Editorial



A New Vitamin D Receptor Agonist, VS-105: A Promising Path to Control of Postmenopausal Osteoporosis

Darlan Gusso*

Pontificia Universidade Católica do Rio Grande do Sul, Escola de Ciências, Programa de Pós-Graduação em Biologia Celular e Molecular, Porto Alegre, RS. Brazil

Received: November 16, 2020 | Revised: January 27, 2021 | Accepted: January 29, 2021 | Published: February 5, 2021

Vitamin D refers to a group of fat-soluble secosteroids that are crucial for bodily functions. Increasingly evidence has shown their importance in regulating the process of cancer,¹ cardiovascular diseases, and immune responses,²,³ as well as postmenopausal osteoporosis (PMO).⁴ PMO is a metabolic bone disorder that is characterized by decreased bone mass and frequent bone fractures, and affects about 49 million women worldwide.⁵ Despite the variation in osteoporosis phenotypes, osteoporosis is attributed to estrogen deficiency. Clinically, osteoporosis can be diagnosed by assessing bone mineral density.⁶ Although there are various drugs, such as calcitriol, paricalcitol, abaloparatide and rommosozumab, available for PMO treatment, the discovery of new therapeutic drugs remains an urgent need.

In recent years, a vitamin D receptor agonist, VS-105, has been developed and shows potential for the intervention of PMO.7 In fact, a recent article published in the Journal Exploratory Research in Pharmacology entitled "A Novel Vitamin D Receptor Agonist, VS-105, Improves Bone Mineral Density without Affecting Serum Calcium in a Postmenopausal Osteoporosis Rat Model". The authors demonstrated that VS-105 was relatively safe and significantly improved bone mineral density, but did not change serum calcium and phosphate levels in rats. In addition, the therapeutic effects of VS-105 on bone mineral density were comparable to calcitriol. These novel findings are important as a small increase in phosphate levels can be harmful.⁸ These findings extend previous observations in that the administration of VS-105 did not cause hypercalcemia, hyperphosphatemia, or vascular calcification in rodents was found to be safer than paricalcitol. Together, these data suggest that VS-105 may be safe and effective for the intervention of PMO, as well as chronic kidney diseases. Currently, there are ongoing clinical trials to test the safety and pharmacokinetics of VS-105 in human subjects. If successful, this drug should be rapidly translated from the bench to the bedside for the intervention of women with PMO. Overall, VS-105 is a promising adjunctive

Abbreviations: VS-105, vitamin D receptor agonist; PMO, postmenopausal osteoporosis.

*Correspondence to: Darlan Gusso, Laboratório de Neuroquímica e Psicofarmacologia, Escola de Ciências, Pontificia Universidade Católica do Rio Grande do Sul. Avenida Ipiranga, 6681, 90619-900, Porto Alegre, RS, Brazil. ORCID: https://orcid.org/0000-0003-0293-239X. Tel: +55-54-99958-5316, E-mail: gusso.d@hotmail.com How to cite this article: Gusso D. A New Vitamin D Receptor Agonist, VS-105: A Promising Path to Control of Postmenopausal Osteoporosis. *J Explor Res Pharmacol* 2021;6(1):3–4. doi: 10.14218/JERP.2020.00037.

therapy or alternative strategy to improve bone mineral density in patients with PMO.

Acknowledgments

None.

Funding

None.

Conflict of interest

None.

References

- [1] Fathi N, Ahmadian E, Shahi S, Roshangar L, Khan H, Kouhsoltani M, et al. Role of vitamin D and vitamin D receptor (VDR) in oral cancer. Biomed Pharmacother 2019;109:391–401. doi:10.1016/j.biopha.2018.10.102.
- [2] Manson JE, Cook NR, Lee IM, Christen W, Bassuk SS, Mora S, et al. Vitamin D Supplements and Prevention of Cancer and Cardiovascular Disease. N Engl J Med 2019;380(1):33–44. doi:10.1056/NEJ-Moa1809944.
- [3] Baeke F, Takiishi T, Korf H, Gysemans C, Mathieu C. Vitamin D: modulator of the immune system. Curr Opin Pharmacol 2010;10(4):482–496. doi:10.1016/j.coph.2010.04.001.
- [4] Zhang L, Yin X, Wang J, Xu D, Wang Y, Yang J, et al. Associations between VDR Gene Polymorphisms and Osteoporosis Risk and Bone Mineral Density in Postmenopausal Women: A systematic review and Meta-Analysis. Sci Rep 2018;8(1):981. doi:10.1038/s41598-017-18670-7
- [5] Wade SW, Strader C, Fitzpatrick LA, Anthony MS, O'Malley CD. Estimating prevalence of osteoporosis: examples from industrialized countries. Arch Osteoporos 2014;9:182. doi:10.1007/s11657-014-0182-3.
- [6] Eastell R, O'Neill TW, Hofbauer LC, Langdahl B, Reid IR, Gold DT, et al. Postmenopausal osteoporosis. Nat Rev Dis Primers 2016;2:16069. doi:10.1038/nrdp.2016.69.
- [7] Wu-Wong JR, Wessale JL, Chen YW, Chen T, Oubaidin M, Atsawasu-

- wan P. A Novel Vitamin D Receptor Agonist, VS-105, Improves Bone Mineral Density without Affecting Serum Calcium in a Postmenopausal Osteoporosis Rat Model. J Explor Res Pharmacol 2020;5(4):73–80. doi:10.14218/JERP.2020.00020.
- [8] Hong SH, Park SJ, Lee S, Kim S, Cho MH. Biological effects of inorganic phosphate: potential signal of toxicity. J Toxicol Sci 2015;40(1):55–69.
- doi:10.2131/jts.40.55.
- [9] Fujii H, Yonekura Y, Nakai K, Kono K, Goto S, Nishi S. Comparison of the effects of novel vitamin D receptor analog VS-105 and paricalcitol on chronic kidney disease-mineral bone disorder in an experimental model of chronic kidney disease. J Steroid Biochem Mol Biol 2017;167:55–60. doi:10.1016/j.jsbmb.2016.11.002.