Genome wide expression profiling of p53 regulated miRNAs in neuroblastoma

Ali Rihani¹, Alan Van Goethem¹, Maté Ongenaert¹, Sara De Brouwer¹, Pieter-Jan

Volders¹, Saurabh Agarwal², Katleen De Preter¹, Pieter Mestdagh¹, Jason Shohet²,

Frank Speleman¹, Jo Vandesompele¹, and Tom Van Maerken^{1,§}

¹Center for Medical Genetics, Ghent University, De Pintelaan 185, B-9000 Ghent,

Belgium

²Texas Children's Cancer Center, Baylor College of Medicine, Houston, Texas

[§]Corresponding author

Figure S1







Figure S2



Figure S1: **Overexpression of miRNAs reduces proliferation.** Neuroblastoma cell lines were transfected with the indicated pre-miRs, and their growth was assessed using the xCELLigence system. Transfection time point is used for normalization (dotted line). IMR-32(A), NGP (B), N206 (C), SKNAS (D), SKNBE(2c) (E).

Figure S2: Survival and expression analysis of miR-432-5p. Kaplan-Meier progression-free survival curve for neuroblastoma patients (n=100) stratified according to miR-432-5p expression levels (quartiles) (log-rank test p-value) (A). Comparison of miR-432-5p expression between neuroblasts and primary neuroblastoma tumors stratified according to MYCN status, data compared using rank product analysis. MNN (*MYCN* non-amplified neuroblastoma), MNA (*MYCN* amplified neuroblastoma) (B).

miR-432-5p expression is associated with progression free survival of neuroblastoma patients and is expressed higher in normal fetal neuroblasts

We analyzed the expression patterns of the aforementioned miRNAs in a cohort of 100 primary neuroblastoma tumors and human fetal adrenal neuroblasts (details in materials and methods), the latter being the cell of origin for neuroblastoma [7,12]. While the miRNAs were not significantly associated to overall survival, low expression of miR-432-5p was significantly associated with poor progression-free survival and shows lower expression in the *MYCN*-amplified neuroblastoma tumors as compared to the tumors without *MYCN* amplification. When comparing the expression of the 4 miRNAs between the neuroblastoma tumors and the neuroblasts, miR-432-5p and miR-203a show higher expression in the neuroblasts as compared to the neuroblastoma tumors (Figure 4).