ABSTRACT

Introduction Osteoarthritis (OA) is a major clinical and public health concern. The primary surgical treatment of knee OA is total knee replacement (TKR), a procedure that aims to alleviate pain and restore physical function. TKR is expensive, however, and based on professional guidelines, inappropriately performed in up to a third of patients. Patient-reported outcome measures (PROMs) help evaluate treatment options by quantifying health outcomes that matter to patients and can thus inform shared decision-making (SDM) between patients and health professionals.

Methods and analysis This is a US-based 2-year, two-site hybrid type 1 study to assess clinical effectiveness and implementation of a machine learning-based patient decision aid (PDA) integrating patient-reported outcomes and clinical variables to support SDM for patients with knee OA considering TKR. Substudy 1: At one study site, a randomised controlled trial is evaluating the clinical effectiveness of the PDA and SDM process on decision quality as measured after the baseline consultation and treatment choice measured 3 and 6 months after the baseline visit among 200 patients with knee OA. Substudy 2: At a second study site, a qualitative assessment using principles of behaviour design and intervention mapping is evaluating the feasibility and acceptability of the PROMs, PDA and SDM process by interviewing seven health professionals and 25 patients before and 25 patients after PDA implementation.

Ethics and dissemination Ethics approval has been obtained from The University of Texas at Austin Institutional Review Board (protocol number: 2018-11-0042). Informed consent will be obtained from all participants. Study results will be disseminated through conference presentations, publications and professional societies. Trial registration number NCT04805554.

INTRODUCTION

Osteoarthritis (OA) of the knee constitutes a major clinical and public health problem. This common and disabling condition has a substantial detrimental impact on affected individuals and society at large, accounting for over $27 billion in US healthcare costs annually. Treatment options for knee OA range from lifestyle changes to pharmacological management to total knee replacement (TKR) surgery. While TKR has a strong track record in alleviating pain and improving functional limitations in individuals with advanced knee OA, there are growing concerns over the escalating volume and cost of these procedures. TKR is one of the most common elective surgical procedures: the estimated number of people living in the USA in 2010 who have had a TKR was 4.7 million, with widespread variation in rates across states. By 2030, 7.4 million are expected to have knee replacement. Thus, appropriate application of TKR for the right patient at the right time is critical, especially...
within existing fee-for-service structures that incentivise performing more procedures. Notably, up to 33% of TKRs have been shown to be inappropriate based on criteria developed by the American Academy of Orthopaedic Surgeons, resulting in a substantial proportion of patients failing to experience improvement in the outcomes that matter to them. Such outcomes can be captured using patient-reported outcome measures (PROMs)—surveys that score aspects of a person’s physical, psychological and social health and well-being, directly from their perspective without interpretation by a clinician or researcher. PROMs have now been used extensively in clinical research to evaluate health status and are increasingly being applied in clinical care to monitor health outcomes and support shared decision-making (SDM).

**SDM, patient decision aids and patient-reported outcomes**

SDM is a ‘process of communication in which clinicians and patients work together to make informed health care decisions that align with what matters most to patients’. SDM and active patient participation in decision-making can be facilitated by patient decision aids (PDAs)—tools that can help people make informed decisions through patient education, knowledge assessment, elicitation of patient preferences and decision support. SDM is most appropriate for ‘preference-sensitive’ conditions, such as OA of the knee, where multiple treatment options exist and the patient preferences and values are critical in making informed treatment choices. Thus, making a decision to undergo TKR should incorporate SDM and understanding of patient preferences, values and goals, rather than objective clinical findings alone. The importance of SDM has been recognised at a national level by the US Centers for Medicare & Medicaid Services (CMS), which ties the concept to coverage of certain services that incentivise learning network in public hospital systems to increase the use of PROMs in rheumatoid arthritis and create scalable natural language processing systems to extract PROs from clinical notes, and a project looking at ways to integrate ‘patient-preferred’ hip and knee PRO scores into the EHR for use at the point of care. PDAs are generally static in the sense that their calculations are not updated with new input data. A machine learning-based tool has a theoretical advantage over a static PDA by continuously refining its prediction algorithms with new input data. Still, studies evaluating the impact of a PRO-driven, machine learning technology-enabled PDA in SDM in patients with knee OA considering TKR are lacking.

In this study, we are evaluating and implementing a tool to guide SDM in two ambulatory orthopaedic surgery practices with different patient populations, levels of expertise with PROMs and SDM, care delivery models and EHR systems. Specifically, the project involves integrating PROs and clinical data within a machine learning-based predictive analytical model, then using its output as part of SDM. Knowledge gained will be critical to scaling the use of PROMs and tools (PDAs) for SDM among patients with knee OA considering surgery.

We have designed a 2-year, two-site study using a hybrid type 1 study design to assess both clinical effectiveness and implementation. Specifically, our two aims are:

**Substudy 1:** In a randomised controlled trial (RCT) at one site, to evaluate the clinical effectiveness of the PRO-guided predictive analytical tool and process in terms of decision quality and treatment choice for patients with knee OA.

**Substudy 2:** In a qualitative assessment at the second site, to implement and evaluate the feasibility and acceptability of the tool and SDM process in a clinical setting with a different clinical population, provider group and EHR by using principles of behaviour design and intervention mapping.

**METHODS AND ANALYSIS**

**Research strategy**

**Overview**

This hybrid effectiveness-implementation study includes a non-blinded RCT of effectiveness outcomes at one site, plus periodic reflections and semistructured interviews with providers and patients to evaluate implementation processes and outcomes (eg, feasibility and acceptability) at a second site. Data will be integrated following recommended principles for mixed methods research to inform ongoing refinements to the predictive analytical tool (via formative evaluation) and plans for scaling (via intervention mapping).

**Joint Insights (artificial intelligence–enabled SDM tool) and PROMs**

Joint Insights (OM1, Boston, Massachusetts) is a machine learning-enabled PDA that uses PROMs along with patient clinical and demographic information (age, sex, body mass index, smoking status, comorbidities and number of times the patient has recently visited an emergency department or has been hospitalised) to provide...
personalised estimates of likely benefit or harm from TKR (figure 1).27 The tool is designed to collect PROs or pull in PROs collected through other systems (eg, an EHR or a third-party PROM platform). It also provides condition-specific education to patients with knee OA and allows a patient to reflect on and document their preferences and goals. Patient journeys are drawn from the OM1 Intelligent Data Cloud for patients undergoing TKR who have adequate follow-up for the outcome being evaluated. Approximately 675,000 patient records were used for the original risk model, which continues to be updated. In the modelling population (risk model), 60.8% of patients are male, the mean age is 65 years and the mean body mass index is 31.8 kg/m². The PROMs used with Joint Insights include the Patient-Reported Outcomes Measurement Information System (PROMIS) Global-10 physical and mental health subscores30 and the Knee injury and Osteoarthritis Outcome Score for Joint Replacement (KOOS JR).31 PROMIS subscores are expressed as t-scores, with 50 representing the population mean; higher scores indicate better physical function (physical function subscore) but worse mental health (mental health subscore). The KOOS JR is a seven-item PROM encompassing questions on function, pain and stiffness and scored using a scale from 0 to 100, where 0 represents poorest knee health and 100 represents best knee health.

Study dates and sites
This is a 2-year study planned from September 2020 to August 2022. The recruitment start date of this study was 22 February 2021. The PDA has already been integrated into the workflow of the UT Health Austin Clinic, where the effectiveness trial (substudy 1) is taking place. The study design and choice of different setting (UT Health San Antonio) for the implementation study (substudy 2) is intended to elucidate the feasibility and acceptability of implementing the tool into a clinic with a different population; care delivery team having less familiarity with using PROs routinely in practice; and a different EHR system, which automatically uploads PRO scores for the clinician to view at the point of care (table 1).

Substudy 1 overview
Substudy 1 is projected to take place in year 1 and early in year 2 at the UT Health Austin Musculoskeletal Institute in Austin, Texas. Patients are randomised to one of two arms: intervention, with the full Joint Insights tool
and decision regret (Decision Regret Scale (DRS)). The OA; and, as secondary endpoints, level of SDM from the subscore of the Decision Quality Index (DQI) for knee rating by using the previously validated decision process quality—as assessed at the conclusion of the initial consultation.

Substudy 2 overview

Substudy 2 is being carried out over both years primarily at UT Health San Antonio, Texas. Year 1 has entailed preparing UT Health San Antonio’s EHR to collect PROMs, preparing the EHR for integration of the predictive analytical tool, assessing baseline feasibility and acceptability and working with the clinic site to develop an implementation plan. Baseline interviews conducted with San Antonio providers and staff inquired about acceptability and feasibility of collecting PROs and using the tool, as well as exploring key factors (barriers and facilitators) impacting motivation and ability to implement the tool and SDM process at the individual and clinic levels. Interview guides were tailored to clinical role (eg, surgeon, resident, staff) and reflected implementation concepts based on the Consolidated Framework for Implementation Research (CFIR) and behaviour design, which theorises that any given behaviour is most likely to manifest when motivation, ability and a prompt to carry out the behaviour all occur in the same moment (see online supplemental information). In collaboration with the site’s clinical team, we are identifying each step in the workflow necessary to collect PROs, incorporate the PRO scores and clinical variables into Joint Insights and conduct an SDM consultation for a single patient, and assessing team perspectives on the barriers and facilitators of each step in this workflow being achieved. These data then go into developing a preliminary plan for implementation at the site, which in turn is refined iteratively in collaboration with the clinical team. Post implementation interviews are planned with providers and staff 3 months following tool roll-out to assess reported use of and experiences with the PROs and Joint Insights tool, adaptations to tool use and workflow integration and factors impacting the likelihood of sustainment of the process of care.

Semistructured interviews have also been conducted with 25 patients prior to implementing PROMs and Joint Insights in order to assess: priorities and hopes for treatment (eg, CFIR: patient needs and resources); experience of discussing treatment options with providers; and expectations for next steps in their treatment process. Then, 3 months following implementation, 25 new patients will be interviewed to assess the experiences with and acceptability of the Joint Insights tool.

Finally, periodic reflections are being conducted with members of the Austin and San Antonio implementation teams in order to document implementation processes, adaptations and contextual factors at each site. Periodic reflections are an established, low-burden method for capturing dynamic factors impacting implementation of health interventions.

Substudy 1

Practice settings, patient populations and use of PROs: UT Health Austin Musculoskeletal Institute

The UT Health Austin Musculoskeletal Institute averages about 12 new patients presenting with knee OA per week. Patients are seen by a care team that may include an associate provider (nurse practitioner), physical therapist, social worker, nutritionist and/or surgeon depending on the patient’s needs. Approximately 60% of patients are women; 50% are uninsured but covered by the Medical Access Program (MAP), which provides access to care for uninsured low-income residents of Central Texas; and 32% speak Spanish as their primary language. Musculoskeletal providers collect general and condition-specific PROs from every patient seen in the Musculoskeletal Institute (figure 2). The practice has experience with PROMs, the Joint Insights tool and SDM. PROs are collected for clinical purposes via an electronic interface and results are pulled into the EHR (Athena, Watertown, Massachusetts). Investigators (KJB, PJ) from UT Austin worked with OM1 to codevelop the PDA.

Participant selection

Inclusion criteria

► New patients aged 45–89 with a presumptive diagnosis of knee OA.
► Kellgren-Lawrence Scale joint space narrowing grade 3 or 4 (moderate to severe OA) and KOOS JR scores between 0 and 85.
► Ability to give informed consent for participation in the study.
► Ability to read text at the eighth grade reading level on a tablet in English or Spanish.

Exclusion criteria

► Patients with a prior TKR or prior consultation with another orthopaedic surgeon for TKR.
► Patients having prior experience with the Joint Insights tool.
► Patients undergoing consideration for revision joint replacement.
Patients seeking care for trauma, psoriatic arthritis or rheumatoid arthritis.

Patients with a body mass index less than 20 kg/m² or greater than 46 kg/m².

Participant recruitment and data collection
The UT Health Austin Musculoskeletal Institute sees a mix of patients seeking care for knee OA comprising a range of pathological severity – individuals who are referred from primary or specialty care or are self-referred. Suitable patients for the study are identified during the preclinic meeting. Once the patient has entered the clinic room or private consultation space, they are met by a researcher and invited to participate in the study. If they agree to participate, the researcher obtains informed consent. We are using the randomisation module of REDCap, a HIPAA-compliant research database. We are stratifying patients who enroll in the RCT on three variables: ethnicity (Latino/non-Latino), insurance (MAP/non-MAP) and orthopaedist seen (provider A (author KJB) vs provider B). This stratification ensures balance of these three variables between intervention and control groups over time and within stratum. Patients from each of the resulting eight strata are randomised to intervention or control in randomly sequenced blocks of four or six. Neither provider nor study participant will know the next allocation in the sequence until the participant is consented and it is time to begin the intervention. Due to the nature of the

Figure 2 Outcomes collected at UT Health Austin. The Knee injury and Osteoarthritis Outcome Score for Joint Replacement (KOOS JR) is a seven-item patient-reported outcome measure of knee joint-related stiffness, pain and function; interval scores range from 0 to 100, with 0 representing poorest knee health and 100 best knee health. The Patient-Reported Outcomes Measurement Information System (PROMIS) Global-10 is a 10-item measure assessing health-related quality of life with items about overall physical and mental health including social connections and physical capabilities. The survey is scored using two subscales, one for physical health and one for mental health, where specific items are used for a raw score and then converted to a t-score. Population norm t-scores are 50 on each subscore; higher scores reflect better physical health but worse mental health. The Patient Health Questionnaire (PHQ) is a validated two-item or nine-item survey assessing depressive symptoms and scored categorically as none, mild, moderate, moderately severe and severe. The two-item questionnaire is deployed, and if crossing a score threshold, an additional seven questions are generated. The Generalized Anxiety Disorder Questionnaire (GAD) is a two-item or seven-item survey assessing generalised anxiety disorder and scored categorically as none, mild, moderate and severe. Similar to the PHQ, if a score threshold is crossed on the two-item form, an additional five questions are generated. The Knee Decision Quality Instrument (K-DQI) is a 16-item survey with three specific scores: a total knowledge score, a concordance score and a decision process score. For the purposes of this study, the five questions in the shared decision-making section are used. One point is scored for ‘yes’ or ‘a lot/some’. These points are summed and then divided by 5, resulting in a score from 0% to 100%, with higher scores indicating a greater level of shared decision-making. The Collaborate is a three-item, 10-point anchor scale measuring the level of shared decision-making in a clinical encounter. It yields a continuous score with a possible range from 0 to 100, with higher scores represent a greater degree of shared decision-making. The Decision Conflict Scale (DCS) is a 10-item survey, with each response value summed, divided by the total item number and multiplied by 25. The score ranges from 0 to 100, where 0 is no decisional conflict and 100 is the greatest decisional conflict. Finally, the Decision Regret Scale (DRS) measures distress or remorse after making a healthcare decision. The answer values are summed and converted to a 0–100 scale, where a higher score indicates more regret. BMI, body mass index; ED, emergency department; TKR, total knee replacement.
intervention, patients, researchers and clinicians are not blinded to treatment arm assignment.

Demographic information is collected via tablets after randomisation. Next, patients in the intervention group receive a Joint Insights risk/benefit report. Those randomised to the intervention group may review and discuss the Joint Insights report as part of the clinical visit. The control group does not receive the Joint Insights report. Following the completion of the visit, survey instruments are collected for participants in both arms by using REDCap forms on the tablet. At 3 and 6 months of follow-up, participants are given follow-up surveys on REDCap either in person, by email or by phone. Participants completing follow-up surveys receive a $25 gift card.

Statistical precision and sample size
We calculated the sample size for the RCT by treating the decision process score of the DQI as continuous. We aimed to detect a treatment effect size (ie, Cohen’s $D$) as small as 0.5 (consistent with preliminary data from the first 26 subjects we have studied) with a type I error rate of 0.05 and power of 0.90, assuming equal sample size in intervention and control groups. Given our eight randomisation strata, we estimate a needed sample size of 180 participants, or 90 for each group. With an estimated loss to follow-up rate of 10%, our target enrolment for the RCT is 200 participants, or 100 for each arm.

Quantitative analysis
For the RCT in Austin, formal comparative analysis will follow the intent-to-treat principle. Primary analysis will compare the intervention and control groups by using multiple linear regression analysis. The model will include DQI score as the response variable and, as explanatory variables, a binary indicator for the intervention group and seven binary indicator variables representing the eight strata in order to reflect the stratified randomisation design. Additionally, as a secondary analysis, we will compare treatment decisions between the intervention and control groups by using multiple logistic regression. The model will include the treatment decision as the binary response variable and the same explanatory variables as in the linear regression model. Depending on the uptake of the intervention, additional analyses will follow the per-protocol principle where the main treatment variable will be whether the Joint Insights tool was actually used.

For analysis of the 3- and 6-month data, we will fit the linear mixed models for continuous outcomes and generalised estimating equations logistic regression models for binary outcomes, including indicator variables for time point, for treatment group and for the interaction between the two (yielding treatment effects at 3 and 6 months). Owing to the balanced design, it will be possible to fit an unstructured correlation model to eliminate any sensitivity to correlation model misspecification.

Substudy 2
Practice settings, patient populations and use of PROs: UT Health San Antonio Medical Arts & Research Center
This academic practice in San Antonio currently has one orthopaedist who treats the vast majority of patients with knee OA. This provider and a care team comprising resident physicians and an associate provider (nurse practitioner) see 16–26 new patients with knee OA per week, in addition to returning patients with OA. As in Austin, approximately 61% of patients are women, but in contrast to Austin, only 2% are uninsured and 12% report that Spanish is their primary language. The clinic had not implemented PRO collection prior to this study. The clinic uses Epic (Epic, Verona, Wisconsin) as its EHR. PROs are collected either through Epic’s MyChart patient portal in advance of the patient’s appointment or via tablets in the clinic on the day of the appointment. PRO scores are then transmitted to clinicians through the EHR’s clinician interface.

Participant recruitment
The Medical Arts & Research Center Orthopaedics Clinic in San Antonio sees a mix of patients seeking care for knee OA or considering TKR, and a mix of patients who are referred or self-referred. New patients being seen for possible TKR are contacted by project staff to schedule an interview to be conducted either in person immediately following their clinic appointment or by Zoom within the subsequent 1–2 days. A research associate obtains informed consent from all willing patients; participants who complete an interview receive a $25 gift card as compensation.

Sample size calculation
For staff and provider interviews, we have invited every member of the clinical team to participate in order to have full representation of those involved in implementation. In developing our patient sample, we considered the need to capture heterogeneity in patient demographics, condition severity, need for surgery, health literacy and preferences for treatment planning, while also acknowledging the relative homogeneity of the patient population being evaluated for knee replacement surgery in a single orthopaedic clinic. Following recommendations for ensuring information power, as specified by Malterud and colleagues, we estimated that a sample of 25 patients at each time point would provide adequate information power to represent a broad range of patient experiences and perspectives.

Qualitative analysis
All interviews are audio recorded for transcription and analysis. Interview data will be analysed using established processes for rapid qualitative analysis. We will create structured summaries from transcribed recordings to capture key domains drawn from CFIR, behaviour design and emerging content reflecting provider, staff and patient perspectives. We will then transpose domain
content from summaries into a matrix to allow for structured content comparison across participants and domains (ie, matrix analysis), an effective method for rapid and rigorous summary of findings to aid formative and implementation evaluation.23 In accordance with behaviour design and intervention mapping, we will then identify key factors impacting motivation and ability across each CFIR construct identified, separating out by stakeholder group (clinic staff, providers). For example, Joint Insights-based SDM may be perceived to be relatively advantageous (CFIR domain: intervention characteristics) by comparison with previous practice but may also raise concerns about staff burden. We will create a visual map to summarise staff and provider suggestions and concerns across each step of the Joint Insights tool implementation workflow; this map will aid collaborative implementation planning. Data from periodic reflections will also be analysed by using rapid qualitative methods in order to assess key events occurring during implementation (eg, adaptations) and factors impacting implementation (eg, barriers and facilitators). These findings will be used to support scale-up and spread of Joint Insights-based SDM and the collaboratively developed implementation strategy in future research, should results of substudy 1 suggest that the intervention is clinically beneficial.

Patient and public involvement
Patients and industry stakeholders assisted with design and feedback of the Joint Insights PDA tool for readability and usability prior to the start of this research study. Specifically, the tool was shown to patients with knee pain in the UT Health Austin Musculoskeletal Institute Lower Extremity Clinic and patients were asked a short set of open-ended questions in response to viewing the risk-benefit calculator in order to assess their understanding of the information presented and their preferences for how the information was displayed. Otherwise, no formal patient or public input was involved in designing or planning this study.

EXPECTED RESULTS
Substudy 1: We expect that patients who use the full Joint Insights tool will have higher decision process scores, reflecting better decision quality, compared with those who receive the education and preference modules only. We also expect patients in the intervention group to report higher levels of SDM and lower levels of decision conflict and decision regret. We do not expect a difference in rates of treatment selected (operative vs nonoperative) between the two groups.

Substudy 2 is exploratory and therefore has no formal hypotheses.

ETHICS AND DISSEMINATION
The University of Texas at Austin Dell Medical School Institutional Review Board (IRB) reviewed and approved this study (protocol number: 2018-11-0042). The University of Texas Health Science Center at San Antonio’s IRB has a formal reliance agreement with The University of Texas at Austin IRB. Any modifications to the protocol will be submitted to the UT Austin IRB for approval before implementation.

Patients and clinic staff are enrolled in this study after providing informed consent. During this study, participants complete the questionnaires related to their decision-making process and experience or are interviewed formally about their experiences. Data are kept in strict confidence. No information will be given to anyone without permission from the participant. Confidentiality is assured by use of identification codes, password-protected electronic files on secure servers or hosting applications and paper files stored under lock and key. The assessments are conducted in a private setting, through encrypted email or by telephone. Although we do not anticipate any adverse events, any adverse events will be reported to the local IRB.

Dissemination of results
The project will facilitate developing a learning healthcare system. PRO data will be collected electronically and used to inform clinical decision-making in real time. We will evaluate PRO data to improve clinical decision-making and patient outcomes locally at two sites. We will disseminate results through publications, meeting presentations and professional organisations.

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Competing interests PJ has received personal fees from Johnson & Johnson Medical Devices. KJB has received personal fees from the CMS and Purchaser
Business Group on Health; has stock options from Carrum Health; and has a leadership role with the American Academy of Orthopaedic Surgeons. KJB has royalty agreements with Wolters Kluwer and Slack. KJB and PJ are codevelopers of the Joint Insights tool; they have no personal financial interest in the tool. JT receives royalties from Wolters Kluwer. The University of Texas at Austin has a royalty agreement with OM1.

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