

Figure S1: Simulated plasma concentration of meropenem and sulbactam. Grey shading indicates 95% CI; 0.95, 95<sup>th</sup> percentile; 0.75, 75<sup>th</sup> percentile; 0.5, 50<sup>th</sup> percentile; 0.25, 25<sup>th</sup> percentile; 0.05, 5<sup>th</sup> percentile; q8h, every 8 hours.

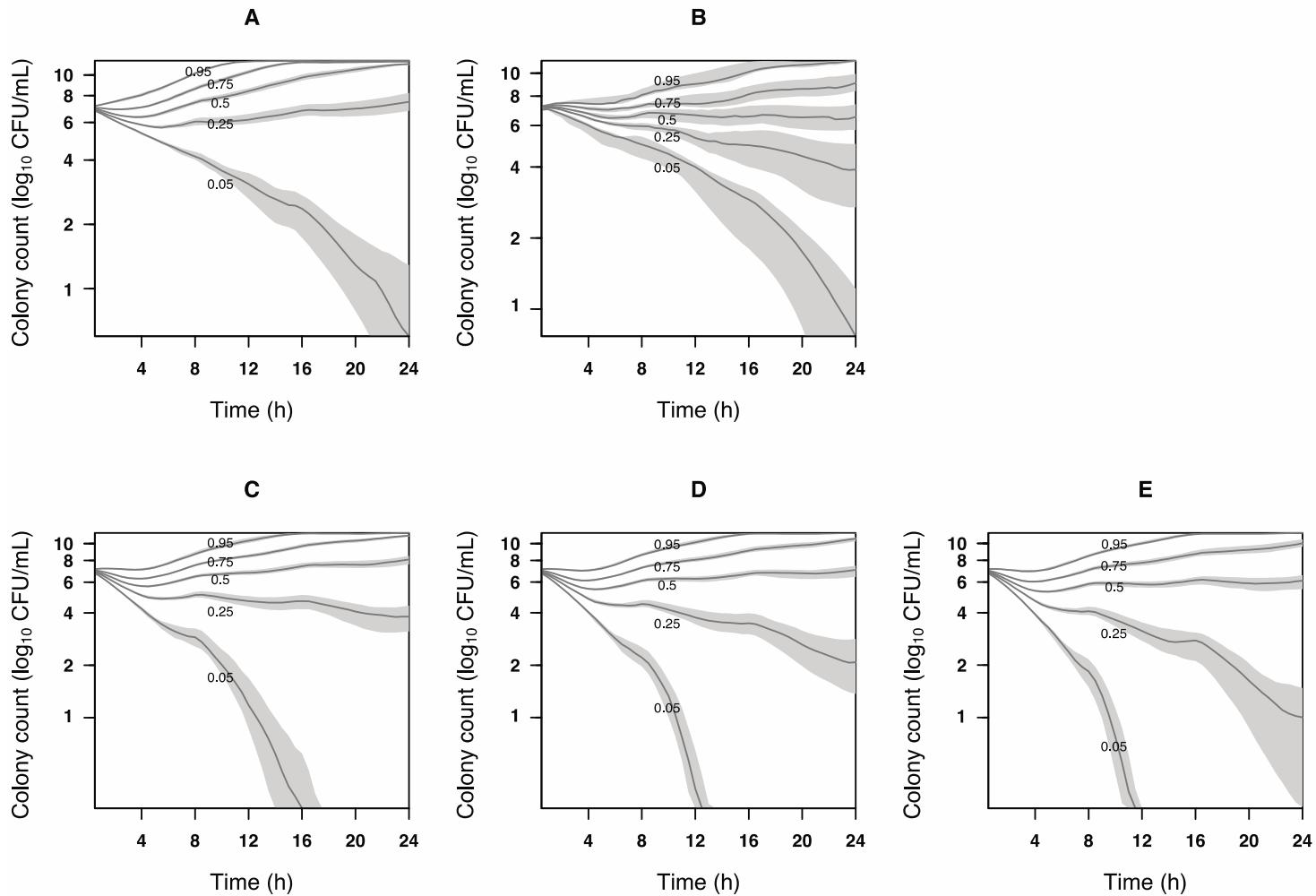


Figure S2: Simulated viable count profiles of carbapenem-resistant *A. baumannii* isolate #79 for monotherapy regimens (A) fosfomycin 8 g q8h; (B) sulbactam 4 g q8h, and combination regimens of sulbactam 2 g q8h with (C) fosfomycin 4 g q8h; (D) fosfomycin 6 g q8h; (E) fosfomycin 8 g q8h. Grey shading indicates 95% CI; 0.95, 95<sup>th</sup> percentile; 0.75, 75<sup>th</sup> percentile; 0.5, 50<sup>th</sup> percentile; 0.25, 25<sup>th</sup> percentile; 0.05, 5<sup>th</sup> percentile; q8h, every 8 hours.

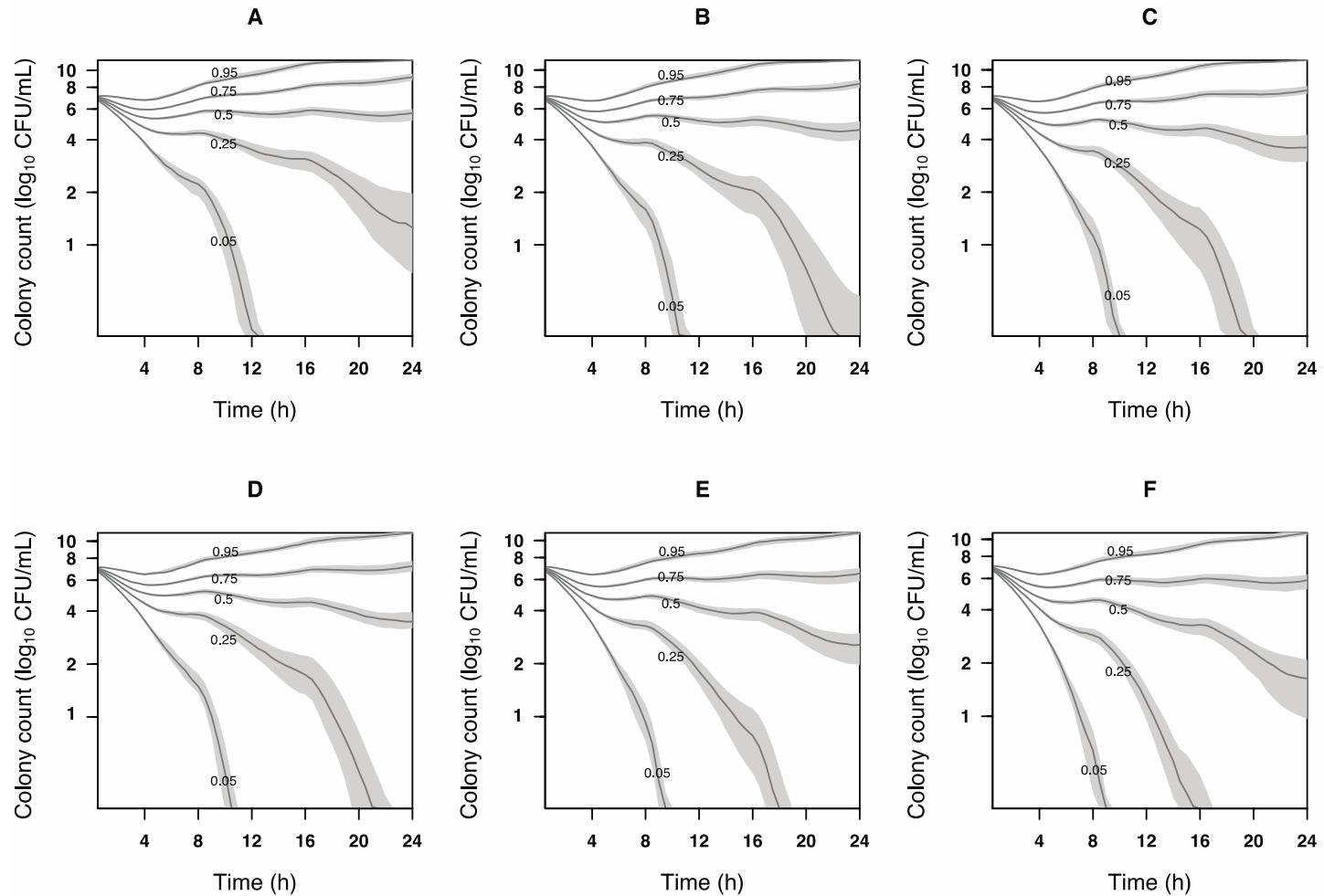


Figure S3: Simulated viable count profiles of carbapenem-resistant *A. baumannii* isolate #79 for combination regimens of sulbactam 3 g q8h with (A) fosfomycin 4 g q8h; (B) fosfomycin 6 g q8h; (C) fosfomycin 8 g q8h, and sulbactam 4 g q8h with (D) fosfomycin 4 g q8h; (E) fosfomycin 6 g q8h; (F) fosfomycin 8 g q8h. Grey shading indicates 95% CI; 0.95, 95<sup>th</sup> percentile; 0.75, 75<sup>th</sup> percentile; 0.5, 50<sup>th</sup> percentile; 0.25, 25<sup>th</sup> percentile; 0.05, 5<sup>th</sup> percentile; q8h, every 8 hours.

Table S1: Population mean parameter estimates for fosfomycin-sulbactam combination models against four carbapenem- resistant *A.baumannii* isolates.

Parameter	#79		#80		#99		#110	
	Mean	SD	Mean	SD	Mean	SD	Mean	SD
K <sub>gs</sub> ( $\log_{10}$ CFU/mL/h)	1.32	0.05	0.79	0.01	1.37	0.57	1.10	0.89
K <sub>gr</sub> ( $\log_{10}$ CFU/mL/h)	1.60	0.33	1.58	0.58	1.72	0.23	1.93	0.07
Bmax ( $\log_{10}$ CFU/mL)	4.76x10 <sup>9</sup>	6.31 x10 <sup>9</sup>	8.55 x10 <sup>9</sup>	8.14 x10 <sup>8</sup>	3.01 x10 <sup>10</sup>	8.41 x10 <sup>9</sup>	6.14 x10 <sup>9</sup>	4.71 x10 <sup>9</sup>
E <sub>maxF</sub> ( $\log_{10}$ CFU/mL/h)	2.33	0.15	1.82	0.77	1.02	0.87	1.19	0.89
E <sub>maxS</sub> ( $\log_{10}$ CFU/mL/h)	2.48	0.35	1.72	0.34	2.60	0.50	1.89	0.86
H <sub>F</sub>	0.78	0.24	0.98	0.47	0.39	0.04	0.86	0.22
EC50 <sub>fs</sub> (mg/L)	6.65	9.26	91.39	55.84	4.27	0.44	82.01	43.61
EC50 <sub>fr</sub> (mg/L)	99.67	36.51	17.21	26.25	127.73	0.15	63.83	20.68
H <sub>S</sub>	1.31	0.49	2.33	1.08	1.47	0.29	2.25	0.86
EC50 <sub>ss</sub> (mg/L)	35.47	12.77	58.98	45.19	36.63	34.68	70.18	48.03
EC50 <sub>sr</sub> (mg/L)	74.39	18.72	81.79	21.02	99.21	4.65	95.99	30.28
INT <sub>fs</sub>	0.89	0.15	-0.40	0.09	-0.53	0.06	-0.13	0.09
HI <sub>fs</sub>	1.17	0.56	1.98	0.60	0.61	0.91	2.66	0.32
EC50 <sub>INTfs</sub> (mg/L)	16.24	21.51	40.91	27.94	4.24	7.25	49.10	30.75
CFU <sub>r</sub> ( $\log_{10}$ CFU/mL)	2.39 x10 <sup>4</sup>	2.52 x10 <sup>4</sup>	3.45 x10 <sup>2</sup>	4.50 x10 <sup>2</sup>	8.51 x10 <sup>3</sup>	4.63 x10 <sup>3</sup>	2.89 x10 <sup>4</sup>	1.83 x10 <sup>4</sup>

SD, standard deviation; CFU<sub>s</sub> and CFU<sub>r</sub> represent the bacterial burden for the sensitive and resistant bacterial subpopulations, respectively; K<sub>gs</sub> and K<sub>gr</sub> represent the growth rate constant for the sensitive and resistant bacterial subpopulations, respectively; Bmax is the maximal bacterial burden; E<sub>maxF</sub> and E<sub>maxS</sub> represent the maximum rate of fosfomycin- and sulbactam-mediated bacterial killing, respectively ( $\log_{10}$  CFU/mL/h); C<sub>F</sub> and C<sub>S</sub> represent the concentration of fosfomycin and sulbactam, respectively; H<sub>F</sub> and H<sub>S</sub> represent the power parameter (Hill coefficient) for fosfomycin and sulbactam effect on both subpopulations, respectively; EC50<sub>fs</sub> and EC50<sub>ss</sub> represent the fosfomycin and sulbactam concentration for which effect is 50% on the sensitive subpopulation, respectively; EC50<sub>fr</sub> and EC50<sub>sr</sub> represent the fosfomycin and sulbactam concentration for which effect is 50% on the resistant subpopulation, respectively; INT<sub>fs</sub> represents the maximum fractional change of the EC50<sub>ss</sub> and EC50<sub>sr</sub> caused by fosfomycin; HI<sub>fs</sub> represents the power parameter (Hill coefficient) for fosfomycin potentiation of sulbactam effect; EC50<sub>INTfs</sub> represents the fosfomycin concentration needed to achieve 50% of INT<sub>fs</sub>.