

Citation: Mersha TT, Mekonnen Wolde B, Shumuye NA, Hailu AB, Mohammed AH, Redda YT, et al. (2021) Prioritization of neglected tropical zoonotic diseases: A one health perspective from Tigray region, Northern Ethiopia. PLoS ONE 16(7): e0254071. https://doi.org/10.1371/journal. pone.0254071

Editor: Martin Chtolongo Simuunza, University of Zambia, ZAMBIA

Received: October 1, 2020

Accepted: June 21, 2021

Published: July 22, 2021

Peer Review History: PLOS recognizes the benefits of transparency in the peer review process; therefore, we enable the publication of all of the content of peer review and author responses alongside final, published articles. The editorial history of this article is available here: https://doi.org/10.1371/journal.pone.0254071

Copyright: © 2021 Mersha et al. This is an open access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

Data Availability Statement: All relevant data are within the manuscript and its S1, S2 Figs, S1–S6 Files.

RESEARCH ARTICLE

Prioritization of neglected tropical zoonotic diseases: A one health perspective from Tigray region, Northern Ethiopia

Tadesse Teferi Mersha¹, Biruk Mekonnen Wolde^{2*}, Nigus Abebe Shumuye^{1,3}, Abrha Bsrat Hailu², Abrahim Hassen Mohammed⁴, Yisehak Tsegaye Redda², Birhanu Hadush Abera¹, Habtamu Taddele Menghistu^{2,5}

1 Department of Veterinary Clinical Medicine and Epidemiology, College of Veterinary Sciences, Mekelle University, Mekelle, Tigray, Ethiopia, 2 Department of Basic and Diagnostic Sciences, College of Veterinary Sciences, Mekelle University, Mekelle, Tigray, Ethiopia, 3 Lanzihou Veterinary Research Institute, CAAS, Lanzhou, China, 4 Health Bureau, Research, Projects, and International Relations, National Regional State of Tigray, Mekelle, Ethiopia, 5 Institute of Climate and Society, Mekelle University, Mekelle, Tigray, Ethiopia

* mbiruk1972@gmail.com, biruk.mekonnen@mu.edu.et

Abstract

Neglected tropical zoonotic diseases (NTZDs) continue to have a major effect on the health of humans and animals. In this study, a one health approach was used to prioritize and rank neglected tropical zoonotic diseases at the regional and zonal levels in Tigray National Regional State, Ethiopia. For prioritization of NTZDs a cross-sectional study through a structured guestionnaire was administered to 313 health experts from human and animal health sectors. In addition, focus group discussions (FGD) were held with purposively selected key informants. Descriptive, and Multivariable analysis was applied to report the results and a ranked list of diseases was developed at the zonal and regional level. In the region, 8 of the 12 World Health Organization listed NTZDs were considered major diseases including anthrax, brucellosis, bovine tuberculosis, taeniasis, leishmaniasis, rabies, schistosomiasis, and soil-transmitted helminths. Considering the zoonotic and socioeconomic importance of the diseases at the regional level, rabies ranked 1stwhereas anthrax, bovine tuberculosis, leishmaniasis, and brucellosis were ranked from 2nd to 5th, respectively. The FGD result also supported the prioritization result. The Multivariable analysis showed a statistically significant difference in the zonal distribution of anthrax (= 0.009, OR = 1.16), taeniasis (p<0.001, OR = 0.82), leishmaniasis (p<0.001, OR = 1.91), rabies (p = 0.020, OR = 0.79)and soil-transmitted helminths (p = 0.007, OR = 0.87) but not for brucellosis (p = 0.585), bovine tuberculosis (p = 0.505), and schistosomiasis (p = 0.421). Anthrax (p < 0.001, OR = 26.68), brucellosis (p<0.001, OR = 13.18), and taeniasis (p<0.001, OR = 6.17) were considered as the major zoonotic diseases by veterinary practitioners than human health practitioners whereas, leishmaniasis was recognized as a major health challenge by human health professionals. Understanding the priority diseases in the region is supportive for informed decision-making and prioritizes the limited resources to use. Furthermore, strengthening the collaboration between human and animal health professions is important to control the diseases.

Funding: This study received financial support from the MSc thesis research support of Mekelle University (MU) through the recurrent budget and MU-NMBU NORAD project (award recipient; TTM).

Competing interests: The authors have declared that no competing interests exist.

Introduction

The increasing interactions of humans and animals within the environment are aggravating the ongoing transmission of zoonoses from cattle to humans and vice versa [1, 2]. Zoonoses are exerting a significant burden on both animal and human health, particularly in developing countries. Zoonotic diseases impose a double burden on the well-being of people by compromising the health and productivity of their livestock; however, they are often neglected by health facility managers and policymakers of the developed and developing world [3].

Neglected tropical zoonotic diseases (NTZDs) are a subset of zoonoses that primarily affect the world's poorest population [2, 4]. The World Health Organization (WHO) has identified NTZDs namely anthrax, bovine tuberculosis (BTB), brucellosis, leptospirosis, rabies, echino-coccosis, food-borne trematodes, human African trypanosomiasis, taeniasis/cysticercosis, and leishmaniasis [2, 3]. Additionally, soil-transmitted helminths and schistosomiasis are included as major neglected zoonotic diseases in Ethiopia [5, 6].

Ethiopia reported the second-highest burden of zoonotic diseases in Africa [7]; yet, NTZDs has not received attention at various levels in the country. Moreover, data on the burden and distribution of these diseases are incomplete and not updated periodically. To date, prevention and control of NTZDs are challenged by the lack of coordinated efforts between human and animal health professionals and other concerned authorities [6]. As a result, NTZDs continued to affect the livelihoods of poor communities. Considering this fact, Ethiopia has developed a multi-year national master plan that enhances the prevention, control, and eradication of neglected tropical diseases [5, 8].

Selection and prioritization of NTZDs are essential to allocate resource-based investment for their control and prevention. Prioritization can also help to identify vulnerabilities not only where zoonosis poses a significant health threat but also where efforts can be focused to improve prevention, communication, and coordination across veterinary and human health [9–11]. In addition to this, understanding the perceptions of human health and veterinary experts on various NTZDs and their risk is a crucial step to properly plan, manage, and monitor any public health system. This facilitates the prediction of zoonotic disease and in turn, guides federal and regional authorities in decision-making and policy planning for cost-effective resource allocation. The one health approach, based on a multi-sectoral collaboration and coordination, plays a significant role in the prevention and control of zoonoses [3].

A study conducted for the prioritization of zoonotic diseases in Ethiopia has identified rabies, anthrax, brucellosis, leptospirosis, and echinococcosis as the top five diseases [12]. However, this study does not show the priority zoonotic diseases at regional and zonal levels where the distribution and burden of the diseases vary with the diverse agro-ecology of the country and potentially different health systems. Moreover, this study was based on the opinion of experts at the federal level and it doesn't take into account the facts on the ground. A regional/zonal level prioritization of zoonotic diseases is a crucial step for informed decision-making and to design and enforce locally feasible disease control and prevention options. Therefore, this study used a one health approach as a tool to prioritize NTZDs and develop a ranked list of these diseases at the regional and zonal administrative levels in Tigray National Regional State, Ethiopia.

Materials and methods

Ethical statement

The study was reviewed and approved by the government of the National Regional State of Tigray, Bureau of Health (Ref.No. 31/1418/17). Permission was also obtained from each study

district and verbal consent was obtained from all volunteer participants. Confidentiality for all collected information was preserved using secret codes for each participant.

Study area description

The study was conducted from November 2017 to May 2018 in Tigray National Regional State, Ethiopia. Tigray Region is in the northernmost of the country bordered by Eritrea to the north, Sudan to the west, the Afar region to the east, and the Amhara region to the south and southwest. The region is situated between 12°30'N and 15°N latitude, and 36°30'E and 40°30'E longitude. Topographically the region has a diversified agro-ecology, having an altitude ranging from 500–3,935 meter above sea level. Moreover, the region is characterized by an arid and semi-arid climate with low and erratic rainfall. The mean annual temperature of the region is between 15°C and 21°C. The estimated population projections for 2017 based on the 2007 census is 5,247,005 (49.27% male, and 50.73% female) [13]. During this period, the region had seven zones, and 35 districts [14] (Fig 1).

According to the Central Statistical Agency (CSA) projection, the estimated population of the region in 2017 was 5,247,005 (49.27% male, and 50.73% female) [15]. The livestock population of the region was estimated at 4,791,341 heads of cattle, 2,041,731 heads of sheep, 4,584,138 heads of goat, 3,815 heads of horse, 7,634 heads of mule, 838,053 heads of donkey, 54,348 heads of camel, 5,735,973 heads of poultry, and 287,135 beehives [16]. The region has one referral hospital, 16 zonal hospitals, 22 primary hospitals, 202 health centers, 712 health posts, and 159 Veterinary clinics [14].

Study design, study population and sample size determination

Cross-sectional study design was employed. Key informants' in-depth interview (KIDI) and Focus Group Discussion (FGD) were conducted with health experts (human health and veterinary) from all zones of the region. Depending on their size, infrastructures, and transport accessibility for data collection, one to three districts (a total of 15 districts) were selected from each zone. In addition, the 13 major urban centers in the study zones were considered. In each district/urban center, health experts from human health facilities and veterinary clinics were

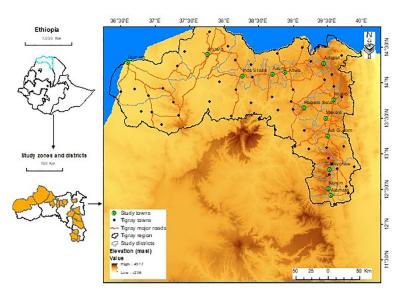


Fig 1. Map of the study zones, districts, and urban centers.

https://doi.org/10.1371/journal.pone.0254071.g001

selected for the KIDI. Some of them participated in the FGD. For the KIDI, the sample size was calculated based on the formula developed by WHO for questionnaire-based studies in health research and previously used by [17], using $n = Z^{2*}P(1-P)/d^2$ where n = number of health experts (both human health and veterinary experts) enrolled in the study, Z = represents a critical value for the 95% confidence interval (1.96), d = the level of precision (5%) and P = proportion used on expected prevalence (p = 0.5). Since the total health experts in the region were less than 10,000, a correction formula [nf = ni/(1 + ni/N)] was used, where N = total health experts in the region (study population, N = 7278). The number of health experts, in each zone, to be involved in the study subjects in each zone, Ni = total number of health experts in each zone, n = total number of study subjects obtained and N = total number of health experts in the region. A 10% non-response rate was also considered to maximize precision. Therefore, 421 health experts were included for the KIDI (Table 1). For the FGD, Health experts from each district were selected purposively based on their understanding and relevance to the study objectives as verified in the KIDI.

Methods of data collection

Key informants' in-depth interview (KIDI). Key informants were selected from diverse backgrounds (multi-sectoral) including public health, epidemiologist, laboratory technologists, nurses, physicians, disease surveillance experts, and veterinary professionals. Participants were informed of the procedures and significance of the study. A pre-test questionnaire was administered to collect relevant data and information regarding the status of the twelve NTZDs and their impact on the community. Key informants were asked about the severity of illness in humans and/or animals, mode of transmission, impact (health and economic burden) of the diseases, inter-sectoral collaborations, control measures (availability of interventions), and major risk factors of NTZDs (S1 File). The questionnaire survey was pre-tested in two selected districts within two selected study zones. From each district ten professionals (veterinary and human health) were involved in the pre-testing. Based on the results of the pre-test, the questionnaire contents were modified. To minimize bias during the discussion with informants, the information was correlated with data generated from the bureau of health and literature [12, 18, 19].

Table 1. Number of health experts by zone an	d sample size determination for questionnaire survey.
Tuble 1. Humber of neurin experts by zone un	a sumple size acter inmation for questionnance survey.

Zones		He	alth experts			Total sample size			
	Total population	ni = (Ni*n/ N)	nf = ni/ (1+ni/ N)	Sample size with 10% NRR	Total population	ni = (Ni*n/ N)	nf = ni/(1+ni/ N)	Sample size with 10% NRR	
Western	981	51	51	56	82	4	4	5	61
Eastern	1331	69	69	76	69	4	4	4	80
Southern	1344	70	69	76	74	4	4	4	80
N/ Western	860	45	45	49	115	6	6	7	56
S/Eastern	641	33	33	37	46	2	2	3	40
Mekelle	448	23	23	26	23	1	1	1	27
Central	1237	64	64	70	122	6	6	7	77
Total	6842	6842 390					31		421

 $NRR = Non-response \ rate; \ N/Western = Northwestern; \ S/Eastern = Southeastern.$

https://doi.org/10.1371/journal.pone.0254071.t001

Focus group discussion (FGD). The FGD aimed to support the disease distribution, burden, and prioritization of NTZDs, and to assess the associated risk factors and the laboratory infrastructures in the respective zones. In each zone, a group having 7–12 participants from the selected study districts and urban centers was formed composing the diverse background of experts who already participated in KIDI. The FGD points were mainly focusing on the impact of NTZDs on socioeconomic and public health importance, distribution, and burden of NTZDs in the study zone. Moreover, questions related to the identification of major NTZDs, the status of laboratory infrastructure, diagnostic techniques and treatment efficacy, disease reporting system, inter-sectoral collaboration, and major risk factors of NTZDs were included (S2 File). The discussion was conducted using the local language (Tigrigna) and the researcher took notes and tape-recorded. Finally, the discussion points were transcribed and translated into the English language.

Prioritization and ranking of neglected tropical zoonotic diseases. The prioritization started with the identification of a specific list of NTZDs [12] followed by the development of structured ranking criteria [6]. The criteria used for the prioritization include severity of illness to humans (max score = 1), transmission potential between humans and animals (max score = 0.85), economic burden of the disease (max score = 0.65), intersectoral collaboration (max score = 0.45) and availability of intervention (max score = 0.30). If a disease is of high priority, it will get a maximum score of 3.25 for a single respondent. The structured questionnaire was distributed to the selected key informants based on the inclusion criteria (S3 File). The participants were eligible if they had two and above years of work experience in the study area and if they were able to answer at least four questions on knowledge assessment criteria [20, 21]. Finally, the score given by the informants was summarized, and diseases were ranked.

Data management and analysis

For the in-depth interviews, all the responses of the informants were assessed in relation to the facts obtained from publications and the general truth of NTZDs. The answer for each question was ranked based on their significance of measurement for NTZDs in the area and scores assigned based on the response to each question. A decision tree was designed using Microsoft Excel and used to determine the final disease ranking. The scores for a single question were multiplied by the number of respondents who correctly answered the questions. Finally, the total scores for each disease at the zonal and regional level were summed up to rank the disease according to its priority. Data compiled during the literature review was used to determine appropriate responses for each question for all NTZDs under consideration.

Data obtained from the questionnaire survey were entered into Microsoft Excel 2010 spreadsheet, exported to STATA version 15.0 for Windows (Stata Corp. College Station, USA), coded and analyzed. Descriptive statistics and Multivariable analysis were applied to present the results. A univariable logistic regression was applied to measure the strength of association between the dependent and independent variables before running the Multivariable logistic regression. The logistic regression model was fitted with individual NTZD result in the zones (whether a major zoonoses or not) as the outcome. The model was built using the forward stepwise (conditional) selection procedure by applying the iterative maximum likelihood estimation procedure, while the statistically significant contribution of individual predictors to the models was tested using the Wald's test and likelihood-ratio tests. Any interaction between variables was assessed by constructing a multivariable model as described previously [22, 23]. The logistic model was checked for goodness-of-fit using the Hosmer and Lemeshow test. The odd ratio at 95% CI was computed and results were considered significant at p-value<0.05. Zone, profession, and experience were considered as independent variables against the existence of major NTZDs in the zones.

Results

Key informants' in-depth interview (KIDI)

A total of 421study participants were enrolled for KIDI, but 21 of them were excluded due to the incompleteness of their data. Thus, the KIDI analysis was performed based on the response from 400 experts.

The burden of NTZDs by expert perspectives

Of the 12 listed NTZDs, eight of them namely anthrax, brucellosis, bovine tuberculosis (BTB), taeniasis, leishmaniasis, rabies, schistosomiasis, and soil-transmitted helminths (STH) were considered as the major zoonotic diseases in the region. Rabies (92%, 368/400) followed by BTB (57.8, 231/400), and leishmaniasis (44.5, 178/400) were considered as the most important major zoonotic diseases in the region (Table 2). The ranking of NTZDs by profession showed that anthrax, brucellosis, taeniasis/cysticercosis and hydatidosis were ranked as major NTZDs by veterinarians whereas leishmaniasis, schistosomiasis, and STH were considered as major priority NTZDs by human health experts (S1 Fig). Professionals above 1 year of experience consider NTZDs as major zoonoses in their zones (S2 Fig).

Variable	Category	Anthrax, Yes (%)	Brucella, Yes (%)	BTB, Yes (%)	Taenia, Yes (%)	Hydatid, Yes (%)	Leishmania, Yes (%)	HAT, Yes (%)	Lepto, Yes (%)	Rabies, Yes (%)	Schisto, Yes (%)	STH, Yes (%)	FBT, Yes (%)
Zone	WZ (n = 59)	27 (45.8)	20 (33.9)	37 (62.7)	14 (23.7)	8 (13.6)	54 (91.5)	1 (1.7)	2 (3.4)	51 (86.4)	8 (13.6)	9 (15.3)	5 (8.5)
	NWZ (n = 53)	25 (47.2)	11 (20.8)	35 (66.0)	21 (39.6)	3 (5.7)	44 (83.0)	0 (0)	0 (0)	48 (90.6)	27 (50.9)	15 (28.3)	2 (3.8)
	CZ (n = 75)	27 (36.0)	21 (28.0)	41 (54.7)	15 (20.0)	4 (5.3)	40 (53.3)	0 (0)	1 (1.3)	68 (90.7)	52 (69.3)	17 (22.7)	3 (4.0)
	EZ (n = 73)	40 (54.8)	19 (26.0)	32 (43.8)	27 (37.0)	5 (6.8)	11 (15.1)	2 (2.7)	3 (4.1)	65 (89.0)	12 (16.4)	40 (54.8)	20 (27.4)
	SEZ (n = 38)	7 (18.4)	13 (34.2)	27 (71.1)	15 (39.5)	12 (31.6)	2 (5.3)	0 (0)	6 (15.8)	38 (100)	13 (34.2)	17 (44.7))	14 (36.8)
	MZ (n = 26)	10 (38.5)	6 (23.1)	19 (73.1)	16 (61.5)	8 (30.8)	16 (61.5)	3 (11.5)	9 (34.6)	26 (100)	15 (57.7)	13 (50.0)	2 (7.7)
	SZ(n = 76)	21 (27.6)	24 (31.6)	40 (52.6)	35 (46.1)	10 (13.2)	11 (14.5)	1 (1.3)	2 (2.6)	72 (94.7)	29 (38.2)	23 (30.3)	14 (18.4)
Region		157(39.3)	114(28.5)	231 (57.8)	143(35.8)	50(12.5)	178(44.5)	7(1.8)	23(5.8)	368 (92.0)	156(39)	134 (33.5)	60(15)
Profession	Vets (n = 43)	40 (93.0)	34 (79.1)	28 (65.1)	31 (72.1)	27 (62.8)	15 (34.9)	1 (2.3)	4 (9.3)	41 (95.3)	12 (27.9)	10 (23.3)	12 (27.9)
	Human (357)	117 (32.8)	80 (22.4)	203 (56.9)	112 (31.4)	23 (6.4)	163 (45.7)	6 (1.7)	19 (5.3)	327 (91.6)	144 (40.3)	124 (34.7)	48 (13.4)
Experience	< 1 yr (n = 73)	21 (28.8)	16 (21.9)	40 (54.8)	24 (32.9)	6 (8.2)	33 (45.2)	0 (0)	2 (2.7)	62 (84.9)	23 (31.5)	23 (31.5)	12 (16.4)
	$\begin{array}{c} 1-3 \text{ yrs} \\ (n=82) \end{array}$	29 (35.4)	26 (31.7)	47 (57.3)	28 (34.1)	7 (8.5)	41 (50.0)	3 (3.7)	4 (4.9)	76 (92.7)	33 (40.2)	22 (26.8)	15 (18.3)
	3–5 yrs (n = 87)	31 (35.6)	23 (26.4)	55 (63.2)	30 (34.5)	12 (13.8)	33 (37.9)	0 (0)	8 (9.2)	86 (98.9)	32 (36.8)	32 (36.8)	14 (16.1)
	> 5 yrs (n = 158)	76 (48.1)	49 (31.0)	89 (56.3)	61 (38.6)	25 (15.8)	71 (44.9)	4 (2.5)	9 (5.7)	144 (91.1)	86 (54.4)	57 (36.1)	19 (12.0)

Table 2. Zonal distribution of major NTZDs and comparisons with profession and working experience.

<u>NB</u>: Brucella = Brucellosis; BTB = Bovine Tuberculosis; Taenia = Taeniasis; Hydatid = Hydatidosis; Leishamania = Leishmaniasis; HAT = Human African Trypanosomiasis; Lepto = Leptospirosis; Schisto = Schistosomiasis; STH = Soil Transmitted Helminths; FBT = Foodborne Trematodes.

WZ = Western zone; NWZ = Northwestern zone; CZ = Central zone; EZ = Eastern zone; SEZ = Southeastern zone; MZ = Mekelle zone; SZ = Southern zone.

https://doi.org/10.1371/journal.pone.0254071.t002

Variable	Category		Anthrax		Brucellosis	Tuberculo	osis	Taeniasis/	Taeniasis/cysticercosis		
		Yes (%)	AOR (95%CI)	Yes (%)	AOR (95%CI)	Yes (%)	OR (95%CI)	Yes (%)	OR (95%CI)		
Zone	WZ (n = 59)	27 (45.8)		20 (33.9)		37 (62.7)		14 (23.7)			
	NWZ (n = 53)	25 (47.2)	1.184 (0.519–2.702)	11 (20.8)	2.768 (1.032–7.422)	35 (66.0)	0.884 (0.402- 1.942)	21 (39.6)	0.461 (0.193–1.099)		
	CZ (n = 75)	27 (36.0)	1.585 (0.738-3.404)	21 (28.0)	1.234 (0.548–2.778)	41 (54.7)	1.371 (0.679– 2.768)	15 (20.0)	1.190 (0.449–2.837)		
	EZ (n = 73)	40 (54.8)	0.676 (0.320-1.426)	19 (26.0)	1.409 (0.613-3.237)	32 (43.8)	2.121 (1.043– 4.315)	27 (37.0)	0.474 (0.210–1.067)		
	SEZ (n = 38)	7 (18.4)	4.832 (1.585–14.728)	13 (34.2)	0.854 (0.332-2.193)	27 (71.1)	0.719 (0.294– 1.757)	15 (39.5)	0.379 (0.148-0.972)		
	MZ (n = 26)	10 (38.5)	1.534 (0.536-4.384)	6 (23.1)	1.758 (0.533–5.799)	19 (73.1)	0.625 (0.224- 1.745)	16 (61.5)	0.164 (0.058–0.464)		
	SZ (n = 76)	21 (27.6)	2.643 (1.180-5.922)	24 (31.6)	1.086 (0.488-2.414)	40 (52.6)	1.572 (0.779– 3.173)	35 (46.1)	0.309 (0.139–0.685)		
Profession	Vets (n = 43)	40 (93.0)		34 (79.1)		28 (65.1)		31 (72.1)			
	Human (357)	117 (32.8)	30.801 (9.065– 104.648)	80 (22.4)	15.163 (6.732– 34.153)	203 (56.9)	1.419 (0.718– 2.806)	112 (31.4)	6.258 (2.986– 13.114)		
Experience	< 1 yr(n = 73)	21 (28.8)		16 (21.9)		40 (54.8)		24 (32.9)			
	1-3 yrs(n=82)	29 (35.4)	0.593 (0.280-1.256)	26 (31.7)	0.513 (0.235-1.121)	47 (57.3)	0.951 (0.494- 1.830)	28 (34.1)	0.959 (0.468–1.964)		
	3-5 yrs(n=87)	31 (35.6)	0.560 (0.262–1.119)	23 (26.4)	0.807 (0.358–1.816)	55 (63.2)	0.802 (0.414- 1.554)	30 (34.5)	1.276 (0.619–2.629)		
	> 5 yrs (n = 158)	76 (48.1)	0.510 (0.261–0.997)	49 (31.0)	0.694 (0.338-1.427)	89 (56.3)	1.016 (0.569– 1.815)	61 (38.6)	0.974 (0.515–1.841)		

Table 3. Multivariable analysis of anthrax, brucellosis, TB, and taeniasis as major zoonoses against the zone, profession, and work experience.

NB: WZ = Western zone; NWZ = Northwestern zone; CZ = Central zone; EZ = Eastern zone; SEZ = Southeastern zone; MZ = Mekelle zone; SZ = Southern zone.

https://doi.org/10.1371/journal.pone.0254071.t003

According to the perception of experts using the Multivariable analysis, anthrax was considered 4.83 and 2.64 times as a major zoonosis in Western zone compared to Southeastern (AOR = 4.832, CI = 1.585-14.728) and Southern (AOR = 4.643, CI = 1.180-5.922) zones, respectively; whereas brucellosis was 2.77 times more important zoonotic diseases in Western zone (AOR = 2.768, CI = 1.032-7.422) than Northwestern zone. Bovine TB was also considered 2.12 times as a major zoonosis in Western zone compared to Eastern zone (AOR = 2.121, CI = 1.043-4.315). Taeniasis was considered as less major zoonoses in Western zone of the region compared to Southeastern, (AOR = 0.379, CI = 0.148-0.972), Mekelle (AOR = 0.164, CI = 0.058-0.464), and Southern (AOR = 0.308, 0.139-0.685) zones (Table 3).

Leishmaniasis was considered as a major zoonosis by 91.5% and 83% of the experts in Western and North-western zones, respectively, and the disease was considered 10.103, 66.464, 229.974, 6.850, and 74.936 times as major zoonosis in the Western zone compared to Central(AOR = 10.103, CI = 3.544–28.804), Eastern(AOR = 66.464, CI = 21.135-209.012), Southeastern(AOR = 229.974, CI = 40.930-1292.138), Mekelle (AOR = 6.850, CI = 1.978-23.728) and Southern(AOR = 74.936, CI = 23.676-237.180) zones, respectively. Rabies was considered as a major zoonosis in all zones of the region and there was no statistically significant variation between zones. According to the perception of experts, the burden of schistosomiasis was less in the Western zone by 0.16, 0.07, 0.30, 0.11, and 0.25 times, respectively, than Northwestern(AOR = 0.159, CI = 0.062-0.404), Central(AOR = 0.069, CI = 0.028-0.170), Southeastern(AOR = 0.302, CI = 0.109-0.840), Mekelle(AOR = 0.112, CI = 0.037-0.337), and

Variable	Category		Leishmaniasis		Rabies	Schistosor	niasis	STH		
		Yes (%)	AOR (95%CI)	Yes (%)	AOR (95%CI)	Yes (%)	OR (95%CI)	Yes (%)	OR (95%CI)	
Zone	WZ (n = 59)	54 (91.5)		51 (86.4)		8 (13.6)		9 (15.3)		
	NWZ (n = 53)	44 (83.0)	2.087 (0.635–6.864)	48 (90.6)	0.869 (0.252–2.999)	27 (50.9)	0.159 (0.062– 0.404)	15 (28.3)	0.467 (0.182–1.97)	
	CZ (n = 75)	40 (53.3)	10.103 (3.544–28.805)	68 (90.7)	0.625 (0.207–1.890)	52 (69.3)	0.069 (0.028– 0.170)	17 (22.7)	0.650 (0.264– 1.597)	
	EZ (n = 73)	11 (15.1)	66.464 (21.135–209.012)	65 (89.0)	0.759 (0.257–2.238)	12 (16.4)	0.840 (0.315-2.242)	40 (54.8)	0.157 (0.067– 0.368)	
	SEZ (n = 38)	2 (5.3)	229.974 (40.930– 1292.138)	38 (100)	1.000	13 (34.2)	0.302 (0.109-0.840)	17 (44.7))	0.238 (0.090-0.628)	
	MZ (n = 26)	16 (61.5)	6.850 (1.978-23.728)	26 (100)	1.000	15 (57.7)	0.112 (0.037- 0.337)	13 (50.0)	0.194 (0.067– 0.560)	
	SZ (n = 76)	11 (14.5)	74.936 (23.676–237.180)	72 (94.7)	0.434 (0.121–1.556)	29 (38.2)	0.248 (0.102-0.603)	23 (30.3)	0.420 (0.176– 1.000)	
Profession	Vets (n = 43)	15 (34.9)		41 (95.3)		12 (27.9)		10 (23.3)		
	Human (357)	163 (45.7)	0.340 (0.140-0.824)	327 (91.6)	2.228 (0.495– 10.034)	144 (40.3)	0.492 (0.230– 1.057)	124 (34.7)	0.540 (0.248– 1.176)	
Experience	< 1 yr(n = 73)	33 (45.2)		62 (84.9)		23 (31.5)		23 (31.5)		
	1-3 yrs(n=82)	41 (50.0)	0.846 (0.354–2.021)	76 (92.7)	0.413 (0.140–1.221)	33 (40.2)	0.605 (0.287- 1.279)	22 (26.8)	1.165 (0.557– 2.437)	
	3-5 yrs(n=87)	33 (37.9)	0.948 (0.400-2.247)	86 (98.9)	0.081 (0.010-0.659)	32 (36.8)	0.791 (0.377– 1.659)	32 (36.8)	0.859 (0.424– 1.742)	
	> 5 yrs (n = 158)	71 (44.9)	1.166 (0.558–2.435)	144 (91.1)	0.570 (0.234–1.392)	86 (54.4)	0.576 (0.295– 1.125)	57 (36.1)	0.852 (0.452- 1.608)	

Table 4. Multivariable analysis of leishmaniasis, rabies, schistosomiasis and STH as major zoonoses against the zone, profession, and work experience.

NB:WZ = Western zone; NWZ = Northwestern zone; CZ = Central zone; EZ = Eastern zone; SEZ = Southeastern zone; MZ = Mekelle zone; SZ = Southern zone.

https://doi.org/10.1371/journal.pone.0254071.t004

Southern(AOR = 0.248, CI = 0.102–0.603) zones. Similarly, the burden of STH was less in the Western zone by 0.16, 0.24, and 0.19 times, respectively, compared to Eastern (AOR = 0.157, CI = 0.067–0.368), Southeastern(AOR = 0.237, CI = 0.090–0.628), and Mekelle(AOR = 0.194, CI = 0.067–0.560) zones (Table 4).

Compared to health professionals, veterinarians indicated anthrax, brucellosis, and taeniasis 30.80(AOR = 30.801, CI = 9.065-104.648), 15.16(AOR = 15.163, CI = 6.732-34.153), and 6.26 (AOR = 6.258, CI = 2.986-13.114) times, respectively, as major health challenges in their professional carrier (Table 3). However, leishmaniasis was considered as less health challenge by veterinary professionals (AOR = 0.340, CI = 0.140-0.824) than human health experts (Table 4), but it was identified as an important zoonosis by human health experts. Except for the case of rabies, the experience of the health experts didn't influence to consider these diseases as major zoonoses in their respective zones. However, health experts with experience less than 1 year considered rabies 0.08 (AOR = 0.081, CI = 0.010-0.659) times less than experts with 3-5 years' experience as a major zoonosis (Table 4).

Prioritization of NTZDs by health experts

Out of the 400 interviewed respondents, considering the identified exclusion criteria, responses from 313 experts were considered for the disease ranking. Rabies ranked 1st with a regional score of 885.83 out of the maximum score of 1017.25. Anthrax, BTB, leishmaniasis, brucellosis, and leptospirosis ranked from 2nd to 6th, respectively (<u>Table 5</u>). Likewise, the three major NTZDs in each zone were rabies, anthrax, and BTB (<u>Table 6</u>).

Diseases	Regional Score	R	ank
Rabies	8	885.83	1^{st}
Anthrax	5	792.08	2 nd
Bovine Tuberculosis (BTB)		785.85	3 rd
Leishmaniasis		724.27	4^{th}
Brucellosis		721.97	5 th
Leptospirosis	6	542.57	6 th
Hydatidosis	6	538.52	7 th
Schistosomiasis	5	592.06	8^{th}
Taeniasis		571.55	9 th
Food-borne Trematodes (FBT)		570.05	$10^{\rm th}$
Soil-Transmitted Helminths (STH)	5	550.68	$11^{\rm th}$
Human African Trypanosomiasis (HAT)		532.86	12 th

Table 5. Regional prioritization of neglected tropical zoonotic disease by experts.

https://doi.org/10.1371/journal.pone.0254071.t005

Focus group discussion (FGDs)

Concerning the impact of NTZDs on public health and socio-economic importance, the discussants from all zones of the region indicated the adverse effects of NTZDs on both human and animal health. According to the discussants, the major negative impacts of these diseases in animals were high morbidity and mortality, chronic illness, low production, and productivity, international trade restriction, and retard genetic improvement programs. Similarly, the public health effects of NTZDs identified by the discussants were overburden on the public health system, massive economic and social troubles, insufficient human resource development, stunting growth, easy way of disease transmission, and compromising the working performance of individuals. One of the key messages forwarded by the discussants was "*as significant numbers of impoverished people may not afford the additional costs for medications, often they are trapped in a never-ending cycle of poverty. Because of the endemicity of these diseases, there is also an impact on the cost of treatment, control and prevention programs*".

Rabies, bovine tuberculosis, anthrax, leishmaniasis, schistosomiasis, brucellosis were identified as major NTZDs with wider distribution across the region. According to the discussants,

Disease	W	Z	NV	NZ	C	Z	E	Z	SI	ΞZ	М	Z	SZ	Z
	Score	Rank	Score	Rank	Score	Rank	Score	Rank	Score	Rank	Score	Rank	Score	Rank
Rabies	135.18	1 st	146.4	1 st	160.75	1 st	164.05	1 st	61.18	1st	51.59	1st	166.64	1st
Anthrax	120.69	3 rd	130.2	2 nd	143.13	2 nd	147.99	3 rd	55.85	2nd	44.67	2nd	149.60	2nd
ВТВ	125.97	2 nd	125.1	3 rd	139.87	3rd	152.27	2 nd	50.90	4th	43.18	3rd	148.54	3rd
Leishmaniasis	113.62	4 th	119.1	4 th	132.54	5th	131.00	5 th	50.00	5th	41.52	4th	136.50	4th
Brucellosis	106.25	5 th	117.9	5 th	135.46	4th	138.63	4^{th}	51.98	3rd	39.37	5th	132.41	5th
Shistosomiasis	87.76	6 th	99.4	6 th	108.88	8th	111.44	8 th	43.54	8th	30.67	8th	110.39	8th
Hydatidosis	85.15	7 th	94.8	8 th	118.63	7th	124.93	7 th	47.51	6th	36.68	7th	130.82	6th
Leptospirosis	84.13	8 th	97.4	7 th	124.68	6th	125.50	6 th	45.48	7th	37.18	6th	128.21	7th
Taeniasis	82.23	9 th	92.8	9 th	106.36	10th	109.90	9 th	42.05	10th	29.85	9th	108.38	9th
STH	81.60	11 th	90.7	11th	104.21	11th	106.67	11 th	38.87	11th	29.66	11th	98.93	12th
HAT	81.59	12 th	88.2	12th	96.72	12th	96.93	12 th	38.37	12th	29.20	12th	101.82	11th
FBT	81.83	10 th	92.6	10th	107.72	9th	107.60	$10^{\rm th}$	42.74	9th	29.40	10th	108.13	10th

Table 6. Zonal prioritization of neglected tropical zoonotic disease by experts.

NB:WZ = Western zone; NWZ = Northwestern zone; CZ = Central zone; EZ = Eastern zone; SEZ = Southeastern zone; MZ = Mekelle zone; SZ = Southern zone.

https://doi.org/10.1371/journal.pone.0254071.t006

Disease rank	Zone											
	WZ	NWZ	CZ	EZ	MZ and SEZ	SZ						
1	Rabies	Rabies	Rabies	Rabies	Rabies	Rabies						
2	Leishmaniasis	Leishmaniasis	Schistosomiasis	Tuberculosis	Anthrax	Tuberculosis						
3	Brucellosis	Anthrax	Anthrax	Anthrax	Helminthiasis	Anthrax						
4	Tuberculosis	Brucellosis	Tuberculosis	Helminthiasis	Tuberculosis	Brucellosis						
5	Anthrax	Tuberculosis	Helminthiasis	Brucellosis	Leishmaniasis	Schistosomiasis						

Table 7. List of top five NTZDs by zone through the FGD.

NB: WZ = Western zone; NWZ = Northwestern zone; CZ = Central zone; EZ = Eastern zone; SEZ = Southeastern zone; MZ = Mekelle zone; SZ = Southern zone.

https://doi.org/10.1371/journal.pone.0254071.t007

rabies was considered as the most important zoonotic disease with wider distribution throughout the zones (Table 7).

Discussion on disease reporting system and institutional collaboration indicated that there was poor veterinary case recording system in all zones of the region characterized by limited recording practice and disease data management. Rather diseases were reported generally as bacterial infection, parasitic infection, viral infection, and so on. On the other hand, the human health recording and reporting system are much better and are computerized through the Health Management Information System (HMIS) from the bottom (district) to top (region). However, HMIS data does not include disease reports of anthrax, brucellosis, hydatidosis, taeniasis, soil-transmitted helminths, and food-borne trematodes that are listed by WHO and the national master plan of Ethiopia for neglected tropical diseases. All the discussants agreed that the collaboration between human and animal health professionals in disease communication, control, and prevention was weak. Collaborations between the two sectors are initiated whenever there is an outbreak of rabies. Lack of awareness about NTZDs, free animal movement and illegal trade, close human-animal interaction, feeding habits of raw animal product, illegal slaughtering practices, presence of suitable environment for some vectorborne NTZDs, and lack of planned control and prevention strategies and laboratory infrastructure were identified as major risk factors in epidemiology and dynamics of NTZDs.

Discussion

In the present study, prioritization and ranking of NTZDs were conducted through KIDI and triangulated with the FGDs. Prioritization of NTZDs permits a region to reconsider priority diseases periodically and guides the direction of the limited resource allocation.

In this study, the top five prioritized NTZDs in Tigray region in descending order of importance were rabies, anthrax, bovine tuberculosis (BTB), leishmaniasis, and brucellosis. These top five priority NTZDs were also included in the list of priority zoonotic diseases at the country level, Ethiopia [12] reflecting the importance of these diseases in both the region and the country. In addition, a mapping study conducted in selected districts of Tigray, Afar, and Amhara regions has shown rabies, TB, leishmaniasis, and schistosomiasis as major NTZDs [8]. Rabies, anthrax, and brucellosis were considered in Kenya and Uganda among the top priority zoonotic diseases [24, 25].

Rabies is a fatal but neglected zoonotic disease, which constitutes a major public health concern globally [26, 27]. In this study, the KIDIand FGD results highlighted the importance of rabies in the region, where it was ranked as the 1st priority zoonoses in all the zones and the region. The high rank of rabies in the region could be due to the poor management of owned dogs, the presence of a high population of unvaccinated stray dogs, and lack of an effective rabies control program [28–31]. This indicates the region is far behind the Global Rabies eradication program by 2030 [32].

Furthermore, anthrax and BTB were prioritized as the other next important NTZDs. Focus group discussion results also support the fact that anthrax and BTB ranked as the top five NTZDs in all zones of the region. Anthrax is among the most prevalent diseases of animals with repeated annual outbreak and is one of the major public health concern in Ethiopia with the highest human case prevalence reported in the Tigray region [5, 33]. Even though there is an effective anthrax vaccine for use in animals, the annual vaccination coverage is challenged by limited vaccine delivery and public reluctance to vaccinate their animals [34]. This suggests the need for increased community awareness on public health importance of anthrax in addition to rising anthrax vaccine coverage in the animals from time to time as human anthrax occurs by contact with infected animals or contaminated animal products [35–37]. The country has no vaccine for use in humans.

Bovine tuberculosis (BTB) is a recognized endemic disease of cattle in Ethiopia and is a frequent cause of zoonotic human tuberculosis. The absence of effective cattle TB control programs and lack of routine milk pasteurization procedures in Ethiopia favors the widespread of human tuberculosis due to *Mycobaterium bovis*(*M. bovis*) [38, 39]. In addition, the close physical contact between farmers and their cattle in the Tigray region [40]promotes aerosol transmission. In human beings, infection with BTB presents with a special challenge for patient treatment and recovery as *M. bovis* is naturally resistant to pyrazinamide, one of the four medications used in the standard first-line anti-tuberculosis treatment regimen leading to long hospitalization, high treatment cost, and low productivity [41].

In this study, leishmaniasis ranked as 4th priority NTZDs in the region but with a statistically significant variation in the zonal distribution. Of the respondents, 91.5%, 83%, and 53.3% of the experts from Western, Northwestern, and Central zones, respectively indicated that the disease is a major zoonosis in their respective zones, which was supported by the FGD results too. This finding is in line with prior researches in different parts of the country [42–44]. The reasons for an increment of the disease in the areas particularly in the Western zone Kafta-Humera could be strongly associated with the endemic nature of the disease, migration of a large number of the labor force for the job opportunity, high temperature, and the environment is very suitable for the survival and replication of the vector [44–46].

Brucellosis is reported in many regions of the world including Ethiopia as endemic with high economic loss and zoonotic potential [47–49]. In the present study, brucellosis ranked as the fifth most important zoonotic disease in the region, which is in line with the findings of previous studies [12, 24, 25]. The high prevalence of the disease in animals and anthropogenic factors such as eating habits, poor hygiene, and practices that expose humans to infected animals or their products play a major role in the epidemiology of the disease [48, 50].

Unlike the reports from neighboring Kenya, Somalia, and Uganda, Human African Trypanosomiasis was ranked as the least priority NTZD in the present study [24, 25, 51]. This could be due to the reason that Tigray region lies outside the tsetse belt areas of the country where there is no risk to acquire the disease [52]. In addition, the disease was not considered among the priority diseases in the national neglected tropical disease in Ethiopia [5] and it was not also among the prioritized diseases according to a previous study at national level [12].

There was a statistically significant difference in the zonal distribution of rabies, anthrax, leishmaniasis, taeniasis, and soil-transmitted helminths. This zonal variation could be due to differences in health infrastructure facilities, agro-ecology that can affect pathogen survival and persistence, availability and implementation of disease management measures, and awareness of communities to these NTZDs.

Anthrax, brucellosis, and taeniasis showed statistically significant variation with the perspectives of professionals. This may be explained by the lack of a reporting system for anthrax, brucellosis, and taeniasis in the HMIS, which downgrades the zoonotic and economic importance of these diseases [8]. On the contrary, these diseases are frequently encountered in veterinary practice. The FGD result also indicated a weak inter-sectoral collaboration between human health and livestock sectors in dealing with the prevention and control of NTZDs which is in agreement with previous studies [2, 3, 12, 53–55].

The negative impacts of NTZDs identified in this study include high morbidity and mortality, chronic illness, low production, and productivity, massive economic and social troubles, high cost of treatment, control and prevention programs, and export-import trade restrictions. These were also indicated as major impacts of zoonoses in previous reports [2, 3, 5, 6, 25, 47, 55]. Lack of awareness about NTZDs, free animal movement and illegal trade, close humananimal interaction, feeding habits of raw animal product, illegal slaughtering practices, suitable environment for some vector-borne NTZDs, lack of planned control and prevention strategies, and lack of laboratory infrastructure were considered as the major risk factors and constraints in relation to NTZDs [2, 5, 35, 55].

This study presented some limitations: First, the absence of context specific to one health zoonotic disease prioritization tool could have influenced the prioritization of diseases. In addition, the lack of country-specific data for the majority of the zoonotic diseases has made it difficult to triangulate the results obtained in this study. Second, despite the fact those participants were from diversified disciplines, the number of animal health experts was fewer than human health experts, which may result in bias on professional opinions. In addition, the difference in the background of professionals involved in the study might have influenced the weighing and scoring results of zoonotic diseases. Third, only 15 accessible districts which had a better road infrastructure and health facility were considered in this study which may not infer the non-accessible districts of the region.

Conclusion

The prioritization result of the present study indicated rabies, anthrax, bovine tuberculosis, leishmaniasis, and brucellosis as the top five major zoonotic diseases in the region. In all the zones of the region, rabies was ranked as the first priority disease. Even though anthrax and brucellosis are identified as a priority NTZDs, unfortunately, these were not included in the HMIS reporting system. The importance of BTB as a priority disease was identified across all zones and was equally recognized by both human and animal health professionals, and individuals with different working experience. The presence of the diseases could be amplified by limited commitment and policy dialogues to contain, socio-cultural practices, poor diagnostic capacity, and lack of coordinated control and prevention programs.

Supporting information

S1 Fig. Rating of neglected tropical zoonotic diseases (NTZD) by profession. (TIF)

S2 Fig. Rating of neglected tropical zoonotic diseases (NTZD) by experience of professionals.

(TIF)

S1 File. Questionnaire for professionals. (PDF)

S2 File. FGD discussion points for health professionals (human health and veterinarians). (PDF)

S3 File. Tool for prioritization of NTZD in Tigray region, Northern Ethiopia. (PDF)

S4 File. (XLSX) S5 File. (XLSX) S6 File. (XLSX)

Acknowledgments

The authors express their sincere appreciation to Tigray National Regional State Bureau of Health and study site veterinary and health facilities for assistance with obtaining the consent and facilitation during data collection. We also thankfully acknowledge study participants (experts) for their participation. Finally, we thank Mrs. Rahel HMIS data officer at the Tigray Region Bureau of Health for her unreserved support on providing HMIS data necessary for the study. Finally, we also gratefully acknowledge Dr. Matthew Thomas (Associate Professor, College of Veterinary Medicine, Iowa State University) for his help in editing the English language.

Author Contributions

Conceptualization: Biruk Mekonnen Wolde, Nigus Abebe Shumuye, Habtamu Taddele Menghistu.

Data curation: Tadesse Teferi Mersha.

Formal analysis: Habtamu Taddele Menghistu.

Funding acquisition: Tadesse Teferi Mersha.

- Methodology: Tadesse Teferi Mersha, Nigus Abebe Shumuye, Yisehak Tsegaye Redda, Birhanu Hadush Abera, Habtamu Taddele Menghistu.
- **Supervision:** Nigus Abebe Shumuye, Abrha Bsrat Hailu, Abrahim Hassen Mohammed, Habtamu Taddele Menghistu.

Writing - original draft: Tadesse Teferi Mersha.

Writing – review & editing: Biruk Mekonnen Wolde, Nigus Abebe Shumuye, Abrha Bsrat Hailu, Abrahim Hassen Mohammed, Yisehak Tsegaye Redda, Birhanu Hadush Abera, Habtamu Taddele Menghistu.

References

- Mbugi EV, Katale BZ, Kendall S, Good L, Kibiki GS, Keyyu JD, et al. Tuberculosis cross-species transmission in Tanzania: Towards a One-Health concept. Onderstepoort Journal of Veterinary Research. 2012; 79(2):01–6. https://doi.org/10.4102/ojvr.v79i2.501 PMID: 23327386
- 2. WHO. Investing to overcome the global impact of neglected tropical diseases: third WHO report on neglected tropical diseases 2015: World Health Organization; 2015.

- 3. WHO. The control of neglected zoonotic diseases: A route to poverty alleviation: Report of a Joint WHO/DFID-AHP Meeting, 20 and 21 September 2005, WHO Headquarters, Geneva, with the participation of FAO and OIE. Geneva, Switzerland. 2006.
- Kariuki T, Phillips R, Njenga S, Olesen OF, Klatser PR, Porro R, et al. Research and capacity building for control of neglected tropical diseases: the need for a different approach. PLoS Negl Trop Dis. 2011; 5(5):e1020. https://doi.org/10.1371/journal.pntd.0001020 PMID: 21655352
- FDRE-MoH. Second Edition of National Neglected Tropical Diseases (NTDs) Master Plan (2015/16-2019/20). Addis Ababa, Ethiopia: Ministry of Health, Federal Democratic Republic of Ethiopia; 2016.
- McFadden A, Muellner P, Baljinnyam Z, Vink D, Wilson N. Use of multicriteria risk ranking of zoonotic diseases in a developing country: case study of Mongolia. Zoonoses and Public Health. 2016; 63 (2):138–51. https://doi.org/10.1111/zph.12214 PMID: 26177028
- Grace D, Mutua F, Ochungo P, Kruska R, Jones K, Brierley L, et al. Mapping of poverty and likely zoonoses hotspots. Zoonoses Project 4. Report to the UK Department for International Development. International Livestock Research Institute, Nairobi, Kenya. 2012.
- Menghistu HT, Hailu KT, Shumye NA, Redda YT. Mapping the epidemiological distribution and incidence of major zoonotic diseases in South Tigray, North Wollo and Ab'ala (Afar), Ethiopia. PloS one. 2018; 13(12). https://doi.org/10.1371/journal.pone.0209974 PMID: 30596744
- Deribe K, Cano J, Newport MJ, Golding N, Pullan RL, Sime H, et al. Mapping and modelling the geographical distribution and environmental limits of podoconiosis in Ethiopia. PLoS neglected tropical diseases. 2015; 9(7):e0003946. https://doi.org/10.1371/journal.pntd.0003946 PMID: 26222887
- Sorrell EM, El Azhari M, Maswdeh N, Kornblet S, Standley CJ, Katz RL, et al. Mapping of networks to detect priority zoonoses in Jordan. Frontiers in public health. 2015; 3:219. <u>https://doi.org/10.3389/ fpubh.2015.00219 PMID: 26528460</u>
- Victoria NG, Sargeant JM. A quantitative approach to the prioritization of zoonotic diseases in North America: a health professionals' perspective. PloS one. 2013; 8(8):e72172. <u>https://doi.org/10.1371/journal.pone.0072172 PMID: 23991057</u>
- Pieracci EG, Hall AJ, Gharpure R, Haile A, Walelign E, Deressa A, et al. Prioritizing zoonotic diseases in Ethiopia using a one health approach. One Health. 2016; 2:131–5. <u>https://doi.org/10.1016/j.onehlt.</u> 2016.09.001 PMID: 28220151
- 13. CSA. 2017 human population projection based on the 2007 Census. Addis Ababa, Ethiopia: Central Statistical Agency (CSA), 2017.
- 14. TBFP. GPS data of health facilities and veterinary clinics in Tigray region. Mekelle, Tigray, Ethiopia: Tigray Bureau of Finance and Plan (TBFP) 2018.
- 15. CSA. Population Projection of Ethiopia for All Regions At Wereda Level from 2014–2017. Addis Ababa, Ethiopia: Central Statistical Authority (CSA) of Ethiopia; 2013.
- CSA. Agricultural Sample Survey 2017/18 (2010 E.C.): Report on Livestock and Livestock Characteristics (Private Peasant Holdings), Statistical Bulletin 587 Addis Ababa, Ethiopia: Central Statistical Agency (CSA); 2018 [cited 2020 March 20]. Available from: http://www.csa.gov.et/survey-report/ category/358-eth-agss-2017?download=938:livestock-report-2010-ec-2017.
- 17. Alemu A, Atnafu A, Addis Z, Shiferaw Y, Teklu T, Mathewos B, et al. Soil transmitted helminths and Schistosoma mansoni infections among school children in Zarima town, northwest Ethiopia. BMC infectious diseases. 2016; 11(1):189.
- Kock R, Croft S, Dixon M, Fletcher C, Good L, Guzman J, et al. DFID ZOONOSES REPORT 6 'Prioritising the need for new diagnostics, medicine, vaccines and management practices of zoonoses which have significant impact in the. Royal Veterinary College, Hatfield, website: http://wwwdfidgovuk/r4d/ Output/190316/Defaultaspx. 2012.
- Schelling E, Grace D, Willingham III AL, Randolph T. Research approaches for improved pro-poor control of zoonoses. Food and Nutrition Bulletin. 2007; 28(2_suppl2):S345–S56. https://doi.org/10.1177/ 15648265070282S214 PMID: 17658081
- Ouma EA, Dione MM, Lule PM, Pezo DA, Marshall K, Roesel K, et al. Smallholder pig value chain assessment in Uganda: results from producer focus group discussions and key informant interviews. 2015.
- Speybroeck D, Huffel S. Risk Ranking in Food Chains. In: Carlos Van Peteghem, Chair Scientific Committee FASFC Xavier Van Huffel, Director Staff Direction for Risk Assessment, FASFC Gil Houins CF (ed) symposium SciCom. Brussels. 1–85. 2013.
- Hailessilasie M, Shewit K, Moses K. Serological survey of bovine brucellosis in barka and arado breeds (Bos indicus) of western Tigray, Ethiopia. Preventive Veterinary Medicine. 2010; 94:28–35. https://doi. org/10.1016/j.prevetmed.2009.12.001 PMID: 20034690

- Haftu R, Taddele H, Gugsa G, Kalayou S. Prevalence, bacterial causes, and antimicrobial susceptibility profile of mastitis isolates from cows in large-scale dairy farms of Northern Ethiopia. Trop Anim Health Prod. 2012; 44(7):1765–71. https://doi.org/10.1007/s11250-012-0135-z PMID: 22476790.
- Munyua P, Bitek A, Osoro E, Pieracci EG, Muema J, Mwatondo A, et al. Prioritization of zoonotic diseases in Kenya, 2015. PloS one. 2016; 11(8):e0161576. https://doi.org/10.1371/journal.pone.0161576 PMID: 27557120
- Sekamatte M, Krishnasamy V, Bulage L, Kihembo C, Nantima N, Monje F, et al. Multisectoral prioritization of zoonotic diseases in Uganda, 2017: A One Health perspective. PLoS One. 2018; 13(5): e0196799. https://doi.org/10.1371/journal.pone.0196799 PMID: 29715287
- Takayama N. Rabies: a preventable but incurable disease. Journal of infection and Chemotherapy. 2008; 14(1):8–14. https://doi.org/10.1007/s10156-007-0573-0 PMID: 18297443
- OIE. Frequently asked questions on rabies Paris, France: World Organization for Animal Health (OIE); 2020 [cited 2020 August 15]. Available from: https://www.oie.int/fileadmin/Home/fr/Animal_Health_in_ the_World/docs/pdf/Portail_Rage/QA_Rage_EN.pdf.
- Habtamu TM, Andrew GT, Midekso B, Jamleck B, Getachew G, Abrha B, et al. Free Roaming Dogs and the Communities' Knowledge, Attitude and Practices of Rabies Incidence/Human Exposures: Cases of Selected Settings in Ethiopia and Kenya. Ethiopian Journal of Health Development. 2018; 32 (1):11–20.
- Hadush H, Mekonnen G, Biruk M. Human Rabies Exposure in Central Zone of Tigray, Ethiopia: A Five-Year Retrospective Study. Ethiopian Journal of Veterinary Sciences and Animal Production. 2020; 3 (2):41–9.
- Teklu GG, Hailu TG, Eshetu GR. High incidence of human rabies exposure in northwestern tigray, Ethiopia: A four-year retrospective study. PLoS neglected tropical diseases. 2017; 11(1):e0005271. https:// doi.org/10.1371/journal.pntd.0005271 PMID: 28060935
- **31.** WHO. Human rabies transmitted by dogs: current status of global data, 2015. Weekly Epidemiological Record = Relevé épidémiologique hebdomadaire. 2016;91(2):13–20.
- **32.** WHO. World Organisation for Animal Health, Food and Agriculture Organization, Global Alliance for Rabies Control. Zero by 30: the Global Strategic Plan to end human deaths from dog-mediated rabies by 2030. 2018.
- Bahiru G, Bekele A, Seraw B, Boulanger L, Ali A. Human and animal anthrax in Ethiopia: A retrospective record review 2009–2013. Ethiopian Veterinary Journal. 2016; 20(2):76–85.
- 34. Hooper P. Review of animal health service delivery in the mixed crop-livestock system in Ethiopia. LIVES Working Paper Nairobi, Kenya: International Livestock Research Institute (ILRI); 2016.
- Cook EAJ, de Glanville WA, Thomas LF, Kariuki S, de Clare Bronsvoort BM, Fèvre EM. Working conditions and public health risks in slaughterhouses in western Kenya. BMC Public Health. 2017; 17(1):14. https://doi.org/10.1186/s12889-016-3923-y PMID: 28056885
- Hörmansdorfer S. Bacterial Zoonotic Pathogens as Bioterroristic Agents. Zoonoses-Infections Affecting Humans and Animals: Springer; 2015. p. 1063–75.
- 37. WHO. Anthrax in humans and animals. Geneva, Switzerland: World Health Organization; 2008.
- Ameni G, Tadesse K, Hailu E, Deresse Y, Medhin G, Aseffa A, et al. Transmission of Mycobacterium tuberculosis between farmers and cattle in central Ethiopia. PLoS One. 2013; 8(10):e76891. <u>https://doi.org/10.1371/journal.pone.0076891</u> PMID: 24130804
- Sibhat B, Asmare K, Demissie K, Ayelet G, Mamo G, Ameni G. Bovine tuberculosis in Ethiopia: a systematic review and meta-analysis. Preventive veterinary medicine. 2017; 147:149–57. https://doi.org/ 10.1016/j.prevetmed.2017.09.006 PMID: 29254713
- Zeru F, Romha G, Berhe G, Mamo G, Sisay T, Ameni G. Prevalence of bovine tuberculosis and assessment of Cattle owners awareness on its public health implication in and around Mekelle, Northern Ethiopia. Journal of Veterinary Medicine and Animal Health. 2014; 6(6):159–67.
- Michel AL, Müller B, Van Helden PD. Mycobacterium bovis at the animal–human interface: A problem, or not? Veterinary microbiology. 2010; 140(3–4):371–81. <u>https://doi.org/10.1016/j.vetmic.2009.08.029</u> PMID: 19773134
- Bantie K, Tessema F, Massa D, Tafere Y. Factors associated with visceral leishmaniasis infection in North Gondar Zone, Amhara Region, North West Ethiopia, case control study. Sci J Pub Health. 2014; 2(6):560–8.
- Bsrat A, Berhe M, Gadissa E, Taddele H, Tekle Y, Hagos Y, et al. Serological investigation of visceral Leishmania infection in human and its associated risk factors in Welkait District, Western Tigray, Ethiopia. Parasite epidemiology and control. 2018; 3(1):13–20. <u>https://doi.org/10.1016/j.parepi.2017.10.004</u> PMID: 29774295

- 44. Leta S, Dao THT, Mesele F, Alemayehu G. Visceral leishmaniasis in Ethiopia: an evolving disease. PLoS Negl Trop Dis. 2014; 8(9):e3131. https://doi.org/10.1371/journal.pntd.0003131 PMID: 25188253
- Dawit G, Girma Z, Simenew K. A review on biology, epidemiology and public health significance of leishmaniasis. Journal of Bacteriology and Parasitology. 2013; 4(166):2.
- 46. Kalayou S, Tadelle H, Bsrat A, Abebe N, Haileselassie M, Schallig HD. Serological evidence of Leishmania donovani infection in apparently healthy dogs using direct agglutination test (DAT) and rk39 dipstick tests in Kafta Humera, north-west Ethiopia. Transbound Emerg Dis. 2011; 58(3):255–62. https://doi.org/10.1111/j.1865-1682.2011.01209.x PMID: 21371289.
- **47.** FAO. Guideline for coordination human and animal brucellosis surveillances. Proceedings of animal productions and health conference paper. Rome, Italy: Food and Agriculture Organization of the United Nations; 2003. p. 1–45.
- Tadesse G. Brucellosis seropositivity in animals and humans in Ethiopia: A meta-analysis. PLoS neglected tropical diseases. 2016; 10(10):e0005006. <u>https://doi.org/10.1371/journal.pntd.0005006</u> PMID: 27792776
- Wakene WZ, Mamo G. Review on Epidemiology of Camel and Human Brucellosis in East Africa, Igad Member Countries. Science. 2017; 6(6):109–15.
- Haileselassie M, Kalayou S, Kyule M, Asfaha M, Belihu K. Effect of Brucella infection on reproduction conditions of female breeding cattle and its public health significance in Western Tigray, northern Ethiopia. Veterinary medicine international. 2011;2011. <u>https://doi.org/10.4061/2011/354943</u> PMID: 21822466
- Onyango D, Fascendini M, Wieland B, Ikiror D, Sircely J, Tefera S. One health policy context of Ethiopia, Somalia and Kenya: One Health Units for Humans, Environment, Animals and Livelihoods (HEAL) Project. 2019.
- 52. Birhanu H, Fikru R, Said M, Kidane W, Gebrehiwot T, Hagos A, et al. Epidemiology of Trypanosoma evansi and Trypanosoma vivax in domestic animals from selected districts of Tigray and Afar regions, Northern Ethiopia. Parasites & Vectors. 2015; 8(1):1–11. <u>https://doi.org/10.1186/s13071-015-0818-1</u> PMID: 25889702
- Narrod C, Zinsstag J, Tiongco M. A one health framework for estimating the economic costs of zoonotic diseases on society. EcoHealth. 2012; 9(2):150–62. <u>https://doi.org/10.1007/s10393-012-0747-9</u> PMID: 22395956
- WHO. High-level technical meeting to address health risks at the human-animal ecosystems interfaces: Mexico city, Mexico 15–17 November 2011. 2012.
- 55. World Bank. Zoonotic disease prevention and control, one health, and the role of the World Bank (English) Washington, D.C.: World Bank Group; 2012 [cited 2018 March 5]. Available from: http://documents.worldbank.org/curated/en/529871468331160494/Zoonotic-disease-prevention-and-control-one-health-and-the-role-of-the-World-Bank.