Supplementary

FIGURE LEGENDS

Figure 1. Flow chart of study selection process

Figure 2. The mixed effect of ACEi and ARBs on all-cause mortality in patients with diabetes and kidney disease

Figure 3. The mixed effect of ACEi and ARBs on the risk of end-stage renal disease in patients with diabetes and kidney disease

Figure 4. The mixed effect of ACEi and ARBs on the risk of hyperkalemia in patients with diabetes and kidney disease

Figure 5. The mixed effect of ACEi and ARBs on the risk of cough in patients with diabetes and kidney disease

Supplementary Tables 1-6

Table 1. Quality of the assessments made in the 8 meta-analyses included in the meta-analysis

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AMSTAR criteria	Suetonia C Palmer	ChengJ	HaoG	Jicheng Lv	P. Vejakama	Sharma P	Strippoli G. F	Strippoli G. F
AMSTAK criteria	2015	2014	2014	2012	2012	2011	2006	2005
1. Was an 'a priori' design provided?	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
2. Was there duplicate study selection and data extraction?	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
3. Was a comprehensive literature search performed?	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
4. Was the status of publication used as an inclusion criterion?	Yes	Yes	Yes	No	No	Yes	No	Yes
5. Was a list of studies provided?	No	Yes	No	Yes	Yes	Yes	Yes	Yes
6. Were the characteristics of the included studies provided?	Yes	Yes	Yes	No	Yes	No	No	No
7. Was the scientific quality of the included studies assessed and documented?	Yes	No	No	Yes	Yes	Yes	Yes	Yes
8. Was the scientific quality of the included studies used appropriately in formulating conclusions?	Yes	Yes	No	No	Yes	Yes	Yes	Yes
9. Were the methods used to combine the findings of studies appropriate?	Yes	Yes	Yes	No	No	Yes	No	No
10. Was the likelihood of publication bias assessed?	No	No	No	Yes	Yes	No	Yes	Yes
11. Was the conflict of interest stated?	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
Total score (out of 11)	9	10	7	8	9	9	8	9
Overall methodological quality (L=low, M=moderate, H=high)	Н	Н	М	М	Н	Н	М	Н

Assessment of Multiple Systematic Reviews (AMSTAR) scale ratings of the 8 included systematic reviews and meta-analysis

			The total number of samples	Intervention(study/patient)		(95%	6 CI)
First author year	Country	Disease	number	ACEi/ARBs vs Placebo	OR/RR	11	ul
Suetonia C Palmer et.al 2015	New Zealand	type 2 diabetes mellitus, chronic kidney disease	12716	ACEi + calcium-channel blocker 1/335 ARBs 5/4443 ACEi 10/7938	0·20 0·91 0.85	0·01 0·71 0.61	4·17 1·16 1.19
Cheng J et.al 2014	China	diabetes mellitus	43457	ACEi 20/25544 ARB 12/17913	0.87 0.94	0.78 0.82	0.98 1.08
Hao G et.al 2014	China	type 2 diabetes mellitus	18369	ARB/ACE inhibitor 7/18369	0.91	0.83	1
Jicheng Lv et.al 2012	Australia	diabetic kidney disease	19003	ACEI 6/11350 ARB 5/7653	0.84 1.12	0.73 0.88	0.97 1.41
Sharma P et.al 2011	England	Chronic kidney disease	2177	ACEi 4/2177	0.71	0.53	0.96
Strippoli G. F et.al 2006	Australia	diabetic kidney disease	10704	ACEi 21/7295 ARBs 5/3409	0.91 0.99	0.71 0.85	1.17 1.17
Strippoli G. F et.al 2005	Australia	diabetic kidney disease	4570	ACEi 10/4570	0.81	0.64	1.02

Table 2. Effect of ACEi and ARBs on all-cause mortality in patients with diabetes and kidney disease

End-stage renal disease										
First author year	Country	Disease	The total number of samples number	Intervention (study/patient)	OR/RR	(95% 11	% CI) ul			
Suetonia C Palmer et.al	New Zealand	type 2 diabetes mellitus,	9807	ACEi 4/6580	0.73	0.47	1·14			
2015	chronic kidney disease	2001	ARBs 3/3227	0.81	0.69	0.96				
Jicheng Lv et.al 2012	Australia	diabetic kidney disease	10504	ACEi 3/10504	1.94	0.66	5.07			
2012				ARBs 3/6217	0.5	0.09	2.71			
P. Vejakama et.al	751 1 1		12729	ACEi/ARBs 6/2147	0.82	0.64	1.05			
2012	Thailand	type 2 diabetes mellitus	12728	ACEi/ARBs 4/10581	0.8	0.69	0.93			
Strippoli G. F et.al	4 4 1		10070	ACEi 10/6819	0.6	0.39	0.93			
2006	Australia	diabetic kidney disease	10070	ARBs 3/3251	0.78	0.67	0.91			

Table 3. Effect of ACEi and ARBs on the risk of end-stage renal disease in patients with diabetes and kidney disease.

Hyperkalaemia										
			The total number of samples	Intervention		(95% CI)				
First author year	Country	Disease	number	(study/patient)	OR/RR	11	ul			
Suetonia C Palmer	tonia C Palmer	type 2 diabetes mellitus,		ARBs 2/1714	2.54	0.94	6.86			
et.al New Zealand 2015	chronic kidney disease	2194	ACEi 3/480	1.63	0.33	7.95				
Jicheng Lv et.al	1.1.2.1.1.1.1.	5974	ACEi 3/2783	2.98	0.47	18.78				
2012	Australia	diabetic kidney disease	3974	ARBs 3/3191	2.3	0.69	7.71			
Strippoli G. F et.al 2006	Australia	diabetic kidney disease	1219	ACEi 2/1219	0.85	0.32	2.21			
Strippoli G. F et.al 2005	Australia	diabetic kidney disease	2594	ACEi 2/2594	2.95	0.31	28.18			

Table 4. Effect of ACEi and ARBs on the risk of hyperkalemia in patients with diabetes and kidney disease

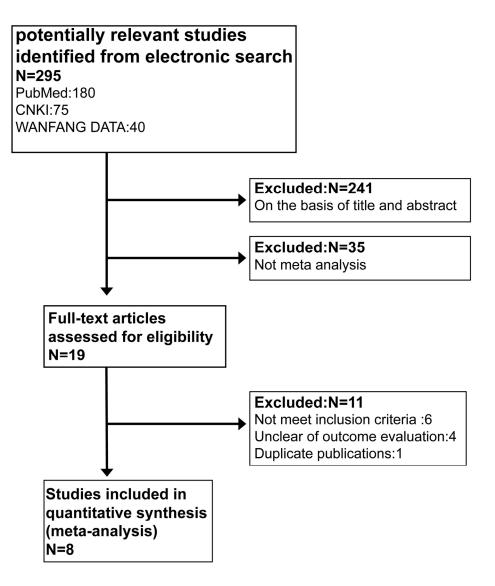
		С	ough				
			The total number of			(959	% CI)
First author year	Country	Disease	samples number	Intervention	OR/RR	11	ul
		type 2 diabetes		ARBs	1.40	0.48	4.1
Suetonia C Palmer et.al 2015	New Zealand	mellitus, chronic kidney disease	22730	ACEi	2.95	1.93	4.5
Jicheng Lv et.al	Australia	diabetic kidney	diabetic kidney 13561 disease	ACEi 6/11791	1.84	1.24	2.7
2012	2012 Australia	disease		ARBs 2/1770	1.01	0.42	2.4
Strippoli G. F et.al	Anstaclic	diabetic kidney	7291	ACEi 10/7087	3.17	2.29	4.3
2006	2006 Australia disease	disease	7281	ARBs 2/194	4.93	1	24.
Strippoli G. F et.al 2005	Australia	diabetic kidney disease	3725	ACEi 4/3725	1.79	1.19	2.6

Table 5. Effect of ACEi and ARBs on the risk of cough in patients with diabetes and kidney disease

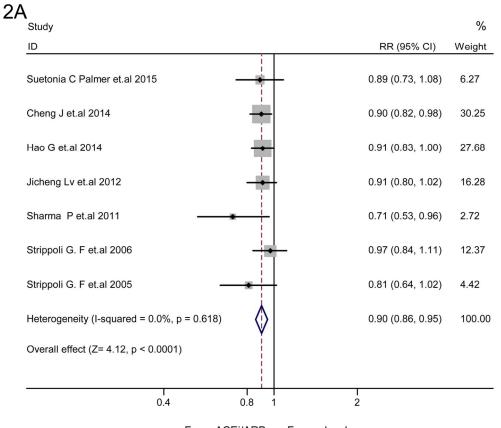
			Headache				
			The total number of samples	Intervention		(95% CI)	
First author year	Country	Disease	number		OR/RR	11	ul
Jicheng Lv et.al	Australia	diabetic kidney	13695	ACEi 3/10504	1.19	0.47	3.02
2012		disease		ARBs 3/3191	0.73	0.52	1.01
Strippoli G. F et.al 2006	Australia	diabetic kidney disease	6186	ACEi 4/6186	0.92	0.33	2.53
Strippoli G. F et.al 2005	Australia	diabetic kidney disease	2438	ACEi 1/2438	1.25	0.44	3.61

 Table 6. Effect of ACEi and ARBs on the risk of headache in patients with diabetes and kidney disease

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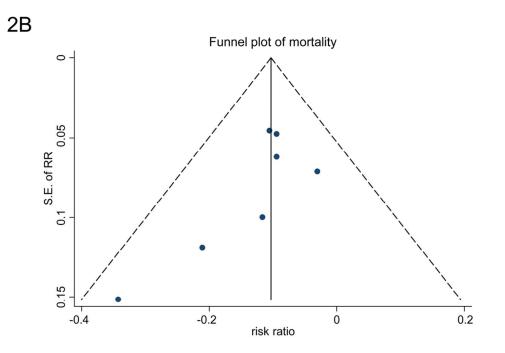


165x206mm (300 x 300 DPI)

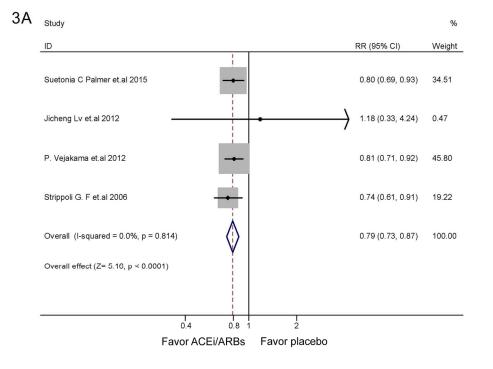


Favor ACEi/ARBs Favor placebo

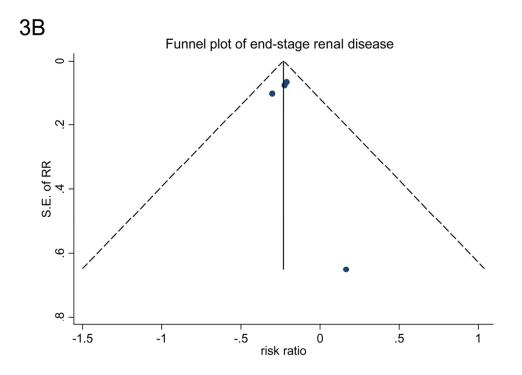
143x127mm (300 x 300 DPI)



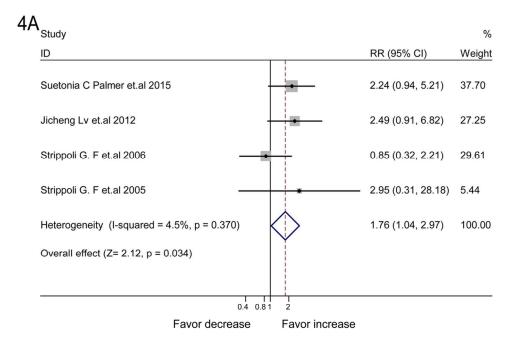
106x74mm (300 x 300 DPI)



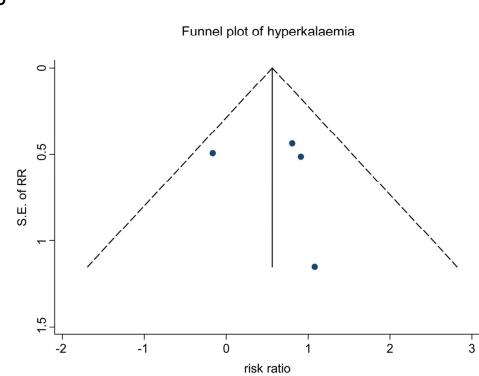
146x107mm (300 x 300 DPI)



105x75mm (300 x 300 DPI)

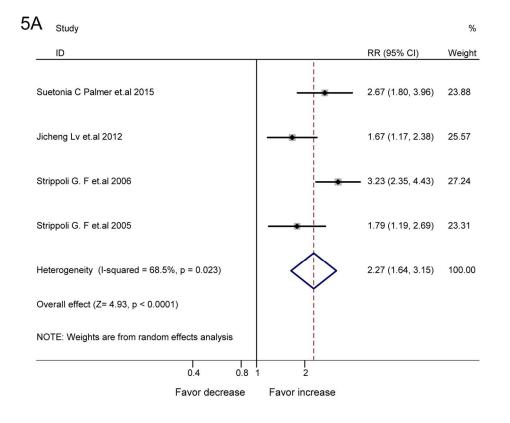


107x73mm (300 x 300 DPI)

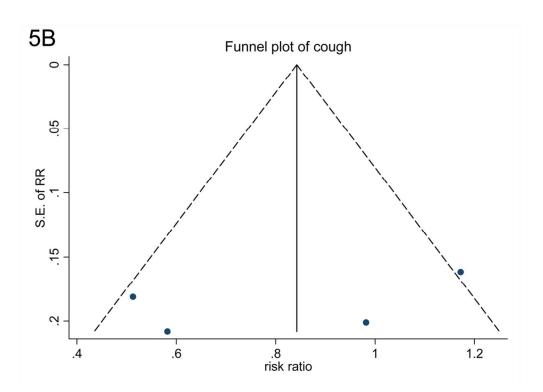


4B

112x87mm (300 x 300 DPI)



139x110mm (300 x 300 DPI)



100x72mm (300 x 300 DPI)