Rural and Remote Health





The International Electronic Journal of Rural and Remote Health Research, Education, Practice and Policy

LETTER TO THE EDITOR

Appropriate nomenclature: angiotensin II receptors

A D'Amore, EKL Mitchell

Monash University, School of Rural Health, Bairnsdale, Victoria, Australia

Submitted: 27 May 2014; Accepted: 4 September 2014; Published: 4 April 2015

D'Amore A, Mitchell EKL

Appropriate nomenclature: angiotensin II receptors Rural and Remote Health 15: 3150. (Online) 2015

Available: http://www.rrh.org.au

Dear Editor

Radford¹ published an interesting clinical case report of rural patients taking angiotensin II type 1 (AT_1) receptor antagonist – commonly also referred to as angiotensin receptor antagonists, AT_1 receptor blockers, angiotensin receptor blockers or ARBs – who experienced a sudden loss of consciousness whilst exercising gently in the heat. The findings in this manuscript support, in a clinical setting, previous experimental observations seen in rats². The article also highlights the potential need for a larger scale controlled study, which includes a rural cohort, to confirm these observations, considering the widespread use of AT_1 receptor antagonists.

However, Radford's¹ article title, 'Sudden collapses in the heat in patients on angiotensin type 2 receptor blockers' and the referral to angiotensin type 2 receptor blocker (AT_2RB) throughout the manuscript is incorrect nomenclature. We were surprised to see that angiotensin II type 2 (AT_2) receptor antagonists were being used by patients; however, when viewing the medication list in Table 1 of their article, it became clear that it was, in fact, AT_1 receptor antagonists

(namely, irbesartan and candesartan) that were being used by these patients. This is certainly not the first time that inappropriate nomenclature has been used for angiotensin II receptors in the literature.

According to the International Union of Basic and Clinical Pharmacology, and British Pharmacological Society³ and the highly regarded and cited manuscript by de Gasparo et al4, there are two main angiotensin II receptors - the type 1 (AT₁) and type 2 (AT₂) receptors. The actions of the AT₁ receptor are well characterised and include vasoconstriction, aldosterone and vasopressin release, renal reabsorption, increased collagen deposition, cellular proliferation, and cardiomyocyte hypertrophy, as reviewed by de Gasparo et al⁴ and D'Amore et al⁵; hence, the beneficial effects of AT₁ receptor antagonists in conditions such as hypertension. However, the role of the AT₂ receptor remains somewhat enigmatic but is thought to be upregulated in pathological remodelling, oppose the actions of the AT₁ receptor, potentially act constitutively and potentially play a cardio-protective role⁴⁻⁷. With this in mind, it has been suggested that there is a potential pharmacological role of AT_2 receptor ligands⁷.

-Rural-and-Remote-Health



The International Electronic Journal of Rural and Remote Health Research, Education Practice and Policy

With a large amount of research (both basic and clinical) in the area of angiotensin II receptors and the potential future pharmacological use of AT_2 receptor ligands, there is a real danger of causing confusion by the use of incorrect angiotensin II receptor nomenclature. We suggest that future manuscripts (such as angiotensin II receptor-related manuscripts) are published using the correct receptor nomenclature and drug classifications as outlined by the International Union of Basic and Clinical Pharmacology, and British Pharmacological Society⁸.

Angelo D'Amore, PhD, Senior Lecturer Eleanor KL Mitchell, PhD, Lecturer School of Rural Health, Monash University, Victoria, Australia

References

- 1. Radford PJ. Sudden collapses in the heat in patients on angiotensin type 2 receptor blockers. *Rural and Remote Health* 14: 2578. (Online) 2014. Available: www.rrh.org.au (Accessed 26 May 2014).
- 2. Kregel KC, Stauss H, Unger T. Modulation of autonomic nervous system adjustments to heat stress by central ANG II receptor antagonism. *American Journal of Physiology. Regulatory, integrative, and comparative physiology* 1994; **266 (6 Pt 2):** R1985-1991.

- **3**. Alexander SPH, Benson HE, Faccenda E, Pawson AJ, Sharman JL, Spedding M, et al., CGTP Collaborators. The concise guide to pharmacology 2013/14: G protein-coupled receptors. *British Journal of Pharmacology* 2013; **170**: 1459-1581.
- 4. de Gasparo M, Catt KJ, Inagami T, Wright JW, Unger T. International Union of Pharmacology. XXIII. The angiotensin II receptors. *Pharmacological Reviews* 2000; **52**: 415-472.
- **5**. D'Amore A, Black MJ, Thomas WG. The angiotensin II type 2 receptor causes constitutive growth of cardiomyocytes and does not antagonize angiotensin II type 1 receptor-mediated hypertrophy. *Hypertension* 2005; **46(6)**: 1347-1354.
- **6.** Porrello ER, D'Amore A, Curl CL, Allen AM, Harrap SB, Thomas WG, Delbridge LMD. Angiotensin II type 2 receptor antagonizes angiotensin II type 1 receptor-mediated cardiomyocyte autophagy. *Hypertension* 2009; **53**: 1032-1040.
- **7**. McCarthy CA, Widdop RE, Denton KM, Jones ES. Update on the angiotensin AT2 receptor. *Current Hypertension Reports* 2013; **15 (1)**: 25-30.
- 8. Pawson AJ, Sharman JL, Benson HE, Faccenda E, Alexander SPH, Buneman OP, et al., NC-IUPHAR. The IUPHAR/BPS guide to pharmacology: an expert-driven knowledgebase of drug targets and their ligands. *Nucleic Acids Research* 2014; **42 (database issue):** D1098-1106.