PLOS Neglected Tropical Diseases Determinantes of Podoconiosis among Residents in Machakle District East Gojjam Zone Amhara Region Ethiopia --Manuscript Draft--

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Short Title:	Determinantes of Podoconiosis
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Keywords:	Determinants; Machakel; Podoconiosis; Neglected Tropical diseases; Field Epidemiology; Ethiopia
Abstract:	Background
	Podoconiosis (endemic non-filarial elephantiasis) is a chronic disease characterized by the development of persistent swelling of plantar foot initially; which progresses to the dorsal foot and lower leg slowly or in a number of acute episodes to reach the knee. About 4 million people are said to be affected by the disease worldwide and it is deemed a serious public health problem in at least 10 African countries including Ethiopia. Therefore this study aimed to identify the determinants of Podoconiosis among residence in Machakel district.
	Method
	Unmatched case control study design was conducted at Machakel district from August 30 to September 30/-2022. The sample size was calculated using Epi-info software yielded 211 controls and 106 cases (317 study participants). Simple random sampling technique was used to select the cases using registration books of the district. Data were entered to Epi info version 7 and exported to SPSS version 22 for statistical analysis. Binary logistic regression was used to identify explanatory variables.
	Result
	A total of 312 study participants (104 cases and 208 controls) were included giving for a response rate of 98.42%. Bare foot (AOR, 5.83 [95% CI: 2.34-14.50]), female sex (AOR, 4.25 [95% CI: 2.22-8.14]), family history of podoconiosis (AOR 3.01(95% CI: 1.41-6.42) and age group 41-60 (AOR 5.05(95% CI: 2.35-10.83), and 61-80 AOR 15.74 95% CI: (5.56-44.55) were determinants of Podoconiosis.
	Conclusion and recommendation
	Barefoot, sex, family history of podoconiosis and age group were determinants of Podoconiosis. District health office should encourage for at risk populations especially for older age and family history of podoconiosis about shoe wearing practice in all their lives and do not exposed their skin and feet.
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Determinantes of Podoconiosis among Residents in Machakle District 1 East Gojjam Zone Amhara Region Ethiopia 2 Teshome Tefera¹, Kassawmar Angaw Bogale¹, Yiteka Tegegn¹, Abebaw Gedef Azene¹, 3 Kebadnew Mulatu¹, Gizachew Tadesse Wassie^{*1} 4 5 ¹Department of Epidemiology and Biostatistics, School of Public Health, College of Medicine 6 and Health Sciences, Bahir Dar University, Bahir Dar, Ethiopia 7 8 * Corresponding author 9 10 Po Box: 79 11 12 13

14 Abstract

Background: Podoconiosis (endemic non-filarial elephantiasis) is a chronic disease characterized by the development of persistent swelling of plantar foot initially; which progresses to the dorsal foot and lower leg slowly or in a number of acute episodes to reach the knee. About 4 million people are said to be affected by the disease worldwide and it is deemed a serious public health problem in at least 10 African countries including Ethiopia. Therefore this study aimed to identify the determinants of Podoconiosis among residence in Machakel district.

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(5.56-44.55) were determinants of Podoconiosis.

33 **Conclusion and recommendation:** Barefoot, sex, family history of podoconiosis and age 34 group were determinants of Podoconiosis. District health office should encourage for at risk 35 populations especially for older age and family history of podoconiosis about shoe wearing 36 practice in all their lives and do not exposed their skin and feet. Key words: Determinants, Machakel, Podoconiosis, Neglected Tropical diseases, Field
Epidemiology, Ethiopia.

39

40 Introduction

Podoconiosis is a chronic disease characterized by the development of persistent swelling of plantar foot initially; which progresses to the dorsal foot and lower leg slowly or in a number of acute episodes to reach the knee. Finally, the disease may end up in a permanent feature of elephantiasis of varying degree. The disease is common in families of barefooted agriculturalists of tropical Africa(1).

46 Podoconiosis (from the Greek word for foot: podos, and dust: konos) is unique in being an 47 entirely preventable non-communicable tropical disease. In local communities, it is often 48 called 'mossy foot disease', because the skin becomes rough and bumpy and its appearance 49 resembles moss(2).

Podoconiosis has a curable pre-elephantiasis phase. However, once elephantiasis is 50 established, podoconiosis persists and may cause lifelong disability(3). Podoconiosis 51 (endemic non- filarial elephantiasis) has been recognized as a specific disease entity for over 52 one thousand years and is widespread in tropical Africa, Central America and north India, yet 53 it remains a neglected and under-researched condition(4). It is non-infective, usually 54 55 crystalline blockage of the limb lymphatic, almost always affecting the lower limbs, 56 It is widely distributed in a number of countries of tropical Africa, especially the feet. 57 Central and South America, North India, Indonesia, Colombia, Ecuador, Brazil and Sri Lanka 58 (5).

59 Podoconiosis (endemic non-filarial elephantiasis) is a geochemical disease occurring in 60 individuals exposed to red clay soil of volcanic origin(6). The disease causes bilateral, but 61 asymmetrical swelling almost invariably of the lower legs(7). Early symptoms of 62 podoconiosis include itching of the skin of the forefoot and recurrent episodes of burning and 63 oedema of the foot or lower leg, especially after periods of intense physical activity(8).

Although the aetiology is not fully understood, existing scientific evidence suggests the important role of exposure to irritant red clay soil in endemic areas as well as the effect of genetic susceptibility(9). Podoconiosis is found in highland areas of tropical Africa, Central 67 America and north-west India. Areas of high prevalence have been documented in Uganda (10), Tanzania (11), Kenya (12), Rwanda, Burundi, Sudan and Ethiopia (13), and in 68 Equatorial Guinea (14), Cameroon (15), the islands of Bioko, Sao Tome & Principe (16) and 69 the Cape Verde islands. And it is related to poverty. Studies have also indicated that 70 podoconiosis exists in areas where the altitude is above 1000meters above sea level and 71 annual rainfall above 1000 millimetres. About 4 million people are said to be affected by the 72 disease worldwide and it is deemed a serious public health problem in at least 10 African 73 74 countries(1,17).

Podoconiosis is a multifactorial disease with evidence of genetic susceptibility and environmental exposure(18). Although the ethology is not fully understood, current evidence suggests genetic susceptibility and the role of mineral particles from irritant volcanic soils supposed to cause podoconiosis(19). The disease only affects some barefoot individuals; not all exposed individuals develop podoconiosis.

Podoconiosis follows a chronic course causing progressively increasing disability with continued exposure to irritant soils. It results in bilateral progressive chronic swelling of the lower legs, usually limited below the level of the knees. The pathogenesis of the disease has not yet been investigated in depth, but it is believed to be caused by fine particles in the soil that penetrate the skin and induce an inflammatory reaction in the lymphatic system(20). However, early stage disease can easily be treated by foot hygiene, bandaging and shoes.

Podoconiosis is classified into five stages where the first stage swelling is limited to below the ankle and is reversible overnight. The second stage swelling is not reversible, and when bumps and knobs are present they remain below the level of the ankle. In the third stage of the disease, bumps and knobs are found above the level of the ankle. The fourth stage entails above knee swelling whereas the fifth stage involves joint fixation as a result of surrounding soft tissue overgrowth(20).

Podoconiosis has recently been included in the World Health Organization's Neglected Tropical Diseases (NTDs) list(21). Areas of high prevalence of podoconiosis have been documented in tropical Africa, Central America and north India(22). Of affected countries, Ethiopia appears to have the highest number of people with podoconiosis(23), with 11 million people at risk through exposure to irritant soil, and an estimated 1 million people affected countrywide(24). In Ethiopia, prevalence estimates range from 2.8 to 7.4% in endemic areas(24-26). Podoconiosis can be prevented, early forms of the disease can be 99 treated, and disease progression can be controlled with simple but effective measures such as 100 washing feet with soap and water on a regular basis and wearing protective shoes 101 consistently(1). Hence this study aimed to study the determinants of podoconiosis in 102 Machakel district Ethiopia.

103 Methods and Materials

104 Study Area

This study was conducted in Machakel district. Machakel district is one of the districts in 105 East Gojjam zone, Amhara national Regional State. It is found at a distance of 328 km North 106 West from Addis Abeba, the capital city of Ethiopia, 237 km far from Bahir Dar, the capital 107 city of Amhara Region. Machakel woreda is bordered by on the North by Bibugn district, on 108 109 the South by Debre Elias district, on the Northwest by Sinan, on the Southwest by Gozamen and on the East by Dembecha (West Gojjam). According to the 2014 E.C projected census 110 2007, the total population is 146,942 from which about 73,618(50.1%) are female and 111 73,324(49.9%) male population. Machakel district has 30 kebeles. In the Machakel district, 112 there are 24 health posts and six health centres all are providing health service. 113

114 Study design and period

115 Unmatched case control study design was conducted from August 30 to September $30 \neq 2022$.

116 Source and Study population

All resident of Machakel district were the source population. Whereas the study population was all adult patients age 18 and older those who identified and registered as podoconiosis case by the district and those neighbouring individuals without podoconiosis.

120 Cases: Were person of age 18 and old, resided in all kebeles of Machakel district and who
121 had diagnosed and registered as podoconiosis case by the district.

122 Controls: Were person with age 18 years and old who had not podoconiosis after diagnosed
123 by clinical nurses elinically and who lives in the neighbouring house to podoconiosis patients
124 (case).

- 125 Inclusion criteria
- 126 Cases: were residents of Machakel district and those registered as podoconiosis cases of age
- 127 18 years and old resident in the study area for at least for the last six months were included.
- 128 Controls: were those age of 18 years and old, resident of Machakel district, who had not
- 129 developed podoconiosis diseases and neighbour to a podoconiosis patient (case).

130 Exclusion criteria

131 Cases: Severely ill and bed riddle cases, age less than 18years and resided less than six132 months was excluded during the data collection period.

133 Controls: Those people without podoconiosis and not being neighbour to the cases were134 excluded.

135 Sample size determination

The sample size was calculated using Epi-info 7 software based on the assumption of 95% confidence interval, 85% power, control to case ratio of 2:1, present of exposed family history among controls 11.4%, odds ratio to be detected as 2.81(15) and non-response rate of 10% yielding 211 controls and 106 cases (317 study participants).

140 Sampling Techniques

141 Simple random sampling technique was used to select the cases from podoconiosis 142 registration books of Machakel woreda health office as a sampling frame. The control groups 143 were included from the community and Simple random sampling technique was used to select the control. Those controls were selected from neighbouring of the cases by lotterymethod.

146 Variables

- 147 Dependent variable
- 148 ✓ Podoconiosis case(yes/no)
- 149 Independent variables
- 150 Socio-demographic Characteristics such as: Age, Sex, Residence, Income,
- 151 Educational status, Occupation,
- 152 Behavioural related factors : Feet washing practice, and Shoe Wearing practice
- **153** ✓ **Family history:**1rst degree, 2nd degree, 3rd degree and others family
- 154 **Operational Definition**
- 155 **Cases:** Those podoconiosis cases registered by the district as a podoconiosis case and found
- 156 on the sampling frame.
- **157 Controls:** A person who had not podoconiosis disease after diagnosed by the clinical nurses
- 158 clinically and neighbours of cases.
- 159 Feet washing: patient daily wash their foot with soap.
- 160 **Barefoot:** not wearing any type of shoe every day.
- 161 **Family history:** History of podoconiosis in the family clustering, such as,1st degree(parents,
- 162 child), 2nd degree(grandparents, siblings), 3rd degree(aunt, uncle, nephew, cousin, niece),
- 163 other (husband and wife)(11).
- 164 **Shoe wearing:** wearing a full covering shoe every day during each activity.
- 165 **Data collection tools and techniques**

Data were collected using interviewer administered questionnaire it was prepared in English and translated into Amharic language for the purpose of community level and back to English for checking language consistency. Data was collected by clinical nurse. The whole data collection process was supervised by health officers.

170 Data quality control

Questionnaire were prepared in English version by reviewing different literatures and 171 translated in to Amharic (local language) and back to English by different language experts 172 for consistency. Steps were taken to ensure the quality of this work. Pre-test was conducted 173 on 16(5%) study participants in another district and necessary corrections were made on the 174 questionnaire. The supervisors and principal investigator closely following the day to day 175 176 data collection process to ensure completeness and consistency of the collected questionnaires on a daily basis. Training was given for the data collectors and supervisors, 177 and the whole data collection process was closely supervised. 178

179 Data processing and analysis

Prior to analysis, the whole data was cleaned and checked for completeness. Errors related to 180 inconsistency was verified using cross tabulation and other data exploration methods. After 181 the completeness of the data checked; it was entered into Epi-Info version 7.2.1.0 software 182 packages and transferred to SPSS version 22 software for analysis. Descriptive statistics was 183 used to give a clear picture of background variables like age, sex, and other variables. The 184 frequency distribution of both dependent and independent variables were done. Binary 185 186 logistic regression was used to identify the determinants of Podoconiosis and the single independent variable with the outcome variable. Hosmer-Lemeshow test was used to check 187 goodness of fit of the model. Variables having an association with the outcome variable a p-188 189 value of less than 0.2 were considered in the multivariable logistic regression analysis.

Adjusted odds ratios (AORs) with 95% confidence intervals were used to show association
between explanatory variables and a dependent variable. Those independent variables with
P-value < 0.05 was considered statistically significant factors associated with outcome
variable.

194 **Results**

195 Socio-demographic characteristics of participants

A total of 312 participants (104 cases and 208 controls) were included, giving for a response rate of 98.42%. The mean age of respondents was 47.21 years (SD 11.47 years). Nearly two third 229(73.32%) were farmers by occupation. The mean age of the cases was 53.8 years (SD=15.43 years). Female constitute 67.30 % of cases, 58(55.76 %) of cases had family history of Podoconiosis, the proportion of bare foot among cases was 30.76% and 76.92% of cases were can not read and write.

The mean age of controls was 43.88 years (SD 9.5 years). Having family history of Podoconiosis was recorded in (57.69%), bare foot (11.53%), Female (44.71%) and of farmer occupation constitutes (71.15%) of controls (Table 1).

Table 1: Socio-demographic characteristics of study participants in Machakel district North-West Ethiopia, 2022.

Characteristics	Category	Cases (n (%))	Controls (n (%))	
Sex	Male	34(54.8)	115(60)	
	Female	70(45.19)	93(44.71)	
Age	21-40	14(13.46)	93(44.71)	
	41-60	63(60.57)	101(48.55)	
	61-80	27(25.96)	14(6.73)	
Residence	Urban	16(15.38)	29(13.9)	
	Rural	88(84.6)	179(86.05)	
Marital status	Unmarried+	32(30.7)	13(6.25)	

9

	Married	72(69.23)	195(93.75)
Occupation	Merchant	4(3.8)	11(5.25)
	Farmer	81(77.88)	148(71.15)
	House wife	16(15.38)	47(22.59)
Education	Not read and wright	80(76.92)	112(53.84)
	Read and wright	21(20.19)	67(32.21)
	primary	2(1.92)	28(13.65)
	Higher	1(0.96)	0(0)
Income	500-1500	48(46.15)	48(23.41)
	1501-3500	43(41.34)	118(57.56)
	>3500	13(12.5)	42(20.2)

207

208 Behavioural related factors of participants

The proportion of bare foot among cases was 32(30.76%) whereas it was 24(11.53%) among controls. About 84(80.76%) of cases and 188(90.38%) of controls did wash their feet 6-8 times per week and 52(50%) of cases, 152(73.07%) of controls wash their feet 6-8 times per week with soap.

- Among cases 26(25%) and 9(8.6%) did wear shoe at work and at home respectively. Whereas 88(42.30%) and 38(18.26%) were among controls, respectively. Regarding mechanism of feet washing, among cases 84(80.76%) washed their legs with water only. About 188(90.38%) controls washed their legs with water only. Among cases 12(11.5%) and controls 120(57.69%) start shoe wearing between 1-20 years of age. About 58(55.76%) of cases and 120(57.69%) controls had family history of podoconiosis (Table 2).
- Table 2: Behavioural related factors of podoconiosis in Machakel district North-West
- **220** Ethiopia, 2022

Characteristics	Category	Cases (n (%))	Controls (n (%))
Having enough		104(100)	208(100)
water for	Yes		
washing			
Washing feet	1-2 times	2(1.92)	0(0)

per week	3-5 times	17(16.34)	13(6.25)	
	6-8 times	84(80.76)	188(90.38)	
	>8	1(0.96)	7(3.36)	
Washing feet	1-2 times	5(4.80)	10(4.80)	
per week with	3-5 times	33(31.73)	35(16.82)	
soap	6-8 times	52(50)	152(73.07)	
	>8	0(0)	2(0.96)	
Barefooted	Yes	32(30.76)	24(11.53)	
	No	72(69.23)	184(88.46)	
Starting year	1-20 year	12(11.53)	120(57.69)	
of shoe	21-40 year	53(50.96)	59(28.36)	
wearing	41-60	7(6.73)	5(2.40)	
Years with	All years	30(28.84)	181(87.01)	
shoe wearing	More than have	22(21.15)	2(0.96)	
	the year			
	Half year	16(15.38)	1(0.48)	
	Less than year	4(3.84)	0()	
Days with shoe	Every day	47(45.19)	194(93.26)	
from the week	5 5	`` '	` '	
	More than 5 day	22(21.15)	4(1.95)	
	2-5 day	35(33.65)	10(4.80)	

221

222 Factors Associated with Podoconiosis

Bare foot, sex, family history of podoconiosis, occupation, marital status, Age and owned a
pair of shoes were found to be associated with Podoconiosis in the uni-variable binary
logistic regression analysis at p values< 0.2.

From multivariable logistic regression analysis, Sex, bare foot, family history of podoconiosis and age were found to be associated with Podoconiosis with 95% CI, at p-value <0.05 statistical significance level. The goodness-of-fit statistics for the model were assessed by using the Hosmer-Lemeshow test, with a p-value of 0.97.

- The odds of having Podoconiosis was 5.83 times higher in bare footed persons (AOR 5.83,
- 231 95% CI: 2.34-14.5) as compared who wore shoes. The odds of having Podoconiosis 4.25
- times higher in females (AOR 4.25: 95% CI: 2.22-8.14) as compared to males. The odds of
- having Podoconiosis was 3.01 times higher among participants who had family history of
- podoconiosis [AOR= 3.01, 95% CI: 1.41-6.42] as compared to that of those who had no
- family history of podoconiosis. The odds of having podoconiosis was 2.05 times higher in
- 236 age group 41-60(AOR=5.05, 95% CI: 2.35-10.83) and 15.74 times higher in age group 61-80
- 237 (AOR=15.74, 95% CI: 5.56-44.55) as compared to age less than 40 (Table 3).
- Table 3: Factors associated with podoconiosis in Machakel district North-West Ethiopia,2022.

Variables	Category	case	Cont	COR [95% CI]	AOR [95% CI]
			rol		
Bare foot	No	72	184	1	1
	Yes	32	24	3.40(1.87-6.17)	5.83(2.34-14.50)*
Sex	Female	70	93	2.54(1.55-4.16)	4.25(2.22-8.14)*
	Male	34	115	1	1
Marital status	Single	3	6	1	1
	Married	80	195	0.82(0.2-3.36)	0.19(0.03-1.11)
	Divorced	11	3	7.33(1.11-48.26)	0.83(0.09-7.5)
	Widowed	10	4	5.00(0.82-30.46)	0.34(0.04-2.94
Age group	21-40	14	93	1	1
	41-60	63	101	4.14(2.17-7.89)	5.05(2.35-10.83)*
	61-80	27	14	12.81(5.44-30.14)	15.74(5.56-44.55)*
Family history	No	46	88	1	1
	Yes	58	120	1.09(1.21-3.11)	3.01(1.41-6.42)*
Occupation	Farmer	81	148	0.36(0.06-2.22)	0.62(0.07-5.35)
	House wife	16	47	0.22(0.03-1.48)	0.27(0.02-2.74)
	Merchant	7	13	1	1
Owned a pair of shoe	No	4	3	2.73(0.6-12.44)	2.5(0.38-16.53)
	Yes	100	205	1	1

240 1=Reference, *=significant at p-value <0.05 in multivariable logistic regression

241

242 **Discussion**

- Even though podoconiosis has been known for more than a millennium, it has been neglected and under-researched. Podoconiosis has recently been designated neglected tropical disease status by the WHO(25).
- 246 Bare footed people were at higher risk to develop podoconiosis as compared to that of their
- 247 counterparts. This implies that bare footed individuals might be exposed to irritant minerals
- found in earth. This leads to Podoconiosis. This finding was agreed with the study conducted
- in Southern Ethiopia(26). This might be due to the reason that barefooted individuals may
- allow mineral particles from the soil to enter in to their feet that can possibly initiate the
- 251 pathology of the disease(27, 28).
- 252 Being female was another variable with higher odds among cases compared to Males. This
- 253 finding agreed with a study done in West and East Gojjam zone shows that; The odds of
- 254 Podoconiosis was lower risk in males (15). This is might be due to variation in exposure in
- 255 males and females in these areas. Traditionally males usually wear shoes frequently than
- 256 females.
- Family history of podoconiosis was also at higher odds among cases than controls. This might be due to the effect of genes in the development of Podoconiosis(29). This study was in line with a study done in West and East Gojjam zone (35).
- Older age, particularly people aged 41-60 years and 61-80 years were at higher risks to develop podoconiosis as compared to those with age group less than 40 years. This implies that older age individuals might have long-term exposure to irritant minerals on the earth that may increase the risk of developing podoconiosis.
- 264
- 265
- 266

267 Limitation of the study

268 The limitation of this study was the possibility of recall bias that might have been introduced269 due to retrospective nature of the study design.

270 Conclusion

271 Podoconiosis is a common but neglected tropical disease leading to dramatic non filarial elephantiasis in the tropics region. This study showed that the 272 determinant factors of Podoconiosis are regular walking on barefoot, family history of 273 274 podoconiosis, female sex and older age. People should be encouraged to wearing shoes all the time. Particularly, females in the study area and other similar settings should be given 275 attention in the existed primary health care packages to promote shoes wearing as males. 276

277 **Declarations**

278 Ethics approval and consent to participate

279 The ethical clearance issues of this study were reviewed and approved by the Ethical Review

280 Committee (IRB) of the College of Medicine and Health Sciences, Bahir Dar University.

281 Permission letter was also obtained from Amhara Public Health Institute (APHI) before the

actual data collection; permission was taken from East Gojjam zone health department and

283 Machakel woreda health office. Written informed consent was taken for each participant.

284 Confidentiality was kept and their name was changed to codes.

- 285 **Consent for publication**
- 286 Not applicable

287 Availability of data and material

The datasets used and/or analysed during the current study are available from the corresponding author on reasonable request.

- 290 Competing interests
- 291 The authors declare that they have no competing interests.
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- 293 No fund was received

294 Authors' contributions

TT conceived and designed the research protocol. KA, YT, AG, KM and GT approved the 295

- 296 proposal with extensive revisions, participated in the data analysis, and had written the
- manuscript. All authors have read and approved the final manuscript. 297
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