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Title: Capsaicin cough sensitivity in Korean adults

Dr. Ju-Young 33814 Kim juice99@gmail.com MD ¹², Dr. Kyoung-Hee 33815 Sohn realtopquark@hanmail.net MD ¹², Dr. Min-Gyu 33816 Kang irreversibly@gmail.com MD ¹², Dr. Mi-Yeong 33817 Kim miyoungonion@hanmail.net MD ³, Dr. So-Hee 33818 Lee lshsophia@hanmail.net MD ¹², Dr. Min-Suk 33821 Yang iatrus13@hanmail.net MD ^{1,2,4}, Dr. Min-Hye 33822 Kim mineyang@hanmail.net MD ⁵, Dr. Woo-Jung 33823 Song swj0126@gmail.com MD ¹², Dr. Sae-Hoon 33829 Kim imimdr@yahoo.co.kr MD ^{1,2,6}, Dr. Hye-Ryun 33830 Kang helenmed@hanmail.net MD ¹², Dr. Heung-Woo 33831 Park guinea71@snu.ac.kr MD ¹², Dr. Yoon-Seok 33833 Chang addchang@snu.ac.kr MD ^{1,2,6}, Dr. Sang-Heon 33839 Cho shcho@snu.ac.kr MD ¹² and Dr. Kyung-Up 33840 Min drmin@snu.ac.kr MD ¹². ¹ Department of Internal Medicine, Seoul National University College of Medicine, Seoul, Korea ; ² Institute of Allergy and Clinical Immunology, Seoul National University Medical Research Center, Seoul, Korea ; ³ Department of Internal Medicine, Pusan Paik Hospital, Inje University College of Medicine, Busan, Korea ; ⁴ Department of Internal Medicine, Seoul National University Boramae Medical Center, Seoul, Korea ; ⁵ Department of Internal Medicine, Kyungpook National University School of Medicine, Daegu, Korea and ⁶ Department of Internal Medicine, Seoul National University Bundang Hospital, Seongnam, Korea .

Body: Background: Cough hypersensitivity is a major factor mediating chronic persistent cough. Capsaicin inhalation cough challenge is a representative tool for assessing cough sensitivity. However, the normal ranges and determinants have not been investigated in healthy Korean adults. Methods: Healthy adult volunteers were recruited by public postings at a single tertiary medical institution. For every volunteer, four age-gender-matched patients with chronic cough (lasting more than 8 weeks) were included. Capsaicin cough challenge tests were performed by dose-response, tidal-breathing method (15 seconds) in 1 minute intervals and the concentration causing five or more coughs (C5) was determined. Results: A total of 65 volunteers (52 females and 13 males) with mean age of 35.5 ± 11.4 (SEM) were recruited. Age, gender, atopy, sputum eosinophilia did not influence capsaicin cough sensitivity. In healthy females, BMI showed moderate negative correlation to C5 ($r = -0.353$, $p = 0.001$). Healthy volunteers had significantly higher C5 values compared to the patients (median C5 (IQR): 32 (112) vs 16 (24) respectively; p -value= 0.000). However, in ROC curve analysis, C5 showed poor discriminating value (AUC= 0.329) in identifying clinical cough. Conclusions: Capsaicin cough reflex was examined in Korean adults. Chronic cough patients had significantly more sensitive capsaicin reflex; however, its wide overlap between volunteers and patients may limit its application in the diagnosis of cough patients. The significant relationship between capsaicin sensitivity and BMI in healthy females warrants further consideration in the interpretation of the results.