



Supplementary material

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## **Diabetes Mellitus and Risks of Dual Blockade of the Renin-Angiotensin-Aldosterone System**

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### Tabla 1

Referencias bibliográficas de los nueve ensayos incluidos

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<p>3. Uresin Y, Taylor AA, Kilo C, Tschöpe D, Santonastaso M, Ibram G, et al. Efficacy and safety of the direct renin inhibitor aliskiren and ramipril alone or in combination in patients with diabetes and hypertension. <i>J Renin Angiotensin Aldosterone Syst</i>. 2007;8:190-8.</p>
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<p>6. Mehdi UF, Adams-Huet B, Raskin P, Vega GL, Toto RD. Addition of angiotensin receptor blockade or mineralocorticoid antagonism to maximal angiotensin-converting enzyme inhibition in diabetic nephropathy. <i>J Am Soc Nephrol</i>. 2009;20:2641-50.</p>
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<p>8. Titan SM, Vieira JM Jr, Dominguez WV, Barros RT, Zatz R. ACEI and ARB combination therapy in patients with macroalbuminuric diabetic nephropathy and low socioeconomic level: a double-blind randomized clinical trial. <i>Clin Nephrol</i>. 2011;76:273-83.</p>

9. Parving HH, Brenner BM, McMurray JJ, De Zeeuw D, Haffner SM, Solomon SD, et al. Cardiorenal end points in a trial of aliskiren for type 2 diabetes (ALTITUDE). *N Engl J Med*. 2012 3 Nov [Epub ahead of print]. DOI: 10.1056/NEJMoa1208799.

10. FDA Drug Safety Communication: New warning and contraindication for blood pressure medicines containing aliskiren (Tekturna). Bethesda: U.S. Food and Drug Administration; 2012 [citado 22 Abr 2012]. Disponible en: <http://www.fda.gov/drugs/drugsafety/ucm300889.htm>

## Tabla 2

Análisis de sensibilidad (eliminación del estudio ALTITUDE)

Variable de seguridad	Bloqueo dual frente a monoterapia, n/N	Modelo de efectos aleatorios, RR (IC95%)	Modelo de efectos fijos, RR (IC95%)	Heterogeneidad, I <sup>2</sup>
Hiperpotasemia (6 estudios)	83/850; 59/1.127	1,63 (1,12-2,38)	1,56 (1,13-2,16)	12,1%
Hipotensión (2 estudios)	13/368; 5/428	2,86 (0,90-9,14)	2,92 (0,96-8,86)	4,6%
Daño renal (6 estudios)	83/1.058; 108/1.615	1,10 (0,68-1,76)	0,98 (0,75-1,28)	48,5%
Mortalidad (2 estudios)	54/549; 19/824	0,92 (0,52-1,65)	0,96 (0,73-1,28)	4,6%