

**CAMILLA, ORESTE E LE CAROTE DI GIANCARLO
(ITALIAN EDITION)**

Jennifer Khawaja

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Eleven patients treated with single-agent vemurafenib for advanced BRAF V mutant melanoma were analyzed in an exploratory monocentric study. Mutant p53 mutp53 proteins undergo massive constitutive stabilization specifically in tumors which is the key requisite for GOF6–8.

BRAF is the most frequently mutated gene in melanoma. We conducted two phase 2, single-group, multicenter studies of vemurafenib at a dose of mg twice daily -- one in Italy and one in the United States. At 21 months m follow-up, clinical outcome wt compared to mut was not significantly different: These proof-of-principle data identify mutp53 as an actionable cancer-specific drug target.

This study demonstrates the relevance of vemurafenib plasma monitoring during treatment. We use a novel mutp53 mouse model expressing an inactivatable RQ hotspot mutation floxQ to show that tumors depend on sustained mutp53 expression.