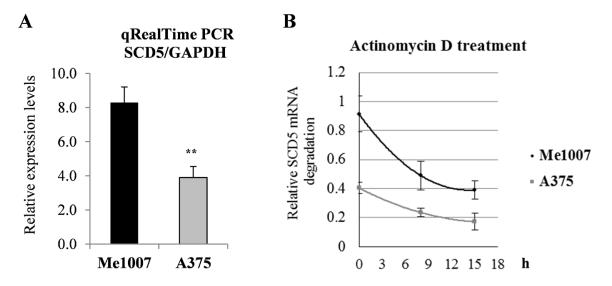
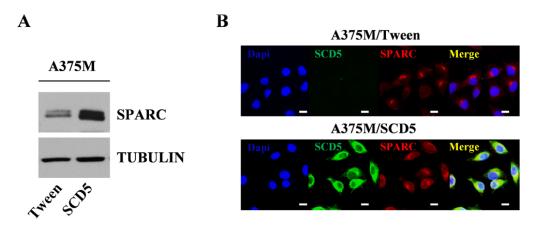
## SCD5 restored expression favors differentiation and epithelialmesenchymal reversion in advanced melanoma

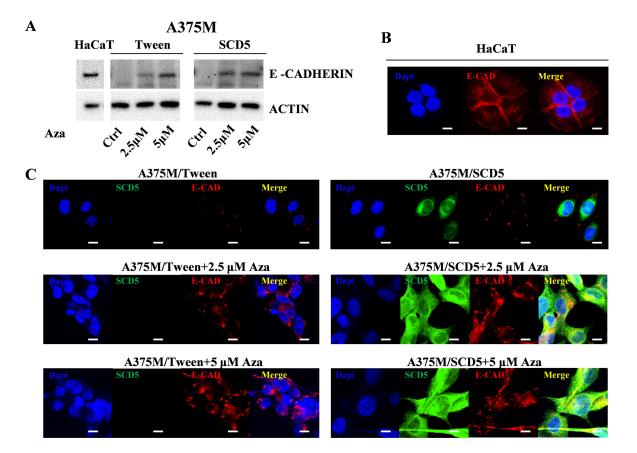
## SUPPLEMENTARY MATERIALS



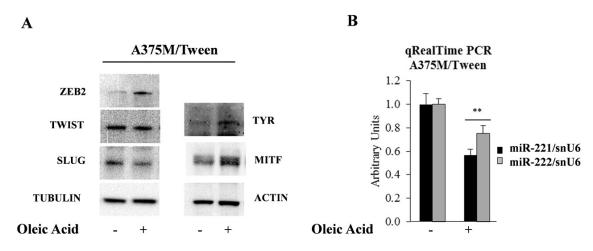
**Supplementary Figure 1:** SCD5 mRNA stability in melanoma cell lines (**A**) The basal level of SCD5 mRNA was quantitated by qRT-PCR in Me1007 early primary and A375 metastatic melanoma cell lines. (**B**) Evaluation of SCD5 mRNA stability in the same cell lines treated with Actinomycin D up to 15 hours. A representative experiment is shown. Although the initial amounts are quite different, no significant differences were detected in their mRNA degradation rates.



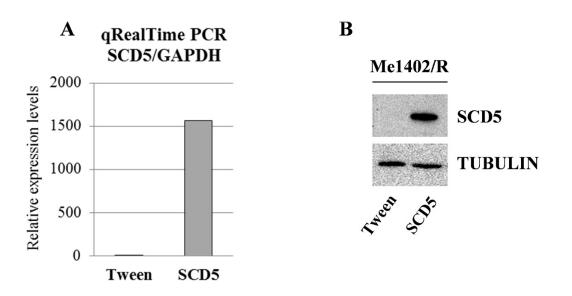
Supplementary Figure 2: SCD5-dependent regulation of SPARC in A375M melanoma. (A) WB and (B) IF analyses show the intracellular accumulation of SPARC. Bar, 10 µm.



**Supplementary Figure 3: E-cadherin epigenetic regulation.** (A) Representative WB analysis demonstrates 5AzaCdR treatment capability to induce E-cadherin full lenght re-expression in both A375M control and A375M/SCD5 cell lines. HaCaT cell line was used as a positive control. (B) IF analysis shows E-cadherin localization at cell membrane junctions in the HaCaT cells and (C) at cell periphery as cytoplasmic spots in A375M/SCD5 compared to the perinuclear distribution in Tween control cells. According to E-cadherin demethylation, 5AzaCdR treatment induces E-cadherin expression, particularly in A375M/SCD5 cells. Bar, 10 µm.



**Supplementary Figure 4: Oleic acid reduces the aggressive phenotype of A375M/Tween cell line.** (A) Western blot evaluation of representative EMT-TFs (left) and differentiation factors (right) in oleic acid (OA) treated A375M/Tween cells, and (B) qRealTime PCR showing miR-221&222 reduced expressions.



Supplementary Figure 5: SCD5 enforced expression in Me1402/R human melanoma cell line. (A) qRealTime PCR and (B) western blot analysis comparison between lentiviral transduction of SCD5 and Tween empty vector control.

Supplementary Table 1: Melanoma cell lines analyzed in the current study

Cell line	<b>B-RAF</b>	N-RAS	Туре	References
Me1007	wt	wt	Primary Tumor VGP*	1,2
Mel501	wt	wt	Primary Tumor VGP*	3, our unpublished results
WM983A	V600E	wt	Primary Tumor VGP*	4, 3
Me1402/R	V600E	wt	Recurrence of Primary Tumor	1, 2
A375	V600E	wt	Metastatic melanoma	5, 6
A375M	V600E	wt	Metastatic melanoma	7, 8

\*VGP vertical growth phase.

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