Modifying the health system to maximize voluntary medical male circumcision uptake: a qualitative study in Botswana

Background: In 2007, the World Health Organization and the Joint United Nations Programme on HIV/AIDS endorsed voluntary medical male circumcision (VMMC) as an add-on HIV-prevention strategy. Similar to many other sub-Saharan countries, VMMC uptake in Botswana has been low; as of February 2016, only 42.7% of the program target had been achieved. Previous work has examined how individual-level factors, such as knowledge and attitudes, influence the update of VMMC. This paper examines how factors related to the health system can be leveraged to maximize uptake of circumcision services, with a focus on demand creation, access to services, and service delivery.

Methods: Twenty-seven focus group discussions with 238 participants were conducted in four communities in Botswana among men (stratified by circumcision status and age), women (stratified by age), and community leaders. A semi-structured guide was used by a trained same-gender interviewer to facilitate discussions, which were audio recorded, transcribed, translated to English, and analyzed using an inductive analytic approach.

Results: Participants felt demand creation activities utilizing age- and gender-appropriate mobilizers and community leaders were more effective than mass media campaigns. Participants felt improved access to VMMC clinics would facilitate service uptake, as would designated men’s clinics with male-friendly providers for VMMC service delivery. Additionally, providing comprehensive pre-procedure counseling and education, outlining the benefits and disadvantages of the surgical procedure, and explaining the differences between the surgical and non-surgical procedures, were suggested by participants to increase understanding and uptake of VMMC.

Conclusion: Cultural acceptability of circumcision services can be improved by engaging age- and gender-appropriate community mobilizers. Involving influential community leaders, providing a forum for men to discuss health issues, and bringing services closer to people can increase VMMC utilization. Service delivery can be improved by communicating the pros and cons of the procedure to the clients for informed decision-making.

Keywords: HIV, demand creation, service delivery, sub-Saharan Africa

Introduction

In 2007, the World Health Organization (WHO) and the Joint United Nations Programme on HIV/AIDS endorsed voluntary medical male circumcision (VMMC) as an add-on HIV-prevention strategy following the results of 3 randomized control trials demonstrating the procedure reduces risk of acquisition of HIV by 40%–60%. Subsequently, multiple WHO priority countries in sub-Saharan Africa with high HIV...
prevalence rates and low levels of circumcision implemented VMMC programs. The ambitious targets were to circumcise 80% of HIV-negative men by 2016. While some countries such as Ethiopia, Kenya, and Tanzania were able to make substantive progress toward meeting targets, the majority found that VMMC uptake was substantially below targeted levels.

In Botswana, ~1 in 5 adults are living with HIV, a prevalence second only to Uganda and Swaziland worldwide. Historically, the country has been a leader in HIV prevention, care and treatment; in 2002, it became the first country in sub-Saharan Africa to provide antiretroviral therapy (ART) free of charge to its citizens and its prevention of mother-to-child transmission program has reduced the rate of infant-acquired infection to below 2%, a rate comparable with that observed in the USA. More recently, in 2016, Botswana adopted the WHO HIV treatment guidelines recommending ART for all persons diagnosed with HIV infection, regardless of their CD4 cell count or viral load.

As a WHO priority country, Botswana, committed to circumcising 385,000 HIV-negative men (80% of the negative men in the country). While there has been a steady increase in VMMC uptake, as of February 2016, only 163,911 circumcisions have been conducted, 57.4% below the target.

There have been a number of studies investigating barriers and facilitators to the uptake of VMMC, with the majority of these focusing on individual-level or micro-level factors such as clients’ knowledge, attitudes, and beliefs. For example, barriers noted in the literature include fear of procedure-related complications and pain as well as unfavorable perceptions about the 6-week abstinence from sexual intercourse in the period following circumcision. Data from Botswana indicated that improved hygiene and cleanliness were major determinants in men’s decision to get circumcised. Increased sexual pleasure and perceived attractiveness were also influencers. Additionally, the desire to fit in and be fashionable as well as protection against HIV and other infections were reported as triggers to male circumcision. Fear of HIV testing acted as a barrier for some men’s participation in circumcision.

This substantial body of literature focuses on individual-level motivators and barriers for VMMC. However, little has been published about modifiable health system factors that affect uptake of VMMC. In fact, a recent review article calls for countries to critically examine these systems-level factors that influence circumcision rates. To our knowledge, only 1 study in Botswana has previously examined health systems factors related to uptake of VMMC; the study, which was designed to evaluate a specific 6-month communication strategy to increase community-level awareness of VMMC, identified long distances to treatment centers, long waiting periods, and shortage of staff and equipment as key infrastructural impediments. Further research, not limited to an evaluation of a specific intervention, is needed to identify the modifiable systems-level factors that influence uptake of VMMC generally. Therefore, we sought to identify potential changes within the health system that would maximize uptake of VMMC through improvements in demand creation, access to services, and/or service delivery.

Materials and methods

This study was conducted by the International Training and Education Center for Health (I-TECH) in Botswana, which is a collaboration between the University of Washington and University of California, San Francisco. The study received ethical approvals by the Health Research and Development Committee at the Botswana Ministry of Health and Wellness as well as the University of Washington Institutional Review Board.

Focus group discussions (FGDs) were conducted in 4 communities in Botswana among men, women, and community leaders. These communities were purposefully selected to represent varying circumcision rates. Specifically, 2 of the communities (Gaborone and Palapye) represented high-volume VMMC clinics, which ranked in the top 25% of the total number of circumcisions performed nationwide. The remaining 2 communities (Maun and Molepolole) represented low-volume VMMC clinics, which ranked in the bottom 25% of the total number of circumcisions performed nationwide.

Twenty-seven FGDs were conducted with 238 participants between July and November 2013. Participant characteristics are included in Table 1. Men and women aged 18–49 years were eligible to participate. An upper age limit was not applied to the community leaders. The age limit in this study coincides with that of the national VMMC program. Purposeful sampling of study participants was conducted according to age, and for men, circumcision status. As a result, FGDs were held with 2 age groups, 18–30 years and 31–49 years, of men (circumcised), men (uncircumcised), and women. Participants were recruited from places of public gathering (e.g. bus stations, shopping malls). Male circumcision status was assessed through self-report. The recruitment of community leaders was done through the government district bureaus and religious and education institutions. A letter describing the study and inviting leaders to participate was sent out to each of the bureaus and institutions in the
4 districts. Subsequently, follow-up calls or face-to-face meetings were held to confirm interest in participation and schedule the discussions. Focus group participants were provided with refreshments and travel costs.

The FGDs were conducted in Setswana, the local language, by trained interviewers who were the same gender as the participants. A semi-structured guide was used to facilitate the discussion on the modifications that could be made in the health system to maximize VMMC. Both open-ended and probing questions were used to facilitate the discussions, which were audio-recorded, transcribed, and translated into English. Written consent was obtained from all participants. Basic demographic data on each participant was captured and entered into a Research Electronic Data Capture database.

An inductive analytic approach was used to identify and describe the modifications that can be made in the health system to maximize VMMC uptake. Two investigators (KEW, JHL) identified concepts and key themes related to the topic and developed a coding scheme for major concepts. Using the coding scheme, 2 members of the research team (KEW, NR) independently coded a set of randomly selected transcripts to assess consistency between coders. Areas of inconsistency were discussed, noting why a particular code was assigned, and if appropriate, the operational definition was modified. The data set was coded using Atlas.ti v7.0 software by two authors (KEW and NR), discrepancies related to the assignment of codes were resolved by a third member of the research team (JHL). Thematic

Table 1 Descriptive characteristics of N=238 focus group discussion participants in four communities in Botswana

<table>
<thead>
<tr>
<th>Characteristics</th>
<th>Circumcised men</th>
<th>Uncircumcised men</th>
<th>Women</th>
<th>Community leaders*</th>
</tr>
</thead>
<tbody>
<tr>
<td>N (%)*</td>
<td>N (%)*</td>
<td>N (%)*</td>
<td>N (%)*</td>
<td>N (%)*</td>
</tr>
<tr>
<td>Location</td>
<td>N (%)</td>
<td>N (%)</td>
<td>N (%)</td>
<td>N (%)</td>
</tr>
<tr>
<td>Gaborone</td>
<td>18 (28%)</td>
<td>16 (26%)</td>
<td>27 (31%)</td>
<td>8 (31%)</td>
</tr>
<tr>
<td>Maun</td>
<td>13 (20%)</td>
<td>18 (30%)</td>
<td>20 (23%)</td>
<td>0 (0%)</td>
</tr>
<tr>
<td>Molepolole</td>
<td>17 (26%)</td>
<td>12 (20%)</td>
<td>20 (23%)</td>
<td>9 (35%)</td>
</tr>
<tr>
<td>Palapye</td>
<td>17 (26%)</td>
<td>15 (25%)</td>
<td>19 (22%)</td>
<td>9 (35%)</td>
</tr>
<tr>
<td>Age, years, median (IQR)</td>
<td>31 (24, 35)</td>
<td>30 (22, 34)</td>
<td>31 (25, 38)</td>
<td>4 (4%)</td>
</tr>
<tr>
<td>Current relationship status</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Never married</td>
<td>21 (32%)</td>
<td>20 (33%)</td>
<td>34 (40%)</td>
<td>3 (12%)</td>
</tr>
<tr>
<td>Dating and living together</td>
<td>17 (26%)</td>
<td>11 (18%)</td>
<td>12 (14%)</td>
<td>2 (8%)</td>
</tr>
<tr>
<td>Dating, but not living together</td>
<td>23 (35%)</td>
<td>27 (44%)</td>
<td>27 (31%)</td>
<td>2 (8%)</td>
</tr>
<tr>
<td>Married</td>
<td>3 (5%)</td>
<td>2 (3%)</td>
<td>9 (10%)</td>
<td>15 (58%)</td>
</tr>
<tr>
<td>Divorced, separated, or widowed</td>
<td>1 (2%)</td>
<td>1 (2%)</td>
<td>4 (5%)</td>
<td>4 (15%)</td>
</tr>
<tr>
<td>Religious affiliation</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Christianity</td>
<td>51 (78%)</td>
<td>49 (80%)</td>
<td>81 (94%)</td>
<td>26 (100%)</td>
</tr>
<tr>
<td>Badimo</td>
<td>4 (6%)</td>
<td>5 (8%)</td>
<td>0</td>
<td>0 (0%)</td>
</tr>
<tr>
<td>No religious affiliation</td>
<td>8 (12%)</td>
<td>5 (8%)</td>
<td>4 (5%)</td>
<td>0 (0%)</td>
</tr>
<tr>
<td>Other</td>
<td>2 (3%)</td>
<td>2 (3%)</td>
<td>1 (1%)</td>
<td>0 (0%)</td>
</tr>
<tr>
<td>Ethnic affiliation</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Bakgatla*</td>
<td>6 (9%)</td>
<td>3 (5%)</td>
<td>7 (8%)</td>
<td>1 (4%)</td>
</tr>
<tr>
<td>Bakwena</td>
<td>9 (14%)</td>
<td>15 (25%)</td>
<td>25 (29%)</td>
<td>7 (27%)</td>
</tr>
<tr>
<td>Bangwato</td>
<td>17 (26%)</td>
<td>16 (26%)</td>
<td>12 (14%)</td>
<td>8 (31%)</td>
</tr>
<tr>
<td>Bangwaketse</td>
<td>5 (8%)</td>
<td>1 (2%)</td>
<td>4 (5%)</td>
<td>3 (12%)</td>
</tr>
<tr>
<td>Batatawa</td>
<td>8 (12%)</td>
<td>7 (11%)</td>
<td>4 (5%)</td>
<td>0 (0%)</td>
</tr>
<tr>
<td>Bakalanga</td>
<td>7 (11%)</td>
<td>2 (3%)</td>
<td>12 (14%)</td>
<td>1 (4%)</td>
</tr>
<tr>
<td>Batswapong</td>
<td>1 (2%)</td>
<td>3 (5%)</td>
<td>6 (7%)</td>
<td>1 (4%)</td>
</tr>
<tr>
<td>Other</td>
<td>12 (18%)</td>
<td>14 (23%)</td>
<td>16 (19%)</td>
<td>5 (19%)</td>
</tr>
<tr>
<td>Employment status</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Employed and receiving a salary</td>
<td>21 (32%)</td>
<td>11 (18%)</td>
<td>14 (16%)</td>
<td>18 (69%)</td>
</tr>
<tr>
<td>Self-employed</td>
<td>18 (28%)</td>
<td>16 (26%)</td>
<td>29 (34%)</td>
<td>4 (15%)</td>
</tr>
<tr>
<td>Unemployed</td>
<td>17 (26%)</td>
<td>24 (39%)</td>
<td>31 (36%)</td>
<td>4 (15%)</td>
</tr>
<tr>
<td>Student</td>
<td>9 (14%)</td>
<td>10 (16%)</td>
<td>12 (14%)</td>
<td>0 (0%)</td>
</tr>
<tr>
<td>Occupation</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Professional or managerial</td>
<td>11 (28%)</td>
<td>3 (11%)</td>
<td>10 (23%)</td>
<td>17 (77%)</td>
</tr>
<tr>
<td>Skilled worker</td>
<td>21 (53%)</td>
<td>18 (67%)</td>
<td>7 (16%)</td>
<td>2 (9%)</td>
</tr>
<tr>
<td>Unskilled worker</td>
<td>8 (20%)</td>
<td>6 (22%)</td>
<td>25 (58%)</td>
<td>2 (9%)</td>
</tr>
<tr>
<td>Other</td>
<td>0 0</td>
<td>0 0</td>
<td>1 (2%)</td>
<td>1 (5%)</td>
</tr>
</tbody>
</table>

Notes: *A focus group discussion among community leaders in Maun was not conducted due to unresolvable scheduling constraints. Percentages may not sum to 100% due to rounding. Traditionally circumcising ethnic group. Data not collected.
categories, which were used to form larger concepts, were derived from the coded data. Operational definitions were established for the thematic categories and examples of data for each of the thematic categories were assembled.

**Results**

**Age- and gender-appropriate mobilizers and community leaders are more effective than mass media campaigns for creating demand**

Cultural norms around discussion of sex, respect for elders, and generational differences affect who is able to influence circumcision decisions, making peers of the target audience or community leaders the most effective mobilizers. One younger circumcised male in Gaborone stated:

> If I was to tell someone older than me to circumcise, say he is like 50 years old, he will say, ‘even my father did not circumcise. Young as you are, how can you tell me to circumcise? What do you know about life?’

Similarly, a community leader in Palapye stated, “Our culture here does not permit any person younger to discuss issues of sexuality with the older ones.”

Face-to-face interactions with peers were seen as being particularly effective at building demand for the procedure in order to encourage men to communicate freely on issues surrounding VMMC. An older, uncircumcised male in Gaborone commented:

> I am in support of re-introduction of initiation schools, even though they cannot be the same as the old ones. I feel there should be a time when men meet to talk about and discuss issues affecting men.

In addition to peer mobilizers, participants frequently mentioned the potential role of community leaders. A young circumcised male in Gaborone stated: “People believe a lot in community leaders, chiefs, ministers, when they speak people feel that someone important has spoken.” Another young circumcised male in Molepolole stated that:

> I feel they are not doing enough […] our leaders, especially traditional leaders are dragging their feet and though the government is playing its role, I still feel there is a gap that needs to be closed.

In contrast to peers and community leaders, participants generally felt media campaigns were less effective in demand creation, given one-way communication and less immediacy. One community leader in Palapye stated there was a serious problem of knowledge and education inadequacy. I can’t remember at any time when a seminar, workshop, or anything of the like was held to sensitize men about any issues affecting them, let alone [VMMC]. They only hear about these things on the radio […] hearing from the radio is far different than hearing first hand, from an expert, sitting with them, face to face as that accords one the platform to ask for clarity.

One female in Gaborone shared:

> There is a billboard with a goalkeeper. It took me a while to get what it is all about. Every time I will pass by it I will see the goalkeeper and promise myself I will read it next time but I never get to read it.

Mobile entertainment trucks used to advertise circumcision services were also viewed as having limited utility, with a respondent in Gaborone saying they: “Just entertain people, the moment they leave, people forget about them.”

**Maximizing access to VMMC services through reduction of logistic hurdles**

Another barrier to circumcision uptake was limited access due to logistic concerns, including distance to facilities, wait times, and fees. One older uncircumcised male in Gaborone said:

> People are scared to travel long distances to be circumcised. Men keep wondering how they will cope [with] it for example, they take public transport, go circumcise and travel back in public transport.

Another older uncircumcised man in Gaborone shared: “Sometimes you want to avoid long queues at the government hospital but again you cannot afford the exorbitant fees charged at the private hospitals.” Multiple respondents advised that it was better to bring circumcision services closer to people, as currently they are only offered at select health facilities. Participants recommended either using mobile trucks or: “Maybe, if possible, every clinic should offer circumcision so that those without money and really want to do [get circumcised] can access services easily.”

**Providing VMMC services at designated men’s clinics and allowing men to select gender of service provider**

Participants discussed many other barriers to circumcision, including typical health care-seeking behaviors among men and feelings about gender, age, and professionalism of providers. Participants also discussed the health care-seeking
behavior of men, indicating they rarely access services at hospitals. A community leader in Gaborone mentioned:

If you see a man going to the hospital, we call him a sissy because we believe that the hospital is for women. We need to instill into men that the hospital is for everybody. If we realize that the hospital is where we get treated and get better health, we can be able to utilize circumcision services.

Several men advised that to increase circumcision uptake, preferences about the age and gender of the provider must be considered, recommending seeking client preferences in settings where providers of both genders are present. One older circumcised male in Gaborone stated:

If the [VMMC] services were offered by male practitioners only, men would come in large numbers. It would work without women. Where there are women, men are reluctant to be engaged.

Another young, uncircumcised male in Maun stated,

They never ask whether you want to be circumcised by a woman or man, they will send you to the theater then you will see women coming there. Some of us are sensitive. It’s like you are being insulted or something because this woman may tell her friend that she has seen your manhood.

A young, uncircumcised male in Maun unequivocally added that: “Being circumcised by a woman is not right.” Aside from gender, age was considered an important trait among providers. Older circumcision providers, regardless of gender, were preferred to young providers because they were perceived as being better at upholding confidentiality.

Respondents also stated that they felt providers could act uncaring, rushed, and unconcerned about their comfort. A positive attitude among circumcision providers is seen as encouraging. A young circumcised male in Palapye relayed his impressions of VMMC providers:

I also feel that sometimes the people who are performing those surgeries are overwhelmed by their work such that the surgery room is not a friendly place, because you go there and sometimes you are met by someone who just wants to make sure the queue finishes such that even when they touch you it’s not in a loving way and I feel that the surgeons have a role in making men not go to circumcise in large numbers. It would be nice to find smiling people there not the ones who just want to remove your skin and let you go.

Providing comprehensive VMMC pre-procedure counseling and education, including describing differences between surgical and nonsurgical options

Participants suggested pre-procedure counseling could benefit from a more objective review of procedure advantages, disadvantage, and options. Clients going for VMMC services currently receive counseling, but participants felt the sessions focused on procedure benefits. One young circumcised male in Gaborone stated that:

[...] it’s like they are more concerned about advantages and try to hide disadvantages. They do not want people to know what complications of circumcision [exist], only advantages. I feel if this can be explained to people, they will be in a better position to choose what to do.

Additionally, a community leader in Palapye stated:

[...] one of the things I have realized is that all the discussions about [VMMC] centers on its advantages, they hardly touch on the procedure itself [...] whether it is painless or painful. I suggest that nothing be left out, let us be truthful when talking about VMMC.

Male participants had mixed views about the newly rolled-out non-surgical circumcision devices. Many who feared the complications of surgical circumcision were positive toward non-surgical devices, but an older circumcised male in Maun noted: “[…] one of the things I have realized is that all the discussions about [VMMC] centers on its advantages, they hardly touch on the procedure itself […] whether it is painless or painful. I suggest that nothing be left out, let us be truthful when talking about VMMC.

Discussion

This study has highlighted areas of the health system that can be modified to maximize uptake of circumcision services. In relation to demand creation, participants felt that using age- and gender-appropriate mobilizers and community leaders was a more effective strategy than mass media campaigns. Participants felt that improved access to VMMC clinics would facilitate service uptake, as would designating men’s clinics for VMMC service delivery. Providing comprehensive pre-procedure counseling and education, outlining the benefits and disadvantages of the surgical procedure, and describing the differences between both the surgical and non-surgical procedures were suggested. These health systems modifications to demand creation, access to services, and service delivery hold promise for increasing uptake of VMMC.

Similar findings have been reported in Botswana related to engaging age- and gender-appropriate community mobilizers. Additionally, studies have found that cultural acceptability can
be improved through the involvement of community and political leaders. Influential leaders can play an important role in convincing men that medical circumcision does not conflict with culture. The role of community leaders and influential persons in sensitizing their communities in health care programs has been successfully executed in other resource-limited settings. Kenya, which has had a successful VMMC program, has benefited from the support of politicians. Several members of parliament and cabinet ministers publicly shared that they were circumcised in order to promote VMMC. In Malawi, leaders of traditionally non-circumcising communities were influential in creating demand for VMMC and encouraging males to undergo circumcision. Based on these findings, VMMC uptake in Botswana can be maximized by support from government and community leadership and using age-and gender-appropriate community mobilizers.

Data from this study indicate that the effectiveness of media campaigns in creating demand varies and the type of campaign should be given ample consideration. Some participants questioned the effectiveness of these activities, highlighting the need to actively monitor efficacy as activities are initiated. There are multiple examples from the literature of highly successful VMMC campaigns. For example, a multimedia campaign in Uganda paired with community mobilization motivated 89% of exposed men to participate in at least 1 health promotion activity such as HIV testing as a result of the campaign, and over 10% of these men reported getting circumcised. In addition, a 30-day national mass-media campaign in Malawi was responsible for an increase in circumcision services. Botswana could apply some of the lessons learned from successful campaigns to develop a strategy to maximize VMMC impact.

This study yielded important findings related to service delivery and the need for “men’s clinics” to support VMMC as well as general men’s health services. Poor health-seeking behavior by men has been well documented, particularly for preventative health services. Men have historically had lower HIV testing rates across sub-Saharan Africa; data from Botswana show they often comprise less than one-third of all testers. In the present study, there was recognition among participants that a discussion of men’s health issues was lacking. There is a clear need for future work to explore the challenges African men face in seeking health care services. A recent study from rural Zimbabwe on the role of masculinity in African men’s use of HIV services found that uptake of services was related to dominant masculinities, socio-cultural influences, and global expectations of HIV-infected patients. The present study, as well as 2 others in the region highlight that male health care providers were viewed as most desirable for delivery of VMMC services. Globally, patient preferences about their health care provider’s gender has been observed broadly in non-HIV-related fields, including urology, obstetrics and gynecology, and gastroenterology. This suggests male health care workers could be preferentially assigned to support these initiatives. This study also highlights the need for staff conducting VMMC counseling to have good knowledge about the pros and cons of the procedure and to ensure this information is communicated to the clients for informed decision-making. Future interventions could develop male-centered clinics, use male providers, and train health care staff to present the full range of information on circumcision to better address the needs of the population in this setting.

While this article has important implications for work with men in Botswana and elsewhere, it has limitations. Findings from this study were from 4 of 28 districts of Botswana and may not be generalizable to the whole country. Additionally, circumcision status of male participants was assessed by self-report. Given that men may misreport their circumcision status, the ensuing discussion and corresponding findings could have been influenced by men assigned to an inappropriate focus group. Furthermore, among the circumcised men, reasons for undergoing circumcision were not assessed. Some participants may have been circumcised as infants or as a result of a medical complication (i.e., phimosis). These participants’ perceptions of systems-level factors about VMMC likely differ from those circumcised during the national VMMC campaign. Additionally, participants may have been inhibited to speak freely due to the traditional cultural and social norms related to discussing issues related to sexual health. Skilled, same gender moderators were used to mitigate this potential limitation.

Conclusion
The uptake of VMMC programs is gradually improving in Botswana and in several priority countries in sub-Saharan Africa. This study has highlighted modifications related to the health system in the areas of demand creation and service delivery that can be employed to optimize VMMC uptake, including using age- and gender-appropriate mobilizers and community leaders for demand creation, improving access to VMMC services, particularly through men’s clinics, and providing comprehensive pre-procedure counseling and education that outline the benefits and disadvantages of the surgical and non-surgical procedures.

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Disclosure
The authors report no conflicts of interest in this work.

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