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Custom triflange acetabular components for large acetabular defect reconstruction in revision total hip arthroplasty: a systematic review and meta-analysis on 1218 patients

HIP

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Purpose: This is a systematic review and meta(regression) analysis to assess the performance
of custom triflange acetabular components (CTAC) in total hip arthroplasty (THA) revision
surgery. Implant-related complications, failure rate, functional outcomes and implant and
surgical technique-related predictors for outcome were assessed.

- Methods: This systematic review was performed according to PRISMA guidelines and registered with PROSPERO (2020 CRD42020209700). PubMed, Embase, Web of Science, COCHRANE Library and Emcare were searched. Studies on Paprosky type 3A and 3B or AAOS type 3 and 4 acetabular defects with a minimum follow-up of 12 months and cohorts > 10 patients were included.
- *Results:* Thirty-three studies were eligible for inclusion (*n* = 1235 hips, 1218 patients). The methodological quality of the studies was moderate (AQUILA: 7.4/11 points). Considerable heterogeneity was observed in terms of complications, re-operations and implant failure reporting. The total incidence of implant-related complications was 24%. The incidence of re-operation for any reason was 15%, and the implant failure rate was 12% at a mean of 46.9 months and the post-operative Harris Hip Score improved by a mean of 40 points. Several predictors for outcome were found, such as implant generation, follow-up length and study start date.
- *Conclusions:* The use of CTAC in revision THA has satisfactory complication and implant failure rates. The CTAC technique improves post-operative clinical outcomes and the meta-regression analysis showed that there is a clear association between improvements in the CTAC performance and the evolvement of this technique over time.

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### **Keywords**

- custom triflange acetabular component
- total hip arthroplasty revision
- large acetabular defects

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## Introduction

The management of large acetabular defects in total hip arthroplasty (THA) revision surgery is challenging. Large defects with and even without pelvic discontinuity are notoriously difficult to reconstruct. Paprosky type 3A and 3B and AAOS type 3 and 4 defects demonstrate severe bone loss of the acetabular rim and supporting structures, such as the medial wall and the anterior/posterior column with possible pelvic discontinuity (1, 2, 3). Over the past decades, multiple reconstruction techniques have been proposed, including structural allografts, bone impaction grafting, jumbo cups, anti-protrusion cages, trabecular metal augments, cup-cage combinations, oblong cups and custom-made cages (4, 5, 7, 6, 7, 8, 9, 10, 11). Nevertheless, none of these surgical techniques has been proven to be superior. For that matter, custom triflange acetabular components (CTAC) aimed at reconstructing large defects with a mono-block, patient-specific implant which fits into the defect and can also address stabilisation of pelvic discontinuity, have also been proposed. These CTAC implants have evolved over the past years with improvements in rapid prototyping and three-dimensional (3D) printing techniques (12). Improvements have also been made in surgical pre-planning including 3D visualisation of the defects, interaction between surgeon and manufacturer on the design and intra-operative (navigation) tools, as well as the implant characteristics



itself. Thus, these developments may have an effect on the outcome of this technique. Several studies have reported on the use of CTAC for hip revision arthroplasty, reporting on small numbers, with large heterogeneity in patient characteristics, outcome measures, complication reporting and end-point definition (e.g. implant survival, failure). This heterogeneity makes valid comparisons between data unreliable (13, 14).

Therefore, our primary aim was to perform a systematic review, meta-analysis and meta-regression analysis on the outcome of CTAC for pelvic reconstructions in THA revision surgery for Paprosky type 3A and 3B and AAOS type 3 and 4 acetabular defects. The main objectives were to assess CTAC performance by means of identifying implant-related complications, implant failure and functional outcomes. The secondary aim was to evaluate the predictors of outcome for CTAC surgical techniques and implant characteristics.

## **Methods**

The reporting of this systematic review is in accordance with the PRISMA 2020 guidelines (15) and was registered under PROSPERO number 2020 CRD42020209700. All studies on THA revision surgery in patients older than 18 years using a CTAC to reconstruct Paprosky type 3A and 3B and AAOS type 3 and 4 acetabular defects, with or without revision of the femoral component, were included. For analysis, inclusion was limited to studies with a minimum follow-up of 12 months and reporting on implant survival or complication rates. Exclusion criteria were as follows: studies on non-custom acetabular implants, reviews or expert opinion articles and case series including less than ten patients. Articless in a language other than English, such as Dutch or German were not considered.

We adopted a search strategy up to 28th of January 2022 using the following bibliographic databases: PubMed, Embase (OVID version), Web of Science, Cochrane Library and Emcare (OVID version). Meeting abstracts were searched in Embase, Web of Science and Cochrane Library. The full search strategy (composed by DB, ZV and JS) for all databases can be found in Appendix 1 (see section on supplementary materials given at the end of this article). Additionally, reference lists of papers included in the systematic review and systematic review reports on similar topics were examined.

Two independent reviewers (ZV and DB) reviewed all titles and abstracts of the retrieved articles. Next, full texts of the selected papers were screened by two independent reviewers (RT and DB) for inclusion, applying the predefined in- and exclusion criteria. Disagreement on inclusion was resolved by discussion, and if no consensus was reached, a third independent reviewer was consulted (RN) until consensus was reached.

Two reviewers (RT and DB) individually extracted data from the selected studies into two separate electronic databases (IBM SPSS Statistics, Version 23.0., IBM Corp). Next, both databases were compared and inconsistencies were identified and resolved by re-evaluating the specific studies until consensus was reached. Variables/study-level factors collected in the database included: study title, year of publication, author, study design, number of patients and hips included, study period, patient demographics (age, gender, BMI), follow-up in months, amount of previous surgical revisions on the affected hip, acetabular defect type (Paprosky/AAOS), clinical data (surgical time, time to full weight bare), implant-related complications, re-operations (for any reason), implant failure (defined as partial or full surgical revision of the CTAC or associated THA components; that is, open reduction without component revision for dislocation will not classify as implant failure), functional outcome scores, surgical approach, implant manufacturer, liner type and fixation method into the CTAC, bone interface type (hydroxyapatite, porous, combinations), screw type (locking/non-locking) and CTAC generation. CTAC generation was defined as either old or new. New generation was defined as 3D titanium printed implants, developed with rapid prototyping and/or the availability of patient-specific drill guides and digital or 3D printed bone models to aid the surgical implantation. Old-generation implant was defined as older CTAC manufacturing techniques such as machining or casting without the use of rapid prototyping, also lacking patient-specific drill guides and digital or 3D printed bone models to aid the surgical procedure.

### Quality assessment

The methodological quality of all included studies was assessed using the Assessment of Quality In Lower Limb Arthroplasty (AQUILA) checklist, a tool specifically designed to appraise the quality of observational studies concerning total hip and knee arthroplasty (16, 17). Two authors (RT and DB) independently assessed the quality of all included studies using predefined data extraction sheets. Inconsistencies between the two authors were resolved by consensus.

### Statistical analysis

The data were combined for the meta-analysis and a random effects model was applied using the metafor package for R version 3.5 (18). Heterogeneity between studies was tested using the  $l^2$  statistic, which describes the variation across studies as a result of heterogeneity. Possible sources of heterogeneity were explored using meta-regression: the random-effects regression model, which has previously been used to study the effectiveness of the BCG vaccine against tuberculosis, was employed

(19). The random-effects regression model identifies modifying variables that affect the outcome of interest between studies, therefore helping to resolve contradictory outcomes of different studies. To assess publication bias, a funnel plot for studies reporting the primary outcome was constructed. In the case of asymmetry in the funnel plot, a trim-and-fill method and cumulative meta-analysis were used to explore the magnitude and direction of the publication bias.

### Results

### Literature review

The flow chart of study selection is shown in Fig. 1. The search strategy resulted in a total of 1428 references. One additional study was identified through screening of study reference lists. After removing duplicates, a total of 623 studies remained. Screening of title and abstract identified 140 papers eligible for inclusion. Three meeting abstracts that matched the inclusion criteria were not published and therefore could not be reviewed/included, leaving 137 papers for full-text assessment. In total, 104 studies were excluded for the following reasons: 36 papers were review papers or expert opinions; 27 papers concerned other techniques or the cohort existed of CTAC and standard implants without sufficient individual patient data to extract the CTAC patients; 27 papers included a cohort of less than 10 patients and 13 papers did not meet the language requirements. One paper was excluded after verification with the author that the cohort



Figure 1 Flow chart for study selection.

was included in a more recent published study by the same group (20). A total of 33 papers remained for final analysis, including a total of 1218 patients and 1235 hips. Details of the included studies are summarized in Table 1.

### Quality assessment

The mean AQUILA methodological quality score was 7.4 points, ranging from 4 to 10 points (out of a maximum of 11 points). The majority of methodological flaws were related to the follow-up period (AQULA item 4): in 8 of 33 studies the follow-up was predefined; in 22 of 33 studies the follow-up was performed when patients had complaints or a chart review (of non-predefined follow-up) and in 3 of 33 studies it was unclear how the follow-up was performed. Methodological quality related to follow-up (AQUILA item 4) was not an effect modifier. Table 2 shows the methodological scores for each item.

### Surgical details, implant characteristics and CTAC generation

Sixteen of 33 studies reported the number of prior THA revisions before CTAC usage. The mean prior revision amount was 2.8 (s.p. 1.3). The surgical approach was described in the majority of the selected studies (30 of 33). The posterolateral approach was used in the majority of patients (22 studies, 720 patients, 731 hips). Others described a mix of approaches used (six studies). The direct lateral and triradiate-transtrochantaric approaches were used in one study each. Different fixation techniques for the THA liner into the CTAC were described. The liner was cemented into the CTAC in 390 hips (15 studies, 386 patients) and the liner attached/combined into the CTAC in an uncemented fashion in 468 hips (11 studies, 461 patients). The type of liner fixation into the CTAC was not reported in five studies. The majority of studies (17 studies, 772 patients, 782 hips) reported a mix of standard, bipolar or constrained liners used in their cohorts. Constrained liners were mainly applied if the revision of the liner in the CTAC was needed due to THA instability. Six studies (119 patients, 120 hips) reported the exclusive use of standard liners. Only four studies reported the sole use of bipolar liners (61 patients, 62 hips) and six studies (266 patients, 271 hips) did not report liner type. In most cases, non-locking screws were used for CTAC primary fixation (18 studies, 514 patients, 519 hips), whereas a combination of locking and non-locking screws were used in 381 hips (6 studies, 374 patients). Screw type was unknown in nine studies (330 patients, 335 hips). The CTAC had porous bone interfaces in 383 hips (13 studies, 382 patients); hydroxyapatite coatings in 194 hips (3 studies, 189 patients) and a combination of porous surfaces and hydroxyapatite coating in 347 hips (8 studies, 340 patients). Nine studies did not report bone interface type (307 patients, 311 hips). In 448 hips (16

Study	Publication year	Study design	Patients (n)	<b>Hips</b> ( <i>n</i> )	Mean age (years)*	Male/female	<b>BMI</b> (kg/m²)*	Follow-up (months)*	Year study start	CTAC generation
Christie et al. (12)	2001	RCS	76	78	59 (29–87)	20/56	NA	53 (24–107)	1992	old
Joshi et al. (21)	2002	RCS	27	27	68 (55–77)	9/18	NA	40 (25–56)	1993	old
Dennis (22)	2003	RCS	24	24	68 (44–82)	7/17	NA	48 (24–78)	NA	old
Holt & Dennis (23)	2004	RCS	26	26	69 (44–82)	8/18	NA	54 (24–85)	NA	old
DeBoer et al. (24)	2007	RCS	18	20	56 (30–77)	3/15	NA	123 (89–157)	1992	DIO
Taunton et al. (25)	2012	RCS	57	57	61 (35–81)	6/51	27.0 (21–40)	76 (24–215)	1992	DIO
Wind <i>et al.</i> (26)	2013	RCS	19	19	58 (42–79)	7/12	NA	31 (16–59)	2001	old
Friedrich et al. (27)	2014	RCS	18	18	68 (26–79)	7/11	NA	30 (17–62)	2007	New
Berasi et al. (28)	2015	RCS	23	24	67 (47–85)	7/17	28.0 (23–39)	57 (28–108)	2003	Old
Mao <i>et al.</i> (29)	2015	RCS	22	23	61 (38–80)	NA	NA	82 (NA)	2001	New
Barlow et al. (30)	2016	RCS	63	63	63 (NA)	NA	27.0 (NA)	52 (NA)	NA	New
Li et al. (31)	2016	RCS	24	24	65 (54–79)	8/16	NA	67 (24–120)	2003	New
Baauw et al. (32)	2017	RCS	12	12	66 (33–79)	3/9	NA	25 (18–39)	2011	New
Myncke <i>et al.</i> ( <b>33</b> )	2017	RCS	21	21	67 (50–83)	6/15	NA	25 (NA)	2009	New
Berend et al. (34)	2018	RCS	94	95	66 (38–85)	34/61	29.0 (18–51)	43 (4–128)	2004	DIO
Gladnick et al. (35)	2018	RCS	73	73	59 (32–83)	21/52	29.6 (30–48)	90 (60–144)	2000	DIO
Gorianov et al. (20)	2018	RCS	11	11	72 (48–82)	5/6	NA	20 (6–37)	NA	New
Kieser <i>et al.</i> ( <b>36</b> )	2018	RCS	20	20	67 (50–89)	8/12	NA	38 (34–110)	2007	New
Moore et al. (37)	2018	RCS	35	35	60 (38–80)	21/14	33.0 (23–42)	Min. 120 (NA)	2001	Old
Weber et al. (38)	2018	RCS	11	11	76 (60–85)	3/8	27.6 (24–40)	21 (6–36)	2013	New
Jones et al. (39)	2019	RCS	91	96	62 (31–85)	22/74	27.0 (NA)	25 (0–116)	2004	Old
Burastero et al. (40)	2020	RCS	19	20	(0 (NA)	11/8	25.1 (NA)	42 (NA)	2014	New
Durand-Hill et al. (41)	2020	RCS	20	20	(NA) 99	7/13	28.0 (NA)	26 (12–40)	2016	New
Fröschen et al. (42)	2020	RCS	68	68	70 (27–89)	ΝA	27.3 (19–39)	43 (2–120)	2008	Unknown
Matar <i>et al.</i> (43)	2020	RCS	17	17	72 (61–83)	3/14	NA	43 (24–84)	2013	Combination
Walter et al. (44)	2020	RCS	54	58	70 (NA)	14/44	27.3 (NA)	56 (24–120)	2008	Unknown
Zampelis & Flivik (45)	2020	PCS	10	10	64 (36–87)	5/5	26.0 (19–34)	12 (NA)	2014	New
Fröschen et al. (46)	2021	RCS	70	70	70 (27–89)	19/51	27.4 (19–27)	42 (1–120)	2008	Combination
Goriainov et al. (47)	2021	RCS	19	19	77 (52–91)	8/11	NA	53 (17–88)	2014	New
Kawalkar <i>et al.</i> (48)	2021	RCS	13	13	69 (57–86)	5/8	27 (NA)	50 (18–70)	2011	New
Scharff-Baauw et al. (49)	2021	PCS	49	49	68 (41–89)	8/41	27.0 (19–44)	24 (NA)	2013	New
Sershon et al. (50)	2021	RCS	50	50	60 (33–83)	16/34	28.4 (17–48)	NA	2000	Unknown
von Hertzberg-Boelch et al. (51)	2021	RCS	14	14	70 (55–83)	5/9	28 (25–31)	35 (14–94)	2010	New
Tikhilov et al. (52)	2022	RC	61	61	57 (NA)	9/52	28.1 (15–44)	37 (12–59)	2013	New
Total			1218	1235	65.4	29/71%	27.8			
*Values in parentheses are range.	rocoactiva casa sarias	· DC retrochect	ive cohort. BC	C retrocher	tivo caso series					
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 Table 1
 Details of included studies.

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Table 2	Overview	of AQUILA	study	methodological	scores
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AQUILA methodological quality item	Number of studies
<ol> <li>Is there a clear primary research question/hypothesis?</li> <li>How were the cohorts constructed?</li> </ol>	Yes: 25 of 33
A: Consecutively	21 of 33
B: Non-consecutively	5 of 33
C: Unknown	7 of 33
3. How adequate was the follow-up?	
A: Fully completed FU	21 of 33
B: $\leq$ 5% lost-to-FU or FU quotient is $\leq$ 1	6 of 33
C: >5% lost-to-FU or FU quotient is >1	6 of 33
D: Unknown	0 of 33
4. How was the follow-up performed?	
A: Predefined (e.g. yearly)	8 of 33
B: When the patient had complaints or chart review (of non-predefined FU)	22 of 33
C: Unknown	3 of 33
5. How many arthroplasties are at risk at the FU of interest?	
A: ≥20	20 of 33
B: <20	13 of 33
C: Unknown	0 of 33
6. Has a worst-case analysis or competing risk analysis for competing endpoints been performed?	Yes: 0 of 33

studies, 442 patients), 3D metal printing was used to manufacture the CTAC, whereas in 491 hips (12 studies, 485 patients) the CTAC was machined. After applying the criteria for old and new generation as stated earlier, 574 hips (12 studies, 563 patients) were classified as old-generation CTAC and 398 hips (16 studies, 396 patients) were classified as new-generation CTAC. Due to insufficient or missing information on implant/technique characteristics in several papers, it was not possible to classify implant generation (old or new) for 263 hips (5 studies, 259 patients).

### Complications

All complications extracted from the included studies are summarised in Table 3. Meta-analysis on 31 studies including 1069 hips showed a total incidence of implantrelated complications of 24% (95% CI: 18–29%; I<sup>2</sup> of 81%), indicating considerable heterogeneity. Two studies did not report on complications. THA dislocation was the most frequently reported implant-related complication: the meta-analysis on 30 studies, comprising 1006 hips, showed an incidence of hip dislocation of 6.6% (95% CI: 4.7-8.6%; I<sup>2</sup> of 39%). The meta-regression did not indicate any study-level factors, such as liner type or liner fixation method, CTAC generation or surgical approach associated with the incidence of dislocation. Results of the meta-regression indicated that CTAC generation did have an association with the incidence of total implant-related complications: 29% (95% CI: 21-37%) complications for the old generation compared to 16% (95% CI: 9.2-22%) for the new-generation CTAC. Meta-regression did not identify any other study-level factors that influenced complication rates.

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#### *Re-operations and implant failure*

The meta-analysis of 33 studies including 1235 hips found an incidence of re-operation for any reason of 15% (95% CI 10–19%; l<sup>2</sup> of 85%) at a mean follow-up of 46.9 months (range 12-123 months). The meta-analysis found a large heterogeneity between studies. All reasons for re-operations extracted from the included studies are summarised in Table 3. The results of the meta-regression indicated that the mean follow-up (months) and study start date (i.e. the time frame during which the implants were probably introduced) were associated with the incidence of re-operation for any reason. For every month increase in mean study follow-up time, the incidence of re-operation for any reason increased by 0.24% (95% CI 0.04–0.44%). The incidence of implant failure decreased by 0.64% (95% CI 0.1-1.3%) each subsequent year after 1992 (the start date of the first study that was included in this meta-analysis). There were no other study-level factors associated with re-operation for any reason, such as bone interface type, mean number of prior revisions, implant manufacturing process, CTAC generation, screw type, study publication year or methodological quality.

The meta-analysis on CTAC failure included 33 studies and 1235 hips and showed a mean implant failure rate of 12% (95% CI 8.2–16%; *l*<sup>2</sup> of 83%) at a mean follow-up of 46.9 months (range 12–123 months). The results of the meta-regression indicated that follow-up time, number of prior revisions, CTAC generation and start year of the study were associated with the incidence of implant failure. For every month increase in mean follow-up time, the incidence of failure increased by 0.2% (95% CI 0.03– 0.4%). For every additional prior THA revision, the CTAC failure rate increased by 4.2% (95% CI 1.0% to 7.5%).

 Table 3
 Summary of implant-related complications, re-operations and failure rates.

	Number of			
	Studies	Hips	Incidence	95% CI
Implant-related				
complications				
Prosthetic joint infection	31	1069	4.9%	3.3-6.5%
Dislocation	30	1006	6.6%	4.7-8.6%
Nerve lesion	21	743	2.0%	0.9-3.1%
Screw failure	23	714	2.1%	1.0-3.1%
Aseptic loosening	24	810	2.2%	1.3-3.3%
Periprosthetic fracture	27	902	2.5%	1.5-3.5%
Total implant-related complications	31	1069	24%	18–29%
Re-operations				
Prosthetic joint infection	32	1157	5.3%	3.6–7.1%
Dislocation	31	1148	2.9%	1.8-4.0%
Screw failure	25	877	1.1%	0.4-1.8%
Aseptic loosening	26	976	2.5%	1.5-3.4%
Periprosthetic fracture	25	914	1.4%	0.7-2.2%
Total re-operations	33	1235	15%	10–19%
Total CTAC failures	33	1235	12%	8.2–16%

Regarding the evolution of the CTAC technique, the year in which the study started was positively associated with implant survival: implant failure rate decreased by 0.6% (95% CI: 0.04–1.1%) for each year the study started after 1992. Regarding CTAC generation, implant failure rates were higher for the old generation; 16% (95% CI: 10–21%), compared to 6.4% (95% CI: 1.6–11%) for the new generation. This result remained similar after correcting for study follow-up time. There were no other study-level factors associated with a change in the incidence of implant failure, such as bone interface type, mean number of prior revisions, implant manufacturing process, screw type, liner type or liner fixation method, study publication year or methodological quality.

### Functional outcomes

The majority of the included studies reported on clinical outcome scores assessed by the physicians (29 of 33 studies). Harris hip score (HHS) was the most frequently used outcome measure (21 of 33 studies). Seven studies reported on the Oxford hip score (OHS) as a patient outcome measure. For 676 hips (18 studies, 664 patients) the pre- and post- operative HHS were available. A mean 40 points (95% CI: 35-44 points; 12 of 93%) increase in the post-operative HHS was found (Fig. 2). Meta-regression indicated that bone interface type influenced the post-operative gain in HSS, showing favourable HHS results for hydroxyapatite-coated implants over porous surfaced implants or implants with combinations of porous surfaces and hydroxyapatite coatings. For hydroxyapatite coated implants, the postoperative HHS increased by 62 points (95% CI: 48-76 points), compared to a 44-point (95% CI: 38-51 points) increase for porous coated implants and a 38-point (95% CI: 32-44 points) increase for implants with porous and hydroxyapatite coatings.





As for the surgical approach, the posterolateral approach resulted in a mean HHS improvement of 44 points (95% CI: 39–49 points), compared to a mean increase in HHS of 37 points (95% CI: 29–46 points) for the studies reporting various surgical approaches. The use of only the direct lateral approach was reported in one study, which showed a mean increase of HHS of 23 points (95% CI: 4–42 points). Age, follow-up time, mean number of prior revisions, implant manufacturing process, screw type, liner type and fixation method or CTAC generation were not associated with the post-operative HSS.

## Discussion

Our meta-analysis showed an overall CTAC implantrelated complication rate of 24% in reconstructing large acetabular defects (Paprosky type 3A and 3B and AAOS type 3 and 4 acetabular defects). Dislocation and prosthetic joint infection were the most frequent reported complications. The new CTAC generation of implants was associated with a lower overall implant-related complication rate (16%) compared to the old generation (29%). This difference remained, also after correction for possible modifiers such as follow-up time. The latter stresses the important positive evolution over time of this complex acetabular revision technique with CTAC. We could not identify any other study-level predictors (such as CTAC generation, liner type, surgical approach, surgical time, number of prior revisions) that influenced the rates of dislocation, prosthetic joint infection or any other type of implant-related complication. Furthermore, our analysis shows that at a mean follow-up of nearly 4 years, the overall CTAC implant failure rate was 12%. Interestingly, implant failure rates decrease as the start date of the study increases, and the incidence of CTAC

### Figure 2

Forrest plot of studies analysing gained postoperative HHS. The black diamond shows overall effect size.

failure is higher for older generation implants (16%) as compared to new-generation CTACs (6.4%). We were unable to identify any other associations between studylevel factors and the incidence of implant failure. Finally, the result of our meta-analysis showed that the CTAC technique resulted in a clinically relevant increase in postoperative functional outcomes, with an overall HHS gain of 40 points, although with large variability between studies. No relation between HHS and study-level factors was found, apart from bone interface type; for hydroxyapatite-coated implants, the post-operative HHS increased the most, by 62 points (95% CI: 48–76 points).

We are not aware of any other meta-analysis performed on this specific subject on extensive acetabular defect reconstruction. Two systematic reviews focussing on CTAC for THA revision in the presence of Paprosky 3A, 3B, AAOS III or IV defects are available in the literature (13, 14). However, these systematic reviews lack appropriate meta-analytic techniques, do not take into account the between-study variation and have not evaluated sources of possible heterogeneity. The systematic review by de Martino et al. (13) published in 2019, reports on 17 papers including 579 hips. They report a 17.3% overall re-operation rate, which is in line with our study results (15%). The overall complication rate of 29% reported in their study is higher compared to our analysis (24%). CTAC revision-free survival reported in this study was 82.7% at a mean follow-up of 57 months. It was unclear, however, how the authors classified implant failure. We classified implant failure as full or partial hip revision and have found a 12% failure rate, resulting in an 88% survival rate at 47 months. Martino et al. present a mean gain in post-operative HHS of 38.6 points, based on eight studies (293 hips), which is consistent with our results that show a mean of 40 points gained based on 676 hips. A systematic review by Chiarlone et al. (14) published in 2020, including 18 papers and 634 hips, is very similar to the review published by de Martino et al., with almost identical included papers (15/18) and comparable study results. In contrast to our study, both the abovementioned reviews lack a meta-analysis and the presented outcomes of pooled data do not account for between-study variation (heterogeneity), nor do they try to explain it. In the period after these reviews, a significant additional amount of studies have been published which we were able to include in our systematic review, to a total of 33 papers and 1235 hips.

Our results revealed substantial between-study variation, indicating that it is important to use the appropriate meta-analytic and meta-regression techniques. We found that the following study-level factors were associated with the study's primary and secondary outcomes: CTAC generation, study start date, follow-up time and bone interface type.

The new CTAC generation was associated with a lower overall implant-related complication rate compared to the old generation. It is possible that improved patient selection, increased experience and improved techniques in general (surgical tools, implant designs, 3D printing, etc.) could (partially) explain this improvement, but the available data did not allow to explore these factors in more detail. CTAC generation was also associated with implant failure, favouring the new-generation implants. Since studies with new-generation implants were published more recently and had a shorter follow-up, a correction for the duration of follow-up was performed and the results remained similar with a remarkable lower implant failure rates for new-generation implants as compared to the old-generation implants. Therefore, based on our data, we observed that CTAC technique performance (i.e. lower failure rates) has improved over time.

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We observed an association between the type of bone interface and improved functional outcomes, favouring hydroxyapatite-coated implants over porous surfaced implants and implants combining porous and hydroxyapatite interfaces. Nevertheless, care must be taken to interpret these results, as this association does not imply causality. Data on bone interface type supplied in the included studies were heterogeneous and often lacked specifics on details such as total contact area, HA/porous ratios, pore geometry, pore diameter or porosity percentage. Also, new-generation CTACs often incorporate highly porous 3D printed titanium interfaces, which could be seen as a completely separate bone interface entity, but this could not be differentiated in our analysis. Other unidentified factors may have played a role in the superior clinical outcome of hydroxyapatitecoated CTAC. Hydroxyapatite promotes osteointegration, and it might be that hydroxyapatite produces a more stable implant, resulting in better functional outcomes, as has been shown in primary THA literature (53, 54, 55). Nevertheless, the latter is also seen in early and mid-term results of the latest generation non-custom highly porous 3D printed titanium acetabular components used for primary and revision THA (56, 57).

We should consider some limitations. First, most papers in this analysis are observational studies with a high between-study heterogeneity and a moderate level of methodologic quality. Identified associations between outcomes and factors such as CTAC generation, study start date and bone interface type that were found in our metaregression analysis are based on study-level associations and do not imply association on a patient level. Secondly, many studies lacked detailed information on CTAC designs, manufacturing processes, surgical techniques or peri-operative tools such as patient-specific instruments. Therefore, for several studies, only a distinction between an old- and new-generation CTAC could be made based

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on general or indirect implant characteristic information based on, implant manufacturer.

Based on this study, several recommendations for future research can be made. In general, it is important to provide detailed CTAC specifications and surgical techniques in future studies in order to perform more valid comparisons. Secondly, considering the innovation phase of the CTAC technique as such, it is important to evaluate implant safety. One of the areas of potential safety concern for custom acetabular components would be long-term implant fixation/osseointegration. Since no long-term data on the fixation of any new type of implant can be available, radiostereometric analysis (RSA) can be of value for CTAC implant evaluation. RSA can be used as an adjunct to identify early migration patterns in order to predict long-term implant fixation and survival, as demonstrated before in primary total hip arthroplasty RSA studies (58, 59). Lastly, the custom shape of a CTAC necessitates precise implant positioning for biomechanical hip/pelvis reconstruction as well as an adequate implantto-bone apposition and screw placement. Intra-operative difficulties can arise when the pre-operative plan and the custom implant shape do not match the peri-operative situation. Nevertheless, positioning accuracy studies on new-generation CTAC do demonstrate very high levels of accuracy between the pre-operative plan and the achieved position in regards to cup anteversion, cup inclination, implant rotation and hip centre of rotation restoration (38, 41, 45, 60, 61). As a sequelae to implant positioning analysis, studying the usage of intra-operative navigation can be beneficial to assess if navigation could improve implant positioning and thereby implant performance.

The results of this study are of value for orthopaedic hip surgeons in clinical decision-making and in informing patients. Identifying which aspects of this technique, and if changes to this technique seen over the past decade, have resulted in improvements in CTAC performance and patients' functional outcomes is of importance for clinicians and implant manufacturers, seeking to use and/ or improve the CTAC technique.

## Conclusion

The use of CTAC in revision THA for large peri-acetabular bone defects demonstrates satisfactory complication and implant failure rates and the CTAC technique improves post-operative clinical outcomes. Our meta-regression analysis showed that there is a clear association between improvements in CTAC performance (such as lower complication and implant failure rates) and the evolution of this technique over time. Nevertheless, it is still unclear which part of the technique evolution contributed most to this improvement. The quality of evidence of the included studies was moderate and further research is important, either to assess long-term outcomes and in understanding which innovations have, and will lead to further optimization of this technique, for the better of patient outcomes.

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#### **Supplementary materials**

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This is linked to the online version of the paper at https://doi.org/10.1530/EOR-22–0081.

#### **ICMJE conflict of interest statement**

The authors declare that there is no conflict of interest that could be perceived as prejudicing the impartiality of the research reported.

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