

**Additional file 1. Studies on the interactions between worms and malaria in Humans.**

Study date	site	Age group	Design & Sample size	Worm species	Malaria	Remarks
1977	Comoro Islands [1]	Children 0-14	Cross sectional & ecological 869	<i>Ascaris lumbricoides</i>	Decreased prevalence/incidence	It is not clear if prevalence or incidence is considered  The comparison is not with absence of <i>Ascaris</i> but between high burden and low burden
1978	Comoro Islands[2]	Children 2-14	Randomized controlled trial 122	<i>Ascaris lumbricoides</i>	Increased 'incidence' between 6-14 days after piperazine treatment	Given the short interval, malaria was probably present but asymptomatic before piperazine
2000	Thailand [3]	Adults	Case control 537	<i>Ascaris lumbricoides</i>	Protection from cerebral malaria  Renal failure  Pulmonary edema	Dose dependent protection  Protection increases with the number of GI species involved
2001	Thailand[28]	Adults	Case control 179	pooled	Protection from renal failure  Protection from jaundice	Linear trend between egg count and odds of renal failure  Fewer mature schizonts in GI nematode-infected patients

<b>Thailand</b> [33] 2001	Adults	Cross sectional	hookworm	Decreased admission temperature	
<b>Thailand</b> [49] 2001	Adults	Cross sectional	Pooled (excluding hookworm)	Increased anemia	
<b>Thailand</b> [36] 2001	Adults	Cross sectional	<i>Ascaris lumbricoides</i>	More mixed Pf-PV infections	
<b>Thailand</b> [22] 2002	Adults	Cohort	Pooled	Increased incidence	Incidence tends (P=0.07) to increase with the number of worm species  Mostly hookworm (57%) significant linear trend between incidence and hookworm egg burden
<b>Thailand</b> [27] 2002	Adults	Case control	pooled	Protection from cerebral malaria	<i>Ascaris</i> only individual species significantly associated with protection (AOR=0.15)  Controlling for body mass index
<b>Thailand</b> [34] 2002	Adults	Cross sectional	pooled	Increased gametocyte carriage	Association is confounded by lower hemoglobin counts  Linear trend between egg count and odds of gametocyte carriage

<b>Senegal</b> [23] 2003	Children 1-14	Cohort  80	pooled	Increased incidence	
<b>Thailand</b> [35] 2003	Adults	Cross sectional  248	<i>Trichuris trichiura</i>	Increased multiplicity of infection	
<b>Senegal</b> [16] 2004	Children	Case control  128	<i>Ascaris lumbricoides</i>	Increased severe malaria	Case definition includes vomiting (exposure can cause vomiting=>bias)  Case classification not performed by physician (39% of the severe malaria diagnoses in fact not malaria)  Controls do not have malaria
<b>Senegal</b> [30] 2004	Children	Cohort  512	<i>Schistosoma mansoni</i>	Increased <i>falciparum</i> malaria incidence	No linear trend between egg burden and malaria but heavy worm burdens had highest malaria incidence
<b>Senegal</b> [32] 2004	Children 7-15& adults>30	Cross sectional  79 children + 49 adults	<i>Schistosoma haematobium</i>	No difference in parasitaemia	NB. Patients with clinical mild or severe malaria excluded
<b>Thailand</b>	Adults	Cross sectional	<i>Ascaris lumbricoides</i>	Negative correlation between proportion of fertilized <i>Ascaris</i> eggs and admission	

[37]		119		temperature in <i>vivax</i> malaria	
<b>2005</b>					
<b>Uganda</b>	Children+ Adults	Cross sectional	Pooled	No association	
[4]		856	+individually		
<b>2005</b>					
<b>Senegal</b>	Children	Longitudinal	<i>Schistosoma haematobium</i>	Decreased parasitaemia	Association with decreased parasitaemia observed in low egg burdens
[26]		523			
<b>2005</b>			<i>GI nematodes</i> (pooled)		Non significant trend for negative association  For GI nematodes & malaria no association with parasite densities. Pooled but mainly <i>Ascaris</i> .
<b>Mali</b>	Children (4-14)	Cohort	<i>Schistosoma haematobium</i>	Decreased incidence of <i>clinical</i> malaria	IL-6 and IL-10 levels blunted by <i>S. haematobium</i>
[31]		676			[50]
<b>2005</b>					
<b>Madagascar</b>	Children	Randomized controlled trial	<i>Ascaris</i>	Increased <i>falciparum</i> parasitaemia after levamisole treatment of <i>Ascaris</i> in children > 5 years	No apparent effect before 5 years of age
[6] 2006		350			
<b>Madagascar</b>	Children	Randomized controlled trial	<i>Ascaris</i>	Increased <i>falciparum</i> parasitaemia after levamisole treatment of <i>Ascaris</i> in children > 5 years	No apparent effect before 5 years of age
[7] 2007		212			

<b>Kenya [14]</b> <b>2008</b>	Children	Cohort	Pooled	No increased incidence	
		387			
<b>Uganda [19]</b> <b>2008</b>	Pregnant women	Cross sectional	Hookworm	Increased malaria prevalence	<i>Mansonella perstans</i> associated with hookworm and malaria
		2507			
<b>Zimbabwe [20]</b> <b>2008</b>	Children	Cross sectional	Hookworm	Increased <i>falciparum</i> malaria prevalence	<i>S. mansoni</i> also associated with increased <i>falciparum</i> malaria prevalence
		1303			
<b>Kenya [8]</b> <b>2009</b>	Pregnant women	Cross sectional	<i>Ascaris lumbricoides</i>	Lower malaria prevalence	Gravida 2 & 3
		390			
<b>Ethiopia [9]</b> <b>2009</b>	Children & adults	Cross sectional	Hookworm <i>Ascaris lumbricoides</i> Pooled	Intensity of hookworm infection correlates with malaria parasitaemia Lower malaria parasitaemia in <i>Ascaris</i> heavy infections Less severe malaria in helminth- infected persons	
		458			
<b>Zanzibar [25]</b>	Children 6-23 months	Cross sectional+Case control	Pooled individual nematodes	and Less malaria in nematode-infected children	Nematode-infected children had higher hemoglobin concentration and mid-upper arm circumference than children without nematodes

<b>2009</b>		2322 + 690			
<b>Ghana</b> [12] <b>2010</b>	Pregnant women	Cross sectional 746	Pooled individual nematodes	& Increased malaria prevalence in hookworm-infected women & <i>Ascaris</i> -infected women	
<b>Senegal</b> [43] <b>2010</b>	Children 1-14	Cohort 203	pooled	Increased malaria incidence	Mostly hookworm (43%) Then <i>Ascaris</i> (10 %)
<b>Brazil</b> [10] <b>2010</b>	Children 5-14	Cohort 216	<i>Ascaris lumbricoides</i> <i>Trichuris trichiura</i> hookworm	Lower drop in haematocrit during <i>vivax</i> malaria in patients with <i>Ascaris</i> , <i>trichuris</i> , or hookworm	
<b>Thailand</b> [11] <b>2010</b>	Pregnant women	Cross sectional 829	hookworm <i>Ascaris lumbricoides</i>	Increased malaria ( <i>vivax</i> & <i>falciparum</i> ) in hookworm-infected women Decreased malaria ( <i>vivax</i> & <i>falciparum</i> ) in <i>Ascaris</i> -infected women	
<b>Nigeria</b> [24] <b>2010</b>	Children 12-59 months	Randomized control trial 320	All worms	Decreased malaria prevalence on 4-monthly screenings in patients receiving albendazole Non significant trend to have higher parasitaemia in the placebo group Non significant increase in haemoglobin concentration in children receiving	Authors conclude that parasite clearance and immunity may be delayed. <i>Ascaris</i> singled out but other helminths also treated by albendazole. Increased incidence and role of hemoglobin concentration not discussed.

albendazole					
<b>Gabon</b> <b>[13]</b> <b>2010</b>	Pregnant women	Longitudinal survey  388	All worms	<i>Ascaris</i> associated with increased malaria incidence	Not clear if <i>Ascaris</i> -infected women were treated before malaria
<b>Global</b> <b>2010</b>	Country prevalences for different geohelminths  Malaria incidence data for each country	Classification analysis regression trees/ Ecological data from 108 countries	<i>Ascaris lumbricoides</i> / hookworm	<i>Ascaris</i> negatively associated with malaria incidence (10 fold reduction)  Hookworm associated with increased malaria	Ecological data regression analysis trees.  Hookworm effect not observed in multivariable analysis
<b>Uganda</b> <b>[21]</b> <b>2011</b>	Children & adults	Cross sectional	hookworm	Positive association between Plasmodium and hookworm among preschool-aged children and adults, but not school -aged children.  Spatial and household clustering of coinfections.	Link between malaria and other gastrointestinal nematodes not reported in the study.  Study controlled for socioeconomic and microgeographic factors.