



## Microvascular vasodilator properties of the angiotensin II type 2 receptor in a mouse model of type 1 diabetes

Submitted by Daniel Henrion on Mon, 09/24/2018 - 15:22

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| Titre                   | Microvascular vasodilator properties of the angiotensin II type 2 receptor in a mouse model of type 1 diabetes   |
| Type de publication     | Article de revue   |
| Auteur                  | Begorre, Marc-Antoine [1], Dib, Abdallah [2], Habchi, Khalil [3], Guihot, Anne-Laure [4], Bourreau, Jennifer [5], Vessieres, Emilie [6], Blondeau, Bertrand [7], Loufrani, Laurent [8], Chabbert, Marie [9], Henrion, Daniel [10], Fassot, Céline [11]   |
| Editeur                 | Nature Research (part of Springer Nature)  |
| Type                    | Article scientifique dans une revue à comité de lecture  |
| Année                   | 2017   |
| Langue                  | Anglais  |
| Date                    | 2017 03 31   |
| Pagination              | 45625  |
| Volume                  | 7  |
| Titre de la revue       | Scientific reports   |
| ISSN                    | 2045-2322  |
| Résumé en anglais       | <p>Diabetes Mellitus is associated with severe cardiovascular disorders involving the renin-angiotensin system, mainly through activation of the angiotensin II type 1 receptor (AT1R). Although the type 2 receptor (AT2R) opposes the effects of AT1R, with vasodilator and anti-trophic properties, its role in diabetes is debatable. Thus we investigated AT2R-mediated dilatation in a model of type 1 diabetes induced by streptozotocin in 5-month-old male mice lacking AT2R (AT2R<sup>-/-</sup>). Glucose tolerance was reduced and markers of inflammation and oxidative stress (cyclooxygenase-2, gp91phox p22phox and p67phox) were increased in AT2R<sup>-/-</sup> mice compared to wild-type (WT) animals. Streptozotocin-induced hyperglycaemia was higher in AT2R<sup>-/-</sup> than in WT mice. Arterial gp91phox and MnSOD expression levels in addition to blood 8-isoprostane and creatinine were further increased in diabetic AT2R<sup>-/-</sup> mice compared to diabetic WT mice. AT2R-dependent dilatation in both isolated mesenteric resistance arteries and perfused kidneys was greater in diabetic mice than in non-diabetic animals. Thus, in type 1 diabetes, AT2R may reduce glycaemia and display anti-oxidant and/or anti-inflammatory properties in association with greater vasodilatation in mesenteric arteries and in the renal vasculature, a major target of diabetes. Therefore AT2R might represent a new therapeutic target in diabetes.</p> |
| URL de la notice        | <a href="http://okina.univ-angers.fr/publications/ua17600">http://okina.univ-angers.fr/publications/ua17600</a> [12]   |
| DOI                     | 10.1038/srep45625 [13]   |
| Titre abrégé            | Sci Rep  |
| Identifiant (ID) PubMed | 28361992 [14]  |

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## Liens

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- [13] <http://dx.doi.org/10.1038/srep45625>
- [14] <http://www.ncbi.nlm.nih.gov/pubmed/28361992?dopt=Abstract>

Publié sur *Okina* (<http://okina.univ-angers.fr>)