

Appendix for manuscript: The role of the environment in transmission of antimicrobial resistance between humans and animals

Table S1: Parameter definitions

Parameter	Definition and units
β_{HH}	Per capita rate at which humans acquire antibiotic resistant bacteria as a result of exposure to other humans harbouring resistant bacteria per time step
β_{AA}	Per capita rate at which animals acquire antibiotic resistant bacteria as a result of exposure to other animals harbouring resistant bacteria per time step
β_{AH}	Per capita rate at which humans acquire antibiotic resistant bacteria as a result of exposure to animals harbouring resistant bacteria per time step
β_{HA}	Per capita rate at which animals acquire antibiotic resistant bacteria as a result of exposure to humans harbouring resistant bacteria per time step
β_{HE}	Per environmental unit rate* at which the environment acquires resistant bacteria as a result of exposure to humans harbouring resistant bacteria per time step
β_{EH}	Per capita rate at which humans acquire antibiotic resistant bacteria as a result of exposure to environmental units harbouring resistant bacteria per time step
β_{AE}	Per environmental unit rate* at which the environment acquires resistant bacteria as a result of exposure to animals harbouring resistant bacteria per time step
β_{EA}	Per capita rate at which animals acquire antibiotic resistant bacteria as a result of exposure to environmental units harbouring resistant bacteria per time step
Λ_H	Per capita rate at which humans acquire antibiotic resistant bacteria as a result of direct exposure to antibiotics per time step
Λ_A	Per capita rate at which animals acquire antibiotic resistant bacteria as a result of direct exposure to antibiotics per time step
γ_H	Proportion of Λ_H that reaches the environment as antibiotics (a scalar parameter)
γ_A	Proportion of Λ_A that reaches the environment as antibiotics (a scalar parameter)
$\gamma_H \Lambda_H$	Per environmental unit rate* at which the environment acquires resistant bacteria as a result of exposure to a proportion of antibiotics given to humans per time step
μ_H	Per capita rate at which humans with resistant bacteria revert to have only sensitive bacteria per time step
μ_A	Per capita rate at which animals with resistant bacteria revert to have only sensitive bacteria per time step
μ_E	Per environmental unit rate* at which environmental units harbouring resistant bacteria revert to having only sensitive bacteria per time step

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Table S2: Parameter values

Table S2A: Transmission coefficients

Unbounded model

Parameter	Balanced	Balanced (low β_{HA})	Environment- driven	Animal- driven	Human- driven
β_{HH}	0.074	0.074	0.001	0.001	0.202
β_{AA}	0.074	0.074	0.001	0.202	0.001
β_{HA}	0.074	0.0007	0.001	0.001	0.202
β_{AH}	0.074	0.074	0.001	0.202	0.001
β_{EH}	0.074	0.074	0.142	0.001	0.001
β_{EA}	0.074	0.074	0.142	0.001	0.001
β_{HE}	0.074	0.074	0.142	0.001	0.202
β_{AE}	0.074	0.074	0.142	0.202	0.001

Bounded model

Parameter	Balanced	Balanced (low β_{HA})	Environment- driven	Animal- driven	Human- driven
β_{HH}	0.081	0.081	0.001	0.001	0.202
β_{AA}	0.081	0.081	0.001	0.202	0.001
β_{HA}	0.081	0.0008	0.001	0.001	0.202
β_{AH}	0.081	0.081	0.001	0.202	0.001
β_{EH}	0.081	0.081	0.231	0.001	0.001
β_{EA}	0.081	0.081	0.231	0.001	0.001
β_{HE}	0.081	0.081	0.231	0.001	0.202
β_{AE}	0.081	0.081	0.231	0.202	0.001

Table S2B: Other parameters

Parameter	Value	
	Fig 1. B	Fig 2. And 3.
Λ_H	0.1	Beta(1.7, 15.3) (mean 0.1)
Λ_A	0.1	No intervention: 0.1 or U(0.000001,1.); intervention: 0.0.
γ_H	0.001	0.001
γ_A	0.001	0.001
μ_H	0.1	Beta(1.7, 15.3) (mean 0.1)
μ_A	0.1	0.1
μ_E	0.2	Beta(3, 12) (mean 0.2)

Additional methods information

Methods for finding transmission parameter coefficients

Transmission parameters were chosen by the following method: some parameters were fixed (p_f) whilst the transmission parameters of interest varied (p_v) to reach a human resistance level of 54.1% (rate of ampicillin resistance in clinical isolates of *E. coli* in the EU in 2020, based on EARS-Net data¹):

$$\min_{p_v \in (0,1)} |0.71 - f(p_v, p_f)|$$

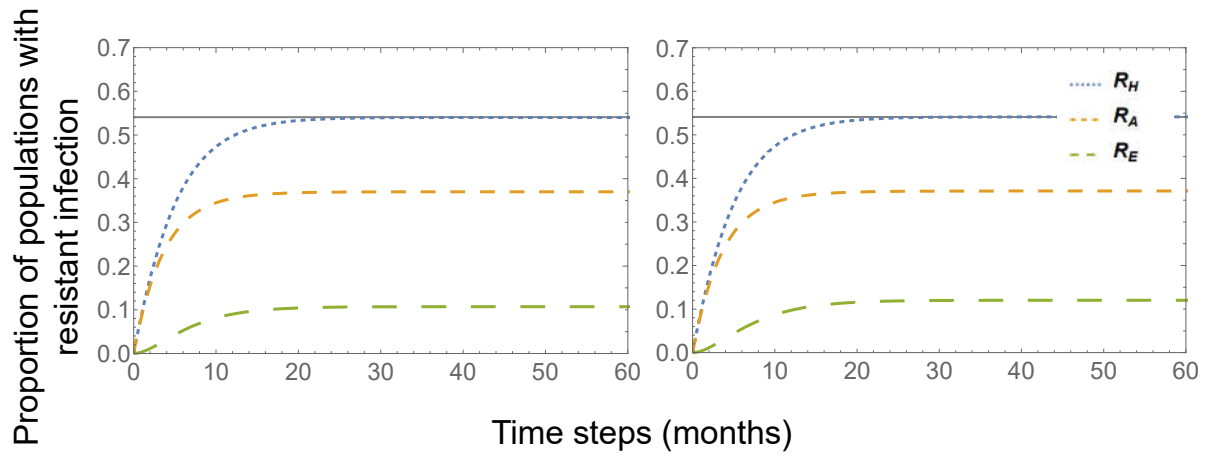
For example, for the human-driven transmission scenario, rates of transfer of resistance from humans to any other population were varied ($\beta_{HH} = \beta_{HA} = \beta_{HE}$), and all other transmission parameter values were fixed at a low value, 0.001.

Model timesteps

To ensure equilibrium values were obtained for all experiments, we initially numerically solved the model to 500 timesteps, and if there was a difference of more than 0.0000001 between the R_H values for the final two timesteps, we solved to 10,000 timesteps.

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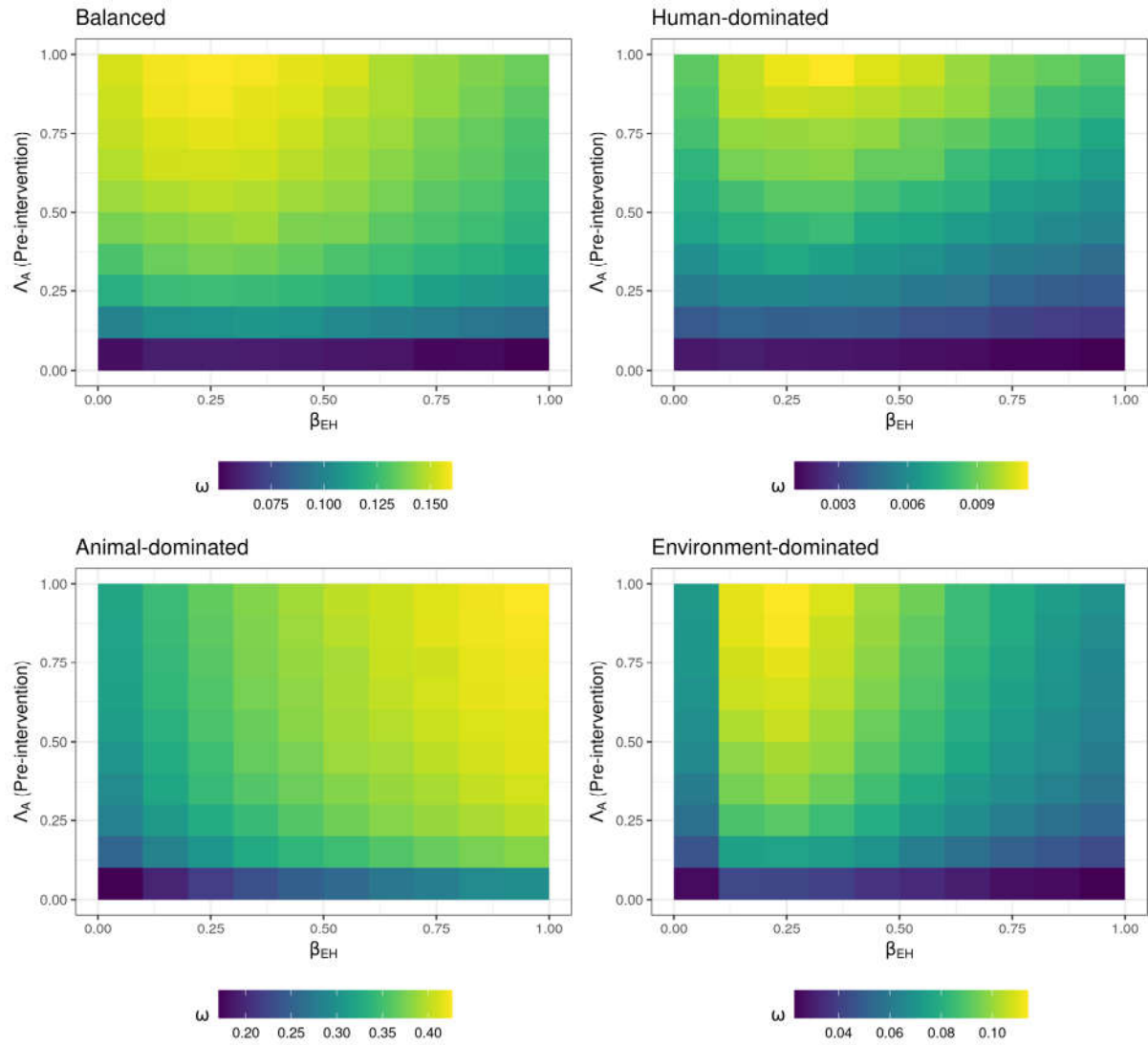
Figure S1: Trajectory plot of the fraction of human and animal populations carrying resistant bacteria (R_H , R_A), and the amount of resistant material in the environment (R_E). For bounded environment model (left) and unbounded (right). Uses the balanced transmission scenario.



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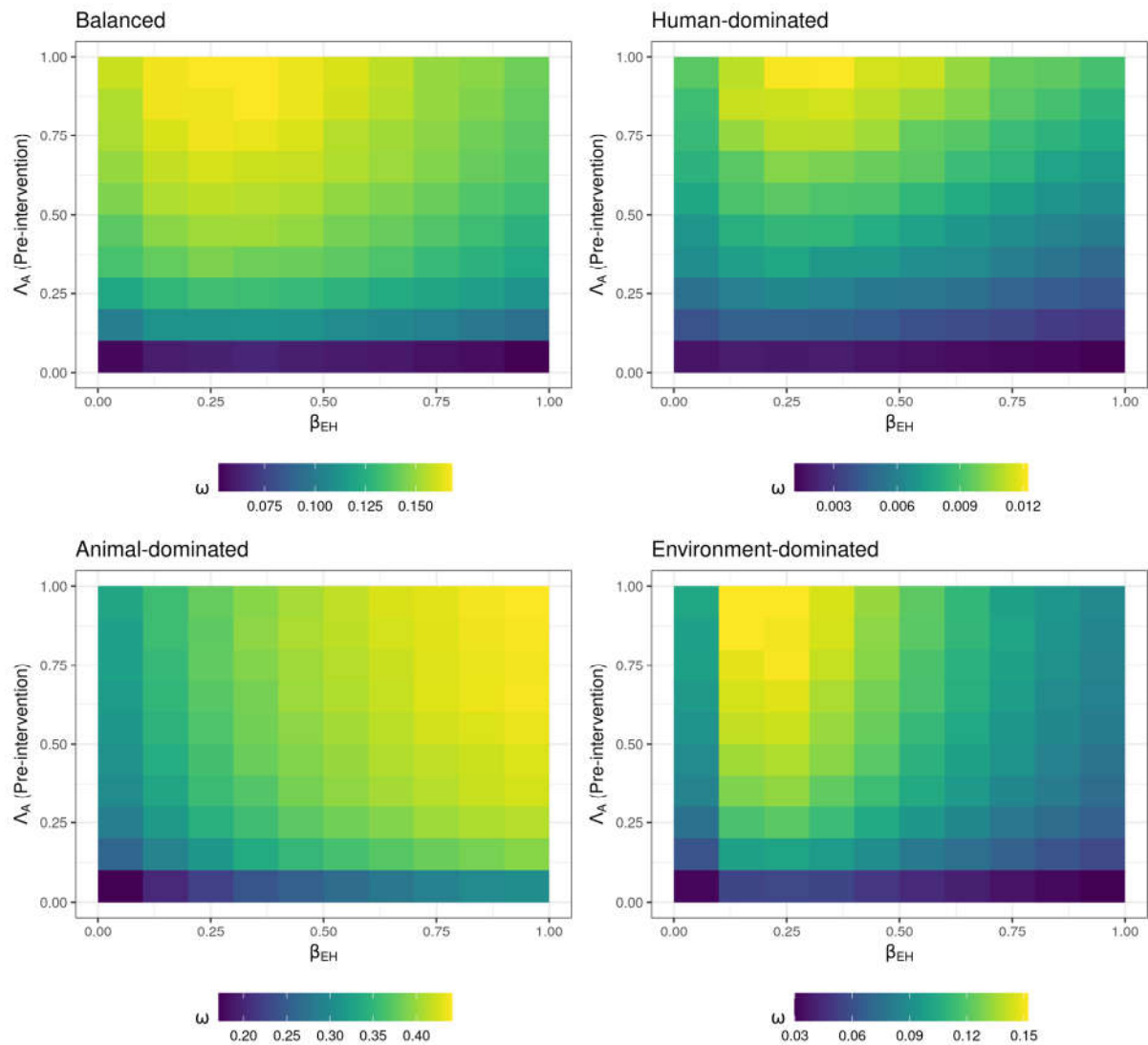
Figure S2: Heatmaps of the impact of reducing Λ_A , for different pre-intervention levels of Λ_A (Y axis) and β_{EH} (X axis), in all transmission scenarios. Plot A) uses the bounded, and plot B) uses the unbounded versions of the model.

A



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B



Appendix references

1. WHO Regional Office for Europe/European Centre for Disease Prevention and Control. *Antimicrobial Resistance Surveillance in Europe 2022 - 2020 Data*. WHO Regional Office for Europe; 2022.