preoperative PROMIS, ODI, and NDI scores were recorded. Hierarchical multiple linear regressions were used to assess the relationship between total number of allergies and the preoperative pain and function scores.

Results: A total of 570 patients were included (476 lumbar, 94 cervical). The mean number of allergies reported was 1.89 \pm 2.32. The mean preoperative ODI and NDI scores were 46.39 \pm 17.67 and 43.47 \pm 16.51, respectively. The mean preoperative PROMIS Physical Health and PROMIS Mental Health scores were 37.21 + 6.54 and 43.89 + 9.26, respectively. Hierarchical multiple linear regression showed that total number of reported allergies shared a statistically significant negative relationship with all of the following scores: ODI (B = 0.83, P = .02), NDI (B = 1.45, P = .02), PROMIS Physical Health (B = -0.29, P = .013), and PROMIS Mental Health (B = -0.38, P = .024).

Conclusions: Patient-reported allergies share a statistically significant negative relationship with preoperative pain and function scores; as patients have increasing total number of allergies, the ODI/NDI scores become worse (increase) and the PROMIS scores become worse (decrease).

Keywords

patient-reported allergies, preoperative pain, preoperative function, spine surgery

Introduction

Over the past few years, there has been increasing literature demonstrating that patient-reported medication and environmental allergies and sensitivities can be used as a marker of increased pain, disability, and poor outcomes in elective surgery. Various subspecialties, such as hip and knee arthroplasty, have evaluated this and shown that patients with multiple reported allergies have less improvement in patient-reported outcomes postoperatively compared with patients with fewer allergies.¹⁻³ This topic has also been investigated in shoulder arthroplasty; however, the results have been varied. A recent study by Rosenthal et al revealed that the presence of one or more self-reported allergies did not correlate with worse

Corresponding Author:

Keith Lyons, Dartmouth-Hitchcock Medical Center, Department of Orthopaedics, One Medical Center Drive, Lebanon, NH 03756, USA. Email: keith.w.lyons@hitchcock.org



Dartmouth-Hitchcock Medical Center, Lebanon, NH, USA

Lyons et al 593

function, satisfaction, pain, or range of motion postoperatively. On the contrary, another recent study demonstrated that a greater number of patient-reported allergies correlated with severe postoperative pain in patients undergoing elective total shoulder arthroplasty. 5

To date, there are very few studies in spine surgery that examine patient allergy lists, and their corresponding function and pain scores. Graves et al found that patients with more than 3 allergies had less improvement in their Oswestry Disability Index (ODI) postoperatively compared with patients with less than 3 allergies; they concluded that the number of allergies may serve as a predictive marker for patients at risk for poorer outcomes after elective spine surgery. Similarly, Kay et al observed that patients with more than 5 reported allergies had worse pain and disability outcome scores. It has also been shown that the number of reported allergies is associated with both depression and anxiety in patients with lumbar spine disease.

On the contrary, the work of Xiong et al demonstrated that patient-reported allergies actually correlated with subjective improvement in pain and disability postoperatively. However, they found that this relationship did not persist longer than 1 year postoperatively, and they also found that a greater number of allergies corresponded with higher admissions costs and lengths of stay.⁹

As seen from these studies, the evidence is mixed regarding the association or correlation of patient-reported allergy lists and their pain and function scores on presentation for and after elective spine surgery. At this point, the evidence tends to point to the fact that patients with increasing numbers of allergies may have worse pain and function scores, as well as less improvement postoperatively. Another important factor is the role of underlying psychiatric comorbidities. Patientreported allergies correlate with underlying Diagnostic and Statistical Manual of Mental Disorders (DSM) Axis 1 psychological disorders, such as major depression, bipolar disorder, phobias. 10 Various Axis 1 diagnoses (depression, anxiety, somatization, panic disorders, etc) are also associated with poorer outcomes postoperatively and less satisfaction.¹¹ Patients with multiple drug intolerances also tend to have higher levels of anxiety, depression, and somatization. 12,13 Therefore, allergies may be associated with, or even serve as a surrogate for, Axis 1 disorders.

Patient-reported allergy lists can certainly be used as an important piece of data in the elective surgery setting. As reimbursement methods tend to transition to quality-driven models, outcome scores and patient satisfaction become increasingly important. Identifying those at risk and subsequently considering them for and optimizing them prior to elective surgery is key. Therefore, we sought to evaluate patient allergy lists to see how they correlated with preoperative pain and function scores. We hypothesized that patients with multiple reported allergies would have lower preoperative Patient-Reported Outcomes Measurement System (PROMIS) scores, as well as worse preoperative Neck Disability Index (NDI) and ODI scores.

Materials and Methods

Institutional review board approval was obtained for this project; formal consent was waived as this was a retrospective study with minimal risk to subjects. Furthermore, the patientreported outcome surveys that we used all involved obtaining consent at the time of the initial survey. All patients who had undergone elective lumbar, thoracolumbar, and cervical spine surgery from May 1, 2017, through October 18, 2018, were included in this study, which was conducted at the Spine Center at the Dartmouth-Hitchcock Medical Center, a spine referral center. This time frame was determined based on when the Spine Center started using PROMIS scores. We excluded all fracture, malignancy, and infection cases. We excluded patients who did not complete the preoperative function/pain surveys. A retrospective chart review was then performed of the preoperative association of patient-reported allergies and sensitivities and patient-reported outcomes at baseline.

The PROMIS is a questionnaire for patients that assess their pain and function. PROMIS is a 2-part (Physical Health and Mental Health) questionnaire for patients that assess their pain and function. Numerous studies in the spine literature have demonstrated that both are valid and responsive questionnaires. ¹⁴⁻¹⁶ The ODI and the NDI are 2 other commonly used questionnaires that have been in use since the early 1980s and are validated as useful clinical tools representing pain and disability. ^{17,18}

The preoperative PROMIS scores were recorded, as well as the preoperative ODI and NDI scores. Baseline demographic information (age, sex, body mass index [BMI], American Society of Anesthesiologists [ASA] score, and smoking status) as well as any documented *DSM* Axis I diagnoses were collected through a retrospective chart review. The total number of listed allergies/adverse drug reactions was recorded. This was broken down into 2 groups, medication allergies and environment/food allergies.

All analyses were conducted in version 3.5.1 of the R environment (R Development Core Team, 2013). Hierarchical multiple linear regression was used to assess the relationship between total number of allergies and the preoperative ODI, NDI, PROMIS Mental Health, and PROMIS Physical Health scores after controlling for relevant demographics. Hierarchical multiple linear regression was used to demonstrate how an independent variable of interest (in this case allergies) explains or relates to variance in a dependent variable (PROMIS scores and ODI/NDI scores). 19,20 In this statistical method, a model is built that includes numerous independent variables, such as the various patient demographics (age, sex, BMI, etc). Then, a second model is created that introduces another variable (allergies). The models are then compared, and if the difference in \mathbb{R}^2 between the 2 models is statistically significant, the variable that was introduced in the second model is assumed to explain the difference.

Results

A total of 923 patients underwent elective spine surgery between May 2017 and October 2018. All cases of infection, tumor, and fracture were excluded. Patients who did not 594 Global Spine Journal 10(5)

Table 1. Baseline Characteristics of Patients Undergoing Elective Spine Surgery from May 2017-October 2018.

Characteristics	Mean (\pm SD) or Sum (%)		
N	570		
Age	58.06 (±15.02)		
BMI	30.20 (±6.12)		
ASA	$2.38\ (\pm 0.59)$		
Sex	, ,		
Male (%)	329 (57.7)		
Female (%)	241 (43.3)		
Current smoker (%)	92 (16.1)		
Surgery			
Thoracolumbar (%)	476 (83.5)		
Cervical (%)	94 (16.5)		
Allergies			
Total	1.89 (±2.32)		
Medications	1.49 (±2.00)		
Environmental/food	$0.43 \ (\pm 0.91)$		
Presence of Axis I disorder(s) (%)	191 (33.5)		
Preoperative ODI	46.39 (± 17.67)		
Preoperative NDI	43.47 (<u>+</u> 16.51)		
Preoperative PROMIS Physical Health	37.21 $(\pm 6.54)^{'}$		
Preoperative PROMIS Mental Health	43.89 (±9.26)		

Abbreviations: BMI, body mass index; ASA, American Society of Anesthesiologists score; ODI, Oswestry Disability Index; NDI, Neck Disability Index; PROMIS, Patient-Reported Outcomes Measurement System.

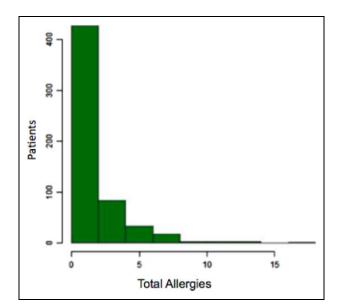


Figure 1. Distribution of allergies.

complete preoperative PROMIS, ODI, or NDI questionnaires were also excluded. A total of 570 patients met the inclusion criteria, and their baseline characteristics are presented in Table 1. The distribution of the number of allergies per patient is demonstrated in Figure 1. Hierarchical multiple linear regression and then ANOVA were used to assess the relationship between allergies and preoperative ODI (Table 2), NDI (Table 3), PROMIS Mental Health (Table 4), and PROMIS

Table 2. Hierarchical Multiple Regression Models with Dependent Variable (DV) ODI Scores.

DV: Preoperative ODI Scores				
Predictors	Estimates	CI	Estimates	CI
(Intercept)	47.88***	37.12 to 58.65	46.97***	36.23 to 57.71
Age	-0.21***	-0.32 to -0.09	-0.22***	-0.33 to -0.11
Sex ^a	-1.58	-4.75 to 1.59	-0.49	-3.78 to 2.80
BMI	0.15	-0.11 to 0.41	0.16	-0.10 to 0.42
ASA	2.4	-0.56 to 5.37	2.15	-0.81 to 5.11
Current smoker (Yes ^b)	9.89***	5.36 to 14.41	9.95***	5.45 to 14.45
Axis I disorders (Yes ^b)	0.77	-2.58 to 4.12	0.32	-3.03 to 3.67
Total allergies			0.83*	0.13 to 1.52
Observations		472		4 72
$R^{2} (\Delta R^{2})$ $\Delta \chi^{2} = 5.4067^{*}$		0.09		0.1

Abbreviations: DV, dependent variable; ODI, Oswestry Disability Index; CI, confidence interval; BMI, body mass index; ASA, American Society of Anesthesiologists score.

Physical Health (Table 5). All regressions demonstrated that there was a statistically significant relationship between total number of allergies and these pain/function scores. As the number of allergies increased, the preoperative ODI and NDI scores increased (patients had increased pain and disability) and the preoperative PROMIS scores decreased (patients had worse mental and physical health). Since the magnitude of the effect was small, as demonstrated by the ΔR^2 metric, a χ^2 test for nested models was employed to investigate whether model fit significantly improved with the introduction of the allergy predictor. These results showed that allergies did account for the variance and that the variance was statistically significant (P < .05).

As mentioned above, 4 nested models were analyzed to understand the impact of allergies on preoperative scores over and above age, sex, BMI, ASA score, smoking status, and the presence of DSM-IV Axis I disorders. The first included preoperative ODI as the dependent variable (Table 2). ODI is a function score that assesses low back pain and disability; therefore, it was used for the subset of patients undergoing lumbar and thoracolumbar procedures. As the number of reported allergies increased by 1, the ODI score increased by 0.83, which was statistically significant (B = 0.83, P = .02). The inclusion of number of allergies significantly increased the variance explained in the ODI dependent variable ($\chi^2 = 5.4067$, P = .02049).

The second model included preoperative NDI as the dependent variable (Table 3). This was based on the subset of patients who had undergone cervical procedures. The NDI is a version of the ODI, but it is focused on neck pain and disability, rather than low back pain. Therefore, it was used for

^aFemale is referent.

^bNo is referent

^{*}P < .05. **P = .001. ***P < .001.

Lyons et al 595

Table 3. Hierarchical Multiple Regression Models with Dependent Variable (DV) NDI Scores.

DV: Preoperative NDI Scores				
Predictors	Estimates	CI	Estimates	CI
(Intercept)	51.6***	31.95 to 71.24	53.23***	33.96 to 72.50
Age	-0.46***	-0.70 to -0.22	-0.48***	-0.72 to -0.24
Sex ^a	3.14	-2.99 to 9.28	4.22	-1.85 to 10.29
BMI	0.13	-0.33 to 0.59	0.11	-0.34 to 0.56
ASA	4.38	-1.11 to 9.87	3.26	-2.19 to 8.72
Current smoker (Yes ^b)	10.59*	3.96 to 17.23	10.57*	4.08 to 17.06
Axis disorders (Yes ^b)	0.38	-5.76 to 6.52	−I.64	-7.91 to 4.62
Total allergies			1.45*	0.17 to 2.73
Observations		94		94
$R^{2} (\Delta R^{2})$ $\Delta \chi^{2} = 4.9436^{*}$		0.242		0.283

Abbreviations: DV, dependent variable; NDI, Neck Disability Index; CI, confidence interval; BMI, body mass index; ASA, American Society of Anesthesiologists score.

Table 4. Hierarchical Multiple Regression Models with Dependent Variable (DV) PROMIS Physical Health Scores.

DV: PROMIS Physical Health				
Predictors	Estimates	CI	Estimates	CI
(Intercept)	46.44***	37.15 to 55.73	46.85***	37.59 to 56.10
Age	0.06**	0.02 to 0.10	0.07**	0.03 to 0.11
Sex ^a	0.19	-0.86 to 1.25	-0.17	-1.27 to 0.92
BMI	-0.12**	-0.21 to -0.03	-0.12**	-0.21 to -0.04
ASA	-1.80***	-2.79 to -0.82	−1. 7 **	-2.68 to -0.72
Current smoker (Yes ^b)	−3.92***	−5.35 to −2.48	-3.93***	−5.36 to −2.51
Thoracolumbar	-4.23	-12.83 to 4.37	-4.41	-12.97 to 4.15
Cervical	-3.68	-12.35 to 5.00	-3.83	-12.47 to 4.81
Axis I disorders (Yes ^b)	-1.01	-2.13 to 0.10	-0.82	-1.94 to 0.30
Total allergies			-0.29*	-0.53 to -0.06
Observations		568		568
R^2 (ΔR^2) $\Delta \chi^2 = 6.1524^*$		0.123	0.	133 (0.01)

Abbreviations: DV, dependent variable; PROMIS, Patient-Reported Outcomes Measurement System; CI, confidence interval; BMI, body mass index; ASA, American Society of Anesthesiologists score.

the subset of patients who underwent cervical procedures. As the number of allergies increased by 1, the NDI increased by 1.45, which was statistically significant (B = 1.45, P = .02). The inclusion of number of allergies significantly increased the variance explained in the NDI dependent variable ($\chi^2 = 4.9436$, P = .02881).

The third model included preoperative PROMIS Physical Health as the dependent variable (Table 4). This included all

Table 5. Hierarchical Multiple Regression Models with Dependent Variable (DV) PROMIS Mental Health Scores.

DV: PROMIS Mental Health Scores				
Predictors	Estimates	CI	Estimates	CI
(Intercept)	43.97***	30.72 to 57.23	44.5***	31.29 to 57.72
Age	0.09**	0.04 to 0.15	0.1**	0.04 to 0.15
Sex ^a	0.28	-1.23 to 1.79	-0.2	-1.76 to 1.36
BMI	0.02	-0.10 to 0.15	0.02	-0.10 to 0.14
ASA	-2.56***	-3.97 to -1.16	-2.42**	-3.83 to -1.02
Current smoker (Yes ^b)	−4.19 ***	−6.24 to −2.15	-4.2I***	−6.25 to −2.17
Thoracolumbar	1.72	-10.55 to 13.98	1.48	-10.74 to 13.70
Cervical	1.22	-11.16 to 13.60	1.02	-11.32 to 13.35
Axis I disorders (Yes ^b)	−3.11****	−4.69 to −1.52	-2.85***	−4.45 to −1.26
Total allergies			-0.38*	-0.71 to -0.05
Observations		568		568
$R^{2} (\Delta R^{2})$ $\Delta \chi^{2} = 5.0934^{*}$		0.109		0.118

Abbreviations: DV, dependent variable; PROMIS, Patient-Reported Outcomes Measurement System; CI, confidence interval; BMI, body mass index; ASA, American Society of Anesthesiologists score.

570 patients in the study (cervical and lumbar surgeries). As the number of allergies increased by 1, the PROMIS Physical Health score decreased by 0.29, which was statistically significant (B = -0.29, P = .013). The inclusion of number of allergies significantly increased the variance explained in the PROMIS Physical Health dependent variable ($\chi^2 = 6.1524$, P = .01342).

The fourth model included preoperative PROMIS Mental Health as the dependent variable (Table 5). This included all 570 patients in the study (cervical and lumbar surgeries). As the number of allergies increased by 1, the PROMIS Mental Health score decreased by 0.38, which was statistically significant (B=-0.38, P=.024). The inclusion of number of allergies significantly increased the variance explained in the PROMIS Mental Health dependent variable ($\chi^2=5.0934$, P=.0244).

Discussion

This study shows that within our patient population, patient-reported allergies shared a statistically significant negative relationship with preoperative pain and disability scores. In other words, with increasing numbers of allergies, patients have worse (higher) ODI and NDI scores and worse (lower) PROMIS mental and physical health scores.

As seen in previous studies, there is an association with increasing amounts of patient reported allergies and poorer pain and disability after elective lumbar spine surgeries.^{6,7} A similar association has also been seen in multiple investigations in hip and knee arthroplasty.^{1,6,21}

Multiple self-reported allergies are also associated with depression and anxiety. ¹⁰ Studies have shown that patients with

^aFemale is referent.

^bNo is referent.

^{*}P < .05. **P = .001. ***P < .001.

^aFemale is referent.

^bNo is referent.

^{*}P < .05. **P = .001. ***P < .001.

^aFemale is referent.

^bNo is referent.

^{*}P < .05. **P = .001. ***P < .001.

596 Global Spine Journal 10(5)

psychological distress undergoing lower extremity arthroplasty have worse self-perceived preoperative and postoperative function and disability scores. ²² This raises the question as to why multiple drug allergies, sensitivities, or multiple drug intolerance syndrome (3 or more unrelated drug class allergies) is associated with poorer preoperative function. One reason may be that patients with multiple allergies tend to be older, overweight, and have higher rates of health care and medication usage. ^{23,24}

As the economic landscape continues to change, reimbursement models are increasingly focusing on quality and patient-reported outcomes. Preoperative screening and identification of risk factors has also become a critical factor in elective surgery; poor pain control and function have strong correlations with numerous psychological and behavioral characteristics that may be evaluated in the preoperative setting. Just as smoking, uncontrolled diabetes, obesity, and liability issues are conditions that can adversely impact surgical outcomes in spine patients, multiple allergies or sensitivities to both medications and environmental factors may play a similar role. An improved understanding of these factors may help establish methods or pathways to counsel and manage these patients prior to considering surgical options.

This study does have some limitations. The distribution of total allergies in our patient population was skewed to the left. Although this is a nonsymmetric distribution, our sample size was large and this distribution is representative of our population. Second, all allergies were included as reported by and confirmed by the patient to the intake nurse, and therefore not all were necessarily confirmed allergies. The patients' perception of the burden of sensitivities and allergies may in fact be the true correlate with outcomes, not whether or not the allergy is real or not. Also, additional analyses evaluating whether or not these relationships also affect postoperative scores will be important in evaluating the role of patient allergies and our assessments of their pain and function.

Declaration of Conflicting Interests

The author(s) declared no potential conflicts of interest with respect to the research, authorship, and/or publication of this article.

Funding

The author(s) received no financial support for the research, authorship, and/or publication of this article.

ORCID iD

Keith W. Lyons, MD https://orcid.org/0000-0002-0518-5847 Kevin J. McGuire, MD, MS https://orcid.org/0000-0003-1069-4790

Ethical Approval

This study was approved by our institution's institutional review board committee, CPHS# STUDY00031299.

References

- Hinarejos P, Ferrer T, Leal J, Torres-Claramunt R, Sánchez-Soler J, Monllau JC. Patient-reported allergies cause inferior outcomes after total knee arthroplasty. *Knee Surg Sports Traumatol Arthrosc.* 2016;24:3242-3246. doi:10.1007/s00167-015-3837-8
- Otero JE, Graves CM, Gao Y, et al. Patient-reported allergies predict worse outcomes after hip and knee arthroplasty: results from a prospective cohort study. *J Arthroplasty*. 2016;31: 2746-2749. doi:10.1016/j.arth.2016.07.040
- Bourne RB, Chesworth BM, Davis AM, Mahomed NN, Charron KDJ. Patient satisfaction after total knee arthroplasty: who is satisfied and who is not? *Clin Orthop Relat Res*. 2010;468: 57-63. doi:10.1007/s11999-009-1119-9
- Rosenthal BD, Knesek MJ, Kahlenberg CA, Mai H, Saltzman MD. Shoulder arthroplasty outcomes in patients with multiple reported drug allergies: does number of drug allergies have an effect on outcome? *Orthop J Sport Med.* 2016;4: 2325967116671501. doi:10.1177/2325967116671501
- Menendez ME, Lawler SM, Ring D, Jawa A. High pain intensity after total shoulder arthroplasty. *J Shoulder Elbow Surg*. 2018;27: 2113-2119. doi:10.1016/j.jse.2018.08.001
- Graves CM, Carreon LY, Gvozdyev B, Riley S, Gum JL, Glassman SD. Multiple patient reported allergies are associated with worse outcomes following lumbar spine surgery. *Spine J.* 2016; 16:S363-S364.
- Kay HF, Chotai S, Wick J, et al. Does number of reported drug allergies affect patient-reported outcomes and satisfaction following operative treatment for degenerative lumbar spine disease? *Spine J.* 2015;15:S94. doi:10.1016/j.spinee.2015.07.031
- Levin JM, Boyle S, Winkelman RD, et al. Patient-reported allergies are associated with preoperative psychological distress and less satisfying patient experience in a lumbar spine surgery population. *Clin Spine Surg*. 2018;31:E368-E374.
- Xiong DD, Ye W, Xiao R, et al. Patient-reported allergies predict postoperative outcomes and psychosomatic markers after spine surgery. Spine J. 2019;19:21-130. doi:10.1016/j.spinee. 2018.05.032
- Patten SB, Williams JVA. Self-reported allergies and their relationship to several axis I disorders in a community sample. *Int J Psychiatry Med.* 2007;37:11-22. doi:10.2190/L811-0738-10NG-7157
- Greenley JR, Young TB, Schoenherr RA. Psychological distress and patient satisfaction. *Med Care*. 1982;20:373-385.
- De Pasquale T, Nucera E, Boccascino R, et al. Allergy and psychologic evaluations of patients with multiple drug intolerance syndrome. *Intern Emerg Med*. 2012;7:41-47. doi:10.1007/s11739-011-0510-1
- Hassel JC, Danner D, Hassel AJ. Psychosomatic or allergic symptoms? High levels for somatization in patients with drug intolerance. *J Dermatol*. 2011;38:959-965. doi:10.1111/j.1346-8138. 2011.01249.x
- Patel AA, Dodwad SM, Boody BS, et al. Validation of Patient Reported Outcomes Measurement Information System (PRO-MIS) computer adaptive tests (CATs) in the surgical treatment of lumbar spinal stenosis. *Spine (Phila Pa 1976)*. 2018;43: 1521-1528. doi:10.1097/BRS.0000000000002648

Lyons et al 597

- Cella D, Riley W, Stone A, et al. The Patient-Reported Outcomes Measurement Information System (PROMIS) developed and tested its first wave of adult self-reported health outcome item banks: 2005-2008. *J Clin Epidemiol*. 2011;63:1179-1194. doi:10. 1016/j.jclinepi.2010.04.011
- Raad M, Jain A, Huang M, et al. Validity and responsiveness of PROMIS in adult spinal deformity: the need for a self-image domain. Spine J. 2019;19:50-55. doi:10.1016/j.spinee.2018.07. 014
- Papuga MO, Mesfin A, Molinari R, Rubery PT. Correlation of PROMIS physical function and pain CAT instruments with Oswestry Disability Index and Neck Disability Index in spine patients. Spine (Phila Pa 1976). 2016;41:1153-1159. doi:10. 1097/BRS.00000000000001518
- 18. Young IA, Cleland JA, Michener LA, Brown C.Reliability, construct validity, and responsiveness of the Neck Disability Index, Patient-Specific Functional Scale, and Numeric Pain Rating Scale in patients with cervical radiculopathy. Am J Phys Med Rehabil. 2010;89:831-839. doi:10.1097/PHM. 0b013e3181ec98e6

- Mccormick JD, Werner BC, Shimer AL. Patient-reported outcome measures in spine surgery. J Am Acad Orthop Surg. 2013;21:99-107.
- O'Connell AA, McCoach DB. Applications of hierarchical linear models for evaluations of health interventions: demystifying the methods and interpretations of multilevel models. *Eval Health Prof.* 2004;27:119-151.
- Mclawhorn AS, Bjerke-Kroll BT, Blevins JL, Sculco PK, Lee Y, Jerabek SA. Patient-reported allergies are associated with poorer patient satisfaction and outcomes after lower extremity arthroplasty: a retrospective cohort study. *J Arthroplasty*. 2015;30: 1132-1136. doi:10.1016/j.arth.2015.01.043
- 22. Lavernia CJ, Alcerro JC, Brooks LG, Rossi MD. Mental health and outcomes in primary total joint arthroplasty. *J Arthroplasty*. 2012;27:1276-1282. doi:10.1016/j.arth.2011.11.015
- 23. Macy E, Ho NJ. Multiple drug intolerance syndrome: prevalence, clinical characteristics, and management. *Ann Allergy Asthma Immunol*. 2012;108:88-93. doi:10.1016/j.anai.2011.11.006
- 24. Omer HMRB, Hodson J, Thomas SK, Coleman JJ. Multiple drug intolerance syndrome: a large-scale retrospective study. *Drug Saf*. 2014;37:1037-1045. doi:10.1007/s40264-014-0236-x