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High-resolution respirometry and cancer



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Article

Mitochondrial MDM2 Regulates Respiratory Complex I Activity Independently of p53

Giuseppe Arena,^{1,2,3} Madi Yann Cissé,^{1,2,13} Samuel Pyrdziak,^{1,2,13} Laurent Chatre,³ Romain Riscal,^{1,2} Maryse Fuentes,^{1,2} Jamie Jon Arnold,⁴ Markus Kastner,⁴ Laurie Gayte,^{1,2} Christelle Bertrand-Gaday,⁵ Kevin Nay,⁵ Claire Angebault-Prouteau,⁶ Kerren Murray,⁷ Beatrice Chabi,⁵ Christelle Koechlin-Ramonatxo,⁵ Béatrice Orsetti,^{1,2} Charles Vincent,^{1,2} François Casas,⁵ Jean-Christophe Marine,^{8,9} Sandrine Etienne-Manneville,⁷ Florence Bernex,^{1,10} Anne Lombès,¹¹ Craig Eugene Cameron,⁴ Hervé Dubouchaud,¹² Miria Ricchetti,³ Laetitia Karine Linares,^{1,2,14,*} and Laurent Le Cam^{1,2,14,15,*}

mtMDM2 depletion in cancer cells increase NADH-linked OXPHOS respiration

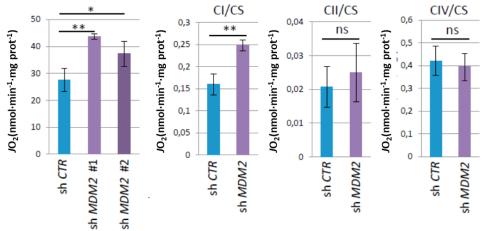


Figure 1. NADH-linked pathway OXPHOS capacity. Lung cancer cells H1299 were transduced with lentiviruses encoding control or two independent MDM2 shRNAs (a) NADH-linked was fuelled with glutamine, malate and pyruvate (b) Complex IV activity, NADH- and succinate pathways were assessed to evaluate the control exerted by MDM2 over respiration. Mean \pm SEM,

MDM2 localization regulates NADH-linked OXPHOS capacity CIV/CS CI/CS CII/CS 0,07 JO₂(nmol·min⁻¹·mg prot⁻¹) prot-1) mg prot-1 25 0,25 0,06 ·mg 0.2 0.2 20 0,15 15 0,04 0,15 JO₂(nmol·min⁻¹ (nmol·min⁻¹ 0,03 10 0,1 0,1 0,02 0.05 0.05 0.01 딮 딮

Figure 2. NADH-linked pathway OXPHOS capacity expressing different MDM2 isoforms. H1299 cells expressing Flag full-length FL-MDM (FL), mitochondrial targeted MTS-MDM2 (MTS), cropped MDM2 1-292 (1-291) or cells transfected with an empty vector (Control). Mean \pm SEM, N=3.

Reference: Arena G, Cissé MY, Pyrdziak S, Chatre L, Riscal R, Fuentes M, Arnold JJ, Kastner M, Gayte L, Bertrand-Gaday C, Nay K, Angebault-Prouteau C, Murray K, Chabi B, Koechlin-Ramonatxo C, Orsetti B, Vincent C, Casas F, Marine JC, Etienne-Manneville S, Bernex F, Lombès A, Cameron CE, Dubouchaud H, Ricchetti M, Linares LK, Le Cam L (2018) Mitochondrial MDM2 regulates respiratory complex I activity independently of p53. Mol Cell 69:594-609.

Figures and texts slightly modified based on the recommendations of the COST Action MitoEAGLE CA15203. Doi:10.26124/mitofit:190001.v3