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5-hydroxymethylcytosine expression in proliferative nodules arising within congenital nevi allows differentiation from malignant melanoma.

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Running title: 5-hydroxymethylcytosine in proliferative nodules.

Key words: giant congenital nevus, proliferative nodules, melanoma, 5-hydroxymethylcytosine, epigenetic modifications.

Abbreviations: α -KG, α -ketoglutarate; 2HG, 2-hydroxyglutarate; 2OG, 2-oxoglutarate; 5-caC, 5-carboxycytosine; 5-fC, 5-formylcytosine; 5-hmC, 5-hydroxymethylcytosine; 5-mC, 5-methylcytosine; 5-MCDG, 5-methylcytosine DNA glycosylase; AID, activation-induced deaminase; APOBEC, apolipoprotein B editing complex; C, cytosine; CGIs, CpG islands; DNMT, DNA methyltransferase; FISH, fluorescence *in situ* hybridization; GADD45A, growth arrest and DNA-damage-inducible protein 45 alpha; GAPDH, glyceraldehyde-3-phosphate dehydrogenase; GCN, giant congenital nevus; H&E, hematoxylin and eosin; IDH -1, -2, isocitrate dehydrogenase -1, -2; qRT-PCR, quantitative real-time PCR; PN, proliferative nodule; PTEN, phosphatase and tensin homolog; TDG, G/T mismatch-specific thymine DNA glycosylase; TDG/BER, TDG-initiated base excision repair; TET, Ten-eleven translocation enzymes; TSG, tumor suppressor gene; SAM, S-adenosylmethionine.

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