

Supplemental

Table S1. Characterization of the clinical isolates from hospitalized patients in Germany (RKI strain collection).

RKI ID	Species	MIC [$\mu\text{g/mL}$] ^{a, b}											
		AMP	MSU	CTX	CAZ	GEN	KAN	AMK	OTE	CIP	SXT	MEM	CST
19/09	<i>A. baumannii</i>	>16	>32	>16	>32	>8	>32	>32	>8	>64	>128	>64	0.5
523/16	<i>A. baumannii</i>	>16	>32	>16	>32	>8	≤2	≤2	>8	>64	>128	32	≤0.5
584/16	<i>A. baumannii</i>	>16	≤2	16	4	1	4	4	8	0.5	8	≤0.5	≤0.5
596/16	<i>A. baumannii</i>	>16	≤2	>16	>32	>8	>32	>32	>8	>64	>128	>64	>64
271/16	<i>A. baumannii</i>	>16	>32	>16	>32	>8	>32	>32	>8	>64	>128	>64	≤0.5
272/16	<i>A. baumannii</i>	>16	>32	>16	>32	>8	>32	>32	>8	>64	>128	>64	≤0.5
30/16	<i>A. baumannii</i>	>16	>32	>16	>32	>8	>32	>32	>8	>64	>128	>64	1
70/16	<i>A. baumannii</i>	>16	32	>16	>32	>8	>32	>32	>8	>64	>128	>64	64
32/16	<i>A. baumannii</i>	>16	>32	>16	8	>8	>32	>32	4	64	>128	64	2
11/16	<i>A. baumannii</i>	>16	>32	>16	>32	8	>32	>32	>8	>64	>128	>64	1
12/16	<i>A. baumannii</i>	>16	>32	>16	>32	>8	>32	>32	>8	>64	>128	64	≤0.5
13/16	<i>A. baumannii</i>	>16	>32	>16	>32	>8	>32	>32	>8	>64	>128	>64	≤0.5
14/16	<i>A. baumannii</i>	>16	≤2	>16	>32	>8	4	8	>8	>64	>128	>64	≤0.5
16/16	<i>A. baumannii</i>	>16	>32	>16	>32	>8	>32	>32	>8	>64	>128	>64	64
230/15	<i>A. baumannii</i>	>16	>32	>16	>32	>8	>32	>32	>8	>64	>128	>64	≤0.5
231/15	<i>A. baumannii</i>	>16	>32	>16	>32	>8	>32	>32	>8	>64	>128	64	≤0.5
377/15	<i>A. baumannii</i>	>16	>32	>16	>32	2	>32	>32	>8	>64	>128	8	1
3/15	<i>A. baumannii</i>	R	nd.	R	R	S	nd.	S	nd.	R	R	32	≤0.5
4/15	<i>A. baumannii</i>	R	nd.	R	R	S	nd.	S	nd.	R	R	32	≤0.5
16/15	<i>A. baumannii</i>	R	nd.	R	R	R	nd.	S	nd.	R	R	>64	≤0.5
588/16	<i>A. pittii</i>	>16	≤2	16	8	≤0.5	≤2	≤2	2	0.5	8	≤0.5	≤0.5
146/09	<i>E. cloacae</i>	>16	>32	>16	>32	2	32	≤2	4	≤0.063	>128	16	2
33/16	<i>E. cloacae</i>	>16	32	16	4	≤0.5	≤2	≤2	4	0.25	>128	8	≤0.5
34/16	<i>E. cloacae</i>	>16	>32	>16	>32	4	>32	16	8	32	>128	32	≤0.5
113/16	<i>E. cloacae</i>	>16	>32	>16	>32	1	4	4	8	4	8	2	≤0.5
460/16	<i>E. cloacae</i>	>16	>32	>16	>32	>8	>32	32	2	>64	>128	64	≤0.5
136/15	<i>E. cloacae</i>	>16	>32	>16	>32	4	>32	4	8	8	64	>64	≤0.5
141/15	<i>E. cloacae</i>	>16	32	>16	>32	≤0.5	≤2	≤2	4	0.125	≤4	2	4
210/15	<i>E. cloacae</i>	>16	32	>16	4	2	8	8	2	≤0.063	≤4	≤0.5	1
243/15	<i>E. cloacae</i>	>16	>32	>16	>32	1	4	≤2	2	≤0.063	≤4	≤0.5	4
348/15	<i>E. cloacae</i>	>16	>32	>16	>32	8	>32	4	8	64	>128	1	4
506/15	<i>E. cloacae</i>	>16	>32	>16	>32	2	>32	16	>8	≤0.063	>128	16	1
766/15	<i>E. cloacae</i>	>16	>32	>16	>32	1	4	4	4	≤0.063	≤4	≤0.5	≤0.5
76/15	<i>E. cloacae</i>	>16	>32	>16	>32	≤0.5	32	4	>8	8	>128	≤0.5	≤0.5

5/14	<i>E. cloacae</i>	> 16	> 32	> 16	> 32	2	> 32	8	> 8	> 64	> 128	4	≤ 0.5
37/14	<i>E. cloacae</i>	> 16	4	16	> 32	4	32	≤ 2	8	0.25	> 128	≤ 0.5	≤ 0.5
733/14	<i>E. cloacae</i>	> 16	> 32	> 16	> 32	> 8	> 32	> 32	> 8	> 64	> 128	64	≤ 0.5
287/13	<i>E. cloacae</i>	> 16	4	> 16	≤ 2	1	4	≤ 2	2	≤ 0.063	≤ 4	64	> 64
308/13	<i>E. cloacae</i>	> 16	4	16	> 32	2	32	≤ 2	4	0.5	> 128	8	≤ 0.5
361/13	<i>E. cloacae</i>	> 16	> 32	> 16	> 32	4	32	≤ 2	> 8	0.5	> 128	≤ 0.5	≤ 0.5
466/13	<i>E. cloacae</i>	> 16	16	> 16	> 32	4	> 32	≤ 2	4	4	> 128	16	≤ 0.5
546/13	<i>E. cloacae</i>	> 16	> 32	16	≤ 2	4	32	≤ 2	> 8	2	> 128	≤ 0.5	2
571/13	<i>E. cloacae</i>	> 16	> 32	> 16	> 32	4	16	8	8	≤ 0.063	≤ 4	≤ 0.5	≤ 0.5
RKI ID	Species	AMP	MSU	CTX	CAZ	GEN	KAN	AMK	OTE	CIP	SXT	MEM	CST
215/15	<i>E. cloacae</i>	16	8	≤ 1	≤ 2	1	4	4	2	≤ 0.063	16	≤ 0.5	≤ 0.5
1026/14-1	<i>E. aerogenes</i>	> 16	> 32	> 16	> 32	1	4	4	4	0.5	128	16	8
131/08	<i>E. coli</i>	> 16	32	≤ 1 (2)	≤ 2	> 8	4	4	> 8	≤ 0.063	> 128	32	0.5
6A-6	<i>E. coli</i>	> 16	8	8	16	1	8	8	4	≤ 0.063	≤ 4	≤ 0.5	8
602/16	<i>E. coli</i>	≤ 4	nd.	≤ 1	≤ 2	≤ 0.5	nd.	≤ 2	nd.	≤ 0.063	≤ 4	≤ 0.5	16
521/16	<i>E. coli</i>	> 16	nd.	≤ 1	≤ 2	> 8	32	≤ 2	nd.	> 2	≤ 4	≤ 0.5	8
314/16	<i>E. coli</i>	> 16	4	≤ 1	≤ 2	2	4	8	4	≤ 0.063	≤ 4	≤ 0.5	8
317/16	<i>E. coli</i>	> 16	16	≤ 1	≤ 2	2	8	8	8	1	≤ 4	≤ 0.5	4
330/16	<i>E. coli</i>	> 16	16	≤ 1	≤ 2	2	> 32	4	> 8	> 64	> 128	≤ 0.5	8
524/16	<i>E. coli</i>	> 16	nd.	> 16	> 32	≤ 0.5	nd.	≤ 2	nd.	> 2	≤ 4	≤ 0.5	8
525/16	<i>E. coli</i>	≤ 4	nd.	≤ 1	≤ 2	≤ 0.5	nd.	≤ 2	nd.	≤ 0.063	≤ 4	≤ 0.5	8
462/16	<i>E. coli</i>	> 16	> 32	≤ 1	≤ 2	1	4	≤ 2	> 8	32	8	≤ 0.5	8
443/16	<i>E. coli</i>	> 16	> 32	> 32	> 32	4	8	8	> 8	> 64	> 128	4	4
581/16	<i>E. coli</i>	> 16	32	4	8	1	4	4	nd.	1	32	≤ 0.5	≤ 0.5
582/16	<i>E. coli</i>	> 16	32	≤ 1	4	1	4	4	nd.	≤ 0.063	32	≤ 0.5	≤ 0.5
583/16	<i>E. coli</i>	> 16	32	> 16	> 32	> 8	> 32	8	nd.	> 64	> 128	≤ 0.5	≤ 0.5
590/16	<i>E. coli</i>	> 16	> 32	> 16	> 32	> 8	8	4	4	0.5	> 128	≤ 0.5	≤ 0.5
592/16	<i>E. coli</i>	> 16	> 32	> 16	≤ 2	1	≤ 2	4	> 8	≤ 0.063	8	≤ 0.5	≤ 0.5
593/16	<i>E. coli</i>	> 16	> 32	4	4	2	4	4	4	≤ 0.063	16	≤ 0.5	≤ 0.5
607/16	<i>E. coli</i>	> 16	> 32	> 16	16	> 8	> 32	4	> 8	2	> 128	8	≤ 0.5
513/16	<i>E. coli</i>	> 16	32	2	8	2	8	8	1	≤ 0.063	> 128	≤ 0.5	≤ 0.5
514/16	<i>E. coli</i>	> 16	8	16	> 32	2	8	4	4	1	32	≤ 0.5	≤ 0.5
515/16	<i>E. coli</i>	> 16	32	8	8	2	8	4	2	64	≤ 4	≤ 0.5	≤ 0.5
95/15	<i>E. coli</i>	> 16	> 32	> 16	32	1	> 32	32	2	≤ 0.063	≤ 4	≤ 0.5	≤ 0.5
97/15	<i>E. coli</i>	> 16	> 32	> 16	> 32	> 8	> 32	32	> 8	2	≤ 4	≤ 0.5	≤ 0.5
75/15	<i>E. coli</i>	R	nd.	R	R	R	nd.	R	nd.	R	R	> 64	≤ 0.5
1023/14	<i>E. coli</i>	> 16	≤ 2	> 16	8	1	8	8	2	64	> 128	≤ 0.5	≤ 0.5
867/14	<i>E. coli</i>	> 16	> 32	> 16	> 32	≤ 0.5	≤ 2	≤ 2	1	> 64	8	8	≤ 0.5
93/10	<i>K. pneumoniae</i>	> 16	> 32	> 16	> 32	> 8	> 32	> 32	> 8	> 64	> 128	64	1
600/16	<i>K. pneumoniae</i>	8	nd.	> 16	> 32	≤ 0.5	nd.	≤ 2	nd.	> 2	≤ 4	≤ 0.5	32
393/16	<i>K. pneumoniae</i>	8	nd.	> 16	> 32	≤ 0.5	nd.	≤ 2	nd.	> 2	≤ 4	≤ 0.5	64
373/16-1	<i>K. pneumoniae</i>	16	nd.	> 16	> 32	≤ 0.5	nd.	≤ 2	nd.	> 2	≤ 4	≤ 0.5	32
320/16	<i>K. pneumoniae</i>	≤ 4	nd.	≤ 1	≤ 2	≤ 0.5	nd.	≤ 2	nd.	≤ 0.063	≤ 4	≤ 0.5	32

328/16	<i>K. pneumoniae</i>	>16	>32	>16	>32	4	>32	≤2	nd.	>64	>128	64	64
164/16	<i>K. pneumoniae</i>	>16	>32	>16	>32	>8	>32	>32	>8	>64	>128	>64	8
138/16	<i>K. pneumoniae</i>	>16	4	≤1	≤2	1	4	4	2	≤0.063	≤4	≤0.5	64
95/16	<i>K. pneumoniae</i>	>16	8	≤1	≤2	≤0.5	4	4	2	≤0.063	≤4	≤0.5	>64
68/16	<i>K. pneumoniae</i>	>16	>32	8	8	≤0.5	≤2	≤2	>8	4	>128	≤0.5	64
19/16	<i>K. pneumoniae</i>	>16	>32	>16	16	>8	>32	16	>8	>64	>128	≤0.5	64
544/13	<i>K. pneumoniae</i>	>16	nd.	>16	16	4	nd.	32	nd.	>64	>128	>64	16
461/16	<i>K. pneumoniae</i>	16	nd.	>16	>32	≤0.5	nd.	≤2	nd.	>2	≤4	≤0.5	32
908/15	<i>K. pneumoniae</i>	>16	nd.	2	≤2	>8	nd.	>32	nd.	>2	>128	32	32
80/15	<i>K. pneumoniae</i>	>16	>32	>16	32	>8	32	8	>8	8	>128	≤0.5	1
94/15	<i>K. pneumoniae</i>	>16	>32	>16	32	>8	>32	16	>8	4	≤4	≤0.5	≤0.5
107/15	<i>K. pneumoniae</i>	>16	>32	≤1	≤2	≤0.5	≤2	≤2	>8	1	≤4	1	≤0.5
268/15	<i>K. pneumoniae</i>	>16	nd.	>16	>32	1	nd.	4	nd.	>2	>128	16	32
577/15	<i>K. pneumoniae</i>	>16	>32	>16	4	>8	32	8	>8	0.125	>128	≤0.5	≤0.5
503/15	<i>K. pneumoniae</i>	>16	>32	>16	>32	>8	>32	>32	>8	4	>128	32	64
229/15	<i>K. pneumoniae</i>	>16	>32	>16	>32	>8	>32	>32	>8	>64	>128	16	≤0.5
RKI ID	Species	AMP	MSU	CTX	CAZ	GEN	KAN	AMK	OTE	CIP	SXT	MEM	CST
52/07	<i>K. oxytoca</i>	>16	>32	>16	>32	2	>32	≤2	>8	64	>128	32	2
609/16	<i>K. oxytoca</i>	>16	>32	>16	16	>8	32	8	1	2	>128	16	1
93/12	<i>P. aeruginosa</i>	>16	>32	>16	>32	>8	>32	>32	>8	0.5	>128	16	≤1
628/16	<i>P. aeruginosa</i>	>16	>32	>16	4	4	>32	8	>8	0.5	>128	8	1
<i>Serratia</i>													
517/16	<i>fonticola</i>	>16	>32	>16	32	4	4	4	>8	4	>128	32	>64
532/14-1	<i>S. enterica</i>	>16	>32	>16	>32	4	>32	4	>8	0.5	128	≤0.5	8
436/14	<i>S. enterica</i>	>16	>32	>16	16	≤0.5	≤2	≤2	2	≤0.063	>128	≤0.5	≤0.5
270/14	<i>S. enterica</i>	>16	>32	>16	8	1	4	≤2	4	≤0.063	>128	≤0.5	≤0.5
205/14	<i>S. enterica</i>	>16	>32	>16	16	1	≤2	≤2	2	≤0.063	≤4	≤0.5	8
583/14	<i>S. enterica</i>	>16	>32	>16	>32	>8	4	≤2	>8	2	8	≤0.5	≤0.5
277/14	<i>S. enterica</i>	>16	32	>16	8	>8	>32	4	>8	0.25	>128	≤0.5	≤0.5
419/14	<i>S. enterica</i>	>16	>32	>16	8	≤0.5	≤2	≤2	2	≤0.063	>128	≤0.5	≤0.5
533/14	<i>S. enterica</i>	>16	>32	>16	>32	≤0.5	4	4	>8	0.5	≤4	≤0.5	≤0.5
437/14	<i>S. enterica</i>	>16	>32	>16	8	1	4	4	>8	0.125	>128	≤0.5	≤0.5
206/14	<i>S. enterica</i>	>16	>32	>16	>32	>8	>32	4	2	≤0.063	≤4	≤0.5	≤0.5
729/15	<i>St. maltophilia</i>	>16	32	16	4	>8	>32	32	>8	8	>128	>64	>64
136/09	<i>St. maltophilia</i>	8	8	2	≤2	2	>32	4	>8	4	≤4	>64	8
599/16	<i>M. morgannii</i>	>16	>32	>16	>32	2	4	4	>8	16	>128	>64	>64
81/15	<i>M. morgannii</i>	>16	4	16	16	≤0.5	≤2	≤2	>8	≤0.063	≤4	≤0.5	>64
93/15	<i>M. morgannii</i>	>16	4	4	16	>8	>32	≤2	>8	16	≤4	≤0.5	>64
659/13	<i>M. morgannii</i>	>16	nd.	8	8	nd.	nd.	nd.	nd.	≤0.063	≤4	≤0.5	>64
604/16	<i>C. freundii</i>	>16	>32	>16	32	>8	32	4	4	8	>128	>64	≤0.5

^a AMP, ampicillin; MSU, mezlocillin/sulbactam (0.8 µg/mL); CTX, cefotaxim; CAZ, ceftazidim; GEN, gentamicin; KAN, kanamycin; AMK, amikacin; OTE, oxytetracycline, CIP, ciprofloxacin; SXT,

sulfameracin-trimethoprim; MEM, meropenem; CST; colistin. ^b nd represents values which were not determined.

Table S2. Distribution of the MIC values of EtCec1-a in the panel of Gram-negative clinical isolates.^a

Species and resistance phenotype ^b (no. isolates)	MIC of EtCec1-a (µg/mL)							
	2	4	8	16	32	64	128	>128
<i>E. coli</i> (26)	1	15	9	1				
CST ^R MEM ^R (1)			1					
CST ^R (9)		5	4					
MEM ^R (4)		3	1					
S (12)	1	7	3	1				
<i>E. cloacae</i> (23)	1	15	3	4				
CST ^R MEM ^R (1)				1				
CST ^R (3)	1	1		1				
MEM ^R (10)		7	1	2				
S (9)		7	2					
<i>E. aerogenes</i> (1)		1						
CST ^R MEM ^R (1)		1						
<i>K. pneumoniae</i> (21)		6	7	7	1			
CST ^R MEM ^R (6)		1	4		1			
CST ^R (9)		1	1	7				
MEM ^R (2)		1	1					
S (4)		3	1					
<i>K. oxytoca</i> (2)		2						
MEM ^R (2)		2						
<i>S. enterica</i> (10)				9	1			
CST ^R (2)				2				
S (8)				7	1			
<i>C. freundii</i> (1)			1					
MEM ^R (1)			1					
<i>A. baumannii</i> (20)		10	9	1				
CST ^R MEM ^R (3)		1	2					
MEM ^R (16)		9	6	1				
S (1)			1					
<i>A. pittii</i> (1)		1						
S (1)		1						
<i>P. aeruginosa</i> (2)							2	
MEM ^R (2)							2	
<i>S. maltophilia</i> (2)						1	1	
CST ^R MEM ^R (2)						1	1	
<i>M. morgani</i> (4)								4
CST ^R MEM ^R (1)								1
CST ^R (3)								3

<i>S. fonticola</i> (1)	1
CST ^R MEM ^R (1)	1

^a Table shows the numbers of isolates for which the corresponding MIC value was determined. ^b CST^R, resistant to colistin; MEM^R, resistant to meropenem; S, sensitive to colistin and meropenem.

Table S3. Distribution of the MIC values of EtCec2-a in the panel of Gram-negative clinical isolates.^a

Species and resistance phenotype ^b (no. isolates)	MIC of EtCec2-a (µg/mL)							
	2	4	8	16	32	64	128	>128
<i>E. coli</i> (26)		1	8	13	4			
CST ^R MEM ^R (1)					1			
CST ^R (9)		1	3	3	2			
MEM ^R (4)			1	3				
S (12)			4	7	1			
<i>E. cloacae</i> (23)		1	1	5	10		4	2
CST ^R MEM ^R (1)							1	
CST ^R (3)		1			1		1	
MEM ^R (10)				2	5		1	2
S (9)			1	3	4		1	
<i>E. aerogenes</i> (1)				1				
CST ^R MEM ^R (1)				1				
<i>K. pneumoniae</i> (21)				1	7		4	9
CST ^R MEM ^R (6)				1	4			1
CST ^R (9)							3	6
MEM ^R (2)								2
S (4)					3		1	
<i>K. oxytoca</i> (2)			1		1			
MEM ^R (2)			1		1			
<i>S. enterica</i> (10)					3		6	1
CST ^R (2)					2			
S (8)					1		6	1
<i>C. freundii</i> (1)							1	
MEM ^R (1)							1	
<i>A. baumannii</i> (20)		1	9	8	1			1
CST ^R MEM ^R (3)			2	1				
MEM ^R (16)		1	7	8				
S (1)								1
<i>A. pittii</i> (1)			1					
S (1)			1					
<i>P. aeruginosa</i> (2)								2
MEM ^R (2)								2
<i>S. maltophilia</i> (2)								2
CST ^R MEM ^R (2)								2
<i>M. morgani</i> (4)								4

CST ^R MEM ^R (1)	1
CST ^R (3)	3
<hr/>	
<i>S. fonticola</i> (1)	1
CST ^R MEM ^R (1)	1
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^a Table shows the numbers of isolates for which the corresponding MIC value was determined. ^b CST^R, resistant to colistin; MEM^R, resistant to meropenem; S, sensitive to colistin and meropenem.

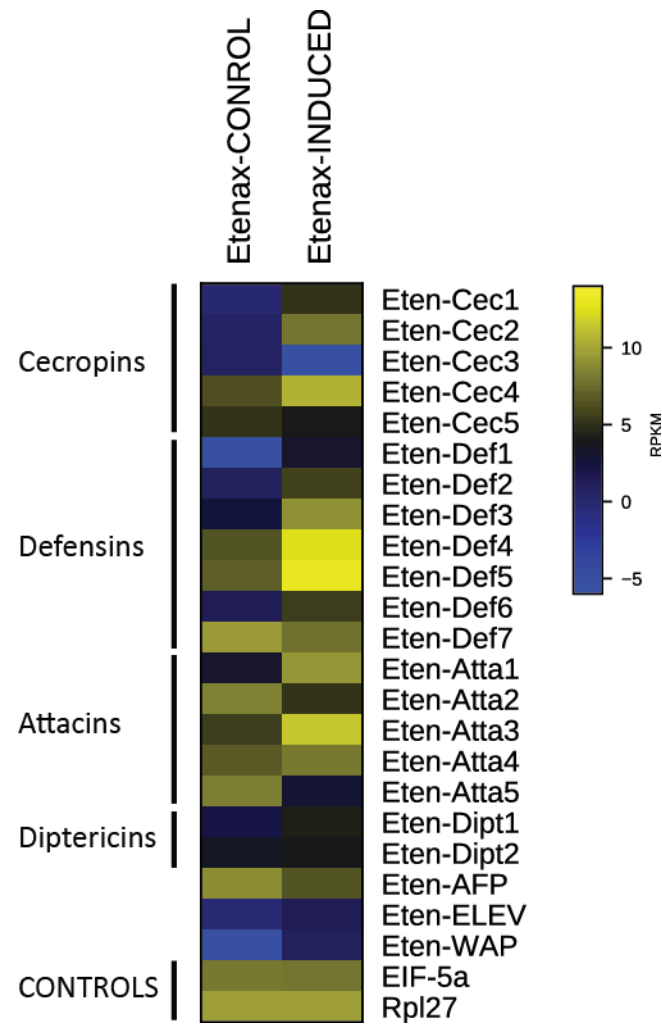


Figure S1. Heatmap of the 22 *Eristalis tenax* AMPs discovered.

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Cecropin A      1 ---KWKLFKKIEKVGQNI RDGI I KAGPAVAVVGGATQIAK--
EtCec1         1 -GFLKKI GKKLE GAVQRT RDATIQTI IAVAQAAANVAATAKQ-
EtCec2         1 -GWL RDFGKRIERTGQNI RDATIQTI IGAQEAANVAATLK--
EtCec3         1 -GFLKVVGKKLE GASDLTRDATIQTI IAVAQAAANVAATAKQ-

Defensin 1     1 VTCDLLSAAEAKG----VKVNHAA CAAHCLLKRKRGGYCNKRRICVCRN-
Et-Def1        1 AACSLGSLINVG-----CNSCA CAAHCLATRGKNGACNSQRRRCVCK-
Et-Def4        1 ATCDLLSEFN-----VKDAACA AHCLAKGYRGGYCDGRKVCNCRK-

Hymenoptaecin 1 -----HADPQ GSLVINGKKPLSGPDRRPSLDVDYHQRVYDRNQMNA DAYGGLNIRPG
EtDip          1 IN -----QFNMQG-----GGSPRQ GFDVNA NARFPIWQ SQNARNSVHG TAS-----

Hymenoptaecin 53 QPAQPHLG VQIQREYKNGFIRGYSQAERGPGRISPSFVGGGFRF--
EtDip          42 ---IN -----YAQHLGG EYGN SRPN EGGGLQFT

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Figure S2. Alignment of *Eristalis tenax* derived peptides to their analogues from other insect species. Shown are similarities of the cecropin-like AMPs to the sequence of Cecropin A from *Hyalophora cecropia*, of the defensin-like AMPs to defensin 1 from *Tribolium castaneum*, and of the dipterocin-like AMP to Hymenoptaecin from *Bombus pascuorum*.