EXTINCTION TIMES IN THE SUBCRITICAL STOCHASTIC SIS LOGISTIC EPIDEMIC

GRAHAM BRIGHTWELL, THOMAS HOUSE, AND MALWINA LUCZAK

ABSTRACT. Many real epidemics of an infectious disease are not straightforwardly super- or sub-critical, and the understanding of epidemic models that exhibit such complexity has been identified as a priority for theoretical work. We provide insights into the near-critical regime by considering the stochastic SIS logistic epidemic, a well-known birth-and-death chain used to model the spread of an epidemic within a population of a given size N.

We study the behaviour of the process as the population size N tends to infinity. Our results cover the entire subcritical regime, including the "barely subcritical" regime, where the recovery rate exceeds the infection rate by an amount that tends to 0 as $N \to \infty$ but more slowly than $N^{-1/2}$. We derive precise asymptotics for the distribution of the extinction time and the total number of cases throughout the subcritical regime, give a detailed description of the course of the epidemic, and compare to numerical results for a range of parameter values. We hypothesise that features of the course of the epidemic will be seen in a wide class of other epidemic models, and we use real data to provide some tentative and preliminary support for this theory.

1. INTRODUCTION

1.1. Epidemiological motivation. Models of the dynamics of disease spread are widely used throughout infectious disease epidemiology, and inform an increasing number of health policy domains (Heesterbeek et al. 2015). Typically, epidemic models have a quantity called the basic reproduction ratio \mathcal{R}_0 such that, if $\mathcal{R}_0 > 1$, then the epidemic is supercritical and grows exponentially and, if $\mathcal{R}_0 < 1$, then the epidemic is subcritical and shrinks exponentially; see for instance Diekmann, Heesterbeek and Britton (2013).

Increasingly, however, epidemiologists are confronted with epidemics where the dynamics cross the threshold from supercritical to subcritical (see for instance Klepac et al (2013) and other papers in that journal issue) due

Date: January 19, 2018.

²⁰⁰⁰ Mathematics Subject Classification. 60J27, 92D30.

Key words and phrases. stochastic SIS logistic epidemic; birth-and-death chain; time to extinction; near-critical epidemic.

Thomas House is supported by the EPSRC, grant reference EP/N033701/1.

The research of Malwina Luczak was largely carried out at Queen Mary, University of London, and was supported by an EPSRC Leadership Fellowship, grant reference EP/J004022/2.

to control measures, or from subcritical to supercritical due, for example, to mutation (Antia et al. 2003; Bull and Dykhuizen 2003; Scheffer et al. 2009; O'Regan and Drake 2013), or where the behaviour is not exponential (Chowell et al. 2016). This has led to the understanding of near-critical epidemics being highlighted as a key challenge for disease-dynamic models of infectious diseases (Britton et al. 2015).

In this paper, we are interested in understanding the course of subcritical epidemics, especially where \mathcal{R}_0 is close to 1. We shall give a detailed analytical study of a particularly simple epidemic process, the *stochastic SIS logistic process* (also called the SIS model, the contact process, or the logistic model) in the subcritical regime. In this model, each member of a population of fixed size N is either susceptible or infective. Infective individuals encounter a random other member of the population at rate λ , and infect them if they are susceptible: infective individuals recover at rate μ , and once recovered they are immediately susceptible again. The state of the epidemic is effectively determined by the number $X_N(t)$ of infectives, or by the *prevalence* $X_N(t)/N$, at time t. We defer the formal definitions to the next section. For most purposes, this process is too simple to reflect the full behaviour of a real-world epidemic, but it can be suitable for modelling sexually transmitted and hospital-acquired infections (Eames and Keeling 2002; Ross and Taimre 2007).

In our model, the basic reproduction ratio \mathcal{R}_0 is equal to λ/μ , and we are especially interested in the regime where \mathcal{R}_0 tends to 1 from below as N tends to infinity. When also $(1 - \mathcal{R}_0)N^{1/2} \to \infty$, we are in the *barely* subcritical regime. We note the following features of the typical course of the epidemic in this regime, all of which distinguish regimes near criticality from those where \mathcal{R}_0 is fixed and less than 1. We make the hypothesis that these features are common to a wide class of epidemic models (not only SIS models) in barely subcritical regimes. We express our statements in terms of the basic reproduction ratio \mathcal{R}_0 , the population size N, and a speed parameter μ , where $1/\mu$ is roughly the expected duration of an individual case.

- The time to extinction is of order much larger than $\log N/\mu$ (typically it is of order $\log N/\mu(1-\mathcal{R}_0)$).
- There is a period of time before extinction, of order $1/\mu(1 \mathcal{R}_0)$, where the number of infectives follows a track resembling a random walk, remaining of order at most $1/(1 \mathcal{R}_0)$ throughout. The duration of this period is not well-concentrated.

The *cut-off phenomenon*, in the case of a stochastic epidemic process, is that the typical time to extinction is much greater than the window of time over which the probability of extinction goes from near 0 (at this stage, the number of infectives will typically be larger than $1/(1 - \mathcal{R}_0)$, but much smaller than the population size) to near 1. The SIS logistic process exhibits this phenomenon in the subcritical regime, with the expected time to extinction around $\log[N(1 - \mathcal{R}_0)^2]/\mu(1 - \mathcal{R}_0)$, and the window where extinction typically occurs having width of order $1/\mu(1 - \mathcal{R}_0)$: thus cut-off becomes less pronounced as we approach the *critical regime* where $|1 - \mathcal{R}_0|$ is of order at most $N^{-1/2}$. We hypothesise that this weakening of cut-off is also a feature of many barely subcritical epidemic processes.

Figure 1 shows three real examples of the courses of epidemics after they became subcritical due to control efforts. These concern: the recent Ebola epidemic in Sierra Leone (top plot), smallpox (middle plot), and polio (bottom plot). For all three examples, we superimpose curves showing smooth decay (in the top plot the curves are solutions of an ODE: in the others, they are exponential decay curves at various rates r), illustrating that these provide a poor fit for the observed detailed dynamics of the disease over time, especially in the end stages of the epidemic (see in particular the right hand panels in the bottom two plots, which are rescaled versions of the plots showing the behaviour more clearly). However, at a glance, the plots do exhibit behaviour very broadly in line with our expectations for a barely subcritical epidemic. We discuss the data in more detail in Appendix A.

For comparison, Figure 2 shows the behaviour of the stochastic SIS logistic process for a population of size 10^6 with 10^3 initial cases as λ is increased towards $\mu = 1$ from below. These sample paths are compared to "smooth decay" curves of the form $e^{(\lambda-\mu)t}$; the figure illustrates our analytic results for the model: individual realisations of the stochastic SIS logistic process deviate significantly from smooth exponential decay in the final stages, and exhibit the behaviour we described above.

The potential implications of our work for more general epidemic models are discussed in more detail in Section 9.

Our results on the stochastic SIS logistic process are the first to incorporate the barely subcritical regime, where \mathcal{R}_0 approaches 1 from below as the population size tends to infinity. We provide analytic methods for studying the SIS logistic process in the large-population limit, and thereby obtain precise asymptotic results for the distribution of extinction times and the total number of infection events. We also give attention to numerical approaches that are well adapted to the near-critical regime and allow the exploration of model behaviour at a given value of the population size. We expect that our methodology can be generalised to apply to other models of epidemiological and biological interest.

1.2. Technical background and outline of paper. The stochastic SIS logistic process is defined as follows. Given a "size parameter" N, and two further parameters λ and μ , let $X_N = (X_N(t))_{t\geq 0}$ be the continuous-time Markov chain with state space $\{0, \ldots, N\}$, and transitions as follows:

$$X \rightarrow X+1$$
 at rate $\lambda X(1-X/N)$, (1.1)
 $X \rightarrow X-1$ at rate μX .

This is the most basic stochastic model of the spread of an SIS (susceptibleinfective-susceptible) epidemic within a population of size N. In this context, $X_N(t)$ represents the number of infective individuals at time t. Each infective encounters a random other member of the population at rate λ ; if the other individual is currently susceptible, they become infective. Also, each infective recovers at rate μ ; once they are recovered they become susceptible again. The stochastic SIS logistic process is also used as a model for a metapopulation process, where N represents the number of available patches, $X_N(t)$ is the number of patches that are populated at time t, λ represents the rate at which one existing colony attempts to colonise another patch, and μ represents the rate at which an entire patch becomes depopulated due to some catastrophe. The model was first formulated by Feller (1939), and further studied by Bartlett (1957). It was rediscovered by Weiss and Dishon (1971), and has since been investigated by many authors. A recent thorough treatment of the model is the book of Nåsell (2011), who in particular mentions a number of other application areas and gives a large list of further references.

Suppose to begin with that λ and μ are fixed constants. The key parameter is the ratio λ/μ , which, for a small-scale epidemic, approximates the basic reproduction ratio \mathcal{R}_0 of the epidemic, defined as the mean number of individuals infected by a single infective in a large population of susceptibles. The behaviour of the logistic process is radically different depending on whether the quantity \mathcal{R}_0 is greater or less than 1. In the case $\mathcal{R}_0 > 1$, the process typically takes a time exponential in N to die out, spending most of its duration near to the value $(\lambda - \mu)N/\lambda$ where the upward and downward transition rates are equal: this models a *supercritical* epidemic, where an initially small number of infectives may generate an outbreak that becomes endemic in the population for a very long period. In the case $\mathcal{R}_0 < 1$, the process is always drifting downwards, and even an initially very large epidemic dies out with high probability in time of order $\log N$. In this paper, we give precise results about the distribution of the extinction time in this subcritical case. In the special case $\mathcal{R}_0 = 1$, the expected extinction time is of order \sqrt{N} .

To study behaviour in the transition between the supercritical and subcritical regimes, we regard λ and μ as functions of the size parameter N, and we pay special attention to cases where $\mathcal{R}_0 = \lambda/\mu$ tends to 1 as N tends to infinity.

Earlier work (Nåsell 1996; Dolgoarshinnykh and Lalley 2006; Kessler 2008) has demonstrated that there is a *critical window* for the logistic process where $|\mathcal{R}_0 - 1| = O(N^{-1/2})$; the change in the nature of the process occurs as \mathcal{R}_0 crosses this window. Our main aim in this paper is to give the distribution of the extinction time throughout the entire subcritical range, i.e., as long as $(1 - \mathcal{R}_0)N^{1/2} \to \infty$. Our results show exactly how the extinction time changes from order log N to order \sqrt{N} as \mathcal{R}_0 approaches 1 from below.

Our results for the case where λ and μ are fixed real numbers with $\lambda < \mu$ are new, but our main interest is in the case where $\lambda = \lambda(N)$ and $\mu = \mu(N)$ are functions of N with $\mu(N) - \lambda(N) \to 0^+$. We shall always assume that $\mu(N)$ and $\lambda(N)$ are bounded away from both 0 and ∞ . We choose to state all our results in terms of the two parameters λ and μ for ease of comparison to earlier results: however, apart from a constant factor determining the speed of the epidemic, all of our results could be restated in terms of \mathcal{R}_0 . Note that, under our assumptions, $1 - \mathcal{R}_0 = (\mu - \lambda)/\mu$ is of the same order as $\mu - \lambda$, and to say that our sequence of parameter values is in the *subcritical regime* means that $(\mu - \lambda)N^{1/2} \to \infty$.

We also allow the initial state $X_N(0)$ to depend on N. One case of natural interest is where $X_N(0) \simeq \alpha N$ for some $\alpha \in (0, 1]$, but our results also cover the case where $X_N(0)/N \to 0$. We set T_N to be the time to extinction (i.e., the hitting time of the absorbing state 0) for $X_N(t)$, with infection rate $\lambda = \lambda(N)$, recovery rate $\mu = \mu(N)$, and initial state $X_N(0)$. Our interest is in the asymptotic distribution of T_N , as $N \to \infty$.

There is an exact expression for $\mathbb{E}T_N$ as a double summation, due to Weiss and Dishon (1971) in the case where $X_N(0) = N$, and in general to Leigh (1981) and Norden (1982). The asymptotics of this sum have been determined in some cases, e.g., by Doering, Sargsyan and Sander (2005). Our methods give precise information about the distribution of T_N , not just its expectation.

The logistic process is naturally associated with the differential equation

$$\frac{\mathrm{d}x}{\mathrm{d}t} = \lambda x (1-x) - \mu x = \lambda x (1-\mu/\lambda - x), \qquad (1.2)$$

where x(t) represents the proportion of infective individuals at time t. This equation was first studied by Verhulst (1838), and it is known as the Verhulst equation or logistic equation. It follows from the general theory of Kurtz (1971) that, as $N \to \infty$, $X_N(t)/N$ is well concentrated around the solution x(t) of the differential equation (1.2), uniformly over fixed time intervals, as long as $X_N(0)/N$ is well approximated by its initial condition x(0). For our purposes, we need to show concentration for longer periods, and this is possible thanks to the special structure of the logistic equation when $\mu \ge \lambda$.

The behaviour of the deterministic process x(t) also depends on whether \mathcal{R}_0 is greater than, equal to, or less than 1. In the case where $\lambda > \mu$ (i.e., $\mathcal{R}_0 > 1$), there is a stable fixed point of the drift equation (1.2) at $x = 1 - \mu/\lambda$ (and an unstable fixed point at x = 0). If there are a large number of infective individuals at time 0, then with high probability $X_N(t)/N$ heads rapidly towards the stable fixed point, then spends most of its time in the neighbourhood of that fixed point, making excursions into the rest of the state space until eventually one of these excursions reaches the absorbing state 0. Precise results are known about the distribution of the time to

extinction, which is exponential in N, and about the quasi-stationary distribution, which is centred around the stable fixed point of (1.2). See, for instance: Barbour (1976), Kryscio and Lefèvre (1989), Nåsell (1996), Andersson and Djehiche (1998) and the book of Nåsell (2011).

If $\lambda \leq \mu$, then the differential equation (1.2) has a single stable fixed point at x = 0, and all its solutions converge to zero as $t \to \infty$. For the corresponding Markov chain, it is also known that the epidemic dies out rapidly with high probability whenever λ and μ are fixed constants with $\lambda \leq \mu$.

Doering, Sargsyan and Sander (2005) give an asymptotic formula for the mean extinction time, in the case where $\lambda < \mu$ are fixed constants and the initial state $X_N(0)$ is of order N:

$$\mathbb{E} T_N = \frac{1}{\mu - \lambda} (\log N + O(1)).$$
(1.3)

In the case where $\lambda = \mu$, they obtain:

$$\mathbb{E} T_N = \frac{1}{\lambda} \left[\left(\frac{\pi}{2} \right)^{3/2} \sqrt{N} + \log N \right] + O(1).$$

Doering, Sargsyan and Sander (2005) also study the mean time to extinction starting from a state with a single infective individual, and Kessler (2008) extends these results to cover the whole of the "transition region", where $\mu - \lambda$ is of order $N^{-1/2}$.

A formula for the asymptotic distribution of the time T_N to extinction, in the case where $\lambda < \mu$ and $X_N(0)/N$ tends to a constant, is presented by Kryscio and Lefèvre (1989) with a heuristic argument, and then reproduced by Andersson and Djehiche (1998). However, the formula is erroneous. It was noted by Doering, Sargsyan and Sander (2005) that the formula given by Kryscio and Lefèvre (1989) and Andersson and Djehiche (1998) is inconsistent with their result (1.3), and with their numerical results. As far as we are aware, no correct explicit formula for the asymptotic distribution of the time T_N to extinction when $\lambda < \mu$ has appeared in the literature, even in the case where λ and μ are fixed constants. In his book, Nåsell (2011) identifies two distinct regimes: one "critical regime", where $\mu - \lambda$ is of order at most $N^{-1/2}$, and another (subcritical) where $\mu - \lambda$ is constant or tends to zero more slowly than $N^{-1/2}$. For both regimes, Nåsell (2011) poses as an open problem the determination of the mean extinction time $\mathbb{E} T_N$. Our results have some similarity with Theorem 2(ii) of Sagitov and Shaimerdenova (2013), who study the distribution of the extinction time for a different version of the logistic model in a completely different limit.

Barbour, Hamza, Kaspi and Klebaner (2015) study a very general class of population models, which includes this one. The distribution of the extinction time T_N , in the case where λ and μ are fixed with $\lambda < \mu$, can, with some effort, be derived from their Theorem 1.2. Our results cover the case of fixed λ and μ , as well as the near-critical case (which Barbour et al (2015) do not cover), and our proof for this model is significantly simpler than the general argument given by Barbour et al (2015).

Here we obtain the asymptotic distribution of T_N throughout the subcritical regime, for general initial conditions. Our main result is as follows. We recall that a random variable W has the *standard Gumbel distribution* if $\mathbb{P}(W \leq w) = e^{-e^{-w}}$ for all $w \in \mathbb{R}$. The mean of W is equal to Euler's constant $\gamma \approx 0.5772$.

Theorem 1.1. Suppose that $\mu = \mu(N)$ and $\lambda = \lambda(N)$ are bounded away from both 0 and infinity. Suppose also that $(\mu - \lambda)N^{1/2} \to \infty$ as $N \to \infty$, that $X_N(0)$ is non-random and that $X_N(0)(\mu - \lambda) \to \infty$ as $N \to \infty$. Then, as $N \to \infty$,

$$(\mu - \lambda)T_N - \left(\log N + 2\log(\mu - \lambda) - \log\left(1 + \frac{(\mu - \lambda)N}{\lambda X_N(0)}\right) - \log \mu - \log \lambda\right) \to W,$$
(1.4)

in distribution, where W is a standard Gumbel variable. Hence, as $N \to \infty$,

$$\mathbb{E}T_N = \frac{\log N + 2\log(\mu - \lambda) - \log\left(1 + \frac{(\mu - \lambda)N}{\lambda X_N(0)}\right) - \log\mu - \log\lambda + \gamma + o(1)}{\mu - \lambda}$$

Observe that

$$\log N + 2\log(\mu - \lambda) - \log\left(1 + \frac{(\mu - \lambda)N}{\lambda X_N(0)}\right) = -\log\left(\frac{1}{N(\mu - \lambda)^2} + \frac{1}{\lambda X_N(0)(\mu - \lambda)}\right),$$

which tends to infinity under the hypotheses of the theorem. The remaining terms, $\log \mu$ and $\log \lambda$, in the expression in (1.4) are of constant order, and so Theorem 1.1 implies that, for any fixed $\varepsilon > 0$, the probability of extinction before time

$$(1-\varepsilon)\frac{\log N + 2\log(\mu-\lambda) - \log\left(1 + \frac{(\mu-\lambda)N}{\lambda X_N(0)}\right)}{\mu-\lambda}$$

tends to zero, while the probability of extinction by time

$$(1+\varepsilon)\frac{\log N + 2\log(\mu-\lambda) - \log\left(1 + \frac{(\mu-\lambda)N}{\lambda X_N(0)}\right)}{\mu-\lambda}$$

tends to 1. This is an instance of the cut-off phenomenon (see for instance: Diaconis 1996; Levin, Peres and Wilmer 2009), where, for a Markov chain $(X_N(t))$, the total variation distance between the distribution of $X_N(t)$ and the stationary distribution moves rapidly from 1 to 0 over a time interval much smaller than the time to stationarity. In our instance, the support of the stationary distribution is $\{0\}$, so the total variation distance at time t is the probability that the epidemic is not yet extinct. This probability goes from near 1 to near 0 over a time interval of length of order $1/(\mu - \lambda)$, whereas the expected extinction time, from a large enough initial state, is of order $\log[N(\mu - \lambda)^2]/(\mu - \lambda)$. Thus the cut-off phenomenon becomes less

pronounced as $N(\mu - \lambda)^2$ tends more slowly to infinity, i.e., as we approach the critical regime. See Figures 3 and 4.

Another point to note is how the expected extinction time changes as $\mu - \lambda$ decreases, showing the transition between the subcritical and critical regimes. When $\mu - \lambda$ is of constant order, the expected extinction time is $(1 + o(1)) \log N/(\mu - \lambda)$; as $(\mu - \lambda)N^{1/2}$ tends to infinity more and more slowly, the expected extinction time grows almost as large as $N^{1/2}$.

We next give versions of (1.4) valid when $X_N(0)/N$ lies in certain ranges, assuming always that $(\mu(N) - \lambda(N))N^{1/2} \to \infty$ and $X_N(0)(\mu(N) - \lambda(N)) \to \infty$. One important special case is when $X_N(0)/N \to \alpha$, with $\alpha \in (0, 1]$, when Theorem 1.1 gives that, for a standard Gumbel random variable W,

$$(\mu - \lambda)T_N - \left(\log N + 2\log(\mu - \lambda) + \log \alpha - \log(\lambda \alpha + \mu - \lambda) - \log \mu\right) \to W,$$
(1.5)

in distribution, as $N \to \infty$.

In general, (1.4) is the most that can be said if $\mu - \lambda$ is of the same order as $X_N(0)/N$ (e.g., if both are constants). On either side of this regime, the formula in (1.4) can be simplified.

For instance, if $X_N(0)/N(\mu - \lambda) \to \infty$ (so $X_N(0)/N$ is asymptotically of larger order than $\mu - \lambda$), then

$$(\mu - \lambda)T_N - \left(\log N + 2\log(\mu - \lambda) - \log \mu - \log \lambda\right) \to W, \tag{1.6}$$

in distribution, as $N \to \infty$, where W has the standard Gumbel distribution. In (1.6), necessarily $\mu - \lambda \to 0$, so either of the terms $\log \mu$ and $\log \lambda$ could be replaced by the other.

Note that, for any X,

$$\log N + \log(\mu - \lambda) - \log\left(1 + \frac{(\mu - \lambda)N}{\lambda X}\right) - \log \lambda = \log X - \log\left(\frac{\lambda X}{(\mu - \lambda)N} + 1\right),$$

and so an equivalent form of (1.4) is

$$(\mu - \lambda)T_N - \left(\log X_N(0) + \log(\mu - \lambda) - \log\left(1 + \frac{\lambda X_N(0)}{(\mu - \lambda)N}\right) - \log\mu\right) \to W.$$
(1.7)

It follows that, if $X_N(0)/N(\mu - \lambda) \to 0$ (so $X_N(0)/N$ is asymptotically of smaller order than $\mu - \lambda$) and $X_N(0)(\mu - \lambda) \to \infty$, then

$$(\mu - \lambda)T_N - \left(\log X_N(0) + \log(\mu - \lambda) - \log\mu\right) \to W, \tag{1.8}$$

in distribution, as $N \to \infty$.

We observe that the asymptotic formula for the distribution of T_N in (1.8) is independent of N, while that in (1.6) is independent of $X_N(0)$. An explanation for the first of these phenomena is that, in this regime, the quadratic terms in the drift are smaller than the linear ones, and the logistic process behaves essentially identically to a linear birth-and-death chain with birth rate λ and death rate μ . In Section 2, we show that the logistic correction to the birth rate (i.e., the term $-\lambda X(t)^2/N$), does not affect the asymptotics of the remaining time to extinction. If $\mu - \lambda \to 0$, then, for large enough $X_N(0)$ (such that $X_N(0) \gg (\mu - \lambda)N$), we are in the regime covered by (1.6), where the asymptotic distribution of the time to extinction does not depend on the starting state. To explain this, we give an informal description of the typical course of the epidemic in the case where $\mu - \lambda \to 0$, and we start in some large state, say with $X_N(0) = \lceil \alpha N \rceil$ and $0 < \alpha \le 1$.

For such a regime, in the initial phase of the epidemic, the number $X_N(t)$ of infectives very rapidly drops – in time $o(1/(\mu - \lambda))$ – until it reaches states of the same order as $(\mu - \lambda)N$. The majority of the duration of the epidemic – asymptotically $\log(N(\mu - \lambda)^2))/(\mu - \lambda)$ – is spent getting from there to states of the same order as $(\mu - \lambda)^{-1}$; the time taken to cross this gap is very well concentrated around the value derived from the approximating differential equation. Most of the variability of the time to extinction comes from the end stages of the epidemic, when $X_N(t)$ is of order below $(\mu - \lambda)^{-1}$; for this phase, the differential equation is no longer an adequate guide to the behaviour of the stochastic process, and instead $X_N(t)$ is well approximated by a linear birth-and-death chain. The expected remaining time to extinction, from a state of order $(\mu - \lambda)^{-1}$, is on the order of $1/(\mu - \lambda)$, and the standard deviation is of the same order. (The choice of where to draw the line between the "intermediate" and "final" phases is somewhat arbitrary, and a substantial part of the duration of the epidemic could be assigned to the final phase. Our results essentially show that the approximation by a linear birth-and-death chain is good starting from any state below the order of $(\mu - \lambda)N$.)

Note that the description above relies on having $(\mu - \lambda)N \gg (\mu - \lambda)^{-1}$. If $\mu - \lambda = o(N^{-1/2})$, then the situation is completely different: the time to extinction is essentially distributed as in the case $\mu = \lambda$.

The assumption in Theorem 1.1 that $X_N(0)(\mu - \lambda) \to \infty$ is necessary for the conclusion to hold; otherwise the variability in the extinction time is not as large as is given by the Gumbel distribution. We give more details for the case where this assumption is not satisfied at the end of Section 2.

Sections 2-5 are devoted to the proof of Theorem 1.1. We track the epidemic process through three phases, roughly corresponding to the three phases mentioned in the informal description above. For some regimes, not all the phases are necessary, and we tackle them in reverse order, starting with the final phase of the epidemic. Our intermediate results are stated in terms of a function $\omega(N)$, which tends to infinity suitably slowly; for convenience, we specify throughout that

$$\omega(N) = (\mu(N) - \lambda(N))^{1/4} N^{1/8}.$$
(1.9)

We treat the final phase in Section 2. Here, we start from a state below $N^{1/2}\omega(N)$, and show that, from this point on, $X_N(t)$ is well approximated by a linear birth-and-death chain with the same parameters. Since the distribution of the extinction time for a linear birth-and-death chain is known

explicitly, this enables us to analyse very precisely the behaviour of the logistic chain. This phase covers the stage of the epidemic giving rise to the randomness in the time to extinction. An alternative way to view the final phase of the epidemic is to approximate it by a branching process, where each initially infected individual sparks a brief small epidemic within the population, and these various small epidemics do not interact significantly. The time to extinction is then the maximum of the durations of these small epidemics, and this explains the appearance in our formulae of the standard Gumbel distribution, which typically arises as the maximum of a number of independent samples from a given distribution. Note again that the exact break point between the final and intermediate phases is somewhat arbitrary. We do need to start the final phase in a state well below $N(\mu - \lambda)$, so that the logistic effects can be ignored, and it is helpful to us to start slightly smaller yet; on the other hand we do need the initial state larger than $(\mu - \lambda)^{-1}$ for the formula involving the Gumbel distribution to apply.

The intermediate phase is covered in Section 3. Here, effectively, we prove Theorem 1.1 under the additional assumption that $X_N(0) \leq (\mu - \lambda) N \omega(N)$. We show that the scaled process $(X_N(t)/N)$ stays close to the solution of the differential equation (1.2) for a (deterministic) period of time until there are about $N^{1/2}\omega(N)$ infective individuals (from which point the analysis for the final phase can be invoked).

In Section 4, we provide a fairly crude upper bound on the duration of the initial phase of the epidemic, starting from any state and reaching a state of order about $N(\mu - \lambda)\omega(N)$. The length of this phase is negligible compared to the overall duration of the epidemic, or even the fluctuations in the overall duration, so greater precision is not necessary.

In Section 5, we combine our results to prove Theorem 1.1.

Our results have some bearing on the critical regime, where $|\mu - \lambda| = O(N^{-1/2})$. In particular, the methods of Section 4 can be used to show that the expected time for the epidemic starting from an arbitrary state to reach a state of size about $N^{1/2}$ is of order at most $N^{1/2}$. Dolgoarshinnykh and Lalley (2006) show that, in this regime, the scaled logistic process starting from a state of order $N^{1/2}$ converges in law to an "attenuated" Feller diffusion. One consequence is that the time to extinction from states of size about $N^{1/2}$ is of order $N^{1/2}$ (and is not well-concentrated). We discuss the critical regime briefly in the short Section 6, but make no attempt to provide precise results.

In Section 7, we consider the total number C_N of new cases (i.e., infection events) over the duration of the epidemic. Theorem 7.1 provides a precise estimate, valid throughout the subcritical regime, of the expectation of C_N of new cases, and states that C_N is well-concentrated around its mean, via an estimate of the variance of C_N . One consequence of this result is that, if $X_N(0) \gg (\mu - \lambda)N$, then most of the new cases occur during the short first phase of the epidemic, i.e., before $X_N(t)$ has dropped to around $(\mu - \lambda)N$. The total number of new cases in the SIS logistic epidemic is studied in detail by Kessler (2008), for the full range of parameter values. In the subcritical regime, Kessler (2008) gives an asymptotic formula agreeing with ours for the expectation of C_N when $X_N(0)$ is of order N, and discusses other cases, including ones where $\mu - \lambda = \delta N^{-1/2}$ and δ is large. He also estimates the asymptotic distribution of C_N in subcritical, critical and supercritical regimes, but only in the case $X_N(0) = 1$. Our results show that, provided $X_N(0)(\mu - \lambda)$ tends to infinity, C_N is well-concentrated around its expectation.

In Section 8, we present numerical methods to treat fixed values of N. All our theoretical results concern limiting behaviour as the population size N tends to infinity, and in many places it is important that terms such as $\log N$ (or indeed functions potentially growing more slowly) are much larger than constants. It is not apparent that our results have any bearing on "human-size" populations: we address this issue by performing numerical calculations for a range of values of N and appropriate values of λ and μ . As we explain in Section 8, it is more efficient to estimate the distribution of (for instance) the extinction time by numerical integration, as opposed to using Monte Carlo methods. We see good agreement between asymptotic results and simulation for temporal behaviour even for N = 10 and N = 100, when typically one does not see this until N = 1000 for epidemic models (see, e.g., Demiris and O'Neill 2006).

We conclude, in Section 9, by expanding on our observations about the behaviour of a barely subcritical epidemic. We discuss in particular those features that we expect to carry over to more complex models, or to realworld epidemics, especially the resemblance to a random walk for a period before extinction, and the weakening of the cut-off phenomenon as we approach criticality. In Appendix A, we present some data from real epidemics, and make some very tentative connections between our hypotheses and the observations.

2. FINAL PHASE: APPROXIMATION BY LINEAR BIRTH-AND-DEATH CHAINS

Suppose that $\mu = \mu(N)$ and $\lambda = \lambda(N)$ are bounded away from both 0 and infinity, and that $X_N(0)$ is non-random with $X_N(0)(\mu - \lambda) \to \infty$ and $X_N(0) \le \omega(N)N^{1/2}$, where $\omega(N) = (\mu - \lambda)^{1/4}N^{1/8}$, as in (1.9). We will show that, with such an initial state, $X_N(t)$ is well approximated until extinction by a pair of linear birth-and-death chains. The assumption that $X_N(0)(\mu - \lambda) \to \infty$ ensures that the randomness in the extinction time of the approximating linear birth-and-death chains has a Gumbel distribution.

We will prove the following lemma.

Lemma 2.1. Suppose that $(\mu - \lambda)N^{1/2} \to \infty$. Set $\omega(N) = ((\mu - \lambda)N^{1/2})^{1/4}$, and assume that $X_N(0)(\mu - \lambda) \to \infty$ and $X_N(0) \le 2N^{1/2}\omega(N)$. Then

$$(\mu - \lambda)T_N - (\log X_N(0) + \log(\mu - \lambda) - \log \mu) \rightarrow W_N$$

in distribution, as $N \to \infty$, where W has the standard Gumbel distribution.

Note that this result is the same as the special case of Theorem 1.1 covered by (1.8), under the more restrictive hypothesis that $X_N(0) \leq 2N^{1/2}\omega(N) =$ $2N^{5/8}(\mu - \lambda)^{1/4}$ instead of $X_N(0) = o((\mu - \lambda)N)$. Later results will supply the conclusion of Theorem 1.1 with no upper bounds on the starting state.

For a birth-and-death chain (B(t)) on \mathbb{Z}_+ with a unique absorbing state at 0, let T^B be the extinction time, that is $T^B = \inf\{t : B(t) = 0\}$ is the time when (B(t)) gets absorbed at 0. Thus the extinction time of $(X_N(t))$ is $T_N = T^{X_N}$.

Let $(Y(t))_{t\geq 0}$ be a linear birth-and-death chain with birth rate λ and death rate μ , so its transition rates from state $Y \in \mathbb{Z}_+$ are given by

$$Y \rightarrow Y+1$$
 at rate λY ,
 $Y \rightarrow Y-1$ at rate μY .

Assume that Y(0) is non-random. It is known – see for instance (2.4.23) in the book of Renshaw (2011) – that, for $t \ge 0$ and $\mu \ne \lambda$,

$$\mathbb{P}(T^Y \le t) = \mathbb{P}(Y(t) = 0) = \left(\frac{\mu - \mu e^{-(\mu - \lambda)t}}{\mu - \lambda e^{-(\mu - \lambda)t}}\right)^{Y(0)} = \left(1 - \frac{(\mu - \lambda)e^{-(\mu - \lambda)t}}{\mu - \lambda e^{-(\mu - \lambda)t}}\right)^{Y(0)}.$$
(2.1)

We will write $(Y_N(t))$ to denote a linear birth-and-death chain with birth rate $\lambda(N)$ and death rate $\mu(N)$. Suppose that $Y_N(0) = X_N(0)$, where $X_N(0)(\mu - \lambda) \to \infty$ as $N \to \infty$. For each fixed $w \in \mathbb{R}$, we set

$$t_w = t_w(\mu, \lambda, X_N(0)) = \frac{\log X_N(0) + \log(\mu - \lambda) - \log \mu + w}{\mu - \lambda}$$

and note that $t_w > 0$ for sufficiently large N. Restricting to those N for which t_w is indeed positive, we have that $e^{-(\mu-\lambda)t_w} = \mu e^{-w}/(\mu-\lambda)X_N(0)$. Hence, from (2.1), the extinction time T^{Y_N} satisfies

$$\mathbb{P}(T^{Y_N} \le t_w) = \left(1 - \frac{\mu e^{-w} / X_N(0)}{\mu - \lambda \mu e^{-w} / (\mu - \lambda) X_N(0)}\right)^{X_N(0)} \\
= \left(1 - \frac{e^{-w}}{X_N(0) - \lambda e^{-w} / (\mu - \lambda)}\right)^{X_N(0)} \to e^{-e^{-w}},$$

as $N \to \infty$, since $X_N(0)(\mu - \lambda) \to \infty$. This can be written as

$$(\mu - \lambda)T^{Y_N} - \left(\log X_N(0) + \log(\mu - \lambda) - \log\mu\right) \to W, \qquad (2.2)$$

in distribution, as $N \to \infty$, where W has the standard Gumbel distribution.

The plan of the proof of Lemma 2.1 is to sandwich the logistic process $(X_N(t))$ between two linear birth-and-death chains, the upper of which is $(Y_N(t))$. The upper bound on $X_N(0)$ ensures that $X_N(0)/N(\mu - \lambda) \to 0$, so "logistic effects" in the drift become negligible and so the linear birthand-death chains approximate $(X_N(t))$ well. Our argument is a little crude, in that the birth rate of the lower of the two birth-and-death chains is significantly below that of the logistic process for most of the phase, and this is why we need the stronger hypothesis $X_N(0) \leq 2N^{1/2}\omega(N)$, rather than just $X_N(0)/N(\mu - \lambda) \to 0$, and the precise form of ω matters here.

We will use the following result about birth-and-death chains with a higher rate of deaths than births: we omit the routine proof.

Lemma 2.2. Let (B(t)) be a birth-and-death chain on \mathbb{Z}_+ , with B(0) nonrandom. Suppose that, from any state, the probability that the next transition is upwards is at most p, and the probability that the next transition is downwards is at least q > p.

For any state B > B(0), the probability that the chain (B(t)) reaches B before it reaches 0 is at most

$$\frac{(q/p)^{B(0)} - 1}{(q/p)^B - 1} \le \left(\frac{p}{q}\right)^{B - B(0)} \le \exp\left(-(1 - p/q)(B - B(0))\right).$$

Proof of Lemma 2.1. We couple three Markov chains: one is the logistic process $(X_N(t))$, another is the linear birth-and-death chain $(Y_N(t))$ with the same parameters $(\lambda(N), \mu(N))$ as $(X_N(t))$, and the third is a linear birth-and-death chain $(Z_N(t))$ with parameters $(\lambda'(N), \mu(N))$ where $\lambda'(N) = \lambda(N)(1 - 2X_N(0)/N)$. We let $Y_N(0) = X_N(0) = Z_N(0)$. Let τ_N be the first time that either $X_N(t) = 0$ or $X_N(t) = 2X_N(0)$. The birth rate of $(X_N(t))$ when in state X is $\lambda X(1-X/N)$, which for $t \leq \tau_N$ is sandwiched between the birth rates of the two linear birth-and-death chains in the same state. For each N, we may thus construct a coupling such that $Z_N(t) \leq X_N(t) \leq Y_N(t)$ for all $t \leq \tau_N$. The rule is that, if any two chains are in the same state, then they make jumps together as far as possible; otherwise two chains in different states make jumps independently according to their given transition rates, and so they a.s. do not jump simultaneously (so they do not cross). With this coupling, on the event that $X_N(\tau_N) = 0$ (i.e., $(X_N(t))$ reaches 0 before it reaches the upper boundary $2X_N(0)$), $T^{Z_N} \leq T_N \leq T^{Y_N}$.

For each fixed w and N, we choose v = v(w, N) so that $t_w(\mu, \lambda, X_N(0)) = t_v(\mu, \lambda', X_N(0))$, i.e.,

$$\frac{\log X_N(0) + \log(\mu - \lambda) - \log \mu + w}{\mu - \lambda} = \frac{\log X_N(0) + \log(\mu - \lambda') - \log \mu + v}{\mu - \lambda'}$$

This translates to

$$v(w,N) - w = \frac{\lambda - \lambda'}{\mu - \lambda} \left(\log X_N(0) + \log(\mu - \lambda) - \log\mu + w \right) - \log\left(1 + \frac{\lambda - \lambda'}{\mu - \lambda}\right)$$
(2.3)

We observe that

$$\frac{\lambda - \lambda'}{\mu - \lambda} = \frac{2X_N(0)\lambda}{N(\mu - \lambda)} \le \frac{4\lambda\omega(N)}{N^{1/2}(\mu - \lambda)} = \frac{4\lambda}{\omega(N)^3}.$$
 (2.4)

Also we have, for N sufficiently large,

 $\log(X_N(0)(\mu-\lambda)) \le (X_N(0)(\mu-\lambda))^{1/2} \le (2N^{1/2}\omega(N)N^{-1/2}\omega(N)^4)^{1/2} = 2\omega(N)^{5/2}.$ Therefore, for each fixed w, we have both

$$\frac{\lambda - \lambda'}{\mu - \lambda} \left(\log X_N(0) + \log(\mu - \lambda) - \log\mu + w \right) \le \frac{2\lambda}{\omega(N)^3} \left(2\omega(N)^{5/2} + O(1) \right) = o(1)$$
and by (2.4)

and, by (2.4),

$$\log\left(1 + \frac{\lambda - \lambda'}{\mu - \lambda}\right) = o(1),$$

and so, from (2.3), |v(w, N) - w| = o(1). In other words, $v(w, N) \to w$ as $N \to \infty$, for each fixed w.

Thus

$$|\mathbb{P}(T^{Z_N} \le t_w(\mu, \lambda, X_N(0))) - e^{-e^{-w}}|$$

= $|\mathbb{P}(T^{Z_N} \le t_v(\mu, \lambda', X_N(0))) - e^{-e^{-w}}|$
 $\le |\mathbb{P}(T^{Z_N} \le t_v(\mu, \lambda', X_N(0))) - e^{-e^{-v(w,N)}}| + |e^{-e^{-v(w,N)}} - e^{-e^{-w}}| \to 0$

as $N \to \infty$. Here we used (2.2) applied to $(Z_N(t))$ (note that $X_N(0)(\mu - \lambda') > X_N(0)(\mu - \lambda)$, which tends to infinity).

Also by (2.2), as $N \to \infty$,

$$\mathbb{P}(T^{Y_N} \le t_w(\mu, \lambda, X_N(0))) \to e^{-e^{-w}}$$

As $(X_N(t))$ is sandwiched between $(Y_N(t))$ and $(Z_N(t))$ for all times t, on the event $A := \{X_N(\tau_N) = 0\}$, we see that

$$\mathbb{P}(\{T^{Y_N} \le t\} \cap A) \le \mathbb{P}(\{T_N \le t\} \cap A) \le \mathbb{P}(\{T^{Z_N} \le t\} \cap A),$$

and in particular this holds with $t = t_w(\mu, \lambda, X_N(0))$. By Lemma 2.2 with $p = \lambda/(\lambda + \mu) = 1 - q$ and $B(0) = X_N(0)$, $\mathbb{P}(\overline{A}) \leq e^{-(\mu - \lambda)X_N(0)/\mu} = o(1)$, since $(\mu - \lambda)X_N(0) \to \infty$. Hence, as $N \to \infty$,

$$\mathbb{P}(T_N \le t_w(\mu, \lambda, X_N(0))) \to e^{-e^{-w}}.$$

Equivalently, as $N \to \infty$,

$$(\mu - \lambda)T_N - (\log X_N(0) + \log(\mu - \lambda) - \log \mu) \to W,$$

in distribution, where W has the standard Gumbel distribution, as claimed. $\hfill \Box$

In the case where $(\mu - \lambda)X_N(0)$ does not tend to infinity, we can use a similar argument to show that the distribution of the extinction time of $(X_N(t))$ is asymptotically the same as that of the linear birth-and-death chain with the same parameters. We give a brief sketch of the argument in the case where $(\mu - \lambda)X_N(0) \rightarrow 0$.

For a linear birth-and-death chain $(Y_N(t))$ with parameters (λ, μ) and $Y_N(0) = X_N(0) = o(1/(\mu - \lambda))$, Lemma 2.2 shows that the probability of the event A that $(Y_N(t))$ never reaches $N^{1/2}$ before extinction is 1 - o(1). Accordingly, we consider also a linear birth-and-death chain $(Z_N(t))$ with

birth rate equal to $\lambda' = \lambda(1 - N^{-1/2})$. As in the proof of Lemma 2.1, we may couple our three processes so that $Z_N(t) \leq X_N(t) \leq Y_N(t)$ for all t, on the event A.

It can be seen from (2.1) that, if $(\mu - \lambda)X_N(0) \to 0$ and $X_N(0) \to \infty$, then for any $v \in (0, \infty)$,

$$\mathbb{P}\left(T^{Y_N} \le vX_N(0)/\mu\right) \to e^{-1/\nu} \quad \text{as } N \to \infty.$$
(2.5)

Note that (2.5) does not depend on $\mu - \lambda$, provided that $(\mu - \lambda)X_N(0) \to 0$. Note also that $(\lambda - \lambda')X_N(0) = \lambda N^{-1/2}X_N(0) = o(N^{-1/2}/(\mu - \lambda)) = o(1)$, and so $(\mu - \lambda')X_N(0) \to 0$ whenever $(\mu - \lambda)X_N(0) \to 0$. Therefore (2.5) holds with T^{Y_N} replaced by T^{Z_N} , and hence also $\mathbb{P}(T_N \leq vX_N(0)/\mu) \to e^{-1/v}$.

We can also consider the case where the epidemic starts with a single infective: if $\mu - \lambda \to 0$ and $X_N(0) = 1$, then, for any $u \in (0, \infty)$,

$$\mathbb{P}\left(T^{X_N} \le u/\mu\right) \to \frac{u}{1+u} \quad \text{as } N \to \infty.$$

3. INTERMEDIATE PHASE: DIFFERENTIAL EQUATION APPROXIMATION

For any $\alpha \in [0, 1]$, the differential equation (1.2) subject to initial condition $x(0) = \alpha$ has an explicit solution

$$x(t) = \frac{\alpha(\mu - \lambda)e^{-(\mu - \lambda)t}}{\mu - \lambda + \alpha\lambda(1 - e^{-(\mu - \lambda)t})}, \qquad t \ge 0.$$
(3.1)

For fixed $0 < \alpha \leq 1$, the inverse of the function x(t) is given by

$$t_{\alpha}(x) = \frac{s(x) - s(\alpha)}{\mu - \lambda}; \quad \text{where } s(x) = \log\left(1 + \frac{\lambda}{\mu - \lambda}x\right) - \log x, \quad (3.2)$$

for $0 < x \leq \alpha$.

We also note for future reference that $x(t) \leq x(0)e^{-(\mu-\lambda)t}$, and therefore, for any $t \geq 0$,

$$\int_0^t x(s) dt \le \frac{x(0)}{\mu - \lambda}.$$
(3.3)

As in (1.9), we set $\omega(N) = (\mu - \lambda)^{1/4} N^{1/8}$, and suppose that $N^{1/2}\omega(N) \leq X_N(0) \leq (\mu - \lambda)N\omega(N)$. Let $X^* = X^*(N) = N^{1/2}\omega(N)$. We will show that $X_N(t)/N$ is well approximated by the solution x(t) to the differential equation (1.2) with $x(0) = X_N(0)/N$, at least until the time $t^* = t_{X_N(0)/N}(X^*/N)$ when $x(t^*) = X^*/N$. It will then follow that $X_N(t^*)$ is close to X^* with probability 1 - o(1) as $N \to \infty$. The total time to extinction will then be obtained by adding t^* to the time to extinction from a state very near to X^* , which is covered in Lemma 2.1.

To be precise, we will prove the following result.

Lemma 3.1. Suppose $(\mu - \lambda)N^{1/2} \to \infty$ as $N \to \infty$. Set $\omega(N) = (\mu - \lambda)^{1/4}N^{1/8}$, and $X^* = X^*(N) = N^{1/2}\omega(N)$. Suppose $X^* \leq X_N(0) \leq N^{1/2}\omega(N)$.

$$\omega(N)(\mu - \lambda)N$$
. Then

$$\mathbb{P}\left(|X_N(t^*) - X^*| > \omega(N)^{-1/3} X^*\right) = o(1),$$

where

16

$$t^* = t_{X_N(0)/N}(X^*/N) = \frac{s(X^*/N) - s(X_N(0)/N)}{\mu - \lambda}.$$

Moreover, we have

$$t^* = \frac{1}{\mu - \lambda} \left(\log X_N(0) - \log(X^*) - \log\left(1 + \frac{\lambda}{\mu - \lambda} \frac{X_N(0)}{N}\right) + o(1) \right).$$
(3.4)

The final assertion in the statement follows immediately from the expression in (3.2) for s(x), since, with the assumptions given, $\log\left(1 + \frac{\lambda}{\mu - \lambda} \frac{X^*}{N}\right) = o(1)$.

To prove Lemma 3.1, we will use standard martingale techniques, taking advantage of the special error-correcting nature of the drift in process $X_N(t)$, thanks to which errors in the approximation do not accumulate much over time. Let x(t) be as in (3.1), with $\alpha = x(0) = X_N(0)/N$. Now set

$$T = \inf\left\{t \ge 0: |N^{-1}X_N(t) - x(t)| > 2\sqrt{X_N(0)(\omega(N))^{1/4}(\lambda + \mu)/N^2(\mu - \lambda)}\right\}.$$

Note that it will suffice to show that $\mathbb{P}(T \leq t^*) = o(1)$. This is because, if $T > t^*$, then

$$\begin{aligned} |X_N(t^*) - X^*| &\leq N \sup_{t \leq t^*} \left| \frac{X_N(t)}{N} - x(t) \right| \leq 2\sqrt{\frac{X_N(0)\omega^{1/4}(\lambda + \mu)}{\mu - \lambda}} \\ &\leq \sqrt{\omega^{5/4}N(\mu + \lambda)} = (\mu + \lambda)^{1/2}N^{1/2}\omega^{5/8} \\ &= o(X^*\omega^{-1/3}), \end{aligned}$$
(3.5)

since we have assumed that $X_N(0) \leq \omega(N)(\mu - \lambda)N$ and since $X^* = N^{1/2}\omega(N)$.

We write, as is standard,

$$x(t) = x(0) + \int_0^t f(x(s)) \, ds$$

where $f(x) = \lambda x(1-x) - \mu x = -(\mu - \lambda)x - \lambda x^2$. Also by standard theory,

$$\frac{X_N(t)}{N} = \frac{X_N(0)}{N} - (\mu - \lambda) \int_0^t \frac{X_N(s)}{N} \, ds - \lambda \int_0^t \left(\frac{X_N(s)}{N}\right)^2 \, ds + M_N(t), \quad (3.6)$$

where $(M_N(t))_{t\geq 0}$ is a zero-mean martingale.

Setting
$$e_N(t) = X_N(t)/N - x(t)$$
, it follows that

$$e_N(t) = -\int_0^t e_N(s) \left[(\mu - \lambda) + \lambda \left(\frac{X_N(s)}{N} + x(s) \right) \right] ds + M_N(t).$$
 (3.7)

To bound $e_N(t)$, we use the following simple lemma. For future applications (e.g., in forthcoming work by Lopes and Luczak on the SIS logistic competition model), we state it in a slightly more general form than needed here.

Lemma 3.2. Fix a time τ_0 , and let $m : [0, \tau_0] \to \mathbb{R}$ and $r, v : [0, \tau_0] \to \mathbb{R}^+$ be càdlàg functions, where v is decreasing, and suppose that $u : [0, \tau_0] \to \mathbb{R}$ is a càdlàg function satisfying

$$u(t) = m(t) - v(t) \int_0^t r(s)u(s) \, ds,$$

for $0 \leq t \leq \tau_0$. Then

$$\sup_{t \le \tau_0} |u(t)| \le 2 \sup_{t \le \tau_0} |m(t)|.$$

Proof. Let $M = \sup_{t \leq \tau_0} |m(t)|$. Choose any $\tau \in [0, \tau_0]$, and suppose without loss of generality that $u(\tau) \geq 0$. If $u(t) \geq 0$ for all $t \leq \tau$, then we certainly have $u(\tau) \leq m(\tau) \leq M$. Otherwise, let $\sigma = \sup\{t \leq \tau : u(t) < 0\} > 0$, and observe that $\lim_{s \to \sigma^-} u(s) \leq 0$ and $u(s) \geq 0$ for $\sigma < s \leq \tau$, and so $\int_{\sigma}^{\tau} r(s)u(s) ds \geq 0$. We may therefore write

$$\begin{aligned} u(\tau) &= m(\tau) - v(\tau) \int_0^\sigma r(s)u(s) \, ds - v(\tau) \int_\sigma^\tau r(s)u(s) \, ds \\ &= m(\tau) - \lim_{t \to \sigma-} (m(t) - u(t)) \frac{v(\tau)}{v(t)} - v(\tau) \int_\sigma^\tau r(s)u(s) \, ds \\ &\leq M + M \lim_{t \to \sigma-} \frac{v(\tau)}{v(t)} + \lim_{t \to \sigma-} u(t) \frac{v(\tau)}{v(t)} - v(\tau) \int_\sigma^\tau r(s)u(s) \, ds \\ &\leq M + M + 0 + 0 = 2M. \end{aligned}$$

Hence $|u(t)| \leq 2M$ for all $t \leq \tau_0$, as required.

We apply Lemma 3.2 with $\tau_0 = t^*$, $u(t) = e_N(t)$, $m(t) = M_N(t)$, v(t) = 1, and $r(s) = (\mu - \lambda) + \lambda(X_N(s)/N + x(s))$. The hypotheses of the lemma are satisfied since $\mu > \lambda$, and so we have

$$\sup_{t \le t^*} |e_N(t)| \le 2 \sup_{t \le t^*} |M_N(t)|.$$
(3.8)

Therefore,

$$\mathbb{P}(T \le t^*) \le \mathbb{P}\left(\sup_{t \le t^*} |M_N(t)| > \sqrt{X_N(0)(\omega(N))^{1/4}(\lambda+\mu)/N^2(\mu-\lambda)}\right).$$
(3.9)

To bound $|M_N(t)|$, we use a standard exponential martingale argument. Let $q_N^1(x) = \lambda N x(1-x)$ and $q_N^{-1}(x) = \mu N x$ denote the rates of transition of $N^{-1}X_N(s)$, by 1/N and -1/N, respectively. For $\theta \in \mathbb{R}$, we define $V_N^{\theta}(t)$ by

$$V_N^{\theta}(t) = \exp\left(\theta N^{-1}(X_N(t) - X_N(0)) - \int_0^t \sum_j q_N^j (N^{-1}X_N(s))(e^{\theta N^{-1}j} - 1) \, ds\right)$$

=
$$\exp\left(\theta M_N(t) - \int_0^t \sum_j q_N^j (N^{-1}X_N(s))(e^{\theta N^{-1}j} - 1 - \theta N^{-1}j) \, ds\right) (3.10)$$

The process $(V_N^{\theta}(t))$ is a mean 1 martingale. Using that $e^z - 1 - z = z^2 \int_0^1 e^{rz} (1-r) dr \leq \frac{1}{2} z^2 e^{|z|}$, we see that

$$V_N^{\theta}(t) \ge \exp\left(\theta M_N(t) - \frac{\theta^2}{2N^2} e^{|\theta|/N} \int_0^t \sum_j q_N^j(N^{-1}X_N(s)) \, ds\right).$$

Assume that $|\theta| \leq N \log 2$. Let $T_1 = \inf\{t \geq 0 : X_N(t) > 2Nx(t)\}$; then for $t \leq T_1$,

$$V_{N}^{\theta}(t) \geq \exp\left(\theta M_{N}(t) - \frac{2\theta^{2}}{N}(\lambda+\mu)\int_{0}^{t}x(s)\,ds\right)$$

$$\geq \exp\left(\theta M_{N}(t) - \frac{2\theta^{2}(\lambda+\mu)X_{N}(0)}{N^{2}(\mu-\lambda)}\right), \qquad (3.11)$$

by (3.3).

For $\delta \in \mathbb{R}$, let $T^+(\delta) = \inf\{t \ge 0 : M_N(t) > \delta\}$, and let $T^-(\delta) = \inf\{t \ge 0 : M_N(t) < -\delta\}$. On the event $\{T^+(\delta) \le T_1\}$,

$$V_N^{\theta}(T^+(\delta)) \ge \exp\left(\theta\delta - \frac{2\theta^2(\lambda+\mu)X_N(0)}{N^2(\mu-\lambda)}\right)$$

By optional stopping and the Markov inequality,

$$\mathbb{P}(T^+(\delta) \le T_1) \le \exp\Big(-\theta\delta + \frac{2\theta^2 X_N(0)(\lambda+\mu)}{N^2(\mu-\lambda)}\Big).$$

Choosing $\theta = \frac{1}{4} \delta N^2(\mu - \lambda) / X_N(0)(\lambda + \mu)$, we have $|\theta| \leq N \log 2$ for sufficiently large N, provided $\delta = o(X_N(0)/N(\mu - \lambda))$. We then obtain

$$\mathbb{P}(T^+(\delta) \le T_1) \le e^{-\delta^2 N^2(\mu-\lambda)/8X_N(0)(\lambda+\mu)},$$

and, similarly,

$$\mathbb{P}(T^{-}(\delta) \leq T_1) \leq e^{-\delta^2 N^2(\mu-\lambda)/8X_N(0)(\lambda+\mu)}.$$

It follows that

$$\mathbb{P}(\sup_{t \le t^* \land T_1} |M_N(t)| > \delta) \le 2e^{-\delta^2 N^2(\mu - \lambda)/8X_N(0)(\lambda + \mu)}$$

Take $\delta = \sqrt{X_N(0)\psi(\lambda+\mu)/N^2(\mu-\lambda)}$, for some $\psi \leq \sqrt{N}$; this choice guarantees that $\delta = o(X_N(0)/N(\mu-\lambda))$, since it is equivalent to $\psi =$

18

 $o(X_N(0)/(\mu-\lambda))$, and we have assumed that $X_N(0)/N^{1/2} \to \infty$. We thus obtain

$$\mathbb{P}\left(\sup_{t \le t^* \wedge T_1} |M_N(t)| > \sqrt{\frac{X_N(0)\psi(\lambda+\mu)}{N^2(\mu-\lambda)}}\right) \le 2e^{-\psi/8}.$$

If we choose $\psi = (\omega(n))^{1/4}$, then $\psi \leq N^{1/2}$ for N large enough, since, by (1.9), $\omega(N) = (\mu(N) - \lambda(N))^{1/4} N^{1/8} = O(N^{1/8})$. Let $\delta_0 = \sqrt{\frac{X_N(0)\omega^{1/4}(\lambda+\mu)}{N^2(\mu-\lambda)}}$, and let $T_0 = T^+(\delta_0) \wedge T^-(\delta_0)$. Then, using (3.9),

$$\mathbb{P}(T \le t^*) \le \mathbb{P}(T_0 \le t^*) \le \mathbb{P}(T_0 \le t^* \land T_1) + \mathbb{P}(T_1 \le t^* \land T_0) \le 2e^{-\omega(N)^{1/4}/8},$$

since we showed in (3.5) that $\delta_0 = o(X^*/N) = o(x(t))$ for all $t \leq t^*$, and so $\mathbb{P}(T_1 \leq t^* \wedge T_0) = 0$, provided N is sufficiently large.

This completes the proof of Lemma 3.1.

4. INITIAL PHASE: UPPER BOUNDS

In this section, we show that, if $X_N(0) > (\mu - \lambda)N\omega(N)$, then by the time $t_0 = 1/(\omega(N)\lambda(\mu - \lambda))$, $X_N(t)$ with high probability will have dropped down below $(\mu - \lambda)N\omega(N)$.

To this end, we give a lemma showing that $\mathbb{E}(X_N(t)/N)$ is always bounded above by the solution x(t) of the differential equation (1.2). This result has earlier been proved by Allen (2008, p94), and in a more general setting by Simon and Kiss (2013).

Lemma 4.1. Suppose that $X_N(0) \in \{0, 1, \dots, N\}$ is non-random, and let x(t) be the solution to (1.2) with initial condition $x(0) = X_N(0)/N$. Then, for all $t \geq 0$,

$$\mathbb{E} X_N(t) \le Nx(t).$$

Proof. (Sketch) It is easy to calculate that, for all $t \geq 0$,

$$\frac{\mathrm{d}}{\mathrm{d}t} \left(\mathbb{E} X_N(t) - Nx(t) \right) = \left(\mathbb{E} X_N(t) - Nx(t) \right) \left\{ -(\mu - \lambda) - \lambda \left(\frac{\mathbb{E} X_N(t)}{N} + x(t) \right) \right\} - \frac{\lambda}{N} \mathbb{E} \left(X_N(t) - \mathbb{E} X_N(t) \right)^2.$$

Using the integrating factor $\exp\left(t(\mu - \lambda) + \lambda \int_0^t (N^{-1} \mathbb{E} X_N(s) + x(s)) \, ds\right)$, and the fact that $\mathbb{E} X_N(0) - Nx(0) = 0$, it follows that $Y_N(t) = \mathbb{E} X_N(t) - V_N(t)$ Nx(t) satisfies

$$Y_N(t) = -\frac{\lambda}{N} \int_{s=0}^t \mathbb{E}(X_N(s) - \mathbb{E} X_N(s))^2 e^{-(t-s)(\mu-\lambda) - \lambda \int_s^t (N^{-1} \mathbb{E} X_N(u) + x(u)) \, du} \, ds,$$

and so is non-positive for all $t \ge 0$.

and so is non-positive for all $t \ge 0$.

The previous result implies the following lemma.

Lemma 4.2. Let $\omega(N)$ be any function tending to infinity, and set $t_0 = t_0(N) = \frac{1}{\omega(N)^{1/2}\lambda(\mu-\lambda)}$. Then, for any initial state $X_N(0)$,

$$\mathbb{P}\left(X_N(t_0) \ge N(\mu - \lambda)\omega(N)\right) = o(1).$$

Proof. We note that, for any value of x(0), and any $t \ge 0$,

$$x(t) = \frac{x(0)(\mu - \lambda)e^{-(\mu - \lambda)t}}{(\mu - \lambda) + x(0)\lambda(1 - e^{-(\mu - \lambda)t})} \le \frac{(\mu - \lambda)e^{-(\mu - \lambda)t}}{\lambda(1 - e^{-(\mu - \lambda)t})} \le \frac{1}{\lambda t}.$$

Here we used the inequality $e^{-u} \leq (1 - e^{-u})/u$, valid for all u > 0.

Therefore we have $x(t_0) \leq (\mu - \lambda)\omega(N)^{1/2}$, for any value of x(0). Hence, by Lemma 4.1, we have $\mathbb{E} X_N(t_0) \leq (\mu - \lambda)N\omega(N)^{1/2}$, for any initial state $X_N(0)$, and it follows that

$$\mathbb{P}\left(X_N(t_0) \ge (\mu - \lambda)N\omega(N)\right) \le \frac{1}{\omega(N)^{1/2}} = o(1).$$

5. Proof of Theorem 1.1

In this section, we assemble the lemmas from the preceding three sections into a proof of Theorem 1.1.

Given two copies (B(t)) and $(\tilde{B}(t))$ of a continuous-time birth-and-death chain, with $B(0) \leq \tilde{B}(0)$, we can couple them in a monotone way. If $B(t) = \tilde{B}(t)$, then they make the next jump (and all subsequent jumps) together. For as long as $B(t) \neq \tilde{B}(t)$, (B(t)) and $(\tilde{B}(t))$ evolve independently, so that a.s. they do not jump simultaneously. This ensures that a.s. the two copies of the chain never cross, so that $B(t) \leq \tilde{B}(t)$ a.s. for all t.

Proof. Recall from (1.9) that $\omega(N) = (N^{1/2}(\mu - \lambda))^{1/4}$. We distinguish three ranges for the starting state $X_N(0)$, assuming always that $(\mu - \lambda)X_N(0) \rightarrow \infty$:

- (a) $X_N(0) \le 2N^{1/2}\omega(N)$,
- (b) $2N^{1/2}\omega(N) < X_N(0) \le (\mu \lambda)N\omega(N),$
- (c) $X_N(0) > (\mu \lambda) N \omega(N)$.

It could be that $X_N(0)$ falls into different ranges for different values of N: we partition the set of natural numbers into three sets depending on which of (a), (b), (c) holds. It suffices to prove the result separately for whichever subsequence(s) are infinite, and so we may treat each of the three ranges in turn, working (tacitly) with an infinite sequence of values of N for which the inequalities defining the range hold.

(a) Suppose
$$(\mu - \lambda)X_N(0) \to \infty$$
 and $X_N(0) \le 2N^{1/2}\omega(N)$. By Lemma 2.1,
 $(\mu - \lambda)T_N - (\log X_N(0) + \log(\mu - \lambda) - \log \mu) \to W,$

in distribution, as $N \to \infty$, where W has the standard Gumbel distribution. This is (1.8), which is equivalent to (1.4) in this range. (b) Let $X^* = X^*(N) = N^{1/2}\omega(N)$. Suppose that $2N^{1/2}\omega(N) < X_N(0) \le (\mu - \lambda)N\omega(N)$. We run $(X_N(t))$ for a time $t^* = t_{X_N(0)/N}(X^*/N)$, as in the statement of Lemma 3.1. Let E be the event that $X^*(1 - \omega(N)^{-1/3}) \le X_N(t^*) \le X^*(1 + \omega(N)^{-1/3})$; by Lemma 3.1, $\mathbb{P}(\overline{E}) = o(1)$.

Let $(Y_N(t))$ be a copy of the logistic process with $Y_N(0) = X^*(1 - \omega(N)^{-1/3})$, and $(Z_N(t))$ be a copy with $Z_N(0) = X^*(1 + \omega(N)^{-1/3})$. On the event E, we couple these with $(X_N(t))$ from time t^* onwards in a monotone way, so that $Y_N(t) \leq X_N(t^*+t) \leq Z_N(t)$ for all $t \geq 0$. Then, on the event E, $T^{Y_N} + t^* \leq T_N \leq T^{Z_N} + t^*$. By Lemma 2.1, as $N \to \infty$, in distribution,

$$(\mu - \lambda)T^{Y_N} - \left(\log X_N(0) + \log(1 - \omega(N)^{-1/3}) + \log(\mu - \lambda) - \log\mu\right) \to W,$$

where W is a standard Gumbel random variable. Since $\log(1 - \omega(N)^{-1/3}) = o(1)$, we then have from the asymptotic formula (3.4) for t^* , as $N \to \infty$,

$$(\mu - \lambda)(T^{Y_N} + t^*) - \left(\log X_N(0) + \log(\mu - \lambda) - \log\left(1 + \frac{\lambda X_N(0)}{(\mu - \lambda)N}\right) - \log\mu\right) \to W,$$

in distribution, and the same holds when T^{Y_N} is replaced by T^{Z_N} . Hence, as $N \to \infty$,

$$(\mu - \lambda)T_N - \left(\log X_N(0) + \log(\mu - \lambda) - \log\left(1 + \frac{\lambda X_N(0)}{(\mu - \lambda)N}\right) - \log\mu\right) \to W,$$

in distribution. This is (1.7), which we have seen is equivalent to (1.4).

(c) Suppose $X_N(0) > (\mu - \lambda)N\omega(N)$. Let κ_N be the hitting time of $\lfloor (\mu - \lambda)N\omega(N) \rfloor$. Let $t_0 = 1/\omega(N)^{1/2}\lambda(\mu - \lambda)$, as in Lemma 4.2; by Lemma 4.2, $\kappa_N \leq t_0$ with probability 1 - o(1). Then T_N is the sum of κ_N and the time to extinction from state $\lfloor (\mu - \lambda)N\omega(N) \rfloor$. So T_N is bounded below by

$$\frac{1}{\mu - \lambda} \Big(\log \big((\mu - \lambda) N \omega(N) \big) + \log(\mu - \lambda) - \log \big(1 + \lambda \omega(N) \big) - \log \mu + W_N \big),$$

and, with probability 1 - o(1), bounded above by

$$\frac{1}{\mu - \lambda} (t_0(\mu - \lambda) + \log ((\mu - \lambda)N\omega(N)) + \log(\mu - \lambda) - \log (1 + \lambda\omega(N)) - \log \mu + W_N))$$

where W_N converges in distribution to a standard Gumbel random variable W. Since $(\mu - \lambda)t_0 = o(1)$ and $\omega(N) \to \infty$, it follows that, as $N \to \infty$,

$$(\mu - \lambda)T_N - (\log N + 2\log(\mu - \lambda) - \log \lambda - \log \mu) \to W,$$

in distribution. This is (1.6), which is equivalent to (1.4) in this range.

This completes the proof.

6. The critical regime

Our methods can also be applied in the critical regime, where $|\mu - \lambda| = O(N^{-1/2})$. In this case, there exist constants $\delta, c > 0$ (depending on $\limsup_{N\to\infty} (\lambda - \mu) N^{1/2}$), such that, regardless of the value taken by

 $X_N(0), \mathbb{P}(T_N \leq cN^{1/2}) > \delta$. One way to prove this is as follows: (a) apply Lemma 4.1 for $t = N^{1/2}$, and so $x(t) \leq 1/(\lambda t) = O(N^{-1/2})$, to show that, for some constant c_1 , with positive probability, $X_N(t) \leq c_1 N^{1/2}$, uniformly in $X_N(0)$; (b) compare $(X_N(t))$ with a linear birth-and-death chain with the same parameters, with initial state $c_1 N^{1/2}$, and show that, for some constant c_2 , with positive probability, $(X_N(t))$ reaches 0 in a further time $c_2 N^{1/2}$. Thus there is a positive probability of extinction by time $(c_1 + c_2)N^{1/2}$, whatever the initial state. It now follows, by repeated trials, that $\mathbb{P}(T_N > \omega(N)N^{1/2}) \to 0$ whenever $\omega(N) \to \infty$. Throughout the critical regime, a lower bound on the extinction time of the form $\mathbb{P}(T_N \leq \varepsilon(N)N^{1/2}) \to 0$ whenever $\varepsilon(N) \to 0$ can again be obtained by comparing with a suitable linear birth-and-death chain. Much more precise results concerning the process in the critical regime with initial state of order $N^{1/2}$ are given by Dolgoarshinnykh and Lalley (2006). In this regime, both the expected extinction time and the fluctuations are of order $N^{1/2}$, so that we do not have cut-off. This is in line with our results for the barely subcritical regime showing that cut-off becomes less pronounced as we approach the critical regime from below.

7. Total number of cases

We now turn our attention to the total number C_N of new cases (infection events) from the start of the epidemic until its extinction.

We will prove that, provided $X_N(0)(\mu - \lambda) \to \infty$, the total number C_N of cases is concentrated around its expectation, which is close to

$$N\frac{\mu}{\lambda}\log\left(1+\frac{\lambda X_N(0)}{N(\mu-\lambda)}\right)-X_N(0).$$

In the case where $X_N(0)/N(\mu - \lambda) \to 0$, our results imply that the expectation of C_N is close to $\frac{\lambda X_N(0)}{\mu - \lambda}$ and the variance is of the order at most $X_N(0)(\mu - \lambda)^{-3}$. (This can be interpreted as saying that the epidemic behaves as $X_N(0)$ independent outbreaks from a single initial infective.) If $X_N(0)/N(\mu - \lambda) \to \infty$, then our results show that C_N has expectation approximately $N \log \left(X_N(0)/N(\mu - \lambda) \right)$, and variance of order at most $N(\mu - \lambda)^{-2}$.

Theorem 7.1. Suppose that $\mu = \mu(N)$ and $\lambda = \lambda(N)$ are bounded away from both 0 and infinity. Suppose also that $(\mu - \lambda)N^{1/2} \to \infty$ as $N \to \infty$, and that $X_N(0)$ is non-random. Let $v_N(x) = \min\left(\frac{x^{1/2}}{(\mu - \lambda)^{3/2}}, \frac{N^{1/2}}{\mu - \lambda}\right)$. Then, for any $\varepsilon > 0$, there exists $K(\varepsilon)$ such that, for N sufficiently large,

$$\mathbb{P}\left(\left|C_N - \frac{\mu}{\lambda}N\log\left(1 + \frac{\lambda X_N(0)}{N(\mu - \lambda)}\right) + X_N(0)\right| \ge K(\varepsilon)v_N(X_N(0))\right) \le \varepsilon.$$

Proof. For X = 0, ..., N, let $\ell_X = \lambda(1 - X/N)$, so that the birth rate when $X_N(t) = X$ is equal to $X\ell_X$. Then C_N has the same distribution

as the number of births before extinction in a corresponding discrete-time birth-and-death chain $(\hat{X}_N(t))$, where the probability of a birth in state Xis $\ell_X/(\mu + \ell_X)$, and the probability of a death is $\mu/(\mu + \ell_X)$, so we can work with $(\hat{X}_N(t))$ instead.

Given $X_N(0) = X$, the number of births before extinction can be represented as the sum of independent random variables $C_{N,Y}$, for $Y = X, X - 1, \ldots, 1$, where $C_{N,Y}$ is the number of new cases starting in state Y until hitting state Y - 1. We will show that, for all $Y = 1, \ldots, N$,

$$\left(1+\frac{\mu\lambda}{N(\mu-\ell_Y)^2}\right)^{-1}\frac{\ell_Y}{\mu-\ell_Y} \le \mathbb{E}\,C_{N,Y} \le \frac{\ell_Y}{\mu-\ell_Y}.$$

Conditioning on the first step in a standard way, we see that $\mu \mathbb{E} C_{N,Y} = \ell_Y(\mathbb{E} C_{N,Y+1} + 1)$, for $1 \leq Y \leq N - 1$. We now proceed by downward induction. Since $\ell_N = 0$, both our upper and lower bounds on $\mathbb{E} C_{N,N}$ are equal to zero, which is the true value. Suppose that we have the stated upper bound on $\mathbb{E} C_{N,Y+1}$, so that, using the fact that $\mu \geq \lambda$ implies $\mu - \ell_Z \geq 0$ for all Z, $\mathbb{E} C_{N,Y+1} \leq \frac{\ell_{Y+1}}{\mu - \ell_{Y+1}} \leq \frac{\ell_Y}{\mu - \ell_Y}$. Then

$$\mathbb{E} C_{N,Y} = \frac{\ell_Y}{\mu} (\mathbb{E} C_{N,Y+1} + 1) \le \frac{\ell_Y}{\mu} \left(\frac{\ell_Y}{\mu - \ell_Y} + 1 \right) = \frac{\ell_Y}{\mu - \ell_Y},$$

which is the required upper bound on $\mathbb{E}C_{N,Y}$.

Suppose now that we have the stated lower bound on $\mathbb{E}C_{N,Y+1}$, so that

$$\mathbb{E} C_{N,Y+1} \ge \left(1 + \frac{\mu\lambda}{N(\mu - \ell_{Y+1})^2}\right)^{-1} \frac{\ell_{Y+1}}{\mu - \ell_{Y+1}}$$

.

Then, since $\ell_Y \ge \ell_{Y+1}$,

$$\begin{split} \mathbb{E} C_{N,Y+1} + 1 &\geq \frac{\mu}{\mu - \ell_{Y+1}} - \frac{\ell_{Y+1}}{\mu - \ell_{Y+1}} \left(1 - \frac{1}{1 + \frac{\mu\lambda}{N(\mu - \ell_{Y+1})^2}} \right) \\ &\geq \frac{\mu}{\mu - \ell_{Y+1}} - \frac{\ell_Y}{\mu - \ell_{Y+1}} \left(1 - \frac{1}{1 + \frac{\mu\lambda}{N(\mu - \ell_Y)^2}} \right) \\ &= \frac{\mu}{\mu - \ell_{Y+1}} - \frac{\ell_Y}{\mu - \ell_{Y+1}} \frac{\frac{\mu\lambda}{N(\mu - \ell_Y)^2}}{\left(1 + \frac{\mu\lambda}{N(\mu - \ell_Y)^2}\right)} \\ &= \frac{\mu}{\mu - \ell_{Y+1}} \left(\frac{1 + \frac{\mu\lambda}{N(\mu - \ell_Y)^2} - \frac{\lambda\ell_Y}{N(\mu - \ell_Y)^2}}{1 + \frac{\mu\lambda}{N(\mu - \ell_Y)^2}} \right) \\ &= \frac{\mu}{\mu - \ell_{Y+1}} \left(\frac{1 + \frac{\lambda}{N(\mu - \ell_Y)^2}}{1 + \frac{\mu\lambda}{N(\mu - \ell_Y)^2}} \right) = \frac{\mu}{\mu - \ell_Y} \frac{1}{\left(1 + \frac{\mu\lambda}{N(\mu - \ell_Y)^2}\right)} \end{split}$$

In the last step, we also used that $\mu - \ell_{Y+1} = \mu - \ell_Y + \lambda/N$, and so $1 + \frac{\lambda}{N(\mu - \ell_Y)} = \frac{\mu - \ell_{Y+1}}{\mu - \ell_Y}$. It follows, as required for the induction step, that

$$\mathbb{E} C_{N,Y} = \frac{\ell_Y}{\mu} (\mathbb{E} C_{N,Y+1} + 1) \ge \frac{\ell_Y}{\mu - \ell_Y} \left(1 + \frac{\mu \lambda}{N(\mu - \ell_Y)^2} \right)^{-1}$$

.

It follows from the above bounds that

$$\mathbb{E} C_N \leq \sum_{Y=1}^{X_N(0)} \frac{\ell_Y}{\mu - \ell_Y} \leq \int_{x=0}^{X_N(0)} \left(\frac{\mu}{\mu - \lambda(1 - x/N)} - 1\right) dx$$
$$= \frac{\mu N}{\lambda} \log \left(1 + \frac{\lambda X_N(0)}{N(\mu - \lambda)}\right) - X_N(0),$$

and, since $N(\mu - \lambda)^2 \rightarrow \infty$, also that, for N sufficiently large,

$$\mathbb{E} C_N \ge \sum_{Y=1}^{X_N(0)} \frac{\ell_Y}{\mu - \ell_Y} - \frac{2\mu\lambda^2}{N} \sum_{Y=1}^{X_N(0)} \frac{1}{(\mu - \ell_Y)^3}$$

Noting that

$$\begin{split} &\int_{x=0}^{X_N(0)} \left(\frac{\mu}{\mu - \lambda(1 - x/N)} - 1 \right) \, dx \\ &\leq \sum_{Y=0}^{X_N(0)-1} \frac{\ell_Y}{\mu - \ell_Y} = \sum_{Y=1}^{X_N(0)} \frac{\ell_Y}{\mu - \ell_Y} + \frac{\lambda}{\mu - \lambda} - \frac{\ell_{X_N(0)}}{\mu - \ell_{X_N(0)}} \\ &\leq \sum_{Y=1}^{X_N(0)} \frac{\ell_Y}{\mu - \ell_Y} + \frac{\mu(\lambda - \ell_{X_N(0)})}{(\mu - \lambda)(\mu - \ell_{X_N(0)})} \\ &\leq \sum_{Y=1}^{X_N(0)} \frac{\ell_Y}{\mu - \ell_Y} + \frac{\mu\lambda X_N(0)}{N(\mu - \lambda)^2}, \end{split}$$

that

$$\sum_{k=1}^{X} \frac{1}{(\mu - \ell_k)^3} \leq \min\left\{\frac{X}{(\mu - \lambda)^3}, \int_{x=0}^{\infty} \frac{1}{(\mu - \lambda + \lambda x/N)^3} dx\right\}$$
$$= \min\left\{\frac{X}{(\mu - \lambda)^3}, \frac{N}{2\lambda(\mu - \lambda)^2}\right\},$$

and that $X_N(0) \leq N$, we see that, for N large enough,

$$\left| \mathbb{E} C_N - \left(\frac{\mu N}{\lambda} \log \left(1 + \frac{\lambda X_N(0)}{N(\mu - \lambda)} \right) - X_N(0) \right) \right| \\ \leq \frac{\mu \lambda X_N(0)}{N(\mu - \lambda)^2} + \min \left(\frac{2\mu \lambda^2 X_N(0)}{(\mu - \lambda)^3 N}, \frac{\mu \lambda}{(\mu - \lambda)^2} \right) \\ \leq 3 \min \left(\frac{\mu^2 \lambda X_N(0)}{(\mu - \lambda)^3 N}, \frac{\mu \lambda}{(\mu - \lambda)^2} \right).$$
(7.1)

We now estimate the variance of C_N , noting that $\operatorname{Var} C_N = \sum_{Y=1}^{X_N(0)} \operatorname{Var} C_{N,Y}$. Starting from Y and until the hitting time $\tau_{N,Y}$ of Y - 1 by $(\hat{X}_N(t))$,

Starting from Y and until the hitting time $\tau_{N,Y}$ of Y - 1 by $(X_N(t))$, we can couple $(\hat{X}_N(t))$ with a discrete chain $(\hat{X}_{N,Y}(t))$ where, in any state, the probability of a birth is $\ell_Y/(\mu + \ell_Y)$ and the probability of a death is $\mu/(\mu + \ell_Y)$, in such a way that $\hat{X}_N(t) \leq \hat{X}_{N,Y}(t)$ for $0 \leq t \leq \tau_{N,Y}$. Letting $D_{N,Y}$ be the number of births in $(\hat{X}_{N,Y}(t))$ starting from Y until hitting Y - 1, we thus see that, under the coupling, $C_{N,Y} \leq D_{N,Y}$. It follows that

$$\operatorname{Var} C_{N,Y} = \mathbb{E} C_{N,Y}^2 - (\mathbb{E} C_{N,Y})^2 \le \mathbb{E} D_{N,Y}^2 = \operatorname{Var} D_{N,Y} + (\mathbb{E} D_{N,Y})^2.$$

For a given value of Y, consider a discrete random walk, starting at 1, with probability $p = \mu/(\ell_Y + \mu)$ of a down-step and probability $q = \ell_Y/(\ell_Y + \mu)$ of an up-step. Then $D_{N,Y}$ has the same distribution as $(T_{N,Y} - 1)/2$, where $T_{N,Y}$ is the hitting time of the origin for this walk. Standard arguments imply that the generating function $G_{N,Y}$ of $T_{N,Y}$ satisfies the recurrence $G_{N,Y}(z) = pz + qz(G_{N,Y}(z))^2$, and so $G_{N,Y}(z) = (1 - \sqrt{1 - 4pqz^2})/2qz$. Differentiating, we obtain $\mathbb{E} D_{N,Y} = \ell_Y/(\mu - \ell_Y)$ and $\operatorname{Var} D_{N,Y} = \ell_Y \mu (\ell_Y + \mu)/(\mu - \ell_Y)^3$, and hence $\operatorname{Var} C_{N,Y} \leq \frac{2\lambda\mu^2}{(\mu - \ell_Y)^3}$. Summing over Y,

$$\operatorname{Var} C_N \le 2\mu^2 \min\left(\frac{\lambda X_N(0)}{(\mu - \lambda)^3}, \frac{N}{(\mu - \lambda)^2}\right).$$
(7.2)

Suppose that $X_N(0)/N(\mu - \lambda) \to \infty$. Then for N large enough, the upper bound in (7.1) is equal to $\frac{3\mu\lambda}{(\mu-\lambda)^2} \leq N^{1/2}/(\mu-\lambda)$, and $\operatorname{Var} C_N \leq 2\mu^2 N/(\mu-\lambda)^2$ in this case. If $X_N(0)/N(\mu-\lambda)$ is bounded, then the upper bound in (7.1) is at most of the order $\frac{X_N(0)}{(\mu-\lambda)^3 N}$, and, for N large enough,

$$\frac{X_N(0)^{1/2}}{(\mu-\lambda)^{1/2}N^{1/2}} \cdot \frac{1}{(\mu-\lambda)N^{1/2}} \cdot \frac{X_N(0)^{1/2}}{(\mu-\lambda)^{3/2}} \le \frac{X_N(0)^{1/2}}{(\mu-\lambda)^{3/2}},$$

while $\operatorname{Var} C_N$ is of the order at most $\frac{\lambda X_N(0)}{(\mu - \lambda)^3}$. In both cases, the theorem follows by Chebyshev's inequality.

8. Numerical methods

The stochastic SIS logistic process $(X_N(t))$, as defined by events and rates (1.1), can be analysed through use of its corresponding Kolmogorov forward equations, as we now explain. We will not index all terms by Nexplicitly here for notational simplicity, but the method of analysis is for a population of size N. We start by writing $p_X(t) = \mathbb{P}(X_N(t) = X)$, and let $\mathbf{p}(t)$ be a column vector whose X-th entry is $p_X(t)$ – our convention is that such a vector starts at its 0-th element and has length N + 1. (We follow the more applied literature in treating $\mathbf{p}(t)$ as a column vector.) The Kolmogorov forward equations then take the form of a linear system of differential equations

$$\frac{\mathrm{d}p_X}{\mathrm{d}t} = -\left(\mu X + \lambda X \left(1 - \frac{X}{N}\right)\right) p_X + \lambda (X - 1) \left(1 - \frac{X - 1}{N}\right) p_{X-1} \\
+ \mu (X + 1) p_{X+1} , \quad 0 < X < N , \\
\frac{\mathrm{d}p_0}{\mathrm{d}t} = \mu p_1 , \\
\frac{\mathrm{d}p_N}{\mathrm{d}t} = -\mu N p_N + \lambda \frac{N - 1}{N} p_{N-1} .$$
(8.1)

These can be expressed in the form

$$\frac{\mathrm{d}\mathbf{p}}{\mathrm{d}t} = \boldsymbol{M}\mathbf{p} \;, \tag{8.2}$$

where M is an $(N+1) \times (N+1)$ matrix. Quantities of interest include:

$$\mathbb{E} X_N(t) = \mathbf{X} \cdot \mathbf{p}(t) , \quad F_{T_N}(t) = \mathbb{P}(T_N \le t) = p_0(t) , \quad f_{T_N}(t) = \mu p_1(t) ,$$
(8.3)

where **X** is a vector whose X-th element is X, F_{T_N} is the distribution function of the extinction time and f_{T_N} is the probability density function of the extinction time, which has the form above due to the second equation in (8.1). To integrate (8.2) numerically, we make use of the implicit Euler scheme, as has been advocated for stochastic epidemic models by Jenkinson and Goutsias (2012). We now sketch the arguments and approach presented in that paper. First, note that the solution of (8.2) is given by a matrix exponential,

$$\mathbf{p}(t) = \mathrm{Exp}(\boldsymbol{M}t)\mathbf{p}(0)$$

and therefore over a time interval [t, t+h] we can write

$$\mathbf{p}(t+h) = \operatorname{Exp}(\boldsymbol{M}h)\mathbf{p}(t) = (\boldsymbol{I} + \boldsymbol{M}h + O(h^2))\mathbf{p}(t) ,$$

$$\mathbf{p}(t) = \operatorname{Exp}(-\boldsymbol{M}h)\mathbf{p}(t+h) = (\boldsymbol{I} - \boldsymbol{M}h + O(h^2))\mathbf{p}(t+h) ,$$

(8.4)

where I is the $(N+1) \times (N+1)$ identity matrix. The implicit Euler numerical scheme is based on the second equation in (8.4) and involves solving the matrix equation

$$(I - Mh)\mathbf{p}_{+} = \mathbf{p} \tag{8.5}$$

to obtain an approximation \mathbf{p}_+ to $\mathbf{p}(t+h)$, in terms of an approximation \mathbf{p} to $\mathbf{p}(t)$, at each timestep, for example by using MATLAB's \backslash operator. We see from the first equation of (8.4) that the error introduced at each timestep is $O(h^2)$, and so the global error over the interval [0,t] is O(th) as $h \to 0$. This means that, in practice, for given choices of N, $X_N(0)$, λ and μ , we can tune h to achieve any desired accuracy. Suppose that \mathbf{p} is a probability vector; we now show that \mathbf{p}_+ generated by (8.5) is also a probability vector. First, premultiplying (8.5) by a row vector of ones, $\mathbf{1}^{\top}$, we obtain

$$1 = \mathbf{1}^{\top} \mathbf{p} = \mathbf{1}^{\top} I \mathbf{p}_{+} + h \mathbf{1}^{\top} M \mathbf{p}_{+} = \mathbf{1}^{\top} \mathbf{p}_{+} , \qquad (8.6)$$

which holds because M generates a Markov chain and so its columns must sum to zero: $\mathbf{1}^{\top}M = \mathbf{0}$. Secondly, from its definition, the off-diagonal elements of M are non-negative, meaning that the off-diagonal elements of I - Mh are non-positive and so, after checking for non-singularity, all elements of $(I - Mh)^{-1}$ are non-negative.

Often, solution of equations such as (8.2) for $\mathbf{p}(t)$ is more numerically efficient (particularly for calculating distributions of quantities like extinction times) than Monte Carlo methods that use (pseudo-)random number generation; see Keeling and Ross (2008). To see why this should be so for our case, note that one of the quantities we wish to calculate is the probability distribution function for the extinction time of an epidemic with rates λ and μ of order 1 with $\mu - \lambda \approx 10^{-3}$, population size $N = 10^7$, and $X_N(0) = N$. For these parameter values, our asymptotic results from Theorem 7.1 give that we expect to see over 10^8 events. Using Monte Carlo methods, we would need to simulate each of these to achieve one extinction, and would need to simulate many realisations of the entire epidemic to control the Monte Carlo error. In contrast, use of the forward Euler method as above requires just one realisation with the step-size h at an appropriate value relative to required numerical error, and solution of an N-dimensional sparse linear system at each step.

We performed a comparison of the simulation results based on implicit Euler solution of the Kolmogorov forward equations with our asymptotic results for a range of values of N from 10^1 to 10^7 , keeping $\mu = 1$ throughout, for two different scenarios. In the first scenario, we do not scale $\mathcal{R}_0 = \lambda$ with N, but instead leave it constant at 0.9. In the second scenario, we scale $1 - \mathcal{R}_0$ approximately like $N^{-1/3}$, starting with 0.9 for N = 10. The results for these two scenarios are pictured in Figures 3 and 4 respectively. These demonstrate that the asymptotic results can be a good approximation to the system behaviour for large population sizes (e.g., on the scale of a town, city or country) with regard to the extinction times, and, for any population of more than a hundred, also for the mean number of infectives. Furthermore, they show that for the unscaled case, extinctions happen relatively quickly for all values of N, but that, as \mathcal{R}_0 tends to 1 with N, extinctions can take an extremely long time to occur despite an initial fast decline in $\mathbb{E}X_N(t)$.

We also give numerical results for the total number of cases C_N . For this, we adapt the path sum numerical method introduced by Ross (2011); we shall show that this is suited to rapid calculation not only of the probability mass function (as in Ross (2011)) but also of the mean variance of C_N (see (8.9) and (8.11) below). To implement this method, we track the current state as $X_N \times B \in \{0, \ldots, N\} \times \{0, 1\}$, where the Bernoulli random variable B is defined to be 1 if the last event was an infection, and 0 if the last event was a recovery. Since we are interested in the final number of cases, we only need to consider the jump chain for this process. We will start the system in state $(X_{N,0}, B_0)$. Writing the state after u events as $(X_{N,u}, B_u)$, we see that

28

$$C_N = \sum_{u=0}^{\infty} B_u . aga{8.7}$$

(For our model, it is possible to recover C_N from the total number of events without the need for the auxiliary variable B; in more complex epidemic models, the auxiliary variable aids easy calculation of the quantities of interest.)

The transition probabilities for the jump chain are:

$$\mathbb{P}(X_{N,u+1} = X + 1, B_{u+1} = 1 | X_{N,u} = X, B_u) = \begin{cases} \frac{\lambda(1 - X/N)}{\lambda(1 - X/N) + \mu} & \text{if } X > 0, \\ 0 & \text{otherwise.} \end{cases}$$

$$\mathbb{P}(X_{N,u+1} = X - 1, B_{u+1} = 0 | X_{N,u} = X, B_u) = \begin{cases} \frac{r}{\lambda(1 - X/N) + \mu} & \text{if } X > 0, \\ 0 & \text{otherwise.} \end{cases}$$
(8.8)

Clearly, the state space decomposes into an absorbing class $\{(0,0), (0,1)\}$ and a transient class \mathcal{T} of states with a positive number of infectives. For a state i = (X, B), we set b_i equal to B. We write $P_{i,j}$ for the probability of moving from state i to state j as defined in (8.8). We write c_i for the expected value of the random variable C_N given that the initial state is i. A standard calculation conditioning on the first step then shows that, for each $i \in \mathcal{T}$, $c_i = \sum_j P_{ij}(c_j + b_j)$. Let $g_i = c_i + b_i$; then, for each $i, g_i = b_i + \sum_j P_{ij}g_j$. It follows that

$$g = (I - P)^{-1} \mathbf{b}$$
 (8.9)

We note that g_i differs from c_i by at most 1, and only in the case where $b_i = 1$. Note further that: (i) the inverse in this equation does not need to be calculated explicitly, and instead a system of linear equations can be solved, for example using the backslash operator $\$ in MATLAB, and (ii) we have restricted attention to the transient states so that the inverse in (8.9) is well-defined.

To study the variability of the distribution of C_N about its mean, we let $h_i = \mathbb{E}[C_N(C_N - 1) \mid X_N(0) = i]$. From Ross (2011), if $\varphi_i(z) := \mathbb{E}[z^{C_N}|X_N(0) = i]$, then

$$\varphi_i(z) = z^{b_i} \sum_j P_{i,j} \varphi_j(z) \qquad (i \in \mathcal{T}) .$$
(8.10)

We note that $h_i = \varphi_i''(1)$ and hence

$$\sum_{j \in \mathcal{T}} (\delta_{i,j} - P_{i,j}) h_j = 2b_i \sum_{j \in \mathcal{T}} P_{i,j} c_j \qquad (i \in \mathcal{T}) .$$
(8.11)

This equation for h_i can also be evaluated by solving a system of linear equations and used to calculate the standard deviation of C_N .

The results of comparing the path sum to the asymptotic formula for the mean, as well as the asymptotic variance bound (7.2), using the same parameter choices as previously, are shown in Figure 5. These results exhibit

rapid convergence of the (scaled) mean to its asymptotic value as the population size N gets large, and also rapid reduction in the variability of the distribution.

9. The relationship between the deterministic process and time of extinction

One feature that becomes apparent by studying the numerical results is that, especially in the barely subcritical regime, there is a clear distinction between the time that the size of the epidemic first becomes "small" (which is in practice often taken as a proxy for the end of the epidemic) and the time that extinction occurs with high probability.

In a situation where control measures have brought an epidemic into a subcritical regime, but observations of the prevalence of the epidemic are only partial, it is potentially important to infer the likely time of extinction from the existing observations and/or fits to models governed by differential equations, so that control measures can be maintained for long enough that the epidemic has died out with high probability.

For the SIS logistic process, our results indicate that an appropriate "guide time" to the extinction time is the time \hat{t} at which the deterministic process, given by (3.1) and starting from $X_N(0)/N$, reaches $\hat{x} = \frac{\mu}{(\mu-\lambda)N}$ (i.e., when the number of infectives is projected to be $\mu/(\mu - \lambda)$). Note that \hat{t} occurs significantly later than the time when the deterministic process reaches εN , for ε a small constant, but (in the near-critical regime) considerably earlier than the time when the deterministic process reaches 1/N, corresponding to a single remaining infective.

To see that the time \hat{t} has the desired property, we re-write our result (1.4) in terms of the function s(x) introduced in (3.2). We note that

$$s(X_N(0)/N) = \log\left(1 + \frac{\lambda}{\mu - \lambda} \frac{X_N(0)}{N}\right) - \log(X_N(0)/N)$$

=
$$\log\left(1 + (\mu - \lambda)N/\lambda X_N(0)\right) - \log(\mu - \lambda) + \log\lambda,$$

and deduce that

$$(\mu - \lambda)T_N - \left(s(\hat{x}) - s(X_N(0)/N) - \log\left(1 + \frac{\mu\lambda}{(\mu - \lambda)^2 N}\right)\right) \to W,$$

in distribution, as $N \to \infty$. As $(\mu - \lambda)^2 N \to \infty$, and (from (3.2)) $\hat{t} = (s(\hat{x}) - s(X_N(0)/N))/(\mu - \lambda)$, this implies that

$$(\mu - \lambda)(T_N - \hat{t}) \to W.$$
 (9.1)

Thus the distribution of T_N is concentrated in a window of width of order $1/(\mu - \lambda)$ around the guide time \hat{t} .

We now explain briefly how to express the probability of extinction by time t, asymptotically, in terms of the deterministic process x(t). For w a fixed constant, set $t_w = \hat{t} + w/(\mu - \lambda)$. From (9.1), we have that $\mathbb{P}(T_N \leq t_w) = e^{-e^{-w}} + o(1)$. From (3.2), we see that, for any α , and any constant w,

$$t_{\alpha}(\widehat{x}e^{-w}) - t_{\alpha}(\widehat{x}) = \frac{s(\widehat{x}e^{-w}) - s(\widehat{x})}{\mu - \lambda} = \frac{w + o(1)}{\mu - \lambda}$$

noting that $\hat{x}/(\mu - \lambda) = o(1)$. It follows that, for any value of $\alpha = x(0)$, and any fixed w,

$$x(t_w) = \hat{x}e^{-w}(1+o(1)),$$

and therefore

$$\exp(-Nx(t_w)(\mu - \lambda)/\mu) = e^{-e^{-w}} + o(1) = \mathbb{P}(T_N \le t_w) + o(1).$$

One can now see that

$$\sup_{t} \left| \mathbb{P}(T_N \le t) - \exp\left(-Nx(t)(\mu - \lambda)/\mu\right) \right| \to 0 \quad \text{as } N \to \infty.$$
 (9.2)

By the time $\tilde{t} \simeq \hat{t}$ when the number $X_N(\tilde{t})$ of infectives has dropped to $\mu/(\mu-\lambda)$, the epidemic is well within its final phase, and, as we have shown, after time \tilde{t} it is well-approximated by a linear birth-and-death chain, or by a subcritical branching process. The behaviour of such a process is well-understood: once it reaches a level of order $1/(\mu - \lambda)$, it fluctuates through states of that order until it goes extinct.

This is also an illustration of the effect of parameter choice on cut-off. Away from criticality, we have a strong cut-off phenomenon: the extinction time is concentrated within a window of time much shorter than the overall extinction time, reflecting the idea that, before the window, the process is "large" with high probability, and it is unlikely to drop to 0 very quickly. As we approach criticality, once the process drops to order $\mu/(\mu - \lambda)$, it can (but does not always) stay around that level for a relatively long time, giving a weaker cut-off.

As set out in the Introduction, we expect these findings to extend to a wide class of epidemic models (not only SIS models). In their final stages, many models will be well-approximated by a barely subcritical branching process independent of population size, and the nature of this approximating branching process will govern the final stages of the epidemic, for suitable parameter values. So we expect the prevalence curve of an epidemic to follow the solution of a differential equation closely until the number of infectives becomes small, and the actual time of extinction to fall in a window of time of width of order $1/\mu(1 - \mathcal{R}_0)$, containing within it the point where the differential equation predicts the number of infectives to be $1/(1 - \mathcal{R}_0)$. We expect a strong cut-off for epidemics away from criticality – once the process approaches extinction, it goes extinct very quickly – and a weaker cut-off as we approach criticality.

A typical sample path for a barely subcritical epidemic will resemble the sample paths of the SIS logistic process in Figure 2: they reach states of order $1/(1 - \mathcal{R}_0) = \mu/(\mu - \lambda)$ following a fairly smooth trajectory, but then fluctuate around that level for a period of time of order $1/\mu(1 - \mathcal{R}_0) =$

 $1/(\mu - \lambda)$, possibly nearing extinction several times, until finally the epidemic does die out.

To illustrate (9.2), we simulated the relationship between the mean prevalence of infection $\mathbb{E}X_N(t)/N$ (as computed from the Kolmogorov forward equations), which is very close to the deterministic process x(t)), and the probability of extinction $\mathbb{P}(X_N(t) = 0)$ (as computed from Theorem 1.1). We used the parameter values $N = X_0 = 10^7$, $\mu = 1$ and a variety of different values of $\mathcal{R}_0 = \lambda$: the results are shown in Figure 6.

As a first step towards more realistic models, there has been recent interest in a variant of an SIS epidemic where the durations of each case of infection are iid random variables Q_i with mean 1, not necessarily having an exponential distribution. Ball, Britton and Neal (2016) show that, if the epidemic starts with a single individual, the expected duration of the epidemic does not depend on the distribution of the Q_i . It would be interesting to investigate whether our results can be extended to this more general setting.

References

- Allen LJS (2008) An introduction to stochastic epidemic models. In Mathematical Epidemiology (Lecture Notes Math 1945) Springer, Berlin pp 81–130
- Andersson H, Djehiche B (1998) A threshold limit theorem for the stochastic logistic epidemic. J Appl Probab 35:662–670
- Antia R, Regoes RR, Koella JC, Bergstrom CT (2003) The role of evolution in the emergence of infectious diseases. Nature 426:658–661
- Ball F, Britton T, Neal P (2016) On expected durations of birth-death processes, with applications to branching processes and SIS epidemics. J Appl Probab 53:203–215
- Barbour AD (1976) Quasi-stationary distributions in Markov population processes. Adv Appl Probab 8:296–314
- Barbour AD, Hamza K, Kaspi H, Klebaner FC (2015) Escape from the boundary in Markov population processes. Adv Appl Probab 47:1190– 1211
- Bartlett MS (1957) On theoretical models for competitive and predatory biological systems. Biometrika 44:27–42
- Britton T, House T, Lloyd AL, Mollison D, Riley S, Trapman P (2015) Five challenges for stochastic epidemic models involving global transmission. Epidemics 10:54–57
- Bull J, Dykhuizen D (2003) Epidemics-in-waiting, Nature 426:609–610
- Camacho A, Kucharski A, Aki-Sawyerr Y, White MA, Flasche S, Baguelin M, Pollington T, Carney JR, Glover R, Smout E, Tiffany A, Edmunds WJ, Funk S (2015) Temporal Changes in Ebola Transmission in Sierra Leone and Implications for Control Requirements: a Real-time Modelling Study. PLOS Currents Outbreaks. doi:10.1371/currents.outbreaks.406ae55e83ec0b5193e30856b9235ed2

Chowell G, Sattenspiel L, Bansal S, Viboud C (2016) Mathematical models to characterize early epidemic growth: a review. Phys Life Rev 18:66–97

- Demiris N, O'Neill PD (2006) Computation of final outcome probabilities for the generalised stochastic epidemic. Stat Comput 16:309–317
- Diaconis P (1996) The cutoff phenomenon in finite Markov chains. P Natl Acad Sci USA 93:1659–1664
- Diekmann O, Heesterbeek H, Britton T (2013) Mathematical Tools for Understanding Infectious Diseases Dynamics. Princeton University Press
- Doering CR, Sargsyan KV, Sander LM (2005) Extinction times for birthdeath processes: Exact results, continuum asymptotics, and the failure of the Fokker-Planck approximation. Multiscale Model Sim 3:283–299
- Dolgoarshinnykh RG, Lalley SP (2006) Critical scaling for the SIS stochastic epidemic. J Appl Probab 43:892–898
- Eames KDT, Keeling MJ (2002) Modeling dynamic and network heterogeneities in the spread of sexually transmitted diseases. P Natl Acad Sci USA 99:13330–13335
- Feller W (1939) Die Grundlagen der Volterraschen Theorie des Kampfes ums Dasein in wahrscheinlichkeitstheoretischer Behandlung. Acta Biotheor 5:11–40.
- Heesterbeek H, Anderson RM, Andreasen V, Bansal S, De Angelis D, Dye C, Eames KTD, Edmunds WJ, Frost SDW, Funk S, Hollingsworth TD, House T, Isham V, Klepac P, Lessler J, Lloyd-Smith JO, Metcalf CJE, Mollison D, Pellis L, Pulliam JRC, Roberts MG, Viboud C, Isaac Newton Institute IDD Collaboration (2015) Modeling infectious disease dynamics in the complex landscape of global health. Science 347:6227:aaa4339. doi:10.1126/science.aaa4339
- Jenkinson G, Goutsias J (2012) Numerical integration of the master equation in some models of stochastic epidemiology. Plos One 7(5):e36160. doi:10.1371/journal.pone.0036160
- Keeling MJ, Ross JV (2008) On methods for studying stochastic disease dynamics. J Roy Soc Interface 5:171–181. doi:10.1098/rsif.2007.1106
- Kessler DA (2008) Epidemic size in the SIS model of endemic infections. J Appl Probab 45:757–778
- Klepac P, Metcalf JE, McLean AR, Hampson K (2013) Towards the endgame and beyond: complexities and challenges for the elimination of infectious diseases. Philos T Roy Soc B 368:1623:1–12
- Kryscio RJ, Lefèvre C (1989) On the extinction of the SIS stochastic logistic epidemic. J Appl Probab 27:685–694
- Kurtz TG (1971) Limit theorems for sequences of jump Markov processes approximating ordinary differential processes. J Appl Probab 8:354–366
- Leigh EJ (1981) The average lifetime of a population in a varying environment. J Theor Biol 90:213–239
- Levin D, Peres Y, Wilmer E (2009) Markov chains and mixing times. American Mathematical Society, Providence, RI.

- Nåsell I (1996) The quasi-stationary distribution of the closed endemic SIS model. Adv Appl Probab 28:895–932
- Nåsell I (2011) Extinction and Quasi-stationarity in the Stochastic Logistic SIS Model. Lecture Notes in Mathematics 2022. Springer-Verlag, Berlin, Heidelberg
- Norden RH (1982) On the distribution of the time to extinction in the stochastic logistic population model. Adv Appl Probab 14:687–708
- O'Regan SM, Drake JM (2013) Theory of early warning signals of disease emergence and leading indicators of elimination. Theor Ecol 6:333–357
- Renshaw E (2011) Stochastic Population Processes. Oxford University Press Ross JV (2011) Invasion of infectious diseases in finite homogeneous popu
 - lations. J Theor Biol 289:83–89
- Ross JV, Taimre T (2007) On the analysis of hospital infection data using Markov models. In: Kulasiri D, Oxley L (eds) Proceedings of the 17th Biennial Congress on Modelling and Simulation (MODSIM07), Modelling and Simulation Society of Australia and New Zealand, pp 2939–2945.
- Sagitov S, Shaimerdenova A (2013) Extinction times for a birth-death process with weak competition. Lith Math J 53:220–234
- Scheffer M, Bascompte J, Brock WA, Brovkin V, Carpenter SR, Dakos V, Held H, van Nes EH, Rietkerk M, Sugihara G (2009) Early-warning signals for critical transitions. Nature 461:53–59. doi:10.1038/nature08227
- Simon PL, Kiss IZ (2013) From exact stochastic to mean-field ODE models: a new approach to prove convergence results. IMA J Appl Math 78:945– 964
- Verhulst P-F (1838) Notice sur la loi que la population suit dans son accroisement. Corresp Math Phys 10:113–121
- Weiss GH, Dishon M (1971) On the asymptotic behavior of the stochastic and deterministic models of an epidemic. Math Biosci 11:261–265

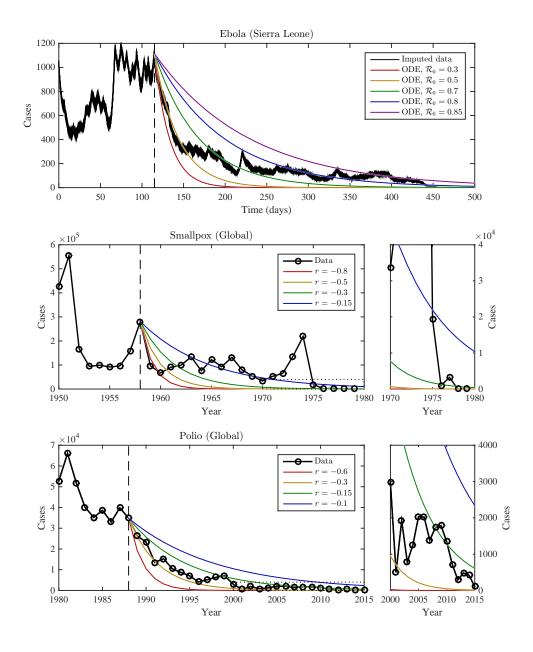


FIGURE 1. Data relating to diseases placed under control. For smallpox and polio the years when official eradication efforts started are indicated with dashed vertical lines. For Ebola, the effective start of control is estimated by eye and indicated by a dashed vertical line. Exponential decay curves with different rates are superimposed on the later part of the data. See Appendix for more details.

35

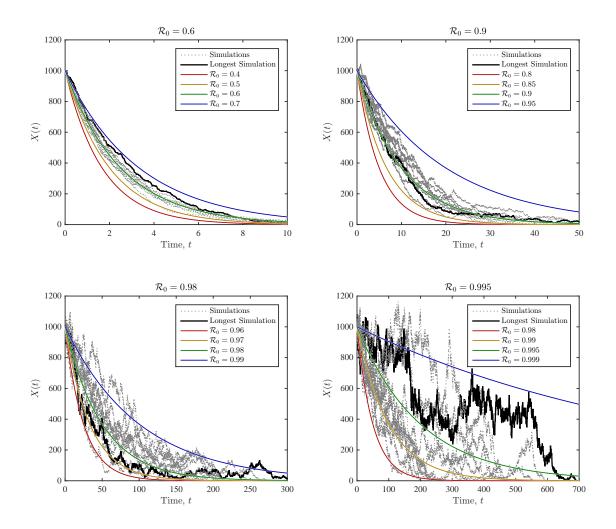


FIGURE 2. The SIS logistic model at different values of \mathcal{R}_0 . Other parameter choices are $\mu = 1$, $N = 10^6$ and $X_N(0) = 10^3$. Exponential decay curves with different rates are superimposed on ten realisations, with the longest one emphasised.

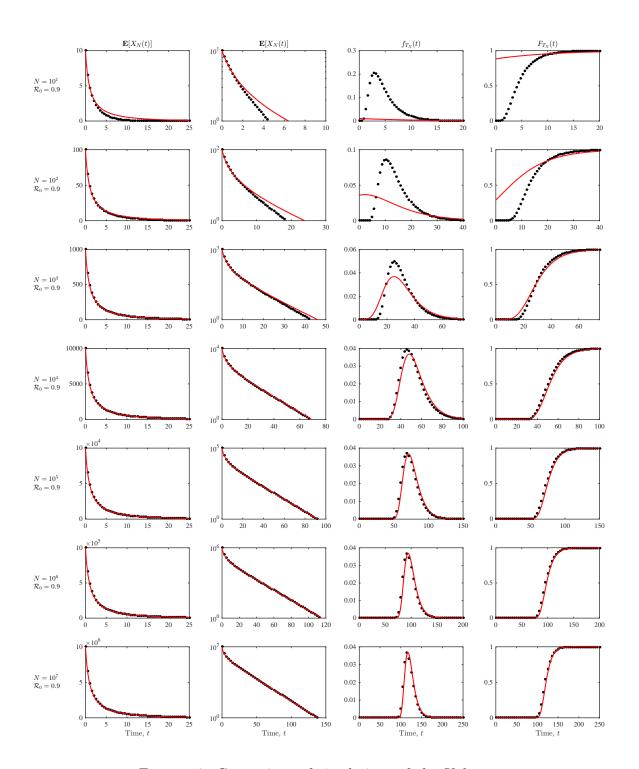


FIGURE 3. Comparison of simulations of the Kolmogorov forward equations (black dots) with asymptotic results (red solid lines). First column: Expected number of infectives with a linear y-axis. Second column: Expected number of infectives with a logarithmic y-axis. Third column: probability density function for the extinction time. Fourth column: distribution function for the extinction time. Rows represent different values of N from 10^1 to 10^7 with \mathcal{R}_0 unscaled.

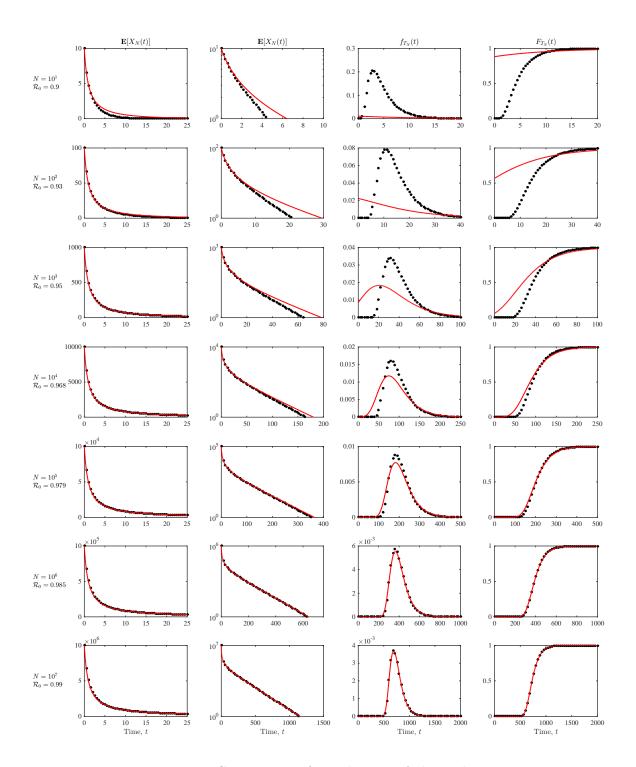


FIGURE 4. Comparison of simulations of the Kolmogorov forward equations (black dots) with asymptotic results (red solid lines). First column: Expected number of infectives with a linear y-axis. Second column: Expected number of infectives with a logarithmic y-axis. Third column: probability density function for the extinction time. Fourth column: distribution function for the extinction time. Rows represent different values of N from 10^1 to 10^7 with $1 - \mathcal{R}_0$ scaling approximately like $N^{-1/3}$.

38

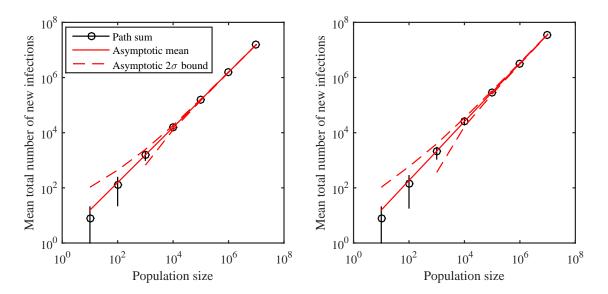


FIGURE 5. Comparison of total number of infections C_N with N for (left) the parameter choices in Figure 3 above and (right) the parameter choices in Figure 4 above. Calculations for the path sum are shown as black circles, with vertical black lines showing ± 2 standard deviations. The asymptotic formula for large N is shown as red lines, with the asymptotic bounds on ± 2 standard deviations shown as red dashed lines.

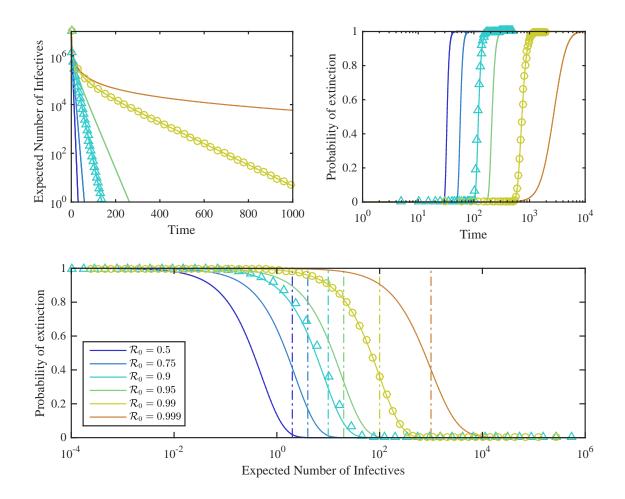


FIGURE 6. Comparison of asymptotic mean number of infectives $\mathbb{E} X_N(t)$ with asymptotic probability of extinction $\mathbb{P}(X_N(t) = 0)$ (from Theorem 1.1) for $N = X_N(0) = 10^7$, at a variety of different values of \mathcal{R}_0 for $\mu = 1$. The nonasymptotic results due to numerical integration of the Kolmogorov forward equations from Figures 3 and 4 are shown as markers next to the appropriate asymptotic curves. The vertical dot-dash lines in the bottom figure show the point where $Nx = \mu/(\mu - \lambda)$.

40

APPENDIX A. NUMERICAL EVIDENCE FROM REAL EPIDEMICS

We present here data from three real epidemics, pictured in Figure 1, where the behaviour near extinction fits our description of a barely subcritical epidemic. We make no claim that the real epidemics are well-modelled by any particular stochastic process. In reality, the available data will give only a partial picture of the true spread of disease, and the parameter values of any process will vary widely in time and geographical location.

Our first example is a simulation of the Ebola epidemic in 2014/15, based on the real-time study on Ebola in Sierra Leone performed by Camacho et al (2015).

The model is a stochastic compartmental model, with individuals in six different classes: S (susceptible), E_1 and E_2 (two non-infectious latent classes), I_c (infectious cases in the community), I_h (hospitalised infectious cases) and R (removed). The transitions can be encoded as:

$$\begin{array}{rcl} (S,E_1) & \rightarrow & (S-1,E_1+1) & \text{at rate} & \beta(I_c+I_h), \\ (E_1,E_2) & \rightarrow & (E_1-1,E_2+1) & \text{at rate} & 2\nu E_1, \\ (E_2,I_c) & \rightarrow & (E_2-1,I_c+1) & \text{at rate} & 2\nu E_2, \\ (I_c,I_h) & \rightarrow & (I_c-1,I_h+1) & \text{at rate} & \tau I_c, \\ (I_h,R) & \rightarrow & (I_h-1,R+1) & \text{at rate} & \gamma I_h. \end{array}$$

For instance, the first line represents an individual moving from class S to class E_1 at rate proportional to the number of infectives: we have simplified the model for our purposes by assuming that the number of susceptibles is constant throughout, and that the infection rate β is constant throughout the epidemic.

The parameter values were fitted to data by Camacho et al (2015) using a computationally intensive statistical framework in which β varies over time: the overlaid curves in Figure 1 show that fixed β – and therefore fixed $\mathcal{R}_0 = \beta(\gamma^{-1} + \tau^{-1})$ – does not work. The values derived from the data are $\nu^{-1} = 9.4$ days (so the latent period has a 2-Erlang distribution with mean ν^{-1}), $\tau^{-1} = 4.3$ days and $\gamma^{-1} = 6.9$ days.

For the simplified model, the population means in each compartment obey the ODEs

$$\frac{\mathrm{d}\,\mathbb{E}\,X_{E_1}}{\mathrm{d}t} = \beta(\mathbb{E}\,X_{I_c} + X_{I_h}) - 2\nu\,\mathbb{E}\,X_{E_1} , \quad \frac{\mathrm{d}\,\mathbb{E}\,X_{E_2}}{\mathrm{d}t} = 2\nu(\mathbb{E}\,X_{E_1} - \mathbb{E}\,X_{E_2}) ,$$
$$\frac{\mathrm{d}\,\mathbb{E}\,X_{I_c}}{\mathrm{d}t} = 2\nu\,\mathbb{E}\,X_{E_2} - \tau\,\mathbb{E}\,X_{I_c} , \qquad \qquad \frac{\mathrm{d}\,\mathbb{E}\,X_{I_h}}{\mathrm{d}t} = \tau\,\mathbb{E}\,X_{I_c} - \gamma\,\mathbb{E}\,X_{I_h} .$$
(A.1)

Based on this model, we generated estimates for the total number of infectives of Ebola over time. We took publicly available data on cumulative incidence for Ebola in Sierra Leone, and assumed: (i) that individual cases moved from I_c to I_h at some time t with uniform probability density in the day before they are reported; (ii) that the latent period ended at a time $t - t_1$, where t_1 is exponentially distributed with rate τ ; (iii) that the

infectious period ended at a time $t+t_2$, where t_2 is exponentially distributed with rate γ .

Simulating from this process ten times gives the black lines for the number of cases over time in Figure 1 (top plot), which are also compared to numerical solutions to (A.1) for different values of \mathcal{R}_0 , shown as coloured lines.

Our other two examples are broad-brush pictures of the courses of wellknown epidemics, where we present simply the number of recorded cases in each year. These examples are smallpox (Figure 1, middle plot), which was subject to a successful global eradication campaign, and polio (Figure 1, bottom plot), which is currently subject to a global eradication campaign that will hopefully be successful soon. In these cases, we do not compare to specific ODE models, but instead to exponential decay curves of the form e^{-rt} , for various values of r, which are shown as coloured lines.

For all three real examples, we see that the duration of the epidemics after they have been brought under control is longer than might be predicted from the smooth curves, which would be associated with straightforwardly subcritical epidemics with much smaller extinction times, of order $\log N/\mu$. We also see that the prevalence does not decay smoothly in time in its final stages, but goes up and down several times before extinction.

We take the smallpox data as an example, to show how the data might be seen to fit our predicted behaviour for a barely subcritical process in a rough quantitative sense. The highly infectious period for smallpox is on the order of a week, which corresponds to a value of μ^{-1} around 0.02 years. The middle graph in Figure 1 shows the total number of cases in each year, which is consistent with the number of cases remaining of order about 2000 over the period 1958-1973. Aggregating cases over full years obscures any erratic behaviour within each year. It seems clear that the effective value of \mathcal{R}_0 for the smallpox epidemic must have varied considerably over that period, due to seasonal effects, changes in control measures both globally and in response to local outbreaks, and other factors. However, the total number of cases of smallpox over the 15-year period is around two million, and the total number of cases at any time in 1973 is within a few thousand of the number in 1958, indicating that the average value of \mathcal{R}_0 over this period is within about 0.001 of 1. For a postulated average value of \mathcal{R}_0 of 0.999, our hypothesis would suggest a period of time of order 20 years before

42 GRAHAM BRIGHTWELL, THOMAS HOUSE, AND MALWINA LUCZAK

extinction, during which the number of cases would resemble a random walk remaining of order at most 1000, which is a reasonable fit to the data shown.

DEPARTMENT OF MATHEMATICS, LONDON SCHOOL OF ECONOMICS E-mail address: g.r.brightwell@lse.ac.uk URL: http://www.maths.lse.ac.uk/Personal/graham/

SCHOOL OF MATHEMATICS, UNIVERSITY OF MANCHESTER E-mail address: thomas.house@manchester.ac.uk URL: http://personalpages.manchester.ac.uk/staff/thomas.house/

School of Mathematics and Statistics, University of Melbourne $E\text{-}mail\ address: mluczak@unimelb.edu.au$