

# Agreement of the MIC Test Strip versus Etest in MIC determination of *Streptococcus pneumoniae*

Bjørg C. Haldorsen<sup>1</sup>, Anne-Sofie Furberg<sup>2</sup>, Ørjan Samuelsen<sup>1</sup>, Didrik F. Vestheim<sup>3</sup>, Martin Steinbakk<sup>3</sup>, and Arnfinn Sundsfjord<sup>1,4</sup>.

<sup>1</sup>Reference Centre for Detection of Antimicrobial Resistance, Department of Microbiology and Infection Control, University Hospital of North Norway, Tromsø, Norway. <sup>2</sup>Department of Community Medicine, Faculty of Health Sciences, University of Tromsø, Tromsø, Norway. <sup>3</sup>Department of Bacteriology and Immunology, Div. Infect. Dis. Control, Norwegian Institute of Public Health, Oslo, Norway. <sup>4</sup>Research Group for Host-Microbe Interactions, Department of Medical Biology, Faculty of Health Sciences, University of Tromsø, Tromsø, Norway.

## Objective

We have examined the agreement between MIC-values obtained by the MIC Test Strip (Liofilchem, Roseto degli Abruzzi, Italy) and the Etest (bioMérieux, Marcy l'Etoile, France) for commonly used antibiotics using a well characterized collection of *Streptococcus pneumoniae* strains.

## Bacterial strains

A collection of *S. pneumoniae* strains ( $n=93$ ) from the National Institute of Public Health, Norway, which included: wild type (susceptible) strains ( $n=32$ ), penicillin non-susceptible pneumococci (PNSP) ( $n=31$ ), and macrolide resistant strains ( $n=30$ ), was used. The *S. pneumoniae* ATCC 49619 reference strain was also included.

## Methods

A total of nine antibiotics were tested: penicillin G (PG), ampicillin (AM), cefotaxime (CT), meropenem (MP), ciprofloxacin (Cl), erythromycin (EM), clindamycin (CM), tetracycline (TC), and trimethoprim-sulphamethoxazole (TS).

A 0.5 McFarland bacterial suspension in Mueller-Hinton (MH) broth (BBL, Becton, Dickinson and Company, LePont de Claix, France) were inoculated on MH agar (Oxoid, Basingstoke, Hampshire, UK) with 5% defibrinated horseblood (TCS Biosciences Ltd, Botolph Claydon Buckingham, UK) and incubated for 20-24 hours at  $35 \pm 2$  °C in 5% CO<sub>2</sub>.

## Evaluation of results

The numbers of strains with a MIC Test Strip MIC-value > 2 dilution steps, 2 dilution steps, or 1 dilution step difference from, or 100% agreement with that obtained by the Etest were recorded. The SIR (S-susceptible, I-intermediate, or R-resistant) categorization was compared for the strains with > 2 or 2 dilution steps disagreement, using clinical breakpoints as defined by EUCAST.

Differences in MIC Test Strip MIC-values compared to that obtained by Etest were categorized as very major (R → S), major (S → R), or minor errors (S ↔ I, I ↔ R) if the MIC Test Strip MIC resulted in a change in clinical categorization (SIR).

## Results

The overall agreement between MIC-values for MIC Test strips compared to Etest, is presented in Table 1 and Figure 1.

A > 2 dilution steps disagreement between MIC Test Strips and Etest was found for three antibiotics (PG, CT, and TS) in three different strains. This did not change the SIR-categorization for two of the strains, while one minor error (S → I) were observed for PG in one strain.

A 2 step divergence was found in altogether 47 observations for seven antibiotics (PG, AM, CT, MP, Cl, CM, and TS). This did not cause any change in SIR-categorization in 43 of the observations. In one of the strains two minor errors; R → I ( $n=1$ ) and S → I ( $n=1$ ) for AM and MP, were observed, while one minor error S → I ( $n=1$ ) and I → R ( $n=1$ ) was observed in two strains for MP and Cl, respectively.

The numbers of errors are listed in Table 2.

Table 1. The overall agreement in absolute numbers and percentage between MIC-values obtained by the MIC Test Strips compared to Etest.

	PG	AM	CT	MP	Cl	EM	CM	TC	TS
	n= %								
100 % agreement	41 44	41 44	30 32	34 37	5 5	53 57	31 34	59 64	6 6
1 dilution step	49 53	51 55	57 62	55 59	64 70	40 43	57 61	34 36	80 87
2 dilution steps	2 2	1 1	5 5	4 4	24 26	0 0	5 5	0 0	6 6
>2 dilution steps	1 1	0 0	1 1	0 0	0 0	0 0	0 0	0 0	1 1

PG: penicillin G; AM: ampicillin; CT: cefotaxime; MP: meropenem; Cl: ciprofloxacin; EM: erythromycin; CM: clindamycin; TC: tetracycline; and TS: trimethoprim-sulphamethoxazole.

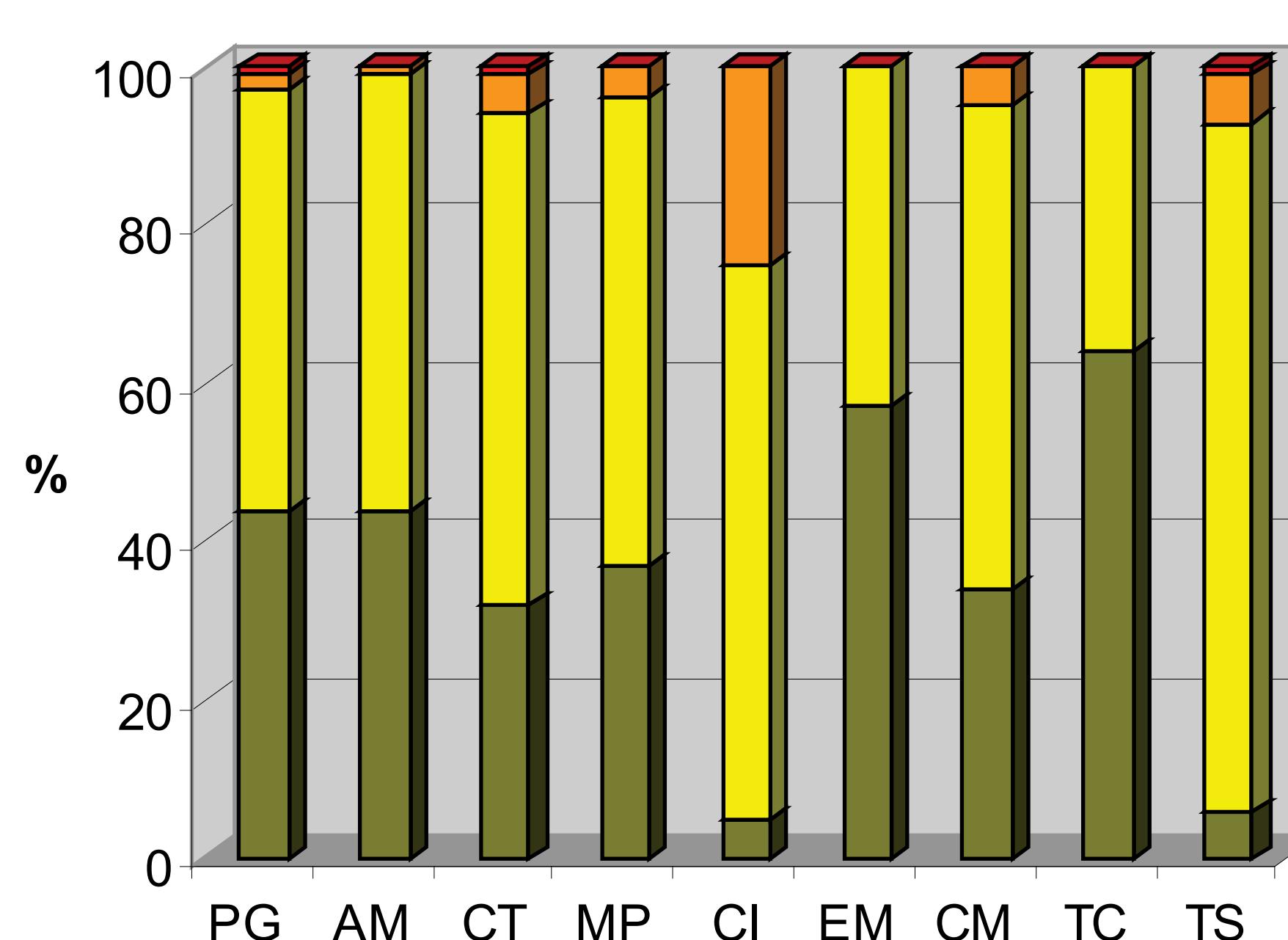


Figure 1. The overall agreement in percentage between MIC-values for the MIC Test Strip compared to the Etest. PG: penicillin G; AM: ampicillin; CT: cefotaxime; MP: meropenem; Cl: ciprofloxacin; EM: erythromycin; CM: clindamycin; TC: tetracycline; and TS: trimethoprim-sulphamethoxazole.

Table 2. The numbers of ≥ 2 and 2 dilution steps disagreement between MIC Test Strip and Etest for the different antibiotics tested, and their respective categorization of error.

	PG	AM	CT	MP	Cl	EM	CM	TC	TS
>2 dilution steps disagreement, n	1	0	1	0	0	0	0	0	1
No error, n	0	0	1	0	0	0	0	0	1
Minor error, n	1	0	0	0	0	0	0	0	0
2 dilution steps disagreement, n	2	1	5	4	24	0	5	0	6
No error, n	2	0	5	2	23	0	5	0	6
Minor error, n	0	1	0	2	1	0	0	0	0

PG: penicillin G; AM: ampicillin; CT: cefotaxime; MP: meropenem; Cl: ciprofloxacin; EM: erythromycin; CM: clindamycin; TC: tetracycline; and TS: trimethoprim-sulphamethoxazole.

## Conclusions

Overall, we observed good agreement between the two gradient methods. In total only 5 minor errors in SIR-categorization were observed, with respect to observations with > 2 and 2 dilution steps disagreement.

### Acknowledgements

We are grateful to Anne Ramstad Alme, Gunnhild Rødal, Bettina Aasnæs, and Runa Wolden for excellent technical assistance.

### Conflict of interests

The MIC Test and Etest strips were kindly donated by Montebello Diagnostics, Oslo, Norway.