

A voluntary use of insecticide-treated cattle can eliminate African sleeping sickness

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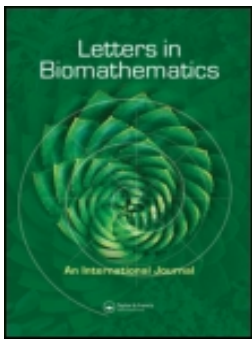
Abstract:

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Keywords: Game theory | vector control | African sleeping sickness | insecticide

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A voluntary use of insecticide-treated cattle can eliminate African sleeping sickness

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ABSTRACT

African sleeping sickness is a vector-borne disease caused by the parasite *Trypanosoma brucei*. It is transmitted by tsetse flies and one of the most effective strategies to combat the disease is the use of insecticide-treated cattle (ITC). In this paper, we present a game-theoretical model, in which individual farmers choose their own level of ITC in order to maximize their own benefits, effectively balancing the cost of ITC and the risk of their cows contracting the disease. We find that even when the usage of ITC is strictly voluntary, the optimal ITC usage will eliminate the disease when the cost of ITC is not prohibitively large. This is in a sharp contrast with similar scenarios of vaccination games where a voluntary vaccination never eliminates a disease entirely.

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1. Introduction

African sleeping sickness (African Trypanosomiasis) is a vector-borne disease caused by the parasite *Trypanosoma brucei* (Simarro, Jannin, & Cattand, 2008). The disease is fatal in humans if left untreated (Coleman & Welburn, 2004; Davis, Aksoy, & Galvani, 2011) and the treatment of livestock has been hindered by drug resistance (Bourn, Grant, Shaw, & Torr, 2005; Hargrove, Omolo, Msalilwa, & Fox, 2000; Kajunguri et al., 2014). The disease is transmitted by tsetse flies and the most effective strategy to combat the disease is to eliminate the vectors (Rowlands et al., 2001). The use of insecticide-treated cattle (ITC) as baits for the tsetse flies is an effective vector control method causing little environmental damage (Hargrove, Ouifki, Kajunguri, Vale, & Torr, 2012; Kabayo, 2002; Peter, Van den Bossche, Penzhorn, & Sharp, 2005; Vale & Torr, 2005). The original ITC protocol was to treat the whole body of the adult cattle but an application of insecticide to only legs where the flies bite most often (Torr et al., 2001) and to the ears where ticks accumulate, improved the cost and efficacy of tsetse control (Torr, Maudlin, & Vale, 2007). Still, to be effective, the ITC technique should be applied to a large area to prevent re-infestation by the controlled region (Torr & Vale, 2011). It requires participation and cooperation by a large number of farmers who are encouraged to control tsetse themselves rather than relying on governments (Kajunguri et al., 2014).

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Game-theoretical models of individuals facing a decision to either do a disease-preventive action (such as to vaccinate or to use ITC) or to risk contracting the disease have become increasingly common (Fenichel, Horan, & Hickling, 2010; Reeling & Horan, 2015; Wang & Hennessy, 2015). Game theory has been applied to vaccination against major public health threats, including smallpox (Bauch, Galvani, & Earn, 2003), measles (Shim et al., 2012b), rubella (Shim, Kochin, & Galvani, 2009), childhood diseases (Bauch, 2005), influenza (Galvani, Reluga, & Chapman, 2007) and many others. There is no human vaccine against African Trypanosomiasis (La Greca & Magez, 2011; Tabel, Wei, & Bull, 2013). The ITC technique is not a vaccination and does not prevent cattle from contracting the disease (Vale, Mutika, & Lovemore, 1999). Still, ITC can be seen as a disease-preventive action as the use of ITC reduces the incidence of the disease in the population and it can thus be modelled in a similar way as vaccination (Bauch & Earn, 2004).

From the game theoretic perspective, an individual takes an action (such as to use ITC) that will maximize personal pay-off, taking into account the disease incidence and risk of infection which is determined by actions taken by the rest of the population (Shim, Chapman, Townsend, & Galvani, 2012a). In theory, individuals are driven by self-interests rather than by the interest of the group (Shim, Meyers, & Galvani, 2011). A usual outcome is that as the frequency of preventive actions increases, the incentive to take such an action decreases because of indirect protection by other protected individuals (Bauch & Earn, 2004). This makes the eradication of a disease very difficult and practically impossible, even when the cost of the preventive action is very low (Geoffard & Philipson, 1997).

In this paper, we follow a general approach of previous game theoretical models such as Bauch and Earn (2004) and apply the approach to a situation where individuals choose to use ITC to reduce the tsetse vector prevalence. Unlike in previously studied vaccination games, the use of ITC does not prevent the cows contracting the disease (Kajunguri et al., 2014), but rather helps to reduce the number of tsetse flies and consequently helps to increase the number of healthy cows. Thanks to this indirect but large benefit, our model shows that even when the use of ITC is strictly voluntary, the optimal level of protection yields the level needed for tsetse elimination, i.e. effectively achieving the herd immunity that was not possible in vaccination games.

2. Mathematical model

2.1. Transmission model with ITC

Our model relies heavily on the results and analysis of Kajunguri (2013) and Kajunguri et al. (2014). In Kajunguri et al. (2014), the authors present a SIR model involving all the variables related to tsetse vector control by ITC in a multi-host population. A slightly modified model for one host only (cattle) is shown in Figure 1 and described in this section. The notation and model parameters are summarized in Table 1.

The adult cattle population is divided into three classes, susceptible (S), infectious (I) and recovered (R), whereas the tsetse population is divided into two classes, susceptible (S_V) and infectious (I_V). To include ITC in the model, we split the cattle population into treated (with superscript t) and untreated with (with superscript u). The ITC does not prevent the tsetse fly from biting but each fly that bites or touches a treated cow dies within a few hours (Vale, Mutika, & Lovemore, 1999). It means that I and R classes are also divided into treated and untreated

Table 1. Notations and parameter values, adapted from Kajunguri et al. (2014).

	Meaning	Value	Source
Λ	Host recruitment rate	22	WHO (2012)
μ	Host natural mortality	.00055	WHO (2012)
a	Biting rate	.25	Coleman and Welburn (2004)
β	Prob. of transmission from fly to cattle	.62	Rogers (1988)
α	Prob. of transmission from cattle to fly	.065	Rogers (1988)
T_H	Incubation period in host	7 days	Coleman and Welburn (2004)
T_V	Incubation period in flies	18 days	Coleman and Welburn (2004)
σ	Mortality of infected hosts	.006	Davis et al. (2011)
g	Host recovery rate	.014	Davis et al. (2011)
ν	Rate of immunity loss in hosts	1	Coleman and Welburn (2004)
B_V	Tsetse birth rate	1440	Davis et al. (2011)
Λ_V	Tsetse recruitment rate	Equation (5)	Kajunguri et al. (2014)
μ_V	Tsetse natural mortality rate	.03	Rogers (1988)
m	Tsetse additional mortality due to ITC	.57	Torr et al. (2001)
m_V	Tsetse mortality rate with ITC	Equation (4)	Kajunguri et al. (2014)
d^{-1}	Duration of insecticide efficacy	28 days	Torr et al. (2007)
ψ	Proportion of cattle being treated	variable	
π	Stable proportion of treated cattle	$\psi/(\psi + d)$	
λ	Force of infection in hosts	Equation (14)	
λ_V	Force of infection in flies	Equation (15)	
C	Normalized cost of ITC per cow per day	variable	
x	Percentage of cattle owned by a focal farmer	variable	

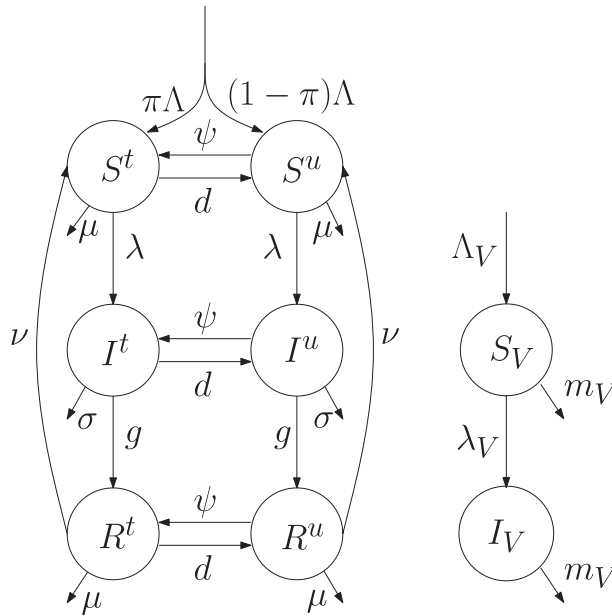


Figure 1. Compartmental model of African sleeping sickness transmission in cattle host and Tsetse flies, adapted from Kajunguri et al. (2014). There is a time lag due to the incubation period and so at time t , $\lambda(t - T_H)S(t - T_H)$ of cattle leave the susceptible and enters the infections class; here $\lambda(t - T_H)$ is the force of infection at time $t - T_H$ and $S(t - T_H)$ is the susceptible population at time $t - T_H$. A similar situation occurs with tsetse flies where the force of infection is λ_V and the incubation period is T_V .

$$S = S^t + S^u, \quad (1)$$

$$I = I^t + I^u, \quad (2)$$

$$R = R^t + R^u. \quad (3)$$

We assume the cattle is treated at rate ψ , thus moving from untreated classes S^u, I^u or R^u to the treated classes, S^t, I^t or R^t . The insecticide effect lasts on average for $1/d$ days after which the cattle moves from the treated classes, S^t, I^t or R^t to the untreated classes S^u, I^u or R^u .

The tsetse flies feed on the adult cattle. The feeding rate is a , i.e. each fly takes a new blood meal on average every $1/a$ days. During the blood meal, the disease is transmitted from an infected cow to a fly with probability α and from an infected fly to a cow with probability β . As in Kajunguri et al. (2014), the time delays T_H and T_V are representing the incubation period in the host and tsetse vector populations, respectively.

The cattle recruitment rate Λ ; the rate is given by a combination of the immigration and reaching adult size. A newly recruited cow is assumed to be susceptible. However, unlike Kajunguri et al. (2014), we assume that the proportion $\pi = \psi/(\psi + d)$ of the recruited cattle is treated, i.e. the proportion of treated versus untreated newly recruited cattle is ψ/d . This assumption is the only deviation from the model described in Kajunguri et al. (2014) and we do it in order to make the flow chart in Figure 1 more symmetric which allows us to get an analytical solution and postpone the use of computation by one step. The assumption can be easily satisfied by treating the newly recruited cattle.

The mortality of the tsetse flies is given by

$$m_V = \mu_V + am \frac{S^t + I^t + R^t}{N} \quad (4)$$

where μ_V is the natural mortality, $N = S + I + R$ is the total number of cattle and $(S^t + I^t + R^t)/N$ is the probability a flies bites a treated cow and m is the additional mortality due to the insecticides.

The recruitment rate of tsetse vectors is derived in Kajunguri et al. (2014) as

$$\Lambda_V = \frac{B_V}{a + m_V} (1 - \exp(-a - m_V)) \quad (5)$$

where B_V is the tsetse birth rate.

The flow chart on Figure 1 then yields the following equations

$$\frac{d}{dt} S^u = (1 - \pi)\Lambda + \nu R^u + dS^t - (\psi + \mu)S^u - \lambda(t - T_H)S^u(t - T_H) \quad (6)$$

$$\frac{d}{dt} I^u = \lambda(t - T_H)S^u(t - T_H) + dI^t - (\psi + g + \sigma)I^u \quad (7)$$

$$\frac{d}{dt} R^u = gI^u + dR^t - (\psi + \mu + \nu)R \quad (8)$$

$$\frac{d}{dt} S^t = \pi\Lambda + \psi S^u + \nu R^t - (d + \mu)S^t - \lambda(t - T_H)S^t(t - T_H) \quad (9)$$

$$\frac{d}{dt}I^t = \psi I^u + \lambda(t - T_H)S^t(t - T_H) - (d + g + \sigma)I^t \quad (10)$$

$$\frac{d}{dt}R^t = \psi R^u + gI^t - (d + \mu + \nu)R^t \quad (11)$$

$$\frac{d}{dt}S_V = \Lambda_V - \exp(-m_V T_V)\lambda_V(t - T_V)S_V(t - T_V) - m_V S_V \quad (12)$$

$$\frac{d}{dt}I_V = \exp(-m_V T_V)\lambda_V(t - T_V)S_V(t - T_V) - m_V I_V \quad (13)$$

where

$$\lambda = a\beta \frac{I_V}{N}, \quad (14)$$

$$\lambda_V = a\alpha \frac{I}{N} \quad (15)$$

are the forces of infection. As we will later see from (21), λ plays a critical role in determining the equilibrium value of N , the total count of the cattle. In turn, N is critical in evaluating the farmer's benefits as seen in (20).

2.2. Game-theoretical model

Let us assume that an individual farmer owns a proportion x of all of the cattle in a given region that is large enough to prevent an immediate tsetse re-introduction from regions with no tsetse control (Torr & Vale, 2011). Let ψ_0 be the average ITC usage among other farmers in the area. We want to optimize ψ_f , the level of ITC usage of the focal farmer. The average usage of ITC in the population is given by

$$\psi = x\psi_f + (1 - x)\psi_0. \quad (16)$$

If there are N cattle in total, the focal farmer will have to treat $\psi_f xN$ of the existing cattle as well as proportion $x\pi$ of the newly recruited cattle (assuming the recruitment of cattle is uniform among the farmers). When the system (6)–(13) is in equilibrium, $N\mu = \Lambda$ and thus the total number of treated cattle per day, N_f^t , is given by $xN(\psi_f + \pi\mu)$. Assuming x is relatively small, we get

$$N_f^t \approx xN \left(\psi + \mu \frac{\psi_0}{\psi_0 + d} \right). \quad (17)$$

Let C be the normalized cost of treatment of one cow per day (assuming benefits of having such a cow are equal to 1). We can then evaluate $E(\psi_f, \psi_0)$, the focal farmer's net benefits of using strategy ψ_f in the population where everybody else uses strategy ψ_0 by

$$E(\psi_f, \psi_0) = xN - CN_f^t \quad (18)$$

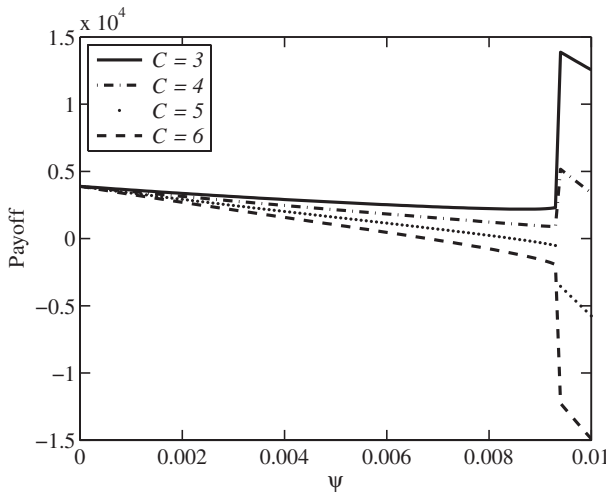


Figure 2. Dependence of pay-off $E(\psi, \psi)$ on the ITC level of treatment ψ for different relative costs of treatments C . Parameter values as in Table 1. Since the farmer uses strategy ψ as everybody else in the population, the value of x is irrelevant in this case.

$$= xN \left(1 - C\psi - C\mu \frac{\psi_0}{\psi_0 + d} \right) \tag{19}$$

$$= xN \left(1 - Cx\psi_f - C(1-x)\psi_0 - C\mu \frac{\psi_0}{\psi_0 + d} \right).$$

3. Results

When we rewrite system (6)–(11) for S, I, R , we get exactly the system (4)–(8) from Kajunguri et al. (2014) (for the model formally without ITC).

Following Kajunguri et al. (2014), we have

$$N = \frac{G\lambda + H}{D\lambda + F} \tag{21}$$

where the force of infection λ satisfies

$$\lambda(a_0\lambda^2 + b_0\lambda + c_0) = 0 \tag{22}$$

and the auxiliary variables $a_0, b_0, c_0, A, B, D, F, G, H$ and the basic reproduction number R_0 are given by

$$R_0^2 = \frac{\exp(-m_V T_V)}{m_V^2} \cdot \frac{\alpha a^2 \beta \mu \Lambda_V}{\Lambda(g + \sigma)} \tag{23}$$

and

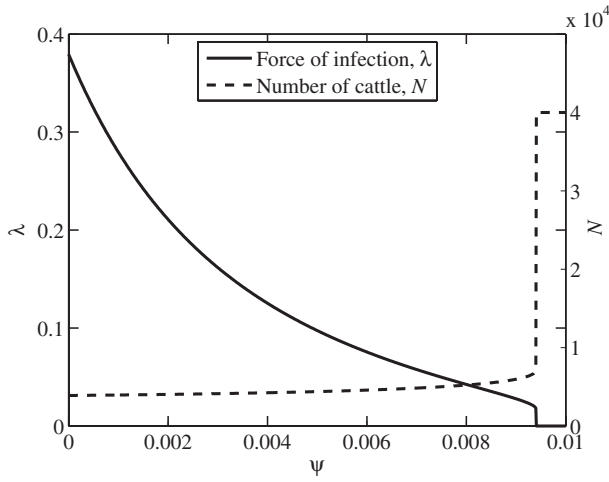


Figure 3. Dependence of force of infection λ and the total number of cattle N on the ITC level of treatment ψ (this is independent of the relative costs of treatments C). Parameter values as in Table 1.

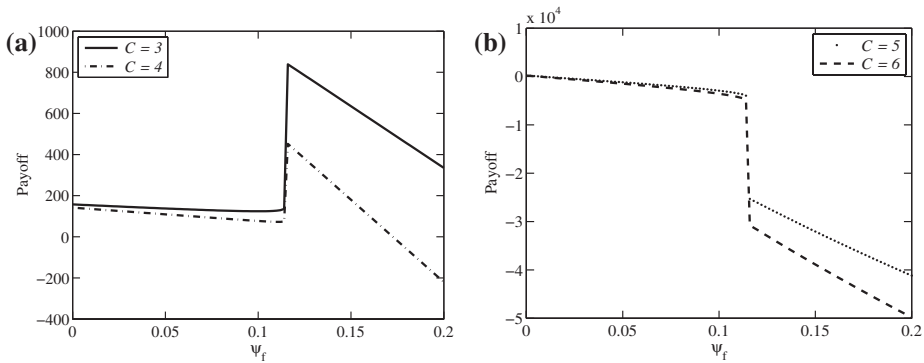


Figure 4. Dependence of pay-off $E(\psi_f, \psi_0)$ on the individual ITC level of treatment ψ_f for different relative costs of treatments C , $\psi_0 = .003$, $x = .05$, and other parameter values as in Table 1. The jump occurs exactly when the average level of ITC usage ψ is at the critical value yielding the eradication of the disease.

$$a_0 = Gm_v(A\alpha aB + Gm_v) \tag{24}$$

$$b_0 = AH\alpha aBm_v + 2HGm_v^2 - A\alpha a^2\Lambda_vBD\beta \tag{25}$$

$$c_0 = \Lambda m_v^2(g + \sigma)(1 - R_0^2) \tag{26}$$

$$A = \exp(-m_v T_v) \tag{27}$$

$$B = \Lambda(\mu + \nu) \tag{28}$$

$$D = g\mu + \sigma(\mu + \nu) \tag{29}$$

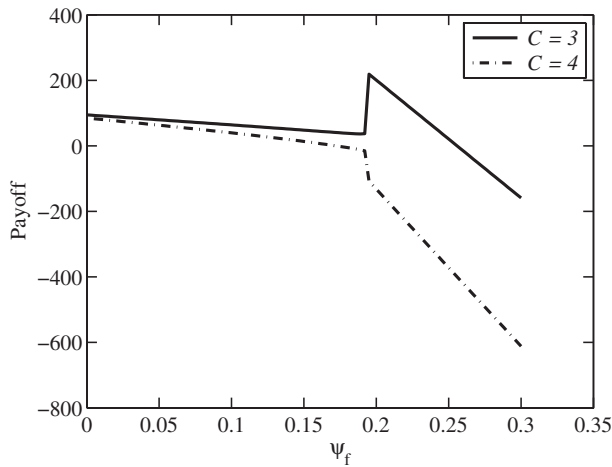


Figure 5. Dependence of pay-off $E(\psi, \psi_0)$ on the individual ITC level of treatment ψ_f for different relative costs of treatments C , $\psi_0 = .003$, $x = .03$, and other parameter values as in Table 1. The jump occurs exactly when the average level of ITC usage ψ is at the critical value yielding the eradication of the disease. However, the value at which the jump occurs is not always optimal. Compare to Figure 4.

$$F = \mu(\nu + \mu)(g + \sigma) \quad (30)$$

$$G = \Lambda(g + \nu + \mu) \quad (31)$$

$$H = \Lambda(\nu + \mu)(g + \sigma). \quad (32)$$

Substituting (21) into (20) then yields the explicit formula for the focal farmer's net benefits, $E(\psi_f, \psi_0)$.

First, assume that everybody uses the same level of ITC. Figure 2 shows the pay-off $E(\psi, \psi)$ for various costs of treatment C , depending on the ITC level ψ . Note the sharp jump at pay-offs that happens exactly at the value of ψ where λ becomes 0 and where the number of cattle jumps from a relatively low to relatively large value, see Figure 3. For relatively small values of C , such a critical value of ψ is optimal from the population perspective (as it yields the highest pay-off for the population as a whole) and it is also a value of ITC usage that eradicates the tsetse flies and the disease. The disease is eradicated at the same value of ψ even when the cost of ITC is large. However, for such a large C , the critical level of ITC is no longer optimal and the pay-off (from the population perspective) is the highest when no ITC is used at all.

We note that while the critical value ψ is relatively small (less than 1%), the cost of using ITC may still be too large for some poorer farmers or some farmers may simply choose not to use it. In order to complete the analysis, we thus need to consider a focal farmer using ψ_f in the population of other farmers using ψ_0 on average and determine the optimal strategy for such a farmer. Figure 4 shows that when C is sufficiently small (below the threshold from Figure 2), the optimal ITC usage for a focal farmer is exactly such that the overall average ITC usage is at the critical value, resulting in the elimination of the disease. However, Figure 4(b) illustrates that when the cost of using ITC is large, the optimal usage for the individual farmer is 0. The same is true when the cost of using ITC is relatively small, but the farmer does not own enough cattle, see Figure 5.

4. Conclusions

Following Bauch and Earn (2004), we provided a game-theoretical model to study the optimal usage of ITC to combat African sleeping sickness. Our analysis is built on the transmission model of sleeping sickness by Kajunguri et al. (2014) that studied the efficiency of ITC. For numerical simulations, we focused on the restricted application of ITC, the formulae and results for whole body applications are similar.

Not surprisingly, our results indicate that the optimal usage of ITC depends on the cost of ITC. From the population perspective (for example, when the usage is mandated/subsidied by the government), when the cost of ITC is not prohibitive, the optimal usage of ITC yields a complete eradication of the disease; otherwise, when the cost of ITC treatments is above a certain threshold, it is optimal not to use ITC at all.

We found that eradication of African sleeping sickness by voluntary usage of ITC is possible (again, when the cost of treatment is not prohibitively large). This finding is in stark contrast with previous work on voluntary disease prevention, see for example (Bauch, 2005; Bauch & Earn, 2004; Geoffard & Philipson, 1997). As discussed in Hargrove, Torr, & Kindness (2003) and Kajunguri et al. (2014), the real-world success of ITC depends on many factors such as the size and shape of the controlled region and the number or density of treated cattle. Nevertheless, application of ITC can indeed lead to reduction of the incidence of sleeping sickness cases (Magona & Walubengo, 2011).

The main reason for such a difference between findings of our analysis and the previous work such as Bauch and Earn (2004) lies in the highly bimodal distribution of the benefits. In the presence of the disease (i.e. when ITC usage is not enough to eradicate it), the equilibrium level of cattle and consequently the farmer benefits are much smaller than when the ITC usage is at or above the eradication levels. We note that the same feature is present in the original Kajunguri et al. (2014) model from which we differed only in treating the newly recruited cattle directly. When the cost of ITC is not large and when the farmer has a sufficiently large population of cattle that his decision can significantly affect the average usage level, the individual interests of the farmer aligns well with the interest of the population and the disease can be eradicated.

We note that our model can be further improved. We used a simple pay-off function (20) that can be made more realistic by including exact benefits of owning a cow (a per day benefit as well as per cow benefit) as well as exact cost of the treatment (set-up costs and per cow cost). Moreover, the added benefits of ITC include tick control (Bourn et al., 2005; Torr, Maudlin, & Vale, 2007; Vale & Torr, 2005) are not included in our model. For the sake of simplicity, we also used only the single-host transmission model from Kajunguri et al. (2014) while multi-host models (including humans) are also available and we did not include the benefits of smaller disease incidence among humans in our model either. Nevertheless, we speculate that adding extra benefits or extending the model for the multi-host situation will not qualitatively change the results. The situation may be different if tsetse flies start to develop resistance against the treatment. As suggested in Magona and Walubengo (2011), a combination of different treatments is likely to minimize the occurrence and spread of the resistant parasites.

Also, individuals can have different perceptions of the real situation (Poletti, Ajelli, & Merler, 2011) and can base their decision also on different social aspects (Xia & Liu, 2013). Providing proper information may thus play an important role in disease eradication. As

noted in Kajunguri et al. (2014), the model presented here does not work when cattle populations are migratory and travel widely through tsetse infested country. Nevertheless, game-theoretical models, in general, can still capture the interplay between disease prevalence, protective action coverage and individual behaviour (Bauch, 2005) and we can argue that since the real benefits to an individual farmer exceeds the benefits considered in our simple model, the optimal usage of ITC yields an eradication of the disease even when the ITC is voluntary.

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Disclosure statement

No potential conflict of interest was reported by the authors.

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