

BMJ Open Effects of female genital cutting on physical health outcomes: a systematic review and meta-analysis

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ABSTRACT

Objective: Worldwide, an estimated 125 million girls and women live with female genital mutilation/cutting (FGM/C). We aimed to systematically review the evidence for physical health risks associated with FGM/C.

Design: We searched 15 databases to identify studies (up to January 2012). Selection criteria were empirical studies reporting physical health outcomes from FGM/C, affecting females with any type of FGM/C, irrespective of ethnicity, nationality and age. Two review authors independently screened titles and abstracts, applied eligibility criteria, assessed methodological study quality and extracted full-text data. To derive overall risk estimates, we combined data from included studies using the Mantel-Haenszel method for unadjusted dichotomous data and the generic inverse-variance method for adjusted data. Outcomes that were sufficiently similar across studies and reasonably resistant to biases were aggregated in meta-analyses. We applied the instrument Grading of Recommendations Assessment, Development and Evaluation to assess the extent to which we have confidence in the effect estimates.

Results: Our search returned 5109 results, of which 185 studies (3.17 million women) satisfied the inclusion criteria. The risks of systematic and random errors were variable and we focused on key outcomes from the 57 studies with the best available evidence. The most common immediate complications were excessive bleeding, urine retention and genital tissue swelling. The most valid and statistically significant associations for the physical health sequelae of FGM/C were seen on urinary tract infections (unadjusted RR=3.01), bacterial vaginosis (adjusted OR (AOR)=1.68), dyspareunia (RR=1.53), prolonged labour (AOR=1.49), caesarean section (AOR=1.60), and difficult delivery (AOR=1.88).

Conclusions: While the precise estimation of the frequency and risk of immediate, gynaecological, sexual and obstetric complications is not possible, the results weigh against the continuation of FGM/C and support the diagnosis and management of girls and women suffering the physical risks of FGM/C.

Trial registration number: This study is registered with PROSPERO, number CRD42012003321.

Strengths and limitations of this study

- Based on the studies in this systematic review, there is clear evidence that female genital mutilation/cutting (FGM/C) entails harms to women's physical health throughout their life, from the moment of cutting as an infant or child, to sexuality and childbirth in adulthood.
- The precise estimation of the frequency and risk of complications was not feasible because of the small sample sizes and varying methods of the studies.
- However, the consistency of results with increased risk of several physical harms in women with genital modification is robust.
- Therefore, efforts should be expended in safeguarding girls and women against the physical risks of FGM/C and caring for those who suffer from its consequences.

INTRODUCTION

The centuries-old practice of female genital mutilation/cutting (FGM/C), also known as female circumcision, is a culturally sanctioned practice^{1 2} that consists of “all procedures involving partial or total removal of the female external genitalia or other injury to the female genital organs for non-medical reasons.”¹ According to the WHO typology, there are three main types: type I (clitoridectomy), type II (excision), type III (infibulation or pharaonic circumcision), and type IV, which is used to describe all other harmful procedures to the female genitalia in the absence of medical necessity.¹ Types I–III and unaltered external female genitalia are depicted in figure 1. The nomenclature for the practice varies across regions, ideological perspectives and research frames, and we use the expression preferred by UNICEF and UNFPA, two central policymakers in the global effort to end the practice, ‘female genital mutilation/cutting’ (FGM/C).² Wade explains that Western efforts to end FGM/C since the early 1970s has relied primarily on

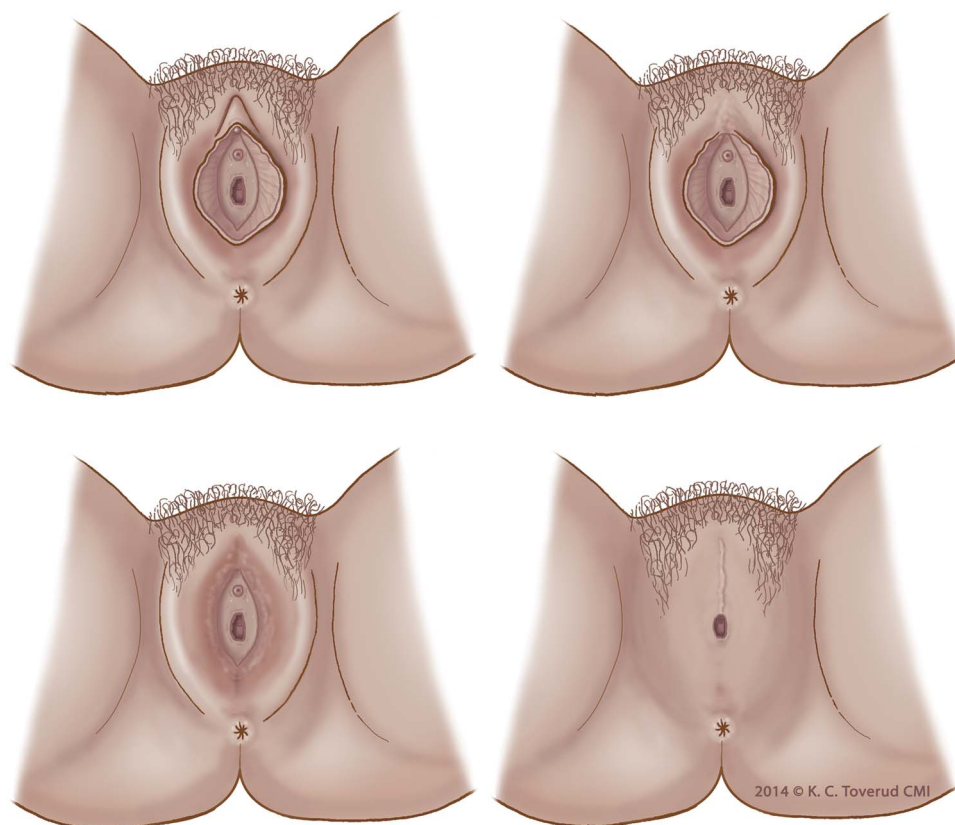


Figure 1 Illustration of unaltered external female genitalia and female genital mutilation/cutting (FGM/C) types I–III. From top left: unaltered external female genitalia, type I (clitoridectomy), type II (excision) and type III (infibulation).

two frames that have influenced the discourse of FGM/C and, in turn, the ideological contestation over the practice.³ In addition to the women's right frame, a dominant frame has been that the practice involves physical and mental harm.

Indeed, for close to a century, observational studies, supported by biological theories, have suggested a negative association between FGM/C and various health outcomes. Such studies have been summarised in a handful of reviews. Epidemiologist and medical anthropologist Obermeyer first reviewed and critiqued the available literature on FGM/C until 1996, and next summarised the subsequent literature from 1997 to 2002.^{4 5} Another early review examined primary data on health complications after FGM/C with particular emphasis on sequelae in childbirth.⁶ Recently, Iavazzo *et al's*⁷ review explored the clinical evidence for an association between FGM/C and infections. The first systematic review in this field—addressing the social, psychological and sexual consequences of FGM/C—critically reviewed and extracted data and analysed 17 comparative studies.^{8 9} While much of this research suggests a harmful effect of FGM/C on women's health, the findings from observational studies and non-systematic reviews are equivocal. Additionally, some commentators have questioned the evidence for many of the claimed short-term and long-term medical consequences of FGM/C, such as quality of sex life, obstetric complications and infections.¹⁰

We aimed to systematically review the evidence for the range of physical health risks associated with FGM/C by summarising the findings from three technical systematic review reports detailing the association.^{11–13}

METHODS

We conducted a systematic review of the physical health risks and consequences of female genital modification in accordance with the Cochrane Handbook for Systematic Reviews of Interventions and PRISMA (Preferred Reporting Items for Systematic reviews and meta-Analyses) guidelines,^{14 15} using a predetermined protocol (PROSPERO, number CRD42012001915). The full details of the systematic review including the search strategies, risk of bias assessment and analysis are found in three technical reports.^{11–13}

Search strategy

The search strategy was developed and undertaken by an experienced search specialist. Fifteen international electronic literature databases were searched to identify research published between inception and January 2012, with Medical Subject Headings descriptors and keywords such as infibulation, applying neither methodology search filters nor language or other search restrictions. The search strategy for MEDLINE (Ovid MEDLINE In-Process & Other Non-Indexed Citations 1946 to 20 January 2012) is shown:

1. Circumcision, Female/
2. ((female\$ or wom#n or girl\$1) adj3 (mutilation\$ or circumcis\$ or cutting\$)).tw.
3. "fgm/c".tw.
4. ((removal\$ or alteration\$ or excision\$) adj6 female genital\$).tw.
5. pharaonic circumcision\$.tw.
6. sunna.tw.
7. (clitoridectom\$ or clitorectom\$).tw.
8. (infibulat\$ or reinfibulat\$ or deinfibulat\$).tw.
9. or/1–8

One reviewer (RCB) manually screened the bibliographies of published reviews on FGM/C and all included studies for additional qualifying studies. RCB did additional searches for the relevant grey literature and unpublished studies in OpenGrey, OpenSigle, OAIster, browsed websites of six international organisations that are engaged in projects regarding FGM/C, and communicated with experts in the field.

Selection of studies and extraction of data

Studies retrieved were eligible for inclusion if they satisfied all our criteria: Be an empirical quantitative study with or without a comparison group published in any language that presented original quantitative data for physical health outcomes in women who had undergone any type of FGM/C as defined by the WHO.¹ All physical health outcomes were eligible, including but not limited to death, infections, infertility, fistula, pain, urinary complications, shock (primary outcomes), and bleeding/haemorrhage, menstrual complications, obstetric complications, vaginal calculus formation, cysts, tissue injury, fractured/displaced bones, urethral meatal stenosis/ urethral stricture, abscesses, keloid and other scarring (secondary outcomes). We applied the following exclusion criteria: Qualitative studies, studies without a quantitative measure of a physical consequence of FGM/C, and all genital modifications not captured by the WHO stated FGM/C definition.

Screening, quality appraisal and data extraction were independently undertaken by two investigators (RCB and VU), with discrepancies resolved by consensus. The two investigators confirmed the eligibility of first titles and abstracts and then full texts. Quality assessment of the identified studies was undertaken as recommended in the Cochrane Handbook, using design specific checklists based on the User's Guide framework.¹⁶ This was done at the study level. The investigators extracted study information and data onto a standardised data collection form, which had been piloted. Data extracted included publication details, study design, sample characteristics, FGM/C characteristics, methods of outcome measurement and health consequences. We contacted authors for additional data or clarification where needed.

Statistical analysis

We grouped the data according to outcomes across the studies, keeping the outcome categories or labels as

reported in each individual study. We estimated associations for dichotomous unadjusted variables in terms of relative risks (RR) with 95% CIs. ORs and 95% CIs were used for case-control studies and adjusted analyses. Outcomes that were sufficiently similar across studies, and reasonable resistant to biases and relatively homogeneous in this respect, were aggregated in meta-analyses. When available, we pooled adjusted estimates; otherwise, we pooled the unadjusted estimates based on crude data from the individual studies. ORs and RRs greater than one indicate an increased risk of complications with FGM/C; if less than one, they indicate a decreased risk.

We anticipated heterogeneity between studies due to different study methodologies and geographical and population differences. Heterogeneity was examined using the χ^2 test and I^2 statistic. We used a random-effects model to account for within-study and between-study heterogeneity.

In random-effects meta-analysis, the weight assigned to each included study is adjusted to include a measure of variation (τ^2) in the effects reported between studies. We used the Mantel-Haenszel method for unadjusted dichotomous data, and for adjusted data we used the generic inverse-variance method, in which weight is given to each study according to the inverse of the variance of the effect, to minimise uncertainty about the pooled effect estimates. Analyses were done with Review Manager (V.5.2.8).

We applied the instrument Grading of Recommendations Assessment, Development and Evaluation (GRADE) to assess the extent to which we have confidence in the effect estimates.¹⁷ GRADE is a transparent and systematic approach to grading our confidence in the evidence. For resource reasons, we used GRADE only for outcomes eligible for meta-analysis.

Those of us who did the systematic review were not masked to the authors, institution or journal of publication. The use of non-masked reviewers is accepted practice in meta-analyses and has been shown not to bias results.¹⁸ In line with recommendations,¹⁴ results from the studies deemed to have the highest internal validity were given preference. In this communication, we present all studies that reported outcomes for differentially FGM/C exposed groups of women, that is, studies with a comparison group.

Role of the funding source

Norad and the WHO commissioned the study and the latter contributed some funding (\$10 000). The commissioners of the systematic review had no role in the study design, data collection, data analysis, data interpretation or writing of the report. RCB had final responsibility for the decision to submit for publication.

RESULTS

Our search strategy identified 5109 unique publications, the titles and abstracts of which were screened for

inclusion. The full text of 12 publications could not be located, while 431 articles were retrieved, of which 185 met the inclusion criteria (figure 2).

The 185 included studies were of varying publication types, published between 1927 and 2011, and from 42 different countries (see online supplementary appendix 1). There were 13 studies from the Middle East, 43 from a Western country, and 129 from Africa. Twenty different African countries were represented. The FGM/C participants in studies from a Western country originated in the majority of cases from Somalia, and in the remaining cases they originated from another African country where FGM/C is commonly practised. Overall, the 185 studies involved 3.17 million female participants, from infants to women in their 70s, with a mean age of approximately 30. With respect to the FGM/C characteristics, the majority of women had genital alteration that involved the cutting and removal of portions of the external female genitalia, without stitching, corresponding to either type I or type II. The procedure had in the absolute majority of cases been undertaken in early childhood, usually before the age of 10, by a traditional circumciser. A total of 75 different outcomes were extracted.

In this overview, we present key physical health complications of FGM/C in a life course perspective. Except for some immediate outcomes, these key outcomes derive from comparative studies, that is, women with FGM/C are compared to women without FGM/C with respect to an outcome in a cohort, case-control or cross-sectional study (table 1). We prioritise the presentation of studies with clinically measured and adjusted outcome data, but note also the best available evidence for additional key outcomes, largely immediate complications. Table 1 shows the 57 studies with the best

available evidence regarding the physical health sequelae of FGM/C (comparative cohort, case-control, cross-sectional studies).^{19–72} About 40% of the outcomes were self-reported primarily by adult women, although the great majority of the obstetric and some genitourinary outcomes were clinically measured. The meta-analytical results that are based on unadjusted estimates are presented in figure 3, and those based on adjusted estimates are shown in figure 4.

Immediate complications

In most cases of FGM/C, a girl's clitoris and labia are cut away, often with a crude unsterile instrument and without anaesthetics by a traditional practitioner who has little knowledge of female anatomy.² Thus, it is reasonable to assume that physiological harms such as bleeding ensue during the cutting process and the short-term postprocedure period. We identified no studies that analysed the potential statistical differences in the risk of direct, procedure-related complications between types of FGM/C. However, 56 observational studies reported on eight main types of immediate medical harms (bleeding, shock, genital tissue swelling, fever, infections and problems with urination and wound healing) on 133 515 females of various ages and types of FGM/C. The rate of immediate complications varied greatly across the studies. There were strong indications of under-reporting of immediate complications from the procedure, with some studies reporting that 90% of the girls undergoing FGM/C experienced no bleeding at all.^{30 64} However, representative studies (ie, where the participants can be assumed to represent the larger population) of moderate and high methodological quality indicated that the most common immediate

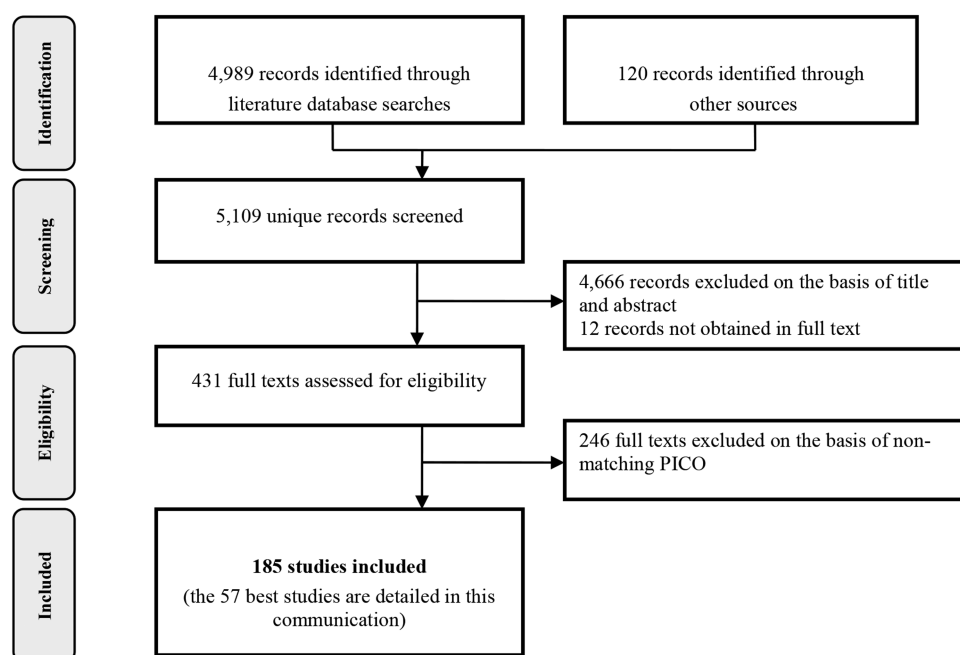


Figure 2 PRISMA flow diagram for selection of literature.

Table 1 Summary of included comparative studies (N=57)

Author, year	Study design	Quality	N	Country	Outcomes (assessment type)
Adinma, 1997 ¹⁹	Cross-sectional	Low	256	Nigeria	Obstetric (self-report)
Almroth, 2005a ²⁰	Case-control	High	279	Sudan	Infertility (clinical)
Almroth, 2005b ²¹	Cross-sectional	High	255	Sudan	Genitourinary (clinical)
Alsibiani, 2010 ²²	Cross-sectional	Moderate	260	Saudi Arabia	Sexual (self-report)
Balk, 2000 ²³	Based on the Sudan DHS 1990	Moderate	5856	Sudan	Infertility (self-report)
Berardi, 1985 ²⁴	Prospective	Low	852	France	Obstetric (clinical)
Bohoussou, 1986 ²⁵	Cross-sectional	Low	4935	Ivory Coast	Obstetric (unclear)
Brewer, 2007 ²⁶	Based on the Kenya DHS 2003	High	539	Kenya	HIV/STIs (clinical)
Browning, 2010 ²⁷	Retrospective cohort	Moderate	492	Ethiopia	Genitourinary, obstetric (clinical)
Chibber, 2011 ²⁸	Prospective	Low	4800	Unclear	Obstetric (clinical)
De Silva, 1989 ²⁹	Prospective	Low	2157	Saudi Arabia	Genitourinary, obstetric (clinical)
El Dareer, 1983 ³⁰	Cross-sectional	Low	3210	Sudan	Immediate, genitourinary (self-report)
el Defrawi, 2001 ³¹	Cross-sectional	Moderate	250	Egypt	Genitourinary, sexual (self-report)
Elmusharaf, 2006 ³²	Case-control	High	222	Sudan	HIV/STIs (clinical)
Elnashar, 2007 ³³	Cross-sectional	Low	264	Egypt	Genitourinary, sexual, obstetric, (self-report)
Essén, 2005 ³⁶	Registry study	Moderate	2554	Sweden	Obstetric (clinical)
Fillo, 2007 ³⁷	Based on the Burkina Faso DHS 2003	Moderate	12 477	Burkina Faso	Genitourinary (unclear)
Hakim, 2001 ³⁸	Unclear if prospective or retrospective	Low	1481	Ethiopia	Obstetric (clinical)
Holmgren, 2003 ³⁹	Cross-sectional	Moderate	857	Guinea-Bissau	HIV/STIs (clinical)
Inhorn, 1993 ⁴⁰	Case-control	High	190	Egypt	Infertility (clinical)
Jackson, 2005 ⁴¹	Cross-sectional	Low	?	Ghana	Infertility (self-report)
Johnson 2005 ⁴²	Registry study	Low	5416	USA	Obstetric (clinical)
Jones, 1999-I ⁴³	Cross-sectional	Low	1920	Burkina Faso	Genitourinary (clinical), obstetric (self-report)
Jones, 1999-II ⁴³	Prospective	Moderate	5337	Mali	Genitourinary, obstetric (clinical)
Kanki, 1992 ⁴⁴	Cross-sectional	Moderate	1710	Senegal	HIV/STIs (clinical)
Kaplan, 2011 ⁴⁵	Cross-sectional	Moderate	871	Gambia	Immediate (self-report), genitourinary (clinical)
Klouman, 2005 ⁴⁶	Cross-sectional	Moderate	396	Tanzania	Genitourinary, HIV/STIs, infertility (clinical)
Larsen, 2002 ⁴⁷	Based on the Sudan DHS 1990	Moderate	5849	Sudan	Infertility (self-report)
Larsen 2002 ⁴⁸	Cross-sectional	Low	1836	Nigeria	Obstetric (self-report)
Larsen, 2000-I ⁴⁹	Based on the CAR DHS 1995	Moderate	4388	Central African Republic	Infertility (self-report)
Larsen, 2000-II ⁴⁹	Based on the Ivory Coast DHS 1995	Moderate	5930	Ivory Coast	Infertility (self-report)
Larsen, 2000-III ⁴⁹	Based on the Tanzania DHS 1997	Moderate	6043	Tanzania	Infertility (self-report)
Larsen, 1989 ⁵⁰	Cross-sectional	Low	2183	Sudan	Infertility (self-report)
Lupo, 1999 ⁵¹	Registry study	Low	114	USA	Obstetric (unclear)
Maslovskaya, 2009 ⁵²	Based on the Kenya DHS 2003	High	3114	Kenya	HIV/STIs (clinical)
Millogo-Traore, 2007 ⁵³	Prospective	Low	454	Burkina Faso	Obstetric (clinical)
Morison, 2001 ⁵⁴	Cross-sectional	High	1157	Gambia	Genitourinary, sexual, HIV/STIs, infertility (clinical)
Msuya, 2002 ⁵⁵	Cross-sectional	High	379	Tanzania	Genitourinary, HIV/STIs (clinical)
Ndiaye, 2010 ⁵⁶	Cross-sectional	Low	354	Burkina Faso	Obstetric (clinical)
NSEO, 2002 ³⁴	Cross-sectional	Low	7765	Eritrea	Sexual, obstetric (self-report)
NSO, 1995 ³⁵	Cross-sectional	Low	4775	Eritrea	Sexual, obstetric (self-report)

Continued

Table 1 Continued

Author, year	Study design	Quality	N	Country	Outcomes (assessment type)
Nwajei, 2003 ⁵⁷	Cross-sectional	Low	400	Nigeria	Genitourinary (self-report)
Odoi, 1997 ⁵⁸	Cross-sectional	Low	195	Ghana	Sexual (self-report)
Oduro, 2006 ⁵⁹	Registry study	High	5071	Ghana	Obstetric (clinical)
Okonofua, 2002 ⁶⁰	Cross-sectional	Moderate	1836	Nigeria	Genitourinary, sexual (clinical/self-report)
Orji, 2006 ⁶¹	Cross-sectional	Low	500	Nigeria	Obstetric (self-report)
Pépin, 2006 ⁶²	Cross-sectional	Moderate	1026	Guinea-Bissau	HIV (clinical)
Pépin, 1991 ⁶³	Cross-sectional	Low	345	Gambia	HIV (clinical)
Rushwan, 1983 ⁶⁴	Cross-sectional	Low	2502	Sudan	Immediate, genitourinary, sexual, infertility (self-report)
Shandall, 1967 ⁶⁵	Cross-sectional	Low	4487	Sudan	Immediate, genitourinary (clinical/self-report)
Slanger, 2002 ⁶⁶	Cross-sectional	Moderate	1107	Nigeria	Obstetric (self-report)
Small, 2008 ⁶⁷	Registry study	Low	2 179 322	Multiple	Obstetric (clinical)
Vangen, 2002 ⁶⁸	Registry study	Low	703 925	Norway	Obstetric (clinical)
WHO study group, 2006 ⁶⁹	Prospective	High	28 393	Multiple	Obstetric (clinical)
Wuest, 2009 ⁷⁰	Prospective	Low	232	Switzerland	Obstetric (clinical)
Yount, 2007 ⁷¹	Based on the Kenya DHS 2003	Moderate	3167	Kenya	HIV/STIs (clinical), obstetric (self-report)
Yount, 2006 ⁷²	Cross-sectional	Low	1700	Egypt	Infertility, obstetric (self-report)

Two studies are presented in Jones 1999⁴³ (Jones 1999-I is from Mali, and Jones 1999-II is from Burkina Faso); three studies are presented in Larsen 2000⁴⁹ (Larsen 2000-I is from the Central African Republic, Larsen 2000-II is from the Ivory Coast, and Larsen 2000-III is from Tanzania).

CAR, Central African Republic; DHS, Demographic and Health Survey; Quality, Methodological study quality at study level; STIs, sexually transmitted infections.

complications were: excessive bleeding (median 32%, range 5–62%), urine retention (median 31%, range 8–53%), genital tissue swelling (median 15%, range 2–27%), problems with wound healing (13%) and pain (11%).^{30 73 74} Girls generally suffered more than one immediate complication. We identified three clinical reports on deaths directly attributed to FGM/C.^{75–77} Fourteen studies reported the number of events for different types of FGM/C separately, allowing us to estimate differences in risk across exposure groups.^{30 45 64 65 78–87} Our results indicated that there might be a greater risk of immediate harms with FGM/C type III relative to types I–II. We found few, and small, differences in risk of immediate complications with FGM/C types I–II compared to type IV (generally ‘nick’).

Genitourinary problems

With respect to the genitourinary sequelae of FGM/C, reported years and sometimes decades following the procedure, we identified 17 comparative studies.^{21 27 29–33 37 43 45 46 54 55 57 60 64 65} In total, the studies included 38 390 women. The most frequently measured outcomes were genital tissue damage, vaginal discharge and itching, urological complications and infections. Many sequelae were examined in only one or a few studies and/or they were relatively rare events, such as keloids and abscesses. Analyses were thus often unable to establish whether there were statistically significant differences between the groups being compared and the CIs were wide. As a result, there was insufficient

information available from the studies to assess difference in risk relative to FGM/C exposure. The results were inconclusive with respect to: scarring, keloids, abscesses, fistulae, damaged tissue (perineum, anal sphincter), disfigurement, vaginal obstruction and cysts.

According to four cross-sectional studies (n=3657), there was a trend for a greater risk of vaginal discharge and itching with FGM/C (adjusted ORs (AOR) from 0.94 to 2.81).^{33 54 57 60} Urological long-term complications were reported in four comparative studies (n=3611), none of which could establish a statistically significant difference, either in unadjusted analyses (RRs from 0.85 to 1.78) or in adjusted analyses (AORs from 0.80 to 1.29), between women with FGM/C and women with no FGM/C.^{27 33 54 60} However, results from two studies of moderate to high methodological study quality indicated a trend for a greater risk of burning or painful urination with FGM/C (RR=2.56, 95% CI 0.80 to 8.22; RR=1.66, 95% CI 0.96 to 2.85).^{33 60} Menstrual problems were reported in five studies (n=6564).^{30 31 33 54 64} They showed a trend towards a greater risk of menstrual problems with FGM/C: dysmenorrhoea (RR=1.44, 95% CI 1.11 to 1.86), difficulty in menstruation (RR=1.02, 95% CI 0.06 to 16.28), menstrual problems (RR=0.77, 95% CI 0.61 to 0.97), irregular menses (RR=2.56, 95% CI 1.48 to 3.45) and difficulty in passing menstrual blood (RR=1.75, 95% CI 0.78 to 3.93).

Ten comparative studies (n=28 940) reported results concerning long-term genitourinary infections.^{21 29 30 37 43 46 54 55 64 65} Owing to few studies and

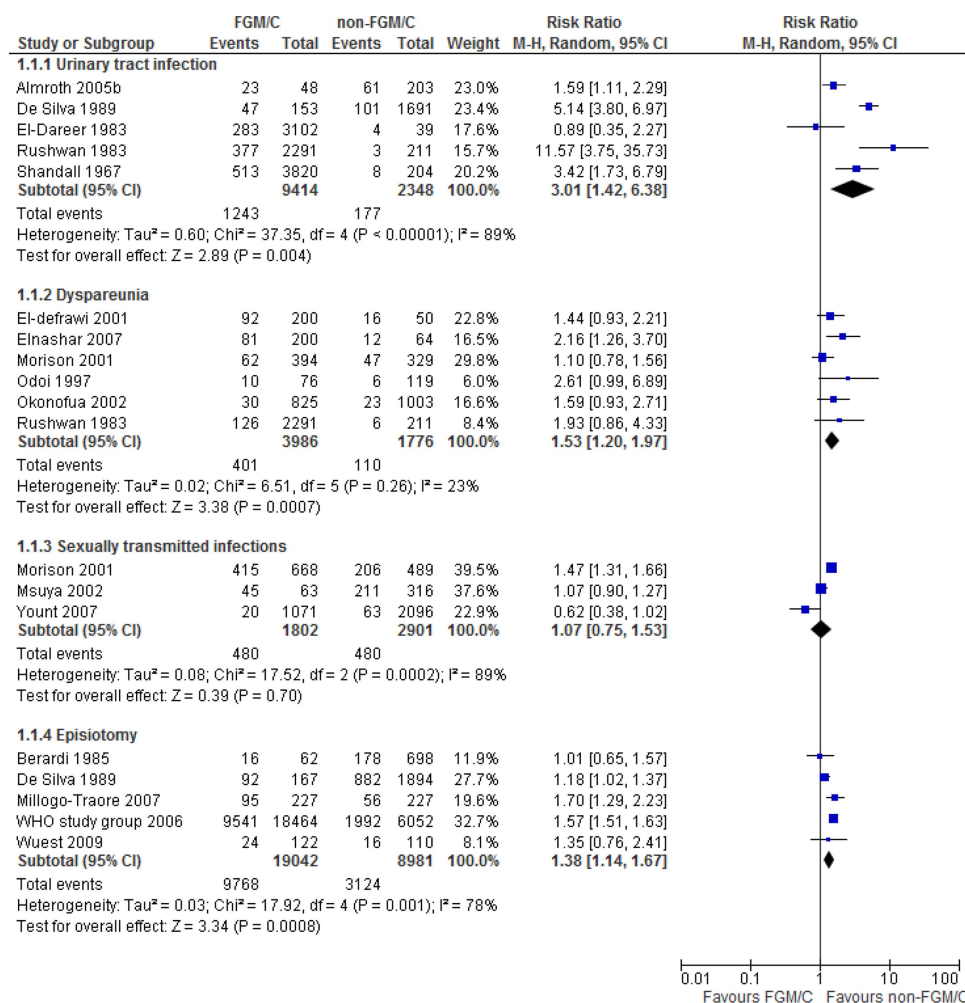


Figure 3 Meta-analyses of urinary tract infection, dyspareunia, sexually transmitted infections, episiotomy (unadjusted effect estimates).

the low number of events, the findings were inconclusive for the following outcomes: chronic pelvic infections, reproductive tract infections, genital infections and vaginitis. However, in adjusted analyses, two studies of low to moderate methodological quality found a statistically higher risk of reproductive tract infections (AOR=1.54, 95% CI 1.08 to 2.21) and genital infections (AOR=1.72, 95% CI 1.02 to 2.92) with FGM/C.^{37 43} Meta-analyses showed a greater risk of urinary tract infections (RR=3.01, 95% CI 1.42 to 6.38; GRADE: very low; figure 3)^{21 29 30 64 65} and bacterial vaginosis (AOR=1.68, 95% CI 1.28 to 2.22; GRADE: very low) with FGM/C (figure 4).^{46 54}

Painful sexual intercourse

Dyspareunia (painful sexual intercourse) was reported in six studies (n=6204).^{31 33 54 58 60 64} The meta-analysis, presented in figure 3, showed an increased risk of dyspareunia with FGM/C (RR=1.53, 95% CI 1.20 to 1.97; GRADE: very low). Correspondingly, results from two nationally representative studies from Eritrea (n=12 540) indicated a 'dose-response' relationship, with a lower risk of problems during sexual relations with FGM/C

types I–II relative to type III (RR=0.19, 95% CI 0.16 to 0.24; RR=0.44, 95% CI 0.27 to 0.72).^{34 35}

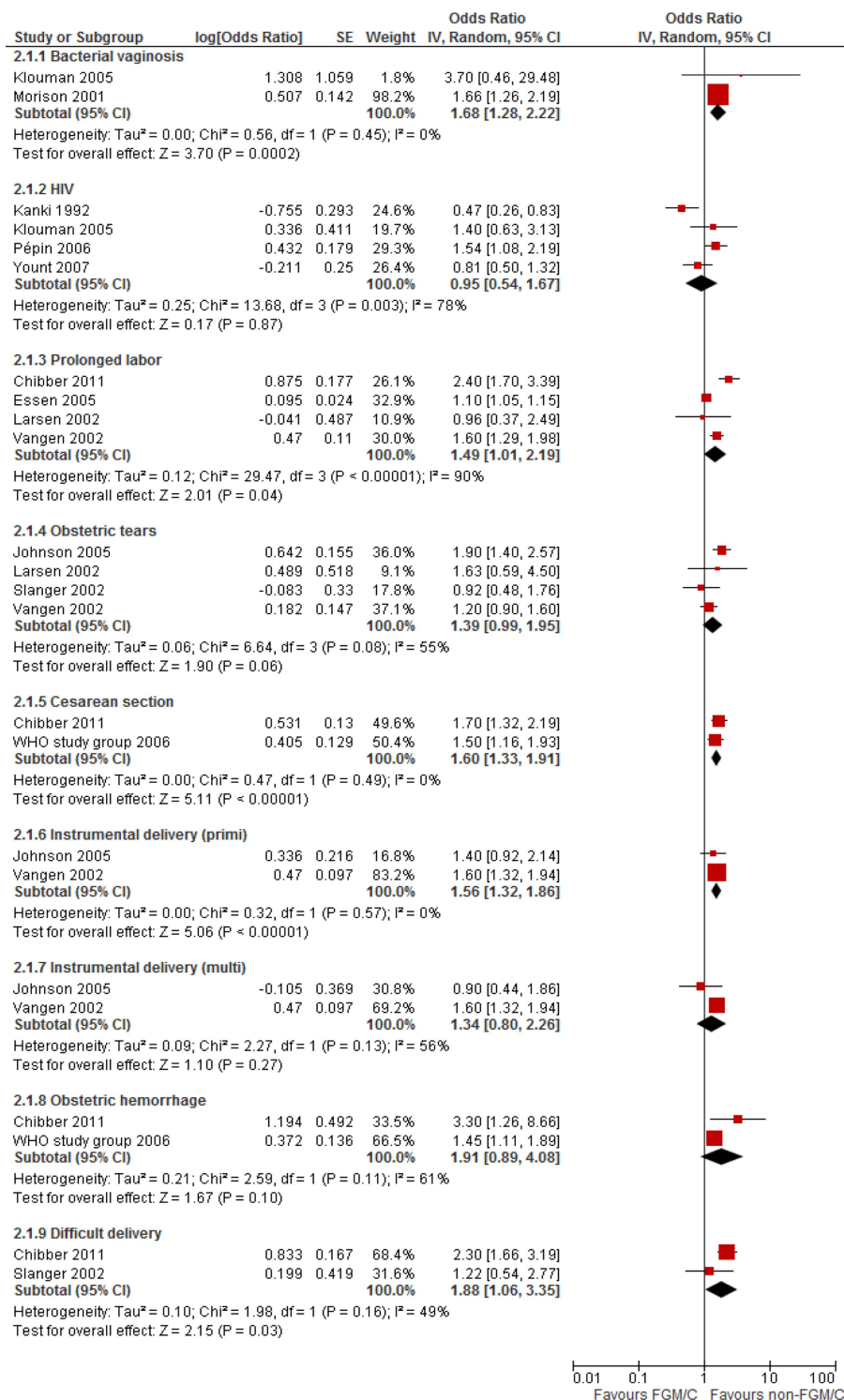
HIV and sexually transmitted infections

HIV and sexually transmitted infections (STIs) were clinically examined in one case-control study and 10 cross-sectional studies (n=12 912).^{26 32 39 44 46 52 54 55 62 63 71} The case-control study could not establish a difference between FGM/C and no FGM/C regarding STIs (AOR=1.13, 95% CI 0.73 to 1.77).³² Similarly, the meta-analysis of cross-sectional studies failed to establish a difference (RR=1.07, 95% CI 0.75 to 1.53; GRADE: very low; figure 3).^{54 55 71} As shown in figure 4, also the meta-analysis for HIV, based on four studies which presented adjusted data, failed to establish a difference relative to FGM/C (AOR=0.95, 95% CI=0.54 to 1.67; GRADE: very low).^{44 46 62 71}

Infertility

Twelve studies presented data on infertility (n=36 473).^{20 23 40 41 46 47 48 50 54 64} Two case-control studies of high methodological quality examined whether FGM/C was a predictor for infertility. The

Figure 4 Meta-analyses of bacterial vaginosis, HIV, prolonged labour, obstetric tears, caesarean section, instrumental delivery, obstetric haemorrhage, difficult delivery (adjusted effect estimates).



case-control study from Sudan could not establish an association between infertility and FGM/C (AOR=1.77, 95% CI 0.52 to 7.10).²⁰ Similarly, the case-control study from Egypt could not establish a difference between FGM/C type II and type I with respect to tubal factor infertility (AOR=1.9, 95% CI 0.8 to 4.2).⁴⁰ The available evidence did not allow us to conclude whether women were more likely to be infertile if they were cut by a traditional than a medical circumciser (AOR=2.1, 95% CI 0.8 to 5.7).⁴⁰ Ten cross-sectional studies examined infertility in women with

and without FGM/C. The association between FGM/C and infertility varied, both in unadjusted analyses (effect estimates ranged from 0.34 to 3.67) and adjusted analyses (AORs from 0.99 to 2.76). Specifically, the results of the two clinical studies of moderate to high methodological quality did not establish a greater risk with FGM/C (OR=1.3, 95% CI 0.7, 2.7; OR=1.05, 95% CI 0.65 to 1.67. Of eight adjusted estimates, two reached significance. These are not shown here because the CIs were not provided in the publications).^{46 54}

Obstetric outcomes

Obstetric events (prolonged labour, tears/lacerations, caesarean section, episiotomy, instrumental delivery, haemorrhage, difficult labour) were reported in 26 comparative studies (2.97 million women).^{19 24–29 33–36 38 42 43 48 51 53 56 59 61 66–69 71 72} Seven of these were prospective.^{24 28 29 43 53 69 70} Eight studies reported adjusted data, with the number and types of confounders varying greatly across studies (detailed in the study under review).^{28 36 42 43 48 66 68 69}

Data on prolonged labour were reported in six studies.^{28 29 36 48 53 68} The meta-analysis of adjusted estimates from four studies showed a significantly greater risk of prolonged labour with FGM/C (AOR=1.49, 95% CI 1.01 to 2.19; GRADE: low; figure 4).^{28 36 48 68} There was one prospective study, of low to moderate methodological quality, that reported an adjusted estimate for prolonged labour, the result of which was concordant with the meta-analysis (AOR=2.40, 95% CI 1.40 to 2.80).²⁸ Regarding obstetric tears/lacerations, the meta-analysis of four studies which presented adjusted data showed an AOR of 1.39 (95% CI 0.99 to 1.95; GRADE: very low; figure 4).^{42 48 66 68} No prospective studies presented adjusted estimates for obstetric tears.

There were 15 studies with data on caesarean section.^{24 25 28 29 33 42 48 56 59 66–71} Five studies reported adjusted estimates, the pooled estimate of which resulted in an AOR of 1.32 (95% CI 0.97 to 1.80; GRADE: very low). Restricting the meta-analysis to the two prospective studies established a significant difference between groups (AOR=1.60, 95% CI 1.33 to 1.91; GRADE: low), indicating a greater risk of caesarean section among women with FGM/C (figure 4).^{28 69} As with the other obstetric outcomes, the study-level results were inconsistent regarding episiotomy. Eleven studies reported on episiotomy,^{19 24 29 33 38 48 53 56 66 69 70} but there was only one (retrospective) study with adjusted data.⁴⁸ The result from the most comprehensive model in this study (ie, adjusting for the highest number of confounders) showed an AOR of 1.18 (95% CI 0.76 to 1.84). No prospective studies presented adjusted estimates for episiotomy, but we aggregated the unadjusted results from five prospective studies (n=32 088 women), finding an increased risk with FGM/C (RR=1.38; 95% CI 1.14 to 1.67; GRADE: very low; figure 3).^{24 29 53 69 70}

There were nine studies with data on instrumental delivery.^{24 25 29 42 53 66–68 70} Two (registry based) studies reported adjusted data.^{42 68} The meta-analysis for primiparous women suggested a greater risk of instrumental delivery with FGM/C (AOR=1.56, 95% CI 1.32 to 1.86; GRADE: very low), which could not be firmly established for multiparous women (AOR=1.34, 95% CI 0.80 to 2.26; GRADE: very low; figure 4). We also included nine studies with data on obstetric or postpartum haemorrhage.^{28 29 38 42 56 66 68–70} Five studies reported adjusted results, which we combined in a meta-analysis. The result indicated a greater risk with FGM/C (AOR=1.50,

95% CI 1.22 to 1.84; GRADE: very low). However, the pooled adjusted estimate based on the two prospective studies that reported adjusted data for haemorrhage failed to establish a convincing difference relative to FGM/C (AOR=1.91, 95% CI 0.89 to 4.08; GRADE: very low; figure 4).^{28 69} Lastly, we included six comparative studies with data on difficult delivery.^{28 42 43 56 66} The pooled result based on adjusted estimates from the two studies that could be combined resulted in an AOR of 1.88 (95% CI 1.06 to 3.35; GRADE: low; figure 4).^{28 66} A third study compared women without FGM/C with women who had FGM/C type I. The AORs were 0.17 (95% CI=0.06 to 0.52) and 0.32 (95% CI=0.19 to 0.54), which favoured not having FGM/C.⁴³ There was one prospective study with data on difficult delivery.²⁸ The estimate showed a greater risk with FGM/C (AOR=2.30, 95% CI 1.3 to 2.5).

DISCUSSION

This systematic review provides clear evidence that FGM/C entails harms to women's physical health throughout their life, from the moment of cutting as an infant or child, to sexuality and childbirth in adulthood. Predictably, the most common direct, procedure-related complication includes haemorrhage, most likely resulting from laceration of the internal pudendal artery or the clitoral artery. It is difficult to determine the number of females who die from procedure-related complications. Only a few studies reported death, but highly publicised fatalities from FGM/C heighten the awareness of the possible harms posed by the procedure, such as three recent cases in Egypt and Kenya.^{88–90}

We found several long-term consequences of FGM/C, including increased risks of urinary tract infections, bacterial vaginosis, dyspareunia and obstetric complications. Studies have been published since we conducted our search, and they corroborate our findings.^{91–98} The identified risks from FGM/C are also supported by biological rationales for the associations. As explained by experts,^{21 54 92} any alteration of the natural anatomy of the vulva, such as removal of the protective labia minora, can lead to structural and physiological changes, including trauma to the urethra, adjacent tissues and nerves at the time of the procedure as well as formation of scars and flaps of skin during the healing process.

FGM/C is a non-therapeutic procedure, which by definition does not treat an underlying pathological process, but rather forms an actionable risk factor for several morbidities. As public policy discussions gain momentum on how to prevent FGM/C, our findings about the health complications could be communicated to practising communities, health practitioners, government bodies, international health organisations and other stakeholders. Global policy efforts should be expended in safeguarding females against the physical risks of FGM/C and, as emphasised by the United Nations agencies,¹ in upholding their bodily integrity

and furthering their human rights. Worldwide, about 125 million girls and women alive today live with FGM/C, and every year another three million girls in the countries where the practice is concentrated are at risk of undergoing the practice.^{2 99} Caring for girls and women who suffer from its consequences and research into the best and most acceptable treatment and care are important.

This systematic review offers the most comprehensive and scientific evidence presently available on the range of physical health complications from FGM/C—other reviews are non-systematic and limited in scope^{4–7}—but a complete understanding is hindered by data gaps. The search was executed in 2012 and it is possible that our systematic review is subject to publication bias. The discrepancy between the large number of records identified in our search and the 185 studies included attest to the literature on FGM/C being longer on opinion and polemic and considerably shorter on data. We identified only three case-control studies and seven prospective studies. Many included studies had low methodological study quality; many outcomes were reported in just a few studies, and event rates in several studies were low. Thus, most data are of low quality, which partly results from the nature of the study question, which is difficult to answer by means of rigorous research methods. Additionally, the measurement and reporting of exposure, outcome and confounders were done with varying precision in the included studies, and although we did subgroup and random-effects analyses, these could not fully account for the observed heterogeneity across studies. Thus, the summary estimates for all outcomes should be interpreted cautiously. Some of the above possible sources of biases would tend to bias our summary estimates towards the null, and are likely to lead to too conservative estimates of the true harmful association of FGM/C with physical health. For example, research on male circumcision has found that adverse events are recorded less frequently in retrospective than prospective studies.¹⁰⁰ Owing to gaps in the evidence base, precise estimation of frequency and risk of complications is not possible. However, irrespective of the exact size of the greater risk from FGM/C, the consistency of results with increased risk of several physical harms in women with genital modification is robust, and even the lowest increase in risk of complications is undesirable from a woman's health perspective.

CONCLUSION

The evidence base on the physical health complications of FGM/C, which covers over half a century of research from more than 20 countries in Africa and beyond, shows that FGM/C is associated with an increased risk of health complications, especially urinary tract infections, bacterial vaginosis, painful sexual intercourse and obstetric difficulties. Further research into this question is unlikely to produce practical value. Rather, efforts

should be expended in safeguarding girls and women against the physical risks of FGM/C and caring for those who suffer from its consequences.

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Appendix 1: Reference list of the 185 studies meeting the eligibility criteria for the systematic review on the range of physical health risks associated with FGM/C

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