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What are patients' preferences for revision surgery after periprosthetic joint infection? A discrete choice experiment

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Abstract

Objectives: Understanding patients' preferences for treatment is crucial to provision of good care and shared decisions, especially when more than one treatment option exists for a given condition. One such condition is infection of the area around the prosthesis after hip replacement, which affects between 0.4 and 3% of patients. There is more than one treatment option for this major complication and our study aimed to assess the value that patients place on aspects of revision surgery for periprosthetic hip infection.

Design: We identified four attributes of revision surgery for periprosthetic hip infection. Using a discrete choice experiment (DCE) we measured the value placed on each attribute by 57 people who had undergone either one- or two-stage revision surgery.

Setting: The DCE was conducted with participants from 9 National Health Service hospitals in the United Kingdom.

Participants: Adults who had undergone revision surgery for periprosthetic hip infection (n=57).

Results: Overall, the strongest preference was for a surgical option that resulted in no restrictions on engaging in valued activities after a new hip is fitted. Less valued, but still important attributes, included a shorter time taken from the start of treatment to return to normal activities, few or no side effects from antibiotics, and having only one operation.

Conclusions: Results highlight that people who have had revision surgery for periprosthetic hip infection most value aspects of care that affect their ability to engage in normal everyday activities. These were the most important characteristics in decisions about revision surgery.

Keywords

Discrete choice experiment; hip; periprosthetic infection; orthopaedic surgery

Strengths and limitations of this study

- This is the first study to quantify the value that patients place on different aspects of revision surgery for prosthetic hip joint infection.
- Using a rigorous process involving empirical qualitative research we identified four attributes for the DCE.
- It was feasible for participants to complete the questionnaires, meaning the results were as
 expected with patients selecting a combination of options reflecting their preferred choice.
- While, the sample size of 57 participants sufficiently powered the analyses presented here, it
 is too small to conduct any subgroup analyses.
- The majority of the study sample were white, male and educated, so results may not reflect the preferences of the wider surgical population.

Introduction

For people with osteoarthritis, hip replacement is a common procedure that aims to improve function and reduce pain. In 2016 over 100,000 hip replacement procedures were conducted in the UK. With an ageing population, rates are predicted to rise ¹. Although successful for many people, approximately 0.4 -1% of patients who have undergone primary hip replacement ^{2,3} and 2-3% of patients undergoing revision hip replacement ⁴ develop deep periprosthetic joint infection (PJI) severe enough to warrant surgical revision. Patients with PJI find it devasting. Symptoms include severe pain, inflammation, discharge from the surgical wound, fever, nausea, malaise, reduction in or loss of function, dislocation, and if left untreated can lead to disability and/or death ⁵.

implant retention' (DAIR), used in 7.6% of cases, or more commonly major revision surgery, which

involves removal of the infected prosthesis, radical debridement of infected tissue and

reimplantation of a new prosthesis with subsequent antibiotic treatment ⁴. Revision surgery can be provided in a single operation (one-stage) or as a staged operation (two-stage) where the infected implant is removed and the patient is left without an implant (with or without a temporary spacer) while receiving antibiotic treatment. In the staged process reimplantation of the new implant is delayed, commonly for up to 6 months, but in some instances over 12 months later ^{5, 6}. There is no clear evidence that either a one or two-stage strategy is superior in eradicating infection ^{7, 8}, but qualitative research has shown that two-stage revision places a greater burden on patients and families than one-stage revision. This burden is due to the extended period of immobility in between operations, complications associated with the period of immobility, and deep psychological distress, with some patients reporting depression and suicidal thoughts ⁶.

Surgeons' decisions about which type of revision surgery is most appropriate for an individual patient take into account many factors. Our previous work indicates that decisions are based on a combination of a surgeon's own training and clinical experience of different techniques; the availability of hospital infrastructure such as microbiology services to quickly identify the infecting organism; characteristics of the infecting organism and duration of infection; patient characteristics such as age, comorbidities, frailty, and extent of damaged tissue; and published evidence of revision techniques, or reports by senior colleagues. Surgeons also considered patients' preferences for surgery, although this could often involve the choice between long-term suppressive antibiotics or surgery ⁹. There is no quantitative evidence that characterises patients' preferences for one-stage or two-stage revision surgery in this context and this is an important area of work to be investigated. The aim of this study is to assess the surgical preferences of patients who underwent revision surgery for prosthetic hip joint infection.

This study is part of a larger programme of research which aims to improve outcomes for patients after periprosthetic joint infection (NIHR PGfAR: RP-PG-1210-12005). The programme includes a

randomised clinical trial comparing one-stage and two-stage revision surgery for prosthetic hip joint infection ¹⁰ within which this study was embedded.

Methods:

Discrete choice experiments (DCEs)

We undertook a discrete choice experiment (DCE) to quantify the surgical preferences of patients who underwent revision surgery for prosthetic hip joint infection. The DCE was embedded within a randomised clinical trial that evaluated clinical and cost effectiveness of revision surgery for prosthetic hip joint infection and compared one-stage revision surgery with two-stage revision surgery. Patients who were taking part in the trial were eligible for the DCE. Ethical approval for the study was granted by NRES Committee South West – Frenchay on 31 December 2014 (14/SW/1166). All participants provided written informed consent to take part in the DCE study.

DCEs are an established method in health services research and have been used to explore a range of health-related services and treatments ¹¹⁻¹³. DCEs involve asking respondents to choose between hypothetical scenarios which describe goods or services, where a service may mean an intervention or an approach to care. The method aims to establish what attributes of that service influence their decision-making and to what extent. This enables quantification of the marginal impact of these attributes. Scenarios within a DCE describe the service of interest (in this case, revision joint replacement) using the same set of attributes, but at different levels in each scenario. Choices between scenarios, or whether to accept or reject a scenario, are used to estimate the influence and value of the different attribute levels.

Questionnaire development

When a patient faces revision surgery for prosthetic hip infection, a one-stage or a two-stage operation is required, and the decision about which is undertaken is made largely at the discretion of the surgeon, taking into account patient preferences for treatment. The DCE study was designed to engage patients in their preferences for the features associated with these two surgical options, given neither is currently known to be clinically superior in terms of patient outcomes.

Qualitative methods are recommended for DCE attribute development because they enable conceptual development of attributes directly from people's experiences and so better reflect the issues that are likely to matter most to people when making a decision ¹⁴. For this study we developed attributes from our previous qualitative study which explored impact of prosthetic joint infection and its treatment on patients and their recovery process; one-to-one interviews were conducted with 19 patients with PJI, focusing on the impact of PJI and surgical treatment. The interviews were audio-recorded, transcribed and the qualitative data set was analysed thematically ⁶. Levels of attributes were assigned by refining the language used to convey the meaning of the attributes, and particularly where some quantification of an attribute was mentioned by participants during qualitative interview, as illustrated in Table 1 ¹⁴.

Table 1 describes the attribute and level selection based on the earlier qualitative work, including illustrative quotations and rationale.

The final questionnaire comprised a single scenario task in which respondents were asked to imagine this was the first time that they had been offered the two surgical options presented and were asked to consider them carefully before selecting the one (from a pair) they would prefer.

With a $4 \times 4 \times 2 \times 2$ design (two four-level and two two-level attributes), a total of 64 different combinations of attribute levels (profiles) is possible. This converts to 32 pairs of profiles, which as a 'full factorial' was considered to be too large for participants to complete ¹⁵. An orthogonal main effects plan was therefore used to reduce the number of choice sets to 16 ¹⁶. Each profile was presented with its pair, and participants were required to select which option they preferred (see Figure 1).

Patient and Public Involvement

The questionnaire was piloted and refined in collaboration with 5 patient and public involvement representatives. At an initial meeting of representatives, group members were involved in questionnaire development and suggested improvements to its formatting to aid readability, clearer phrasing of the questions to avoid ambiguity, and shortening of the instruction leaflet for clarity. At a subsequent meeting, members of the group completed the questionnaire and fed back on their experience. They completed the questionnaire without assistance and felt the instructions were clear but suggested that key points in the questions should be highlighted and that a contact telephone number should be added to enable participants to seek assistance if needed. For participants in the study, a summary of findings will be sent to those who indicated that they wished to be informed.

Participants

Patients who were recruited into the INFORM randomised controlled trial (ISRCTN10956306) received the discrete choice task after completing the 18-month primary outcome measure. The questionnaire was either posted to participants or completed in person with the assistance of a research nurse during a hospital clinic visit.

Data analysis

Sample size calculations for DCEs are challenging due to dependence on the true parameter values estimated in the choice model ¹⁷. However, reliable statistical analysis has been demonstrated with sample sizes of 40–120 respondents and combined with the rarity of periprosthetic joint infection (1% of those undergoing hip replacement), a sample size above 50 participants for the DCE was deemed adequate to obtain sufficient data for analysis and interpretation ^{18, 19}.

Paper questionnaires were distributed to participants as part of the follow-up data collection in the INFORM trial. Questionnaires were returned to the study team between January 2017 and November 2018 and data entered onto a Microsoft Excel spreadsheet. DCE data were effects coded and analysed using STATA SE 15 ^{20, 21}. The influence of the four attributes on patient choices was analysed using a conditional logit model. As attribute levels are effects coded, the mean of all coefficients is 0 across each attribute. The coefficients indicate the strength of preference for each level, where more positive values indicate a greater likelihood that the patient would select a surgical option where that particular attribute level is present.

Results:

Participants

Of the 80 discrete choice questionnaires provided to trial participants, 57 were returned fully complete (71%) from patients from 9 trial sites. Data from 6 questionnaires was not used in analysis as they were partially or totally incomplete. Responding participants had a mean age of 70 (range 51 to 90), 21 (37%) were female, 26 (46%) had undergone 1-stage revision, 14 (25%) lived alone and 41 (72%) were retired from work. Table 2 provides further demographic information.

Table 2 – Demographic and Clinical characteristics of the DCE Respondents

Characteristic	Participants (N=57)
Age – Yrs [mean] (range)	70 (51-90)
Gender [number] (%)	
Male	36 (63)
Female	21 (37)
Ethnicity [number] (%)	
White	55(96)
Black	1 (2)
Mixed	1 (2)
Marital Status [number] (%)	
Married / Partner	42 (74)
Divorced / Separated / Widowed	12 (21)
Single	3 (5)
Living Arrangements [number] (%)	
With Partner / somebody else	43 (75)
Alone	14 (25)
Schooling/Education [number] (%)	
Left at normal school leaving age	35 (61)
Left after normal school leaving age	15 (26)
Left before normal school leaving age	7 (12)
Work Situation [number] (%)	
Retired	41 (72)
Working / Sick leave	14 (25)

Unemployed	2 (4)
Surgery Received for Prosthetic Hip Joint Infection [number] (%)	
2-stage revision	31 (54)
1-stage revision	26 (46)

Discrete choice experiment

Table 3 shows the regression coefficients and the results from the 57 patients who had fully completed the discrete choice questionnaire.

Table 3 – discrete choice task results from conditional logistic regression

Attribute	Level	Coefficient	Standard	95%	P value
			error	confidence	
		2.		interval	
Ability to engage in	Can do	0.70			
valued activities after	everything*				
new hip is fitted.					
	Can do most	0.49	.08	.33 to .64	<0.001
	things				
	Cannot do	-0.39	.07	53 to24	<0.001
	most things				
	Cannot do	-0.80	.13	-1.05 to -	<0.001
	anything			0.55	
Antibiotic side effects	Don't affect	0.22			
	me much*				

	Affects me a	-0.22	.05	33 to12	<0.001
	lot				
Number of operations	1*	0.20			
	2	-0.20	.07	35 to06	<0.001
Time taken after	3m*	0.20			
surgical treatment					
starts, to return to					
normal activities.	10				
	6m	0.31	.09	.14 to .48	<0.001
	12m	-0.06	.05	15 to .04	0.22
	18m	-0.45	.10	64 to26	<0.001
		4:			

*Indicates reference category within attribute

Analysis indicates that participants had the strongest preference for a surgical option that resulted in the least restrictions on engaging in valued activities after the new hip is fitted, illustrated by the largest positive coefficient. Other less valued but important preferences were for a surgical strategy that would result in a shorter time after surgical treatment starts to return to normal activities, few or no side effects from antibiotics, and only one operation. The results also suggest that the least restrictions on engaging in valued activities, and the shortest time taken to return to normal activity are the individual attributes most valued by patients in this sample. This is indicated by the larger spread of coefficients (i.e. more 'value' is placed on changes in these attributes). The most acceptable option was a time period of between 3 to 6 months to return to normal activity, however

there is no clear preference up to 12 months, although 18 months appeared to be significantly disfavoured.

Discussion:

This study aimed to investigate and understand patients' preferences for aspects of revision surgery for periprosthetic joint infection. Four relevant attributes were identified through earlier qualitative work, and quantitatively, patients in this study most value the ability to engage in valued activities and the time taken to return to normal daily activities. This reflects the findings of our previous qualitative work which shows that although both revision strategies impacted greatly on patients and their families, patients receiving two-stage revision surgery experience particularly long periods of immobility and social isolation. This was often followed by a protracted recovery period, which could leave patients much less able than before their primary operation, and some patients experienced profoundly negative psychological effects associated with physical suffering, loss of dignity and independence ⁶. It appears that for patients in our sample, 3-6 months to return to normal activity was preferable, although there was no significant difference up to 12 months, but 18 months was disfavoured. This suggests that the acceptable margin of recovery for patients is up to 12 months after their receipt of a new hip joint.

Discrete choice methodology can be challenging for participants because the format of questions is different to standard surveys and items can seem repetitive. We collected feedback from the first 11 participants who completed the questionnaire. We found that those participants who were supported by a research nurse when completing the questionnaire were more likely to complete it and return it, compared with those who received the questionnaire by post and completed it alone. Participant feedback suggested that the questionnaire was difficult to complete, as the scenarios were similar and appeared to be repetitive. To address this, we amended the questionnaire format and instructions and offered participants support either face-to-face or by phone with one of the

study research nurses. Nurses were then able to answer queries about the questionnaire and offer support if needed. The results suggest the group completed the questionnaire in a rational and logical way, meaning the results were as expected with patients selecting an optimal combination of options as their preferred choice. This is an important methodological finding because although our study demonstrates the feasibility of the DCE method with this population, others conducting similar studies with older, ill populations could consider in advance the need for professional support in the completion of discrete choice questionnaires.

Participants were all individuals who participated in a clinical trial and had already undergone revision surgery for periprosthetic joint infection. This meant that the choices that participants were asked to make in the questionnaire were based on scenarios unlikely to reflect their real-life experiences as in reality such choices would not be available to them, since decisions about surgical strategies are based on a wider variety of clinical, surgeon, patient and organisational factors ⁹. Also, patients were not being faced with these decisions at the time of questionnaire completion as we decided that it would be unethical to ask patients awaiting treatment to complete a DCE about surgical options in a hypothetical context, at a time when they may be particularly vulnerable. To do so would not meet an ethical standard of protection from harm as it might mean that patients were inadvertently led to believe that there were more or different options available to them than were clinically indicated at that time.

We also found that some participants reported difficulty separating their own recent personal experience of revision surgery from the hypothetical scenarios presented in the questionnaire, as they found it hard to imagine receiving a treatment option that differed from the one that they had received. In terms of methodology, many DCE's are conducted with participants who already have some experience of the treatment attributes under investigation ^{12, 18, 19, 22}. In our study previous experience meant that participants had some appreciation of the attributes being tested. While, the

sample size of 57 participants provided sufficient data for the analyses presented here, the sample size was too small to conduct any subgroup analysis to identify whether preferences would differ between participants who had received one or two stage revision. Similarly, patients who are older, still working or live alone may have had stronger preferences for a one-stage operation than those who are younger, retired and have support at home to cope with a two-stage procedure. The majority of the study sample were also white, male and educated, which means that results may not reflect the preferences of the wider surgical population. Further research could explore preferences in a more diverse population.

This work has provided an initial and important first step in understanding patients' preferences for characteristics associated with revision surgery for periprosthetic infection. It is important that orthopaedic healthcare professionals discuss these attributes with patients when discussing options for surgical and antibiotic treatment for periprosthetic infection.

Implications

Results of this study offer insight into the preferences of patients for revision surgery and provide valuable information to surgeons from all disciplines. Although factors affecting patient preferences for surgery differ to those valued by medical professionals, consideration should be given to such factors in order to aid shared decision-making where clinical equipoise between options exists.

Previous research using discrete choice approaches has explored patient preferences in a surgical context. This has included examination of preferences for surgical versus non-surgical interventions in a range of conditions, such as oesophageal cancer and ulcerative colitis ^{22, 23}. Some research has explored preferences for conditions in which there are two surgical options, including for ectopic pregnancy, vaginal wall prolapse, and osteoarthritis ^{24, 25}. Across all of these conditions, evidence suggests that patients choose options that reduce the need for further surgery or operations, have a

shorter recovery time, lower risk of symptom recurrence and improve ability to preserve existing joint motion in the case of osteoarthritis. Findings from the current study are similar in that patients prefer a surgical option that reduces the number of operations, recovery time and the side effects of antibiotics. However, in the setting of infected joint replacement, patients placed highest value on restoration of function. This was more important to the patients in our study than the number of operations they would have to undergo. Although our study focused on the preferences between one and two-stage surgery surgeons may need to consider these preferences during shared decision-making about all options for revision surgery for prosthetic hip infection, including the role of debridement with retention of implants (DAIR). Although DAIR is only efficacious in approximately 60% of cases, it is associated with a quicker return to valued activities and improved joint function ²⁶.

Conclusions

Our results show that the most valued characteristics in decisions about revision surgery for prosthetic hip infection were the ability to engage in valued activities and time taken to return to normal activity. This builds on the findings of our previous qualitative work which shows that although both revision strategies impact greatly on patients and their families' everyday lives, patients receiving two-stage revision surgery experience particularly long periods of immobility and social isolation ⁶. The desire to return to everyday activities should be taken into account when surgeons are discussing options with patients, particularly when there is equipoise from a surgical perspective about the options available, and when the decision is 'preference sensitive'.

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Author Contributions

FEC: Contributed to design, analysis and interpretation of data, and drafting the manuscript. RG-H: Contributed to conception, design, interpretation of data, drafting and revising the manuscript. SS: Contributed to design, data collection and revising the manuscript. AWB: Contributed to the conception, design, interpretation of data and revising the manuscript. AJM: Contributed to the design, interpretation of data, drafting and revising the manuscript. All authors gave final approval of the submitted manuscript.

Competing Interests

None declared

Data Availability

Anonymised research data will be made available at the University of Bristol Research Data Repository (data.bris) beginning 1 year after publication of the results. Access will be granted to bona fide researchers after the University of Bristol Data Access Committee has approved their request. Please contact data-bris@bristol.ac.uk in the first instance.

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References

- 1. National Joint Registry. National Joint Registry 14th Annual Report 2017: National Joint Registry for England, Wales, Northern Ireland and the Isle of Man; 2017 [Available from: www.njrreports.org.uk.
- 2. Blom AW, Taylor AH, Pattison G, Whitehouse S, Bannister GC. Infection after total hip arthroplasty. The Avon experience. The Journal of bone and joint surgery British volume. 2003;85(7):956-9.
- 3. Phillips JE, Crane TP, Noy M, Elliott TS, Grimer RJ. The incidence of deep prosthetic infections in a specialist orthopaedic hospital: a 15-year prospective survey. The Journal of bone and joint surgery British volume. 2006;88(7):943-8.
- 4. Lenguerrand E, Whitehouse MR, Beswick AD, Jones SA, Porter ML, Blom AW. Revision for prosthetic joint infection following hip arthroplasty: Evidence from the National Joint Registry. Bone & joint research. 2017;6(6):391-8.
- 5. Zimmerli W, Trampuz A, Ochsner PE. Prosthetic-joint infections. The New England journal of medicine. 2004;351(16):1645-54.
- 6. Moore AJ, Blom AW, Whitehouse MR, Gooberman-Hill R. Deep prosthetic joint infection: a qualitative study of the impact on patients and their experiences of revision surgery. BMJ open. 2015;5(12):e009495.
- 7. Kunutsor SK, Whitehouse MR, Blom AW, Beswick AD. Re-Infection Outcomes following Oneand Two-Stage Surgical Revision of Infected Hip Prosthesis: A Systematic Review and Meta-Analysis. PloS one. 2015;10(9):e0139166.
- 8. Kunutsor SK, Whitehouse MR, Blom AW, Board T, Kay P, Wroblewski BM, et al. One- and two-stage surgical revision of peri-prosthetic joint infection of the hip: a pooled individual participant data analysis of 44 cohort studies. European journal of epidemiology. 2018;33(10):933-46.
- 9. Moore AJ, Blom AW, Whitehouse MR, Gooberman-Hill R. Managing uncertainty a qualitative study of surgeons' decision-making for one-stage and two-stage revision surgery for prosthetic hip joint infection. BMC musculoskeletal disorders. 2017;18(1):154.
- 10. Strange S, Whitehouse MR, Beswick AD, Board T, Burston A, Burston B, et al. One-stage or two-stage revision surgery for prosthetic hip joint infection--the INFORM trial: a study protocol for a randomised controlled trial. Trials. 2016;17:90.
- 11. Louviere J, Street DJ, Burgess L. A 20+ years' retrospective on choice experiments. Marketing research and modeling: Progress and prospects: Springer; 2004. p. 201-14.
- de Bekker-Grob EW, Hofman R, Donkers B, van Ballegooijen M, Helmerhorst TJ, Raat H, et al. Girls' preferences for HPV vaccination: a discrete choice experiment. Vaccine. 2010;28(41):6692-7.
- 13. Liede A, Mansfield CA, Metcalfe KA, Price MA, Snyder C, Lynch HT, et al. Preferences for breast cancer risk reduction among BRCA1/BRCA2 mutation carriers: a discrete-choice experiment. Breast cancer research and treatment. 2017;165(2):433-44.
- 14. Coast J, Al-Janabi H, Sutton EJ, Horrocks SA, Vosper AJ, Swancutt DR, et al. Using qualitative methods for attribute development for discrete choice experiments: issues and recommendations. Health economics. 2012;21(6):730-41.

- 15. Marshall D, Bridges JF, Hauber B, Cameron R, Donnalley L, Fyie K, et al. Conjoint Analysis Applications in Health How are Studies being Designed and Reported?: An Update on Current Practice in the Published Literature between 2005 and 2008. The patient. 2010;3(4):249-56.
- 16. Hedayat AS, Sloane NJA, Stufken J. Orthogonal arrays: theory and applications: Springer Science & Business Media; 2012.
- 17. Lancsar E, Louviere J. Conducting discrete choice experiments to inform healthcare decision making: a user's guide. PharmacoEconomics. 2008;26(8):661-77.
- 18. Salkeld G, Solomon M, Butow P, Short L. Discrete-choice experiment to measure patient preferences for the surgical management of colorectal cancer. The British journal of surgery. 2005;92(6):742-7.
- 19. de Bekker-Grob EW, Essink-Bot ML, Meerding WJ, Koes BW, Steyerberg EW. Preferences of GPs and patients for preventive osteoporosis drug treatment: a discrete-choice experiment. PharmacoEconomics. 2009;27(3):211-9.
- 20. Bech M, Gyrd-Hansen D. Effects coding in discrete choice experiments. Health economics. 2005;14(10):1079-83.
- 21. StataCorp. Stata Statistical Software: Release 15. College Station, TX: StataCorp LLC; 2017.
- 22. Noordman BJ, de Bekker-Grob EW, Coene P, van der Harst E, Lagarde SM, Shapiro J, et al. Patients' preferences for treatment after neoadjuvant chemoradiotherapy for oesophageal cancer. The British journal of surgery. 2018;105(12):1630-8.
- 23. Bewtra M, Kilambi V, Fairchild AO, Siegel CA, Lewis JD, Johnson FR. Patient preferences for surgical versus medical therapy for ulcerative colitis. Inflammatory bowel diseases. 2014;20(1):103-14.
- 24. van Mello NM, Mol F, Opmeer BC, de Bekker-Grob EW, Essink-Bot ML, Ankum WM, et al. Salpingotomy or salpingectomy in tubal ectopic pregnancy: what do women prefer? Reproductive biomedicine online. 2010;21(5):687-93.
- 25. Notten KJ, Essers BA, Weemhoff M, Rutten AG, Donners JJ, van Gestel I, et al. Do patients prefer mesh or anterior colporrhaphy for primary correction of anterior vaginal wall prolapse: a labelled discrete choice experiment. BJOG: an international journal of obstetrics and gynaecology. 2015;122(6):873-80.
- 26. Kunutsor SK, Beswick AD, Whitehouse MR, Wylde V, Blom AW. Debridement, antibiotics and implant retention for periprosthetic joint infections: A systematic review and meta-analysis of treatment outcomes. The Journal of infection. 2018;77(6):479-88.

Table 1 – qualitative support for attributes included in the discrete choice questionnaire

Attribute	Evidence of attribute	Levels	Rationale for
	inclusion, with pseudonym		levels
	and surgery type		
Number of operations	"There's no way I want two	1) 1 operation	Two types of
	more big operations now at		revision surgery

	my time of life. You do it all	2)	2 operations	are currently
	or not at all"I said there			provided in
	was no way I wanted two			healthcare, and
	ops" (Harriet, 1-stage)			involve either
				one or two
	"Of course, emotionally,			operations
	you want it over and done			
	with as soon as			
	possiblebut ultimately			
	that has to be done in the			
	correct way. There's the			
	tortoise and hare situation.			
	There's absolutely no point			
	in rushing ahead if			
	ultimately it's going to fail."	4		
	(Maggie, 2-stage)			
Ability to engage in	"Fourteen months without	1)	Can do	Following
valued activities after	a hip joint so it meant that I		everything	revision
new hip is fitted.	couldn't drive a car, I	2)	Can do most	surgery, the
	couldn't do anything that		things	ability to
	I'd been used to doing,	2)	Cannot da	engage in
	playing golf or doing	3)	Cannot do	valued
	anything. Well, I gave up		most things	activities can be

golf actually after the first	4)	Cannot do	reduced in a
revision" (Don, 2-stage)		anything	major, or
			somewhat
4 2			more minor
"But when I for example			way. These
went to, on holiday			levels capture
recently and I had serious			variation in
problems getting into the			ability
bath to stand in the			identified by
shower. Because I, I			patients.
couldn't get in. And in a			
wet – on a wet surface and			
that, I'm very conscious of			
not falling in. I can't afford			
to fall. So, pain I've got			
none, stiffness none.	4		
Physical function			
limitations, and that's one			
of them." (Rory, 1-stage)			
"My aim has always been			
to get back on my feet as			
soon as I can, and to walk			
as good as I can, and that's			
a big disappointment. I'm			

	not where I think I should			
	have been" (Robert, 2-			
	stage)			
Time taken after	"I didn't want to go 14	1)	3 months	These time
surgical treatment	weeks with effectively one	2)	6 months	intervals
starts, to return to	leg. What was worse was	2)	10	demonstrate
normal activities.	not knowing that I had to	3)	12 months	best
	endure all of those weeks	4)	18 months	approximations
	not knowing that I was ever			and a reflection
	going to get another hip			of the need to
	joint back." (Maggie, 2-			ensure normal
	stage)			expectations of
				time taken for
		•		soft tissue
	"If I had known how hard it	0		recovery as
	was going to be for her to	7		expected by
	walk in that interim six			surgeons. 18
	months, and if there			months is the
	seemed to have been a			maximum end
	reasonable, or a good			point that
	possibility that the			surgeons would
	infection would be nuked in			suggest for
	a one stage, then that			recovery time.
	might have been a better			

	outcome for her" (Amelia,		
	2-stage)		
	"I would've thought that if		
	you wanted to go back to		
	workyou wouldn't be very		
	happy [having a 2-stage		
	operation] because you		
	wouldn't be able to do		
	nothing for those six weeks		
	again. Then you'd have to		
	go all through it again after		
	three months of having it		
	done. Six weeks is only a	•	
	month and a half, and then	4	
	in another month and a		
	half you're having it all		
	done again." (Jim, 1-stage)		
Antibiotic side effects	"the nightmare on heavy	1) Affects me a	Antibiotics are
	antibiotics, toiletry wise.	lot	an essential
	Now I've had to move into		method of
	a, another bed. My wife		attempting to
	and I are married 50 plus		ensure the

veges and they are being	2)	Don't offers	
years, and I have to have	2)	Don't affect	periprosthetic
my own room because I'm		me much	joint infection
getting up in the night."			is treated and
(Rory, 1-stage)			subsequently
			clear. For some
"I stayed on antibiotics			patients, the
			impact of these
then for, for everand			antibiotics is
after the first [week of			significant,
antibiotics] I was just dying.			while for others
I just wanted to lie on the			
floor and die. I felt so sick.			there are less
			severe side
So ill." (Lottie, 2-stage)			effects.
	•		
"I felt fine. When I was	0		
going to see the surgeon	2		
They'd say, 'How are you			
today?' I'd say, 'I feel fine,			
fit! [laughs] I feel well in			
myself, I eat well.'" (Ray, 2-			
stage)			
	<u> </u>		

OPTION H1
I take antibiotics and the side effects don't affect
me much
and
I have two operations

and

After my new hip is fitted, I can do most of the things that I want to do

and

After my surgical treatment starts, it takes 6 months to get back to the things that I normally do

OR

I have one operation
and

After my new hip is fitted, I cannot do most of the things that I want to do

OPTION H2

I take antibiotics and the side effects affect me a

lot

and

and

After my surgical treatment starts, it takes 12 months to get back to the things that I normally do

Please only tick ONE box:

OPTION H1 OPTION H2

Figure 1 - example profile

338x190mm (96 x 96 DPI)

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What are patients' preferences for revision surgery after periprosthetic joint infection? A discrete choice experiment

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Abstract

- 2 Objectives: Understanding patients' preferences for treatment is crucial to provision of good care
- 3 and shared decisions, especially when more than one treatment option exists for a given condition.
- 4 One such condition is infection of the area around the prosthesis after hip replacement, which
- 5 affects between 0.4 and 3% of patients. There is more than one treatment option for this major
- 6 complication and our study aimed to assess the value that patients place on aspects of revision
- 7 surgery for periprosthetic hip infection.
- 8 Design: We identified four attributes of revision surgery for periprosthetic hip infection. Using a
- 9 discrete choice experiment (DCE) we measured the value placed on each attribute by 57 people who
- 10 had undergone either one- or two-stage revision surgery for infection.
- 11 Setting: The DCE was conducted with participants from 9 National Health Service hospitals in the
- 12 United Kingdom.
- 13 Participants: Adults who had undergone revision surgery for periprosthetic hip infection (n=57).
- 14 Results: Overall, the strongest preference was for a surgical option that resulted in no restrictions on
- engaging in valued activities after a new hip is fitted (β =0.7). Less valued, but still important
- 16 attributes, included a shorter time taken from the start of treatment to return to normal activities (6
- months; β =0.3), few or no side effects from antibiotics (β =0.2), and having only one operation
- 18 (β =0.2).
- 19 Conclusions: Results highlight that people who have had revision surgery for periprosthetic hip
- 20 infection most value aspects of care that affect their ability to engage in normal everyday activities.
- 21 These were the most important characteristics in decisions about revision surgery.

Keywords

2 Discrete choice experiment; hip; periprosthetic infection; orthopaedic surgery

3 Strengths and limitations of this study

- This is the first study to quantify the value that patients place on different aspects of revision surgery for prosthetic hip joint infection.
- Using a rigorous process involving empirical qualitative research we identified four attributes
 for the DCE.
 - It was feasible for participants to complete the questionnaires, meaning the results were as expected with patients selecting a combination of options reflecting their preferred choice.
 - While, the sample size of 57 participants sufficiently powered the analyses presented here, it
 is too small to conduct any subgroup analyses.
 - The majority of the study sample were white, male and educated, so results may not reflect the preferences of the wider surgical population.

Introduction

For people with osteoarthritis, hip replacement is a common procedure that aims to improve function and reduce pain. In 2016 over 100,000 hip replacement procedures were conducted in the UK. With an ageing population, rates are predicted to rise ¹. Although successful for many people, approximately 0.4 -1% of patients who have undergone primary hip replacement ^{2,3} and 2-3% of patients undergoing revision hip replacement ⁴ develop deep periprosthetic joint infection (PJI) severe enough to warrant surgical revision. Patients with PJI find it devasting. Symptoms include

severe pain, inflammation, discharge from the surgical wound, fever, nausea, malaise, reduction in or loss of function, dislocation, and if left untreated can lead to disability and/or death 5. PJI is extremely challenging to treat. Treatment can include 'debridement, antibiotic treatment and implant retention' (DAIR), used in 7.6% of cases, or more commonly major revision surgery, which involves removal of the infected prosthesis, radical debridement of infected tissue and reimplantation ("fitting") of a new prosthesis with subsequent antibiotic treatment ⁴. Revision surgery can be provided in a single operation (one-stage) or as a staged operation (two-stage) where the infected implant is removed and the patient is left without an implant (with or without a temporary spacer) while receiving antibiotic treatment. In the staged process reimplantation of the new implant is delayed, commonly for up to 6 months, but in some instances over 12 months later 5, ⁶. There is no clear evidence that either a one or two-stage strategy is superior in eradicating infection ^{7,8}, but qualitative research has shown that two-stage revision places a greater burden on patients and families than one-stage revision. This burden is due to the extended period of immobility in between operations, complications associated with the period of immobility, and deep psychological distress, with some patients reporting depression and suicidal thoughts 6. Surgeons' decisions about which type of revision surgery is most appropriate for an individual patient take into account many factors. Our previous work indicates that decisions are based on a combination of a surgeon's own training and clinical experience of different techniques; the availability of hospital infrastructure such as microbiology services to quickly identify the infecting organism; characteristics of the infecting organism and duration of infection; patient characteristics such as age, comorbidities, frailty, and extent of damaged tissue; and published evidence of revision techniques, or reports by senior colleagues. Surgeons also considered patients' preferences for surgery, although this could often involve the choice between long-term suppressive antibiotics or surgery 9. There is no quantitative evidence that characterises patients' preferences for one-stage or

two-stage revision surgery in this context and this is an important area of work to be investigated.

- 1 The aim of this study is to assess the surgical preferences of patients who underwent revision
- 2 surgery for prosthetic hip joint infection.
- 4 This study is part of a larger programme of research which aims to improve outcomes for patients
- 5 after periprosthetic joint infection (NIHR PGfAR: RP-PG-1210-12005). The programme includes a
- 6 randomised clinical trial comparing one-stage and two-stage revision surgery for prosthetic hip joint
- 7 infection ¹⁰ within which this study was embedded.

9 Methods:

10 Discrete choice experiments (DCEs)

- We undertook a discrete choice experiment (DCE) to quantify the surgical preferences of patients
- who underwent revision surgery for prosthetic hip joint infection. The DCE was embedded within a
- randomised clinical trial that evaluated clinical and cost effectiveness of revision surgery for
- prosthetic hip joint infection and compared one-stage revision surgery with two-stage revision
- 15 surgery. Patients who were taking part in the trial were eligible for the DCE. Ethical approval for the
- 16 study was granted by NRES Committee South West Frenchay on 31 December 2014 (14/SW/1166).
- 17 All participants provided written informed consent to take part in the DCE study.
- 19 DCEs are an established method in health services research and have been used to explore a range
- of health-related services and treatments ¹¹⁻¹⁴. DCEs involve asking respondents to choose between
- 21 hypothetical scenarios which describe goods or services, where a 'service' may mean an intervention
- or an approach to care. The method aims to establish what attributes of that service influence their
- decision-making and to what extent. This enables quantification of the marginal impact of these

- 1 attributes. Scenarios within a DCE describe the service of interest (in this case, revision joint
- 2 replacement) using the same set of attributes, but at different levels in each scenario. Choices
- 3 between scenarios, or whether to accept or reject a scenario, are used to estimate the influence and
- 4 value of the different attribute levels.

Questionnaire development

- 7 When a patient faces revision surgery for prosthetic hip infection, a one-stage or a two-stage
- 8 operation is required, and the decision about which is undertaken is made largely at the discretion
- 9 of the surgeon, taking into account patient preferences for treatment. The DCE study was designed
- 10 to engage patients in their preferences for the features associated with these two surgical options,
- given neither is currently known to be clinically superior in terms of patient outcomes.

- 13 Qualitative methods are recommended for DCE attribute development because they enable
- 14 conceptual development of attributes directly from people's experiences and so better reflect the
- issues that are likely to matter most to people when making a decision ¹⁵. For this study we
- developed attributes from our previous qualitative study which explored impact of prosthetic joint
- infection and its treatment on patients and their recovery process; one-to-one interviews were
- conducted with 19 patients with PJI, focusing on the impact of PJI and surgical treatment. The
- 19 interviews were audio-recorded, transcribed and the qualitative data set was analysed thematically
- ⁶. Levels of attributes were assigned by refining the language used to convey the meaning of the
- 21 attributes, and particularly where some quantification of an attribute was mentioned by participants
- during qualitative interview, as illustrated in Table 1 15.

- Table 1 describes the attribute and level selection based on the earlier qualitative work, including
- 25 illustrative quotations and rationale.

2 The final questionnaire comprised a single scenario task in which respondents were asked to imagine

this was the first time that they had been offered the two surgical options presented and were asked

to consider them carefully before selecting the one (from a pair) they would prefer.

6 With a $4 \times 4 \times 2 \times 2$ design (two four-level and two two-level attributes), a total of 64 different

7 combinations of attribute levels (profiles) is possible. This converts to 32 pairs of profiles, which as a

'full factorial' was considered to be too large for participants to complete 16. An orthogonal main

effects plan was therefore used to reduce the number of choice sets to 16 ¹⁷. Each profile was

presented with its pair, and participants were required to select which option they preferred (see

11 Figure 1).

Patient and Public Involvement

The questionnaire was piloted and refined in collaboration with 5 patient and public involvement representatives. At an initial meeting of representatives, group members were involved in questionnaire development and suggested improvements to its formatting to aid readability, clearer phrasing of the questions to avoid ambiguity, and shortening of the instruction leaflet for clarity. At a subsequent meeting, members of the group completed the questionnaire and fed back on their experience. They completed the questionnaire without assistance and felt the instructions were clear but suggested that key points in the questions should be highlighted and that a contact telephone number should be added to enable participants to seek assistance if needed. For participants in the study, a summary of findings will be sent to those who indicated that they wished to be informed.

2 Participants

- 3 Patients who were recruited into the INFORM randomised controlled trial (ISRCTN10956306)
- 4 received the discrete choice task after completing the 18-month primary outcome measure. The
- 5 questionnaire was either posted to participants or completed in person with the assistance of a
- 6 research nurse during a hospital clinic visit.

8 Data analysis

- 9 Sample size calculations for DCEs are challenging due to dependence on the true parameter values
- 10 estimated in the choice model ¹⁸. However, reliable statistical analysis has been demonstrated with
- sample sizes of 40–120 respondents and combined with the rarity of periprosthetic joint infection
- 12 (1% of those undergoing hip replacement), a sample size above 50 participants for the DCE was
- deemed adequate to obtain sufficient data for exploratory analysis and interpretation ^{19, 20}.
- 15 Paper questionnaires were distributed to participants as part of the follow-up data collection in the
- 16 INFORM trial. Questionnaires were returned to the study team between January 2017 and
- 17 November 2018 and data entered onto a Microsoft Excel spreadsheet. DCE data were effects coded
- and analysed using STATA SE 15 ^{21, 22}. The influence of the four attributes on patient choices was
- analysed using a conditional logit model. As attribute levels are effects coded, the mean of all
- coefficients is 0 across each attribute. The effects coded preference weights (coefficients) produced
- by the conditional logit model are estimated relative to the mean effect of the attribute, with the p
- value indicating the statistical significance of "the difference between the estimated preference
- weight and the mean effect of the attribute" ²³.

3 Participants

Results:

- 4 Of the 80 discrete choice questionnaires provided to trial participants, 57 were returned fully
- 5 complete (71%) from patients from 9 trial sites. Data from 6 questionnaires was not used in analysis
- 6 as they were partially or totally incomplete. Responding participants had a mean age of 70 (range 51
- 7 to 90), 21 (37%) were female, 26 (46%) had undergone 1-stage revision, 14 (25%) lived alone and 41
- 8 (72%) were retired from work. Table 2 provides further demographic information.

10 Table 2 – Demographic and Clinical characteristics of the DCE Respondents

Characteristic	Participants (N=57)
Age – Yrs [mean] (range)	70 (51-90)
Gender [number] (%)),
Male	36 (63)
Female	21 (37)
Ethnicity [number] (%)	2/
White	55(96)
Black	1 (2)
Mixed	1 (2)
Marital Status [number] (%)	
Married / Partner	42 (74)
Divorced / Separated / Widowed	12 (21)
Single	3 (5)

Living Arrangements [number] (%)	
With Partner / somebody else	43 (75)
Alone	14 (25)
Schooling/Education [number] (%)	
Left at normal school leaving age	35 (61)
Left after normal school leaving age	15 (26)
Left before normal school leaving age	7 (12)
Work Situation [number] (%)	
Retired	41 (72)
Working / Sick leave	14 (25)
Unemployed	2 (4)
Surgery Received for Prosthetic Hip Joint Infection [number] (%)	
2-stage revision	31 (54)
1-stage revision	26 (46)

2 Discrete choice experiment

- 3 Table 3 shows the regression coefficients and the results from the 57 patients who had fully
- 4 completed the discrete choice questionnaire.

6 Table 3 – discrete choice task results from conditional logistic regression

Attribute	Level	Coefficient	Standard	95%	P value
			error	confidence	
				interval	

Ability to engage in	Can do	0.70			
valued activities after	everything*				
new hip is fitted.					
	Can do most	0.49	.08	.33 to .64	<0.001
	things				
	Cannot do	-0.39	.07	53 to24	<0.001
	most things				
	Cannot do	-0.80	.13	-1.05 to -	<0.001
	anything			0.55	
	100				
Antibiotic side effects	Don't affect	0.22			
	me much*	4			
	Affects me a	-0.22	.05	33 to12	<0.001
	lot	7.			
Number of operations	1*	0.20	7		
	2	-0.20	.07	35 to06	<0.001
				>	
Time taken after	3months*	0.20			
surgical treatment					
starts, to return to					
normal activities.					
	6months	0.31	.09	.14 to .48	<0.001
	12months	-0.06	.05	15 to .04	0.22
	18months	-0.45	.10	64 to26	<0.001

*Indicates reference category within attribute

Analysis indicates that participants had the strongest preference for a surgical option that resulted in the least restrictions on engaging in valued activities after the new hip is fitted, illustrated by the largest preference weight. Other less valued but important preferences were for a surgical strategy that would result in a shorter time after surgical treatment starts to return to normal activities, few or no side effects from antibiotics, and only one operation. The results also suggest that the least restrictions on engaging in valued activities, and the shortest time taken to return to normal activity are the individual attributes most valued by patients in this sample. This is indicated by the larger spread of coefficients (i.e. more 'value' is placed on changes in these attributes). The most acceptable option was a time period of between 3 to 6 months to return to normal activity, however

there is no clear preference up to 12 months, although 18 months appeared to be significantly

Discussion:

disfavoured.

This study aimed to investigate and understand patients' preferences for aspects of revision surgery for periprosthetic joint infection. Four relevant attributes were identified through earlier qualitative work, and quantitatively, patients in this study most value the ability to engage in valued activities and the time taken to return to normal daily activities. This reflects the findings of our previous qualitative work which shows that although both revision strategies impacted greatly on patients and their families, patients receiving two-stage revision surgery experience particularly long periods of immobility and social isolation. This was often followed by a protracted recovery period, which could leave patients much less able than before their primary operation, and some patients

1 experienced profoundly negative psychological effects associated with physical suffering, loss of

2 dignity and independence ⁶. It appears that for patients in our sample, 3-6 months to return to

normal activity was preferable, although there was no significant difference up to 12 months, but 18

months was disfavoured. This suggests that the acceptable margin of recovery for patients is up to

12 months after their receipt of a new hip joint.

7 Discrete choice methodology can be challenging for participants because the format of questions is

different to standard surveys and items can seem repetitive. We collected feedback from the first 11

participants who completed the questionnaire. We found that those participants who were

supported by a research nurse when completing the questionnaire were more likely to complete it

and return it, compared with those who received the questionnaire by post and completed it alone.

Participant feedback suggested that the questionnaire was difficult to complete, as the scenarios

were similar and appeared to be repetitive. To address this, we amended the questionnaire format

and instructions and offered participants support either face-to-face or by phone with one of the

study research nurses. Nurses were then able to answer queries about the questionnaire and offer

support if needed. The results suggest the group completed the questionnaire in a rational and

logical way, meaning the results were as expected with patients selecting an optimal combination of

options as their preferred choice. This is an important methodological finding because although our

study demonstrates the feasibility of the DCE method with this population, others conducting similar

studies with older, ill populations could consider in advance the need for professional support in the

completion of discrete choice questionnaires.

Participants were all individuals who participated in a clinical trial and had already undergone

revision surgery for periprosthetic joint infection. This meant that the choices that participants were

25 asked to make in the questionnaire were based on scenarios unlikely to reflect their real-life

experiences as in reality such choices would not be available to them, since decisions about surgical

strategies are based on a wider variety of clinical, surgeon, patient and organisational factors ⁹. Also,
patients were not being faced with these decisions at the time of questionnaire completion as we
decided that it would be unethical to ask patients awaiting treatment to complete a DCE about
surgical options in a hypothetical context, at a time when they may be particularly vulnerable. To do
so would not meet an ethical standard of protection from harm as it might mean that patients were
inadvertently led to believe that there were more or different options available to them than were

clinically indicated at that time.

We also found that some participants reported difficulty separating their own recent personal experience of revision surgery from the hypothetical scenarios presented in the questionnaire, as they found it hard to imagine receiving a treatment option that differed from the one that they had received. In terms of methodology, many DCE's are conducted with participants who already have some experience of the treatment attributes under investigation ^{12, 19, 20, 24}. In our study previous experience meant that participants had some appreciation of the attributes being tested. While, the sample size of 57 participants provided sufficient data for the analyses presented here, the sample size was too small to conduct any subgroup analysis to identify whether preferences would differ between participants who had received one or two stage revision. Similarly, patients who are older, still working or live alone may have had stronger preferences for a one-stage operation than those who are younger, retired and have support at home to cope with a two-stage procedure. The majority of the study sample were also white, male and educated, which means that results may not reflect the preferences of the wider surgical population. Further research could explore preferences in a more diverse population.

This work has provided an initial and important first step in understanding patients' preferences for characteristics associated with revision surgery for periprosthetic infection. It is important that

- 1 orthopaedic healthcare professionals discuss these attributes with patients when discussing options
- 2 for surgical and antibiotic treatment for periprosthetic infection.

Implications

- 5 Results of this study offer insight into the preferences of patients for revision surgery and provide
- 6 valuable information to surgeons from all disciplines. Although factors affecting patient preferences
- 7 for surgery differ to those valued by medical professionals, consideration should be given to such
- 8 factors in order to aid shared decision-making where clinical equipoise between options exists.

Previous research using discrete choice approaches has explored patient preferences in a surgical context. This has included examination of preferences for surgical versus non-surgical interventions in a range of conditions, such as oesophageal cancer and ulcerative colitis ^{24, 25}. Some research has explored preferences for conditions in which there are two surgical options, including for ectopic pregnancy, vaginal wall prolapse, and osteoarthritis ^{26, 27}. Across all of these conditions, evidence suggests that patients choose options that reduce the need for further surgery or operations, have a shorter recovery time, lower risk of symptom recurrence and improve ability to preserve existing joint motion in the case of osteoarthritis. Findings from the current study are similar in that patients prefer a surgical option that reduces the number of operations, recovery time and the side effects of antibiotics. However, in the setting of infected joint replacement, patients placed highest value on restoration of function. This was more important to the patients in our study than the number of operations they would have to undergo. Although our study focused on the preferences between one and two-stage surgery surgeons may need to consider these preferences during shared decision-making about all options for revision surgery for prosthetic hip infection, including the role of debridement with retention of implants (DAIR). Although DAIR is only efficacious in approximately

60% of cases, it is associated with a quicker return to valued activities and improved joint function 28.

Conclusions

Our results show that the most valued characteristics in decisions about revision surgery for prosthetic hip infection were the ability to engage in valued activities and time taken to return to normal activity. This builds on the findings of our previous qualitative work which shows that although both revision strategies impact greatly on patients and their families' everyday lives, patients receiving two-stage revision surgery experience particularly long periods of immobility and social isolation ⁶. The desire to return to everyday activities should be taken into account when surgeons are discussing options with patients, particularly when there is equipoise from a surgical

perspective about the options available, and when the decision is 'preference sensitive'.

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Author Contributions

- FEC: Contributed to design, analysis and interpretation of data, drafting and revising the manuscript.
- RG-H: Contributed to conception, design, interpretation of data, drafting and revising the
- manuscript. SS: Contributed to design, data collection and revising the manuscript. AWB:
- Contributed to the conception, design, interpretation of data and revising the manuscript. AJM:

- 1 Contributed to the design, interpretation of data, drafting and revising the manuscript. All authors
- 2 gave final approval of the submitted manuscript.

3 Competing Interests

4 None declared

5 Data Availability

- 6 Anonymised research data will be made available at the University of Bristol Research Data
- 7 Repository (data.bris) beginning 1 year after publication of the results. Access will be
- 8 granted to bona fide researchers after the University of Bristol Data Access Committee has
- 9 approved their request. Please contact data-bris@bristol.ac.uk in the first instance.

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- 17 of the questionnaire.

Table 1 – qualitative support for attributes included in the discrete choice questionnaire

Attribute	Evidence of attribute	Levels	Rationale for
	inclusion, with pseudonym		levels
	and surgery type		
Number of operations	"There's no way I want two	1) 1 operation	Two types of
	more big operations now at	2) 2 operations	revision surgery
	my time of life. You do it all		are currently
	or not at all"I said there		provided in
	was no way I wanted two		healthcare, and
	ops" (Harriet, 1-stage)	•	involve either
		4	one or two
			operations
	"Of course, emotionally,		
	you want it over and done	7/	
	with as soon as		
	possiblebut ultimately		
	that has to be done in the		
	correct way. There's the		
	tortoise and hare situation.		
	There's absolutely no point		
	in rushing ahead if		

	ultimately it's going to fail."			
	(Maggie, 2-stage)			
Ability to engage in	"Fourteen months without	1)	Can do	Following
valued activities after	a hip joint so it meant that I		everything	revision
new hip is fitted.	couldn't drive a car, I	2)	Can do most	surgery, the
	couldn't do anything that		things	ability to
	I'd been used to doing,			engage in
	playing golf or doing	3)	Cannot do	valued
	anything. Well, I gave up		most things	activities can be
	golf actually after the first	4)	Cannot do	reduced in a
	revision" (Don, 2-stage)		anything	major, or
				somewhat
				more minor
	"But when I for example			way. These
	went to, on holiday	7		levels capture
	recently and I had serious			variation in
	problems getting into the			ability
	bath to stand in the			identified by
	shower. Because I, I			patients.
	couldn't get in. And in a			
	wet – on a wet surface and			
	that, I'm very conscious of			
	not falling in. I can't afford			
	to fall. So, pain I've got			

stiffness none.		
cal function		
tions, and that's one		
em." (Rory, 1-stage)		
im has always been		
back on my feet as		
as I can, and to walk		
od as I can, and that's		
disappointment. I'm		
here I think I should		
been" (Robert, 2-		
n't want to go 14	1) 3 months	These time
s with effectively one	2) 6 months	intervals
/hat was worse was	2) o months	demonstrate
nowing that I had to	3) 12 months	best
re all of those weeks	4) 18 months	approximations
nowing that I was ever		and a reflection
to get another hip		of the need to
oack." (Maggie, 2-		ensure normal
		expectations of
		time taken for
		soft tissue
		recovery as
	aim has always been back on my feet as as I can, and to walk od as I can, and that's disappointment. I'm here I think I should been" (Robert, 2-) n't want to go 14 s with effectively one was mowing that I had to be all of those weeks mowing that I was ever to get another hip back." (Maggie, 2-)	aim has always been back on my feet as as I can, and to walk od as I can, and that's disappointment. I'm here I think I should been" (Robert, 2-) n't want to go 14 s with effectively one //hat was worse was nowing that I had to re all of those weeks nowing that I was ever to get another hip back." (Maggie, 2-

"If I had known how hard it expected by was going to be for her to surgeons. 18 walk in that interim six months is the months, and if there maximum end seemed to have been a point that reasonable, or a good surgeons would possibility that the suggest for infection would be nuked in recovery time. a one stage, then that might have been a better outcome for her" (Amelia, 2-stage) "I would've thought that if you wanted to go back to work...you wouldn't be very happy [having a 2-stage operation] because you wouldn't be able to do nothing for those six weeks again. Then you'd have to go all through it again after three months of having it done. Six weeks is only a month and a half, and then

	in another month and a			
	half you're having it all			
	done again." (Jim, 1-stage)			
Antibiotic side effects	"the nightmare on heavy	1)	Affects me a	Antibiotics are
	antibiotics, toiletry wise.		lot	an essential
	Now I've had to move into			method of
	a, another bed. My wife			attempting to
	and I are married 50 plus	2)	Don't affect	ensure the
	years, and I have to have		me much	periprosthetic
	my own room because I'm			joint infection
	getting up in the night."			is treated and
	(Rory, 1-stage)			subsequently
				clear. For some
				patients, the
	"I stayed on antibiotics			impact of these
	then for, for everand			antibiotics is
	after the first [week of			significant,
	antibiotics] I was just dying.			while for others
	I just wanted to lie on the			there are less
	floor and die. I felt so sick.			severe side
	So ill." (Lottie, 2-stage)			effects.
	"I felt fine. When I was			
	going to see the surgeon			
	James and Sungarium			

They'd say, 'How are you	
today?' I'd say, 'I feel fine,	
fit! [laughs] I feel well in	
myself, I eat well.'" (Ray, 2-	
stage)	

References

- 1. National Joint Registry. National Joint Registry 14th Annual Report 2017: National Joint Registry for England, Wales, Northern Ireland and the Isle of Man; 2017 [Available from: www.njrreports.org.uk.
- 2. Blom AW, Taylor AH, Pattison G, Whitehouse S, Bannister GC. Infection after total hip arthroplasty. The Avon experience. The Journal of bone and joint surgery British volume. 2003;85(7):956-9.
- 3. Phillips JE, Crane TP, Noy M, Elliott TS, Grimer RJ. The incidence of deep prosthetic infections in a specialist orthopaedic hospital: a 15-year prospective survey. The Journal of bone and joint surgery British volume. 2006;88(7):943-8.
- 4. Lenguerrand E, Whitehouse MR, Beswick AD, Jones SA, Porter ML, Blom AW. Revision for prosthetic joint infection following hip arthroplasty: Evidence from the National Joint Registry. Bone & joint research. 2017;6(6):391-8.
- 5. Zimmerli W, Trampuz A, Ochsner PE. Prosthetic-joint infections. The New England journal of medicine. 2004;351(16):1645-54.
- 6. Moore AJ, Blom AW, Whitehouse MR, Gooberman-Hill R. Deep prosthetic joint infection: a qualitative study of the impact on patients and their experiences of revision surgery. BMJ open. 2015;5(12):e009495.
- 7. Kunutsor SK, Whitehouse MR, Blom AW, Beswick AD. Re-Infection Outcomes following Oneand Two-Stage Surgical Revision of Infected Hip Prosthesis: A Systematic Review and Meta-Analysis. PloS one. 2015;10(9):e0139166.
- 8. Kunutsor SK, Whitehouse MR, Blom AW, Board T, Kay P, Wroblewski BM, et al. One- and two-stage surgical revision of peri-prosthetic joint infection of the hip: a pooled individual participant data analysis of 44 cohort studies. European journal of epidemiology. 2018;33(10):933-46.
- 9. Moore AJ, Blom AW, Whitehouse MR, Gooberman-Hill R. Managing uncertainty a qualitative study of surgeons' decision-making for one-stage and two-stage revision surgery for prosthetic hip joint infection. BMC musculoskeletal disorders. 2017;18(1):154.
- 10. Strange S, Whitehouse MR, Beswick AD, Board T, Burston A, Burston B, et al. One-stage or two-stage revision surgery for prosthetic hip joint infection--the INFORM trial: a study protocol for a randomised controlled trial. Trials. 2016;17:90.
- 11. Louviere J, Street DJ, Burgess L. A 20+ years' retrospective on choice experiments. Marketing research and modeling: Progress and prospects: Springer; 2004. p. 201-14.

- de Bekker-Grob EW, Hofman R, Donkers B, van Ballegooijen M, Helmerhorst TJ, Raat H, et al. Girls' preferences for HPV vaccination: a discrete choice experiment. Vaccine. 2010;28(41):6692-7.
- 13. Liede A, Mansfield CA, Metcalfe KA, Price MA, Snyder C, Lynch HT, et al. Preferences for breast cancer risk reduction among BRCA1/BRCA2 mutation carriers: a discrete-choice experiment. Breast cancer research and treatment. 2017;165(2):433-44.
- 14. van Dijk JD, Groothuis-Oudshoorn CG, Marshall DA, MJ IJ. An Empirical Comparison of Discrete Choice Experiment and Best-Worst Scaling to Estimate Stakeholders' Risk Tolerance for Hip Replacement Surgery. Value in health: the journal of the International Society for Pharmacoeconomics and Outcomes Research. 2016;19(4):316-22.
- 15. Coast J, Al-Janabi H, Sutton EJ, Horrocks SA, Vosper AJ, Swancutt DR, et al. Using qualitative methods for attribute development for discrete choice experiments: issues and recommendations. Health economics. 2012;21(6):730-41.
- 16. Marshall D, Bridges JF, Hauber B, Cameron R, Donnalley L, Fyie K, et al. Conjoint Analysis Applications in Health How are Studies being Designed and Reported?: An Update on Current Practice in the Published Literature between 2005 and 2008. The patient. 2010;3(4):249-56.
- 17. Hedayat AS, Sloane NJA, Stufken J. Orthogonal arrays: theory and applications: Springer Science & Business Media; 2012.
- 18. Lancsar E, Louviere J. Conducting discrete choice experiments to inform healthcare decision making: a user's guide. PharmacoEconomics. 2008;26(8):661-77.
- 19. Salkeld G, Solomon M, Butow P, Short L. Discrete-choice experiment to measure patient preferences for the surgical management of colorectal cancer. The British journal of surgery. 2005;92(6):742-7.
- 20. de Bekker-Grob EW, Essink-Bot ML, Meerding WJ, Koes BW, Steyerberg EW. Preferences of GPs and patients for preventive osteoporosis drug treatment: a discrete-choice experiment. PharmacoEconomics. 2009;27(3):211-9.
- 21. Bech M, Gyrd-Hansen D. Effects coding in discrete choice experiments. Health economics. 2005;14(10):1079-83.
- 22. StataCorp. Stata Statistical Software: Release 15. College Station, TX: StataCorp LLC; 2017.
- 23. Hauber AB, Gonzalez JM, Groothuis-Oudshoorn CG, Prior T, Marshall DA, Cunningham C, et al. Statistical Methods for the Analysis of Discrete Choice Experiments: A Report of the ISPOR Conjoint Analysis Good Research Practices Task Force. Value in health: the journal of the International Society for Pharmacoeconomics and Outcomes Research. 2016;19(4):300-15.
- 24. Noordman BJ, de Bekker-Grob EW, Coene P, van der Harst E, Lagarde SM, Shapiro J, et al. Patients' preferences for treatment after neoadjuvant chemoradiotherapy for oesophageal cancer. The British journal of surgery. 2018;105(12):1630-8.
- 25. Bewtra M, Kilambi V, Fairchild AO, Siegel CA, Lewis JD, Johnson FR. Patient preferences for surgical versus medical therapy for ulcerative colitis. Inflammatory bowel diseases. 2014;20(1):103-14.
- 26. van Mello NM, Mol F, Opmeer BC, de Bekker-Grob EW, Essink-Bot ML, Ankum WM, et al. Salpingotomy or salpingectomy in tubal ectopic pregnancy: what do women prefer? Reproductive biomedicine online. 2010;21(5):687-93.
- 27. Notten KJ, Essers BA, Weemhoff M, Rutten AG, Donners JJ, van Gestel I, et al. Do patients prefer mesh or anterior colporrhaphy for primary correction of anterior vaginal wall prolapse: a labelled discrete choice experiment. BJOG: an international journal of obstetrics and gynaecology. 2015;122(6):873-80.
- 28. Kunutsor SK, Beswick AD, Whitehouse MR, Wylde V, Blom AW. Debridement, antibiotics and implant retention for periprosthetic joint infections: A systematic review and meta-analysis of treatment outcomes. The Journal of infection. 2018;77(6):479-88.

Legend: Figure 1. Example profile of DCE



OPTION H1		OPTION H2		
I take antibiotics and the side effects don't affect		I take antibiotics and the side effects affect me a		
me much		lot		
and		and		
I have two operations		I have one operation		
and	OR	and		
After my new hip is fitted, I can do most of the		After my new hip is fitted, I cannot do most of the		
things that I want to do		things that I want to do		
and		and		
After my surgical treatment starts, it takes 6		After my surgical treatment starts, it takes 12		
months to get back to the things that I normally		months to get back to the things that I normally		
do		do		
Please only tick ONE box:				

Figure 1. Example profile of DCE 160x90mm (300 x 300 DPI)

OPTION H2

OPTION H1