## Supplemental Methods

## Cell lines and culture

NB4, U937 and C1498 AML cells (mouse non-APL acute myeloid leukemia cells) were purchased from Shanghai Bioleaf Biotech Co., Ltd, where they were recently authenticated by STR profiling and characterized by mycoplasma and cell vitality detection. These cells were cultured and maintained in RPMI 1640 medium supplemented with $10 \%$ fetal bovine serum (FBS, Invitrogen, CA, USA) under 5\% carbon dioxide.

## Leukemia cell transplantation

Mouse APL-like spleen cells from $h M R P 8-P m l-R a r \alpha$ transgenic mice or normal spleen cells were intravenously injected into nonirradiated FVB mice ( $2 \times 10^{6}$ cells per mouse) aged between 6 and 10 weeks. To compare the effects of the mouse APL cells and non-APL leukemia cells on the lipid metabolism of the recipient mice, APL cells from hMRP8-Pml-Rar $\alpha$ mice or C1498 (non-APL) leukemia cells were intravenously injected into NSG mice $\left(2 \times 10^{6}\right.$ cells per mouse) aged between 4 and 6 weeks. One week later, transplanted mice were sacrificed and analyzed for lipid levels once a week for 4 weeks.

## ATRA, arsenic and FN treatment in vivo

APL cells from hMRP8-Pml-Rar $\alpha$ transgenic mice were intravenously injected into nonirradiated FVB mice ( $2 \times 10^{6}$ cells per mouse) aged between 6 and 10 weeks. Mice implanted with leukemic cells were randomly assigned to either type of treatment. One day later, when transplanted cells had established, $400 \mathrm{mg} / \mathrm{kg} \mathrm{FN}(100 \mu \mathrm{~L}$, p.o., once a day), 2
$\mathrm{mg} / \mathrm{kg}$ ATRA ( $100 \mu \mathrm{~L}$, p.o., once a day) and $2 \mathrm{mg} / \mathrm{kg} \mathrm{As}_{2} \mathrm{O}_{3}$ ( $100 \mu \mathrm{~L}$, i.p., once a day), or 400 $\mathrm{mg} / \mathrm{kg}$ FN ( $100 \mu \mathrm{~L}$, p.o., once a day), $2 \mathrm{mg} / \mathrm{kg}$ ATRA ( $100 \mu \mathrm{~L}$, p.o., once a day) and $2 \mathrm{mg} / \mathrm{kg}$ $\mathrm{As}_{2} \mathrm{O}_{3}(100 \mu \mathrm{~L}$, i.p., once a day) were administered for 3 consecutive weeks. To evaluate the survival rate, these mice were monitored for 28 days.

## Chromatin immunoprecipitation (ChIP)-qPCR assay

ChIP assays were performed according to the manufacturer's protocol using a SimpleChIP ${ }^{\circledR}$ Plus Sonication Chromatin IP Kit (Cell Signaling Technology, Danvers, MA, USA, \#56383). Briefly, NB4 cells transfected with Myc-tagged PML-RARa plasmids were fixated with $1 \%$ formaldehyde for 8 min , incubated with glycine ( 50 mM final) for 10 min and washed three times with PBS. After cell lysis and chromatin extraction, chromatin was sonicated to 100-500 bp using a BioRuptor sonicator (Diagenode), followed by centrifugation at $16,000 \mathrm{~g}$ for 10 min at $4{ }^{\circ} \mathrm{C}$. The lysates were incubated overnight at $4{ }^{\circ} \mathrm{C}$ with ChIP-grade antibodies specific for Myc (Abcam, \#ab9132), which were coupled to magnetic beads. Precipitated material was eluted (input chromatin was used as a control), the crosslink was reverted, and DNA was purified by chloroform/phenol extraction and resuspended in DNA elution buffer. qPCR analysis was performed using specific primers corresponding to the TRIB3 and PPARG promoters. The primers used for the ChIP-qPCR assays are shown below: TRIB3-1-F: 5'-CCCCACAACTTATATCTAGTGCAGG-3', TRIB3-1-R: 5'-CAGCTGGCATTTAG GGAGCATGTCT-3'; TRIB3-2-F: 5'-TCCCTAAATGCCAGCTGTTATTATG-3', TRIB3-2R: 5'-CATGGACAAGCGCTTGTCTCTCACT-3'; TRIB3-3-F: 5'-TGAGAGACAAGCGC

TTGTCCATGCC-3', TRIB3-3-R: 5'-AGAATCTGTGTGGTCATTTCTG CAT-3'; TRIB3-4F: 5'-AATGACCACACAGATTCTACCAATG-3', TRIB3-4-R: 5'-TGTC TGTGAGGTCTT CAGAGCTGAT-3'; TRIB3-5-F: 5'-AGCTCTGAAGACCTCACAGACA CAT-3', TRIB3-5R: 5'-GCTGAGATTACAGGCGTGAGCCACA-3'; TRIB3-6-F: 5'-TGTAATCTCAG CGCTTTGGGAGGCC-3', TRIB3-6-R: 5'-GCTGGAGTGCAATGGCGTGATCTTG-3'; TRIB3-7-F: 5'-TTGCACTCCAGCCTGGGTGACAGAA-3'; TRIB3-7-R: 5'-GGCCATGC CATGTCCAAGGTCACAG-3'; TRIB3-8-F: 5'-ATTCCCTGTGAC CTTGGACATGGCA-3', TRIB3-8-R: 5'-AGAGCTTGGTTTTGAGCCATGTGCT-3'; TRIB3-9-F: 5'-ACATGGCTCA AAACCAAGCTCTGGG-3', TRIB3-9-R: 5'-TGCCACACCTGGTCCATGGACCCTG-3'; TRIB3-10-F: 5'-AGGGTCCATGGACCAGGTGTGGCAG-3', TRIB3-10-R: 5'-AGGATGG AAGCAAAGCTGCAGCCCT-3'; PPARG-1-F: 5'-ACATTGCTGGTGGGATTGTAAA ATG-3', PPARG-1-R: 5'-TCCATTATACACATATACCACATTT-3'; PPARG-2-F: 5'-ATGTGGTATATGTGTATAATGGAAT-3', PPARG-2-R: 5'-ACCCCCTCTCCATTA ACAGTCATTC-3'; PPARG-3-F: 5'-ATGACTGTTAATGGAGAGGGGGTTC-3', PPARG-3R: 5'-ATGACTGTTAATGGAGAGGGGGTTC-3'; PPARG-4-F: 5'-AGGGTCAAGCG ATTCTACTGCCTCA-3', PPARG-4-R: 5'-TACAATTCAGGCCGGGTATGGCAGC-3'; PPARG-5-F: 5'-ACCCGGCCTGAATTGTACATTT TAC-3', PPARG-5-R: 5'-TAATTTTAA TTGTTTAGTAGAGACT-3'; PPARG-6-F: 5'-AACATGTCAAGACACAGTCTCTACT-3', PPAR G-6-R: 5'-TCTTTTCTTTCTTTCTTCCATGAGA-3'; PPARG-7-F: 5'-AAAGAAAGAAAAGAAAGGAAGAAAG-3', PPARG-7-R: 5'-GTCCTTCCTCCAC AGCCCCTAAGAT-3'; PPARG-8-F: 5'-CTGTGGAGGAAGGACATGATTATGT-3',

PPARG-8-R: 5'-TTCTGGGCCTGATCCTCTTTGGGGA-3'; PPARG-9-F: 5'-AGGATCAGG

CCCAGAACAGTATGCT-3', PPARG-9-R: 5'-AGGCA CGAGAAACAGTTTCTCATGT-3'; PPARG-10-F: 5'-AAACTGTTTCTCGTGCCTCACGTCC-3', PPARG-10-R: 5'-ACATGG TTATTCACAAGTCA CTGAC-3'.

## In vivo study of PPAR $\boldsymbol{\gamma}$ function

To generate cells stably expressing PPAR $\gamma$-shRNA, PPAR $\gamma$-shRNA viral particles were purchased from Genecopoeia, and the target sequence of the shRNA was $5^{\prime}$ '-CAACAGGCCTCATGAAGAA-3'. After 24 h of infection, PPAR $\gamma$-shRNA stable C1498 cells were selected in medium containing $1 \mu \mathrm{~g} / \mathrm{mL}$ puromycin for 14 days. After 2-3 passages in the presence of puromycin, the cultured cells were used for experiments without cloning. ControlshRNA or PPAR $\gamma$-shRNA C1498 cells ( $2 \times 10^{6}$ ) were injected intravenously into C57BL/6 mice aged between 6 and 8 weeks. One week later, transplanted mice were sacrificed and analyzed for lipid levels once a week for 4 weeks.

## Measurement of liver lipids

The fresh mouse liver was isolated and ground uniformly, and 100 mg of liver tissue was accurately weighed into the EP tube. Then, 1 mL of a chloroform/methanol solution (chloroform: methanol $=2: 1, \mathrm{~V} / \mathrm{V}$ ) was added, followed by vigorous vortexing of the mixture for 1 min and letting it stand for 2 h on ice. Then, the mixture was centrifuged at 1650 g for 10 minutes at $4^{\circ} \mathrm{C}$. The mixture was separated into 3 layers: the upper methanol layer, the middle
protein disc, and the bottom chloroform and lipids. The bottom phase was transferred to a new EP tube and stored at $-80^{\circ} \mathrm{C}$ before use. The total TG and TC levels were measured by using commercial kits according to the manufacturers' protocols.

## Isolation of primary mouse hepatocytes

Primary hepatocytes were isolated from normal mouse livers as described in JoVE ${ }^{1}$. Briefly, mouse livers were perfused in situ with EGTA solution followed by pronase (Sigma, P5147) solution and collagenase D (Roche, 11088882001 ) solution at $37^{\circ} \mathrm{C}$ for primary hepatic cell isolation. After perfusion, the liver was transferred into a sterile Petri dish containing protease and collagenase D solution. After being gently minced, the liver and cell suspension were filtered through a $70-\mu \mathrm{m}$ cell strainer and centrifuged at 300 g for 3 min at $4{ }^{\circ} \mathrm{C}$ to isolate hepatocytes. Primary hepatocytes were cultured in DMEM (Gibco, 10564029) supplemented with $10 \%$ FBS, insulin (Sigma, I9278), dexamethasone (Sigma, D4902) and penicillinstreptomycin (Gibco, 15070063) on collagen-coated dishes (Corning, 354236).

## Coculture of APL cells with hepatocytes

For coculture, primary mouse hepatocytes were preseeded in 24 -well plates and cultured with DMEM supplemented with $10 \%$ FBS, insulin, dexamethasone and penicillin-streptomycin on collagen-coated dishes overnight. Primary isolated APL or AML cells ( $5 \times 10^{4}$ cells $/$ well in $10 \%$ FBS in RIPM 1640) were added to the transwell (STEMCELL, \#38024) upper chamber, and hepatocytes and leukemia/normal spleen cells were added at a ratio of $10: 1$. The
leukemia/normal cells and hepatocytes were cocultured together at $37^{\circ} \mathrm{C}$ for 48 h and then separated for further analysis.

## ELISA and TC/TG measurement

Human resistin and leptin levels in plasma or culture supernatant were determined by enzymelinked immunosorbent assay (ELISA) according to the manufacturer's instructions (Beijing 4A Biotech Co., Ltd, China). Mouse PCSK9 levels in plasma or culture supernatant were determined by ELISA according to the manufacturer's instructions (Sino Biological, Beijing, China). Cholesterol and triglycerides (TGs) in the cell culture were assessed by an enzymatic assay according to the supplier's protocols (Applygen Technologies Inc., Beijing, China).

## Stable cell lines

To generate cells stably expressing TRIB3-shRNA1/2 (T3 sh) or control-shRNA (CTRL), plasmids were transfected into NB4 cells with Lipofectamine LTX with Plustransfection reagent (Invitrogen, CA, USA) according to the manufacturer's instructions. After 24 h of transfection, stable transfectants were selected in medium containing $1 \mu \mathrm{~g} / \mathrm{mL}$ puromycin (Gibco, CA, USA) for 14 days. After 2-3 passages in the presence of puromycin, the cultured cells were used for experiments without cloning. To generate cells stably expressing TRIB3 ${ }^{\text {Cas9 }}$, TRIB3 ${ }^{\text {Cas } 9}$ viral particles were purchased from TransOMIC Technologies Inc., and the gRNA target sequence was 5'-GTGCTGGTGACAGTGCGCCA-3'. Cas9 then cut at this sequence, and nonhomologous end-joining (NHEJ) took place. TRIB3-C (CRISPR Clone ID) was used
to construct stable $T R I B 3^{\text {Cas } 9}$ NB4 cells. After 24 h of infection, stable cells were selected in medium containing $1 \mu \mathrm{~g} / \mathrm{mL}$ puromycin for 14 days. After 2-3 passages in the presence of puromycin, the cultured cells were used for experiments without cloning.

## Flow cytometry

Fluorescently labeled antibodies against the following surface proteins were used for human cell staining: CD11b (human), annexin-V FITC, and proprium iodide (PI). Data were acquired using a FACS ${ }^{\text {Canto }}$ II flow cytometer (BD, CA, USA). FCS Express software was used for data analysis.

## Immunoprecipitation, immunoblotting, and immunostaining

Co-IP experiments were performed as described previously ${ }^{23}$. Briefly, cells were collected and lysed for 30 min on ice. Soluble lysates were incubated with the indicated antibodies at $4^{\circ} \mathrm{C}$ overnight, followed by incubation with Protein A/G Plus-Agarose (Santa Cruz Biotechnology, TX, USA) at $4{ }^{\circ} \mathrm{C}$ for 2 h . Immunocomplexes were separated from the beads and then boiled for 10 min . The precipitated proteins were subjected to SDS-PAGE and blotted with specific antibodies. For immunoblotting assays, proteins were extracted from cells using RIPA buffer (Cell Signaling Technology, MA, USA). A BCA Protein Assay Kit was used to determine protein concentrations. Protein extracts were separated by SDS-PAGE, transferred onto PVDF membranes, and subjected to immunoblot analysis. Western blot images were captured by a Tanon 5200 chemiluminescent imaging system (Tanon, Shanghai, Beijing).

## Real-time PCR and RNA interference

Total RNA was extracted using TRIzol (Invitrogen, CA, USA) according to the manufacturer's instructions. Reverse transcription of the total cellular RNA was carried out using oligo (dT) primers and M-MLV reverse transcriptase (Transgen Biotech, Beijing, China). PCR was performed using a Mycycler thermal cycler and analyzed using agarose gels. The sequences of the PCR primers were as follows: PPARG forward, $5^{\prime}$-TCTCTCCGTAATGGAAGACC-3'; PPARG reverse, 5'-CCCCTACAGAGTATTACG-3'; RETN forward, 5'-AGCCATCAATGAGAGGATCCAG-3'; RETN reverse, 5'-TCCAGGCCAATGCTGCTT-3'; GPHN forward, 5'-TCGCCTCTCTACAGCTTCCT-3'; GPHN reverse, 5'-CTGCACCTGGACTGGACATT-3'; ME1 forward, 5'-CGGAACCCTCACCTCAACAA-3'; ME1 reverse, 5'-GTTGAAGGAAGGTGGC AACA-3'; LEP forward, 5'-ATGCATTGG GGAACCCTGTGCGG-3'; ME1 reverse, 5'-TGAGGTCCAGCTGCCACAGCATG-3'; LTC4S forward, 5'-ACGAGGTAGCTCTACTGGCTG-3'; LTC4S reverse, 5'-ACCTGCAGGGAGAAGTAGGC-3'; DHCR7 forward, 5'-GCCATGGTCAAGGGCTACTT3'; DHCR7 reverse, 5'-ACTTCCCGA TCCGAGGGTTA-3'; NKIRAS1 forward, 5'-GCTGCAAGGTTGTGGTTTGT-3'; NKIRAS1 reverse, 5'-CACGCCTTCCTGTAGACCTC3'; MOSC2 forward, 5'-GCAAGCAGCCTTCCTCAAAC-3'; MOSC2 reverse, 5'-GCCTCATTGCCACAGTCTCT-3'; ABCA1 forward, 5'-ATGGCTTGTTGGCCTCAGC-3'; ABCA1 reverse, 5'-GCAGCAGCTGACATGTTTGT-3'; GAPDH forward, 5'-GTGGACATCCGCAAAGACC-3'; and GAPDH reverse, 5'-CCTAGAAGCATTTGCGGTG-

3'. TRIB3 siRNAs were produced by RiboBio (Guangzhou, China) and transfected using Lipofectamine RNA interference MAX Transfection Reagent (Life Technologies, CA, USA) according to the manufacturer's instructions.

## Gene set enrichment analysis (GSEA)

We ranked the 11,378 genes by their association with the APL $(\mathrm{n}=50)$ and non-APL $(\mathrm{n}=50)$
groups (GSE13204) using the signal-to-noise measure in the GSEA program according to $\log 2$ -
fold changes. Lipid metabolic gene sets were collected from the database (http://software.broadinstitute.org/gsea/msigdb/index.jsp).

## Antibody table

| ANTIBODY | SOURCE | IDENTIFIER |
| :--- | :--- | :--- |
| Anti-TRIB3 antibody | Abcam | ab75846 |
| Anti-TRIB3 antibody | Abcam | ab137526 |
| PPAR $\gamma$ antibody | CST | C26H12 |
| Anti-leptin antibody | Abcam | Ab3583 |
| Anti-resistin antibody | Abcam | Ab124681 |
| Gapdh | Zsjqbio | Ta-08 |
| Anti-PML protein antibody | Abcam | Ab96051 |
| PML protein antibody | Novus | NB100-59787 |
| RXR $/ \beta / \gamma$ antibody | Santa Cruz | sc-46659 |
| Anti-Myc-tag mAb | MBL BIOTECH | M047-3 |
| Anti-Flag-tag mAb | MBL BIOTECH | PM020 |
| Ubiquitin antibody | CST | 3933 S |
| Anti-retinoic acid receptor alpha antibody | Abcam | ab28767 |
| $\beta$-Actin (D6A8) rabbit mAb | CST | 8457 S |
| Anti-Myc magnetic beads | Biotool | B26302 |
| Anti-Flag magnetic beads | Biotool | B26102 |
| ALEXA FLUOR(R) 488 RABBIT | Invitrogen | A21210 |
| Mounting medium with DAPI | Zsbio | ZLI-9557 |


| Donkey anti-rabbit IgG $(\mathrm{H}+\mathrm{L})$ highly cross- <br> adsorbed secondary antibody, Alexa Fluor 647 | Invitrogen | A31573 |
| :--- | :--- | :--- |
| Donkey anti-mouse IgG (H+L) highly cross- <br> adsorbed secondary antibody, Alexa Fluor 555 | Invitrogen | A31570 |
| Alexa Fluor® 647 anti-mouse/human CD11b | Biolegend | 101220 |

## Supplementary Figures and Legends



Supplementary Figure S1. (A) qRT-PCR was performed to analyze the mRNA levels of $G P H N, D H C R 7, A B C A 1, L T C 4 S$, MOSC2, and ME1 in primary APL cells and non-APL AML cells. (B) Serum leptin levels in newly diagnosed APL patients $(\mathrm{n}=34)$ and non-APL AML patients $(\mathrm{n}=13)$. $(\mathrm{C})$ The effects of PML-RAR $\alpha$ overexpression on the protein levels of PPAR $\gamma$, resistin, leptin, and TRIB3 in U937 cells. (D) The effects of PML-RAR $\alpha$ overexpression on the mRNA levels of RETN and LEP in U937 cells. The indicated mRNA levels in U937 cells with or without PML-RAR $\alpha$ overexpression were detected by qRT-PCR.


Supplementary Figure S2. TRIB3 and PPARG genes are direct targets of PML-RARa. (A) The TRIB3 promoter was analyzed. Fragments located upstream of ATG were used as the promoter region. (B) ChIP-qPCR assays were performed using different specific primers corresponding to the TRIB3 promoter region. (C) The PPARG promoter was analyzed. Fragments located upstream of ATG were used as the promoter region. (D) ChIP-qPCR assays were performed using different specific primers corresponding to the PPARG promoter region.

## References

1. $\mathrm{Au}-\mathrm{Cui} \mathrm{A}, \mathrm{Au}-\mathrm{Hu} \mathrm{Z}, \mathrm{Au}-\mathrm{Han} \mathrm{Y}, \mathrm{Au}-\mathrm{Yang} \mathrm{Y}, \mathrm{Au}-\mathrm{Li} \mathrm{Y}$. Optimized Analysis of In Vivo and In Vitro Hepatic Steatosis. JoVE. 2017(121):e55178.

Table S1. Clinical information of patients samples used in Figure 1

| Patient No | $\begin{aligned} & \mathrm{APL}= \\ & 1 ; \text { non- } \\ & \mathrm{APL} \\ & \mathrm{AML}=2 \\ & \hline \end{aligned}$ | Gender | Age | $\begin{aligned} & \text { Height } \\ & (\mathrm{cm}) \end{aligned}$ | Body weight (kg) | $\begin{aligned} & \hline \mathrm{BMI}>25= \\ & 1 ;<25=0 \\ & \hline \end{aligned}$ | BMI |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 1 | 1 | Male | 45 | 168 | 77 | 1 | 27.281746 |
| 2 | 1 | Female | 51 | 162 | 53 | 0 | 20.1950922 |
| 3 | 1 | Female | 37 | 163 | 50 | 0 | 18.8189243 |
| 4 | 1 | Male | 19 | 176 | 70 | 0 | 22.5981405 |
| 5 | 1 | Female | 30 | 163 | 47 | 0 | 17.6897889 |
| 6 | 1 | Female | 29 | 156 | 65 | 1 | 26.7094017 |
| 7 | 1 | Female | 25 | 160 | 54 | 0 | 21.09375 |
| 8 | 1 | Male | 38 | 172 | 69 | 0 | 23.3234181 |
| 9 | 1 | Female | 61 | 162 | 55 | 0 | 20.9571712 |
| 10 | 1 | Female | 52 | 161 | 65 | 1 | 25.076193 |
| 11 | 1 | Female | 34 | 160 | 60 | 0 | 23.4375 |
| 12 | 1 | Female | 15 | 160 | 50 | 0 | 19.53125 |
| 13 | 1 | Male | 52 | 179 | 90 | 1 | 28.089011 |
| 14 | 1 | Female | 61 | 157 | 60 | 0 | 24.3417583 |
| 15 | 1 | Female | 44 | 162 | 60 | 0 | 22.8623685 |
| 16 | 1 | Male | 34 | 180 | 85 | 1 | 26.2345679 |
| 17 | 1 | Male | 27 | 178 | 107 | 1 | 33.7709885 |
| 18 | 1 | Male | 66 | 173 | 70 | 0 | 23.3886866 |
| 19 | 1 | Female | 26 | 160 | 55 | 0 | 21.484375 |
| 20 | 1 | Female | 34 | 160 | 62 | 0 | 24.21875 |
| 21 | 1 | Female | 13 | 156 | 45 | 0 | 18.4911243 |
| 22 | 1 | Male | 29 | 170 | 65 | 0 | 22.4913495 |
| 23 | 1 | Female | 24 | 153 | 72 | 1 | 30.757401 |
| 24 | 1 | Female | 21 | 165 | 60 | 0 | 22.0385675 |
| 25 | 1 | Female | 29 | 170 | 73 | 1 | 25.2595156 |
| 26 | 1 | Male | 33 | 168 | 72 | 1 | 25.5102041 |
| 27 | 1 | Male | 41 | 175 | 75.5 | 0 | 24.6530612 |
| 28 | 1 | Male | 39 | 168 | 69 | 0 | 24.4472789 |
| 29 | 1 | Male | 57 | 170 | 74 | 1 | 25.6055363 |
| 30 | 1 | Male | 34 | 170 | 81 | 1 | 28.0276817 |
| 31 | 1 | Female | 44 | 162 | 54 | 0 | 20.5761317 |
| 32 | 1 | Female | 30 | 160 | 54 | 0 | 21.09375 |
| 33 | 1 | Female | 61 | 154 | 60 | 1 | 25.2993759 |
| 34 | 1 | Male | 49 | 165 | 71 | 1 | 26.0789715 |
| 35 | 1 | Female | 62 | 156 | 76 | 1 | 31.2294543 |
| 36 | 1 | Female | 63 | 172 | 70 | 0 | 23.6614386 |
| 37 | 1 | Male | 43 | 180 | 78 | 0 | 24.0740741 |


| 38 | 1 | Male | 47 | 170 | 77 | 1 | 26.6435986 |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 39 | 1 | Female | 45 | 162 | 75 | 1 | 28.5779607 |
| 40 | 1 | Male | 45 | 170 | 80 | 1 | 27.6816609 |
| 41 | 1 | Female | 28 | 160 | 75 | 1 | 29.296875 |
| 42 | 1 | Female | 29 | 165 | 57.5 | 0 | 21.1202938 |
| 43 | 1 | Male | 44 | 162 | 67.5 | 1 | 25.7201646 |
| 44 | 1 | Male | 54 | 167 | 65 | 0 | 23.3066801 |
| 45 | 1 | Female | 16 | 152 | 65 | 1 | 28.1336565 |
| 46 | 1 | Male | 41 | 168 | 74 | 1 | 26.2188209 |
| 47 | 1 | Male | 33 | 173 | 59 | 0 | 19.7133215 |
| 48 | 1 | Female | 19 | 163 | 68 | 1 | 25.5937371 |
| 49 | 1 | Male | 40 | 165 | 62 | 0 | 22.7731864 |
| 50 | 1 | Male | 30 | 174 | 94 | 1 | 31.0476945 |
| 51 | 1 | Male | 47 | 160 | 61 | 0 | 23.828125 |
| 52 | 1 | Male | 27 | 177 | 95 | 1 | 30.3233426 |
| 53 | 1 | Male | 46 | 160 | 72 | 1 | 28.125 |
| 54 | 1 | Female | 17 | 165 | 62 | 0 | 22.7731864 |
| 55 | 1 | Male | 50 | 170 | 78 | 1 | 26.9896194 |
| 56 | 1 | Male | 32 | 174 | 78 | 1 | 25.7629806 |
| 57 | 1 | Male | 59 | 174 | 81 | 1 | 26.7538644 |
| 58 | 1 | Male | 26 | 179 | 95 | 1 | 29.6495116 |
| 59 | 1 | Male | 17 | 171 | 80 | 1 | 27.3588455 |
| 60 | 1 | Male | 28 | 175 | 80 | 1 | 26.122449 |
| 61 | 2 | Male | 50 | 168 | 77 | 1 | 27.281746 |
| 62 | 2 | Male | 37 | 178 | 77 | 0 | 24.3024871 |
| 63 | 2 | Female | 29 | 166 | 49 | 0 | 17.7819713 |
| 64 | 2 | Male | 39 | 178 | 79 | 0 | 24.9337205 |
| 65 | 2 | Female | 48 | 165 | 58 | 0 | 21.3039486 |
| 66 | 2 | Male | 48 | 176 | 91 | 1 | 29.3775826 |
| 67 | 2 | Male | 55 | 168 | 70 | 0 | 24.8015873 |
| 68 | 2 | Female | 57 | 160 | 48 | 0 | 18.75 |
| 69 | 2 | Female | 53 | 165 | 49 | 0 | 17.9981635 |
| 70 | 2 | Male | 43 | 170 | 70 | 0 | 24.2214533 |
| 71 | 2 | Male | 59 | 169 | 75.5 | 1 | 26.4346486 |
| 72 | 2 | Male | 26 | 181 | 65 | 0 | 19.8406642 |
| 73 | 2 | Male | 43 | 172 | 81 | 1 | 27.3796647 |
| 74 | 2 | Male | 17 | 183 | 100 | 1 | 29.8605512 |
| 75 | 2 | Male | 24 | 165 | 61 | 0 | 22.405877 |
| 76 | 2 | Male | 45 | 174 | 73.5 | 0 | 24.2766548 |
| 77 | 2 | Male | 24 | 180 | 95 | 1 | 29.3209877 |
| 78 | 2 | Male | 42 | 172 | 90 | 1 | 30.4218496 |
| 79 | 2 | Female | 28 | 150 | 53 | 0 | 23.5555556 |


| 80 | 2 | Male | 54 | 170 | 77 | 1 | 26.6435986 |
| :--- | :--- | :--- | :--- | :--- | :--- | :--- | :--- |
| 81 | 2 | Male | 38 | 176 | 95 | 1 | 30.668905 |
| 82 | 2 | Female | 17 | 150 | 61 | 1 | 27.1111111 |
| 83 | 2 | Female | 49 | 158 | 55 | 0 | 22.0317257 |
| 84 | 2 | Female | 26 | 162 | 55 | 0 | 20.9571712 |
| 85 | 2 | Male | 54 | 183 | 97 | 1 | 28.9647347 |
| 86 | 2 | Female | 39 | 160 | 66 | 1 | 25.78125 |
| 87 | 2 | Male | 41 | 165 | 65 | 0 | 23.8751148 |
| 88 | 2 | Male | 39 | 174 | 72 | 0 | 23.7812128 |
| 89 | 2 | Male | 57 | 155 | 54 | 0 | 22.4765869 |
| 90 | 2 | Male | 51 | 165 | 65 | 0 | 23.8751148 |
| 91 | 2 | Male | 45 | 170 | 85 | 1 | 29.4117647 |
| 92 | 2 | Female | 45 | 166.5 | 63 | 0 | 22.7254281 |
| 93 | 2 | Male | 21 | 167 | 56 | 0 | 20.0796013 |
| 94 | 2 | Female | 43 | 159 | 59 | 0 | 23.3376844 |
| 95 | 2 | Male | 42 | 159 | 48.5 | 0 | 19.1843677 |
| 96 | 2 | Female | 25 | 165 | 70 | 1 | 25.7116621 |
| 97 | 2 | Female | 22 | 160 | 54 | 0 | 21.09375 |
| 98 | 2 | Male | 52 | 176 | 95 | 1 | 30.668905 |
| 99 | 2 | Male | 49 | 165 | 58.5 | 0 | 21.4876033 |
| 100 | 2 | Female | 33 | 165 | 66 | 0 | 24.2424242 |
| 101 | 2 | Male | 24 | 172 | 57 | 0 | 19.2671714 |
| 102 | 2 | Female | 59 | 168 | 55 | 0 | 19.4869615 |
| 103 | 2 | Male | 25 | 188 | 95 | 1 | 26.8786781 |
| 104 | 2 | Male | 29 | 175 | 80 | 1 | 26.122449 |
| 105 | 2 | Female | 23 | 165 | 60.5 | 0 | 22.2222222 |
| 106 | 2 | Female | 55 | 168 | 55 | 0 | 19.4869615 |
| 107 | 2 | Male | 31 | 181 | 81 | 0 | 24.72452 |
| 108 | 2 | Female | 26 | 164 | 60 | 0 | 22.3081499 |
| 109 | 2 | Male | 37 | 168 | 62.5 | 0 | 22.1442744 |
| 110 | 2 | Female | 43 | 159 | 70 | 1 | 27.6887781 |
| 111 | 2 | Male | 43 | 165 | 55 | 0 | 20.2020202 |
| 112 | 2 | Female | 31 | 167 | 50.5 | 0 | 18.1074976 |
| 113 | 2 | Female | 61 | 158 | 52 | 0 | 20.8299952 |
| 114 | 2 | Female | 28 | 170 | 56 | 0 | 19.3771626 |
| 115 | 2 | Male | 45 | 170 | 75 | 1 | 25.9515571 |
| 116 | 2 | Male | 44 | 172 | 62 | 0 | 20.9572742 |
| 117 | 2 | Female | 38 | 172 | 64 | 0 | 21.6333153 |
| 118 | 2 | Female | 25 | 170 | 59 | 0 | 20.4152249 |
| 119 | 2 | Female | 62 | 159 | 52 | 0 | 22.7731868066 |
| 120 | 2 | Female | 19 | 165 | 62 | 0 |  |
|  |  |  |  |  | 0 |  |  |

Table S2. APL patient treatment history and sample details, related to Figure 2, 3, 5,
6 , and 7.

| Name | AML types | Sample type | Gender | Age | Treatment history |
| :---: | :---: | :---: | :---: | :---: | :---: |
| 1 | APL | PB+BM | M | 27 | ATRA + Arsenic |
| 2 | APL | PB+BM | F | 24 | Ara-C + ATRA |
| 3 | APL | PB+BM | M | 62 | ATRA + Arsenic |
| 4 | APL | PB+BM | M | 41 | ATRA + Arsenic |
| 5 | APL | PB+BM | M | 20 | ATRA + Arsenic |
| 6 | APL | PB+BM | F | 49 | ATRA + Arsenic |
| 7 | APL | PB+BM | M | 40 | ATRA + Arsenic |
| 8 | APL | PB+BM | M | 34 | ATRA + Arsenic |
| 9 | APL | PB+BM | M | 34 | ATRA + Arsenic |
| 10 | APL | PB+BM | F | 49 | ATRA + Arsenic |
| 11 | APL | PB+BM | M | 34 | ATRA + Arsenic |
| 12 | APL | PB+BM | M | 44 | ATRA + Arsenic |
| 13 | APL | PB+BM | M | 33 | ATRA + Arsenic |
| 14 | APL | PB+BM | M | 38 | ATRA + Arsenic |
| 15 | APL | PB+BM | F | 41 | IA, CAG |
| 16 | APL | PB+BM | M | 31 | CAG |
| 17 | APL | PB+BM | M | 20 | CAG |
| 18 | APL | PB+BM | M | 38 | ATRA |
| 19 | APL | PB+BM | F | 33 | IA |
| 20 | APL | PB+BM | M | 18 | IA |
| 21 | APL | PB+BM | M | 38 | ATRA |
| 22 | APL | PB+BM | F | 61 | HAA |
| 23 | APL | PB+BM | M | 27 | ATRA |
| 24 | APL | BM | F | 54 | HAA |
| 25 | APL | PB | M | 53 | ATRA |
| 26 | APL | PB | F | 61 | HAA |
| 27 | APL | BM + PB | M | 27 | ATRA |
| 28 | APL | BM | F | 48 | ATRA |
| 29 | APL | BM | F | 16 | ATRA |
| 30 | APL | BM | F | 50 | ATRA |
| 31 | APL | BM | F | 35 | ATRA |
| 32 | APL | BM | M | 58 | ATRA |
| 33 | APL | BM | F | 17 | ATRA |
| 34 | APL | BM | M | 23 | Dasatinib |
| 35 | APL | BM | F | 30 | ATRA |
| 36 | APL | BM | F | 48 | ATRA |
| 37 | APL | BM | M | 58 | ATRA |
| 38 | APL | BM | M | 55 | ATRA + Sorafenib |
| 39 | APL | BM | F | 17 | ATRA |
| 40 | APL | BM | F | 35 | ATRA |
| 41 | APL | BM | F | 16 | ATRA |
| 42 | AML-M2 ETO+K | PB | M | 48 | HAA |
| 43 | AML-ETO | BM | M | 20 | IA, HAA |
| 44 | AML | PB | F | 19 |  |
| 45 | AML-ETO | BM | M | 21 |  |
| 46 | AML-ETO | BM | M | 48 | HAA |
| 47 | B-ALL | PB | M | 46 | Dasatinib |
| 48 | AML (FLT3-ITD) | PB | M | 50 | Sorafenib |
| 49 | AML (FLT3-ITD) | PB | F | 68 | Sorafenib |


| 50 | AML | PB | F | 55 | HAA |
| :--- | :--- | :--- | :--- | :--- | :--- |
| 51 | non-APL | PB | M | 18 | CAG |
| 52 | non-APL | PB | F | 54 | HAA |
| 53 | T-ALL | PB | M | 29 | CAG |
| 54 | non-APL (FLT3-I | BM | F | 15 | Dasatinib |

APL, Acute Promyelocytic Leukemia; AML, Acute Myelogenous Leukemia; PB, Peripheral Blood; BM, Bone Marrow; F, Female; M, Male; ATRA, All-trans retinoic acid; HAA, Homoharringtonine combined aclarubicin and cytarabine; CAG, Cytarabine, Aclarubicin and G-CSF; IA, Idarubicin.

