

ORIGINAL ARTICLE

## Self-medication practice and associated factors among adults in Wolaita Soddo town, Southern Ethiopia

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### Abstract

**Background:** Self-medication is the selection and use of medicines by individuals to treat their self-recognized illnesses or symptoms. Self-medication can decrease costs and enable health professionals to concentrate on more serious health problems.

**Aim:** To assess self-medication practice and associated factors among adults in Wolaita Soddo town, Southern Ethiopia, 2017.

**Methods:** An institution-based cross-sectional study was conducted from September 30 to October 30, 2017. A multi-stage sampling technique of drug retail outlets in Wolaita Soddo town was employed to identify 623 individuals that came to buy drugs in the past three months. Data was collected using a structured questionnaire.

**Results:** About 33.7% of the respondents had practiced self-medication in the past 3 months. Multivariate analysis revealed that female sex (adjusted odds ratio (AOR) = 2.22, 95% confidence interval (CI): 1.47–3.36), low income (AOR = 3.95, 95% CI: 2.32–6.73) and higher educational level (AOR = 5.79, 95% CI: 2.47–13.58) were the independent factors significantly affecting the practice of self-medication with drugs. Headache/fever (32.4%), respiratory tract infections (31.4%) and gastrointestinal diseases (16.2%) were the most frequently reported illnesses or symptoms of illnesses that prompted self-medication of study participants.

**Conclusion:** Health education campaigns, strict legislations on dispensing drugs from private pharmacies, and improving accessibility and affordability of health care are among the important interventions required to change people's health-seeking behavior and prevent the potential risks of self-medication.

**Keywords:** *Self-medication; drug retail outlets; over-the-counter; prescription; Ethiopia.*

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The World Health Organization (WHO) defines self-medication as the selection and use of medicines by individuals to treat self-recognized illnesses or symptoms (1). It is an element of self-care in which people utilize traditional or modern medicine without consulting a physician, either for diagnosis, prescription or surveillance of treatment. Self-medication comprises obtaining medicines without authorized prescription, resubmitting old prescriptions to buy medicines, using medicines in share with relatives or social life members, or utilizing leftover medicines stored at home (2).

Self medication is practiced commonly in the world in both developed and developing countries, and its utilization may be even more than prescribed medication (3). The prevalence of self-medication practice (SMP) was found to be: Pakistan 85% (4, 5), Saudi Arabia 35.4% (3),

Iran 53% (6), Yemeni 85% (7), India 92.8% (8); Brazil 16.1%, (9) Latin American immigrants in Seville, Spain 77.4% (10), Italy 69.2% (11), Greece 54.7% (12) and Spain 20.2% (13); Nigeria 95% – 98%, Southwestern Nigeria, 81.8% (16), and Kenya 76.9% (17).

A study conducted in Addis Ababa showed that the magnitude of SMP was 50% (18), with other studies showing the following prevalence of SMP in Ethiopia: Kolladiba town, Northwest Ethiopia 62.8% (19), Assendabo town, Southwestern Ethiopia 39.2% (20), Sire town, West Ethiopia 27.16%,<sup>2</sup> and the Silte zone, South Ethiopia 24.4% (21). Another study conducted at Gondar University showed that 38.5% of the participants practiced self-medication (22).

According to a study conducted in South Karnataka, India, the most common symptoms prompting self-medication were: gastric symptoms (72.10%), joint pain

(65.89%), headache (63.02%), fever (47.87%), and common cold (37.95%) (23). The study also showed that the main groups of diseases in which patients self-medicated were respiratory diseases (39.9%), common cold (39.5%) and headache (37.3%) (6).

In a study conducted in Mekelle town to assess SMP, the most frequent illnesses or symptoms of illnesses that prompted study participants to self-medicate were headache/fever (20.7%), gastrointestinal diseases (17.3%) and respiratory tract infections (15.9%) (24). In Addis Ababa, commonly reported illnesses or symptoms of illnesses for which respondents self-medicated were gastrointestinal (25.1%), headache/fever (24.9%) and respiratory problems (21.4%) (18).

In a study conducted on self-medication practices in Italy, non-steroidal anti-inflammatory drugs (NSAIDs) (83.5%) were most commonly used, whereas antibiotics (26.7%), antacids (4.2%), and corticosteroids (3.4%) have been used less frequently (11). Another study on SMP among Iraqi patients in Baghdad city showed that antibiotics (74.7%) were the most frequently requested drugs followed by paracetamol (55.1%), antihistamines (36.2%), and NSAIDs (27%) (25).

A study conducted on self-medication in Alexandria, Egypt, showed that analgesics (96.7%), cough and common cold preparations (81.9%), vitamins and minerals (63.2%), antibiotics (53.9%), gastrointestinal drugs (51.4%), and antihypertensives (16.1%) were the most commonly used drugs in self-medication by participants (26).

In a study conducted on SMP in Addis Ababa, the most common types of drugs were analgesics/antipyretics (33.1%), antimicrobials (26.4%) and gastrointestinal drugs (17.7%) (18). A study on SMP in Jimma town, Ethiopia, indicated that the most common drugs used for self-medication were analgesics/antipyretics (28.94%), antimicrobials (28.13%) and antihelmintics (17.56%) (27).

A cross-sectional study conducted on prevalence and pattern of SMP in an urban area of Delhi, India showed that the major sources of information for self-medication were past experience (46.5%) followed by a physician's old prescription (43.8%), recommendation by chemists (35.6%) and family members' opinion (35.6%) (8). Similarly in Alexandria, Egypt, the most common source of advice or information about self-medication with drugs was an old prescription (73.9%), pharmacists (42.7%), and an older person in the household or friend (30.5%) (26).

A study conducted in Addis Ababa showed that 39% of drug consumers received advice from health professionals like physicians, nurses and health assistants but without formal prescriptions. About 23.5% of participants were advised by friends, relatives or neighbors, who themselves had no background in the health profession. Pharmacists or other personnel working in pharmacies were found to be sources

of advice for 15.4% of respondents, whereas 20% of respondents received no advice but had information on the drugs from labels, leaflets or promotional materials (18).

Studies conducted in different countries have shown that age, sex, poor health status, inconvenient access or services (3), along with education (9), place of residence, marital status, occupation, old age group (6), and presence of chronic diseases (26), were factors for self-medication. Convenience of pharmacy, friend/peer pressure and advertisements (28), unavailability of physicians, prior experience treating the same illness and requirement of emergency care were other factors for self-medication (29).

In a study on self-medication practice in Mekelle, Ethiopia, the main reasons for self-medication were belief that the illness is mild (21.7%), previous experience of illness/ medicine (20.7%), inexpensive and time-saving (20.2%), belief that it was an emergency situation (17%), and prevention of known or unknown illness (16.9%) (24).

Studies indicate that self-medication is the first response to illness among people with low income (30). A study conducted in Addis Ababa indicated that the most common reasons for self-medication were non-seriousness of illness (36.6%), emergency situations (19.8%), prior experience (18.2%), and less expensive in terms of time and money (12.6%) (18).

## Methods

### *Study setting and population*

This study was conducted in Wolaita Soddo town, 327 km from the capital city of Addis Ababa. Wolaita Soddo town has an estimated population of 140,105 number. It has 30 drug retail outlets (DROs), two hospitals, three health centers, 11 health posts and 21 private health institutions that provide health services to the community.

### **Study design**

An institution-based cross-sectional study was conducted in nine DROs from September 30 to October 30, 2017 in the Wolaita zone of Soddo town.

### *Study population*

All adult people >18 years of age who attend DROs to buy drugs.

### *Inclusion criteria*

Adults (age >18 years) who buy drugs with or without a prescription from the selected DROs in Wolaita Soddo town during the study period.

### *Exclusion criteria*

People who were not able or willing to sign informed consent.

### Sample size determination

The sample size was calculated using single population proportion formula.

$$n = \frac{Z_{1-\alpha/2}^2 (p)(1-p)}{d^2}$$

where  $n$ : sample size;  $p$ : estimated prevalence of self-medication;  $Z$ : percentile of standard normal distribution corresponding to 95% confidence level;  $d$ : the margin of error which is 0.05. By using the anticipated population proportion of 24.4% from a previous study (21), the sample size calculated as

$$n = \frac{(1.96)^2 0.244(1-0.244)}{(0.05)^2} = 283$$

Hence, the design effect of the study was 2 and the sample become 566. The total sample size when 10% non-response was added became 623.

### Sampling procedure

A multistage sampling technique was conducted to select study participants. All types of drug retail outlet such as pharmacies, drug stores and drug vendors were used. About 9 drug retail outlets were selected randomly from the total drug retail outlet and then individuals were selected systematically.

### Data collection and Analysis procedure

A face to face interview was conducted with individuals coming to selected pharmacies to buy drugs during the data collection period. After gaining informed consent from the participant, background information and possible factors were collected.

### Data collection tool

A pre-structured questionnaire was developed and translated into the local language, Amharic. To ensure consistency, it was then retranslated to English.

### Data Quality control

Four health professionals who had previous experience on data collection were selected and trained for one day. The questionnaire was pre-tested on 5% of the actual sample size before data collection. The data was checked for completeness and consistency. The Hosmer and Lemeshow test for Goodness-of-fit of the model was done ( $p = 0.698$ ).

### Data processing and analysis

Data was coded and entered into the computer with EPI-data version 3.1 and exported to SPSS version 20 for

Table 1. Socio-demographic characteristics of respondents, Wolaita Soddo town, 2017 ( $n = 623$ )

Variables	Frequency (N)	Percentage (%)
<b>Sex</b>		
Male	319	51.2
Female	304	48.8
<b>Age</b>		
18–24	159	25.5
25–34	243	39
35–44	143	23
>44	78	12.5
<b>Religion</b>		
Protestant	351	56.3
Orthodox	205	32.9
Muslim	56	9
Others <sup>a</sup>	11	1.8
<b>Ethnicity</b>		
Wolaita	404	64.9
Amhara	66	10.6
Gurage	52	8.3
Oromo	41	6.6
Gamo	33	5.3
Others <sup>b</sup>	27	4.3
<b>Educational status</b>		
No formal education	45	7.2
Primary education	66	10.6
Secondary education	145	23.3
Higher education	367	58.9
<b>Marital status</b>		
Married	328	52.6
Not married	242	38.8
Widowed	19	3.1
Divorced	34	5.5
<b>Income (in Ethiopian Birr)<sup>d</sup></b>		
0–500	141	22.6
501–1000	98	15.7
1001–1500	55	8.8
1501–2000	55	8.8
2001–3000	52	8.4
≥3001	222	35.7

<sup>a</sup>Catholic and pagan; <sup>b</sup>Hadya, Kambata, and Sidama; <sup>c</sup>private employee and daily laborer; <sup>d</sup>1 USD = 27 ETB.

analysis. Descriptive statistics such as frequencies, percentage and tables were used to present the respondent's result. Independent variables with  $p < 0.25$  on bivariate analysis became a candidate for multivariable analysis. Univariate and multivariate logistic regression was fitted to identify associated factors with 95% CI using  $p$ -value  $< 0.05$ .

## Results

### Socio-demographic characteristics

Out of the 623 individuals who participated in this study, 319 (51.2%) were males, 243 (39%) were aged 25–34 years, 404 (64.9%) were Wolaita ethnicity group, 367 (58.9%) had graduated from college or University, 351 (56.3%)

were Protestant in their religion and 328 (52.6%) were married (Table 1).

**Self-medication practice**

In this study, the prevalence of SMP was 210 (33.7%) (95% CI: 37.2, 44.9). The most frequent illnesses reported by respondents that prompted them to self-medicate were headache 68 (32.4%), cough/common cold 66 (31.4%), gastrointestinal disease 34 (16.2%) and malaria 32(15.2%) (Table 2).

**Source of advice and way of requesting drugs**

The most frequent source of advice for self-medication in the present study were pharmacists 117 (55.7%), other healthcare providers such as nurses and health assistants 41 (19.5%) and friends 34 (16.2%). Drug consumers were also asked how they requested the drugs they used. About 101 (48.1%) of drug consumers made their requests by relating the symptoms of their illness to a pharmacy professional, 82 (39%) of them by telling the specific name of drug, whereas 22 (10.5%) mentioned the class to which the drug belongs. Nevertheless, 2.4% of respondents requested the drug by showing an old sample of the drug or drug products (Table 3).

**Reasons for self-medication**

About 165 (26.5%) of the study participants believed that the disease was mild. However, 164 (26.3%) of the participants opinion was inexpensiveness and 95 (15.2%) to save their time (Table 4).

The most frequent types of drugs used for self-medication in this study were analgesics/antipyretics 118 (56.2%), antibiotics 84 (40%), and antimalarials 34 (16.2%) (Table 5).

**Factors associated with self-medication practice**

The multivariable logistic regression model showed that sex, educational status and income were statistically associated with self-medication practice. Female respondents were about twice more likely to practice self-medication than males (AOR=2.22, 95% CI: 1.47–3.36) ( $p < 0.05$ ). Respondents with higher educational level were 5.8 times more likely to practice self-medication compared to those having no formal education (AOR = 5.79, 95% CI: 2.47–13.58) ( $p < 0.05$ ). Study participants with low monthly income were found to be 4 times more likely to practice self-medication compared to those with higher income levels (AOR=3.95, 95% CI = 2.32–6.73)( $p < 0.05$ ) (Table 6).

**Discussion**

Self-medication has an important role in the care of minor illness. It can decrease cost and enable health professionals to concentrate on more serious health problems. (3, 31) On

*Table 2.* Types of illness or symptoms of illnesses that necessitated self-medication

Illness/symptom	Frequency (N)	Percentage (%)#
Fever	23	11
Headache	68	32.4
Diarrhea	13	6.2
Cough/common cold	66	31.4
Gastrointestinal disorder	34	16.2
Menstrual problem	12	5.7
Back pain	18	8.6
Eye infection	10	4.8
Skin infection	13	6.2
Malaria	32	15.2
Peptic ulcer disease	15	7.1
Others(typhoid and acne)	08	3.8

#Numbers do not add to 100% as patients might have more than one answer.

*Table 3.* Source of advice and way of requesting drugs for self-medication

Variable	Frequency (N)	Percentage (%)
<b>Sources of advice</b>		
Pharmacists	117	55.7
Other health professionals	41	19.5
Friend	34	16.2
Neighbour	10	4.8
Others	08	3.8
<b>Way of requesting drug</b>		
Mentioning name of the drug	82	39
Mentioning class of the drug	22	10.5
Telling symptoms of illness	101	48.1
Showing old samples of the drug	05	2.4

*Table 4.* Reasons of participants for self-medication practice, Wolaita Soddo Town, Sothern Ethiopia, 2017 (n = 623)

Variable	Frequency (N)	Percentage (%)
Emergency use (quick relief)	46	7.4
Mild illness	165	26.5
Inexpensiveness	164	26.3
To save time	95	15.2
Previous experience	36	5.8
Advertisement	25	4
Family/peer pressure	41	6.6
Easy access to drugs	65	10.4
Dissatisfaction with health service	54	8.7
High cost of consulting physician	43	6.9
Lack of health facility in nearby	57	9.1
Lack of trained health professionals	59	9.5

Table 5. Types of drugs used for self-medication

Drug type	Frequency (N)	Percentage (%)#
Analgesics/antipyretics	118	56.2
Antibiotics	84	40
Drugs for respiratory disease	03	1.4
Antihelminthes	14	6.7
Drugs for PUD	15	7.1
Antimalarials	34	16.2
Drugs for eye disease	10	4.8
Drugs for skin disease	13	6.2
Others (antiemetic and laxatives)	09	4.3

# Numbers do not add to 100% as patients might have used more than one drug type.

the other hand, self-medication is a potential contributor to human pathogen resistance to antibiotics. Its potential risks include incorrect self-diagnosis, delay in medical treatment, incorrect treatment choice, severe adverse reactions, incorrect dosage, and drug dependence and abuse (32).

In the present study the prevalence of self-medication was 33.7% (95% CI: 37.2, 42.2). This finding is in line with other studies carried out in different parts of the world such as Saudi Arabia (35.4%) (3) and South India (35.9%) (33). This result is also similar to studies carried out in different parts of Ethiopia such as Assendabo, Southwest Ethiopia (39.2%) (20) and Nekemte (36.7%) (34, 19). On the other hand, the prevalence of SMP in this study is lower than those conducted in India (92.8%),<sup>8</sup> Sudan

Table 6. Logistic regression analyses of the factors associated with practicing self-medication practice among participants, Wolaita Soddo town, 2017 (n = 623)

Respondent's characteristics	Self-medication		COR (95% CI)	AOR (95% CI)
	Yes (%)	No (%)		
<b>Sex</b>				
Male	81 (38.6%)	238 (57.6%)	1	1
Female	129 (61.4%)	175 (42.4%)	<b>2.166 (1.542–3.042)***</b>	<b>2.223 (1.472–3.355)***</b>
<b>Age</b>				
18–24	51 (24.3%)	108 (26.2%)	1	1
25–34	91 (43.3%)	152 (36.8%)	1.268 (0.831–1.934)	1.53 (0.88–2.659)
35–44	49 (23.3%)	94 (22.8%)	1.104 (0.683–1.783)	1.748 (0.835–3.658)
>44	19 (9%)	59 (14.3%)	<b>0.682 (0.369–1.261)*</b>	1.235 (0.517–2.95)
<b>Ethnicity</b>				
Wolaita	147 (70%)	257 (62.2%)	1.411 (0.749–2.658)	1.123 (0.531–2.376)
Amhara	13 (6.2%)	53 (12.8%)	<b>0.605 (0.258–1.42)*</b>	0.452 (0.168–1.22)
Oromo	13 (6.2%)	28 (6.8%)	1.145 (0.47–2.79)	1.194 (0.424–3.361)
Gamo	12 (5.7%)	21 (5.1%)	1.41 (0.557–3.569)	1.525 (0.507–4.588)
Gurage	10 (4.8%)	17 (4.1%)	1.451 (0.542–3.885)	1.491 (0.442–5.032)
Others	15 (7.1%)	37 (9%)	1	1
<b>Educational status</b>				
No formal education	9 (4.3%)	36 (8.7%)	1	1
Primary education	2 (1%)	64 (15.5%)	<b>0.125 (0.026–0.610)**</b>	0.171 (0.033–0.882)
Secondary education	19 (9%)	126 (30.5%)	0.603 (0.251–1.447)	0.859 (0.325–2.269)
Higher education	180 (85.7%)	187 (45.3%)	<b>3.85 (1.803–8.221)***</b>	<b>5.786 (2.465–13.581)***</b>
<b>Income</b>				
0–500	76 (36.2%)	65 (15.7%)	<b>3.157 (2.024–4.923)***</b>	<b>3.949 (2.316–6.734)***</b>
501–1000	39 (18.6%)	59 (14.3%)	<b>1.785 (1.081–2.947)**</b>	<b>1.87 (1.046–3.34)**</b>
1001–1500	12 (5.7%)	43 (10.4%)	0.753 (0.372–1.525)	1.038 (0.464–2.325)
1501–2000	14 (6.7%)	41 (9.9%)	0.922 (0.469–1.811)	1.035 (0.481–2.229)
2001–3000	9 (4.3%)	43 (10.4%)	<b>0.565 (0.26–1.229)*</b>	0.48 (0.207–1.116)
≥3001	60 (28.6%)	162 (39.2%)	1	1
<b>Marital status</b>				
Married	107 (51%)	221 (53.5%)	1	1
Not married	89 (42.4%)	153 (37%)	1.201 (0.848–1.703)	1.16 (0.69–1.95)
Widowed	8 (3.8%)	11 (2.7%)	1.502 (0.587–3.844)	1.82 (0.606–5.469)
Divorced	6 (2.9%)	28 (6.8%)	<b>0.443 (0.178–1.101)*</b>	0.576 (0.202–1.646)

\* $P < 0.25$ ; \*\* $P < 0.05$ ; \*\*\* $P < 0.001$ .

COR, crude odds ratio; 95% CI, 95% confidence interval; AOR, adjusted odds ratio.



(81.8%) (16), Kolladiba (62.8%) (19), Pakistan (85%) (4), Karachi, Pakistan (84.8%) (5), Iran (53%) (6), Yemeni (85%) (7), Latin American immigrants (77.4%) (10) Nigeria (91.4%) (15) and Kenya (76.9%) (17). However, the current finding is greater than those of studies conducted in Brazil (16.1%) (9), Bahir Dar (23.3%) (35), Sire town (27.16%) (2) and Silte (24.4%) (21). This might be due to sociodemographic factors such as educational status, socio-economic status and living status of the study participants. Moreover, the prevalence of self-medication could not be compared across different studies due to the varying nature of definitions used, recall period considered, region selected and methodology adopted, the role of country culture, health-care systems and the perceived role of health professionals.

Data from the present study showed that sex, income and educational status had significant association with self-medication practice. Female respondents were more likely to practice self-medication than males. This finding is consistent with studies conducted in Brazil (9), Iran (36), Italy (11), Greece (12), Serbia (37), Arsi University (38), Kolladiba (19). The fact that females have significant social roles, are more strongly self-care oriented than male and have more active involvement in household activities making them tend not to seek other medical service are other reasons for the increased practice of self-medication among females (12). However, contradicting literature showing that men self-medicate more than women were found in South Karnataka (23), and Saudi Arabia (3).

This study showed that respondents with low socio-economic status were more likely to practice self-medication compared to those with higher income, which is in line with studies conducted in Kharthoum (16) and Kolladiba (19). In the majority of economically deprived countries, nearly 60–80% of health related problems are treated through self-medication as a lower cost alternative. As WHO indicated, self-medication provides a cheap alternative to people who cannot afford to pay healthcare costs, and thus, self-medication is often the first response to illness among people with low income (20, 39). Most people prefer self-medication than going to health facilities because of high costs of laboratory diagnosis and other services.

In this study, participants with higher educational status practice self-medication higher than those with no formal education, which is consistent with previous studies conducted in Iran (36), Spain (13), Greece (12), Rajasthan (40), and India (8). Respondents with college and university educational level have more knowledge about common diseases and drugs for self-treatment than those who are less educated. Moreover, people with higher educational status have more self-confidence in self-diagnosis and self-medication, especially where there is acute illness and as a result they are capable of identifying appropriate drugs to treat their symptoms, whereas less educated people prefer a medical consultation (12).

Headache was the primary symptom for practicing self-medication in the present study, which is supported by studies conducted in Karachi (5), Pakistan (4), Mekelle (24), Kolladiba (19), Silte (21). However, this finding contradicts results from a study conducted in Addis Ababa (18), in which gastrointestinal disease was the most frequently reported illness for self-medication. In this study, cough/common cold, gastrointestinal disease and malaria were the most frequently reported illnesses following headache that necessitated self-medication, consistent with findings in Kolladiba, Ethiopia (19)

In the present study, analgesics/antipyretics were the most frequently utilized drug classes as indicated in other studies done in Brazil (9), India (41), Kolladiba (19), Mekelle (24), Jimma (27) and Southwest Ethiopia (42). This is because these drugs are used to treat mild common symptoms such as headache, fever and pain. It confirms that analgesics/antipyretics are the most commonly utilized over the counter drugs for self-medication. Similarly, antibiotics and antimalarials were frequently utilized drug classes next to analgesics in the present study, which is in line with studies conducted in Iraq (25), Jimma (27), India (41), Kolladiba (19), Southwest Ethiopia (42), Arsi University (38) and Addis Ababa (18).

In the present study, mildness of disease, inexpensiveness of self-medication, and saving time were pointed out as major reasons for self-medication, supported by studies conducted in Mekelle (24), Addis Ababa (18), Rajasthan (40), Sudan (16) and Sire town (2).

## Conclusion

According to our study, self-medication is practiced for both over-the-counter and prescription only drugs. Respondents who were female, having low socio-economic status, divorced and higher educated were more likely to practice self-medication. Headache/fever, RTIs and GITs were the most frequently reported symptoms of illnesses that prompted self-medication. The major reasons for SMP were mildness of disease, inexpensiveness of self-medication and saving time. Analgesics/antipyretics, Antibiotics, and Antimalarials were the most frequently requested drug classes. Health professionals, especially community pharmacists, need to educate people on the benefits and risks of self-medication to encourage responsible self-medication. Future research to assess the practice of Self-medication should be conducted.

## Declaration

### *Ethics approval and consent to participate*

An ethical approval was obtained from the Ethical Review Committee of Wolaita Soddo University, College of Health Sciences, Research and Community Service Office. Official support letter was obtained from Wolaita Soddo University and Wolaita zone health department for

undertaking the study. An informed consent was obtained from each of the study participants. Each participant took part in the study after verbal consent was approved.

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### Author's contribution

TM: conceived and designed the study, performed the data collection, analyzed the data, KD: involved in protocol development, data analysis and manuscript write-up; SB: conceived and designed the study, data collection; DD: involved in data analysis and manuscript write up. All authors have read and approved the manuscript.

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### Availability of data

The raw data should not be provided in order to protect patient's anonymity. Summary data are available in the main documents.

### Consent for publication

Not applicable.

### Conflicts of interests

The authors declare that they have no competing interests.

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