



The members of the AMR Industry Alliance have developed a unified approach to establishing discharge targets for antibiotic manufacturing, based on Predicted No-Effect Concentrations (PNECs) for use in environmental risk assessments of antibiotics. The discharge target can be derived using these PNECs and site-specific parameters. The publication of the PNEC table fulfills the commitment of the AMR Industry Alliance to publish science-driven, risk-based targets for discharge concentrations of antibiotics. For the first time, the pharmaceutical industry, as member companies of the Alliance, have collated, shared, analyzed, and published existing antibiotic data with respect to antimicrobial resistance and eco-toxicity.

The PNEC table contains two values. PNEC-Environment (PNEC-ENV) values are based on eco-toxicology data generated by Alliance member companies and relevant peer reviewed literature. These values are intended to be protective of ecological species and incorporate assessment factors consistent with standard environmental risk methodologies (Brandt et al., 2015^[1]; Le Page et al., 2017^[2]). The PNEC-Minimum Inhibitory Concentration (PNEC-MIC) values are based on the approach published in Bengtsson-Palme and Larsson (2016)^[3] and are intended to be protective of resistance promotion. This table will be updated periodically as new reliable and robust data become available.

The release of this table is an important step in the journey of evaluating antibiotic discharge concentrations using science-driven, risk-based targets, allowing Alliance member companies to work toward achieving these target values at the receiving water body. The AMR Industry Alliance recommendation is that companies target the lower of these two values (when available) for assessing manufacturing site discharges under a risk-based framework.

If an antibiotic is not listed in the table, read-across to a similar antibiotic based on chemical structure or mode of action can be made.^[4] Alternatively, based on a statistical assessment of all available data, a default PNEC in the absence of both a PNEC-ENV and PNEC-MIC of 0.03 µg/L can be leveraged as a target. When available, a compound specific PNEC-ENV, PNEC-MIC or the lowest of both values should be used. If no data are available, a default PNEC of 0.03 µg/L should be used.

The AMR Industry Alliance believes working toward achieving these antibiotic discharge concentration targets will be both protective of ecological resources and also lower the potential for the evolution and selection of AMR in the environment. It is important to note that these values are recommended based on currently available information, thus, may change as new reliable and relevant information generated to recognized protocols comes to light.

^[1] Brandt, et al., 2015. Ecotoxicological assessment of antibiotics: A call for improved consideration of microorganisms. *Environment International*, 85: 189-205.

^[2] Le Page, et al., 2017. Integrating human and environmental health in antibiotic risk assessment: A critical analysis of protection goals, species sensitivity and antimicrobial resistance. *Environment International*, 109: 155-169.

^[3] Bengtsson-Palme & Larsson, 2016. Concentrations of antibiotics predicted to select for resistant bacteria: Proposed limits for environmental regulation, *Environment International* 86: 140–149.

^[4] Grouping of substances and read-across - ECHA (europa.eu)

AMR Alliance Science-Based PNEC Targets for Risk Assessments

Active Pharmaceutical Ingredient	PNEC-ENV (µg/L)	PNEC-MIC (µg/L)	Lowest Value (µg/L)
Amikacin	N/A	16.00	16.00
Amoxicillin	0.57	0.25	0.25
Ampicillin	0.60	0.25	0.25
Avilamycin	125.00	8.00	8.00
Azithromycin	0.03	0.25	0.03
Aztreonam	N/A	0.50	0.50
Bacitracin	114.59	8.00	8.00
Bedaquiline	0.08	N/A	0.08
Benzylpenicillin	N/A	0.25	0.25
Capreomycin	N/A	2.00	2.00
Cefaclor	N/A	0.50	0.50
Cefadroxil	0.14	2.00	0.14
Cefalonium	21.10	N/A	21.10
Cefaloridine	N/A	4.00	4.00
Cefalothin	N/A	2.00	2.00
Cefazolin	N/A	1.00	1.00
Cefdinir	N/A	0.25	0.25
Cefepime	1.30	0.50	0.50
Cefixime	0.60	0.06	0.06
Cefoperazone	N/A	0.50	0.50
Cefotaxime	0.12	0.13	0.12
Cefoxitin	N/A	8.00	8.00
Cefpirome	N/A	0.06	0.06
Cefpodoxime proxetil	1.76	0.25	0.25
Cefquinome	1.60	N/A	1.60
Ceftaroline	0.12	0.06	0.06
Ceftazidime	1.30	0.50	0.50
Ceftibuten	N/A	0.25	0.25
Ceftiofur	N/A	0.06	0.06
Ceftobiprole	0.23	0.25	0.23
Ceftolozane	1.90	N/A	1.90
Ceftriaxone	0.33	0.03	0.03
Cefuroxime	1.70	0.50	0.50
Cephalexin	0.21	4.00	0.21
Cephradine	0.19	N/A	0.19

Active Pharmaceutical Ingredient	PNEC-ENV (µg/L)	PNEC-MIC (µg/L)	Lowest Value (µg/L)
Chloramphenicol	N/A	8.00	8.00
Chlortetracycline	5.00	N/A	5.00
Ciprofloxacin	0.45	0.06	0.06
Clarithromycin	0.25	0.25	0.25
Clinafloxacin	N/A	0.50	0.50
Clindamycin	0.10	1.00	0.10
Cloxacillin	20.00	0.13	0.13
Colistin (Polymyxin E)	9.00	2.00	2.00
Daptomycin	510.00	1.00	1.00
Delamanid	0.03	N/A	0.03
Doripenem	0.46	0.13	0.13
Doxycycline	25.10	2.00	2.00
Enramycin	4.80	N/A	4.80
Enrofloxacin	1.91	0.06	0.06
Ertapenem	14.00	0.13	0.13
Erythromycin	0.50	1.00	0.50
Ethambutol	N/A	2.00	2.00
Faropenem	N/A	0.02	0.02
Fidaxomicin	891.00	0.02	0.02
Florfenicol	38.00	2.00	2.00
Flucloxacillin	26.80	N/A	26.80
Fluconazole	N/A	0.25	0.25
Flumequine	N/A	0.25	0.25
Fosfomycin	52.40	2.00	2.00
Fusidic acid	N/A	0.50	0.50
Framycetine	N/A	0.06	0.06
Gamithromycin	0.24	N/A	0.24
Gatifloxacin	N/A	0.13	0.13
Gemifloxacin	N/A	0.06	0.06
Gentamicin	0.15	1.00	0.15
Imipenem	0.41	0.13	0.13
Isoniazid	N/A	0.13	0.13
Kanamycin	1.05	2.00	1.05
Levofloxacin	1.52	0.25	0.25
Lincomycin	0.81	2.00	0.81
Linezolid	3.50	8.00	3.50

Active Pharmaceutical Ingredient	PNEC-ENV (µg/L)	PNEC-MIC (µg/L)	Lowest Value (µg/L)
Loracarbef	N/A	2.00	2.00
Mecillinam	N/A	1.00	1.00
Meropenem	1.50	0.06	0.06
Metronidazole	N/A	0.13	0.13
Minocycline	1.10	1.00	1.00
Moxifloxacin	N/A	0.13	0.13
Mupirocin	N/A	0.25	0.25
Nalidixic acid	N/A	16.00	16.00
Narasin	N/A	0.50	0.50
Natamycin	210.00	N/A	210.00
Neomycin	0.03	2.00	0.03
Netilmicin	N/A	0.50	0.50
Nitrofurantoin	N/A	64.00	64.00
Norfloxacin	120.00	0.50	0.50
Ofloxacin	10.00	0.50	0.50
Oxacillin	N/A	1.00	1.00
Oxytetracycline	47.00	0.50	0.50
Pefloxacin	N/A	8.00	8.00
Penicillin G Procaine	16.00	N/A	16.00
Phenoxymethylpenicillin	N/A	0.06	0.06
Piperacillin	4.30	0.50	0.50
Polymyxin B	0.06	N/A	0.06
Pristinamycin	71.10	N/A	71.10
Puromycin	31.00	N/A	31.00
Retapamulin	N/A	0.06	0.06
Rifampicin	4.06	0.06	0.06
Rifamycin	1.00	N/A	1.00
Rifaximin	0.11	N/A	0.11
Roxithromycin	6.80	1.00	1.00
Secnidazole	N/A	1.00	1.00
Sparfloxacin	N/A	0.06	0.06
Spectinomycin	N/A	32.00	32.00
Spiramycin	1.09	0.50	0.50
Streptomycin	N/A	16.00	16.00
Sulfadiazine	11.21	13.00	11.21
Sulfamethoxazole	0.60	16.00	0.60
Tedizolid	3.20	N/A	3.20

Active Pharmaceutical Ingredient	PNEC-ENV (µg/L)	PNEC-MIC (µg/L)	Lowest Value (µg/L)
Teicoplanin	12.90	0.50	0.50
Telithromycin	N/A	0.06	0.06
Tetracycline	3.20	1.00	1.00
Thiamphenicol	N/A	1.00	1.00
Tiamulin	N/A	1.00	1.00
Ticarcillin	N/A	8.00	8.00
Tigecycline	0.10	1.00	0.10
Tildipirosin	0.42	N/A	0.42
Tilmicosin	0.80	1.00	0.80
Tobramycin	4.30	1.00	1.00
Trimethoprim	312.45	0.50	0.50
Trovafloxacin	N/A	0.03	0.03
Tulathromycin	0.04	N/A	0.04
Tylosin	0.98	4.00	0.98
Vancomycin	N/A	8.00	8.00
Viomycin	N/A	2.00	2.00
Virginiamycin	N/A	2.00	2.00