Spatial Analysis of Land Cover Determinants of Malaria Incidence in the Ashanti Region, Ghana

Anne Caroline Krefis1,*, Norbert Georg Schwarz1, Bernard Nkrumah3, Samuel Acquah3, Wibke Loag1, Jens Oldeland5, Nimako Sarpong3, Yaw Adu-Sarkodie4, Ulrich Ranft2, Jürgen May1

1 Infectious Disease Epidemiology, Bernhard Nocht Institute for Tropical Medicine, Hamburg, Germany, 2 IUF - Leibniz Research Institute for Environmental Medicine, Heinrich Heine University Düsseldorf, Düsseldorf, Germany, 3 Kumasi Centre for Collaborative Research in Tropical Medicine, Kumasi, Ghana, 4 School of Medical Sciences, Kwame Nkrumah University of Science and Technology, Kumasi, Ghana, 5 Biocentre Klein Flottbek and Botanical Garden, University of Hamburg, Hamburg, Germany

Abstract

Malaria belongs to the infectious diseases with the highest morbidity and mortality worldwide. As a vector-borne disease malaria distribution is strongly influenced by environmental factors. The aim of this study was to investigate the association between malaria risk and different land cover classes by using high-resolution multispectral Ikonos images and Poisson regression analyses. The association of malaria incidence with land cover around 12 villages in the Ashanti Region, Ghana, was assessed in 1,988 children <15 years of age. The median malaria incidence was 85.7 per 1,000 inhabitants and year (range 28.4–272.7). Swampy areas and banana/plantain production in the proximity of villages were strong predictors of a high malaria incidence. An increase of 10% of swampy area coverage in the 2 km radius around a village led to a 43% higher incidence (relative risk [RR] = 1.43, p < 0.001). Each 10% increase of area with banana/plantain production around a village tripled the risk for malaria (RR = 3.25, p < 0.001). An increase in forested area of 10% was associated with a 47% decrease of malaria incidence (RR = 0.53, p = 0.029). Distinct cultivation in the proximity of homesteads was associated with childhood malaria in a rural area in Ghana. The analyses demonstrate the usefulness of satellite images for the prediction of malaria endemicity. Thus, planning and monitoring of malaria control measures should be assisted by models based on geographic information systems.

Citation: Krefis AC, Schwarz NG, Nkrumah B, Acquah S, Loag W, et al. (2011) Spatial Analysis of Land Cover Determinants of Malaria Incidence in the Ashanti Region, Ghana. PLoS ONE 6(3): e17905. doi:10.1371/journal.pone.0017905

Materials and Methods

Introduction

With 250 million estimated malaria cases in 2008 and one million deaths malaria is the most common vector-borne infectious disease with Sub-Saharan Africa carrying most of the burden. In regions of stable transmission children <5 years of age are at highest risk of becoming symptomatic after infection with malaria parasites. The causal protozoon Plasmodium falciparum is transmitted from person to person through the bite of adult female Anopheles mosquitoes [1,2].

Transmission and prevalence of vector-borne diseases such as malaria are highly influenced by spatial and temporal changes in the environment as described during the last 20 years by geographic information systems (GIS) and remotely sensed (RS) data [3,4]. Studies mapping potential mosquito habitats, transmission risk, or disease prevalence have been performed in Africa [5–8], South and Central America [9,10], and Asia [11,12]. However, analyses of the direct correlation between environment and malaria are rare.

Adult vector abundance is positively associated with the availability of aquatic habitats necessary for the deposition of eggs, and the areas with highest malaria risk are often found within just a few hundred meters of such larval habitats [13,14]. It has been suggested that extensive cultivation of maize might influence the larval development of mosquitoes, pupation success, and size of adults in the vicinity [15].

Recent studies from Kenya have shown that highland habitats created by deforestation or cultivation of natural swamps were associated with preferred breeding habitats [16,17].

In Ghana, where the present study was conducted, malaria is prevalent during the entire year and accounts for about 32–42% of all outpatient admissions and for the major in-patient causes of death [18]. The main malaria vectors are mosquitoes of the Anopheles gambiae complex and A. funestus [19].

The aim of the study was to investigate the association between malaria incidence and different classes of land cover that potentially influence the malaria vector abundance as well as human population density. High spatial resolution satellite images as well as statistical modeling was used to assess the influence of land cover classes and the human population at risk on the malaria incidence (per year and 1,000 inhabitants) in children <15 years of age in an area of high endemicity. This information might be of importance to the understanding of environmental determinants of malaria transmission heterogeneity at a micro-geographical scale.

Ethics Statement

Aims and principles of the study were explained in detail to participants and informed consent was obtained by signature or...
thumb print by the caregiver. The study design and the informed
consent form were approved by the Committee on Human
Research, Publications, and Ethics, School of Medical Sciences,
Kwame Nkrumah University of Science and Technology, Kumasi,
Ghana.

Study area and data collection

The hospital-based survey was accomplished at the Child
Welfare Clinic and the Pediatric Ward of the Agogo Presbyterian
Hospital (APH), Asante Akim North District, Ashanti Region,
Ghana. The study area was restricted to the 12 study villages
Agogo, Hwidiem, Akutuase, Amantena, Wioso, Domeabra,
Juansa, Kyekyebiase, Nyaboo, Obenimase, Patriensah and
Pekyerekye and their 2 km surrounding areas (Figure 1). The
total area of our study side covered approximately 170 km². For
more detailed information about the study area see Krefis et al.
2010 [20].

All children <15 years of age visiting the Child Welfare Clinic
of the APH and with permanent residency in one the 12 study
villages were examined for malaria (criterion: fever $\geq 37.5$°C and
positive for asexual \textit{P. falciparum} parasitaemia with $>0$ parasites/
$\mu$L) during the study interval of 18 months (end of May 2007 to
November 2008). Parasite examination was done according to
quality-controlled standardised procedures described elsewhere.

**Figure 1. Map of the 12 included study villages.** Merger of two satellite images (Ikonos) depicting an area with 12 study villages in the Asante
Akim North District, Ashanti Region, central Ghana, West Africa. Areas with a radius of 2 km surrounding the study villages, which were analysed by
supervised maximum likelihood classification, are coloured.

doi:10.1371/journal.pone.0017905.g001
Malaria cases presented within 21 days after the initial malaria diagnosis were considered as a relapse and not counted as a new case.

For the calculation of incidences the population size, the admission rate, the proportion recruited, and the proportion of the population seeking health care in the study hospital were considered. Population figures were taken from the population census 2004 with the estimate that 42% of the individuals were <15 years of age [22]. The proportion of people from each village attending the APH was assessed by a community survey on health seeking behaviour that was carried out in 2007 and the denominator/reference population for the calculation of incidence was corrected for these proportions [23]. After comparing the study population with the hospital admission records it was estimated that 70% of all individuals (range: 67% to 72%) admitted to the hospital were included on average into the study and therefore the reference population was likewise corrected for this factor. Finally, annual malaria incidences per 1,000 children <15 years of age were computed for each of the villages. The human population density per village was computed by using population census and village area data (Table 1).

Mapping land cover classes using remote sensing

In order to map the land cover classes around each village, we acquired two multispectral Ikonos images with 4-meter spatial resolution and four broad spectral bands (wavelengths: blue, 0.45–0.52 μm; green, 0.52–0.59 μm; red, 0.62–0.68 μm; and near-infrared (NIR), 0.77–0.86 μm), along with one panchromatic band with 1 pixel/m. Images were acquired on May 4, 2009, and on November 26, 2009 (it was not possible to obtain two contemporaneous images of high quality from the area due to weather, cloud condition, and other acquisition difficulties).

All pre-processing steps were carried out using ENVI 4.4 (ITTVIS, 2009). For easier computation the images were divided into subsets, each covering one or two village areas. For each subset a Normalized Difference Vegetation Index (NDVI = \([\text{NIR} - \text{red}] / [\text{NIR} + \text{red}]\) image was calculated, which is a commonly used measure of vegetation productivity [24]. Beside the spectral domain, the spatial domain was also considered by calculating a set of textural measures based on a grey level co-occurrence matrix in order to improve the classification [25,26]. Different textural measures (contrast, homogeneity, angular second moment, variance, mean, dissimilarity, entropy, and correlation) were received by moving several windows of different pixel areas (3×3 to 15×15) over the image, leading to a new textural image for each measure. The optimal window size was determined by using a confusion matrix to assess the accuracy of the solely texture based classification [25]. Afterwards, the textural images were combined with the NDVI image and the four spectral bands for further analysis.

In March 2010, field sampling of different land cover classes was conducted by the direct inspection of 490 points randomly selected in the vicinity of the 12 study villages. We marked the points using a Garmin eTrex®H Global Positioning System (GPS) and took notes and photographs on the dominant vegetation or crop type.

By using the ENVI software, these reference areas were digitised as regions of interest and were used to represent one of the following land cover classes: banana or plantain, cacao, palm trees, oranges, swampy area, water, deforested area and roads, built-up areas (houses), and forest. Classes describing the crops “banana/plantain”, “cacao”, “palm trees” producing palm oil fruits, and “oranges” were mostly mixed fields but dominated by one of these crops, respectively. Either the presence of a river or stream nearby or near the ground agricultural crops (such as eggplants, maize, tomatoes, pepper), which was mostly cultivated in the vicinity, characterised the combined variable “swampy area”. “Water” was characterised by a river, stream or lake. “Deforested area” was

| Village   | Village areaa | Total populationb | Population densityc | Proportion with hospital accessd | Population study groupe | Malaria casesf | Incidencea |
|-----------|---------------|-------------------|---------------------|--------------------------------|-------------------------|----------------|------------|
| Agogo     | 5.12          | 13559             | 2648                | 90%                            | 3588                    | 1463           | 271.9      |
| Akutuase  | 0.61          | 1692              | 2774                | 43%                            | 214                     | 9              | 28.1       |
| Amantena  | 0.27          | 890               | 3296                | 55%                            | 144                     | 21             | 97.3       |
| Domeabra  | 1.33          | 3509              | 2638                | 42%                            | 433                     | 73             | 112.3      |
| Hwidiem   | 1.08          | 1402              | 1298                | 95%                            | 392                     | 147            | 250.3      |
| Juansa    | 1.27          | 3992              | 3143                | 40%                            | 469                     | 52             | 73.8       |
| Kekyeklesi | 0.54        | 1801              | 3335                | 46%                            | 244                     | 28             | 76.6       |
| Nyaboo    | 1.02          | 1582              | 1551                | 46%                            | 214                     | 28             | 87.2       |
| Obenimase | 0.51          | 1096              | 2149                | 37%                            | 119                     | 15             | 83.9       |
| Patriensah| 0.54          | 4463              | 8265                | 38%                            | 499                     | 92             | 123.0      |
| Pekyereky | 0.37          | 1692              | 4573                | 45%                            | 224                     | 27             | 80.4       |
| Wioso     | 0.34          | 1783              | 5244                | 52%                            | 273                     | 33             | 80.7       |

*a per km².
*b Population according to the national census 2004 [21].
*c Population of children <15 years of age estimated as a proportion of 42% of the population counted at national census 2004 [22] and an additional proportion of 70% due to inclusion into the study at hospital admittance and by taking into account hospital access.
*d Study period from May 2007 to November 2008 (18 months).
*e Incidence in children <15 years (per year and 1,000 children <15 years).
*f Formula: incidence = 1,000*cases/total_population*0.42*0.70*hospital_access*1.5.

doi:10.1371/journal.pone.0017905.t001
characterised by burned, grassy or bushy underground or open spaces; additionally we assigned roads within this class. “Forest” referred to areas with dense tree cover with a closed canopy.

It was not possible to get images completely free of clouds. Therefore, two additional classes, one for clouds and one for the shadow of a cloud were generated to mask out those particular areas.

All combined bands were classified using a supervised maximum likelihood classifier. Therefore, a random subset of 70% of the pixels for each of the classes was chosen for a basic analysis (“training data”) and 30% were used for assessment of accuracy (“validation data”). In the post-classification process, we applied a majority/minority analysis for generalisation of the classification image to minimise “salt and pepper effects”, a term which describes the existence of dark pixels in bright regions and bright pixels in dark regions, usually causing noise in the validation procedure. Validation of the accuracy of the post-processed classification image was based on the overall accuracy computed from the confusion matrix. The maximum likelihood classification is usually considered to be satisfiable when the overall accuracy is higher than 85%. The final image was transferred to ArcGIS version 9.3, developed by Environmental System Research Institute (ESRI, 2008).

Taking into account that adult mosquitoes remain generally up to 2 km of their breeding side [27–29] a radius of 2 km around each village was created and the percentage of various land cover classes in each radius was computed. Due to the particular size of the village Agogo an oval-shaped radius was used (Figure 2). In order to test the validity of the analyses additional radii of 0.5 km, 1 km, and 1.5 km around each village were used.

Analyses and statistical modeling

The quantitative assessment of associations between proportional land cover and the incidence of malaria was done by Poisson regression analyses with adjustment for overdispersion (STATA/SE software, version 10; Stata Corp LP, College Station, TX). By using Spearman rank correlation we calculated the cross correlation between potential determinants for malaria: population density as a measure for human-mosquito-contact, deforested area and roads, swampy area, respectively prone to the formation of puddles and hence breeding sites, water, houses to look for shelter for mosquitoes during daytime, forest, and vegetation of banana/plantain, oranges, cacao, and palm trees as potential resting and breeding sites or food sources. Land cover proportions were analysed as continuous variables and human population density as per 1,000 inhabitants. The approximated interquartile range was used as unit increase for the continuous variables.

Because of the small sample size (12 village clusters), the influence of each potential determinant was assessed separately in a univariate Poisson regression in a first step. For a measure of association between a determinant and malaria incidence, the relative risk (RR) was calculated and complemented by a 95% confidence interval (CI) and p-value. In a second step as sensitivity analysis and to account for confounding, the high correlated determinants with a p-value less than 0.05 were included in a bivariate Poisson regression analysis.

Results

Malaria incidence and human population density

A total of 1,988 malaria cases were reported in the study hospital during the study interval of 18 months (end of May 2007 to November 2008) and were included in the analysis. Annual malaria incidence ranged from 28.1 to 271.9 per 1,000 children <15 years of age and year in Akutuase and Agogo, respectively. A crude annual malaria incidence of 194.5 per 1,000 children <15 years age and year with a 95% confidence interval of [144.9, 261.3] could be estimated for the whole study area. The total population of the study villages was 37,461 inhabitants (census data 2004). The human village population density ranged from 1,290 inhabitants/km² in Hwidiem to 8,265 inhabitants/km² in Patriensah (Table 1).

Classification and correlation of land cover determinants

All four broad spectral bands (blue, green, red, and near-infrared) from the acquired multispectral Ikonos image along with the NDVI image were considered in our study. A window size of 9×9 pixels (equivalent to 9×9 m) had the highest accuracy of the texture-based classification and respective textural measures were chosen for the analyses accordingly (data not shown).

By using reference areas for all nine land cover classes a maximum likelihood classification was conducted of the combined NDVI image, the spectral, and the textural bands. Overall accuracy of the classification ranged from 87% in Wioso and Akutuase to 95% in Obenimase.

The proportion of areas with banana/plantain vegetation within a village radius of 2 km varied from 4.8% in Pekyerekye to 19.7% in Agogo. The highest proportion of swampy area was found around the village Agogo (37.0%), the lowest proportions (4.7% and 4.9%) around the two villages Akutuase and Juansa, respectively. The proportion of forest coverage varied from 6.3% around Domeabra to 28.7% around Wioso (Table 2, Figure 2).

Spearman rank tests resulted in high positive correlations between the land cover proportions of forest and deforested area/roads ($r = 0.79$, $p = 0.002$), banana/plantain and built-up areas ($r = 0.86$, $p < 0.001$) as well as palm trees and cacao ($r = 0.91$, $p < 0.001$) (Table 3). Highest negative correlations were observed between the land cover proportions of cacao and banana/plantain ($r = -0.73$, $p = 0.007$) and swampy area and oranges ($r = -0.78$, $p = 0.003$).

As expected, the proportions of land cover in the vicinity of villages in the 0.5 km, 1 km, and 1.5 km radii were not exactly the same from what was found in the 2 km village radius. The proportions of built-up areas (houses), deforested areas and roads, and banana/plantain vegetation decreased with distance to the village whereas the proportion of areas with forest, palm trees, orange trees, and cacao trees increased (Tables S1, S2, S3).

Regression modeling

In the univariate Poisson regression analysis, all determinants with the exception of population density, water, and deforested area and road coverage showed a significant influence on malaria incidence, which was positive for banana/plantain cultivation ($RR = 3.25$), swampy areas ($RR = 1.43$), and built-up areas ($RR = 2.24$), but negative for forest ($RR = 0.53$), orange ($RR = 0.63$), cacao ($RR = 0.48$), and palm trees ($RR = 0.59$) plantation (Table 4). However, in the sensitivity analysis by means of bivariate Poisson regression analysis, the univariate results for built-up areas and orange, cacao, and palm tree plantation turned out to be confounded because of high correlation between several determinants (Table 3) and not to be of statistical significance (data not shown).

The association of land cover with malaria incidence in the other 3 radii were similar to those of the 2 km radius (Table S4) and the sensitivity analysis showed similar results (data not shown).

Discussion

The risk for malaria is dependent on a number of individual and environmental factors whereas their impact is dependent on the
endemicity in a certain area [15,16,20]. Recently, it has been shown that the spatial variance of malaria incidence might be pronounced not only in areas with low and seasonal endemicity but also in holoendemic areas [20,30]. It can be assumed that mosquito occurrence, the existence of breeding sites, and human population density are the most important spatial determinants, all significantly linked to land cover and land use [8,13,14,30]. Accordingly, land cover has been associated with entomological measures mainly accumulated as entomological inoculation rates (EIR, infectious mosquito bites per person per year) [3,5]. Studies using high-resolution satellite images in association with vector-borne diseases have already been conducted in other areas [3–12]. However, analyses of the direct relationship between environmental factors and human malaria, especially using high-resolution images and/or subclassification of land cover in such detail are scarce, most probably due to the absence of precise data on malaria incidence and exact description of the land cover in large areas.

The presented analyses have used malaria incidence data over 18 months from a hospital-based survey and high-resolution satellite images of a holoendemic coverage area in the Ashanti Region, Ghana. The analyses demonstrate that an accurate stratification of land cover has been associated with entomological measures mainly accumulated as entomological inoculation rates (EIR, infectious mosquito bites per person per year) [3,5]. Studies using high-resolution satellite images in association with vector-borne diseases have already been conducted in other areas [3–12]. However, analyses of the direct relationship between environmental factors and human malaria, especially using high-resolution images and/or subclassification of land cover in such detail are scarce, most probably due to the absence of precise data on malaria incidence and exact description of the land cover in large areas.
Land cover by satellite images is possible in areas of small-scale cultivation and changing agriculture. Land cover of banana/plantain vegetation and swampy areas significantly increased the malaria risk. In contrast, an increased proportion of forest around villages was associated with decreased malaria risk. These trends remained when conducting the analysis using smaller radii. However, the RRs for banana/vegetation and forest decreased with each diminished radius. This trend may be explained by the increasing proportions of banana vegetation and decreasing proportions of forest near homesteads and hence the reducing divergence among all villages (Tables S1, S2, S3).

The increase of malaria risk in the vicinity of swampy areas, which are preferred mosquito breeding sites, has already been documented and can be considered as an internal control of the validity of the analyses [13,14,16,31]. In the presented model each 10% increase of the proportion of swampy areas around villages increased the malaria risk by 43%.

A higher proportion of all cultivated areas around villages was associated with a slightly increased risk of malaria (data not shown). The main plantations in the study area are with banana/plantains, oranges, cacao, and palm trees [32]. After stratification for the distinct cultivations, a plantation with banana/plantain was found of particular impact and a 10% increase was associated with about 300% higher malaria risk whereas plantations of oranges, cacao, and palm trees showed a negative association. A number of studies that were conducted at a microhabitat scale demonstrated an association between ovipositions of various mosquito species in rainwater retained in tree-holes and the leaf axils of a variety of numerous wild and cultivated plants such as banana or plantain [33–36]. However, none of these studies has directly linked the existence of breeding sites with malaria incidence.

Likewise, deforested areas and dirt roads have been suspected as environmental factors associated with malaria risk in the surrounding areas since both create conditions favourable for the formation of small puddles that are preferred breeding sites for Anopheles spp. [16,17,29,37,38]. However, a significant influence of deforested areas and roads on malaria incidence could not be observed in the presented study. Similarly, population density as a

| Village radius | Banana/Plantain | Cacao | Palm trees | Oranges | Deforested area and roads | Built-up areas (Houses) | Swampy area | Water | Forest |
|----------------|-----------------|-------|------------|---------|--------------------------|-------------------------|-------------|-------|--------|
| Agogo          | 19.7            | 9.5   | 4.5        | 6.9     | 9.2                      | 4.1                     | 37.0        | 0.1   | 6.4    |
| Akutuase       | 11.5            | 10.9  | 4.1        | 28.7    | 9.4                      | 1.4                     | 1.4         | 1.4   | 27.9   |
| Amantena       | 12.1            | 22.2  | 23.3       | 20.4    | 2.6                      | 0.7                     | 7.5         | 0.1   | 10.1   |
| Domeabra       | 9.8             | 26.2  | 23.9       | 22.7    | 3.9                      | 2.1                     | 5.0         | 0.0   | 6.3    |
| Hwidiem        | 18.7            | 13.3  | 12.6       | 13.2    | 5.6                      | 2.6                     | 23.9        | 0.1   | 9.9    |
| Juansa         | 9.2             | 25.9  | 24.5       | 24.0    | 3.8                      | 2.2                     | 4.9         | 0.0   | 5.5    |
| Kyekyebiase    | 7.3             | 28.9  | 24.3       | 26.6    | 1.6                      | 0.6                     | 5.2         | 0.1   | 5.4    |
| Nyaboo         | 17.4            | 15.5  | 13.9       | 2.2     | 19.6                     | 3.9                     | 8.9         | 0.1   | 15.1   |
| Obenimase      | 14.0            | 19.2  | 19.5       | 3.0     | 17.6                     | 2.5                     | 6.9         | 0.3   | 15.0   |
| Pattrensah     | 16.2            | 15.1  | 16.5       | 2.3     | 18.4                     | 3.4                     | 8.9         | 0.0   | 16.0   |
| Pekyerekye     | 4.8             | 26.4  | 22.1       | 13.5    | 9.1                      | 0.6                     | 8.4         | 4.3   | 10.8   |
| Wioso          | 9.3             | 14.4  | 5.0        | 22.5    | 10.2                     | 0.7                     | 7.8         | 1.2   | 28.7   |

*aSize of each radius 2 km².*

| Table 3. Correlation coefficients of the cross correlation function between land cover variables using Spearman rank correlation. |
|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|
| Population density | Built-up areas (Houses) | Deforested area and roads | Forest | Swampy area | Water | Banana/Plantain | Oranges | Cacao | Palm trees |
| Population density | 0.51 | -0.08 | 0.22 | -0.11 | 0.07 | -0.55 | -0.24 | 0.24 | 0.17 |
| Built-up areas (Houses) | 0.09 | 0.55 | 0.08 | 0.52 | -0.38 | 0.86 | -0.66 | -0.62 | -0.41 |
| Deforested area and roads | 0.80 | 0.06 | 0.79 | 0.37 | 0.23 | 0.46 | -0.66 | -0.52 | -0.62 |
| Forest | 0.50 | 0.81 | 0.002 | 0.14 | 0.42 | 0.18 | -0.30 | -0.51 | -0.67 |
| Swampy area | 0.75 | 0.08 | 0.23 | 0.66 | -0.03 | 0.64 | -0.78 | -0.42 | -0.43 |
| Water | 0.84 | 0.22 | 0.48 | 0.17 | 0.92 | -0.24 | 0.13 | -0.14 | -0.46 |
| Banana/Plantain | 0.06 | <0.001 | 0.13 | 0.57 | 0.02 | 0.46 | -0.67 | -0.73 | -0.57 |
| Oranges | 0.46 | 0.02 | 0.02 | 0.34 | 0.003 | 0.70 | 0.02 | 0.27 | 0.26 |
| Cacao | 0.44 | 0.03 | 0.08 | 0.09 | 0.17 | 0.66 | 0.007 | 0.39 | 0.91 |
| Palm trees | 0.60 | 0.19 | 0.03 | 0.02 | 0.16 | 0.13 | 0.05 | 0.42 | <0.001 |

Note: Above the diagonal are the correlation coefficients r, below all p-values.

doi:10.1371/journal.pone.0017905.t003
were most likely detected and included in our analysis.

Climate conditions are suspected to be of importance for the malaria incidence [39]. The satellite images, which were analysed here, were taken during or immediately after the rainy season (May and November), respectively, the assigned land cover might be biased. By interviewing the local population about previous crops and land cover we attempted to minimise this potential misclassification.

Maximum likelihood procedures were used for supervised classification of land cover data, which gave more accurate results than other classification methods such as Decision Tree-, Minimum Distance- or K-Means Classification [40–42]. Even though the overall accuracy of the correlation matrix of the NDVI image, the spectral and the spatial classification in the subsets ranged from 87% to 95%, land cover still might be misclassified to some extent. In the study area as in most areas of Ghana mixed cultivation is widespread [32] which makes it very difficult to unambiguously allocate land covers.

Our study was limited by the inability to sub-classify swampy areas, which are mostly used for near-ground cultivation, into different crops such as maize, eggplants or pepper for the analysis of various influences on malaria risk. In Ethiopia a strong association between maize cultivation in the vicinity of water bodies, used as breeding habitats, and the larvae development was demonstrated [15]. However, because of a high number of classes in relation to 12 village clusters and weak accuracy of data in this classification analyses, this sub-classification was not possible.

Other geo-ecological influence factors for the malaria risk are altitude, slope, geology, and soil types [29,43–46]. However, the intra-radius variation in these measures did not differ significantly.

The consequent next step should be to map individual data in order to link individual spatial patterns and malaria risk. Indeed, in an adjacent study area a continuous and linear reduction of the malaria rate was demonstrated with an increasing distance between children’s households and forest fringe [30]. Other individual factors such as socioeconomic conditions and the access to health facilities could then be included in the model [20,47,48].

The performed analysis demonstrates that satellite images together with appropriate analytical tools are able to predict the risk of malaria in an area of high malarial transmission. Even though only 12 village sides were included in the study a significant association of different land cover classes with the occurrence of malaria incidence could be demonstrated. Human cultivation in the vicinity of homesteads, in particular with banana/plantain, may increase the risk for malaria. On the contrary, forest preservation may decrease malaria risk. In Ghana, the major farming practice is shifting cultivation, often accompanied by deforestation, and crops mostly change twice to three times a year [32]. Due to the fact that a time span of nine and four months occurred between the acquisition of images (May and November 2009) and the conducted field sampling (March 2010), respectively, the assigned land cover might be biased. By interviewing the local population about previous crops and land cover we attempted to minimise this potential misclassification.

Nevertheless, an association of the proportion of open water bodies and risk of malaria could not be demonstrated in the presented study. A limitation of the analysis is that the proportion of open water in the surrounding of the villages was very low and streams and little rivers were mostly located close to forests and, therefore, difficult to detect on our satellite images.

A limitation of our study is that the proportion of children <5 years of age in each village was estimated and could not directly be measured when computing malaria incidences. Moreover, the underlying values for the total population were three years old (census data from 2004) and hence might be not precise. However, it was the best data available for our study population and census data not more than five years old might represent a quite good measure for potential human-mosquito contact did not show an effect on the outcome.

A high proportion of forest coverage was associated with lower malaria incidence with statistical significance. Indeed, the forest floor with a closed canopy tends to be heavily shaded and littered with a thick layer of organic matter that absorbs water and renders it more acidic. Therefore, the proximity of forest could decrease mosquito abundance and hence decrease malaria risk as the preferred habitat of \textit{A. gambiae} larvae are sunlit pools with turbid water and little or no emergent vegetation and that of \textit{A. funestus} are clear water with vertical, emergent vegetation without organic material [31,37].

There was a tendency of an association between an increased proportion of build-up areas and malaria incidence. However, this effect disappeared after adjustment for the highly correlated variable “banana/plantain” what indicates confounding which is, however, difficult to formally test due to the ecological study design. The observation that vicinity to banana/plantain cultivation seems to be a risk factor for malaria may be because of the frequent closeness of this vegetation with homesteads.

A limitation of our study is that the proportion of children <15 years of age in each village was estimated and could not directly be measured when computing malaria incidences. Moreover, the underlying values for the total population were three years old (census data from 2004) and hence might be not precise. However, it was the best data available for our study population and census data not more than five years old might represent a quite good estimate. Additionally, the villages should be comparable in the proportions of children since they have similar social and ethnic structures, are of similar size, and are all situated in a rural area and closely together. Therefore, it is unlikely that a differential bias was created.

Climate conditions are suspected to be of importance for the malaria risk and higher precipitation could be directly linked, with a time lag, to an abundance of vectors and an increase of disease frequency [39]. The satellite images, which were analysed here, were taken during or immediately after the rainy season (May and November). Therefore, most of the open lakes, rivers or streams were most likely detected and included in our analysis.

### Table 4. Influence of determinants on malaria incidence

| Determinant                  | RR   | 95% Confidence Interval | p-value |
|------------------------------|------|-------------------------|---------|
| Population density           | 0.87 | 0.70–1.07               | 0.176   |
| Built-up areas (houses)      | 2.24 | 1.54–3.24               | <0.001  |
| Deforested area and roads    | 1.00 | 0.70–1.44               | 0.988   |
| Forest                       | 0.53 | 0.28–0.99               | 0.029   |
| Swampy area                  | 1.43 | 1.33–1.55               | <0.001  |
| Water                        | 0.70 | 0.37–1.32               | 0.270   |
| Banana/Plantain              | 3.25 | 2.23–4.76               | <0.001  |
| Oranges                      | 0.63 | 0.44–0.91               | 0.012   |
| Cacao                        | 0.48 | 0.33–0.70               | <0.001  |
| Palm trees                   | 0.59 | 0.43–0.81               | <0.001  |

*Poisson regression analysis.

\( ^a \)Unit: 1,000 km\(^2 \).

\( ^b \)Unit = 2%.

\( ^c \)Unit = 5%.

\( ^d \)Unit = 10%.

\( ^e \)Unit = 1%.

doi:10.1371/journal.pone.0017905.t004
Supporting Information

Table S1 Proportion (in %) of land cover around a 1.5 km village centre radius. Swampy area: either the presence of a river or stream nearby or near the ground agricultural crops (such as eggplants, maize, tomatoes, pepper).

Table S2 Proportion (in %) of land cover around a 1 km village centre radius. Swampy area: either the presence of a river or stream nearby or near the ground agricultural crops (such as eggplants, maize, tomatoes, pepper).

Table S3 Proportion (in %) of land cover around a 0.5 km village centre radius. Swampy area: either the presence of a river or stream nearby or near the ground agricultural crops (such as eggplants, maize, tomatoes, pepper).

Table S4 Influence of determinants on malaria incidence. Association of land cover with malaria incidence using Poisson regression analysis in radii of 0.5 km, 1 km, 1.5 km, and 2 km around each village. Land cover proportions were analysed as continuous variables and were scaled by units as percent increase in radii coverage by built-up areas (houses), per increase in open water of 1%, per increase of deforested area and roads of 5%, per increase in forest, swampy area, banana/plantain, oranges, cacao, and palm tree vegetation of 10%, respectively, and human population density as per 1,000 inhabitants.

Acknowledgments
We thank all field workers and interviewees for their participation in this study. We are also grateful for the continuous endeavours of fieldworkers of the Komansi Centre for Collaborative Research in Tropical Medicine (KCCR) without whose efforts this research would not have been possible, and to the members of the Public Health Unit of the Agogo Presbyterian Hospital for their encouraging collaboration. We thank Kai Sondier from the International Institute of Tropical Agriculture (IIITA) for advising us with satellite imagers. The work is part of the PhD thesis of ACK.

Author Contributions
Conceived and designed the experiments: ACK JO JM JO. Performed the experiments: ACK JO UR BN SA. Analyzed the data: ACK JO UR. Contributed reagents/materials/analysis tools: ACK JO BN SA UR WL NS YA-S. Wrote the paper: ACK NGS JO JM UR.

References
1. WHO: World malaria report. Geneva: World Health Organization (2009) Available: http://apps.who.int/malaria/wmr/2008/malaria2008.pdf. Accessed 2010 June 17.
2. Hay SI, Okoro EA, Gehring PW, Patil AP, Tatem AJ, et al. (2010) Estimating the global clinical burden of Plasmodium falciparum Malaria in 2007. PLoS Med Jun 15(7):e1000290.
3. Rogers DJ, Randolph SE, Snow RW, Hay SI (2002) Satellite imagery in the study and forecast of malaria. Nature 415: 710–712.
4. Hay SI, Packer MJ, Rogers DJ (1997) The impact of remote sensing on the study and control of invertebrate intermediate hosts and vectors for disease. Int J Remote Sens 18: 2899–2930.
5. Bogh C, Lindsay SW, Clarke SE, Dean A, Jawaara M, et al. (2007) High spatial resolution mapping of malaria transmission risk in the Gambia, west Africa, using Landsat TM satellite imagery. Am J Trop Med Hyg 76: 875–881.
6. Mushinzimana E, Munga S, Minakawa N, Li L, Feng CC, et al. (2006) Landscape determinants and remote sensing of anopheles mosquito larval habitats in the western Kenya highlands. Malar J 5: 13.
7. Kulkarni MA, Desrochers RE, Kerr JT (2010) High Resolution Models of habitat diversity and ecology of anopheline larvae in Eritrea. J Med Entomol 40: 921–929.
8. de Souza D, Kelly-Hope L, Lawson B, Wilson M, Boakye D (2010) The impact of deforestation on Anopheles gambiae s.s. distributions in Ghana: An Important Vector of Lymphatic Filariasis and Malaria. PLoS ONE 5(3): e9936.
9. Roberts DR, Paris JF, Manguin S, Harbach RE, Woodruff R, et al. (1994) Remote sensing as a landscape epidemiologic tool to identify villages at high-risk for malaria transmission. Am J Trop Med Hyg 51: 271–280.
10. Sharma VP, Nagpal BN, Srivastava A, Adiga S, Manavalan P (1996) Estimation of larval Anopheles gambiae s.s. in Ghana; an Important Vector of Lymphatic Filariasis and Malaria. PLoS ONE 5(3): e9927.
11. Roberts DR, Paris JF, Manguin S, Harbach RE, Woodruff R, et al. (1996) Predictions of malaria vector distribution in Belize based on multispectral satellite data. Am J Trop Med Hyg 54: 304–308.
12. Beck LR, Rodriguez MH, Dieter SW, Rodriguez AD, Rejmankova E, et al. (1994) Remote sensing as a landscape epidemiologic tool to identify villages at high-risk for malaria transmission. Am J Trop Med Hyg 51: 271–280.
13. Kulkarni MA, Desrochers RE, Kerr JT (2010) High Resolution Models of Malaria Vectors in Northern Tanzania: A New Capacity to Predict Malaria Risk? PLoS ONE 5(2): e9396.
14. de Souza D, Kelly-Hope L, Lawson B, Wilson M, Boakye D (2010) Environmental Factors Associated with the Distribution of Anopheles gambiae s.s. in Ghana; An Important Vector of Lymphatic Filariasis and Malaria. PLoS ONE 5(3): e9927.
15. Hopkins GHE (1952) Mosquitoes of the Ethiopian Region I. Larval Bionomics and Human Population Density. London: British Museum (Natural History). London.
16. Hopkins GHE (1952) Mosquitoes of the Ethiopian Region I. Larval Bionomics and Human Population Density. London: British Museum (Natural History). London.
17. Minakawa N, Munga S, Auerl FK, Mushinzimana E, Zhou G, et al. (2005) Spatial distribution of anopheles larval habitats in western Kenya highlands: effects of land cover types and topography. Am J Trop Med Hyg 73: 157–165.
18. de Souza D, Kelly-Hope L, Lawson B, Wilson M, Boakye D (2010) The impact of deforestation on Anopheles gambiae s.s. distributions in Ghana: An Important Vector of Lymphatic Filariasis and Malaria. PLoS ONE 5(3): e9927.
19. Roberts DR, Paris JF, Manguin S, Harbach RE, Woodruff R, et al. (1994) Remote sensing as a landscape epidemiologic tool to identify villages at high-risk for malaria transmission. Am J Trop Med Hyg 51: 271–280.
20. Sharma VP, Nagpal BN, Srivastava A, Adiga S, Manavalan P (1996) Estimation of larval Anopheles gambiae s.s. in Ghana; an Important Vector of Lymphatic Filariasis and Malaria. PLoS ONE 5(3): e9927.
21. Roberts DR, Paris JF, Manguin S, Harbach RE, Woodruff R, et al. (1994) Remote sensing as a landscape epidemiologic tool to identify villages at high-risk for malaria transmission. Am J Trop Med Hyg 51: 271–280.
22. Roberts DR, Paris JF, Manguin S, Harbach RE, Woodruff R, et al. (1994) Remote sensing as a landscape epidemiologic tool to identify villages at high-risk for malaria transmission. Am J Trop Med Hyg 51: 271–280.
23. Marks F, Adu-Sarkodie Y, Hunger F, Sarpong N, Ekuban S, et al. (2010) Typhoid fever among children, Ghana. Emerg Infect Dis Nov;16(11): 1796–7.
24. Pettorelli N, Vik JO, Mysterud A, Gaillard JM, Tucker CT, et al. (2005) Using the satellite-derived NDVI to assess ecological responses to environmental change. Trends in Ecology and Evolution 20: 403–510.
25. Haralick RM, Shanmugan K, Dinstein I (1973) Texture features for image classification. IEEE Trans. Sysms, Man and Cybernetics SMC-3(6): 610–621.
26. de Jong SM, van der Meer FD (2004) Remote Sensing Image Analysis: Including the Spatial Domain. Dordrecht, Boston, London: Kluver Academic Publishers.
27. Russell PF, Santiago D (1934) Flight range of the Anopheles gambiae s.l. in the States of Santiago and CUBA. J Med 14: 139–157.
28. Russell PF, Santiago D (1934) Flight range of the Anopheles gambiae s.l. in the States of Santiago and CUBA. J Med 14: 139–157.
29. HI-STAR: Health improvement using Space technology and Resources (2002) Available: http://www.isunet.edu/index.php?option=com_content&task=view&id=226&Itemid=251. Accessed 2010 August 12.
30. Roberts DR, Paris JF, Manguin S, Harbach RE, Woodruff R, et al. (1994) Remote sensing as a landscape epidemiologic tool to identify villages at high-risk for malaria transmission. Am J Trop Med Hyg 51: 271–280.
31. WHO (1982) Manual on environmental management for mosquito control. Geneva: World Health Organization.
32. Pettorelli N, Vik JO, Mysterud A, Gaillard JM, Tucker CT, et al. (2005) Using the satellite-derived NDVI to assess ecological responses to environmental change. Trends in Ecology and Evolution 20: 403–510.
33. Haralick RM, Shanmugan K, Dinstein I (1973) Texture features for image classification. IEEE Trans. Sysms, Man and Cybernetics SMC-3(6): 610–621.
34. de Jong SM, van der Meer FD (2004) Remote Sensing Image Analysis: Including the Spatial Domain. Dordrecht, Boston, London: Kluver Academic Publishers.
36. Kerr JA (1933) Studies on the abundance, distribution and feeding habits of some West African mosquitoes. Bull Ent Res 24: 493–510.
37. Patz JA, Graczycyk TK, Geller N, Vittert AY (2000) Effects of environmental change on emerging parasitic diseases. Int J Parasitol 30: 1395–1405.
38. Aflano YA, Zhou G, Lawton BW, Githeko AK, Yan G (2007) Life-table analysis of Anopheles arabiensis in western Kenya highlands: effects of land covers on larval and adult survivorship. Am J Trop Med Hyg 77(4): 660–666.
39. Krefi AC, Schwarz NG, Krüger A, Fobil J, Nkumah B, et al. (2011) Modeling the Relationship between Precipitation and Malaria Incidence in Children from a Holoendemic Area in Ghana. Am J Trop Med Hyg Feb;84(2): 285–291.
40. Kalyani S, Swarup KS (2010) Supervised fuzzy C-means clustering technique for security assessment and classification in power systems. IJEST 2: 175–185.
41. Wie W, Zhang X, Chen X, Tang J, Jiang M (2008) Wetland Mapping using Subpixel Analysis and Decision Tree Classification in the Yellow River Delta Area. ISPRS 37: 667–670.
42. Lawrence R, Hurst R, Weaver T, Aquinall R (2006) Mapping Prairie Pothole Communities with Multitemporal Ikonos Satellite Imagery. PE&RS 72: 169–174.
43. Brooker S, Clarke S, Njagi JK, Polack S, Mugo B, et al. (2004) Spatial clustering of malaria and associated risk factors during an epidemic in a highland area of western Kenya. Trop Med Int Health 9: 757–766.
44. Leonardo LR, Rivera PT, Cristo Tomo BA, Sarol JN, Bantayan NC, et al. (2005) A study of the environmental determinants of malaria and schistosomiasis in the Philippines using Remote Sensing and Geographic Information Systems. Parasitologia 47: 105–114.
45. Minakawa N, Sonye G, Mogi M, Githeko A, Yan G (2002) The effects of climatic factors on the distribution and abundance of malaria vectors in Kenya. J Med Entomol 39: 833–841.
46. Warrel DA, Gilles HM (2002) Essential malariology. 4th ed. London: Arnold.
47. Uguru NP, Onwujekwe OE, Tasie NG, Uzochoke BS, Ezereke UE (2010) Do consumers’ preferences for improved provision of malaria treatment services differ by their socio-economic status and geographic location? A study in southeast Nigeria. BMC Public Health 10: 7.
48. Peterson I, Boer, LN, El-Sadr W, Teklehaimanot A (2009) Individual and household level factors associated with malaria incidence in a highland region of Ethiopia: a multilevel analysis. Am J Trop Med Hyg 80: 103–111.