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knee or hip surgery due to

BMJ Open Can obesity and physical activity predict outcomes of elective knee or hip surgery due to osteoarthritis? A metaanalysis of cohort studies

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ABSTRACT

Objective The aim of this study was to systematically review the literature to identify whether obesity or the regular practice of physical activity are predictors of clinical outcomes in patients undergoing elective hip and knee arthroplasty due to osteoarthritis.

Design Systematic review and meta-analysis. Data source and eligibility criteria A systematic search was performed on the Medline, CINAHL, EMBASE and Web of Science electronic databases. Longitudinal cohort studies were included in the review. To be included, studies needed to have assessed the association between obesity or physical activity participation measured at baseline and clinical outcomes (ie, pain, disability and adverse events) following hip or knee arthroplasty.

Data extraction Two independent reviewers extracted data on pain, disability, quality of life, obesity, physical activity and any postsurgical complications.

Results 62 full papers were included in this systematic review. From these, 31 were included in the metaanalyses. Our meta-analysis showed that compared to obese participants, non-obese participants report less pain at both short term (standardised mean difference (SMD) -0.43; 95% CI -0.67 to -0.19; P<0.001) and long term post-surgery (SMD -0.36; 95% CI -0.47 to -0.24; P<0.001), as well as less disability at long term postsurgery (SMD -0.32; 95% CI -0.36 to -0.28; P<0.001). They also report fewer postsurgical complications at short term (OR 0.48: 95% Cl 0.25 to 0.91: P<0.001) and long term (OR 0.55; 95% CI 0.41 to 0.74; P<0.001) along with less postsurgical infections after hip arthroplasty (OR 0.33; 95% CI 0.18 to 0.59; P<0.001), and knee arthroplasty (OR 0.42; 95% CI 0.23 to 0.78; P=0.006).

Conclusions Presurgical obesity is associated with worse clinical outcomes of hip or knee arthroplasty in terms of pain, disability and complications in patients with osteoarthritis. No impact of physical activity participation has been observed.

PROSPERO registration number CRD42016032711.

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INTRODUCTION

Musculoskeletal pain, including pain from knee and hip osteoarthritis (OA), is the leading cause of physical disability in the

Strengths and limitations of this study

- ► The current review is the most comprehensive systematic review on the topic to date.
- The current review is the first review to use a quantitative approach to synthesise the results of pain, disability and surgical complications between non-obese and obese participants who underwent hip or knee arthroplasty due to osteoarthritis.
- The methodological quality of the included studies was in general poor.
- There was a substantial variability of follow-up duration across studies, ranging from 2 weeks to 11 years.

Protected by copyright, including for uses related to text and world and responsible for an increasing burden to patients and society.¹ This problem will increase over time, as the world population ages and physical disability resulting from declining health becomes increasingly . ک prevalent.² The global healthcare expenditure for knee and hip OA is substantial, and most of these costs are incurred by surgical management and associated hospital care.³ For instance, in the UK, the direct costs of **a** OA were estimated at more than £1 billion in 2010, of which £850 million was spent just on surgical procedures.⁴

Although management of the early stages of this condition consists of a combination of one non-pharmacological and pharmacological of therapies (eg, anti-inflammatory and analgesic drugs), surgery has become the most common treatment option for severe cases, especially when non-surgical therapies fail to provide sufficient pain relief.⁵ Osteotomy, mosaicplasty and arthroplasty are some of the existing types of surgery used to manage OA of the hip and knee; with total or partial arthroplasty being the most commonly recommended.^b

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There are multiple risk factors for the development of knee OA. Among the most common of these are increased body weight and muscle weakness; often attributed to a sedentary lifestyle.⁷ Obesity and sedentary lifestyle behaviour have also been associated with serious health conditions such as: coronary heart disease, type 2 diabetes, breast and colon cancers and decreased life expectancy.⁸ Although there is evidence for the role of obesity and physical inactivity in health conditions and quality of life in general,^{9 10} the actual impact of these factors, together or in isolation, on the outcomes of elective surgery of the knee and hip is still controversial.¹¹¹² Although previous attempts to systematically review the literature have been made, these studies^{13–15} have either failed to perform a quantitative summary of the evidence (ie, meta-analysis), have excluded patients undergoing knee arthroplasty¹⁶ or have excluded pain outcomes.¹³ No meta-analyses have been performed considering obesity and physical activity as predictors of surgical outcomes in terms of pain, disability, quality of life and complications after hip or knee arthroplasty for end-stage OA.

Identifying whether obesity and physical activity participation predict surgical outcomes in patients with knee and hip OA will inform clinical practice in terms of prognosis and safety of an increasingly prevalent treatment approach. We have conducted a meta-analysis of cohort studies aiming to quantify the role of obesity and physical activity participation as predictors of clinical outcomes in terms of pain, disability, quality of life and postsurgical complications. This review and meta-analysis focused on patients with knee and hip OA undergoing hip or knee arthroplasty.

METHODS

Data sources and searches

We conducted a systematic review following the PRISMA statement.¹⁷ This review was prospectively registered on PROSPERO, registration number CRD42016032711. A systematic electronic search was performed in the following databases from inception to January 2017: MEDLINE, EMBASE, CINAHL and Web of Science. We used a combination of relevant keywords to construct the search strategy including obesity, physical activity, knee OA, hip OA, arthroplasty and elective surgery (online supplementary appendix 1). The first screening of potentially relevant records was conducted by one author (DP) based on titles and abstract, and two authors (DP and GCM) independently performed the final selection of included trials based on full-text evaluation. A third reviewer arbitrated in case of disagreement (MLF). Moreover, the reference lists of included studies were checked for potential studies. An additional 26 references were screened, but none met our inclusion criteria. No restriction was applied on language.

≥

Study selection

We included only longitudinal studies assessing the role of obesity or physical activity participation on the clinical outcomes following partial or total hip arthroplasty (THA) or partial or total knee arthroplasty (TKA) surgery. Clinical outcomes were defined in terms of pain, disability, quality of life and complications after arthroplasty. To be eligible, studies had to be full reports; include participants who underwent elective arthroplasty of the hip or knee due to OA; include data of presurgical and at least one postsurgical assessment of the clinical outcomes of interest; and assessment of the emilian outcomes of and outcomes of interest. Obesity and physical activity ş participation had to be assessed at baseline. Studies on copyright, inc revision surgery were excluded. Studies were not excluded based on intensity or duration of symptoms.

Data extraction

Using a standardised form, data on study characteristics, predictors and outcome measures of interest were independently extracted from the included studies by two reviewers (DP and GCM). A third author (MLF) resolved any disagreement. Estimates of association between predictors and outcomes of interest were extracted as presented in each study and included ORs, risk ratios (RR), correlations, mean differences (MD) or regression coefficients. When studies reported more than one tool regarding the same topic (eg, Western Ontario and McMaster Universities Osteoarthritis Index text (WOMAC), Hip Disability and Osteoarthritis Outcome Score (HOOS), Oxford Hip Score (OHS), Knee Injury and Osteoarthritis Outcome Score (KOOS), Knee Society and Osteoarthritis Outcome Score (KOOS), Knee Society Score (KSS)), estimates were extracted from the group with the largest sample size.

We contacted the authors to provide further information when there were insufficient data reported in the manuscript. When authors were unavailable we estimated data using the recommendations in the Cochrane Handbook for Systematic Reviews of Interventions.¹⁸

Outcome measures

training, and Data on pain intensity were extracted as visual analogue <u>0</u> scale scores ranging from 0 to 10 and measured directly or as part of the following measurement tools: the WOMAC, the HOOS, the KOOS or the Harris Hip Score (HHS). If studies reported more than one measure of pain intensity or disability for the cohort, the most severe measure at **O** baseline was included in the pooled analyses. Disability & measures included the OHS ranging from 12 to 60, with 12 being the best result; Oxford Knee Score ranging from 0 to 60, with 60 being the best result; the HHS ranging from 0 to 100, with 100 being the best result; KSS ranging from 0 to 100, with 100 being the best result; WOMAC total score ranging from 0 to 96, with 0 being the best result; or WOMAC function subscale ranging from 0 to 10, with 10 being the best result; and were converted into a uniform 0-100 scale where 0 meant less disability. Extracted data on complications included any descriptive

measure of the number of complications or number of patients with a complication reported during the study. Only two of the screened studies had reported specific raw data on quality of life among the participants after joint arthroplasty, but due to differences in follow-up length, any meta-analysis made by merging these data would result in an unreliable measure.

Methodological guality assessment

The methodological quality of included studies was assessed by two independent reviewers using the Newcastle-Ottawa Scale (NOS)¹⁹ recommended by the Cochrane Collaboration.¹⁸ The NOS consists of eight items grouped into three categories, namely: selection, comparability and outcome. A star system, ranging from zero to nine stars, is used to classify the quality of the study being reviewed (the more stars the study receives in each category, the higher its methodological quality). After the independent assessment of included studies by the leading author, each study received the following categorical scores representing its quality: good (three or four stars in selection domain AND one or two stars in comparability domain AND two or three stars in outcome domain), fair (two stars in selection domain AND one or two stars in comparability domain AND two or three stars in outcome domain) or poor (zero or one star in selection domain OR zero star in comparability domain OR zero or one star in outcome domain). A third reviewer (MLF) resolved any disagreement between independent assessors. Methodological quality scores for included studies are presented in table 1.

Data analysis

Data on baseline (ie, presurgical scores) and postoperative outcome scores were weighed by the inverse study variance and used in fractional polynomial regression modelling to build graphs depicting the course of pain and disability over time. STATA V.14 was used for the analyses (StataCorp, College Station, TX).²⁰

Meta-analyses were performed to assess the differences in pain, disability and complications after surgery, between predictor groups (ie, obese and non-obese groups as defined by included studies), using a random effects model. When possible, different analyses were performed for knee and hip arthroplasty and also for different levels of obesity (ie, obesity and morbid obesity). When means and SD of outcomes of interest were presented for multiple predictor groups in the same study (ie, underweight (body mass index (BMI)<18), normal weight (BMI≥18<25), overweight (BMI≥25<30) and obese level I (BMI≥30<35), II (BMI≥35<40) or III (BMI≥40)) these were combined into two groups (non-obese: BMI<30 and obese: BMI≥30) as recommended in the Cochrane Handbook for Systematic Reviews of Interventions¹⁸ before inclusion in the pooled analyses. Results were reported as standardised mean differences (SMD) and 95% CIs. Between-study heterogeneity was calculated using I^2 $(I^2 < 25\%)$: small heterogeneity; $25\% < I^2 < 75\%$: moderate

heterogeneity; $I^2 > 75\%$: large heterogeneity).²¹ We have defined an SMD of 0.2 as small difference, 0.5 as moderate difference and 0.8 as large difference.²²

Assessment of publication bias was performed using funnel plots. The precision (ie, SE) of included studies was plotted against the difference in outcomes between groups (ie, obese or non-obese) and results visually analysed. In the absence of publication bias or small study bias, smaller studies should be evenly spread around the base of the funnel, while the larger studies should be concentrated around the top of the funnel. Plot asymmetry was also quantified using the Egger's tests, for which a null hypothesis represents symmetry of plotted data.²³

All meta-analyses were conducted using Comprehen-All meta-analyses were conducted using Comprehen-sive Meta-Analysis software (Comprehensive Meta-Anal-ysis, Englewood, NJ). For studies not reporting enough data to be included in the meta-analyses, the reported individual associations were tabulated and qualitatively presented in tables 2–5. **RESULTS** Our search strategy identified 11990 studies. After removing 381 duplicates, 11221 studies were screened and excluded based on keywords titles and abstracts All

and excluded based on keywords, titles and abstracts. All relat the remaining 388 studies were written in English and fed were assessed by reading the full text, of which 326 were then excluded, yielding 62 studies to be included in the **ö** systematic review.^{24–85} From these, 31 presented enough text data to be included in at least one of the meta-analyses and (figure 1).

Included studies

data mining, Included studies reported data from 18 different countries: Australia,^{40 47 72 85} Canada,^{38 43 78} China,⁸⁴ Denmark,⁶⁰ England,^{27 30} Finland,^{49–52} France,^{65 73} Germany, ⁵⁵⁷⁵⁸¹ Italy, ²⁸²⁹ Japan, ⁸³ Netherlands, ⁵⁷⁷⁶ Norway, ⁴⁵ Scotland, ^{25 36} South Korea, ⁵⁶ Spain, ^{41 80} Switzerland, ^{61 62 69} UK²⁶³⁵³⁷⁴⁶⁴⁸⁵³⁶³⁶⁷⁶⁸⁷¹⁷⁴ and USA. ^{2431–34394244545859646670777982} ≥ training, and Demographic data from each study are presented in table 1.

Methodological quality

simi An overall quality assessment of the studies showed that 50% (n=31) of the included studies were considered as were considered fair and 48.5% (n=30) were considered fair and 48.5% (n=30) were considered fair and 48.5% (n=30) were considered of poor methodological quality. Of the screened studies, 56 (90%) had a follow-up rate of 80% or greater, and only **g** half (n=32 studies) assessed outcomes via setter half (n=32 studies) assessed outcomes via retrospective analysis of medical records, conducted adjustment for potential confounders (eg, age or sex) or investigated a representative sample of the population (online supplementary appendix 2).

Assessment of publication bias

Inspection of funnel plots and results of Egger's test confirmed no evidence of small study bias for those studies included in our pooled analyses, with P values

Table 1 Included studies and characteristics							
Author, year	Country	Sample size	Predictor	Outcomes	Surgery	Follow- up duration	Quality score
AbdelSalam <i>et al</i> , 2012 ²⁴	USA	210	Obesity	Complications	Total hip and knee arthroplasty	9 years	Fair
Amin <i>et al</i> , 2006 ²⁶	UK	328	Obesity	Complications; disability	Total knee replacement	6, 18, 36 and 60 months	Poor
Amin <i>et al</i> , 2006 ²⁵	Scotland	82	Obesity	Complications	Total knee replacement	38.5 months	Poor
Andrew <i>et al</i> , 2008 ²⁷	England	1059	Obesity	Complications; disability	Total hip arthroplasty	3, 12, 24, 36 and 60 months	Poor
Sadr Azodi <i>et al</i> , 2006 ²⁹	Italy	3309	Obesity	Complications	Total hip replacement	6–9 years	Fair
Sadr Azodi <i>et al</i> , 2008 ²⁸	Italy	2106	Obesity	Complications	Total knee arthroplasty	2 years	Fair
Baker <i>et al</i> , 2012 ³⁰	England	13673	Obesity	Complications; disability	Total hip arthroplasty	6 months	Fair
Belmont <i>et al</i> , 2014 ³¹	USA	17514	Obesity	Complications	Total knee arthroplasty	1 month	Fair
Belmont <i>et al</i> , 2014 ³²	USA	15321	Obesity	Complications	Total knee arthroplasty	1 month	Fair
Bozic <i>et al</i> , 2012 ³⁴	USA	40919	Obesity	Complications	Total hip arthroplasty	10 years	Fair
Bozic <i>et al</i> , 2012 ³³	USA	83011	Obesity	Complications	Total knee arthroplasty	10 years	Fair
Chee <i>et al</i> , 2010 ³⁵	UK	106	Obesity	Complications; disability	Total hip arthroplasty	6, 18, 36 and 60 months	Good
Chesney <i>et al</i> , 2008 ³⁶	Scotland	1278	Obesity	Complications	Total knee arthroplasty	6, 18 and 60 months	Poor
Collins <i>et al</i> , 2012 ³⁷	UK	385	Obesity	Complications; disability	Total knee arthroplasty	6 and 18 months 3, 6 and 9 years	Poor
Davis <i>et al</i> , 2011 ³⁸	Canada	931	Obesity	Pain	Total hip and knee arthroplasty	2 weeks 1, 3, 6 and 12 months	Fair
Dewan <i>et al</i> , 2009 ³⁹	USA	220	Obesity	Complications; disability	Total knee arthroplasty	5.4 years	Poor
Dowsey and Choong, 2008 ⁸⁵	Australia	1207	Obesity	Complications	Hip arthroplasty	1 year	Poor
Dowsey <i>et al</i> , 2010 ⁴⁰	Australia	471	Obesity	Complications; pain; disability	Total hip arthroplasty	1 year	Good
Font-Vizcarra et al, 2011 ⁴¹	Spain	402	Obesity	Complications	Total hip arthroplasty	3 months	Fair
Friedman <i>et al</i> , 2013 ⁴²	USA	12355	Obesity	Complications	Hip and knee arthroplasty	2 months	Poor
Gandhi et al, 2010 ⁴³	Canada	1224	Obesity	Pain; disability	Total hip arthroplasty	1 year	Good
Hamoui <i>et al</i> , 2006 ⁴⁴	USA	63	Obesity	Disability	Total knee arthroplasty	11.3 years	Poor
Heiberg <i>et al</i> , 2013 ⁴⁵	Norway	64	Obesity	Pain	Total hip arthroplasty	3 and 12 months	Good
Ibrahim <i>et al</i> , 2005 ⁴⁶	UK	343	Obesity	Complications	Total hip arthroplasty	1 year	Poor
Jackson <i>et al</i> , 2009 ⁴⁷	Australia	100	Obesity	Complications; pain; disability	Total knee replacement	9.2 years	Poor
						(Continued

Continued

Table 1 Continued							
Author, year	Country	Sample size	Predictor	Outcomes	Surgery	Follow- up duration	Quality score
Jameson <i>et al</i> , 2014 ⁴⁸	UK	5535	Obesity	Disability	Hip arthroplasty	6 months	Fair
Jämsen <i>et al</i> , 2010 ⁵⁰	Finland	2647	Obesity	Complications	Total knee arthroplasty	1 year	Good
Jämsen <i>et al</i> , 2012 ⁴⁹	Finland	7181	Obesity	Complications	Total knee arthroplasty	1 year	Good
Järvenpää <i>et al</i> , 2010 ⁵¹	Finland	100	Obesity	Complications; pain	Total knee arthroplasty	3 months	Poor
Järvenpää <i>et al</i> , 2012 ⁵²	Finland	52	Obesity	Pain; disability	Total knee arthroplasty	10.8 years	Poor
Judge <i>et al</i> , 2010 ⁵³	UK	908	Obesity	Disability	Hip replacement	1 year	Poor
Kandil <i>et al</i> , 2015 ⁵⁴	USA	15770	Obesity	Complications	Unicompartmental knee arthroplasty	3 months	Poor
Kessler and Käfer, 2007 ⁵⁵	Germany	67	Obesity	Disability	Total hip replacement	10 days and 3 months	Good
Kim <i>et al</i> , 2011 ⁵⁶	South Korea	227	Obesity	Complications	Total knee arthroplasty	6 months	Poor
Kort <i>et al</i> , 2007 ⁵⁷	Netherlands	46	Obesity	Complications	Unicompartmental knee replacement	2 years	Poor
Ledford <i>et al</i> , 2014 ⁵⁸	USA	316	Obesity	Complications	Total hip and knee arthroplasty	2 months	Poor
Liabaud <i>et al</i> , 2013 ⁵⁹	USA	273	Obesity	Complications	Total knee arthroplasty	3 and 12 months	Poor
Liljensøe <i>et al</i> , 2013 ⁶⁰	Denmark	197	Obesity	Pain; disability	Total knee arthroplasty	4 years	Poor
Lübbeke <i>et al</i> , 2007 ⁶²	Switzerland	2495	Obesity	Complications; disability	Total hip arthroplasty	5 years	Good
Lübbeke <i>et al</i> , 2007 ⁶²	Switzerland	325	Obesity	Disability	Total hip arthroplasty	5 years	Good
Mackie <i>et al</i> , 2015 ⁶³	UK	1821	Obesity	Complications; pain; disability	Total knee arthroplasty	1 year	Poor
Madsen <i>et al</i> , 2014 ⁶⁴	USA	79	Obesity	Complications	Total knee arthroplasty	10 years	Poor
Maisongrosse <i>et al</i> , 2015 ⁶⁵	France	502	Obesity	Complications	Total hip arthroplasty	58 months	Poor
McLaughlin and Lee, 2006 ⁶⁶	USA	198	Obesity	Complications	Total hip replacement	14.5 years	Poor
Michalka <i>et al</i> , 2012 ⁶⁷	UK	191	Obesity	Complications; pain; disability	Hip arthroplasty	6 weeks	Poor
Murray <i>et al</i> , 2013 ⁶⁸	UK	2438	Obesity	Complications; disability	Unicompartmental knee replacement	1 year	Poor
Naal <i>et al</i> , 2009 ⁶⁹	Switzerland	83	Obesity	Pain; disability	Total knee arthroplasty	6 weeks 3, 12 and 24 months	Poor
Namba <i>et al</i> , 2005 ⁷⁰	USA	1813	Obesity	Complications	Total hip and knee arthroplasty	1 year	Poor
Napier <i>et al</i> , 2014 ⁷¹	UK	100	Obesity	Complications; disability	Total knee arthroplasty	3 and 12 months	Poor
Naylor <i>et al</i> , 2008 ⁷²	Australia	99	Obesity	Pain	Total hip and knee arthroplasty	2, 6, 12, 26 and 52 weeks	Good
						(Continued

Continued

Continued

Table 1

2008⁷⁴

Author, year

Ollivier et al, 201273

Patel and Albrizio,

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				0			
Predic	tor Outcomes	Surgery	Follow- up duration	Quality score			
Physic: activity	-	Total hip arthroplasty	10 years	Fair			
Obesity	y Complications	Total knee replacement	4 weeks, 6 weeks and 1 year	Good			
Physic: activity		Unicompartmental knee arthroplasty	4.2 years	Poor			
Physic: activity	•	Total hip and knee arthroplasty	1 year	Good			
Obesity	y Complications	Total hip and knee arthroplasty	1 year	Fair			
Obesity	y Complications; disability	Total knee arthroplasty	1 year	Fair			
Physic: activity		Total hip arthroplasty	5 years	Poor			
Obesity	y Pain; disability	Total hip arthroplasty	3 months	Good			
Obesity	y Disability	Total hip arthroplasty	6 months	Poor			
Obesity	y Disability	Total hip arthroplasty	3 months 1 and 2 years	Fair			
Obesity	y Complications	Total knee arthroplasty	5 months	Fair			
Obesity	y Complications; disability	Total hip arthroplasty	5 years	Poor			
sociatior	n between postsurgic	al pain and baseline c	besity				
sure	Results						
sure Results VS/KOOS After adjusting for age, gender, joint and presence of back pain, an increased BMI was associated with worst pain outcomes (P<0.02) at long term after THA or TKA.							
	Increased BMI co P=0.018) at short	rrelates significantly to term after TKA.	o VAS pain scale (r=	0.236;			
6	BMI was not asso long term after TK	ociated with SF-36 pa (A.	in scale (OR=0.96; F	⊃=0.1) at			
/IAC		as associated with less i4; P<0.001) at long te		OMAC			

Pietschmann et al, 2013 ⁷⁵	Germany	171	Physical activity	Disability	Unicompartmental knee arthroplasty	4.2 years	Poor
Poortinga et al, 2014 ⁷⁶	Netherlands	658	Physical activity	Disability	Total hip and knee arthroplasty	1 year	Good
Pulido <i>et al</i> , 2008 ⁷⁷	USA	9245	Obesity	Complications	Total hip and knee arthroplasty	1 year	Fair
Rajgopal <i>et al</i> , 2008 ⁷⁸	Canada	760	Obesity	Complications; disability	Total knee arthroplasty	1 year	Fair
Sechriest <i>et al</i> , 2007 ⁷⁹	USA	34	Physical activity	Disability	Total hip arthroplasty	5 years	Poor
Aranda Villalobos <i>et al</i> , 2013 ⁸⁰	Spain	63	Obesity	Pain; disability	Total hip arthroplasty	3 months	Good
Vogl <i>et al</i> , 2014 ⁸¹	Germany	281	Obesity	Disability	Total hip arthroplasty	6 months	Poor
Wang <i>et al</i> , 2010 ⁸²	USA	97	Obesity	Disability	Total hip arthroplasty	3 months 1 and 2 years	Fair
Yasunaga <i>et al</i> , 2009 ⁸³	Japan	3577	Obesity	Complications	Total knee arthroplasty	5 months	Fair
Zhang et al, 2012 ⁸⁴	China	714	Obesity	Complications; disability	Total hip arthroplasty	5 years	Poor

Table 2 Results of individual studies on the associat

Sample

210

527

size

Country

France

UK

Obesity versus pain			
Author, year	BMI: mean (SD)	Measure	Results
Knee			
Davis <i>et al</i> , 2011 ³⁸	NA	HOOS/KOOS	After adjusting for age, gender, joint and presence of back pain, an increased BMI was associated with worst pain outcomes (P<0.02) at long term after THA or TKA.
Järvenpää et al, 2010 ⁵¹	29.7 (NA)	VAS	Increased BMI correlates significantly to VAS pain scale (r=0.236; P=0.018) at short term after TKA.
Liljensøe <i>et al</i> , 2013 ⁶⁰	30 (NA)	SF-36	BMI was not associated with SF-36 pain scale (OR=0.96; P=0.1) at long term after TKA.
Mackie et al, 2015 ⁶³	NA	WOMAC	Increased BMI was associated with less improvement in WOMAC pain scale (t=-2.64; P<0.001) at long term after TKA.
Hip			
Dowsey <i>et al</i> , 2010 ⁴⁰	29.55 (5.64)*	Harris Hip Score	BMI was not associated with pain reduction (P=0.71) at long term after THA.
Heiberg <i>et al</i> , 2013 ⁴⁵	27 (6.27)*	HOOS	BMI was not associated with HOOS pain scale (P>0.05) at short term after THA.

*Calculated following guidelines from Cochrane Handbook for Systematic Reviews of Interventions.

BMI, body mass index; HOOS, Hip Dysfunction and Osteoarthritis Outcome Score; KOOS, Knee Injury and Osteoarthritis Outcome Score; NA, none available; SF-36, Short Form 36 Questionnaire; THA, total hip arthroplasty; TKA, total knee arthroplasty; VAS, visual analogue scale; WOMAC, Western Ontario and McMaster Universities Osteoarthritis Index.

Table 3 Results of individual studies on the association between postsurgical disability and baseline obesit
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Obesity versus disability

	BMI: mean (SD)	Measure	Results
Knee			
Davis <i>et al</i> , 2011 ³⁸	NA	WOMAC/KOOS	After adjusting for age, gender, joint and presence of back pain, an increased BMI was associated with worst outcomes (P<0.02) at long term after TKA or THA.
Dewan <i>et al</i> , 2009 ³⁹	31 (0.5)	Knee Society Score	BMI was not associated with worst knee function (P>0.119) at long term after TKA.
Hamoui <i>et al</i> , 2006 ⁴⁴	27.93 (7.1)*	Knee Society Score	No significant association between BMI and KSS (P>0.05) was found at long term after TKA.
Kort <i>et al</i> , 2007 ⁵⁷	NA	WOMAC	Obesity was not related to disability score (P>0.05) at long term after TKA.
Liljensøe <i>et al</i> , 2013 ⁶⁰	30 (NA)	Knee Society Score	Increased BMI was associated with worst knee scores (OR 0.95; 95% CI 0.9 to 1.0; P=0.04) at long term after TKA. Thes results did not change significantly after adjusting for age, sex, primary disease and surgical approach (OR 0.94; 95% C 0.90 to 0.99; P=0.02).
Mackie et al, 2015 ⁶³	NA	WOMAC	Increased BMI was associated with less improvement in disability scores (WOMAC t=-2.13; P=0.033) at long term after TKA.
Rajgopal <i>et al</i> , 2008 ⁷⁸	32.3 (6.58)*	WOMAC	The morbidly obese group (BMI≥40, n=69) does not present statistically significant difference in improvement in WOMAC score (P=0.669) when compared with other BMI groups at long term after TKA.
Hip			
Heiberg <i>et al</i> , 2013 ⁴⁵	27 (6.27)*	HHS	Increased BMI was associated with lower HHS (P<0.05) at short term after THA.
Jameson <i>et al</i> , 2014 ⁴⁸	NA	OHS	Increased BMI was not associated with changes in OHS (P>0.05) at short term after THA.
Lübbeke <i>et al</i> , 2007 ⁶²	26.4 (4.3)	HHS	Increased BMI was associated with lower hip score (r= -0.4 , 95% CI -0.8 to -0.1) at long term after THA.
McLaughlin and Lee, 2006 ⁶⁶	26 (NA)	HHS	The obese group (BMI≥30; n=95) did not present any statistically significant difference from the non-obese group (BMI<30, n=103) with regard to clinical outcomes assessed by HHS (P>0.05) at long term after THA.
Vogl et al, 2014 ⁸¹	26.9 (4.9)	WOMAC	Obesity was associated with changes in WOMAC score (P<0.05) at short term after THA.
	29.14 (6.23)	WOMAC	Increased BMI was not associated with WOMAC score (P=0.114) at long term after THA.

The course of pain and disability over time

The fractional polynomial regression model resulted in a pooled mean disability score and SD before hip arthroplasty of 59.42 (SD: 10.94; n=5250). At 12 months after surgery it had decreased to a mean of 31.31 (SD: 24.28; n=3017) and a further reduction was observed at 120 months, when the mean disability score after hip

OA, a pooled mean disability score of 56.88 (SD: 10.74; n=17225) was observed for patients undergoing arthroplasty. At 12 months after surgery this value decreased to 21.80 (SD: 13.51; n=2898), while at the 110-month follow-up, the mean disability score was 14.18 (SD: 0.77; n=485). The pooled mean pain score before hip arthroplasty was 54.86 (SD: 10.20; n=2517), decreasing to 13.76 (SD: 1.32; n=1058) 3 months after surgery, 10.8 (SD: 1.69; n=1212) at 6 months and slightly increasing to 13.45 (SD:

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Table 4 Results of individual studies investigating the association between obesity and postsurgical complie	cations
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Obesity versus comp	lications		
Author, year	BMI: mean (SD)	Measure	Results
Ollivier et al, 2012 ⁷³	25.13 (3.14)*	HHS/HOOS	
Pietschmann <i>et al</i> , 2013 ⁷⁵	28.4 (4.62)*	OKS	At long term, physical activities were not related to complications (P<0.01). Physically active patients had less pain and better OKS scores after UKA.
Poortinga <i>et al</i> , 2014 ⁷⁶	28.7 (4.9)	WOMAC	At long term, physical activity was not associated with WOMAC score (P>0.05) after THA or TKA.
Sechriest <i>et al</i> , 2007 ⁷⁹	28.1 (8.3)	UCLA	At long term increased BMI was not correlated to UCLA physical activity score (r=–0.07; P=0.67) after TKA.

*Calculated following guidelines from Cochrane Handbook for Systematic Reviews of Interventions.

BMI, body mass index; HHS, Harris Hip Score; HOOS, Hip disability and Osteoarthritis Outcome Score; OKS, Oxford Knee Score; r, coefficient of association; THA, total hip arthroplasty; TKA, total knee arthroplasty; UCLA, University of California, Los Angeles Activity Questionnaire; UKA, unicompartmental knee arthroplasty; WOMAC, Western Ontario and McMaster Universities Osteoarthritis Index.

7.87; n=2173) at the 12-month follow-up. For patients undergoing knee arthroplasty, the pooled pain score at baseline was 57.78 (SD: 9.28; n=2211), which decreased to 25.67 (SD: 6.61; n=1222) at 6 months, and 14.18 (SD: 0.77; n=1820) at the 12-month follow-up (figure 2).

Association between obesity and postsurgical pain outcomes

Fourteen studies investigated the association between obesity and pain intensity in a total of 5687 patients after hip or knee arthroplasty. Seven of the 14 studies presented enough data to be pooled in a meta-analysis. There was an overall moderate and statistically significant difference in postsurgical pain between obese and non-obese patients after arthroplasty, with non-obese patients having better outcomes at short-term timepoint (SMD -0.44; 95% CI -0.68 to -0.20; P<0.001) and long-term timepoint (SMD -0.36; 95% CI -0.47 to -0.25; P<0.001). The pooled results for separate joints suggest non-obese participants have significantly less short-term (ie, less than 6 months) postsurgical knee pain, compared with obese participants (SMD -0.55; 95% CI -0.90 to -0.20; P=0.002) and

(P>0.05) after THA or TKA. g term increased BMI was **not** correlated to UCLA physical y score (r=-0.07; P=0.67) after TKA. Reviews of Interventions. Osteoarthritis Outcome Score; OKS, Oxford Knee Score; r, iroplasty; UCLA, University of California, Los Angeles Activity iern Ontario and McMaster Universities Osteoarthritis Index. postsurgical hip pain (SMD -0.34; 95% CI -0.67 to -0.02; P=0.039). Obesity was defined as having a BMI over 30 kg/m^2 . At long term (ie, 6 months or longer), there was a significant moderate difference between obese and non-obese groups in terms of knee pain (SMD -0.36; 95% CI -0.48 to -0.25; P<0.001); however, there was no difference between groups for hip pain (SMD -0.32; 95% CI -0.84 to 0.20; P=0.222) (figure 3). The results of individual studies not included in the pooled analyses are presented in table 2. Association between obesity and postsurgical disability outcomes The impact of obesity on disability was investigated

Association between obesity and postsurgical disability outcomes The impact of obesity on disability was investigated by 32 studies which compared postsurgery disability scores in 35286 obese and non-obese participants. Of these, 19 studies presented complete data that were included in the pooled analysis. At short term, no statistically significant difference in overall disability between obese and non-obese participants was observed (SMD –0.16; 95% CI –0.42 to 0.10; P=0.231). Likewise, no statistically significant difference was observed between obese activity and pain or disability sults ong term, high-impact sports were associated with better S (P<0.001) and HOOS (P<0.05) after THA. ong term, physical activities were **not** related to pplications. Physically active patients had less pain and better S, KSS and WOMAC scores (P<0.05) after UKA.

 Table 5
 Individual results on the association between physical activity and pain or disability

Physical activity versus disability						
Author, year	BMI: mean (SD)	Measure	Results			
Ollivier <i>et al</i> , 2012 ⁷³	25.13 (3.14)*	HHS/HOOS	At long term, high-impact sports were associated with better HHS (P<0.001) and HOOS (P<0.05) after THA.			
Pietschmann <i>et al</i> , 2013 ⁷⁵	28.4 (4.62)*	OKS/KSS/WOMAC	At long term, physical activities were not related to complications. Physically active patients had less pain and better OKS, KSS and WOMAC scores (P<0.05) after UKA.			
Poortinga <i>et al</i> , 2014 ⁷⁶	28.7 (4.9)	WOMAC	At long term, physical activity was not associated with WOMAC score (P>0.05) after THA or TKA.			
Sechriest <i>et al</i> , 2007 ⁷⁹	28.1 (8.3)	UCLA	At long term increased BMI was not correlated to UCLA physical activity score (r =-0.07; P=0.67) after TKA.			

*Calculated following guidelines from Cochrane Handbook for Systematic Reviews of Interventions.

BMI, body mass index; HHS, Harris Hip Score; HOOS, Hip disability and Osteoarthritis Outcome Score; KSS, Knee Society Score; OKS, Oxford Knee Score; r, coefficient of association; THA, total hip arthroplasty; TKA, total knee arthroplasty; UCLA, University of California, Los Angeles Activity Questionnaire; UKA, unicompartmental knee arthroplasty; WOMAC, Western Ontario and McMaster Universities Osteoarthritis Index.

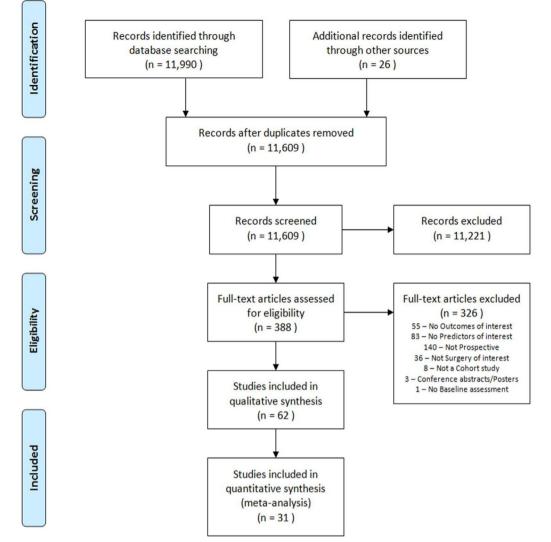


Figure 1 Flow chart of search strategy and screening steps. Detailed steps of references screening process of results from database searches.

and non-obese participants for postsurgical knee or hip disability (SMD -0.42; 95% CI -1.0 to 0.16; P=0.159 and SMD -0.09; 95% CI -0.39 to 0.20; P=0.527, respectively).

At long-term follow-up, however, there was an overall moderate and statistically significant difference in postsurgical disability between obese and non-obese patients regardless of the joint affected (SMD -0.32; 95% CI -0.37to -0.28; P<0.001). That difference was still statistically significant and of moderate magnitude when knee and hip joints were analysed separately (SMD -0.32; 95% CI -0.37to -0.27; P<0.001 and SMD -0.35; 95% CI -0.44to -0.26; P<0.001, respectively, and favouring non-obese patients) (figure 4). The results of individual studies not included in the pooled analyses are presented in table 3.

Association between obesity and postsurgical complications

The association between obesity and complications after joint arthroplasty was assessed by 40 studies including a total of 245 433 patients who underwent knee or hip arthroplasty. Of these, 17 presented enough data and were included in the meta-analyses. The pooled results suggest that at short-term follow-up, non-obese participants are less likely to have postsurgical deep vein thrombosis (DVT) (OR 0.49; 95% CI 0.26 to 0.91; P=0.024) when compared with obese participants (figure 5). A total of 13 studies were pooled (n=22782) showing non-obese patients are also less likely to present any long-term (ie, ≥ 6 months) dislocation (OR 0.50; 95% CI 0.31 to 0.80; P=0.003) and DVT (OR 0.58; 95% CI 0.36 to 0.94; P=0.043). A non-significant difference between groups was observed between non-obese and obese participants for long-term revision surgery (OR 0.66; 95% CI 0.34 to 1.28; P=0.217) (figure 5).

The pooled analysis on short-term postsurgical infection for hip replacement showed that non-obese patients are less likely to develop infections compared with obese participants (OR 0.33; 95% CI 0.19 to 0.59; P<0.001) (figure 6). For knee replacement, separate analyses were conducted for studies comparing obese with non-obese participants and those comparing morbidly obese with non-obese participants (figure 7). The results

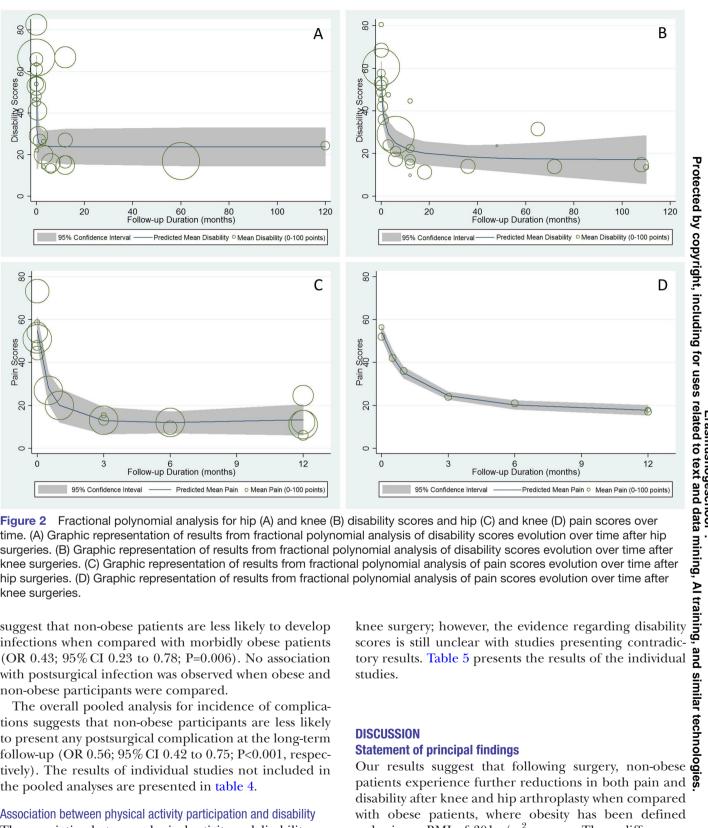


Figure 2 Fractional polynomial analysis for hip (A) and knee (B) disability scores and hip (C) and knee (D) pain scores over time. (A) Graphic representation of results from fractional polynomial analysis of disability scores evolution over time after hip surgeries. (B) Graphic representation of results from fractional polynomial analysis of disability scores evolution over time after knee surgeries. (C) Graphic representation of results from fractional polynomial analysis of pain scores evolution over time after hip surgeries. (D) Graphic representation of results from fractional polynomial analysis of pain scores evolution over time after knee surgeries.

suggest that non-obese patients are less likely to develop infections when compared with morbidly obese patients (OR 0.43; 95% CI 0.23 to 0.78; P=0.006). No association with postsurgical infection was observed when obese and non-obese participants were compared.

The overall pooled analysis for incidence of complications suggests that non-obese participants are less likely to present any postsurgical complication at the long-term follow-up (OR 0.56; 95% CI 0.42 to 0.75; P<0.001, respectively). The results of individual studies not included in the pooled analyses are presented in table 4.

Association between physical activity participation and disability

The association between physical activity and disability was investigated by four studies^{73 75 76 79} or 1033 participants undergoing hip or knee arthroplasty. Included studies have not provided enough data to be pooled. The overall results from these four papers suggest that participants who practise more physical activity before the surgeries were more likely to experience less pain after either hip or

disability after knee and hip arthroplasty when compared with obese patients, where obesity has been defined as having a BMI of 30 kg/m^2 or over. These differences seemed to be more accentuated for knee pain outcomes following arthroplasty than for hip pain or disability outcomes. Non-obese participants also experienced significantly less postsurgical complications, including dislocation, DVT and infection especially following hip arthroplasty. Our analyses also demonstrate that obesity

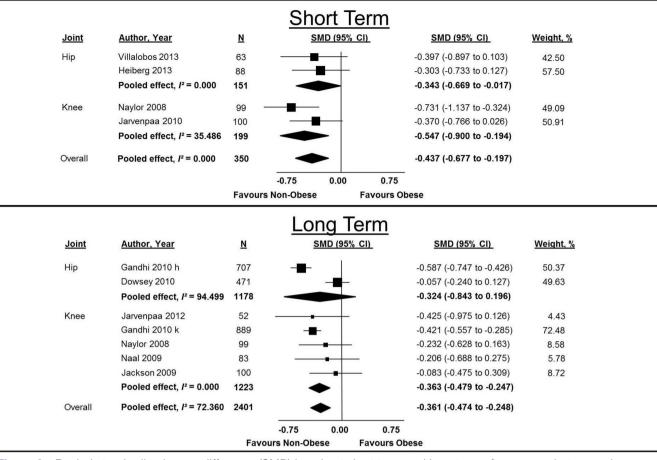


Figure 3 Pooled standardised mean difference (SMD) in pain at short term and long term after surgery between obese and non-obese patients. Results from meta-analysis of included studies presented as SMD of pain scores at short-term (<6 months) and long-term (≥ 6 months) follow-ups between non-obese and obese groups.

is a reliable predictor of complications after THA and TKA, in the short term after the procedure and at longer follow-ups. The evidence regarding preoperative physical activity remains unclear due to conflicting results of included studies, especially in terms of postoperative disability. The four included cohort studies, however, suggest that physical activity participation is associated with better pain outcomes following surgery.

Our results from the fractional polynomial analysis have also shown that all patients experienced an improvement in pain and disability after surgery. We also highlight that although non-obese patients experience further improvements in pain and disability compared with obese participants, both groups improved significantly following surgery as depicted in figure 2. The observed decrease in pain from baseline was approximately 70% at 6 months and 75% at 12 months, with decreases in disability of 55%at 12 months and 67% at 120 months. The interpretation of the postsurgical course of pain and disability, however, needs to be taken in the context of the inclusion criteria we have used in our review, given we have only included data from cohort studies that have assessed the role of obesity or physical activity participation on surgical outcomes.

Strengths and weaknesses in relation to other studies, discussing particularly any differences in results

Protected by copyright, including for uses related to text and data mining, A Our meta-analysis results regarding the association between obesity and postsurgical complications found that obese patients present higher complication rates than non-obese patients. These results are consistent with the findings of previous systematic reviews of Hofstede et al,¹⁴ Samson et al¹⁵ and Liu et al.¹⁶ Our meta-analysis results regarding the association between obesity and postsurgery disability also agreed with the findings of Buirs *et al*¹³ and Samson *et al*,¹⁵ which found that obesity (defined as having BMI over 30 kg/ m²) was associated with worst postsurgical functional score. The only previous review which has performed a meta-analysis on the association between obesity and postarthroplasty pain or disability limited its inclusion criteria to hip joint.¹⁶ That review included a total of 15 studies in their meta-analysis and found that obesity increases the risk of postsurgical complications (RR 1.68; 95% CI 1.23 to 2.30; P=0.0004) and is associated with worse disability scores following surgery (MD -2.75; 95% CI -4.77 to -0.6; P=0.07). Our study has included 33 cohorts of hip arthroplasty participants in the qualitative analysis, 16 in the

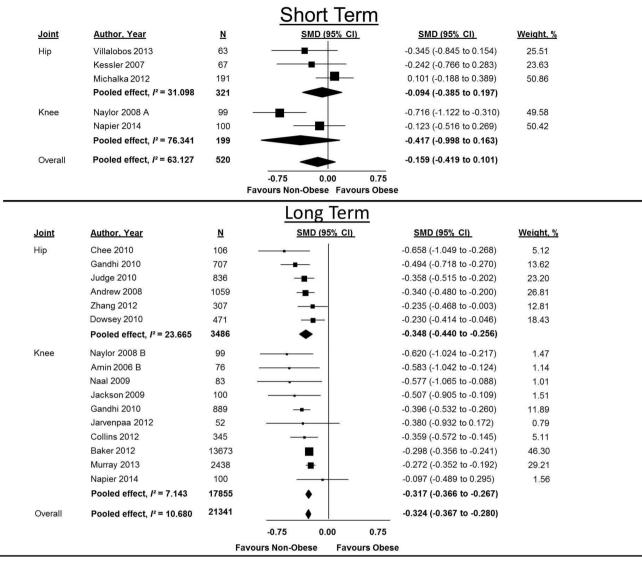


Figure 4 Pooled standardised mean difference (SMD) in disability at short term and long term after surgery between obese and non-obese patients. Results from meta-analysis of included studies presented as SMD of disability scores at short-term (<6 months) and long-term (≥6 months) follow-ups between non-obese and obese groups.

meta-analyses, and confirms past findings that obesity is associated with worse outcomes in terms of disability and complications, and pain at both short-term and long-term periods following surgery. Hofstede *et al*¹⁴ have also conducted a systematic review of the literature on preoperative predictors of surgical outcomes after hip replacement in patients with OA. Although those authors included 35 studies, only five studies investigated the effect of obesity on postsurgical pain, disability and quality of life.¹⁴ No meta-analysis was performed.

Implications for clinicians or policymakers

Our results have a direct impact on clinical practice as the results demonstrate that obese patients have a higher risk of complications and a poorer prognosis in terms of pain and disability postoperatively when compared with non-obese patients. These results also allude to the

Protected by copyright, including for uses related to text and data mining, AI training, and importance of identifying and implementing effective presurgical rehabilitation and weight loss approaches to <u>0</u> optimise postsurgical outcomes and minimise harm to the patient. The importance of weight loss has been highlighted in international clinical guidelines on non-surgical management of knee OA, for instance, given the pain and disability reductions observed following weight loss regimes.⁸⁶ Past research also suggests there is a doseresponse relationship between weight loss and clinical outcome improvement. A recent completer-type analysis of 1383 participants with knee OA showed that a weight loss of 7.7% of body weight or more is associated with clinically important changes in pain and disability, as measured using the Knee Injury and Osteoarthritis Outcome Score (KOOS).⁸⁷ This evidence reinforces the importance of presurgical weight loss programmes and strategies in order to optimise postsurgical recovery.

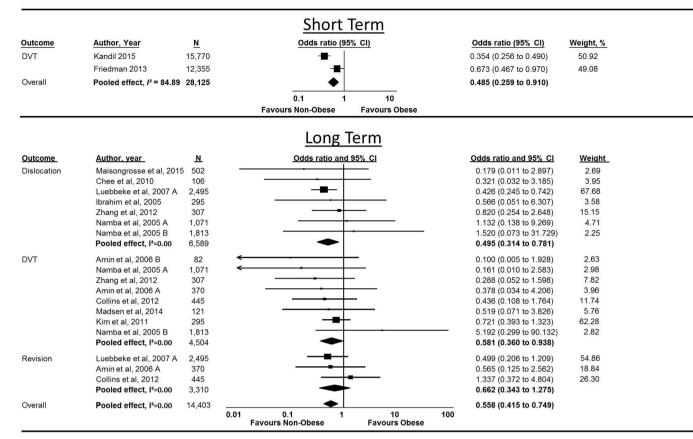


Figure 5 Pooled association between complications and obesity at short-term and long-term follow-ups. Results from metaanalysis of included studies presented as incidence of complications at short-term (<6 months) and long-term (≥6 months) follow-ups between non-obese and obese groups. DVT, deep vein thrombosis.

Strengths and weaknesses of the study

The current review has included 62 cohort studies and a total of 256481 participants and is the most comprehensive systematic review on the topic to date. It is also the first review to use a quantitative approach to synthesise the results of pain, disability and surgical complications between non-obese and obese participants and consider the physical activity level of participants who underwent hip or knee arthroplasty due to OA. Our review has some limitations. The methodological quality of the included studies was in general poor. The most common methodological flaw among included cohorts was not controlling

data mini for confounding factors age, sex or BMI (32 studies, 51%) followed by not using a representative sample (n=30 studies, 48%). Moreover, we have observed great variability of follow-up duration across studies, ranging from 2 weeks to 11 years. We have used a cut-off of 6 . ح months to define short-term follow-up (ie, <6 months) or long-term follow-up (ie, ≥ 6 months), but acknowledge that within each follow-up category there was substantial variation in the duration of follow-up across studies.

b Between-study heterogeneity has also been observed in some of the pooled analysis for obesity presented in this review. A potential source of between-study heterogeneity

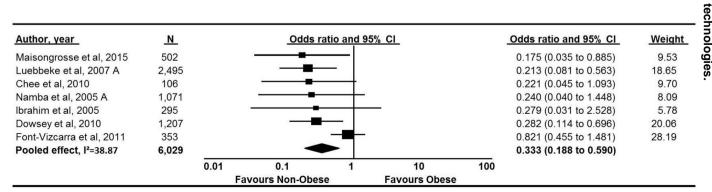


Figure 6 Pooled association between postsurgical infections and obesity for hip surgery. Results from meta-analysis of included studies presented as incidence of infections after hip surgery between non-obese and obese groups.

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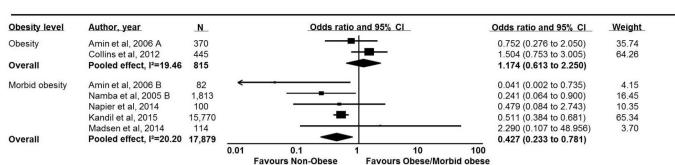


Figure 7 Pooled association of postsurgical infections for knee surgery. Results from meta-analysis of included studies presented as incidence of infections after hip surgery comparing the non-obese group with obese group and the non-obese group with morbidly obese group.

includes the variability in the definition of obesity categories across studies. Although obesity was assessed using BMI scores in all studies, some studies have used only two obesity groups (ie, obese or non-obese) while others used several categories including underweight, normal or overweight, obese and morbidly obese. These needed to be combined for some of our pooled analyses.

Another potential source of between-study heterogeneity across is the difference in surgical procedures used across studies. For instance, in the pooled analysis of risk of postsurgical DVT and obesity, while Kandil *et* $al^{\tilde{p}^4}$ performed unicompartmental knee arthroplasties, Friedman *et al*¹² performed total arthroplasties on both hip and knee joints. That discrepancy might explain the different results reported by these two studies (figure 5). Likewise, the mean physical activity load reported by the included studies varied substantially, ranging from low to high frequency of participation in low and high-impact activities. This should be taken into consideration when interpreting the physical activity results.

CONCLUSION

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Our results have shown that obese patients undergoing hip or knee arthroplasty due to OA have worse outcomes in terms of pain and complications when compared with non-obese patients, with differences more accentuated for patients with knee OA. Likewise, obese patients will have worse surgical outcomes in terms of disability, but only at long-term follow-ups. It is still unclear whether presurgical physical activity participation has an impact on surgical outcomes. However, we acknowledge that the health benefits of physical activity participation for patients with knee and hip OA are multiple and reach beyond those considered in this review.

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Contributors DP, GCM, PHF, FMB and MLF were involved in the conception and design of the review. DP, GCM and MLF developed the search strategy and performed study selection. DP and GCM extracted data from included studies. DP assessed the methodological quality of included studies. DP and MLF were involved in the data analysis. DP, GCM, PHF, FMB and MLF were involved in the interpretation and discussion of results. DP drafted the manuscript, and GCM, PHF, FMB and MLF contributed to the drafting of the review. GCM, PHF, FMB and MLF revised it critically for important intellectual content. All authors approved the final version of the article. All authors had access to all of the data in the study and can take responsibility for the integrity of the data and the accuracy of the data analysis. Theauthors would like to acknowledge the participation of Ms Giovana Visentini inthe independent methodological quality assessment of the included studies.

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Competing interests None declared.

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Provenance and peer review Not commissioned; externally peer reviewed.

Data sharing statement All data extracted from papers and used to write this paper are available to whoever ask. Contact the corresponding author for further information.

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