Supplementary Table 1. Characteristics of the study population.

Characteristics	N (%)
Diagnosis	
De novo AML	150 (85.2)
Secondary AML, including therapy-related AML	26 (14.8)
Sex	
Female	73 (41.5)
Male	103 (58.5)
	59 (16-82)
Median age at diagnosis (years)(range)	
Blood counts at diagnosis:	
Leukocytes per μL (mean)(range)	6900 (900–412000)
Hemoglobin g/dL (mean)(range)	8.8 (1.8–14.4)
Platelets per μL (mean)(range)	82474 (2200–1150000)
Blasts in peripheral blood (%)(range)	36 (0–100)
Blasts in bone marrow (%)(range)	57 (14–99)
LDH μ/L (mean)(range)	600 (97–3842)
Cytogenetic risk	
Low	13 (7.4)
Intermediate	122 (69.3)
High	38 (21.6)
Unknown	3 (1.7)
ELN risk category	
Low	37 (21.0)
Intermediate	66 (37.5)
High	57 (32.4)
Unknown	16 (9.1)
NPM1	
Wild type	128 (72.7)
Mutated	41 (23.3)
Unknown	7 (4)
FLT3-ITD	
Wild type	139 (79)
Mutated	31 (17.6)
Unknown	6 (3.4)
Treatment	
Intensive induction therapy: anthracycline + cytarabine (7+3)	166 (94.3)
Other	9 (5.1)
Exitus prior to induction	1 (0.6)
Response to induction *	
Complete remission	89 (50.6)
Primary refractory disease	67 (38.1)

Did not receive induction	10 (5.7)
Allogeneic HCT in consolidation *	
No	92 (55.4)
Yes	74 (44.6)
Relapse No Yes	119 (67.6) 57 (32.4)
Survival indicators: Progression-free survival, months (median)(range) Event-free survival, months (median)(range) Survival, months (median)(range) Alive at 1-year (n)(%) Exitus	12.5 (0.0–147.0) 13.3 (0.1–148.2) 14.8 (0.2–147.0) 101 (57.4) 117 (66.5)

N: number of observations considered in each analysis; AML: acute myeloid leukemia; ELN: European LeukemiaNet; HCT: hematopoietic cell transplantation.

^{* 10} patients died during induction

Supplementary Table 2. Correlation between gene expression and clinical variables at diagnosis.

Clinical variable	Gene	r	p-value
Age	-	-	ns
Leucocyte count	-	-	ns
Hemoglobin	-	-	ns
Creatinine	-	-	ns
LDH	-	-	ns
Platelets	BCL2	-0.16	0.06
	ASXL1	0.16	0.08
Blasts in PB	BRD4	-0.26	0.01
	EZH2	-0.27	0.01
	ASXL1	-0.2	0.04

R: Pearson coefficient; ns, not significant; LDH: lactate dehydrogenase; PB: peripheral blood.

Supplementary Figure 1. Prediction of blasts in (A) peripheral blood (PB) according to EZH2 and BRD4 expression, and (B) bone marrow (BM) according to EZH2 and ASXL1 expression at diagnosis. Variables selected as predictors of blasts using the Genetic Algorithm model.

