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Running title:

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Word Count:

Keywords: Individual Based Model, Raccoon, Macroparasite, Wildlife disease, Zoonosis, Parasite, Management. . . .

Number of figures:

Article type:

1 Introduction

Rinderpest is eliminated (), rabies is reduced (), and other wildlife microparasite epidemics something. These management feats have been guided by mathematical models. While numerous models exist for managing viral, bacterial and protozoan agents in wildlife (), models directed towards managing wildlife macroparasites are lacking (McCallum 2017). This is because macroparasites, like helminths, require more complex models that incorporate infection load, heterogeneity and complex life cycles. These complexities often make it untenable to use standard modeling approaches (e.g. ordinary differential equations at the population-level) [WC] and individual-based models are often needed when modeling macroparasite systems (McCallum and others). The advantage of using individual-based models for host-macroparasite systems is that it is straight-forward to capture the aforementioned complexities of macroparasite systems and their effects on host-parasite dynamics can be explored. This is useful when the goal is, for example, to use the model to evaluate the efficacy of different management strategies on some measure of macroparasite disease risk [Drawert, others]. However, this additional complexity almost guarantees that the model will not be analytically tractable, such that intensive simulation is necessary to explore the characteristics of the model. Moreover, individual-based models often have a number of unknown parameters and it is unclear whether standard parasitological data, such as age-intensity profiles, is sufficient to estimate these model parameters. Here, we build an individual-based macroparasite model for the widespread zoonotic raccoon parasite, raccoon roundworm (*Baylisascaris procyonis*). We show how this model can be parameterized this model with commonly collected parasitological data and then use this evaluate management strategies that target both host and parasite populations. [General concluding sentence showing the knowledge gap we are filling in disease ecology]

Raccoon roundworm is increasingly recognized as a threat to both human and wildlife health (Page, Graeff, Kazacos), but, like many macroparasites, is difficult to control due to its complex life cycle and resistant environmental infectious stages. Worms mature in the raccoon gut and

infected raccoons can release over a million parasite eggs per day (). Eggs survive for over a year (Shafir) and accumulate at communal raccoon defecation sites termed latrines (). Individual raccoons contribute to multiple latrines and these contaminated sites expose raccoons and other species to eggs. Eggs infect juvenile raccoons, leading to high parasite loads in animals as young as four months (Weinstein 2016, Kazacos, Boyce). Raccoon susceptibility to eggs declines with age and adult raccoons acquire worms by eating infected small mammals and birds (). In these birds and mammals, larval worms do not mature. Instead, larval worms migrate from the gut into other tissues including the brain, often causing fatal neurological damage. Raccoons scavenge these incapacitated animals (), and this trophic transmission maintains infection in older raccoons that are less susceptible to eggs (). Nearly all infected raccoons release eggs; however higher parasite loads in juveniles suggests that this age class contributes disproportionately to human disease risk().

Human risk might be reduced by removing *B. procyonis* eggs from the environment, treating infected raccoons, or reducing raccoon populations. Culling is often considered the simplest and cheapest disease control strategy (). However, although a common component of rabies control (Rosatte et al., 1986), culling is controversial and not always effective (Choisy and Rohani, 2006; Woodroffe et al., 2006; Morders et al., 2013). Birth control is a less controversial method to reduce wildlife populations (smith 2002), but is often slower, more expensive, and less effective than culling (). As an alternative to reducing host populations, parasite populations can be reduced through mass drug administration (Qureshi 1994 et al, deer fascioloides) or vaccination (anthrax antelope De vos 1973 Koedoe, bison; elk brucellosis (rabies). Although no *Baylisascaris* vaccines exist, the mass baiting methods developed for rabies vaccines can also deliver deworming medication (anthelmintics) (Smyser et al., 2015). These anthelmintic baits can reduce *B. procyonis* prevalence in wild raccoons, but raccoons are rapidly re-infected unless latrines are also removed (Page et al., 2011; Page et al., 2014). Latrine, parasite, and raccoon removal all could reduce human *B. procyonis* exposure; however, experiments that compare these strategies are difficult to implement in wild animal populations.

Here we build an individual based macroparasite model to examine raccoon roundworm management strategies. Parameterized with values from literature and field data, this model captures the seasonal and age based infection heterogeneity commonly observed in wild raccoon populations. With this model, we examined how culling, birth control, anthelmintic baiting, and latrine clean-up affected raccoon populations, parasite populations, and human disease risk. We find KEY RESULTS. . . And implication about raccoons/general macroparasite management.

2 Methods

2.1 Model Description

To evaluate raccoon roundworm management strategies, we built an individual-based model in R () for a macroparasite with long-live environmental infectious stages and complex life cycle. We describe this model using the Overview, Design concepts, Details (ODD) protocol (Grimm et al. 2006, 2010), providing first an Overview of the model, then describing the Design Concepts, followed by the Details of submodels, parameterization and initialization. Additional details on submodels and annotated code are available in the Supplementary Material.

2.2 Overview

2.2.1 Purpose

We developed an individual-based macroparasite model for raccoon and raccoon roundworm to compare how host, parasite, and infective stage targeted management affect host demography, parasite demography and human exposure.

2.2.2 Entities, state variables, and scales

In this model, raccoon are the main model entity and exist as an age structured, closed all-female population. Five state variables and attributes describe each raccoon: its identification number, age, alive or dead status, location and parasite load. We track the female parasite load

(infrapopulation) within each raccoon. Raccoons can acquire new worms at each time step and, for each raccoon, we explicitly track each worm cohorts' age. Worms accumulate (and die) over time, and we calculate parasite load by summing parasite cohorts within each raccoon. Infected raccoons contaminate the environment with long-lived parasite eggs, which we model as an environmental egg pool. As even a single worm can produce more than a million eggs per month, we assume that infected raccoons rapidly saturate local latrines with eggs. Thus, we do not explicitly track each egg, and instead use infection prevalence in the local raccoon population as a proxy for local egg production. Eggs decay over time () and we calculate the current environmental egg pool as a weighted sum of past egg production. Eggs infect rodents and we use the environmental egg pool to estimate rodent infection at each time step.

We track hosts, parasites, and environmental egg pools in a spatially implicit 20 km² world with 10 equally sized 2 km² zones. Human population density increases consecutively from zones one through ten (Fig X, x) and zones with higher human density support larger raccoon populations (). We track the raccoon population, parasite population, and egg pool in each zone at monthly time steps for 400 months, using the egg pool in each zone to calculate total human risk within the simulated world.

2.2.3 Process overview and scheduling

We update the model at each monthly time step, first determining individual raccoon survival. Surviving raccoons then reproduce, lose parasites, gain parasites, disperse, age, and contribute to the environmental egg pool. Following the typical raccoon life cycle, raccoons reproduce once per year, with kits entering the population in the following time step. All other events occur monthly in all ten zones.

2.3 Design Concepts

2.3.1 Emergence and Interactions

Although some infectious agents can regulate their hosts (), raccoon roundworm does not regulate raccoons. Unmanaged raccoon populations are limited by resource availability (), which we model by setting a raccoon carrying capacity in each zone. Following the Ricker function (Gurney and Nisbet, Hastings et al 2012), as raccoon population density increases within a zone, reproduction declines. Simulated raccoons reproduce once per year (in the spring), and this pulsed reproduction produces realistic annual population cycles and age distributions for an unmanaged raccoon population (gehr, others).

We add macroparasites to this simulated raccoon population following classic macroparasite modeling theory (eg. Anderson and May, Cornell, McCallum). Individual host-parasite interactions generate an aggregated parasite population with realistic age-intensity and age-prevalence profiles. [is this emergent? given we allow for overdispersion in how many worms raccoons pick up]. The parasite population cycles due to changing host demography, and these emergent patterns match the widely observed fall rise in mean raccoon parasite loads (eg page 2016, weinstein2016, many others).

2.3.2 Stochasticity

We determine model events with a random draw from a parameterized probability distribution using random number generators in R version X (R core Team). As a result, each parameter includes both a mean value and a distribution .

2.3.3 Initialization

We began each simulation with 500 two-year old raccoons stochastically distributed throughout the 20 km² world according to zone specific carrying capacities (average starting zone density:). Initially, each raccoon hosted 10 worms and no eggs contaminated any zones. The mean

rodent worm load (μ) was initially $\mu = 3.49$ based on empirical observations [citation] and the variance of rodent worms was set to 87.08 based on the canonical scaling relationship between log-mean parasite load and log variance (Shaw and Dobson 1995).

2.4 Submodels

2.4.1 Raccoon death

We assume that all raccoons experience a constant per month death probability and then augment this age-independent death rate to account for high nestling mortality (Montgomery 1969) and senescence. Nestling raccoons (<1 month) have constant added risk of death and we model senescence as an increasing age-specific hazard rate, parameterized to ensure that raccoons cannot live past 20 years old. These three mortality processes lead to a “bathtub”-shaped hazard function for raccoon mortality (Fig. X)

Raccoons can also die from *B. procyonis* infection if worms cause an intestinal obstruction (Stone, 1983; Carlson and Nielsen, 1984). Fatal obstructions can occur at loads as low as 141 worms and have been recorded in animals with 636 and 1,321 worms (Stone, 1983; Carlson and Nielsen, 1984), however raccoons can survive with over 200 worms (Kazacos 2001). To model this potential parasite-induced host mortality, we used the logistic parasite-induced mortality function $1 - \frac{e^{\beta + \alpha \log(\text{load} + 1)}}{1 + e^{\beta + \alpha \log(\text{load} + 1)}}$, where α gives the per parasite effect on the log odds raccoon survival probability and β roughly corresponds to the threshold at which parasite-induced mortality begins to occur. We estimated these parameters using the aforementioned data and]. Our parameterization corresponds with the empirical observation that parasite-induced mortality is not a large factor affecting raccoon dynamics.

2.4.2 Raccoon reproduction

Raccoons typically reproduce once a year and can have up to 2 female young per litter (Gehrt 2001, Fritzell 1985, Cowan 1973, Clark et al 1989). Although there is some evidence that popu-

lations in the far south of the range may breed year round (Troyer et al 2014) and late litters do occur (Gehrt and Fritzell 1996, 1978a?), here we assume that all reproduction occurs in a single spring pulse.

The model assumes that average raccoon litter size in a given year decreases with increasing raccoon density. This density-dependence corresponds with reproduction in dense populations being limited by both the availability of nest sites and increased infanticide risk in dense populations (Gehrt and Fritzell 1999; hauer 2010; Wolff 1997). Specifically, we model this density-dependence following a Ricker function (Table X).

2.4.3 Worm transmission

Raccoons acquire *B. procyonis* through two transmission routes: encounter with *B. procyonis* eggs in the environment and by consuming other animals already infected with *B. procyonis* adults (e.g. rodents) (citations).

2.4.4 Worm death

2.5 Fitting Susceptibility and Transmission Parameters with ABC-SMC

As described in Table X, most of the parameters in this model were estimated based on previous *B. procyonis* studies in other systems and previous work by S.W. in the current system. However, there were five transmission and susceptibility parameters in the model that were largely unknown in the raccoon-*B. procyonis* system. These were the mean number of eggs encountered by a raccoon in the environment over a monthly time step, the variability in the number of eggs encountered over a monthly time step, an egg encounter parameter relating the eggs in the environment to the probability of a raccoon encountering an egg, the probability of a raccoon consuming a rodent in a month, and a parameter determining how concomitant immunity increases with raccoon age (Table X).

To estimate these parameters, we used Approximate Bayesian Computing with Sequential

Monte Carlo (ABC-SMC) (Sisson et al. 2007, Beaumont 2010, Kosmala et al. 2015). ABC is typically referred to as a likelihood free parameter estimation method and is often used with simulation-based models in which the likelihood of a data point given the model is intractable to compute. Instead, ABC uses a user-defined distance function to determine how “well” the simulation model parameterized with a set of parameters drawn from some prior distribution can reproduce the observed data. This approach allows one to estimate an approximate posterior distribution for unknown parameters in model based on how well they can reproduce observed data.

To use ABC to parameterize the unknown parameters in the raccoon-*B. procyonis* IBM, we used observed age-abundance and prevalence profiles for raccoons and *B. procyonis* in our system of interest (Weinstein 2017, Fig. X). For many mammal macroparasite systems, this type of intensity and prevalence data is the best available parasitological data as longitudinal, individual-based measures of worm intensity are nearly impossible due to both the difficulty of re-trapping individual hosts and the destructive nature of measuring parasite loads [though I think fecal egg counts can give a good estimate of infection intensity in some cases...Sara?]. Therefore, developing methods to estimate transmission and susceptibility parameters from this type of data is important for a wide-range of host-macroparasite systems.

Using this age-intensity/prevalence data for the raccoon-*B. procyonis* system, we then implemented the ABC-SMC algorithm as follows (citations)

1. We specified uniform priors around our five unknown transmission and susceptibility parameters: [PRIORS HERE]
2. We drew 10,000 samples from each prior distribution such that we had 10,000 vectors of parameters (in the ABC literature each parameter vector is called a particle).
3. For each of the 10,000 particles, we simulated our model for 100 months to remove the effects of the initial conditions.
4. For a given simulation, we then sampled 189 raccoons over the last 24 months of the

simulation which is consistent with the number of raccoons used to build the age- abundance/prevalence profiles given in Weinstein (2017). Specifically, we distributed these 189 sampled raccoons according the eight age classes given in Weinstein (2017).

5. Using this sample, we then constructed the age-abundance/prevalence profile from the simulated data and compared it to the observed age- abundance/prevalence.
6. To compare the simulated and observed data, we first combined the eight observed age-abundance data points and the eight observed age-prevalence data points into a vector $S_{obs} = [age-intensity_1, age-intensity_2, \dots, age-intensity_8, age-prev_1, age-prev_2, \dots, age-prev_8]$. We did the same thing for all 10,000 simulated age-abundance/prevalence profiles such that we had a $10,000 \times 16$ matrix S_{sim} where the columns matched the 16 dimensions in S_{obs} and the rows corresponded to one of the randomly drawn particles.
7. To put the abundance measures and the prevalence data on the same scale, we standardized each column in S_{sim} by subtracting the mean and dividing by the standard deviation of each column. We then performed this identical transformation on S_{obs} *based on the column-wise means and standard deviations from S_{sim}* . This transformation of S_{obs} put the values of S_{obs} in terms of deviations relative to the mean of the simulated data.
8. We then calculated the Euclidean distance between each row in S_{sim} and S_{obs} (i.e. the L2 norm), which resulted in 10,000 distance measures. We then accepted the 1,000 particles that resulted in the 1,000 smallest distances.
9. We then equally weighted each of these 1,000 accepted particles and resampled them with replacement 10,000 times. The upon sampling a particle, we perturbed each parameter in a particle using $\sigma_i U(-1, 1)$ where σ_i is the standard deviation of the i th parameter in the 1,000 accepted parameters. The perturbation helps the algorithm explore nearby parameter space.
10. We then repeated 3- 9 with the 10,000 perturbed parameters [five] times with one important

change to step 9 after the first round of sampling. After identifying 1,000 accepted particles we used the following function to weight each particle: [X]. This is the importance weighting of a particle that accounts for the fact that particles are no longer being sampled from the prior distribution (Beaumont 2010; DOUBLE CHECK WORDING).

11. After the fifth iteration [check], the particles converged on a posterior distribution [How did we determine this...deviation in posterior standard deviation and mean].

This algorithm provided posterior distributions for the five unknown transmission and susceptibility parameters in the raccoon-*B. procyonis* model using commonly obtained age-abundance/prevalence profiles.

[TODO...not sure if this is true yet: We confirmed this algorithm could also recover known transmission and susceptibility parameters based on simulated age-abundance and age-prevalence profiles. (Some parameters might be unidentifiable...)]

2.6 Management Strategies for *B. procyonis*

We used our fully parameterized model to explore of how different management strategies affected the dynamics of worm and raccoon populations and human risk from *B. procyonis*. We considered four distinct management strategies in addition to combinations of these management strategies: culling raccoons, birth control of raccoons, anti-helmenthic baiting, and latrine cleanup. For each of these management strategies, we considered different levels of management effort (see below for definitions) and explored how different management strategy-by-effort combinations affected four major predictions of the model: 1. The total raccoon population 2. The total worm population 3. The level of human risk [define above] and 4. The mean raccoon worm load. For each management strategy, the model was run for 200 months to obtain equilibrium dynamics, at which time the management strategy was enacted monthly for an additional 200 months (FIG. X). Upon completion of the simulation, the four major model predictions listed above were computed based on the mean [max, min, var, too] over the last 24 months of the

simulations.

The different management strategies were implemented as follows

Culling raccoons [This section could probably use some background on culling...]

To implement raccoon culling, we assumed that some number of traps were randomly distributed in the raccoon world each month. We assumed that per trap probability of catching a raccoon increased with raccoon density following a type I functional response: $1 - \exp(-\text{raccoon density} * \gamma)$. γ was estimated based raccoon trapping data from Prange et al. (2003) and Graser et al. (2012) [probably more description here...]. Finally, we assumed that the number of raccoons were caught each month followed a $\text{num_trapped} \sim \text{Binomial}(N, p)$ where the p = probability of success was the per trap probability and the N = number of trials was the number traps deployed that month. Because we assumed that all individual raccoons were equally likely to be trapped, we then randomly selected $\min(\text{num_trapped}, \text{total population size})$ raccoons from the population to cull in that time step.

We also allowed trapping to occur in specific zones of human overlap and only for particular age raccoons. When trapping in a particular zone(s) of human overlap, trapping probability was determined only by the density of raccoons in that zone(s) and only raccoons in that zone could be trapped and culled. When trapping for a particular target age class, trapping proceeded as described above, but raccoons that were not in the target age class were released.

Birth Control [This section could probably use some background on birth control]

Birth control followed the exact same trapping regime as culling. However, upon trapping a raccoon its reproductive ability was set to zero for the remainder of its life [TODO: define reproductive ability above]

Anti-helminthic baiting [This section could probably use some background on baiting]

We implemented anti-helminthic baiting by first randomly distributing some number of baits into the raccoon world. Of the initial number of baits distributed into the world [WC], on average 60% of these baits were degraded or consumed by animals other than raccoons. For the remaining baits, we assumed that all raccoons were equally likely to consume a bait and randomly assigned

baits to raccoons in the world. A given raccoon could consume multiple baits in a month. Upon consuming a bait, all the worms in a raccoon were killed, but the raccoon could immediately start acquiring worms again from the environment and rodents. Consuming one bait or multiple baits in a month had identical effects on raccoon worm loads. Similar to culling and birth control, we also allowed for anti-helminthic baiting in specific zones of human overlap.

Latrine cleanup [again, section needs some background]

The final management strategy we explored was latrine cleanup. [Define latrine here or above?]. This strategy was implemented reducing or eliminating the environmental egg pool each month. Once again, this strategy could be specifically implemented in specific zones of human overlap.

3 Results

Fitting transmission parameters with age-abundance/prevalence profiles

Figure in this section

- Plots of posterior distributions of parameters transmission parameters
- Plots of observed and predicted age-abundance/prevalence profiles
- Plots of model dynamics without any management after fitting

Management strategies for reducing human risk of B. procyonis infection

Figures in this section

- Heat maps of various strategies
- Heat maps of strategy combinations

4 Discussion

When we talk about birth control- There is currently no approved raccoon contraceptive (), although products developed for dogs and cats could be optimized for raccoons given high cost and not great results, strategy doesn't seem worth pursuing.

Culling- potential for a "Hydra effect": It's got a name! Abrams 2009. When does greater mortality increase population size? the long history and diverse mechanisms underlying the hydra effect. Ecology Letters 12:462–474 (named for greek myth, where chopping of one hydra head caused two to grow back. While density dependent birth rates set up possibility, and do lead to increased prev/intensity in animals, overall reduction in rac population still reduces human risk, although not as quickly perhaps as would if no "hydra effect" occurred.

5 Acknowledgments

Acknowledgments, including funding information, should appear in a brief statement at the end of the body of the text. Acknowledgments of specific author contributions to the paper should appear here.

Tables (1 per page)

Figure Legends

Figures (1 per page, labeled "Figure 1", etc)

- Figure sizes should be no more than 6 inches wide and 7 inches high. When possible, submit figures in the size you wish to have them appear in the journal. Most illustrations, except some maps and very wide graphs, should be 1-column width (3 inches) at a resolution of 600 dpi.
- The font size of the x - and y -axis numbers should be slightly smaller than the axis label. A consistent font (Helvetica is preferred) should be used throughout. Use boldface type only if required for journal style. Use sentence case (i.e., only capitalize the first word) for axis

titles, labels, and legends.

- For symbols and lines, avoid very small sizes and line thicknesses (1 point width stroke or greater is preferable). All elements of a figure should appear with the same degree of intensity. If different degrees of intensity need to be conveyed, lines should differ by 1 point width for clarity.

Appendices