

GNAS、Gi 和 Gq 蛋白經由 Huangqi、Hemin 和 HMBA 誘導 K562 細胞分化中所扮演的角色

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摘要

異三元體鳥糞嘌呤核酸結合蛋白(Heterotrimeric guanine nucleotide binding proteins, G protein), 位於細胞雙層膜內部, 負責接收膜上受體(receptor)活化之訊號, 並活化作用體(effector), 將訊號傳遞(signal transduction)至下游, 而G蛋白為訊號傳遞路徑之樞紐, 於細胞生長、分化(differentiation)及發育(development)上極具重要性。而G蛋白是由 α 、 β 和 γ 次單元(subunit)組合而成, 目前共發現27種 α 次單元、5種 β 次單元以及14種 γ 次單元, G蛋白 α 次單元具有GTPase活性, 能夠與GTP結合並釋放GDP, 共可分為四大家族: Gs、Gi/o、Gq/11和G12/13。本研究選用人類慢性骨髓性白血病細胞株(chronic myeloid leukemia cell line, CML)K562細胞作為細胞模式, 利用黃耆(Huangqi)、Hemin及HMBA誘導細胞進行分化, 以合成 α -球蛋白(globin)或 β -球蛋白作為分析指標。利用退化性引子擴增G蛋白, 以觀察誘導過程中不同G蛋白之表現變化, 從中共選殖到四種基因為GNAS isoform(GNASL和GNASS)、G α 2、G11 pseudo gene。於誘導後表現不盡相同, 共歸納出三點: (1)經黃耆誘導後可增加GNASL並同時抑制GNASS之表現量, Hemin和HMBA則皆無特別明顯之變化。(2)G α 2於誘導後皆有較明顯之表現。(3)本研究所選殖之G11 pseudo gene, 經由限制酵素XhoI剪切後, 發現主要表現於經黃耆誘導後之K562細胞當中。以上不同之G蛋白其訊號傳遞調控機制尚未明瞭, 須更進一步探討, 尤其中藥黃耆對於血液疾病之治療具有極大助益, 若能完全了解黃耆誘導分化機制, 或許患者可藉此得到治癒或減緩病情之機會。

關鍵詞: G蛋白; 黃耆

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