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# Historical Perspective and Risk of Multiple Neglected Tropical Diseases in Coastal Tanzania: Compositional and Contextual Determinants of Disease Risk

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# Abstract

# Background

In the past decade, research on neglected tropical diseases (NTDs) has intensified in response to the need to enhance community participation in health delivery, establish monitoring and surveillance systems, and integrate existing disease-specific treatment programs to control overlapping NTD burdens and detrimental effects. In this paper, we evaluated the geographical distribution of NTDs in coastal Tanzania.

# **Methods and Findings**

We also assessed the collective (compositional and contextual) factors that currently determine risks to multiple NTDs using a cross sectional survey of 1253 individuals in coastal Tanzania. The results show that the effect size in decreasing order of magnitude for nonbinary predictors of NTD risks is as follows: NTD comorbidities > poverty > educational attainment > self-reported household quality of life > ethnicity. The multivariate analysis explained 95% of the variance in the relationship between NTD risks and the theoreticallyrelevant covariates. Compositional (biosocial and sociocultural) factors explained more variance at the neighbourhood level than at the regional level, whereas contextual factors, such as access to health services and household quality, in districts explained a large proportion of variance at the regional level but individually had modest statistical significance, demonstrating the complex interactions between compositional and contextual factors in generating NTD risks. collection and analysis, decision to publish, or preparation of the manuscript.

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### Conclusions

NTD risks were inequitably distributed over geographic space, which has several important policy implications. First, it suggests that localities of high burden of NTDs are likely to diminish within statistical averages at higher (regional or national) levels. Second, it indicates that curative or preventive interventions will become more efficient provided they can be focused on the localities, particularly as populations in these localities are likely to be burdened by several NTDs simultaneously, further increasing the imperative of multi-disease interventions.

### Author Summary

Neglected Tropical Diseases (NTDs) are characterized by their high incidence in lowincome countries, thus maintaining the disastrous poverty-disease-poverty cycle. Apart from poverty, however, little is known of the magnitude of importance of both compositional and contextual factors in creating disease risk at the local level, although this knowledge is critical to disease control and policy action. In this study, we show that the order of importance of both sets of factors is as follows: NTD comorbidities > poverty > educational attainment > self-reported household quality of life > ethnicity.

### Introduction

Neglected tropical diseases (NTDs)—a group of seventeen core debilitating infectious diseases [1] that mutually reinforce (act as both a cause and effect of) poverty—have increasingly been receiving cumulative policy, public health attention globally. Neglected tropical diseases affect more than 1 billion people [2, 3] predominantly poor populations living in tropical and subtropical climates. NTDs are endemic in 149 countries with differing populations, economies, resources, political and legal arrangements, health regulations, traditions, cultures, climates, infrastructure and geographies [4]. Yet, NTDs frequently cluster together geographically and individuals are often simultaneously afflicted with more than one parasite or infection (comorbidities). We are still in the early stages of appreciating the full extent of the comorbidity that occurs when the neglected tropical diseases are superimposed on the "Big Three" (HIV/AIDS, tuberculosis, and malaria). Integrated approaches are useful in addressing both NTDs and their comorbidities. For instance, lymphatic filariasis and malaria (NTD comorbidity) are both transmitted by mosquitoes thus distribution of bed nets leads to a decline in both diseases. The pathogens of NTDs have exceedingly complex life-cycles, population dynamics, infection processes and epidemiologies, causing diverse diseases and pathologies [1]. Although they are biomedically heterogeneous, the commonality of NTDs is evidenced in their persistence and prevalence in people and communities living in poverty and social exclusion. It is estimated that more than 70% of countries and territories that report the presence of neglected tropical diseases are low-income or lower middle-income economies especially in sub-Saharan Africa [4]. In particular, those living in remote areas are most vulnerable to infections, and their biological and sociocultural consequences [5]. Notwithstanding this, even within low income countries, there are age- and sex-specific differentials in the health outcomes induced by NTDs. For instance, many NTDs disproportionately affect women and children in sub-Saharan Africa [2, 6].

According to Hotez and Kamath [7], the United Republic of Tanzania has the third highest prevalence of two NTDs namely Lymphatic filariasis and Trachoma in sub-Saharan Africa. Hitherto, research on NTDs in many sub-Saharan African countries including Tanzania, which is endemic to at least 7 NTDs (lymphatic filariasis, schistosomiasis, soil-transmitted helminthiasis, onchocerciasis, trachoma, rabies, trypanosomiasis); have disproportionately focused on clinical aspects [8] especially the use of mass drug administration. Madon et al. [9]show that, as at 2011, functioning NTD coordination units had been set up at national, regional and district levels throughout the United Republic of Tanzania providing a 53% geographical coverage for preventive chemotherapy (PCT). Despite the national- and sub-national level priorities on understanding and eliminating NTDs and its associated risks, we know far less about multiple and differential exposures of individuals to NTDs based on population composition and contextual attributes. According to Singer and Bulled [10], although the role of tropical disease syndemics in contributing to the health burden of the poor is significant, they have been largely unrecognized and generally neglected in the existing literature. Syndemics of NTDs, a relatively new theoretical approach, involves a critical examination of the adverse morbidity-enhancing interactions among NTDs and between NTDs and other diseases. A basis for syndemic analysis is the attention to underlying social conditions and relationships, their causes and contexts, as well as the natural and anthropogenic environmental factors that facilitate the clustering and interaction of constituent diseases. This paper adopts this approach in assessing the risk of multiple neglected tropical diseases in coastal Tanzania.

The complex relationships between environmental factors and NTD-induced human health outcomes, taking into account multiple pathways and interactions, should be seen in a broader spatial, socio-economic and cultural context. Since NTD distribution varies systematically by individual characteristics and place-based attributes it is imperative that the scientific research community push for a greater understanding of the differentials in health outcomes associated with NTD aetiology and distribution. This gap in the literature is a fundamental motivation for this paper. The aim of this paper is threefold. First, we attempted to provide a brief historical overview of NTD prevalence over time in Tanzania. Secondly, we assessed the risk of exposure to multiple NTDs in coastal Tanzania. Thirdly, we evaluated the magnitudes of compositional and contextual determinants of NTD risks in coastal Tanzania. Given that not all the NTDs reported in Tanzania can be found in the coastal zone, for the 3rd aim we restricted the statistical analysis to NTDs that only prevail in three coastal regions of Tanzania (our study area). The study area is part of the Indian Ocean world (IOW), an arena of primary geo-political importance in eastern Africa. The specific study regions were selected based on the assumption that NTDs are co-endemic clusters in endemic geographical areas. Therefore, we expected to see more clustering in these areas. Besides, although these geographical areas are contiguous they have large wealth inequalities. To our knowledge, this paper is the first to characterise and quantify differential risks of NTDs based on population composition and contextual attributes. Based on the extant literature, the following hypotheses were formulated to guide the study. First and foremost, NTDs are rooted in poverty [2,5]; poorer individuals will be associated with higher risks of exposures to multiple NTDs and by extension, experience higher NTD risks than their relatively affluent counterparts. Second, neighbourhood disadvantage (expressed in the lack of or limited access to health, social, water and sanitation services) is more strongly associated with NTD risks such that individuals living in poorer neighbourhoods will likely experience higher NTD risks than their counterparts living in more affluent neighbourhoods. Lastly, social inequities and disadvantage make women in developing countries more vulnerable to negative health outcomes [2, 5]. Therefore, women are more exposed to multiple NTDs and higher NTD comorbidities and so experience higher NTD risks than their male counterparts.

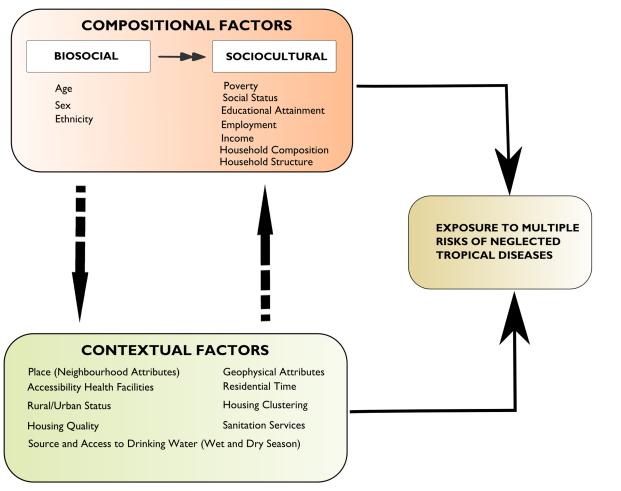
## **Theoretical Context**

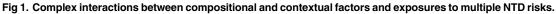
The role of space in shaping health inequities (e.g., differential distribution and by extension differential exposure to and risk of NTDs) is of continuing interest in environmental health geography. In studying the role of space in shaping health outcomes, individual-level (compositional) and place-level (contextual) factors have traditionally been identified [11–15]. Usually, the health outcomes experienced by individuals living proximal to one another are more similar to one another than to those of individuals living in distant neighbourhoods. In theory, three plausible explanations account for this observation. First of all, it may purely be that individuals in the same neighbourhood tend to be more similar to one another than to those in other neighbourhoods in terms of predisposing factors such as age, gender and ethnicity, that is, the *composition* effect [13]. Another viable explanation may be that individuals living in the same neighbourhood are exposed to similar local factors that have impacts on their health outcomes (NTD risks), for example proximity to a river where blackfly proliferate or service provision, that is, the *context* effect [13]. It can also be argued that individuals who live in proximity are more likely to engage in the same types of behavior that may have influences on health outcomes-for example behavior of bathing in rivers among adolescents which are affected by peer pressure-the *collective* effect. Since we live in a complex world, in reality all three elements may be present to varying extents in relation to the distribution of neglected tropical diseases. It is, therefore, imperative to include relevant environmental, behavioural and predisposing factors and to recognise the inherent complexity of composition/context/ collective effects [16].

It is recognized that 'context' and 'place' vary in time and space. Theoretical and empirical approaches (both qualitative and quantitative) require relational notions of space and place that accepts mutually reinforcing and reciprocal relationships between people and place [11]. Further, scale must be included in the analysis of 'contexts' relevant for health. Under this circumstance, place is regarded as complex, socially constructed, unbounded, fluid, and dynamic [11]. It also is multi-scalar, enmeshed in networks, shows social power relations and has cultural meaning [11, 17]. Recent studies also demonstrate that context (e.g. neighbourhoods) is important in determining health outcomes but that compositional factors such as gender, ethnicity, employment status and socio-economic status remain better predictors of inequalities in health [13, 15, 18]. In modelling, the compositional and contextual accounts of health outcomes are usually considered as 'mutually exclusive, competing, and culturally and historically universal' [15]. In fact, from a sustainability perspective, this dichotomous framing is rather problematic. Smyth [15] argues that this supposed difference between people and places, composition and context, is rather artificial. In this study, therefore, we move beyond this polar conceptualization and focus on the collective effect of compositional and contextual attributes. According to Cummins et al. [11], a change from empirical research intended to differentiate between contextual and compositional effects to research that focuses on the processes and interactions occurring between places and people and over time is important for understanding health outcomes (e.g. the distribution of NTDs), and is justified. It is in this milieu that this paper should be understood. In Fig 1 we show how compositional and contextual attributes jointly shape exposures to multiple NTD risks.

In Fig 1, we observe that exposures of individuals to multiple health risks associated with NTDs involve complex and dynamic interplay among biological, environmental, and sociopolitical components spanning multiple time intervals and spatial (geographic) scales. In this context, the question is more about what types, in what places, how they contribute and how they can be addressed.

In this framework, we consider NTD risks and health outcomes as emergent properties of complex interactions between humans and their environment. There are several feedback





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relationships in Fig 1 indicating the multi-factorial nature of both the determinants and the manifestations of NTD health outcomes in coastal populations. For instance, there is a bidirectional linkage between compositional and contextual factors. Similar relationships exist between NTD risks and compositional factors on the one hand and NTD risks and contextual factors, on the other hand. Risks of individuals to NTDs emanate from two mutually-reinforcing factors namely exposure and vulnerability. Predisposing factors to vulnerability include both biosocial and sociocultural dynamics. The latter, together with contextual attributes, also contribute to exposures of individuals to multiple NTD risks. Altogether, this interactivity between variables and processes, which are shaped by multiple factors, warrants a systems approach to addressing exposures and risks to multiple NTDs. A systems approach comprehensively considers all known and measurable aspects of a problem, including feedbacks that cross the boundaries of sub-systems and cut across scales; it acknowledges the nonlinearities and the dynamic nature of underlying processes, uncertainty and surprises [19]. Its potential can be harnessed by policymakers/researchers as they focus more on the social determinants of health when designing NTD interventions (and target interventions).

## Materials and Methods

### Study Area

Tanzania is a coastal country lying between longitude 29° and 49° East and latitude 1° and 12° south of the Equator [20] (Fig 2). The marine waters comprise 64 000 km<sup>2</sup> as territorial waters and 223 000 km<sup>2</sup> as offshore waters (EEZ) [21]. Tanzania's coastline stretches for 800km. It has five coastal regions-Tanga, Pwani, Dar-es-Salaam, Lindi and Mtwara. The five coastal regions cover about 15 percent of the country's total land area and are home to approximately 25 percent of the country's population [22]. According to the 2012 Population and Housing census, the total population was 44,928,923 compared to 12,313,469 in 1967 [23], reflecting an annual growth rate of 2.9 percent. The under 15 age group represented 44.1 percent of the population, with 35.5 percent being in the 15–35 age group, 52.2 percent being in the 15–64 age group, and 3.8 percent being older than 64 [23]. Overall Tanzania on average is sparsely populated with population density of 51 persons per square kilometer, lower significant variation exists across regions. The population density varies from 1 person per square kilometre in arid regions to 51 per square kilometre in the mainland's well-watered highlands to 134 per square kilometre in Zanzibar [24]. The population density for the Dar es Salaam region is 3,133 persons per km<sup>2</sup> (the most densely populated) and that of Lindi is only 13.1 persons per  $\text{km}^2$  [23]. This suggests wide disparities in population density across regions. This study specifically focuses on Dar-es-Salaam, Pwani and Tanga. The 3 coastal regions selected for analysis were chosen for two main reasons. First, the three regions are of historical significance to the Indian Ocean World project. Second, these regions were selected because of the 5 regions, they are the most ethnically diverse (that is, representative of the different geographical locations) and thus, had better prospects of providing heterogeneous survey responses. Dar es Salaam is the capital of the Dar

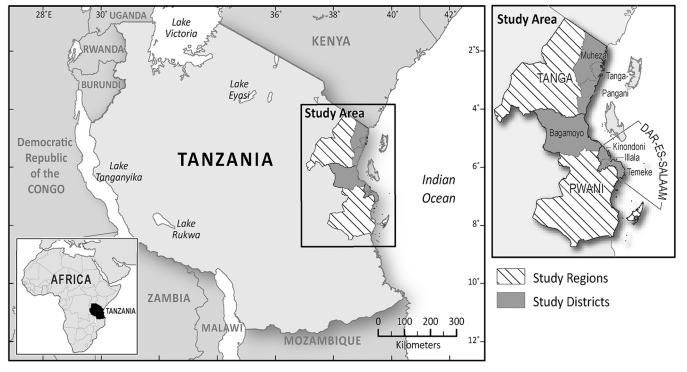


Fig 2. Map of Tanzania showing the study areas.

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es Salaam Region, which is one of Tanzania's 26 administrative regions. The Dar es Salaam Region consists of three local government areas or administrative districts: Kinondoni to the north, Ilala in the center of the region, and Temeke to the south. Pwani (coast) is the 21<sup>st</sup> most densely populated region. It is bordered to the north by the Tanga Region, to the east by the Dar es Salaam Region and the Indian Ocean, to the south by the Lindi Region, and to the west by the Morogoro Region. Tanga region has a population of 2,045,205 [24]. It is bordered by Kenya and Kilimanjaro Region to the north; Manyara Region to the west; and Morogoro and Pwani regions to the south. Its eastern border is formed by the Indian Ocean.

Seven districts namely Kinondoni, Temeke and Illala (in Dar-es-Salaam region), Bagamoyo (in Pwani region), and Tanga Town, Muheza and Pangani (in Tanga region) were considered in this study. According to the Tanzania National Bureau of Statistics [23], Kinondoni municipality has a population of 1,775,049 and density of 3302.8 inhabitants per km<sup>2</sup>. Illala municipality has a population of 1,220,611 and density of 3344.4 inhabitants/km<sup>2</sup> and Temeke municipality has population of 1,368,881 and density of 1878.5 inhabitants/km<sup>2</sup>. The population and density of Bagamoyo are 311,740 and 36.8 inhabitants/km<sup>2</sup> respectively whereas that of Muheza is 204,461 and 136.5 inhabitants/km<sup>2</sup>, respectively [23]. Tanga town has a population of 54,025 and density of 30.8 inhabitants per km<sup>2</sup>. The numbers of participations from each of seven districts were as follows Kinondoni (360), Temeke (101), Illala (140), Bagamoyo (301), Tanga Town (129), Muheza (101) and Pangani (121).

### **Ethics Statement**

The study was approved by the Research Ethics Board of the University of Western Ontario, Canada. Research approval was also granted by the Commission on Science and Technology (COSTECH) in Tanzania. Written informed consent was obtained from participant prior to commencement of the study.

## **Data Collection**

In addressing the first aim (a brief historical perspective on NTD prevalence and distribution across Tanzania) we conducted a literature search on the NTDs that have been reported in Tanzania since the 19<sup>th</sup> century. Using the country- and disease-specific query, we searched the Global Infectious Diseases and Epidemiology Network (GIDEON) database (<u>http://www.gideononline.com/</u>) and the global NTD (GNTD) database (<u>http://www.gntd.org/</u>). We further obtained secondary data from the Tanzania National Institute for Medical Research (NIMR). For triangulation, we interviewed experts from the national office of the Tanzania NTD Control Program (TZNTDCP) as well as the Ministry of Health and Social Welfare's (MOHSW) integrated NTD program.

In addressing the second objective (assessment of the risk of exposure to multiple NTDs along the coastline of Tanzania) we examined most-at-risk populations in coastal Tanzania and NTD profiles at the community level during the last five and ten years. This information was complemented with data from a cross sectional survey.

The third aim was achieved by conducting a cross sectional survey with 1253 individuals in three regions (Dar es Salaam, Tanga, and Pwani) along the coastline of Tanzania. The survey data were collected between March and September 2013. The study population included male (606) and female (647) participants between the ages of 18 and 70+ years. The study used multistage sampling to obtain representative estimates of the population of residents of the three regions. Within each region, a list of villages based on the 2012 Population and Housing Census was divided further into households. The list of villages was also divided into clusters

ensuring that each cluster would provide adequate numbers of eligible respondents to be included in the survey. This approach both corrects for sampling bias and weights the cases to match census percentages of males and females of various age groups and by ethnicity. The enumeration areas (EAs) and their total number of households were listed geographically by urban and rural areas. Where EAs did not include the minimum number of households, geo-graphically adjacent EAs were amalgamated to yield sufficient households. This provided the frame for selecting the clusters to be included in the survey according to a stratified systematic sampling technique in which the probability for the selection of any cluster was proportional to its size. A sampling interval was calculated by dividing the total number households by the number of clusters. A random number between 1 and the sampling interval was computer generated. The EA in which the random number and then progressively until the 20 (urban) and 15 (rural) clusters were identified. These clusters made up the sample for the survey. Households were randomly selected from these clusters for interview.

### Measures

**Outcome variable.** Based on whether the specific NTD had been reported for at least two consecutive years in the past 5 years or not in the communities of the most-at-risk populations, six focal NTDs were considered in this paper: cholera, schistosomiasis (bilharzia), onchocerciasis (river blindness), trachoma (granular conjunctivitis), hookworm and whipworm. For the six NTDs, each of the 1253 individuals in the survey were either assigned a value of 1 (exposed) or 0 (unexposed) depending on whether the specific NTD had been reported in the neighbourhood at least once in the past 5 years or not. Neighbourhood is used to describe an area surrounding a local institution patronized by residents, such as a health centre. It can also be defined by a political ward or precinct. The concept of neighborhood includes both geographic (place-oriented) and social (people-oriented) components. City planning departments often designate neighbourhood boundaries along census tract boundaries. And, in fact, community residents quite frequently have a very different mental map of their neighbourhood than the officially designated neighborhood areas used by planners and policymakers. In the context of this paper, neighbourhoods refer specifically to administrative wards. From the foregoing, we emphasize that the statistical relationship is between survey data on social determinants and reported cases of certain NTDs at the neighbourhood level. The same point applies to the use of the term co-morbidities, which we are assessing at a geographic (and not individual) level. It is in this context that the term *exposure* should be understood.

The exposure to multiple NTDs was calculated as the algebraic sum of the exposure to each of the six NTDs. Several assumptions underlie this exposure summation technique. It assumes homogeneous exposure within the same neighbourhood; thus, likely intra-neighbourhood differentials in exposure to NTDs were considered as random. It also assumes independence of the exposure to each of the six NTDs. It also assumes that the magnitude of an adverse NTD-related health effect will be proportional to the sum of the ratios of the exposures. If these assumptions are incorrect, over- or under-estimation of the actual multiple-NTD risk could result.

**NTD Risk = Function (Exposure, Vulnerabilities).** Vulnerabilities of individuals to NTDs were assigned based on their responses to 10 objective health measures in the survey. These included whether or not respondents had been diagnosed with malaria in the past 12 months, diagnosed with HIV, diagnosed with pneumonia, hepatitis, skin conditions or tuber-culosis in the past 5 years. Also, respondents were asked whether they had been diagnosed with heart disease, cancer, hypertension or diabetes. In each case, respondents who answered in the affirmative were each assigned a value of 1 otherwise 0 was assigned. We hypothesized that

higher number of affirmatives corresponds to more vulnerability of the respondent. Apart from their use as vulnerability indicators, the 10 objective health measures also constituted an additive score for NTD comorbidity.

For each NTD:

Risk =  $\sum$  Respondents Exposure scores  $\times$  Respondents Vulnerability scores

Assuming uniform weight of each NTD based on severity and importance, then

Total NTD Risk = Function (Risk<sub>cholera</sub>, Risk<sub>schistosomiasis</sub>, Risk<sub>onchocerciasis</sub>, Risk<sub>trachoma</sub>, Risk<sub>hookworm</sub>, Risk<sub>whipworm</sub>).

The set of NTD risks was then constituted into a composite score. Cronbach's alpha test of reliability computed for the six NTD risk items was 0.83 showing good internal consistency. Total Exposure NTD Risk was calculated using principal component and factor analyses. Only one factor had an Eigen value >1 and was retained (all the components loaded on a single construct). This factor explained approximately 90% of the variance in the 6 NTD risk scores.

**Compositional and contextual factors.** Theoretically relevant compositional (biosocial and sociocultural) factors used in this study include gender, age of respondents, ethnicity, poverty, educational attainment, marital status, and employment status. The contextual factors include household quality, exposure to single or multiple NTDs, NTD comorbidities, access to health services, rural/urban status, access to good drinking water during the dry season, and access to good drinking water during the wet season.

### Statistical Analysis

Inferential and multivariate techniques were applied to examine associations between NTD risks and theoretically relevant compositional and contextual factors variables using STATA 13SE software. The Ordinary Least Squares technique was employed for the analysis. Analyses were preceded by diagnostic tests to establish whether variables met the assumptions of the regression model. Bivariate analysis was initially performed to examine zero-order correlations between the NTD risks and theoretically-relevant independent variables all of which were significant and in the expected direction, thus supporting good construct validity. Further, multivariate models were estimated to explore the net effects of the predictor variables using the stepwise selection approach. The relative quality of candidate multivariate models (both OLS and ordinal logit based on categorized NTD risks) was tested using Akaike Information Criterion (AIC). After comparison, the best model was chosen based on parsimony, our working hypotheses, and strength of evidence. For analytical purposes, the unstandardized regression coefficients were estimated although standardized regression coefficients were used as indices of effect sizes of the non-binary predictors. Positive coefficients for any of the predictors indicate higher NTD risk, while negative coefficients show lower NTD risk. The ordinary least squares (OLS) regression models in this study are built under the assumption of independence of subjects, but the cross-sectional survey has a hierarchical structure with respondents nested within survey clusters, which could potentially bias the standard errors. STATA 13 (StataCorp, College Station, TX, USA) SE, which has the capacity to address this problem, is used by imposing on our models a 'cluster' variable, that is, the identification numbers of respondents at the cluster level. This in turn adjusts the standard errors (SE) producing statistically robust parameter estimates.

#### Results

# Historical Overview on the Distribution and Epidemiology of Neglected Tropical Diseases (NTDs) in Tanzania

Table 1 shows the various NTDs reported in Tanzania since the 19<sup>th</sup> Century. So far, buruli ulcer, chagas disease and dracuncunliasis have never been reported in Tanzania. However, cholera, schistosomiasis (bilharzia), onchocerciasis (river blindness), trachoma (granular conjunctivitis), hookworm, whipworm, dengue fever, human African trypanosomiasis, leischmaniasis, leprosy and roundworm have been reported at least once during the last 200 years in Tanzania. Cholera outbreaks and cases surpass all cases and outbreaks of other NTDs in Tanzania. In general, prevalence and incidence of all NTDs have declined in the past 5 years in Tanzania. However, there are regional variations in the NTD type, number of cases and frequency of occurrence of the various NTDs.

### Sample Characteristics

Each individual in the survey sample was simultaneously exposed to at least two NTDs. No individual was simultaneously exposed to four or more NTDs as shown in <u>Table 2</u>.

About 75% of 1253 respondents were simultaneously exposed to three NTDs and 25% were exposed to 2 NTDs. Males and females were evenly distributed in terms of simultaneous exposures to 3 NTDs although females (54%) were more exposed to 2 NTDs than their male counterparts. NTD comorbidities was associated with gender ( $x^2(8) = 19.9265$ , pr = 0.011)). Also, there were regional variations in NTD comorbidities ( $x^2(16) = 37.4640$ , pr = 0.002)) and exposures to multiple NTDs ( $x^2(2) = 1.2e+03$ , pr = 0.000)). Although there are no statistically significant differences in NTD comorbidities according to poverty status there are differences in exposures to multiple NTDs by poverty status ( $x^2(1) = 33.5928$ , pr = 0.000)). In this study, the reported comorbidities were malaria (76%), hypertension (25%), tuberculosis (25%), HIV (10%), skin diseases (9%), pneumonia (6%), heart disease (5%), cholera (4%) diabetes (3%), hepatitis (2%) and cancer (1%). Based on the sample, malaria was the most common comorbidity. Based on the responses in the survey sample, inferential statistics (chi-square) did not find any statistically significant relationships between region of residence and each of the following diseases: hepatitis, skin diseases, pneumonia, HIV status, hypertension, cancer and heart disease. This means that these diseases were independent of region of residence. However, region was not independent of malaria ( $x^2(2) = 1.3e+03$ , Pr = 0.000), tuberculosis ( $x^2(2) = 5.9731$ , Pr = 0.045, cholera ( $x^2(2) = 19.1780$ , Pr = 0.000) and diabetes ( $x^2(2) = 11.7872$ , Pr = 0.003).

Exposures to multiple NTDs do not vary according to age even though NTD comorbidities vary by age of respondents. Both NTD comorbidities and exposures to multiple NTDs vary by educational attainment of respondents. In coastal areas of Tanzania, risks of cholera, hookworm and whipworm are higher in Dar es Salaam region than in Tanga and Pwani regions. Risks of schistosomiasis and trachoma are higher in Pwani and Tanga regions than in Dar es Salaam region. Risks of onchocerciasis are higher in Tanga region than in Pwani and Dar es Salaam regions. Based on the magnitude of Cramér's V, the strength of association of theoretically-relevant covariates and exposure to multiple NTDs in decreasing order of magnitude is as follows: region of residence > residential locality > ethnicity > educational attainment > residential time > poverty access to health services > NTD comorbidities > self-rated housing quality.

### **Bivariate Analysis**

<u>Table 3</u> shows zero-order relationships between NTD risks and theoretically relevant covariates. All biosocial factors were significant predictors of NTD risk. Older individuals and

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ž	S/ Name of NTD N		Rep	Reported in Tanzania	zania				Prevalen	Prevalence since 1 <sup>st</sup> reported	eported	ŋ ŋ ŋ	Current prevalence
		Yes No	Yes No Year (ref.)	Region	District	No. of cases in 1 <sup>st</sup> year	Year	Cases (ref.)*	Region	District	Place of residence Sex		SDE
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<u> </u>	Buruli ulcer	>											
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ъ	Cholera	>	1821 (1)	1 <sup>st</sup> records			1977	1671 (4)					
							1977, 1978, 1981, 1983	(5, 6, 7)	TZ Mainland, Dar es Salaam				
							1986	(8)	Mara				
			1974 (2)	TZ Mainland		10 (2)	1992	18526 (2)					
							1996	1,100 (9)	Kigoma				
			1978 (3)	TZ Zanzibar	Tumbatu —Unguja	411 (3)	1997	40'249 (2, 10)	DSM	Illala, Kinondoni			
							1998		Mwanza, Tukwa				
							1999	11,855 (11)					
							2002- 2006	~2000 (2)	DSM, Dodoma, Kigoma, Lindi, Mbeya, Morogoro, Mtwara, Pwani, Tanga, Rukwa, Mwanza, Mara, Shinyanga, s				
							2006	8965 (2)	DSM				
								1507 (2)	Ruvuma				
								1030 (2)	Kigoma				
								315 (3)	Pemba, Unguja				
							2007	1092 (7)	Zanzibar				
							2008	500 (12, 13, 14, 15, 16,	Arusha, Dar es Salaam, Kilimanajaro,				
								17)	Mara, Rukwa				

(Continued)

# PLOS | NEGLECTED TROPICAL DISEASES

Image: constant in the state in th	Vertor         Region         District         No.of in 1         Vertor         Region         District           1 <th>-</th> <th>S/ Name of NTD</th> <th>Ē</th> <th>Reported in Tanzania</th> <th>zania</th> <th></th> <th></th> <th></th> <th>Prevalen</th> <th>Prevalence since 1<sup>st</sup> reported</th> <th>reported</th> <th></th> <th></th> <th>Current</th>	-	S/ Name of NTD	Ē	Reported in Tanzania	zania				Prevalen	Prevalence since 1 <sup>st</sup> reported	reported			Current
Yes No Yes (ref (ref )         Rayotic (ref )         Rayotic (ref )         Report (ref)         Report (ref)         Report (ref)         Report (ref )         Report (ref)         R	Yee No Year (rol)         Rayin         Destrict         Rayin         Rayin </th <th></th> <th>prevalence Status</th>														prevalence Status
Notational         Notation         Notation         Notational<	1         2000         211 (1)         1         1           2001         <			Yes No Year (re		District	No. of cases in 1 <sup>st</sup> year	Year	Cases (ref.)*	Region	District	Place of r	esidence	Sex	
2000         211 (1)         Dare estatam           Party Line         2000         2010         Dare estatam           Dary Line         V         1823 (2)         American         2003         Manercan           Dary Line         V         1823 (2)         Zanchar         2013         Manercan         2013         Manercan           Dary Line         V         1823 (2)         Zanchar         2013         2015         Penha           Dary Line         2001         Manercan         2013         2012         Manercan           Dary Line         2003         Manercan         2013         2012         Manercan           Dary Line         2004         Manercan         2013         2012         Manercan           Dary Line         2014         X         X         X         X         X         X           Unmark Line         2001         Manercan         2013         2012         Manercan         Manercan           Dary Line         2012         Manercan         1892         X         X         X         X         X         X         X         X         X         X         X         X         X         X         X <t< th=""><th>2000         211 (1)         Dare Salaan           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010</th><th></th><th></th><th></th><th></th><th></th><th>(iei.)</th><th></th><th></th><th></th><th></th><th>Rural</th><th>Urban</th><th>Σ</th><th>1</th></t<>	2000         211 (1)         Dare Salaan           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010         2010         2010           2010         2010         2010						(iei.)					Rural	Urban	Σ	1
343(1)         7.130 (1)         343(1)         7.130 (2)         343(1)         7.130 (2)           Degue level         v         1870 (28)         Zarzbar         201         Marazaza           Degue level         v         1870 (28)         Zarzbar         201         Marazaza           Degue level         v         1870 (28)         Zarzbar         201         Penua           Degue level         v         1870 (28)         Zarzbar         201         Penua           Degue level         v         1910 (6)         Kimanjaro         Mora         77% (27)         Penua           Pubman Africa         1920 (28)         Kimanjaro         Mora         77% (27)         Penua           Pubma Africa         1920 (28)         Kimanjaro         Mora         77% (27)         Penua           Pubma Africa         Notein         77% (27)         Penua         Penua         Penua           Pubma Africa         Notein         77% (27)         Penua         Penua         Penua           Pubma Africa         Notein         77% (73)         Penua         Penua         Penua           Pubma Africa         Notein         77% (73)         Penua         Penua         Penua	Math Alian       90,01       Math Alian         Degue fever       v       187,02       Math Alian         Landa Alian       200       Plana         Degue fever       v       187,02       Math Alian         Landa Alian       201       202       Math Alian         Luma Alian       187,02       2012       188,02       Math Alian         Luma Alian       187,02       2013       2013       188,02       Math Alian         Luma Alian       187,02       2014       188,02       Math Alian       188,02       Math Alian         Luma Alian       Math Alian       110,01       Math Alian       110,01       Math Alian       188,02       Math Alian         Luma Alian       Math Alian       110,01       Math Alian       110,01       Math Alian       188,02       Math Alian         Luma Alian       Math Alian       110,01       Math Alian       110,01       Math Alian       110,01       Math Alian         Luma Alian       Luma       192,01       Luma       Luma       Math Alian       110,01							2009– 2010	211 (18)	Dar es Salaam					
For the constraint of the	60 (2)         Manza           Brouce fever         182 (22)         Manza           Brouce fever         182 (22)         Manza           Brouce fever         187 (26)         Zarzibar         201         Brouce           Brouce fever         187 (26)         Zarzibar         201         Brouce           Brouce fever         197 (26)         Zarzibar         201         Brouce           Human Aritean         1987 (26)         Kilmanjao         Note         Note           Brouce fever         1990 (6)         Western         1990 (7)         Shortean           France fever         1910 (6)         Western         (7)         Shortean         Shortean           France fever         1910 (6)         Western         (7)         Shortean         Shortean           France fever         1910 (6)         Western         (7)         Shortean         Shortean           France fever         191         (7)         Shortean         Shortean         Shortean           France fever         191         (7)         Shortean         Shortean         Shortean           France fever         192         192         Shortean         Shorean         Shortean <t< td=""><td></td><td></td><td></td><td></td><td></td><td></td><td></td><td>3454(19, 20)</td><td>Tanga</td><td></td><td></td><td></td><td></td><td></td></t<>								3454(19, 20)	Tanga					
No.         Signation         Sign	Propue forwer       V       1823 (25)       Zanzibar       2013       Ri-Max         Dergue forwer       V       1823 (25)       Zanzibar       2013       Ri-Max         Dergue forwer       V       1823 (25)       Zanzibar       21% (27)       Ri-Max         Dergue forwer       V       1823 (25)       Zanzibar       21% (27)       Ri-Max         Dergue forwer       V       1820 (25)       Zanzibar       21% (27)       Ri-Max         Dergue forwer       2008 (5)       Kimanijaro       Mostin       71% (27)       Rimanijaro         Dergue forwer       2008 (5)       Kimanijaro       Mostin       71% (27)       Rimanijaro       Rimanijaro         Dergue forwer       2008 (5)       Kimanijaro       Mostin       71% (27)       Rimanijaro       Rimanijaro         Dergue forwer       Rimanijaro       Northerm       71% (27)       Rimanijaro       Rimanijaro       Rimanijaro         Dergue forwer       Rimanijaro       Northerm       71% (27)       Rimanijaro       Rimanijaro         Southerm       Southerm       Southerm       Rimanijaro       Rimanijaro       Rimanijaro         Southerm       Rimanijaro       Rimanijaro       Rimanijaro       Rimanijaro								60 (21)	Mwanza					
Dengue fever         V         1823 (26)         Zarxibar         Co13         Ruiwa           1970 (26)         Zarxibar         300 (25)         Ruiwa           1970 (26)         Zarxibar         137% (27)         Pemba           1970 (26)         Kimanjaro         Nostim         71%         Pemba           Umma African         900 (5)         Kimanjaro         Nostim         71%         Pemba           Umma African         900 (5)         Withmanian         Nostim         71%         Pemba           Umma African         910 (6)         Wostem         Nostim         1921         Ruiwa           V         1910 (6)         Wostem         71%         71%         Ruiwa           V         1910 (6)         Wostem         71%         Ruiwa         Ruiwa           V         1921 (75)         71%         71%         Ruiwa         Ruiwa           V         1922 (75)         1	Oncurrent         V         1823 (26)         Carabar         Cond         Ethicat         Ethicat<								3 (22, 23, 24)	Mara					
Dengue fever         1         12/3         2arzbar         7.7%	Dengue fore         V         182 (36)         Zanzlau         T.7% (27)         Imba           187 (32)         2anzlau         Mostia         71(5)         1.8% (27)         Impa           Huma African         2008 (5)         Kimanjaro         Noshi         71(5)         1.8% (27)         Impa           Huma African         2008 (5)         Kimanjaro         Noshi         71(5)         P.4%         Noshi           Huma African         1910 (5)         Westernin         Noshi         71(5)         P.4%         Noshi           Figh         Mosternin         Mosternin         Mosternin         Noshi         Noshi         Noshi         Noshi           Figh         Mosternin         Mosternin         Mosternin         Text         Noshi         Noshi         Noshi           Figh         Mosternin         Text         Text         Text         Noshi         Noshi         Noshi         Noshi           Figh         Mosternin         Text         Text         Text         Noshi         Noshi         Noshi         Noshi           Figh         Figh         Text         Text         Noshi         Noshi         Noshi         Noshi         Noshi           Figh </td <td></td> <td></td> <td></td> <td></td> <td></td> <td></td> <td>2013</td> <td>300 (25)</td> <td>Rukwa</td> <td></td> <td></td> <td></td> <td></td> <td></td>							2013	300 (25)	Rukwa					
1670 (26)         Zanzibar         13% (27)         Inga           2008 (5)         Kimanjaro         0oshi         71(5)         1           Human African         1900 (5)         Kimanjaro         00shi         8           Typanosomissis         1900 (5)         Western         1910 (5)         8         1           V         1910 (6)         Western         1910 (5)         8         1         1           V         1910 (6)         Western         1910 (5)         8         1         1           Southern         Southern         1910 (5)         1         1         1         1           I         1910 (5)         Northern         1         1         1         1         1           I         1910 (5)         1         1         1         1         1         1           I         1         1         1         1         1         1         1         1         1         1           I         1         1         1         1         1         1         1         1         1         1         1         1         1         1         1         1         1         1 <td>Interview         Interview         <t< td=""><td><del></del></td><td>Dengue fever</td><td></td><td></td><td></td><td></td><td></td><td>7.7% (27)</td><td>Pemba</td><td></td><td></td><td></td><td></td><td></td></t<></td>	Interview         Interview <t< td=""><td><del></del></td><td>Dengue fever</td><td></td><td></td><td></td><td></td><td></td><td>7.7% (27)</td><td>Pemba</td><td></td><td></td><td></td><td></td><td></td></t<>	<del></del>	Dengue fever						7.7% (27)	Pemba					
Iman African         2008 (s)         Kitmanjaro         T (s)           Human African         1902 (c3)         Kestminic         1910 (s)         Kestminic	Iman Artical       2006 (5)       Kimanjon       7 (5)         Huma Artical       1902 (5)       1902 (5)       1902 (5)         Typanosonials       1902 (5)       1902 (5)       1902 (5)         V       1002 (5)       Nortem       1921 (7)       Nortem         V       1002 (5)       Nortem       1921 (7)       Nortem         V       1002 (5)       Nortem       1921 (7)       Nortem         V       Nortem       1921 (7)       Nortem       Nortem         V       Nortem       1922 (7)       Nortem       Nortem         V       Nortem       1922 (7)       Nortem       Nortem         V       Nortem       1922 (7)       Nortem       Nortem         Nortem       Nortem       1922 (7)       Nortem       Nortem         Nortem			1870 (26					1.8% (27)						
Human African Typanosomiasis       1902 (29) Gambiense Gambiense Southern Southern       1910 (6) (7,9)       Marvan (7,9)         Yanga Kana Highands       1910 (6) (7,9)       1910 (6) (7,9)       1910 (7) (7,9)       Ninyanga Kana (7,9)       Marvan (7,9)         Yanga Kana Highands       192 (7,9)       192 (7,9)       Indi (7,9)       Ninyanga Kana (9)       Marvan (7,9)         Yanga Kana Highands       192 (7,9)       192 (7,9)       Indi (7,9)       Marvan (7,9)       Marvan (7,9)         Yanga Kana Highands       192 (7,9)       192 (7,9)       Marvan (7,9)       Marvan (7,9)       Marvan (7,9)         Yanga Kana Highands       Yanga Kana (7,9)       192 (7,9)       192 (7,9)       Morvan (7,9)       Morvan (7,9)         Leichmaizet       Yanga Kana (7,9)       192 (7,9)       192 (7,9)       Morvan (7,9)       Morvan (7,9)         Leichmaizet       Yanga Kana (7,9)       192 (7,9)       192 (7,9)       Morvan (7,9)       Morvan (7,9)         Leichmaizet       Yanga Kana (7,9)       192 (7,9)       192 (7,9)       Morvan (7,9)       Morvan (7,9)         Leichmaizet       Yanga Kana (7,9)       192 (7,9)       192 (7,9)       Morvan (7,9)       Morvan (7,9)	Human African       190 (c)       Worksmin       191 (c)       Shinyanga       Mawa         Typanosoniasis       Banbienes       Typanosoniasis       Banbienes			2008 (5)			71 (5)								
V       1910 (6)       Western, Northern       191- (7.9)       Sinivanga       Maswa         Sunda       Northern       192- (7.9)       Inividence       Northern       N	Value       1910 (c) and and souther s	10	Human African Trypanosomiasis	1902 (26 Gambier	() Ise										
Lindi       Kitwa         1925       Lindi       Kitwa         1930, 1	Lindi       Kilva         1325       1925       100       Kilva         1930, 1				es			1919– 1921 (7,9)		Shinyanga	Maswa				
1925       Lindi       Kilwa         1930,       1957,       1957,       Kilwa         1957,       1957,       1957,       1957,       Kilwa         1951,       1925       1922       Kiloma, Taboa       Kaulu         1921,       1922       119       Koma, Ruka       Koma,         1924,       1925       2119       Koma,       Koma,         1924,       1925       2119       Koma,       Koma,         1924,       1925       Kiloma, Rukwa,       Kasulu,       Kasulu,         Listohaniasi       V       1992,       Kiloma, Rukwa,       Kasulu,         Listohaniasi       V       1992,       Kiloma, Rukwa,       Kasulu,	1925       Lindi       Kilva         1930       1930       1930       1930         1931       1930       1930       1930       1930         1931       1932       1932       1933       1933         1932       1935       2119       1933       1934         1935       1935       2119       1934       1934         1935       1935       2119       1934       1934         1934       1934       1934       1934       1934         1934       1934       1934       1934       1934         1934       1934       1934       1934       1934       1934         1934       1934       1934       1934       1934       1934       1934         1934       1934       1934       1934       1934       1934       1934       1934				)					Lindi	Kilwa	Luangwa			
Kigoma, Tabora         1922       (7)         (7)       1925-         (9)       1946         (8)       1979-       6000 (29)         (1992       1992-       1904, Ausha, Hukwa         Leischmaniasis       1       1992       8000 (29)         Leischmaniasis       1       1992       8000 (29)       Tabora, Arusha, 1992	Kigoma, Tabora         1922       Kigoma, Tabora         1925       1925         1926       1925         1926       1926         1926       1926         1926       1926         1926       1926         1926       1926         1926       1926         1926       1926         1926       1926         1926       Kigoma, Arusha,         1927       1922         1928       Kigoma, Rukwa         Leischmaniasis       1 1946         J       1922         1932       Kigoma, Rukwa							1925 (9), 1930, 1957, (9)		Lindi	Kilwa	Matandu			
1922       1925         (7)       1925         (7)       1925         (7)       1925         (7)       1925         (8)       (8)         (8)       (9)         (1925)       1929         (7)       1929         (8)       (9)         (9)       1920         (1926)       Kigoma, Rukwa	1922       1925         (7)       1925         (7)       1925         (7)       1925         (8)       (8)         (9)       1946         (8)       1979         (9)       1929         (7)       1929         (8)       1929         (9)       1924         (8)       1924         (9)       1924         (9)       1924         (9)       1924         (9)       1924         (9)       1924         (9)       1932         (1932)       1932         (1932)       1932									Kigoma, Tabora	Kasulu				
1925-       2119         1946       1946         (8)       (9)         1979-       6000 (29)       Tabora, Arusha, 1992         1992       1992       Kigoma, Rukwa	1925-       2119         1946       (8)         (8)       (9)         (9)       <							1922 (7)			Ikoma				
Leischmaniasis V 1964 (34) Kilimanjaro 1	Leischmaniasis V 1964 (34) Kilimanjaro 1							1925– 1946 (8)	2119		Ikoma				
V 1964 (34) Kilimanjaro     1	Leischmaniasis 🗸 1964 (34) Kilimanjaro 1							1979– 1992	6000 (29)	Tabora, Arusha, Kigoma, Rukwa	Kibondo, Kasulu, Sikonge, Mponda, Urambo				
		6	Leischmaniasis	,		0	-								

# PLOS NEGLECTED TROPICAL DISEASES

Ц Ц	Table 1. (Continued)												
∣ ∕∂ z	S/ Name of NTD N		Rep	Reported in Tanzania	ania				Prevale	Prevalence since 1 <sup>st</sup> reported	eported		Current prevalence
		Yes No	Yes No Year (ref.)	Region	District	No. of cases in 1 <sup>st</sup> year	Year	Cases (ref.)*	Region	District	Place of residence	ssidence Sex	Status
						(rei.)					Rural	Urban M F	
~	Leprosy										>	55% 45%	0.85/10000 (35)
ω	Dracuncunliasis	>											
o	Lymphatic filariasis (elephantiasis)	>	1911 (36)	Lindi		ı	2008	3.3%					33/1000
				Mtwara			2008	3.3%					
				DSM			2008	3.3%					
				Tanga			2008	3.3%					
				Pwani			2008	3.3%					
				Morogoro			2008						
							1990 (37)	1.1%	Pemba				
							2000 (38)	28.5% males	Hale				
							2001	7.2%		Kwahani		>	
							1957 (39)	70% adult males		Kilwa			
				Lake Victoria Zone		ı	2008	ı					
				Lake Nyasa Kyela, zone	Kyela,	ı	2008	4.5%	Mbeya	Kyela,	>		
					Rungwe		2008	4.5%		Rungwe	>		
							2011 (40)	62.9% school children	Morogoro	Mvomero			
10	<ul> <li>Schistosomiasis haematobium &amp; mansoni (snail fever)</li> </ul>	>	1895 (41,42)	Lake Victoria Zone		50%	1903 (43)		Zanzibar				51.5% (48)
							1961 (44)	203 [64%]	Mwanza	Usagara			
							1966 (45)	975 [42%]	Tabora	Bukumbi		664 311	
							1967 (46)	391 [65.2%]	Mwanza	Ukerewe			
													(Continued)

# PLOS NEGLECTED TROPICAL DISEASES

z		Rep(	Reported in Tan:	Tanzania				Prevalei	Prevalence since 1 <sup>st</sup> reported	reported			Current prevalence
	Yes No	Yes No Year (ref.)	Region	District	<b>μ</b>	Year	Cases (ref.)*	Region	District	Place of residence	nce Sex	×	Status
					(ret.)					Rural U	Urban M	L	
						1969 (46)	614 [61.2%]	Zanzibar	Unguja				
						1983 (46)	730 [21%]	Morogoro	Ifakara				
						1985 (46)	483 [19.3%]	DSM					
						1997 (46)	2415 [67%]	Zanzibar	Pemba				
						2005 (46)	393 [62%]	Kilimanjaro	Mwanga				
						2008 (46)	1129 [78%]	Mwanza	llemela, Ukerewe				
						2009 (46)	311 [27.2%]	Tanga	Lushoto	Umba			
						2012 (47)	43 [18.1%]	Pwani	Mafia				
<ol> <li>Onchocerciasis (river blindness)</li> </ol>	>	1958	Tanga	Usambara	ı	1984 (52)	22.7%	Tanga	Usambara				
						1990 (51)	58.6%	Morogoro	Mahenge	Bwakira			
							31.9%	Ruvuma					
							22.4%	Tanga	Amani				
							22.8%	Mbeya	Tukuyu				
						1991 (53)	%09	Central Tanzania					
						2000 (50)	120	Dodoma	Kongwa		32	88	25.4% (49)
							143		Mpwapwa		19	124	
							50	Dodoma	Dodoma		12	38	
							215		Manyoni		50	165	
12 Trachoma (eye infection)	>					2004 (56)	13.9%	Kilimanjaro	Rombo	Kahe			

Table 1. (Continued)											
S/ Name of NTD N	Re	Reported in Tanzania	zania				Prevalen	Prevalence since 1 <sup>st</sup> reported	eported		Current prevalence
	Yes No Year (ref.)	) Region	District	No. of cases in 1 <sup>st</sup> year	Year	Cases (ref.)*	Region	District	Place of residence	ssidence Sex	- Status
				(rei.)					Rural	Urban M F	1
					2006 (55)	44%	Dodoma, Singida, Arusha, Mwanza, Shinyanga, Mtwara, Lindi, Coast and Tanga				
					2006 (54)	119 [9%]	Dodoma	Kongwa	>		
					2013 (57)	20.4%					20.4%
13 Hookworm;	√ 1982–2007 URT (68)	07 URT		20- 50%	1997 (60)	95%	Pemba				
					2000 (61)	72.5% school children	Mafia Island				
					2002 (62)	73.8% male 77.1% female	Tanga				
					2004 (63)	7.7% children	Unguja Island				
					2005 (58)	3145 (40%)	Shinyanga, Mwanza, Tabora and Mara				
					2007 (64)	32.9% pregnant women	Pemba Island				
					2007 (65)	11.9% school children	Zanzibar				
					2007 (66)	21.6% school children	Zanzibar				
					2009	548 [16.2%]	Zanzibar				
					2010 (59)	152 [38%]	Mwanza	Sengerema	Sengerema Nyamatongo	oĝ	

(Continued)

# PLOS | NEGLECTED TROPICAL DISEASES

		•		5						eported			Current prevalence
 -	Yes No Year (ref.)		Region	District	No. of cases in 1 <sup>st</sup> year	Year	Cases (ref.)*	Region	District	Place of residence	sidence Sex		Status
					(rei.)					Rural	Urban M	L	
						2011 (40)	24.7% school children	Morogoro	Mvomero				
14 Roundworm $$	1977	1977 (69)	URT		40%								
	1982- (68)	1982–2007 URT (68)	URT		20- 50%	1997 (60)	72%	Pemba					
						2009	548 [8%]	Zanzibar					
15 Whipworm 🗸	1982- (68)	1982–2007 URT (68)	URT		20 50%	1997 (60)	<b>96</b> %	Pemba					
						2009 (67)	548 [62.8%]	Zanzibar					

Multiple Neglected Tropical Diseases in Coastal Tanzania

Variables	At least 2 NTDs	More than 2 NTDs	Chi-square & Cramér's V
Compositional Attributes			
Biosocial factors	%	%	
Age			
18–35	23.0	77.0	chi2(3) = 1.6642 Pr = 0.645 Cramér's V = 0.0364
36–50	22.9	77.1	
51–65	25.7	74.3	
More than 65	27.7	72.3	
Sex (Gender)			
Male	22.9	77.1	chi2(1) = 0.6513 Pr = 0.420 Cramér's V = -0.0228
Female	24.9	75.1	
Ethnicity			
Zaramo	65.3	34.7	chi2(2) = 292.1794 Pr = 0.000 Cramér's V = 0.4829
Sambaa	5.3	94.7	
Others	15.2	84.8	
Socio-cultural factors			
Poverty			
Poor	19.9	80.1	chi2(1) = 33.5928 Pr = 0.000 Cramér's V = -0.1637
Non-poor	36.1	63.9	
Educational Attainment			
No education	38.3	61.7	chi2(3) = 100.1362 Pr = 0.000 Cramér's V = 0.2827
Primary	33.7	66.3	
Secondary	14.9	85.1	
Tertiary	5.4	94.6	
Employment			
Unemployed	28.3	71.7	chi2(1) = 1.0168 Pr = 0.313 Cramér's V = 0.0285
Employed	23.6	76.4	
Marital Status			
Single	19.5	80.6	chi2(1) = 7.4346 Pr = 0.006 Cramér's V = -0.0770
Married	26.4	73.7	
Contextual Attributes			
Region of Residence			
Dar-es-Salaam	0.0	100.0	chi2(2) = 1.2e+03 Pr = 0.000 Cramér's V = 0.9978
Pwani	99.7	0.3	
Tanga	0.0	100.0	
Self-reported household quality of life relative to others			
The Worst	37.8	62.2	chi2(4) = 13.2716 Pr = 0.010 Cramér's V = 0.1029
Among the Worst	30.0	70.1	
About the Same	23.7	76.3	
Better	17.3	82.7	

(Continued)

Variables	At least 2 NTDs	More than 2 NTDs	Chi-square & Cramér's V
Best in the Community	19.4	80.6	
Residential Locality			
Rural	52.4	47.7	chi2(1) = 381.2196 Pr = 0.000 Cramér's V = -0.5516
Urban	4.4	95.6	
Residential Time			
Up to 5 years	13.7	86.3	chi2(3) = 76.4820 Pr = 0.000 Cramér's V = 0.2471
Up to 10 years	19.5	80.5	
Up to 15 years	17.6	82.4	
20 or more years	38.2	61.8	
Access to Health Services			
No access	16.5	83.5	chi2(1) = 24.1026 Pr = 0.000 Cramér's V = -0.1387
Access	28.7	71.4	
NTD Comorbidities			
0	31.6	68.4	chi2(5) = 19.7724 Pr = 0.001 Cramér's V = 0.1257
1	17.9	82.1	
2	27.2	72.8	
3	24.2	75.8	
4	25	75	
5 or more	13.3	86.7	

#### Table 2. (Continued)

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females were associated with higher NTD risk scores. Age squared was statistically significant indicating possibly that the relationship between age and NTD risks was not linear. Each of the sociocultural factors was a significant predictor of NTD risks except employment (<u>Table 3</u>). Higher NTD comorbidities, higher exposures to multiple NTDs and higher poverty levels were each associated with higher NTD risks. Except residential time and self-reported household quality of life, all contextual factors including sources of drinking water in the wet and dry seasons, rural/urban status and region of residence were significant predictors of NTD risks.

### **Multivariate Analysis**

Table 4 is a nested model that shows the multivariate relationship between NTD risks on the one hand and compositional and contextual factors, on the other hand. The statistically significant zero-order relationship between NTD risks, age, sex and ethnicity remains in the biosocial model in the multivariate analysis. The biosocial model explains only 4% of the variance in the relationship between NTD risks and theoretically relevant covariates. Females and older respondents were associated with higher NTD risks. When the model is adjusted for sociocultural factors, the original zero-order relationship between poverty and NTD risks unexpectedly disappears. Self-reported household quality of life relative to others was a significant predictor of NTD risks in the sociocultural model unlike educational attainment and marital status. The sociocultural model explains about 19% of the variance in the relationship between NTD risks and theoretically relevant covariates.

Variables	Coefficient	Robust Std. Error	95% Confidence Interval
Compositional Attributes			
Biosocial factors			
Age	0.458***	0.090	-0.212-0.094
Sex (Gender)	0.588**	0.056	-0.202 0.019
Ethnicity	0.320**	0.106	0.111 0.529
Age Squared	0.102***	0.020	0.062 0.143
Socio-cultural factors			
Poverty	0.715***	0.056	-0.826-0.603
Educational Attainment	0.489***	0.027	0.436 0.542
Log income	0.369***	0.023	0.323 0.414
Employment	-0.109	0.106	-0.318 0.099
Type of current dwelling	-0.199***	0.017	-0.234–0.164
Marital Status	-0.237***	0.058	-0.352-0.121
Contextual factors			
Source of Drinking Water in Dry Season	0.249*	0.124	0.005 0.492
Source of Drinking Water in Wet Season	-0.345***	0.055	-0.453-0.236
Rural/Urban Status	-1.042***	0.168	-1.372-0.713
Residential Time	0.031	0.021	-0.271–0.187
Region of Residence	-1.111***	0.070	-1.109 0.170
Self-reported household quality of life relative to others	-0.096	0.128	-0.346 0.157
Access to Health Services	-0.447*	0.178	0.145 0.373
NTD Multiple Exposures	1.586***	0.166	1.260 1.911
NTD Comorbidities	2.765***	0.032	2.702 2.827

Table 3. Bivariate relationships of risk of multiple exposures to neglected tropical diseases and
explanatory variables (n = 1253) in coastal Tanzania.

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In the place and neighbourhood model when both compositional and contextual factors (collective effect) are taken into account some interesting results emerge. For instance, the relationship between poverty and NTD risks which disappeared in the sociocultural model reappears. The statistical significance of the relationship between ethnicity and NTD risks, which reduced in the sociocultural model, also becomes stronger in the neighbourhood model. In the sociocultural model, educational attainment was not a significant predictor of NTD risks unlike in the neighbourhood model. Employment status became statistically insignificant when collective effect is controlled. Access to health services, residential locality and NTD comorbidities were significant predictors of NTD risks when collective effect was accounted for in Table 3 unlike self-reported household quality of life and sources of drinking water in wet and dry seasons. Based on the standardised regression coefficients of non-binary predictors in the place and neighbourhood model, the effect size in decreasing order of magnitude is as follows: NTD morbidities > educational attainment > self-reported household quality of life > ethnicity. The place and neighbourhood model in the multivariate analysis explains 72% of the variance in the relationship between NTD risks and theoretically relevant covariates.

Poorer, uneducated, unemployed individuals had higher NTD risk scores compared to their relatively affluent, educated and employed counterparts. This implies hypothesis 1, which suggests that poorer individuals will be associated with higher exposures to multiple NTDs and by extension experience higher NTD risks than their less poor counterparts cannot be rejected.

		Compo	ositiona	I Factors			Contextua	al factors	
		1: Bios factors	ocial	Model 2: cultural		N	Model 3: leighbourh		ors
	Coef.	Beta Coef.	Std. Err.	Coef.	Beta Coef.	Std. Err.	Coef.	Beta Coef.	Std. Err.
Intercept	1.13**	-	0.43	1.46	-	0.69	0.19	-	0.19
Gender	0.68***	0.11	0.17	0.72***	0.12	0.18	0.03	0.005	0.04
Age	0.51***	0.15	0.09	0.56***	0.17	0.09	0.04Ψ	0.013	0.03
Ethnicity	0.35**	0.13	0.10	0.29*	0.07	0.10	0.21***	0.055	0.02
Poverty				0.35	0.05	0.21	0.11*	0.015	0.05
Educational Attainment				0.18	0.05	0.12	0.05***	0.14	0.03
Marital Status				-0.23	-0.12	0.18	0.03	0.004	0.04
Employment Status				-0.75*	-0.06	0.35	0.11	0.096	0.08
Self-reported household quality of life relative to others							0.03	0.066	0.03
Access to Health Services							-0.29***	-0.046	0.04
Residential Locality							-0.69***	-0.109	0.05
Source of Drinking Water in Wet Season							-0.02	-0.005	0.02
Source of Drinking Water in Dry Season							0.01	0.003	0.01
NTD Comorbidities							2.74**	0.960	0.03
R <sup>2</sup>	0.04			0.23			0.95		

Table 4. Compositional and contextual determinants of risk of neglected tropical diseases (n = 1253) in coastal Tanzania.

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Individuals living in poorer neighbourhoods with limited access to health, social, water and sanitation services had higher NTD risk scores than their counterparts living in less-deprived neighbourhoods implying that hypothesis 2 cannot also be rejected. However, on disaggregating the results by gender, we did not find any evidence to support the third hypothesis which posits that women, by dint of social disadvantage, are more exposed to multiple NTDs and higher NTD comorbidities and so experience higher NTD risks than their male counterparts. In the females' model, poverty was not even a significant predictor of NTD risks. Only, ethnicity, access to health services, rural/urban status and NTD comorbidities were significant predictors of NTD risks for women. In the males model however, poverty, age, ethnicity, educational attainment, employment status, access to health services and NTD comorbidities were significant predictors of NTD risks. Poorer older males without access to health services in rural areas had higher NTD risks compared to less poor younger males with access to health services living in urban areas.

### Discussion

In this paper, we present a historical overview on the distribution of NTDs in Tanzania during the last two centuries. We also examined the effect sizes of the compositional and contextual determinants of NTD risks with emphasis on coastal Tanzania.

We found regional variations in the type, number of cases and frequency of occurrence of the various NTDs. This is expected as a defining characteristic of most NTDs is their locality [5, 13]. This implies that NTD-induced morbidity and mortality may vary considerably geographically due to different local factors [5]. This result is not entirely surprising given regional and national policy initiatives and clinical interventions on NTDs in Tanzania. To date, Tanzania's NTD program has distributed over 81 million NTD treatments to approximately 34 million people with USAID support [25]. Tanzania has also established an extensive monitoring and evaluation system, including a national NTD database, to gauge the national program's progress towards NTD control and elimination. With support from USAID, tablet computers are being used to document the burden of trachoma. Also, real-time results of the data collected each day are instantly available online for quick analysis and decision-making. Furthermore, Tanzania has integrated data collection for lymphatic filariasis [26, 27, 28], soiltransmitted helminthes, and schistosomiasis as is a cost effective mechanism to monitor the impact of NTD control activities.

The finding that NTD risks are unequally distributed over geographic space has several important implications. First, it suggests that localities of high burden of NTDs are likely to diminish within statistical averages at higher (regional or national) levels. Second, it presupposes that curative or preventive interventions will become more efficient provided they can be focused on the localities, particularly as populations at these locations are likely to be burdened by several NTDs at the same time. In this sense, exposure and comorbidities offer valuable opportunities for integrated control of the various NTDs in Tanzania. Also, from a sustainability perspective it is imperative to continue targeting the most affected populations in order to reduce the underlying health inequalities that may be driving NTD prevalence among disadvantaged groups [29]. In fact, the mass drug administration approach in Tanzania appears to be doing this. For some, the treatments go to a more focal level depending on disease prevalence (e.g., STH, schistosomiasis, onchocerciasis), while for others (trachoma, lymphatic filariasis) the health district is the implementation unit.

This view is reinforced by the positive relationship (direct) between poverty and NTD risks. Furthermore, poor neighbourhoods, which are characterised by the lack of or limited access to health, social, water and sanitation services, were associated with higher NTD risks. These results are consistent with previous scholarly work on NTDs. For instance, Aagaard-Hansen and Chaignat [5] investigated inequities and social determinants of NTDs and found that of all the predictors in their study, poverty (lack of purchasing power) is the only factor having recognized relationship to all 13 NTDs. They proposed two main mechanisms by which poverty mutually reinforces NTDs. They explained that poverty is a structural social determinant and is intrinsically connected to the intermediate determinants of water, sanitation, housing and clustering. Furthermore, poverty may actually be a consequence of some of the NTDs (for example buruli ulcer, dengue fever, human African trypanosomiasis, leishmaniasis and lymphatic filariasis)-either as a result of very expensive treatment [30], or indirectly through loss of labour capability [5]. In terms of housing, there are clear relationships between housing quality, access to water and sanitation services, clustering and neighbourhood quality of life all of which contribute to differential exposure to NTDs. For instance, the defining characteristics of poor neighbourhoods include low quality housing, clustering and limited access to water, and sanitation and health services [30-33].

All respondents were simultaneously exposed to at least 2 NTDs and there were regional and poverty-related differentials in exposure to NTDs. This finding is consistent with previous studies on NTDs. According to Blas and Kurup [34], the same level of exposure may have different effects on different socioeconomic groups, depending on their social, cultural and economic environments and cumulative life course factors. In this situation, clustering of risk

factors in some population groups, such as social exclusion, low income, overcrowded housing and poor access to health services, may be as important as the individual exposure itself [34].

Source of drinking water in the wet season, access to health services and good housing quality were each negatively (inversely) associated with NTD risks. The fact that poorer, uneducated, unemployed individuals had higher NTD risk scores compared to their relatively affluent, educated and employed counterparts likely suggests a social gradient in NTD risks in coastal Tanzania. The social gradient of NTD risks could possibly emanate from multiple interacting factors, including socio-environmental risk factors, social determinants, comorbid conditions, general health status, health-seeking behaviours and access to NTD health care services.

Exposures to multiple NTDs and NTD comorbidities (co-occurrence of one or more diseases or disorders in an individual) were both positively associated with NTD risks. Similarly, age was robustly associated with higher risks of NTDs. It is argued that comorbidity is related to poorer health outcomes, more complex disease management, and increased health care costs [35]. According to Degenhardt et al. [36] several hypotheses exist concerning the reasons why comorbidity (e.g., in the context of NTDs) might occur, including that: (a) there is a causal relationship between the comorbidities; and (c) that the relationship is spurious (artefactual), resulting from factors such as the methods with which the sample was selected. Comorbidity potentially has implications for theories of aetiology and distribution of neglected tropical diseases. For instance, if comorbidity arises because different health problems (e.g. malaria, pneumonia, etc.) share the same risk factors, then interventions addressing these risk factors should reduce the prevalence of these multiple problems [36].

We did not find evidence that women are more exposed to multiple NTDs and have higher NTD comorbidities and so experience higher NTD risks than their male counterparts. However, several studies highlight gender differentials in exposure to NTDs and risks of NTDs. For instance, Aagaard-Hansen and Chaignat [5] suggest there is substantial difference in morbidity and mortality rates for males and females by neglected tropical disease. Cattand et al. [37] also observed that, for human African trypanosomiasis, men are affected at nearly twice the rate of women. Thus, males are disproportionately affected by human African trypanosomiasis and schistosomiasis due to exposure, whereas women suffer more from leprosy (stigma) and trachoma (blindness). The relationship between ethnicity and NTD risks was robust and remained significant even when compositional and contextual factors were accounted for in the multivariate model. This result supports studies which show association between prevalence of dracunculiasis and particular ethnic groups [38]. Ethnic differences may be attributable to different sociocultural and behavioural practices. For instance, people of diverse cultural backgrounds often make different attributions of illness, health, disease, symptoms and treatment [39]. Afrocentric societies usually make spiritual attributions to ill-health compared to Eurocentric societies [37]. In this sense, indigenous beliefs and practices, which are rooted in specific cultures, might affect health outcomes (NTD risks) and interactions with the NTD prevention and treatment services provided in the health care setting.

This study is not without limitations. First, it focused exclusively on coastal Tanzania although NTDs transcend this geographic region. Perhaps, future research could elucidate NTD risks in both coastal and non-coastal regions in order to obtain a much more comprehensive representation of NTD risks across Tanzania. This study is partly based on a cross-sectional survey, which misses the temporal dimension of NTD risks. Given that NTD risks may vary systematically with time it is imperative to design future studies that take into account both the spatial and temporal aspects of NTD risks.

### Conclusion

The study also presented a brief historical perspective on the distribution of NTDs in Tanzania during the last two centuries. This study also assessed exposure to multiple NTD risks using a cross sectional of 1253 individuals in coastal Tanzania. Both compositional and contextual factors act in complex ways to give rise to NTD risks. In particular, low socioeconomic status, poor neighbourhoods, age, NTD comorbidities and exposures to multiple NTDs are robustly associated with an increased risk of NTDs in coastal Tanzania. The link between poverty and NTD risks is persistent and remains even when compositional and contextual factors are adjusted. Based on the standardised regression coefficients of non-binary predictors in the place and neighbourhood model, the effect size in decreasing order of magnitude (importance) is as follows: NTD morbidities > educational attainment > self-reported household quality of life > ethnicity. On the whole, based on the findings, contextual factors are more important practically than compositional factors in terms of relative contribution to NTDs risks. Contextual factors cumulatively explained 72% of the variance in NTD risks whereas compositional (biosocial and sociocultural) factors jointly explained only 23% of the variance. Given the plethora of sociopolitical, economic and cultural factors that culminate in inequities in NTD exposures, comorbidities and risks between poor and less deprived individuals and groups an integrated approach to addressing NTD risks is warranted.

## **Supporting Information**

**S1 Text. List of references in** <u>Table 1</u>. (DOCX)

**S1 Checklist. STROBE statement-checklist of items included in our study.** (DOC)

## **Author Contributions**

Conceived and designed the experiments: FAA IL RQ. Performed the experiments: FAA GC RC HH RQ. Analyzed the data: FAA IL GC HH. Contributed reagents/materials/analysis tools: FAA IL GC RC RQ. Wrote the paper: FAA IL RQ GC RC HH.

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