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PII: S0891-5849(17)30712-8

DOI: http://dx.doi.org/10.1016/j.freeradbiomed.2017.07.030

Reference: FRB13407

To appear in: Free Radical Biology and Medicine

Received date: 10 February 2017 Revised date: 11 July 2017 Accepted date: 29 July 2017

Cite this article as: Seong-Eun Hong, Jiae Lee, Dong-Hyun Seo, Hye In Lee Doo Ri Park, Gong-Rak Lee, You-Jin Jo, Narae Kim, Minjung Kwon, Hansen Shon, Eun Kyoung Seo, Han-Sung Kim, Soo Young Lee and Woojin Jeong Euphorbia factor L1 inhibits osteoclastogenesis by regulating cellular redoctatus and induces Fas-mediated apoptosis in osteoclast, *Free Radical Biolog and Medicine*, http://dx.doi.org/10.1016/j.freeradbiomed.2017.07.030

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## **ACCEPTED MANUSCRIPT**

Euphorbia factor L1 inhibits osteoclastogenesis by regulating cellular redox status and induces Fas-mediated apoptosis in osteoclast

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#### **Abstract**

Excessive bone resorption caused by increased osteoclast number or activity leads to a variety of bone diseases including osteoporosis, rheumatoid arthritis and periodontitis. Thus, the therapeutic strategy for these diseases has been focused primarily on the inhibition of osteoclast formation and function. This study shows that euphorbia factor L1 (EFL1), a diterpenoid isolated from *Euphorbia lathyris*, inhibited osteoclastogenesis and induced osteoclast apoptosis. EFL1 suppressed osteoclast formation and bone resorption at both initial and terminal differentiation stages. EFL1 inhibited receptor activator of NF-κB ligand (RANKL)-induced NFATc1 induction with attenuated NF-κB activation and c-Fos

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