

Figure S1. PRISMA diagram for a systematic review of apramycin resistance in bacteria isolated from animals.

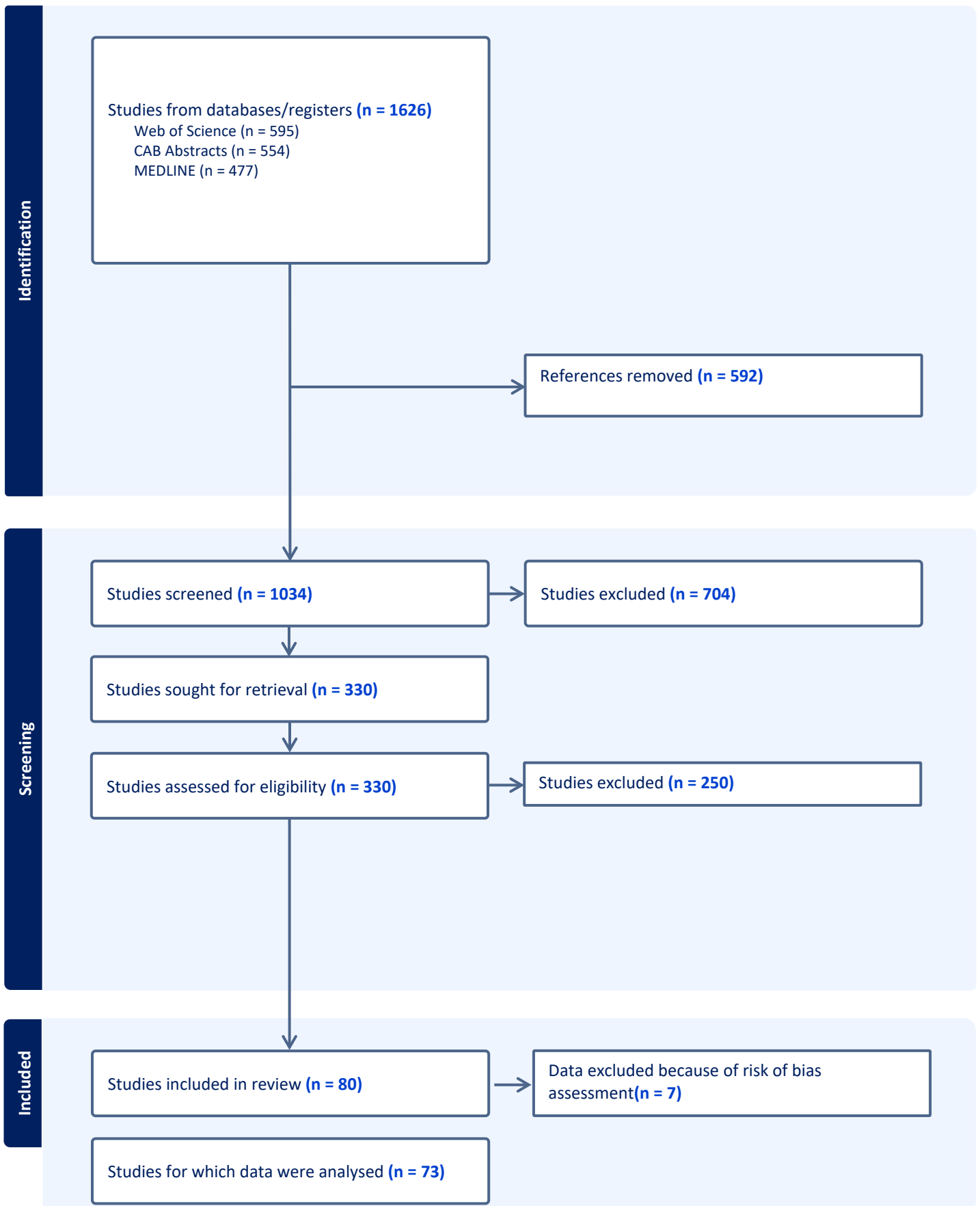


Figure S2. Forest plot of meta-analysis presenting the pooled prevalence and 95% confidence intervals for apramycin resistance in *E. coli* from healthy pigs, poultry, and cattle, by regional subgroup.

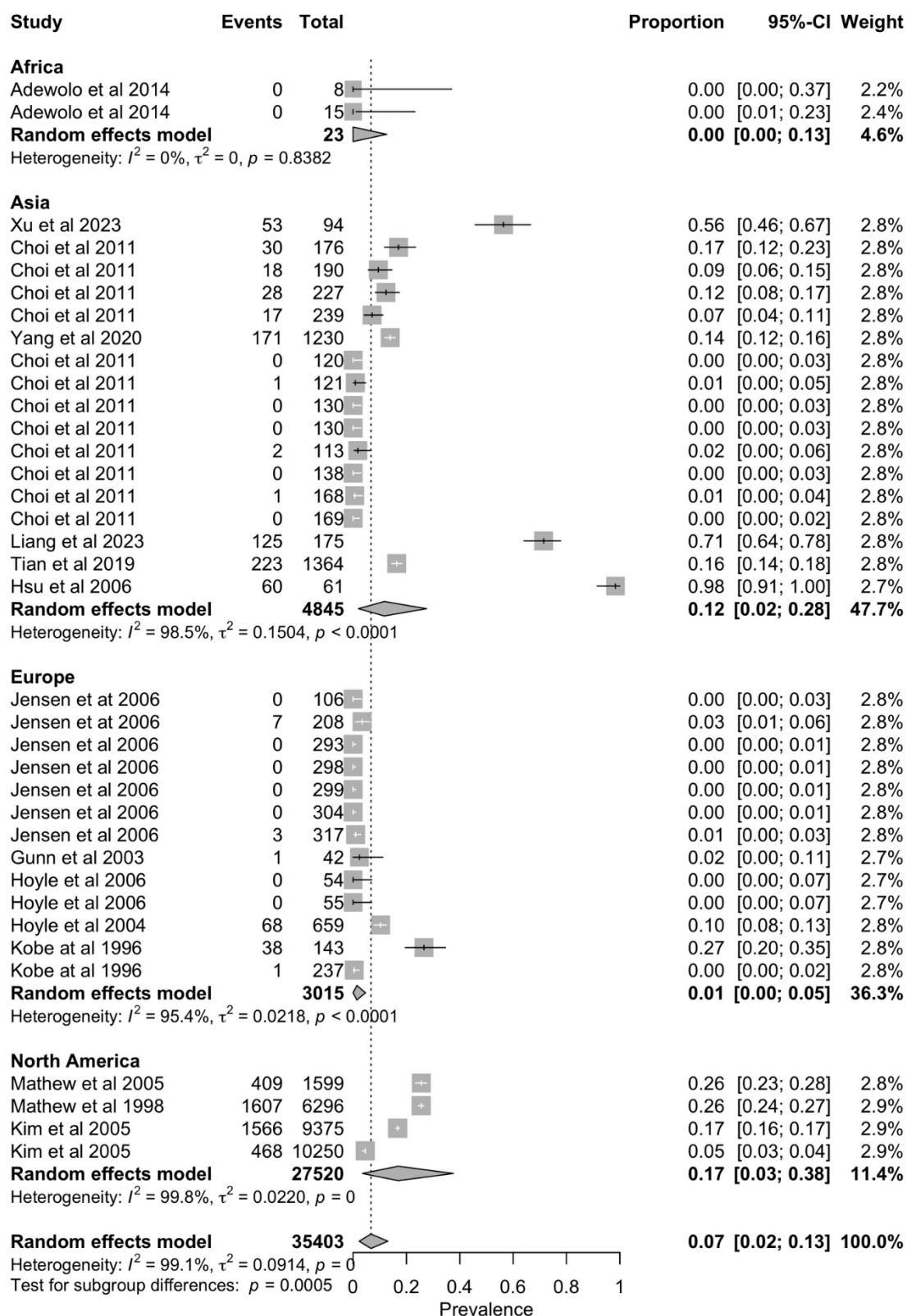


Table S1: Studies that were included in a systematic review of apramycin resistance in bacteria isolated from animals (n=73).

Study (author/year)	Country	Animals	Bacterium
Aarestrup et al 1997(1)	Denmark	Pigs, Cattle, Poultry	<i>Campylobacter</i>
Adewolo et al 2014(2)	Nigeria	Pigs, Poultry	<i>E. coli</i>
Al-Baqir et al 2019(3)	Egypt	Poultry	<i>Salmonella</i>
Amezcuca et al 2002(4)	Canada	Pigs	<i>E. coli</i>
Bischoff et al 2004(5)	US	Cattle	<i>Salmonella</i>
Boerlin et al 2005(6)	Canada	Pigs	<i>E. coli</i>
Borella et al 2020(7)	Italy	Farmed fish	<i>Aeromonas</i> spp
Chao Xi et al 2013(8)	China	Poultry, Dogs, Yaks	<i>Salmonella</i>
Choi et al 2011(9)	Korea	Pigs, Cattle, Poultry	<i>E. coli</i>
da Costa et al 2008(10)	Brazil	Pigs	<i>E. coli</i>
Dipineto et al 2005(11)	Italy	Poultry	<i>Salmonella</i>

Do et al 2006(12)	Vietnam	Pigs	<i>E. coli</i>
Edrington et al 2001(13)	US	Pigs	<i>Salmonella</i>
Edrington et al 2004(14)	US	Cattle	<i>Salmonella</i>
El-Adawy et al 2018(15)	Germany	Poultry	<i>Gallibacterium</i>
Escalante et al 2022(16)	Peru	Pigs	<i>E. coli</i>
Fahmy et al 2023(17)	Egypt	Poultry	<i>Salmonella</i>
Farzan et al 2008(18)	Canada	Pigs	<i>Salmonella</i>
Frank et al 2021(19)	UK	Zoo ungulates	<i>E. coli</i>
Gakuya et al 2001(20)	Kenya	Rats	<i>Aeromonas,</i> <i>Burkholderia</i> <i>cepacia,</i> <i>Chryseomonas, E.</i> <i>coli, Pseudomonas</i>
Garcia-Feliz et al 2008(21)	Spain	Pigs	<i>Salmonella</i>

Giacomelli et al 2014(22)	Italy	Poultry	<i>Campylobacter</i>
Gunn et al 2003(23)	UK	Cattle	<i>E. coli</i>
Gunn et al 2003b(24)	UK	Cattle	<i>E. coli</i>
Hariharan et al 2004(25)	Canada	Pigs	<i>E. coli</i>
Hoyle et al 2004(26)	UK	Cattle	<i>E. coli</i>
Hoyle et al 2006(27)	UK	Cattle	<i>E. coli</i>
Hsu et al 2006(28)	Taiwan	Pigs	<i>E. coli</i>
Intorre et al 2005(29)	Italy	Pigs, Pigeons, Reptiles, Sheep	<i>Salmonella</i>
Jensen et al 2006(30)	Denmark	Pigs	<i>E. coli</i>
Johnson et al 2005(31)	Canada	Pigs, Cattle	<i>Salmonella</i>
Kim et al 2005(32)	US	Pigs	<i>E. coli</i>
Kobe at al 1996(33)	Germany	Poultry	<i>E. coli</i>
Kusumoto et al 2023(34, 35)	Japan	Pigs	<i>E. coli</i>

Lambie et al 2000(35)	Trinidad	Poultry	<i>E. coli</i>
Li et al 2020(36)	China	Poultry	<i>E. coli</i>
Liang et al 2023(37)	China	Poultry	<i>E. coli</i>
Lim et al 2013(38)	Korea	Pigs	<i>Salmonella</i>
Lizarazo et al 2006(39)	Spain	Pigs	<i>Pasteurella multocida</i>
Low et al 1997(40)	UK	Pigs, Cattle, Sheep	<i>Salmonella</i>
Luppi et al 2015(41)	Italy	Pigs	<i>E. coli</i>
Luque et al 2009(42)	Spain	Sheep	<i>Salmonella</i>
Mackie et al 1996(43)	Australia	Cattle	<i>Salmonella</i>
Malgarin et al 2018(44)	Brazil	Pigs	<i>E. coli</i>
Malik et al 2011(45)	US	Pigs	<i>E. coli, Salmonella</i>
Mateu et al 2000(46)	Spain	Pigs	<i>E. coli</i>
Mathew et al 1998(47)	US	Pigs	<i>E. coli</i>
Mathew et al 2005(48)	US	Pigs	<i>E. coli</i>

Moennighoff et al 2020(49)	Germany	Pigs	<i>E. coli</i>
Molla et al 2006(50)	Ethiopia	Pigs	<i>Salmonella</i>
Nadeau et al 2000(51)	Canada	Pigs	<i>E. coli, Salmonella</i>
Oliveira et al 2022 (52)	Portugal	Poultry	<i>E. coli</i>
Orden et al 2000 (53)	Spain	Cattle	<i>E. coli</i>
Pedersen et al 2002 (54)	Denmark	Poultry	<i>Salmonella</i>
Plumb et al 1995 (55)	US	Catfish	<i>Aeromonas spp,</i> <i>Edwardsiella spp,</i> <i>Pseudomonas spp</i>
Radwan et al 2020 (56)	Egypt	Poultry	<i>E. coli</i>
Scullion et al 2010 (57)	UK	Pigeons	<i>E. coli</i>
Sellyei et al 2009 (58)	Hungary	Pigs, Poultry	<i>Pasteurella</i> <i>multocida</i>

Seyfarth et al 1997 (59)	Denmark	Pigs, Cattle, Poultry	<i>Salmonella</i>
Smith et al 2014 (60)	Australia, Vietnam	Pigs	<i>E. coli</i>
Sojka et al 1986 (61)	UK	Cattle	<i>Salmonella</i>
Su Jen et al 1997 (62)	Taiwan	Poultry	<i>E. coli</i>
Sun et al 2012 (63)	China	Poultry	<i>Riemerella anatipestifer</i>
Tian et al 2019 (64)	China	Poultry	<i>E. coli</i>
Varga et al 2018 (65)	Canada	Poultry	<i>E. coli</i>
Wasył et al 2004 (66)	Poland	Pigs, Poultry	<i>Salmonella</i>
White et al 2002 (67)	US	Pigs, Cattle, Poultry	<i>E. coli</i>
Wray et al 1993 (68)	UK	Pigs, Cattle, Poultry, Sheep	<i>Salmonella</i>
Wu et al 1997 (69)	US	Pigs	<i>Mycoplasma hyopneumoniae</i>
Xu et al 2023 (70)	China	Pigs	<i>Pasteurella multocida, Bordetella, Glasserella parasuis,</i>

			<i>Streptococcus suis</i> , <i>E. coli</i>
Yang et al 2020 (71)	China	Pigs	<i>E. coli</i>
Zhang et al 2021 (72)	China	Poultry	<i>E. coli</i>
Zhang et al 2024 (73)	China	Cattle	<i>Proteus mirabilis</i>

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Table S2: Minimum inhibitory concentration (MIC) breakpoints for apramycin in studies included in the systematic review.

MIC breakpoint for resistance	<i>Escherichia coli</i> (n=86)	<i>Salmonella</i> (n=44)
≥ 4 ug/ml	10 (12%)	0
≥ 8 ug/ml	1 (1.2%)	0
≥ 16 ug/ml	3 (3.5%)	2 (4.5%)
≥ 32 ug/ml	18 (21%)	2 (4.5%)
≥ 64 ug/ml	5 (5.8%)	2 (4.5%)
Not specifically reported	46 (53%)	38 (86%)