RESEARCH

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Cost effectiveness and resource 2

- allocation of Plasmodium falciparum malaria 3
- control in Myanmar: a modelling analysis of bed 4 nets and community health workers 5
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- and Yoel Lubell^{1,2} 7

Abstract

8

Background: Funding for malaria control and elimination in Myanmar has increased markedly in recent years. While 9 there are various malaria control tools currently available, two interventions receive the majority of malaria control 10 funding in Myanmar: (1) insecticide-treated bed nets and (2) early diagnosis and treatment through malaria commu-11 nity health workers. This study aims to provide practical recommendations on how to maximize impact from invest-12 ment in these interventions. 13

Methods: A simple decision tree is used to model intervention costs and effects in terms of years of life lost. The 14 evaluation is from the perspective of the service provider and costs and effects are calculated in line with standard 15 methodology. Sensitivity and scenario analysis are undertaken to identify key drivers of cost effectiveness. Standard 16 cost effectiveness analysis is then extended via a spatially explicit resource allocation model. 17

Findings: Community health workers have the potential for high impact on malaria, particularly where there are 18 few alternatives to access malaria treatment, but are relatively costly. Insecticide-treated bed nets are comparatively 19 inexpensive and modestly effective in Myanmar, representing a low risk but modest return intervention. Unlike some 20 healthcare interventions, bed nets and community health workers are not mutually exclusive nor are they necessar-21 ily at their most efficient when universally applied. Modelled resource allocation scenarios highlight that in this case 22 there is no "one size fits all" cost effectiveness result. Health gains will be maximized by effective targeting of both 23 interventions. 24

Keywords: Malaria, Economic, Cost, Cost effectiveness, Policy, Resource allocation 25

Background 26

Malaria in Myanmar is important not only because of 27 the health burden to the country's own population, but 28 29 because of the emergence of artemisinin resistant Plasmodium falciparum parasites in the region [1-3]. The 30 burden of malaria in Myanmar is spatially heterogene-31 ous and seasonal. An estimated 37 % of the population 32 33 live in areas broadly considered at high risk of malaria

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(>1 case per 1000 population) and a further 23 % live in areas of low malaria risk (0–1 cases per 1000 population) [4]. Funds for malaria control and elimination in Myanmar have surged in recent years, including the Myanmar specific Three Millennium Development Goal (3MDG) fund and the Global Fund's Regional Artemisinin Initiative; a US\$ 100 million fund of which US\$ 40 million has been allocated to Myanmar. The financial resources available to Myanmar at this time are both unprecedented in size and potentially time limited. It is critical, therefore, that these resources are allocated efficiently; maximizing impact and improving financially sustainability.



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)	Journal : 12936	Dispatch : 23-9-2015	Pages : 10	
	Article No: 886	🗆 LE	□ TYPESET	
	MS Code : MALJ-D-15-00415	☑ CP	🗹 DISK	

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While there are various malaria control tools cur-46 rently available, two interventions receive the majority 47 of malaria control funding in Myanmar (1) insecticide-48 treated bed nets (ITN), including long-lasting insecti-49 cide-treated nets and (2) early diagnosis and treatment 50 through malaria community health workers (CHW). 51 52 ITN are most effective against mosquitoes which are nocturnal, endophagic blood feeders whereas most 53 species commonly found in Myanmar tend toward cre-54 puscular and exophagic biting [5–7]. The evidence base 55 for the cost effectiveness of ITN against malaria spread 56 by the former type of mosquito is strong [8] and pre-57 vious modelling analysis found that while changes in 58 mosquito biting behaviour could reduce effectiveness, 59 nevertheless ITN could remain a cost effective inter-60 vention [9]. Malaria CHW costs have been estimated 61 in Cambodia [10], Nigeria [11] and across sub-Saharan 62 63 Africa [12].

The malaria policy discourse in Myanmar is frequently 64 framed as a choice between prioritizing universal cov-65 erage of either ITN or CHW. While ITN and CHW 66 can be thought of as competing for limited resources 67 68 they are not mutually exclusive interventions and are in many senses complimentary. It is also the case however 69 that funding is not available for universal access to both 70 interventions, nor has it been demonstrated that such 71 scale-up would be an efficient use of scarce resources in 72 all settings. The factors which determine the costs and 73 effects of both interventions will vary across the country, 74 and context is important in understanding cost effective-75 ness. This study evaluates the costs and effects of these 76 key malaria control interventions in Myanmar with an 77 78 emphasis on sensitivity and scenario analysis rather than a generalized cost effectiveness result. Furthermore, tar-79 geted allocation of these resources is illustrated by an 80 allocation model for a region of Myanmar. 81

Methods 82

Costing 83

Financial costs are included from the perspective of the 84 National Malaria Control Programme or other malaria 85 86 intervention funders. In this analysis ITN distribution is assumed to be conducted though a dedicated distribu-87 tion campaign. ITN cost is comprised of procurement 88 cost (c_p) , direct distribution costs (c_d) and programme 89 management (c_m) . Cost data were obtained from Three 90 Millennium Development Goal (3MDG), a funding 91 92 organization in Myanmar, with crosschecking of components against private sector quotations. A distribution 93 of two nets per household is assumed with 10 % wastage 94 (w) and a mean household size of 5.2 people. The primary 95 time horizon is one year and as such the per person ITN 96 97 cost is annualized according to the lifespan of the net (l),

assumed to be three years, using a discount rate of 5 % (r) [13].

$$c_{ITN} = \frac{(c_p + c_d + c_m)(1+w)}{r^{-1} \left(1 - (1+r)^{-l}\right)} \tag{1}$$

CHW costs are derived from separate detailed cost 101 analysis currently under review. To briefly summa-102 rize, CHW costs are estimated using an ingredients 103 based micro costing of six cost centres: patient ser-104 vices; training; monitoring and supervision, programme 105 management; incentives and overheads. For this cost 106 effectiveness analysis the cost of treatment (c_{ACT}) is sepa-107 rated from the remaining CHW cost per person covered 108 (c_{CHW}) . In addition to intervention costs, diagnosis and 109 treatment direct costs for malaria cases treated by the 110 basic health system are included (c_{ACT}). 111

Model

CHW are an extension of the health system and therefore marginal utility will depend on locally specific access to treatment. The model must define a common metric to quantify the effects of ITN and CHW. The model calculates the number of years of life lost (YLL), a widely used metric for health impact, through treatment of cases or cases directly averted by bed nets. In this case YLL are likely to be similar to disability adjusted life years as the contribution of morbidity will be negligible compared with mortality. The model was developed in both R (version 3.1.2) and TreeAge (TreeAge Pro 2014, USA).

The probability tree (Fig. 1) traces an individual 124 through a chronological series of event possibilities 125 beginning with an annual probability of contracting 126 malaria (m) which is adjusted by the protective effect of 127 ITN (p), if applicable. Individuals with malaria have a 128 probability they will receive treatment from a provider 129 other than a CHW (a). If a CHW is available in the vil-130 lage there is a probability (q) that a malaria case will seek 131 treatment from the CHW, from both those who would 132 have received treatment elsewhere and from those who 133 would not have received any treatment. Each case of 134 malaria has a probability of death in absence of treatment 135 (μ) and a mean number of YLLs lost per death (d). Treat-136 ment is assumed to be with an ACT. The direct reduc-137 tion in mortality is assumed to be the same for ACT (r_1) . 138 The terminal payoffs are scaled by population (ν) and cal-139 culate the net cost and net effects for each intervention 140 arm for one village (or one township when applied in the 141 resource allocation model, see below). Parameter values 142 can be found in Table 1. For the purpose of this model 143 only one provider is attended per person, individuals 144 may seek treatment at a CHW instead of their previous 145 provider. This is intended to reflect the greater marginal 146



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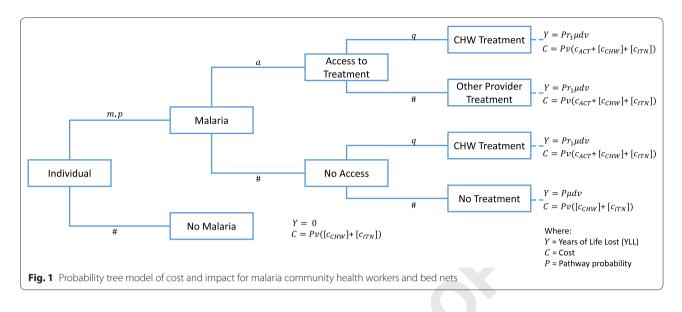
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utility is areas with poor access to treatment, even when 147 uptake at the CHW is equal. 148

149 The model was developed as the simplest structure that incorporates the key relevant data and provides the 150 desired output metrics of cost and years of life lost. The 151 advantages of a simple model are ease of communica-152 tion to end users, speed of development and flexibility of 153 application. 154

Analysis 155

Bed nets and community health workers are not uni-156 157 versally applied interventions and a general estimate of intervention costs and effects misses important 158 159 variation, particularly with respect to the sometimes extreme remoteness of different populations in Myan-160 mar. Instead, intervention cost effectiveness is calcu-161 lated in four illustrative accessibility or remoteness 162 scenarios, whereby more remote settings are character-163 ized by increased cost of programme delivery, increased 164 CHW uptake and decreased baseline access to treat-165 ment (Table 2). Data are not available to support spe-166 167 cific parameterizations for these assumption but the direction of trends are intuitive and supported by policy 168 makers at the national malarial control programme and 169 programme managers at an affiliated non-governmen-170 tal organization, Medical Action Myanmar. In addition 171 172 to the scenario analysis, univariate sensitivity analysis is undertaken to identify key determinants of interven-173 tion cost effectiveness. Probabilistic Sensitivity Analysis 174 (PSA) can be found in the supporting documentation 175 (Additional File 1). Quantified and non-quantified costs 176 and consequences are summarized in Table 3 to aid 177 178 interpretation and to highlight potentially important factors which are not included in the quantitative analysis, 179

as recommended for economic evaluations of public health interventions by Weatherly and colleagues [17].

Cost effectiveness ratios are calculated for each intervention against a common null comparator or "no additional intervention" baseline, which includes the number of YLLs expected in absence of intervention and the cost of treatment for patients who receive it. The marginal benefit of each in the presence of the other is not equal to the marginal benefit of each in isolation. A CHW in a village with good bed net coverage has lower impact than in the same village without bed net coverage because there are fewer cases to treat, and vice versa. For this reason the combined intervention arm is included explicitly as a model output rather than as a sum of separate interventions. Estimates are per year and reflect a village of 500 people with 25 malaria cases per year in absence of interventions.

Resource allocation

An extension to standard cost effectiveness analysis, 198 the second stage of this study applies a spatially explicit 199 resource allocation model for a given budget. The model 200 is applied to the Tier 1 or 'MARC' region of Myanmar, 201 an area in the east of Myanmar identified as a priority 202 area for malaria control. There are 52 townships in Tier 203 1 to which a fixed budget of US\$ 10 million is allocated. 204 Township specific data on population is from the 2014 205 census [18] and malaria incidence is based on routine 206 health system surveillance records, currently managed 207 by WHO Myanmar on behalf of the Ministry of Health 208 (2013, unpublished). The malaria surveillance system in 209 Myanmar is undergoing systemic improvements and data 210 capture is not complete. All other parameter values are as 211 reported in Table 1. 212

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	Model parameter	Symbol	Default value	Lower estimate	Upper estimate	Source
Setting	Baseline access to treatment (% of cases receiving ACT)	a	30 %	1 %	95 %	2011 MARC survey indicates low availability, but recently survey by PSI indicates a substantial increase
	Cost of treatment	CACT	\$3	-	10	Wholesale price of diagnosis and treatment, consumables only.3MDG
	Proportion of malaria cases that die in absence of treatment	μ	1 %	0.1 %	10 %	Expert opinion [14]
	Probability of getting malaria	в	5 %	0.1 %	30 %	Probability of malaria is highly variable but changes do not affect comparative analysis between intervention options
	Probability that a person with malaria uses a CHW (where available)	6	30%	1 %	95 %	Community survey by Department of Medical Research in Myanmar finds 19 % of surveyed first seek treatment at CHW (unpublished). Community survey in Cambodia finds low utilisation of CHW in villages with a CHW (Yeung et al. unpublished)
	Mean number of disability adjusted life years lost per death	P	30	15	45	Assumed based on life expectancy of 65 years and knowing that most malaria deaths in Myanmar are adults
	Village population	>	500	I	I	Village size is based on unpublished unicef data. At the time of the study the village level census data was unavailable
Intervention	Annual cost of ITN per person	C _{ITN}	\$0.70	\$0.50	\$1.5	Estimated
	Annual cost of CHW per person	CCHW	\$2	\$1.10	\$4.50	Kyaw et al. under review
	ITN protective efficacy	р	30 %	% 0	50 %	[5, 9, 15, 16]
	Reduction in mortality after treatment with ACT or ACT + PQ	r.	% 06	50 %	% 66	Expert opinion

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Table 2 Parameter values for four remoteness scenarios

Parameter	Symbol	Remoteness	scenario		
		Easily accessible	Accessible	Difficult to access	Very difficult to access
Annual cost of VHW per person	C _{CHW}	1.10	2.00	3.20	4.50
Annual cost of LLIN per person	C _{ITN}	0.5	0.70	1.2	1.5
Probability that a person with malaria utilises a VHW (where available)	<i>q</i>	0.15	0.3	0.45	0.6
Baseline access to treatment (% of cases receiving ACT)	а	0.5	0.3	0.15	0

Table 3 Cost-consequence summary of insecticide treated nets and malaria community health workers in Myanmar

	ITN	СНШ
Direct costs	One off purchase and distribution costs are annualised over the lifespan of the net Annual equivalent cost per village in modelled scenarios: US\$ 240–750	Annual costs include: training, patient services, monitoring and supervi- sion, programme management and CHW remuneration or incentives. Annual cost range in modelled scenarios (excluding variable drug costs): US\$ 560–2300 Although the effective cost for malaria funds could be reduced through cost sharing
Direct consequences	Modest impact on malaria disease in Myanmar due to crepuscular and exophagic biting	High impact on malaria disease if there is good utilisation of the CHW by people who have malaria
Indirect consequences	Modest impact on malaria transmission in Myan- mar due to crepuscular and exophagic biting	High impact on malaria transmission if there is good utilisation of the CHW by people who have malaria
	Direct effects of ITN result in use of fewer diag- nostics and treatment and therefore save some costs (included in analysis)	CHW can be used to provide other health services, feedback valuable information on malaria burden, provide information and educational messages to the community (not included in analysis)

The allocation model uses the decision tree in Fig. 1 213 to calculate cost effectiveness ratios for all intervention 214 215 options for each geographic patch, in this case a town-216 ship. Once all scenario cost effectiveness ratios are calculated the model allocates the available budget starting 217 with the most cost effective intervention. As the budget 218 is allocated, the most cost effective intervention in a 219 220 particular township may be replaced by a less cost effective, but more effective intervention. Dominated inter-221 vention scenarios, those where any increase in effect 222 can be achieved by a more cost effective alternative, are 223 224 excluded. The allocation process ceases when the remain-225 ing budget is less than the marginal cost of the next most cost effective intervention. It is worth noting that the 226 optimal allocation of resources is not identified through 227 sequential iteration and improvement of budget alloca-228 229 tion options since the cost effectiveness ratios provide 230 sufficient information to identify the allocation result directly. This is more accurate and computationally effi-231 cient than identification of a distribution of resources 232 through iterative optimization or "brute-force" calcula-233 tion of all or a large number of possible distribution sce-234 235 narios. The resource allocation analysis is repeated to examine the impact of variations in bed net protective 236

effectiveness, CHW uptake and cost sharing for integrated CHW programmes.

Results

The cost effectiveness of malaria control in Myanmar is 240 context dependent. CHW have greater potential effects, 241 particularly in more remote settings, but are also more 242 costly. In the scenario analysis, easily accessible village 243 setting CHW avert 0.51 YLLs per year at a cost of US\$ 244 556 (US\$ 1089 per YLL averted). This rises in the very 245 difficult to reach villages to 4.05 YLLs averted at a cost 246 of US\$ 2295 (US\$ 567 per YLL averted), a higher cost but 247 a more cost effective use of CHWs. Bed nets were con-248 sistently less costly and a modestly effective intervention. 249 In the easily accessible village setting bed nets avert 1.24 250 YLLs at a cost of US\$ 238 (US\$ 193 per YLL averted), ris-251 ing to 2.25YLL averted for US\$ 750 (US\$ 333 per YLL 252 averted). In the very difficult to access village setting, a 253 combination of both bed nets and CHW gives the great-254 est impact of 5.08 YLLs averted for a cost of US\$ 3031 255 (US\$ 597 per YLL averted). The above results are sum-256 marized in Table 4 and Fig. 2 and assume that CHW only 257 provide malaria services (this assumption is relaxed in 258 the resource allocation analysis). 259

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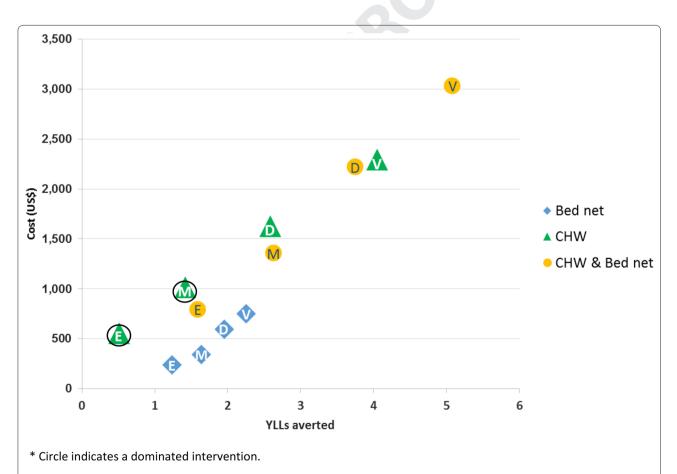
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	Remoteness			
	Easily accessible	Accessible	Difficult to access	Very difficult to access
ITN				
Cost (US\$)	238	343	596	750
Effect (YLLs averted)	1.24	1.64	1.95	2.25
CER*	193	209	306	333
CHW				
Cost (US\$)	556	1016	1629	2295
Effect (YLLs averted)	0.51	1.42	2.58	4.05
CER*	1089	715	631	567
ICER**	Abs dominated	Abs dominated	Ext dominated	Ext dominated
CHW and ITN				
Cost (US\$)	792	1354	2216	3031
Effect (YLLs averted)	1.59	2.63	3.75	5.08
CER*	499	515	591	597
ICER**	1583	1021	503	715

Table 4 Costs and effects of malaria interventions in four remoteness scenarios

* CER here compares costs and effects of an intervention compared with no intervention

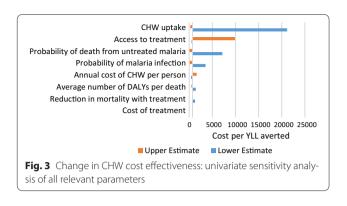
** ICER compares costs and effects of an intervention compared with the next most effective undominated option

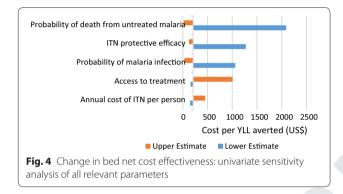


** E: Easily accessible; M: Moderately accessible; D: Difficult to access; V: Very difficult to access.

Fig. 2 Costs and effects of malaria control in different accessibility scenarios. *Circle* indicates a dominated intervention. *E* easily accessible, *M* moderately accessible, *D* difficult to access, *V* very difficult to access

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260 Sensitivity analysis

Univariate sensitivity analysis was conducted for the cost 261 effectiveness of CHW (Fig. 3) and bed nets (Fig. 4) using 262 the wide uncertainty ranges in Table 1. The key determi-263 nants of cost effectiveness for CHW are baseline access 264 to treatment with an ACT and the likelihood that a per-265 son with malaria seeks treatment from the CHW. In real-266 ity these two factors may be related; low baseline access 267 to treatment might be expected to increase treatment 268 seeking at a CHW. Univariate sensitivity analysis treats 269 these values as independent. The key determinants of bed 270 net cost effectiveness are the untreated malaria mortal-271 ity risk and the protective effect of the net. Changes in 272 malaria incidence and mortality affect the magnitude of 273 effects substantially but proportionally for all interven-274 tion options, and therefore do not affect intervention 275 comparison. 276

277 Resource allocation

Figure 5a presents an illustrative optimal allocation of an annual budget of US\$ 10 million to CHW and ITN roll
out in the 52 townships of the MARC region, Myanmar.
Almost half of the townships are allocated both CHW
and ITN, 12 townships receive ITN only and 15 townships are allocated to provide standard health services
without CHW or ITN. Figure 5b-d present the scenario

variations where key assumptions are varied in order 285 to observe the effect on resource allocation. Panel b 286 assumes a low ITN protective effect of 5 %, rather than 287 the default 30 %. Panel c presents resource distribu-288 tion assuming 95 % uptake of CHW by individuals with 289 malaria, rather than 30 %. Panels b and c find that at the 290 margin, CHW rather than ITN should be prioritized. 291 The specific townships receiving these marginal inter-292 ventions is likely to be an artefact of population size and 293 the residual budget amount at the end of the allocation 294 process. Panel d presents a cost-sharing scenario, where 295 the benefits of an integrated CHW programme are repre-296 sented by an assumption that funds allocated for malaria 297 control need only fund 50 % of the total programme cost. 298 Notably, the allocation of both CHW and ITN to the 299 majority of Southern, and Western township and to the 300 Kachin townships in the North, is robust to these sce-301 nario variations. 302

Discussion

Malaria intervention decisions in Myanmar are based on 304 judgement supported by the limited available evidence. 305 The average and incremental cost effectiveness ratios 306 give decision makers a sense of "bang for buck" to inform 307 these judgements while the resource allocation model-308 ling highlights the importance of targeting both interven-309 tions to where they can have the greatest impact. This 310 study finds that CHW have the potential for high impact 311 on malaria, particularly in difficult to access areas, where 312 availability of other services may be low and if CHW 313 use is good. However, CHW are more costly and, if only 314 delivering malaria services, are associated with higher 315 cost-effectiveness ratios. ITN are a robustly cost effec-316 tive intervention but the total health impact is expected 317 to be lower in Myanmar due to the biting habits of the of 318 the main mosquito vector species. The annualization of 319 the ITN cost over the lifespan of the net, conservatively 320 assumed to be three years, means the comparative cost is 321 lower. Although the cost of health gains is low with ITN, 322 in the context of planning for malaria elimination more 323 impactful interventions will need to be considered. 324

The cost effectiveness of both CHW and ITN is sensi-325 tive to the baseline availability of treatment, indicating 326 that services will be most cost effective when targeted 327 to areas with poor access to malaria diagnosis and treat-328 ment. The utilization of CHW is also very important and 329 investment is quality training, CHW supervision and 330 community engagement may be important to imple-331 menting a cost effective CHW programme [19]. A fur-332 ther option available to planners seeking to improve the 333 cost effectiveness of CHW programmes is to expand 334 the package of services offered by CHW. This is already 335

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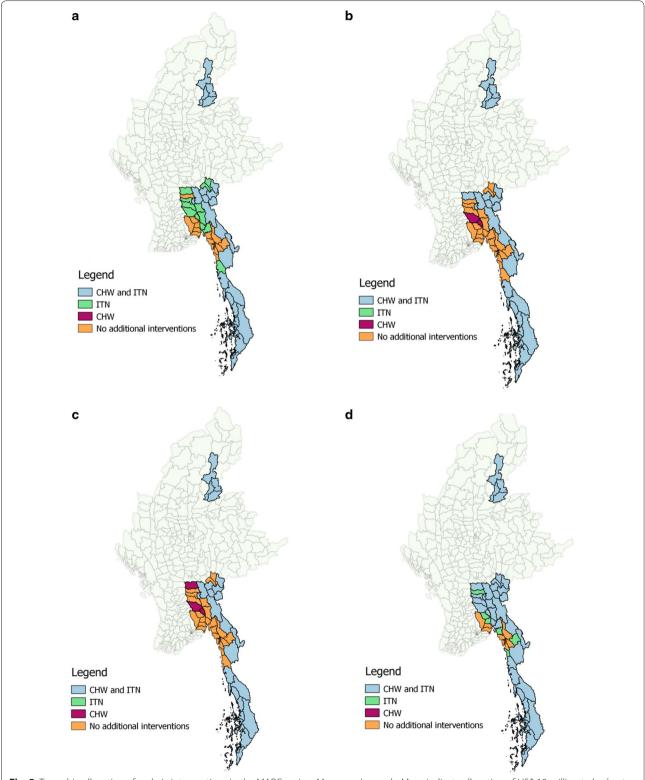


Fig. 5 Township allocation of malaria interventions in the MARC region, Myanmar. Legends: Maps indicate allocation of US\$ 10 million to bed nets and malaria community health workers in the MARC region, Myanmar. **a** Allocation using default parameter values detailed in Table 1. **b** Allocation assuming a lower bed net protective effect of 5 %. **c** Allocation assuming a higher uptake of community health workers; 95 % of malaria infections. **d** Allocation assuming 50 % cost-sharing for community health workers. For *panels* (**b**–**d**) all parameters other than the specified variation are the default values outlined in Table 1

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happening and many CHW are now also providing a 336 basic health care package or providing additional services 337 such as tuberculosis detection and treatment. Measures 338 to improve the cost effectiveness of community health 339 workers include expanding the scope of available ser-340 vices; strategies to improve the likelihood that commu-341 342 nity members seek treatment from the community health worker when they have fever; and targeting community 343 health workers to where they will be most cost effective. 344

For several reasons the main analysis does not apply 345 a cost effectiveness threshold. It is difficult to define an 346 appropriate threshold for the cost per YLL or DALY 347 averted; the budget context in Myanmar is complex with 348 modest NMCP funds being supplemented by interna-349 tional aid. Moreover in the context of a drive towards 350 elimination all interventions will cease to appear cost 351 effective as the malaria burden decreases (in absence of a 352 model for long term benefits). The use of measures such 353 as cost-per DALY averted are, therefore, less relevant 354 and highly uncertain [20, 21]. The most immediately rel-355 evant question is how to maximize impact from malaria 356 funds available in Myanmar and for this no threshold is 357 358 necessary.

An extension of standard cost effectiveness analysis to < 359 spatially (in this case township-wise) specific resource 360 allocation modelling highlights the need for a paradigm 361 shift in policy discussion from prioritizing universal cov-362 erage of the "most cost effective" intervention to targeting 363 of both interventions and presents illustrative township 364 specific recommendations. In this analysis, malaria bur-365 den and to a lesser extent population numbers determine 366 the optimal distribution of resources. Future work will 367 seek to include additional data specific to each township. 368

Part of the aim of this study is to formalize through 369 a cost effectiveness framework the kind of intuitive 370 judgements that many policy makers and influencers in 371 Myanmar are discussing. There has been much debate 372 373 regarding the various merits of bed nets and malaria CHW. This paper does not come down on either side of 374 this debate but seeks to summarize the characteristics of 375 each and highlight the importance of targeting both to 376 areas where impact can be maximized. 377

Limitations 378

This study has several limitations. The model does not 379 include human population movement or malaria trans-380 mission dynamics. A malaria transmission model, incor-381 382 porated into the cost effectiveness model, would be a useful extension. This would allow indirect effects to be 383 incorporated into the analysis and allow provide projec-384 tions of the impact on malaria transmission going for-385 ward. The analysis does not include benefits to the patient 386 387 beyond malaria impact, such as reduced costs to access Page 9 of 10

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care nor are issues of service quality examined here. For 388 CHW there is a strong interest in extending their ability 389 to diagnose and treat other causes of illness and there-390 fore higher health gains than accounted for here. The 391 model considers malaria control in the general popula-392 tion and does not specifically include high-risk groups 393 such as migrant or mobile populations. Resource alloca-394 tion modelling is applied at the township level whereas in 395 Myanmar townships make decisions to allocate malaria 396 interventions on a village-by-village basis. Finally, town-397 ship variation here is characterized by population and 398 malaria burden. Costs, baseline access to treatment and 399 treatment-seeking behaviour are not assumed to vary 400 between townships. 401

Additional file

Additional file 1. Cost effectiveness and resource allocation of malaria control in Myanmar: further sensitivity and scenario analyses.

Authors' contributions

TD and YL conceived of the study. TD and SSK completed the costing sections.	404
TD developed the model and undertook the analyses. FS, ND, LJ, MPK and YL	405
provided critical feedback during several iterations of the analysis and manu-	406
script. All authors read and approved the final manuscript.	407
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Myanmar. ⁴ Medical Action Myanmar, Yangon, Myanmar.	412
Acknowledgements	413
The authors would like to acknowledge the support of the National Malaria	414
Control Programme, the Department of Medical Research and the World	415
Health Organization, Myanmar Country Office.	416
Compliance with ethical guidelines	417
Competing interests	418
The authors declare that they have no competing interests.	419
Funding statement	420
This work was supported by the Three Millennium Development Goal (3MDG)	421
Fund, the Bill and Melinda Gates Foundation (BMGF) and the Wellcome Trust	422
Major Overseas Programme in SE Asia (grant number 106698/Z/14/Z).	423
Received: 23 June 2015 Accepted: 2 September 2015	424
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Journal : 12936	Dispatch : 23-9-2015	Pages : 10	
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