

African Swine Fever Detection and Transmission Estimates Using Homogeneous Versus Heterogeneous Model Formulation in Stochastic Simulations Within Pig Premises

Amos Ssematimba (✉ amos.ssematimba@gmail.com)

Gulu University

Sasidhar Malladi

University of Minnesota

Peter J. Bonney

University of Minnesota

Kaitlyn M. St. Charles

University of Minnesota

Timothy C. Boyer

Animal and Plant Health Inspection Service

Timothy Goldsmith

University of Minnesota

Carol J cardona

University of Minnesota

Cesar A. Corzo

University of Minnesota

Marie R. Culhane

University of Minnesota

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1 **African swine fever detection and transmission estimates using homogeneous versus**
2 **heterogeneous model formulation in stochastic simulations within pig premises**

3 Amos Ssematimba^{1,2,*}, Sasidhar Malladi^{1,\$}, Peter J. Bonney¹, Kaitlyn M. St. Charles¹, Timothy
4 C. Boyer³, Timothy Goldsmith¹, Carol J. Cardona¹, Cesar A. Corzo¹, and Marie R. Culhane¹

5 ¹ Secure Food Systems Team, College of Veterinary Medicine, University of Minnesota, Saint
6 Paul, Minnesota, 55108, USA

7 ² Department of Mathematics, Faculty of Science, Gulu University, P.O. Box 166, Gulu, Uganda

8 ³ U.S. Department of Agriculture, Animal and Plant Health Inspection Service, Veterinary
9 Services, Science Technology and Analysis Services, Center for Epidemiology and Animal
10 Health, Fort Collins, Colorado, USA

11 ^{\$}The authors contributed equally

12 *Corresponding author email: amos.ssematimba@gmail.com

13 **Abstract**

14 This study aimed to assess the impact on within-herd transmission dynamics of African swine
15 fever (ASF) when the models used to simulate transmission assume there is homogeneous
16 mixing of animals within a barn. Barn-level heterogeneity was explicitly captured using a
17 stochastic, individual pig-based, heterogeneous transmission model that considers three types of
18 infection transmission, 1) within-pen via nose-to-nose contact; 2) between-pen via nose-to-nose
19 contact with pigs in adjacent pens; and 3) both between- *and* within-pen via distance independent
20 mechanisms (e.g., via fomites). Predictions were compared between the heterogeneous and the
21 homogeneous Gillespie models. Results showed that the predicted mean number of infectious
22 pigs at specific time points differed greatly between the homogeneous and heterogeneous models
23 for scenarios with low levels of between pen contacts via distance independent pathways and the
24 differences between the two model predictions were more pronounced for the slow contact rate
25 scenario. The heterogeneous transmission model results also showed that it may take
26 significantly longer to detect ASF, particularly in large barns when transmission predominantly
27 occurs via nose-to-nose contact between pigs in adjacent pens. The findings emphasize the need
28 for completing preliminary explorations when working with homogeneous mixing models to
29 ascertain their suitability to predict disease outcomes.

30 **Keywords:** African swine fever; Gillespie algorithm; Heterogeneity; Transmission models;
31 Homogeneous mixing

32 **Introduction**

33 African swine fever (ASF) is a World Organisation for Animal Health-listed, highly fatal, and
34 socioeconomically devastating viral disease of domestic and feral swine that currently has
35 neither an approved vaccine nor treatment. It is endemic in some parts of the world (e.g., sub-
36 Saharan Africa) and there are ongoing outbreaks in both Asian and European countries (Normile,
37 2019). After ASF virus (ASFV)-specific prevention measures fail, the incursion of ASFV causes
38 disease outbreaks that can be far-reaching and long-lasting. Strategies to rapidly control
39 outbreaks include test and removal (with mixed results (Swine Health Information Center,
40 2021)) as well as culling affected farms; although, in some cases culling just the affected farms
41 results in only partial control and hence entire regions have to be depopulated (e.g., see (Council
42 of the European Union, 2002; USDA, 2021)). Such measures negatively impact the swine
43 industry and more generally food security and the livelihoods of farmers and those in the allied
44 industries (Mason-D’Croz et al., 2020; OIE, 2020).

45 To improve outbreak management strategies, proactive risk assessments that include simulation
46 modeling of disease transmission dynamics under varying circumstances can be used to guide
47 policy for effective surveillance of infected farms, deployment of critical activities to prevent
48 further outbreak spread, and continuity of business for farms that are not known to be infected
49 within a region (see (Hayes et al., 2021) for ASF modeling review). When focusing on modeling
50 ASFV transmission within a single swine barn, it is important to include several pathways and
51 mechanisms that can facilitate its spread. These may include direct (e.g., nose-to-nose) or
52 indirect (e.g., fomite-mediated) contact between pigs (Depner et al., 2016; Schulz et al., 2019;
53 Lee et al., 2020). Additional factors that may influence the speed of ASFV spread include those

54 related to virus-host interactions, farm management, and environmental conditions (Schulz et al.,
55 2019).

56 Ideally, in order to improve the accuracy of the model outcomes, all details of the influential
57 disease spread mechanisms would need to be explicitly captured in the disease transmission
58 model. Such details are better captured in heterogeneous models rather than homogeneous
59 models which assume uniform mixing between all animals in the population. Although it is
60 recognized that using a homogenous model with a uniform mixing assumption may result in
61 oversimplification and underestimation, the heterogeneous models are not often deployed due to
62 their computational intensity and necessitation of more refined data to parameterize (Keeling and
63 Rohani, 2008). The rapid results generated from the homogeneous models can also provide quick
64 insights into disease spread dynamics and are therefore useful under time-sensitive
65 circumstances. For highly transmissible and fast-spreading swine diseases like foot and mouth
66 disease, homogeneous mixing within the barn can be a reasonable simplifying assumption
67 (Kinsley et al., 2018). Detailed descriptions of how these approaches may differ have been
68 reported elsewhere (Hethcote, 1996; Bansal et al., 2007; Burr and Chowell, 2008; Keeling and
69 Rohani, 2008; Kong et al., 2016; Andraud and Rose, 2020).

70 A variety of approaches have been used in the literature to model within barn ASFV
71 transmission. For example, Guinat et al. (2016) and Faverjon et al. (2021) model transmission
72 heterogeneously, and both assume that transmission occurs due to direct contacts within-pen and
73 between-pen. A similar approach is used by Nielson et al. (2017) who assume ASFV can be
74 transmitted to non-adjacent pens according to a distance-dependent scaling factor. On the other
75 hand, Barongo et al. (2016) and Malladi et al. (2022) assume homogeneous mixing of the pigs in
76 the population.

77 In this article, we explain a novel heterogeneous approach that was developed and assumes that
78 ASFV transmission occurs due to direct within- and between-pen contacts as well as via distance
79 independent pathways. We explore the effect of heterogeneity on simulated output through
80 comparison to the output from a homogeneous mixing Gillespie algorithm, i.e., a continuous-
81 time transmission model used for fast simulation of stochastic processes (Gillespie, 1977;
82 Vestergaard and Génois, 2015). The developed model captures clustering of infected pens with
83 jumps between pens via distance-independent pathways based on ASF outbreak observations.

84 We evaluated simulated output from the heterogeneous model for a variety of scenarios. These
85 scenarios included variations in a) the number of infectious pigs over time post-virus exposure
86 for slow and fast contact rates; b) contact patterns as informed by barn layout and pen structure,
87 for example, that varied by the relative importance of within-pen and between-pen spread and
88 distance independent transmission; and c) the time to detection based on elevated mortality for
89 different population sizes and the amount of transmission due to distance independent pathways.

90 Since heterogeneity in infection rates can influence epidemic spread (Cai et al., 2013), our
91 endeavors included the imperative step of measuring the impact of the underlying model
92 assumptions in our efforts to improve interpretation and model selection.

93 **Materials and Methods**

94 We used a stochastic individual-based heterogeneous transmission model to simulate ASFV
95 spread within one growing pig production premises. The heterogeneous transmission model
96 incorporates different transmission rates within and between pig subpopulations such as pens and
97 rooms. The model simulates the number of pigs in susceptible (S), latent (E), infectious (I),
98 recovered (R), and dead (D) states in 0.01-day time steps (Δt). The number of pigs with mild

99 clinical signs, severe clinical signs, and detectable viremia were also reported to support
 100 surveillance evaluation; however, these states do not impact the transmission dynamics in the
 101 current model. An infectious pig may transition to the dead state with a probability P_{mort} or
 102 transition to the recovered state otherwise. The disease state durations were all modeled to be
 103 Gamma distributed. In what follows, we provide the equations for various within- and between-
 104 pen transmission mechanisms. Wherever used, I_s and N_s are, respectively, the number of the
 105 infectious and the total number of pigs in a pen s . The presented formulations for the
 106 transmission terms follow derivations described previously (Becker, 1989; Ssematimba et al.,
 107 2018).

108 **Transmission term expression for contacts that exclusively occur within a pen**

109 We assume that transmission via contacts that exclusively occur within a pen is frequency-
 110 dependent (e.g., direct contact with pen mates). The number of contacts each susceptible pig has
 111 with other pigs in the same pen per unit time is assumed to be Poisson distributed with mean B_d
 112 per unit time. The probability P_d that a susceptible pig in a pen k has a contact with at least one
 113 infectious pig within the same pen in time step Δt is given by

114

$$P_d = 1 - e^{-B_d \frac{I_k}{N_k} \Delta t} \quad \text{Eq. 1}$$

115

116 **Between-pen transmission via nose-to-nose contact with adjacent pens**

117 Here we consider a pen k with two adjacent pens $k-1$ and $k+1$ separated by railings where nose-
 118 to-nose contact between pigs may occur. Let η be the mean number of contacts per unit time with

119 pigs in adjacent pens (e.g., nose-to-nose) per pig per railing and β_{ns} be the total number of
 120 contacts with pigs in adjacent pens per unit time. Because there are two adjacent pens, the mean
 121 number of contacts with pigs in adjacent pens per time step would be $2\eta\Delta t$ or equivalently $B_{ns}\Delta t$.
 122 Assuming that the number of contacts is Poisson distributed, then the probability P_{ns} that a
 123 susceptible pig has a nose-to-nose contact with at least one infectious pig in one of the two
 124 adjacent pens in time step Δt is given by,

$$P_{ns} = 1 - e^{-B_{ns} \frac{I_{k-1} + I_{k+1}}{N_{k-1} + N_{k+1}} \Delta t} \quad \text{Eq. 2}$$

125 We now consider an edge pen k with only one adjacent pen $k+1$. In this case, the expected
 126 number of contacts between pigs in pens k and $k+1$ in a time step is $\eta\Delta t$ or equivalently
 127 $0.5*B_{ns}\Delta t$. Note that if the adjacent pen contact rate was not adjusted to $1/2\beta_{ns}$ for an edge pen,
 128 there would be a discrepancy in the number of contacts between the source and recipient pens.

129 **Between- and within-pen transmission via distance independent mechanisms**

130 We consider that contacts for distance independent transmission mechanisms e.g., via fomites,
 131 people, etc., may occur at a similar frequency throughout the barn regardless of whether the pigs
 132 are within the same pen. Assuming that the number of contacts per unit time for these
 133 mechanisms is Poisson distributed with mean β_p , the probability that at least one of the contacts
 134 via distance independent transmission mechanisms in time step Δt is with an infectious pig is
 135 given by:

$$P_p = 1 - e^{-\beta_p \frac{\sum_i I_i}{\sum_i N_i} \Delta t} \quad \text{Eq. 3}$$

136

137 The overall probability that a susceptible pig in pen k at time t becomes infected by $t+\Delta t$ is given
 138 by P_o in equations Eq. 4 and Eq. 5 for a pig in non-edge and edge pens respectively

$$P_o = 1 - e^{-\left(B_p \frac{\sum_i I_i}{\sum_i N_i} + B_d \frac{I_k}{N_k} + B_{ns} \frac{I_{k-1} + I_{k+1}}{N_{k-1} + N_{k+1}}\right) \Delta t} \quad \text{Eq. 4}$$

$$P_o = 1 - e^{-\left(B_p \frac{\sum_i I_i}{\sum_i N_i} + B_d \frac{I_k}{N_k} + \frac{B_{ns} I_{k+1}}{2 N_{k+1}}\right) \Delta t} \quad \text{Eq. 5}$$

139

140 **Reparametrizing to evaluate the relative importance of spread pathways and facilitate**
 141 **translation of transmission rates from published literature**

142 Experimental contact rate estimates for ASF and other diseases from the literature are often
 143 provided separately for contacts that occur exclusively within or between pens. In what follows,
 144 we derive equations to calibrate the contact rates β_d , β_p , β_{ns} in our formulation according to
 145 published contact rates β_w and β_b by equating the force of infection (infection hazard for a
 146 susceptible pig) terms for within- and between-pen transmission components. Let θ be the mean
 147 proportion of the between pen contacts associated with distance independent pathways. Then
 148 equating the force of infection for a pig in pen k via nose-to-nose contacts with pigs in adjacent
 149 pens gives

$$B_b * (1 - \theta) \frac{I_{k-1} + I_{k+1}}{N_{k-1} + N_{k+1}} = B_{ns} \frac{I_{k-1} + I_{k+1}}{N_{k-1} + N_{k+1}} \quad \text{Eq. 6}$$

150

$$B_{ns} = B_b * (1 - \theta) \quad \text{Eq. 7}$$

151 The force of infection term for a pig in pen k for between-pen contacts via distance independent
 152 mechanisms is given by Eq. 8. The right-hand side (RHS) of Eq. 8. is the product of the contact
 153 rate for distance independent pathways, multiplied by the probability that the contact is with a

154 pig in another pen and the probability that the contact is with an infectious pig given that it is in
 155 another pen.

$$B_p \theta \frac{\sum_{i \neq k} I_i}{\sum_{i \neq k} N_i} = B_p \frac{\sum_{i \neq k} N_i}{\sum_i N_i} \frac{\sum_{i \neq k} I_i}{\sum_{i \neq k} N_i} \quad \text{Eq. 8}$$

156

$$B_p = B_b \theta \frac{\sum_i N_i}{\sum_{i \neq k} N_i} \quad \text{Eq. 9}$$

157 Similarly, the force of infection for direct within-pen transmission under the two formulations
 158 would be as given in Eq. 10. The second term on the RHS of Eq. 10 is the contact rate for
 159 distance independent mechanisms multiplied by the probability that the contact occurs within the
 160 same pen and the probability that the contact is with an infectious pig if it occurs within the same
 161 pen.

162

$$B_w \frac{I_k}{N_k} = B_d \frac{I_k}{N_k} + B_p \frac{N_k I_k}{\sum_i N_i * N_k} \quad \text{Eq. 10}$$

$$B_d = B_w - B_p \frac{N_k}{\sum_i N_i} \quad \text{Eq. 11}$$

163

164 Equations 7, 9 and 11 can be used to calibrate the model parameters for the alternative
 165 formulations in the literature that are estimated exclusively for within- and between-pen contact
 166 rates. The parameter θ can be used to control the proportion of between-pen transmission
 167 occurring via nose-to-nose contact with pigs in adjacent pens or through distance-independent
 168 mechanisms.

169 **Comparison with Gillespie direct approach**

170 We performed a simulation evaluation to help identify conditions where the heterogeneous
171 model output differs from the homogeneous Gillespie algorithm. The heterogeneous and
172 homogeneous models were parameterized according to the Genotype II highly virulent Georgia
173 2007/1 ASFV strain and compared by the mean number of infectious pigs over time post virus
174 exposure from 10,000 simulation iterations. The heterogeneous model simulations were based on
175 a 1,200 growing pig barn with two rows of 15 pens each separated by a central alleyway and 40
176 pigs per pen. The heterogeneous model was compared to the homogeneous Gillespie algorithm
177 implementation (Gillespie, 1977) with four disease states (susceptible, latent, infectious, and
178 dead) where the pen structure within the barn was not considered, and all pigs were assumed to
179 die following infection (c.f. 40% in Table 1 for moderately virulent strain). The mean latent and
180 infectious periods were set to 4.0 and 4.5 days, respectively, based on a literature review by
181 Hayes *et al.* (2021). In the heterogeneous model, the relative values of β_d , β_p , β_{ns} were varied
182 while equating their sum to the daily adequate contact rate in the homogeneous model to enable
183 comparison. We evaluated two contact rate scenarios based on literature. The β_w and β_b values in
184 the fast contact rate scenario were, respectively, 2.62 and 0.99 per day based on (Hu et al., 2017).
185 In the slow contact rate scenario β_w of 0.6 per day and β_b of 0.3 per day were applied based on
186 Guinat *et al.* (2016).

187 Let φ be the fraction of transmission from an infected pig that occurs within a pen, i.e.,
188 $\varphi = B_w / (B_w + B_b)$. Then φ was calculated to be 0.73 and 0.67 based on the adequate contact rate
189 estimates from Hu *et al.* (2017) and Guinat *et al.* (2016), respectively. In the first set of
190 comparisons, we evaluated φ values of 0.5, 0.7 and 0.9 while assuming purely distance

191 independent between pen transmission ($\theta = 1$) to evaluate the impact of the relative magnitudes
192 of within- and between-pen transmission.

193 In the next set of comparisons, we compared the Gillespie model with the heterogeneous model
194 for θ values of 0.05, 0.5 and 1 to help infer the impact of the relative contribution of distance
195 independent between pen spread versus spread to adjacent pens via nose to nose contact. The
196 fraction of within pen transmission φ was assumed to be 0.7 for these simulations based on Hu *et al.*
197 *al.* (2017) and Guinat *et al.* (2016).

198 **Impact of heterogeneous within-herd transmission on the predicted time to ASF detection** 199 **via increased mortality**

200 In this section, we evaluate the predicted time to ASF detection under various transmission
201 scenarios to understand how heterogeneous transmission, transmission in clusters due to adjacent
202 pen spread, and barn size impacts the time to detection via daily mortality trigger thresholds. We
203 previously estimated the transmission parameters for moderately virulent ASFV strains using
204 data presented in (de Carvalho Ferreira et al., 2013) in previous work (Malladi et al., 2022).

205 Analysis of mortality data from five flows and 248 finisher herds indicated that a daily mortality
206 trigger threshold of 5 per 1000 finisher pigs results in a low frequency of false triggers (Malladi
207 et al., 2022). The time to detection was then calculated as the earliest day when the simulated
208 daily mortality exceeded a specified fraction of the herd (i.e., daily mortality trigger threshold).

209 The disease state duration and other transmission parameters for moderately virulent ASFV
210 strains applied in time to detection analysis section are summarized in Acknowledgements

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219 **Authors' contributions**

220 AS, SM, PJB, KMS, TCB, TG, CJC, CAC, MRC conceived the ideas of the study; AS, SM, PJB,
221 TB conceived the ideas for the analysis; SM, PJB, AS performed the analyses; SM, AS wrote the
222 manuscript; PJB, MRC were major contributors in writing the manuscript and all other authors
223 commented on the manuscript. All authors read and approved the final manuscript.

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238 **Availability of data and materials**

239 Not applicable.

240 **Ethics approval**

241 The study used already published data.

242 **Competing interests**

243 The authors declare that they have no competing interests.

244 Table 1 and based on Malladi et al. (2022). Two barn configurations and population sizes were
245 evaluated, specifically: 1) a growing pig barn with two rows of pens separated by a central
246 alleyway with each row having 15 pens, each holding 40 pigs with a total population of 1,200
247 pigs; and 2) a hypothetical growing pig barn with two rows of 120 pens each, again separated by
248 a central hallway and each pen holding 40 pigs, but with a larger total population of 4,800 pigs.
249 Although the larger quad barns with 4,800 pigs are typically organized into multiple rooms, we
250 considered a conceptual 4,800 pig barn with a single airspace in this analysis to understand how
251 barn size impacts the relative difference between homogenous and heterogenous spread. The
252 results were estimated from 10,000 simulation iterations.

253 **Results**

254 The results on the number of infectious pigs on various days post-exposure in a 1,200 growing
255 pig barn for the Gillespie algorithm and the heterogeneous transmission model with various
256 fractions of within pen transmission (ϕ) are shown and compared in Figure 1. The predicted
257 number of infectious pigs per the heterogeneous model was similar to the homogeneous
258 Gillespie model when ϕ was 0.5. There was a more gradual increase in the predicted number of
259 infectious pigs from outputs of the heterogeneous model when the fraction of within pen spread
260 (ϕ) was increased to 0.7 and 0.9. The differences between the heterogeneous and homogeneous
261 results were more pronounced in the slow contact rate scenario.

262 The predicted number of infectious pigs on various days post-exposure in a barn with 1,200 pigs
263 per the Gillespie algorithm and the heterogeneous transmission model with various fractions of
264 distance independent between pen transmission (θ) and ϕ set to 0.7 are shown and compared in
265 Figure 2. There was a greater difference between the heterogeneous model and the homogeneous
266 Gillespie model at low θ values. Once again, the differences between the heterogeneous and
267 homogeneous model results were more remarkable in the slow contact rate scenario. For
268 example, and to summarize, the heterogeneous model predicted far fewer than 100 infectious
269 pigs at 40 days post-exposure for slow contact rate scenarios and with either a high ϕ or low θ ;
270 whereas the homogeneous model predicted well over 100 infectious pigs at the same time point
271 post-exposure.

272 The predicted time to detect ASF based on a daily mortality trigger threshold of 5 per 1000 pigs
273 is shown in

274 Table 2. We observe that it may take significantly longer to detect ASF with the heterogeneous
275 transmission model particularly in large populations in the same premises, e.g., barns with a total
276 population of 4,800 pigs or more, especially when the transmission predominantly occurs via
277 nose-to-nose contact between pigs in adjacent pens. This effect is most noticeable when

278 between-pen spread mainly occurs via nose-to-nose contacts (i.e., low θ of 0.05) and where
279 transmission is clustered.

280 **Discussion**

281 Improved preparedness for and response to ASF outbreaks is vitally important given the
282 socioeconomic impact associated with the ongoing epidemics globally. Mathematical models
283 provide a platform to evaluate control strategies like control area surveillance protocols and can
284 thus inform disease management policies and proactive risk assessments. Several approaches to
285 disease dynamics modeling exist and one of the broad classifications is heterogeneous versus
286 homogeneous transmission. These two approaches can have discrepancies in predicted outcomes
287 and the choice of which to use is often informed by factors like the objective of the analysis, data
288 availability, suspected or identified transmission pathways, and computational effort, among
289 others. Comparing and contrasting these approaches can help harmonize and build confidence in
290 their applications, ultimately providing more informative analyses of outbreak response and
291 prevention.

292 From the model results displayed in Figure 1, we observe that the predicted mean number of
293 infectious pigs was similar across the two approaches when the fraction of within pen
294 transmission ϕ was 0.5. The predicted numbers from the two models diverge as ϕ increased. In
295 particular, the predicted mean number of infectious pigs was lower in the heterogeneous model
296 at ϕ of 0.7 and 0.9 during the early stages of herd infection. The differences were more apparent
297 in the slow contact rate scenario. This is likely due to the greater chance of generating higher ϕ
298 values that occurs when the infectious pig makes contact with another infected pig in the same

299 pen thus does not result in disease transmission to a susceptible pig, which is a type of contact
300 that subsequently results in slower between-pen transmission.

301 The results in Figure 2 and Table 2 show that there is a greater difference between the
302 heterogeneous model and the homogeneous Gillespie model when the proportion of between pen
303 contacts due to distance independent pathways θ was low. Note that the parameter θ could be
304 used to capture disease spread patterns observed in field ASF outbreaks that have been described
305 as involving clusters of infected pens with jumps between pens via distance-independent
306 pathways (Yaros, 2019; Nga et al., 2020). We hypothesize that the observed discrepancy is
307 possibly due to the fact that between-pen spread would predominantly occur via nose-to-nose
308 contact with pigs in adjacent pens due to the lower θ , and consequently, a substantial number of
309 contacts may occur with pens that are already infected leading to slower disease transmission
310 overall. Relatedly, Kong et al. (2016) compared heterogeneous and homogeneous mixing models
311 and found that when the disease reproductive number is larger than one, in other words, when
312 disease transmission is occurring at any rate, even low levels of heterogeneity resulted in
313 dynamics similar to those predicted by the homogeneous mixing model. Although the results on
314 time to detection analysis (Table 2) are for moderately virulent strains, the model disease state
315 durations were implemented in a flexible framework to simulate both highly virulent and
316 moderately virulent strains.

317 From Table 2 results, we observe that detecting ASF may take significantly longer when
318 predicted with the heterogeneous transmission model, particularly in large barns, i.e., those with
319 4,800 pigs, and when the transmission predominantly occurs via nose-to-nose contact between
320 pigs in adjacent pens. A potential explanation for the longer time to detection in larger barns is
321 that it would take a higher daily mortality (24 pigs out of a 4,800 pig barn) to exceed the 0.005

322 mortality trigger threshold, by which time of likely occurrence, the infection has probably spread
323 to multiple pens.

324 This analysis showed that homogeneous and heterogeneous model outcomes can match, but only
325 under specific parameter-related conditions. That this matching can indeed occur provides
326 evidence that the discrete individual-based approach can be used to approximate the continuous-
327 time Gillespie approach with adequately small time steps. While the Gillespie approach is well
328 understood and traditionally used in modeling the epidemiology of many diseases (Golightly and
329 Gillespie, 2013; Vestergaard and Génois, 2015; Barongo et al., 2016; Hayes et al., 2021), we
330 present here rationale and results supporting the use of the heterogeneous modeling approach
331 which has the added benefits of being overall more malleable, approximates homogeneous
332 spread, and has greater flexibility in the choice of disease state duration distributions based on
333 experimental data.

334 The heterogeneous approach developed here can approximate the approaches used by Guinat et
335 al. (2016) and Faverjon et al. (2021) by setting θ to zero, thereby forcing all between pen
336 transmission to occur due to direct contacts. When their scaling factor is between zero and one,
337 between pen transmission is distance-dependent in the Nielson et al. (2017) approach, which
338 assumes that there are more and/or higher risk transmission pathways closer to the source pen.
339 We explicitly divide between pen transmission into direct and distance independent pathways,
340 whereas in Nielson et al. (2017) the pathways are expressed only in terms of distance. In both
341 approaches, the pigs directly adjacent to the source pen face the highest infection pressure.
342 However, in our approach the non-adjacent pens have the same transmission risk, whereas in the
343 Nielson et al. (2017) approach the transmission risk can modulate, decreasing as the distance
344 from the source pen increases. The Nielson et al. (2017) approach may be more appropriate for

345 transmission risk from pathways like aerosols, which have been shown by (Olesen et al., 2017)
346 to spread ASFV over short distances within a farm.

347 In our evaluation of the heterogeneous model, we have seen that the simulated outcomes are
348 sensitive to changes in the fraction of within-pen transmission (ϕ) and the mean proportion of the
349 between-pen contacts associated with distance independent pathways (θ). This underlines the
350 importance of parameterizing the model using high-quality, detailed experimental or outbreak
351 data or, in the absence of such data, performing a sensitivity analysis for those parameters with
352 substantial uncertainty.

353 The discrepancies observed in some of the scenarios assessed in this study emphasize the need to
354 perform preliminary explorations on the suitability of the relatively simple disease transmission
355 models that assume homogeneous mixing among individuals. The results of this analysis suggest
356 that homogeneous mixing is a reasonable assumption for outbreaks with a high contact rate and
357 when a large proportion of the disease spread is due to distance independent pathways. If,
358 however, the adequate contact rate is low and the disease spread is dependent on whether or not
359 the pigs are in direct contact with each other within the same or in adjacent pens, then the
360 homogeneous model may overestimate how quickly the virus moves through the population. If
361 the results are overestimated, this can have serious consequences for decision making based on
362 the model output. For example, as observed in Table 2, the time to detection of ASFV by
363 mortality triggers was lower under the homogeneous mixing assumption, especially in large
364 barns. Thus, homogeneous and heterogeneous models should be selectively used depending on
365 the objective of the analysis and the limitations at hand. When intervention strategies and disease
366 surveillance options are developed using the most informative models, there is a potential
367 opportunity to have a realized impact on disease control.

368 Much as we focused on nose-to-nose contact, transmission to adjacent pens might also occur via
369 contact through feces, urine depending on drain design, and fluid flow. Although the current
370 formulation can be parametrized to capture the transmission via these mechanisms to some
371 extent, detailed modeling of pen design and fluid flow needs to be addressed in future research.
372 Note that we assumed that all pens were fully stocked and that the population was closed (i.e.,
373 with no pig introduction into or removal from the pen) during the simulated period. Also, factors
374 such as housing structure, stocking density and production type may all influence model
375 predictions.

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468

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478 **Authors' contributions**

479 AS, SM, PJB, KMS, TCB, TG, CJC, CAC, MRC conceived the ideas of the study; AS, SM, PJB,
480 TB conceived the ideas for the analysis; SM, PJB, AS performed the analyses; SM, AS wrote the
481 manuscript; PJB, MRC were major contributors in writing the manuscript and all other authors
482 commented on the manuscript. All authors read and approved the final manuscript.

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497 **Availability of data and materials**

498 Not applicable.

499 **Ethics approval**

500 The study used already published data.

501 **Competing interests**

502 The authors declare that they have no competing interests.

503 **Table 1.** Disease state duration and transmission parameters used for simulating the time to
 504 detect moderately virulent ASFV

Parameter	Distribution details	Value
Latently infected period (Days)	Gamma (shape= 13.299, scale= 0.3384482)	4.501 (95% P.I., 2.417, 7.223)
Infectious period for pigs that recover	Gamma (shape= 55.42012, scale= 0.7950162)	44.06 (95% P.I., 33.23, 56.394)

(Days)

Infectious period for pigs that die due to ASF (Days)	Gamma (shape= 9.632, scale = 0.862)	8.306 (95% P.I., 3.918, 14.314)
Fraction of infected pigs dying due to ASF	Point estimate	0.4
Within pen adequate contact rate β_w per day	Betapert (min= 1.00, most likely= 1.64, max= 2.74)	1.72 (95% C.I., 1.2 - 2.4)
Between pen adequate contact rate β_b per day	Betapert (min= 0.1, most likely= 0.3, max= 0.5)	0.3 (95% C.I., 0.16 - 0.44)

505

506 **Table 2.** Predicted time to detect ASF based on a daily mortality trigger threshold of 5 per 1000

507 pigs under various heterogeneous and homogeneous within barn transmission scenarios. The

508 homogeneous model results are in italics for emphasis and comparison.

Within-Barn Transmission Model Type	Barn layout (total population)	Fraction of distance independent, between-pen transmission (θ)	Mean predicted days to detection post-exposure (95% prediction interval)
Heterogeneous	120 pens (4800 pigs)	0.05	42 (31 - 65)

Heterogeneous	120 pens (4800 pigs)	0.5	33 (27 - 41)
<i>Homogeneous</i>	<i>120 pens (4800 pigs)</i>	<i>NA</i>	<i>27 (23 - 32)</i>
Heterogeneous	30 pens (1200 pigs)	0.05	25 (12 - 35)
Heterogeneous	30 pens (1200 pigs)	0.5	25 (9 - 32)
<i>Homogeneous</i>	<i>30 pens (1200 pigs)</i>	<i>NA</i>	<i>22 (7 - 27)</i>

509

510 **Figure 1.** Comparison of the number of infectious pigs in a 1,200 growing pig barn based on the
511 homogeneous Gillespie algorithm and the heterogeneous transmission model with various
512 fractions of within pen transmission (ϕ) and distance independent between pen transmission ($\theta =$
513 1)

514 **Figure 2.** Comparison of the number of infectious pigs in a 1,200 growing pig barn based on the
515 homogeneous Gillespie algorithm and the heterogeneous transmission model with various
516 fractions of distance independent between pen transmission (θ) among the total between pen
517 transmission.

518

Figures

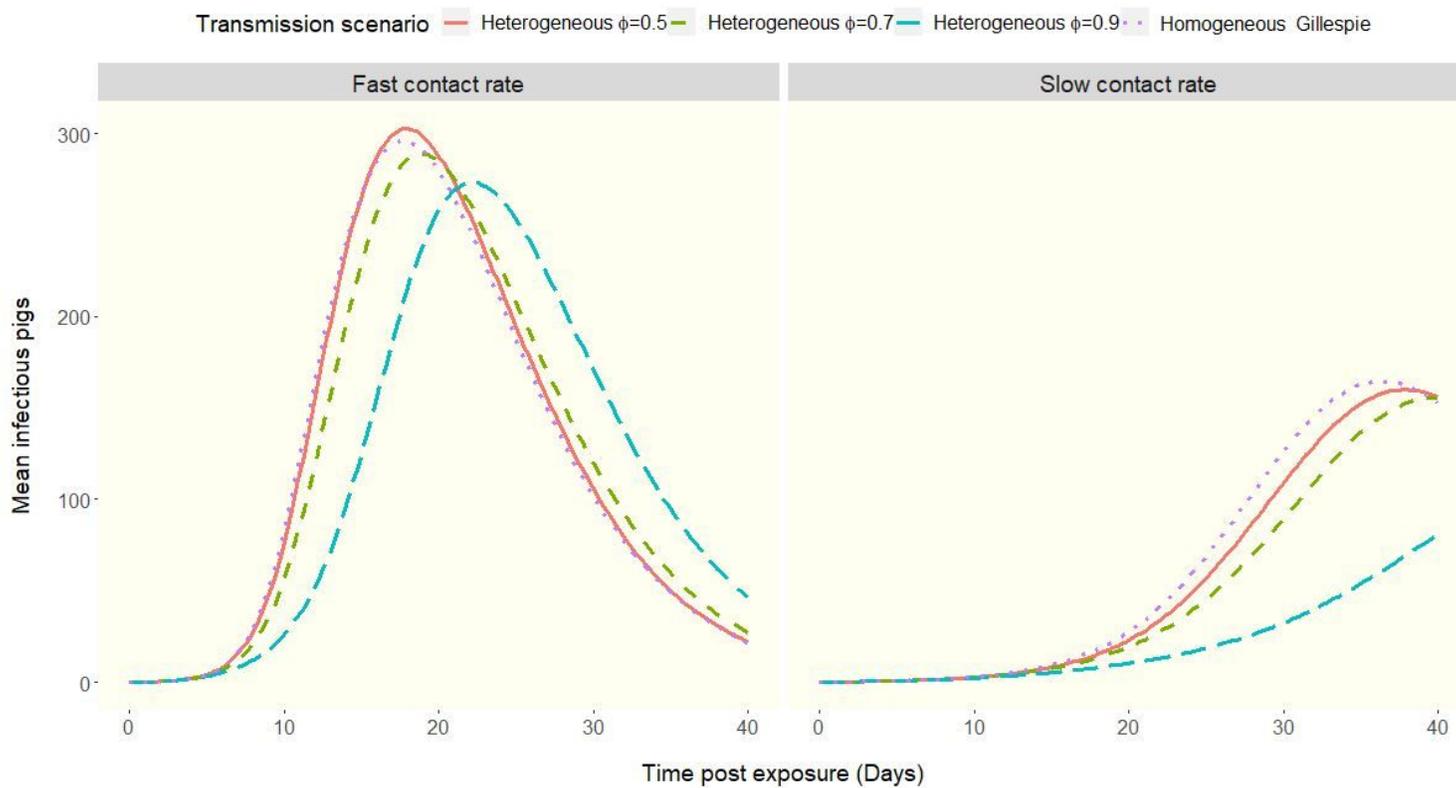


Figure 1

Comparison of the number of infectious pigs in a 1,200 growing pig barn based on the homogeneous Gillespie algorithm and the heterogeneous transmission model with various fractions of within pen transmission (ϕ) and distance independent between pen transmission ($\theta = 1$)

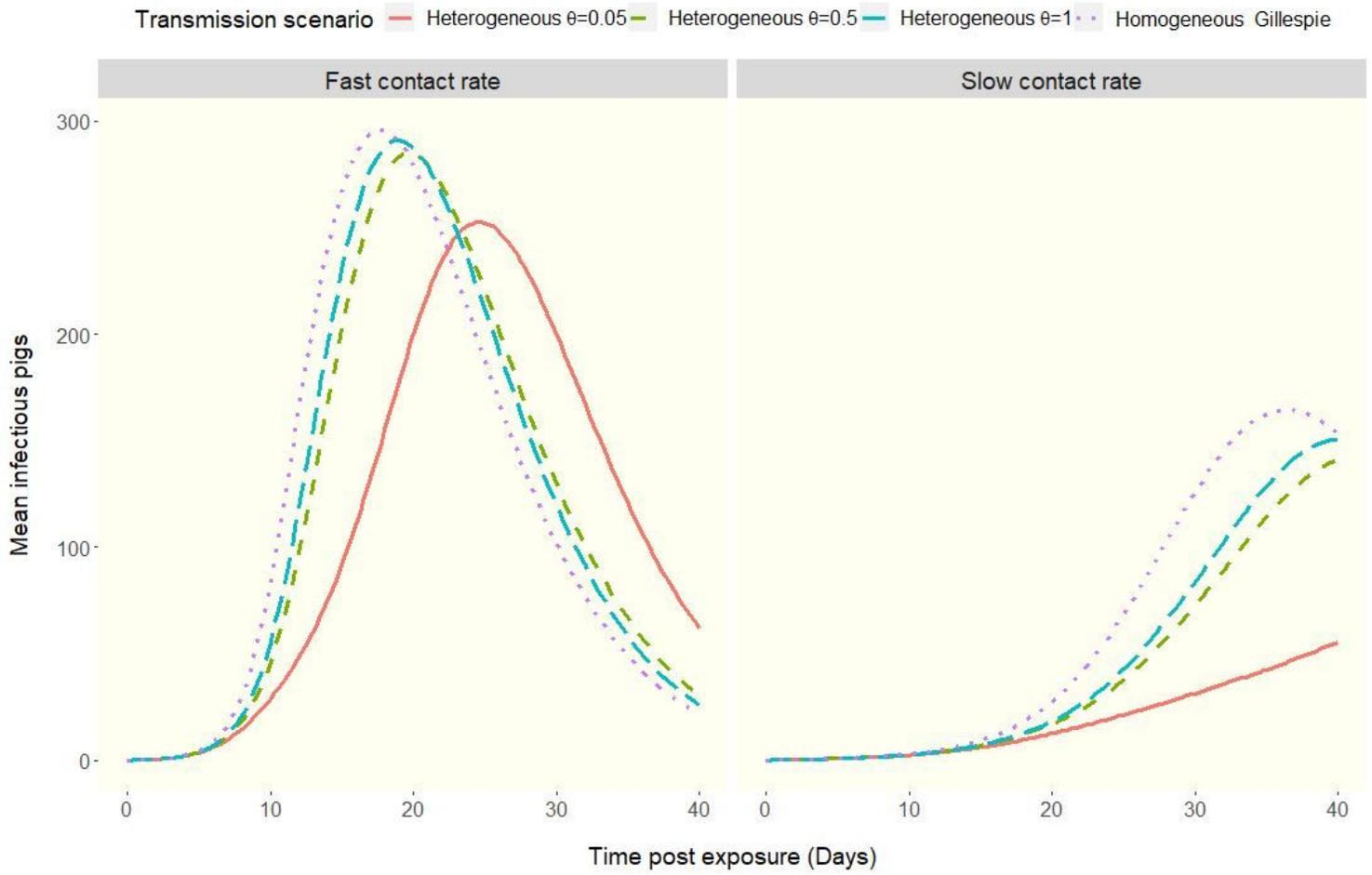


Figure 2

Comparison of the number of infectious pigs in a 1,200 growing pig barn based on the homogeneous Gillespie algorithm and the heterogeneous transmission model with various fractions of distance independent between pen transmission (θ) among the total between pen transmission.