

Supplementary information for:

**MS4A3 Promotes Differentiation in Chronic Myeloid Leukemia Cells by Enhancing
Common β Chain Cytokine Receptor Endocytosis**

Author list:

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SUPPLEMENTARY FIGURE LEGENDS

Figure S1. Expression of resistance classifier genes in CD34⁺ cells from AP/BP-CML versus CP-CML patients. Relative mRNA expression for twenty genes with differential expression in CML disease progression and/or imatinib resistance were selected for validation by qRT-PCR in CD34⁺ cells from an independent cohort of patients with AP/BP-CML (n=16) vs. CP-CML (n=17). (*p<0.05)

Figure S2. Myeloid-specific expression of the six-gene set from the combined meta-analysis. Hierarchical trees were generated with the BloodSpot online tool (<http://servers.binf.ku.dk/bloodspot/>) using the normal hematopoiesis vs. acute myeloid leukemia comparison.

Figure S3. Lentivirus-mediated *MS4A3* KD and overexpression, as well as their effect on colony formation by normal adult BM CD34⁺ cells. (A-B) Relative *MS4A3* expression in LAMA-84 CML cell line **(A)** and in CD34⁺ cells from a CP-CML patient **(B)** transduced with three different dox-shRNAs targeting *MS4A3* after ± 0.1 µg/mL dox treatment for 72 h. shRNA #1659 resulted in the best KD and was selected for use in subsequent experiments. Error bars represent SEM. **(C-D)** Relative *MS4A3* mRNA levels of CML and CB samples after lentiviral transduction and culture in the indicated conditions for 72 h. **(E)** Effect of *MS4A3* KD on colony formation of normal adult BM CD34⁺ cells.

Figure S4. *MS4A3* expression is regulated by DNA methylation in CML and is correlated with granulocytic lineage genes in AML. (A) *MS4A3* co-expression genes in TCGA AML cohort (NEJM, 2013)¹ analyzed by cBioPortal². List shows the top positively correlated genes. Red font: *CEBPE* (C/EBP ϵ) is the most significantly correlated transcription factor. Bold font: genes in the six-gene set identified in the combined meta-analysis. **(B-D)** Primary CML CD34⁺ cells were treated with DNA methylation inhibitors as indicated, and *MS4A3* expression level was analyzed by qRT-PCR.

Figure S5. *MS4A3* does not regulate cell cycle progression or pCDK2 level; nor does it decrease with active cell proliferation in CML cells. (A) K562 cells, LAMA-84 cells, and primary CP-CML CD34⁺ cells were lentivirally transduced for *MS4A3* overexpression or KD, and analyzed for DNA content indicating the cell cycle stages using propidium iodide staining and flow cytometry. No obvious change in cell cycle distribution is observed with alterations of

MS4A3. **(B)** Immunoblot and flow cytometry analyses demonstrate no correlation between MS4A3 protein levels and pCDK2 levels in CML cell lines. **(C)** MS4A3 expression levels were detected in cells with or without active proliferation in CD34⁺ cell cultures, using qRT-PCR. Active proliferation (more divisions) is labelled by reduced CFSE fluorescent intensity. No suppression of MS4A3 is observed in proliferating cells.

Figure S6. Confirmation of MS4A3 flow cytometry antibody specificity, and additional images for Fig.6H. **(A)** Relative mRNA level of *MS4A3* after dox-induced shRNA KD in M07e cells. Flow cytometry detection of MS4A3 protein using the monoclonal antibody specific to MS4A3 and PE-conjugated secondary antibody. Antibody labelling reduces as *MS4A3* mRNA decreases with incremental times of dox induction. **(B)** MS4A3 monoclonal antibody recognizes cell-free MS4A3-EGFP protein. MS4A3-EGFP fusion proteins were ectopically expressed in HEK293T cells, and then pulled down with agarose bead-bound polyclonal GFP antibody. MS4A3-EGFP-bound beads were then labelled by MS4A3 monoclonal antibody, GFP monoclonal antibody (positive control, different species from pulldown antibody), and CD3 monoclonal antibody (negative control), followed by corresponding fluorescent secondary antibodies. MS4A3 monoclonal antibody signal was comparable to the positive control. **(C)** LAMA-84 cells with or without GM-CSF stimulation (5 min) were fixed, stained for MS4A3 and CD116, and imaged by confocal microscopy. Arrows depict clusters of MS4A3 and CD116.

Figure S7. MS4A3 knockdown does not inhibit endocytosis of the βc independent receptor – Kit (CD117) and G-CSFR (CD114). **(A)** Representative flow cytometry plots showing the rapid endocytosis of CD117 after SCF stimulation in LAMA-84 cells. **(B)** Quantification of SCF induced CD117 endocytosis in LAMA-84 cells lentivirally transduced with dox-sh*MS4A3*, with or without dox mediated shRNA induction. **(C-D)** CP-CML CD34⁺ cells were revived and lentivirally transduced for *MS4A3* KD or overexpression, then used for CD114 endocytosis assay. Cells were stimulated with 10 ng/mL G-CSF or PBS control for 5 minutes at 37°C, and surface CD114 were analyzed by flow cytometry. **(E)** Baseline CD114 surface expression on cells used in panel C-D. **(F)** CP-CML CD34⁺ cells were revived and lentivirally transduced with empty GFP vector. GM-CSF-induced CD116 endocytosis was analyzed by flow cytometry. **(G-H)** GM-CSF/IL-3-induced intracellular kinase activation in cells used in panel F.

Figure S8. CD116 (CSF2RA / GM-CSFRα), and CD123 (IL3RA / IL-3Rα) expression in myeloid hematopoiesis. **(A)** CD116 and CD123 levels in various cells as shown by BloodSpot

illustration of the Dataset GSE42519. **(B)** Flow cytometry analysis of cell surface CD116 and CD123 expression in CML patient samples (N = 3).

Figure S9. Evolutionary phylogenetic trees of MS4A3-coding and M-CSF-coding gene orthologs. Gene orthologs were discovered in NCBI Gene database. One protein of each species was included in alignment and clustering using the Constraint-based Multiple Alignment Tool³.

METHODS

Cell lines and patient samples.

All CML cell lines were maintained in RPMI1640 supplemented with 10% fetal bovine serum (FBS), 100 U/mL penicillin-streptomycin, and 2 mM L-glutamine (GIBCO, Thermo Fisher Scientific, Waltham, MA, USA). Mononuclear cells (MNCs) from normal cord blood (CB) or femoral head donors, or peripheral blood (PB) of CML patients were separated by density centrifugation on Ficoll-Hypaque (GE Healthcare, Waukesha, WI, USA). CD34⁺ cells were selected using an autoMACS system (Miltenyi Biotec, Bergisch Gladbach, Germany), confirmed for >90% purity on a Guava easyCyte HT Flow Cytometer (Millipore, Billerica, MA), and cryopreserved. Additional normal adult BM CD34⁺ cells collected from adult donors under 60 years old were purchased from AllCells (Alameda, CA, USA). Fresh or frozen CD34⁺ cells were cultured in IMDM supplemented with 10% BIT9500 (StemCell Technologies, Vancouver, BC), 100 U/mL penicillin-streptomycin, 2 mM L-glutamine, and StemSpan cytokines (CC100; StemCell Technologies) for 24-48 h at 37°C prior to use in biological assays. All cell lines used were authenticated by Short tandem repeat (STR) fingerprinting. CML cell lines were confirmed to harbor exclusively native *BCR-ABL1* by conventional Sanger sequencing in both directions using BigDye terminator chemistry on an ABI3730 instrument⁷¹. A complete list of patient samples used in this study can be found in Supplementary Table S1. All patients gave informed consent in accordance with the Declaration of Helsinki, and all studies with human specimens were approved by the University of Utah Institutional Review Board (IRB).

Scl-tTA⁺;TRE-BCR-ABL1⁺ compound transgenic mice.

Scl-tTA⁺;TRE-BCR-ABL1⁺ double transgenic mice were kindly provided by Emmanuelle Passegue⁴. These mice express tetracycline transactivator (tTA) in BM HSCs and CMPs, as well

as BCR-ABL1 under control of a tetracycline response element (TRE), such that BCR-ABL1 expression is repressed in the presence of doxycycline (tetracycline analog, 2 g/L, 5% sucrose, Gold Biotechnology, St. Louis, MO, USA) and expressed upon dox withdrawal. BM cells from double transgenic mice were used for *ex vivo* culture experiments. Lin⁻ BM cells were isolated from *Scl-tTA⁺;TRE-BCR-ABL1⁺* mice maintained in the continuous presence of dox (0.1 µg/mL, Gold Biotechnology Inc.), and cultured for 72-96 hours without dox to turn on BCR-ABL1 expression. Resulting cells were assessed for *Ms4a3* mRNA expression by qRT-PCR. A list of primer sequences is available in Supplementary Table S2. Lin⁻ BM cells were also used for *ex vivo* shRNA-mediated *Ms4a3* KD experiments with scrambled RNA transfected cells as controls (GE Healthcare Pharmacon, Inc., Chicago, IL, USA). These mouse *Ms4a3* shRNA vectors are not dox-inducible. Lentivirus transduced (GFP⁺) Lin⁻ BM cells were plated in semisolid culture (72 h) or colony formation assays ± doxycycline (0.1 µg/ml). For BM transplant, BM cells were flushed out from donor mice. RBC were lysed, and Lin⁻ BM hematopoietic cells were quantified by flow cytometry. Equal number (6x10⁵) of Lin⁻ BM cells from *Ms4a3* knockout and control donors were i.v. injected into lethally irradiated *Ms4a3⁺* recipient mice. All animal experiments in this manuscript were approved by the University of Utah Institutional Animal Care and Use Committee (IACUC).

Xenograft in NSG mice.

CD34⁺ cells from CP-CML (n=3) patients were lentivirally transduced with dox-inducible shRNA targeting human *MS4A3* (sh*MS4A3*) in tandem with GFP, and the resulting unsorted populations were injected into sub-lethally irradiated 8-week-old NSG recipients (NOD.Cg-*Prkdc*^{scid} *Il2rg*^{tm1Wjl}/SzJ, #005557, Jackson Laboratory, Bar Harbor, ME, USA; 6 mice/sample, 1.2x10⁶ cells/mouse) 72 hours post-infection. Following two weeks of engraftment, half of the mice remained on regular water, and half of the mice were placed on dox-containing water (2 g/L and changed twice a week, with 5% sucrose, Gold Biotechnology) to induce knockdown. All recipient mice received sulfadimethoxine (0.125 mg/mL, Vet One, Boise, ID, USA) in the drinking water for the duration of the study. At 12 weeks, all mice were sacrificed to assess for hCD45⁺ GFP⁺ cells in the PB, BM, and spleen.

mRNA expression analysis by qRT-PCR.

RNA extraction was performed using the RNeasy Mini Kit (Qiagen, Germantown, MD) and converted to cDNA with the iScript cDNA Synthesis Kit (Bio-Rad Laboratories, Hercules, CA). cDNAs of indicated genes were detected by qRT-PCR using gene-specific primers

with SsoAdvanced™ SYBR® Green Supermix (Bio-Rad Laboratories) or by using primer/probes in a CFX96 Real-Time PCR Detection System (Bio-Rad Laboratories). Human genes expression was referenced to human beta-glucuronidase (*GUSB*) or *GAPDH*, and mouse genes to *Gapdh* and analyzed in CFX Manager (Bio-Rad Laboratories) using the “ $\Delta\Delta Ct$ method”. qRT-PCR analysis was done in triplicate for each experiment. All primers used with SYBR® Green Supermix were designed to have consistent amplification efficiency across genes in the same PCR program. A list of primers used can be found in Supplementary Table S2 and additional reagents and resources may be found in the Key Resources Table.

Western blotting analysis.

For LAMA-84 cell line, cells were treated as indicated in the experiments and then immediately chilled on ice-water mixture to stop enzymatic activities before being lysed in RIPA buffer. Protein lysates were denatured at 95 °C for 10 min prior to being electrophoresis on SDS-PAGE gels and being transferred to nitrocellulose membranes. Membranes were blocked in 5% non-fat milk in TBST buffer, and probed with primary antibody, followed by species matched secondary antibody-HRP conjugates. Blots were imaged with Pierce™ enhanced chemiluminescence substrate (Thermo Fisher Scientific, Waltham, MA) on ChemiDoc XRS+ imaging system (Bio-Rad Laboratories, Hercules, CA). A list of antibodies used can be found in Supplementary Table S3 and additional reagents and resources may be found in the Key Resources Table.

Plasmids.

Inducible shRNAs targeting *MS4A3* (sh*MS4A3*) were purchased from Cellecta (Mountain View, CA, USA). Briefly, sh*MS4A3* was inserted into a tetracycline-inducible vector (pRSIT12-U6Tet-CMV-TetR-2A-TagGFP-2A-Puro), containing the wild-type tetracycline repressor (*tetR*); thus, transcription is blocked unless 0.1 µg/mL doxycycline is present in the culture medium. Construct #1659 resulted in the most efficient KD of *MS4A3* in CP-CML CD34⁺ cells, and was chosen for further studies (Suppl. Fig. 3). For ectopic expression, human *MS4A3* was PCR amplified from mononuclear cells of a healthy donor, *CEBPE* from KBM5 cells, *MECOM* (transcript RefSeq NM_005241) from HEL cells, and subcloned into the *Xba*I- and *NotI*-digested pCDH-CMV-MCS-EF1-copGFP vector (System Biosciences, Mountain View, CA, USA). Gene inserts were Sanger sequenced (by Univ. of Utah Genomics Core) to verify authenticity.

Lentivirus.

Lentivirus-producing 293FT (#R70007, Thermo Fisher Scientific) cells were maintained in culture in DMEM plus 10% FBS, 2.0 mM L-glutamine, 100 U/mL penicillin-streptomycin, 1.0 mM sodium pyruvate, and 0.1 mM non-essential amino acids (GIBCO). Lentiviral constructs were packaged in combination with vesicular stomatitis virus glycoprotein (VSV-G) (Clontech, Mountain View, CA, USA) and psPax2 (for lentivirus production) using the ProFection® Mammalian Transfection System as recommended by the manufacturer (Promega, Madison, WI, USA). Lentivirus-containing supernatants were concentrated 100X using polyethylene glycol 8000 (Fisher Scientific, Lenexa, KS, USA). Derivative cells were generated by spinoculation of viral particles into cell lines, mouse BM, or primary CD34⁺ samples, followed by FACS sorting of green fluorescent protein-positive (GFP⁺) cells when indicated. Expression of *MS4A3*, *CEBPE*, *MECOM*, and shRNA-mediated KD were validated by qRT-PCR analysis 48-72 h following sorting or drug selection.

Colony formation assays.

Methylcellulose colony formation assays were performed by seeding CML cell lines or CD34⁺ primary cells in 0.9% MethoCult (H4230; StemCell Technologies). 10³ viable cells were plated in humid chambers at 37°C with 5% CO₂, and primary cells were cultured in the presence of CC100 cytokine cocktail (StemCell Technologies). Where indicated, cells were also incubated with or without 0.1 µg/mL dox and/or imatinib at the indicated concentrations in duplicates. Colony forming unit granulocyte-macrophage colonies (CFU-GM) were scored under an inverted microscope following two weeks in culture.

LTC-IC assays.

CD34⁺ cells from CP-CML (n=3) patients were lentivirally transduced with shRNA targeting *MS4A3* (sh*MS4A3*), and resulting GFP⁺ cells were sorted by FACS. Following 96 h culture ± imatinib and/or doxycycline (0.1 µg/ml), remaining CD34⁺ cells were plated in MyeloCult (H5100; StemCell Technologies) on top of irradiated (80 Gy) M210B4 cells in LTC-IC assays as described⁵⁻⁷. After 6 weeks of culture with weekly half-medium changes, cells were trypsinized, plated into methylcellulose colony assays (H4435, StemCell Technologies), and scored after 21 days. BCR-ABL1⁺ colonies were identified by FISH on individually plucked colonies as previously described ⁸. Briefly, individual colonies were picked and resuspended in 100 µL PBS prior to cytopspin and fixation, and cells were immersed in 10% pepsin for 6 minutes at 37°C prior to hybridization. To detect *BCR-ABL1*, the Vysis LSI BCR/ABL Dual Color Dual Fusion Translocation Probe (Abbott Laboratories, Abbott Park, Illinois, USA) was used according to the

manufacturer's instructions. Fluorescent signals were visualized using an Axioskop 2 mot *plus* equipped with an AxioCam microscope camera (Carl Zeiss Microscopy, LLC, Thornwood, NY, USA).

Digital PCR for BCR-ABL1 genotyping.

To detect low levels of *BCR-ABL1* transcripts in the BM, PB, and spleen of xenograft recipient mice, nanofluidic digital polymerase chain reaction was used ⁹. Briefly, RNA was reverse transcribed followed by a pre-amplification step of 14 cycles. This is partitioned into three separate 765-well sections, followed by qRT-PCR using published primer pairs ⁹. Copy number was calculated based on Poisson distribution.

DNA bisulfite conversion and patch PCR sequencing.

DNA bisulfite and patch PCR sequencing was performed on DNA from CD34⁺ cells from normal donor CB or from CP-CML, or BP-CML patients (n=3 for each sample type), as described ¹⁰. Briefly, the human *MS4A3* promoter CpG islands were analyzed between the transcription start site (TSS) and 2000 bp upstream of the TSS. These sequences were scanned for *A/lul* restriction fragments, and patch oligonucleotides were designed by sequentially including base pairs from the *A/lul* restriction sites into fragment sequences until the Tm of the patch oligo was between 62-67°C. Fragments whose patch oligos contained repetitive elements according to the RepeatMasker track were excluded. Patch oligos were then appended with the complement universal primer sequences, and resulting patch oligonucleotides were synthesized by SigmaGenosys. 149 pairs of patch oligos were ordered in a 96-well plate. Oligonucleotide sequences are listed in Supplementary Table S4. Additional sequences that were omitted due to amplification of repetitive elements are highlighted in grey. DNA bisulfite conversion (Zymo Research, Irvine, CA) was performed by the Molecular Diagnostics Section of the Biorepository and Molecular Pathology Shared Resource at Huntsman Cancer Institute (The University of Utah, Salt Lake City, UT). Resulting DNA sequences were analyzed on a HiSeq 50 Cycle Single-Read Sequencing v4 platform at the High-Throughput Genomics Shared Resource at Huntsman Cancer Institute (The University of Utah, Salt Lake City, UT).

32D-cl3 cells expressing BCR-ABL1.

32D-cl3 cells were cultured in complete RPMI1640 supplemented with 10% WEHI culture supernatant as a source of murine IL-3. To generate stable cells, parental 32D-cl3 cells were infected with retrovirus produced by standard procedures using either pMIG-BCR-ABL1-p210-

WT-IRES-EGFP, pMIG-BCR-ABL1-p210-K271R-IRES-EGFP, or the empty vector pMIG-MCS-IRES-EGFP. After infection, the cells expressing p210^{BCR-ABL1}, the kinase-inactive mutant (p210^{BCR-ABL1-K271R}) or the empty vector were sorted for EGFP using a BD FACSAria™ II flow cytometer (BD Biosciences, Franklin Lakes, NJ). Cells were only used with a few passages to avoid phenotypic drifting. To assess the effect of BCR-ABL1 on *Ms4a3* transcript levels, 1x10⁶ cells/well were plated in a 12-well plate and incubated with and without imatinib (2 μM) for indicated time. Cells were pelleted and qRT-PCR was performed as described above.

ChIP-seq.

CP-CML and BP-CML samples (CD34⁺, n=3 each, 1.2x10⁶ cells each), as well as CB samples (CD34⁺, n=3, 0.9-1.5x10⁶ each) were thawed and immediately resuspended in 1% formaldehyde (diluted in PBS from a 37% stock, with 10-15% methanol, F8775-500ML, Sigma-Aldrich, St. Louis, MO) and incubated at room temperature for 10 minutes without shaking. Glycine was added to 125 mM to quench formaldehyde and cells were centrifuged at 2,400g for 5 minutes. The cell pellet was washed in 1 mL of cold PBS and centrifuged again. The supernatant was discarded and the cells were stored at -80°C until further processing. Chromatin immunoprecipitation was performed as previously described ¹¹. Cells were sonicated for 8 cycles of 30 seconds with 30 seconds of rest on an Epishear probe-in sonicator (Active Motif). An antibody against H3K27me3 (Active Motif, cat # 39155) was used for pulldown and input DNA was used as controls for each sample. Libraries were sequenced on a HiSeq 2500 as single-end 50 base pair reads. Sequence reads were aligned using Bowtie ¹². Peaks were called using MACS2 ¹³ and samples were excluded when less than 10,000 peaks were called. MACS2 was used to create read depth normalized bedgraphs to visualize the *MS4A3* locus on IGV ¹⁴.

Epigenetic drug library screening.

Primary CD34⁺ cells from a CP-CML patient were used to screen an Epigenetics Screening Library (#11076, Cayman Chemicals, Ann Arbor, MI, USA). All cells were incubated in the presence of 1 μM compound or DMSO control for 72 h, followed by assessment of *MS4A3* mRNA by qRT-PCR using the Power SYBR Green RNA-to-C_T™ 1-Step Kit (Thermo Fisher Scientific) according to the manufacturer's instructions. Resulting hits were confirmed with additional CML patient samples as indicated.

FACS and flow cytometry analysis.

Sorting of CD34⁺38⁻ stem cells from magnetically enriched CD34⁺ cells was performed using FITC-anti-CD34 (#11-0341-81, eBioscience, San Diego, CA) and APC-anti-CD38 (#102712, BioLegend, San Diego, CA) antibodies. Antibody stained and GFP⁺ cells were sorted by a FACS Aria™ II flow cytometer. For cell cycle analysis following culture in the indicated conditions, cells were fixed in 70% ethanol, stained with propidium iodide (PI), and analyzed for DNA content. For multi-colored flow cytometry, cells were blocked with Human TruStain FcX Fc Receptor Blocking Solution (BioLegend) and then stained with manufacturer recommended concentrations of antibodies in PBS containing 0.5% BSA on ice for 30 minutes. After washing, cells were analyzed on BD FACSCanto, with corresponding single fluorophore instructed compensations. For FRET signal detection in EGFP-RFP double positive cells, cells were excited at 488 nm laser and detected by the 695/40 nm filter (RFP emission) on BD Fortessa. Data were analyzed using FlowJo V10 data analysis software (FlowJo).

Immunofluorescent (IF) staining and confocal microscopy

For HEK293, cells were cultured on poly-Lysine coated cover glass and IF staining was performed directly on the cover glass. For LAMA-84, IF staining was carried out in suspension and stained cells were pressed onto glass slides by Cytospin (1000 rpm for 2 minutes). For surface protein IF staining, cells were fixed with 2% paraformaldehyde in PBS on ice for 15 minutes. After washing with 0.5% BSA in PBS, cells were blocked with 0.5% normal serum of the secondary antibody matched species in PBS for 30 minutes. Primary antibodies and species-matching fluorescently labelled secondary antibodies were incubated with cells in 0.5% BSA in PBS for 1 hour each, with 3 washed in between. Then cell nucleus was stained with Hoechst 33342 (BD) for 5 minutes. For intracellular protein staining and total cell protein staining, cells were fixed with 2% paraformaldehyde in PBS on ice for 15 minutes. After washing with 0.5% BSA in PBS, cells were permeabilized with 0.2% saponin in PBS for 30 minutes, and washed with perm/wash buffer (0.5% BSA and 0.05% saponin in PBS). Primary antibodies and species-matching fluorescently labelled secondary antibodies were incubated with cells in perm/wash buffer for 1 hour each, with 3 washed in between. Then cell nucleus was stained with DAPI (Biolegend) for 5 minutes. All IF staining procedures were performed on ice. Stained cells on slides were mounted with VECTASHIELD® Antifade Mounting Medium (Vector Laboratories), and imaged with a Leica SP8 405-488-561-633 laser confocal microscopy system (Cell Imaging Core, University of Utah). Images are processed and analyzed using LAS X software (Leica). Pearson's coefficient values between EGFP signal and Alexa Fluor 647 conjugated secondary antibodies were calculated with MS4A3-EGFP positive cells in each field of imaging (N=20). Relative fluorescent intensity (/ Max)

of each marker was calculated by marker protein fluorescent intensity divided by the maximal intensity of each cell.

Cytokine receptor endocytosis analysis.

To assess the regulation of cytokine receptor endocytosis by MS4A3, we utilized the Ph⁺ CML cell line, LAMA-84, or primary CD34⁺ cells from CP-CML patients. Cells were transduced with indicated lentiviruses to overexpress or knockdown *MS4A3*. 48 hours later, LAMA-84 cells were selected by Puromycin (1 µg/mL) to obtain pure populations of transduced cells. After viral transduction, CML cells were cultured with or without dox (0.1 µg/mL) to induce knockdown in dox-sh*MS4A3* group for 72 hours. GFP⁺ cells are positively transduced cells, while GFP⁻ cells serve as un-transduced internal control. To stimulate receptor endocytosis, GM-CSF (10 ng/mL) or IL-3 (40 ng/mL) were added to the cells, and cells were incubated at 37 °C for 5 – 15 minutes. After the stimulation, cells were immediately chilled on ice-water mixture to stop cellular activities, and all operations were performed at 4 °C or on ice from then on. Cells with or without cytokine stimulation were washed with ice-cold PBS (+ 0.5% BSA), and stained with CD116-APC antibody (clone REA211, Miltenyi Biotec) or CD123-BV711 antibody (clone 9F5, BD) on ice. After washing, the surface labelled receptor was analyzed by flow cytometry.

Nanoparticles.

The prototype liposomal nanoparticles used in this study were manufactured from LAMA-84 cells that naturally express CD62L and readily shed granular lipid particles from plasma membrane (unpublished data). For Nano-CD62L/MS4A3, particles were collected from the culture supernatant of LAMA-84 cells overexpressing MS4A3-EGFP. For Nano-CD62L (MS4A3-free), particles were collected from the culture supernatant of LAMA-84 cells after CRISPR-mediated MS4A3 knockout. All nanoparticles were pelleted from the supernatant by adding 10% PEG-8000 and 0.3M NaCl, resuspended in RPMI1640 to achieve 100X concentration, and then stored at -80°C in aliquots. Nanoparticles were added to CD34⁺ cell cultures at 50 µL/mL. To facilitate initial nanoparticle uptake, 10 µg/mL polybrene was added to the cultures for 12h, and then diluted or washed off. Mock treatment is the addition of polybrene only. The standard curve coordinating FSC (flow cytometry) and particle size was generated using commercial reference microbeads of known sizes (BD). To titrate the dosage of nanoparticles for delivery into CD34⁺ cells, Kasumi-1 (CD34⁺ myeloblast, doubling time ~48h) cells were used as target cells.

QUANTIFICATION AND STATISTICAL ANALYSIS

Meta-analysis. Series matrix files for the McWeeney dataset (GSE14671), comparing TKI responders versus non-responders¹⁵, and the Cramer-Morales dataset (GSE47927), comparing LSCs versus progenitor cells¹⁶ were downloaded from the Gene Expression Omnibus (<https://www.ncbi.nlm.nih.gov/geo/>). The Zheng dataset (E-MEXP-480) comparing CP-CML versus BP-CML¹⁷, Yong dataset (E-MIMR-17) comparing long versus short duration of CP-CML¹⁸, and Graham dataset (E-MTAB-2508) comparing cycling versus quiescent CML cells¹⁹ were downloaded from ArrayExpress (<http://www.ebi.ac.uk/arrayexpress>). Probe IDs were converted to gene symbols, and genes in each study having an absolute magnitude fold change ≥ 1.5 (for upregulated genes) and ≤ -1.5 (for downregulated genes) were considered for overlapping gene detection if statistical significance (using non-parametric methods) at $p < 0.05$ was observed. The genes identified as commonly up- or down-regulated compared with the McWeeney dataset were tallied and the statistical likelihood for the number of common genes observed compared to that expected by chance was determined using an exact Binomial test as previously described²⁰.

Kaplan-Meier Survival Curve. Correlation of MS4A3 mRNA levels with survival in CML was established using survival data available for 35 patients from the original microarray training set¹⁵. All CEL files from the original study¹⁵ were imported with Partek software (Partek, Inc.) followed by GC-RMA normalization (which included normalization of intensities to the median sample of the experiment). Expression levels were calculated from the microarray (HG-U133A, Affymetrix, Inc.) and dichotomized into high and low groups based on distribution of the data. OS was assessed with Kaplan-Meier curves generated in Prism version 6.04 (GraphPad Software, La Jolla, CA). All statistical analyses were performed in SAS version 9.3 (SAS Institute, Carey, NC), and P-values are 2-sided from a log-rank test.

MS4A3 analysis by microarray. The microarray data presented here are a combination of two separate experiments from Oehler V. et al. (unpublished data) and Zheng et al.¹⁷, which is available from ArrayExpress under accession number E-MEXP-480. In total, CML CD34⁺ cells were analyzed for 8 CP-CML and 7 BP-CML samples by Oehler et al., and 11 CP-CML and 9 BP-CML samples by Zheng et al.¹⁷, yielding a total of 19 CP-CML and 16 BP-CML samples. Briefly, RNA was extracted using the RNeasy Mini Kit (Qiagen, Hilden, Germany). Preparation of hybridization probes and HG-U133A microarray processing were performed according to the manufacturer's instructions (Affymetrix, Santa Clara, CA).

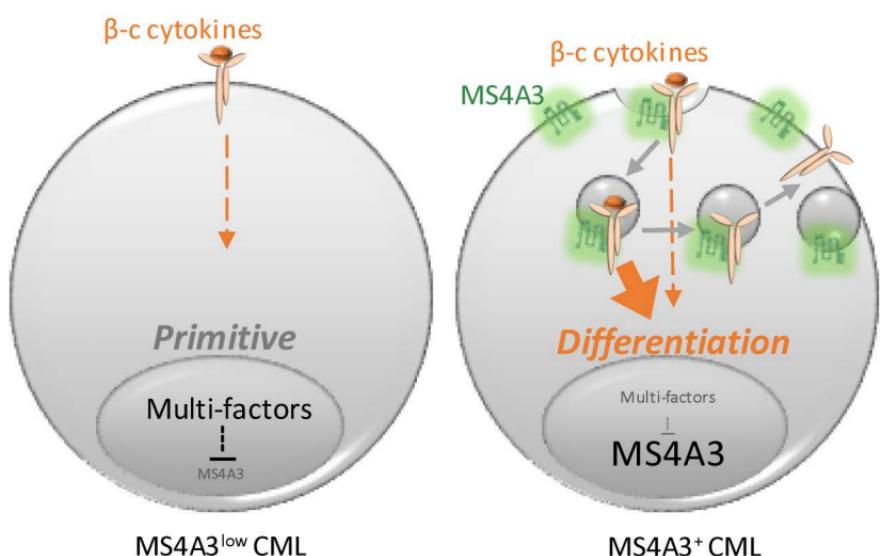
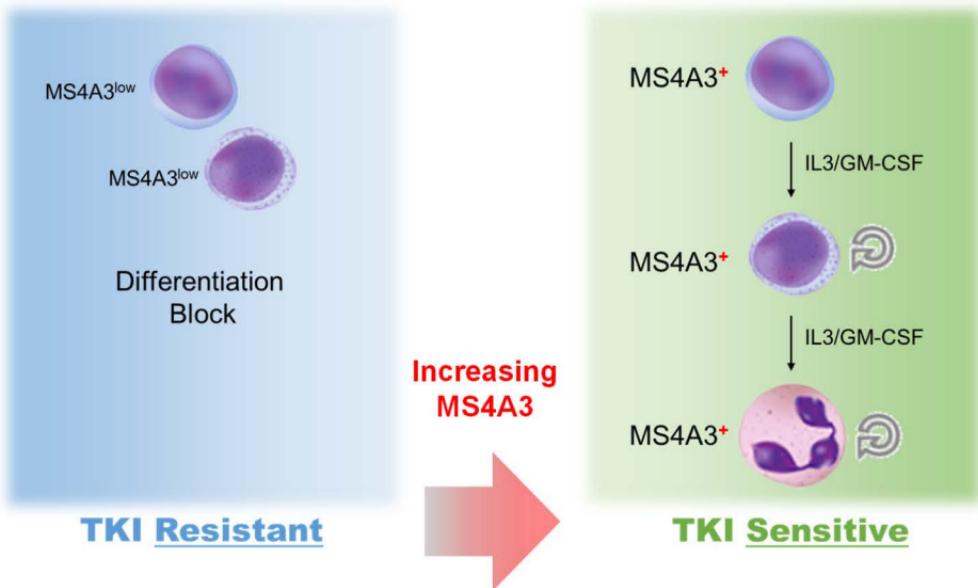
Statistical analysis. Two-tailed *t* test (or with Welch's correction when SD is unequal) was used for experiments in Figure 1C, 1E, 2C-D, 2F-H, 3A-F, 4B-C, 4E-F, 7F-D, 8B-I. When comparing two groups, *t* test is replaced by Mann-Whitney non-parametric test if the sample values do not conform to a normal distribution. One-way ANOVA was used for experiments in Figure 1B, 7E. Fisher's exact test was used for the experiment in Figure 6E. For all assays, three independent experiments were performed unless otherwise noted.

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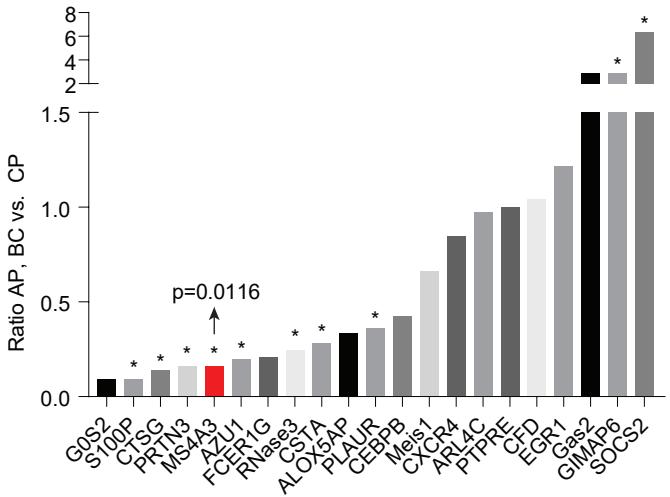
Graphic abstract



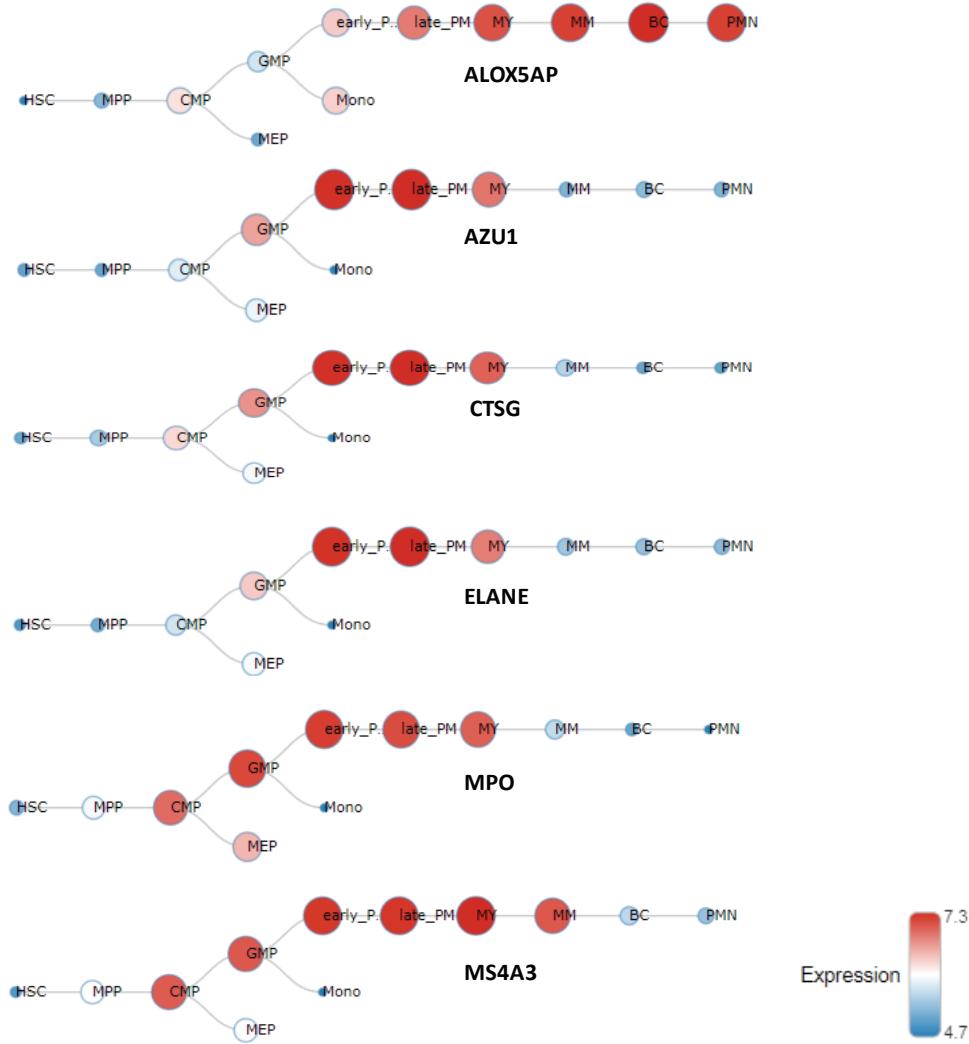
Multi-factors:

BCR-ABL1, MECOM, DNA methylation, EZH2/H3K27me3

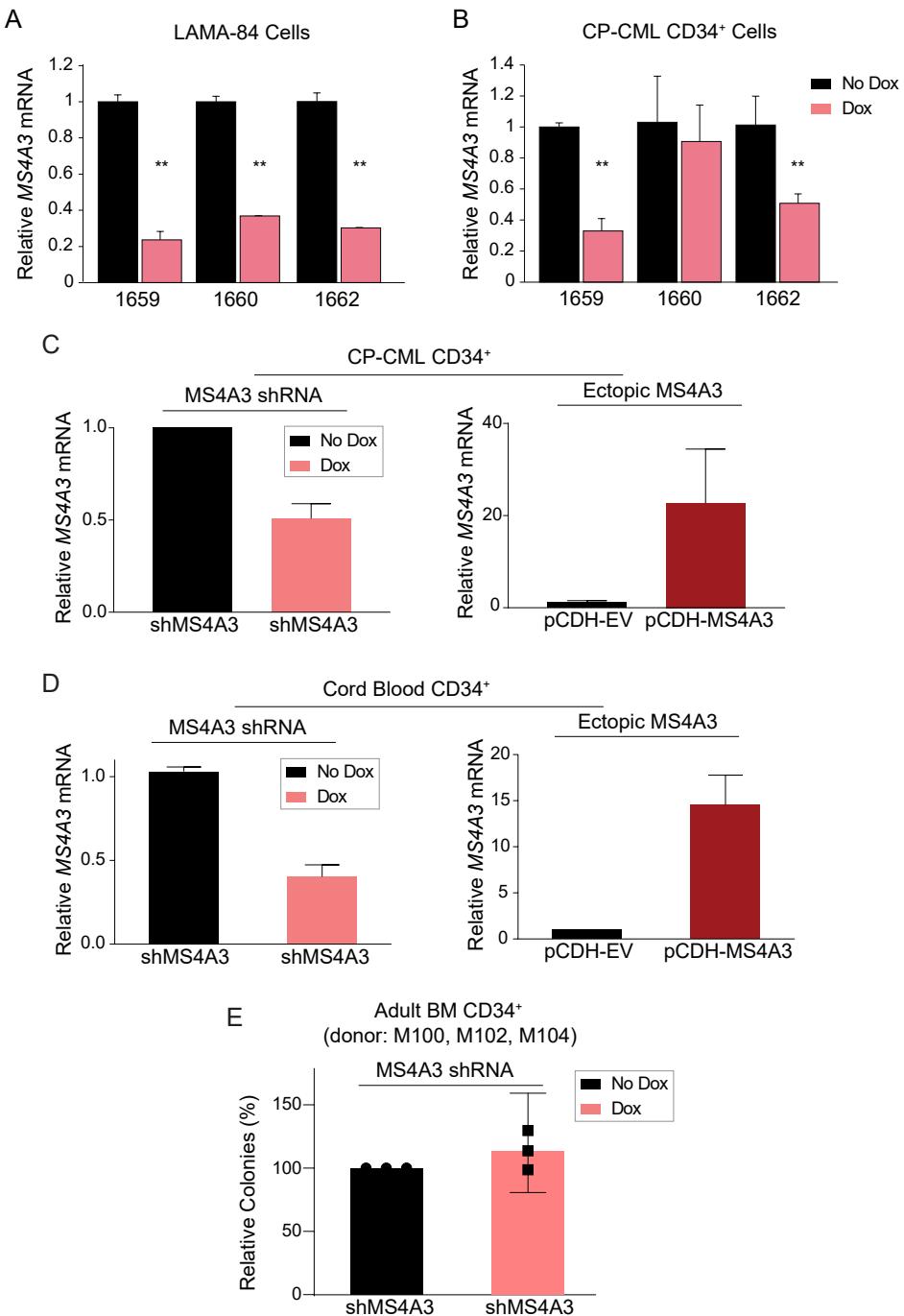
Supplemental Figure 1



Supplemental Figure 2



Supplemental Figure 3



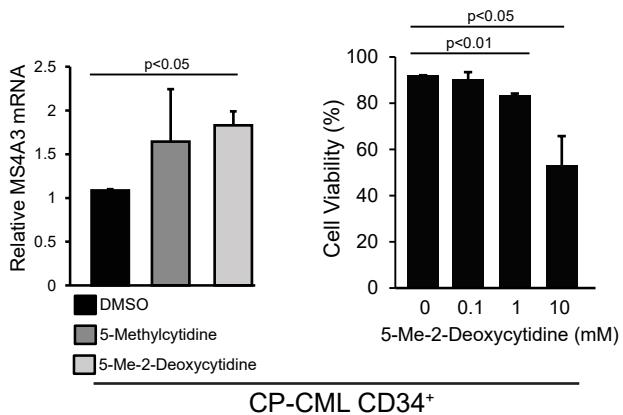
Supplemental Figure 4

A

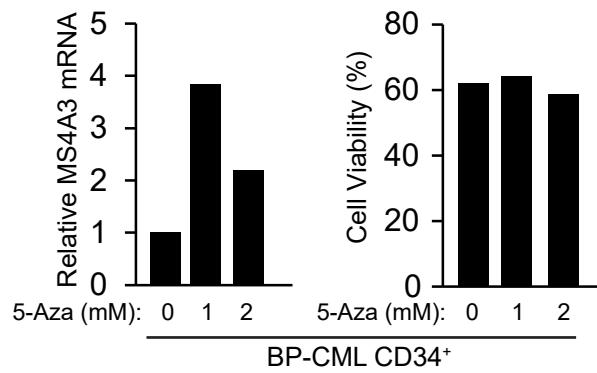
Co-expression with *MS4A3*

Correlated Gene	Spearman's Correlation	-LOG10(q-value)
<i>AFF2</i>	0.695	19.640
<i>CTSG</i>	0.687	19.209
<i>CEBPE</i>	0.678	18.606
<i>AZU1</i>	0.662	17.384
<i>ELANE</i>	0.656	16.991
<i>RNASE3</i>	0.622	14.462
<i>NOCT</i>	0.620	14.427
<i>P4HB</i>	0.620	14.427
<i>CILP2</i>	0.613	14.024
<i>HSP90B1</i>	0.610	13.886
<i>HAL</i>	0.606	13.650
<i>SERPINB10</i>	0.597	13.088
<i>LPO</i>	0.592	12.780
<i>CALR</i>	0.591	12.740
<i>P2RY2</i>	0.587	12.553
<i>RAB32</i>	0.587	12.551
<i>SRGN</i>	0.586	12.551
<i>ERLIN1</i>	0.581	12.221
<i>PLPPR3</i>	0.569	11.511
<i>SLC39A11</i>	0.567	11.455
<i>ATP23</i>	0.565	11.362
<i>RNASE2</i>	0.563	11.243
<i>S100P</i>	0.562	11.194
<i>MPO</i>	0.556	10.863

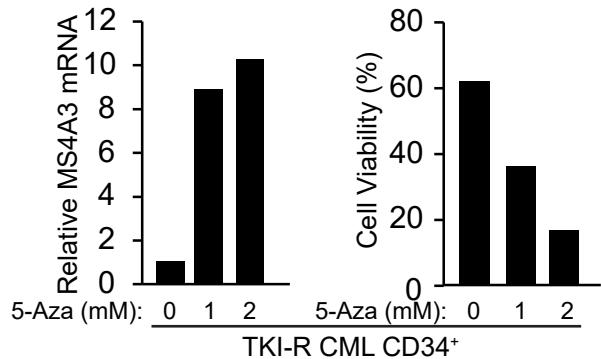
B



C

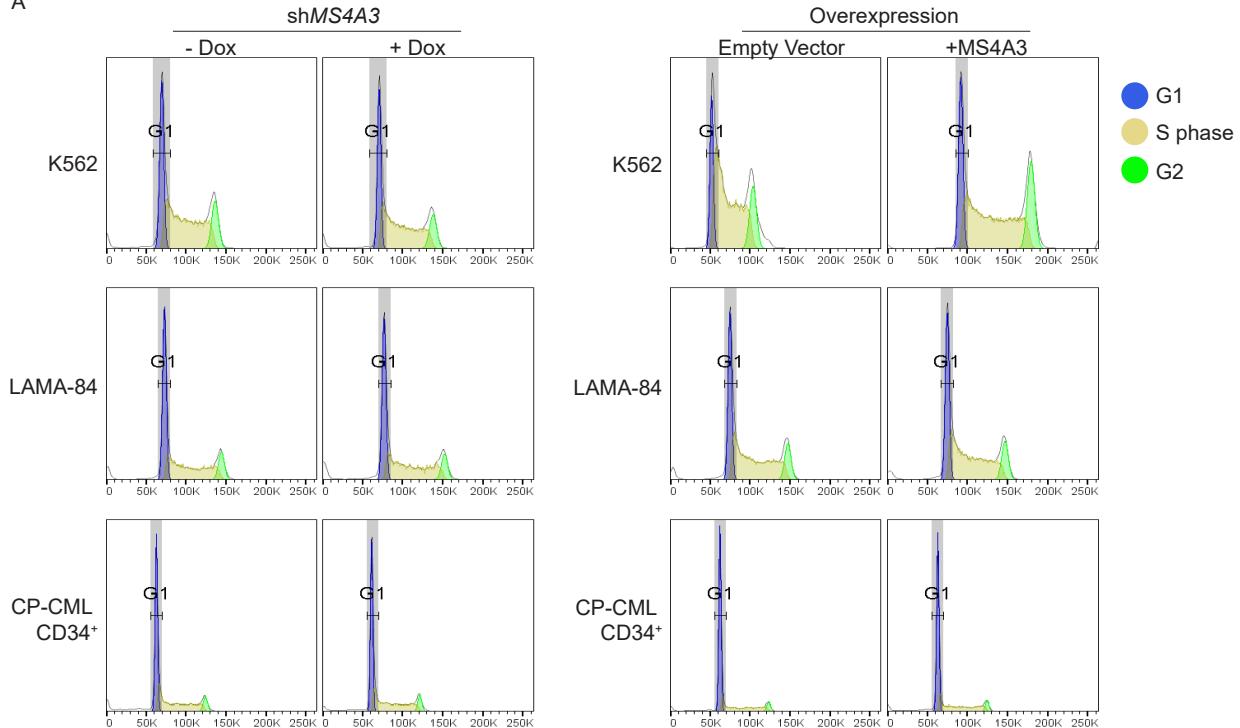


D

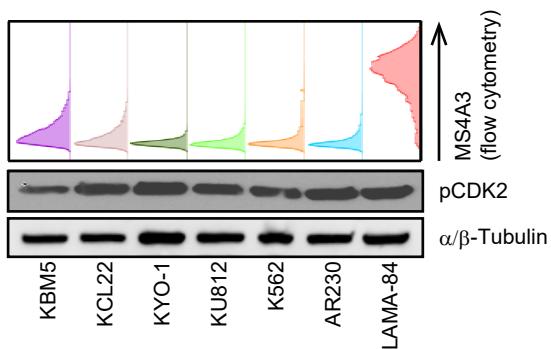


Supplemental Figure 5

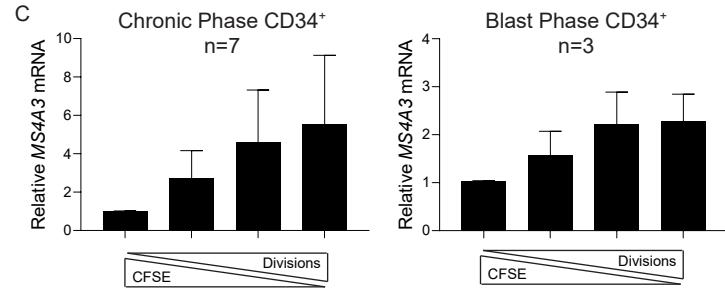
A



B

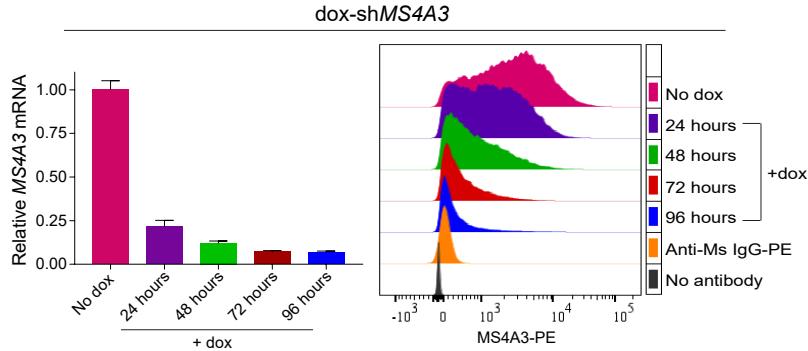
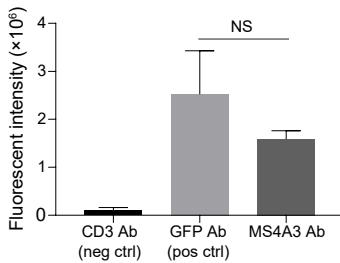


C

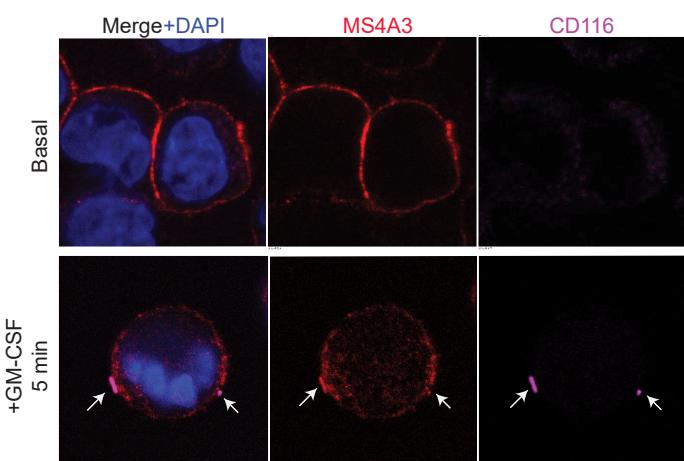


Supplemental Figure 6

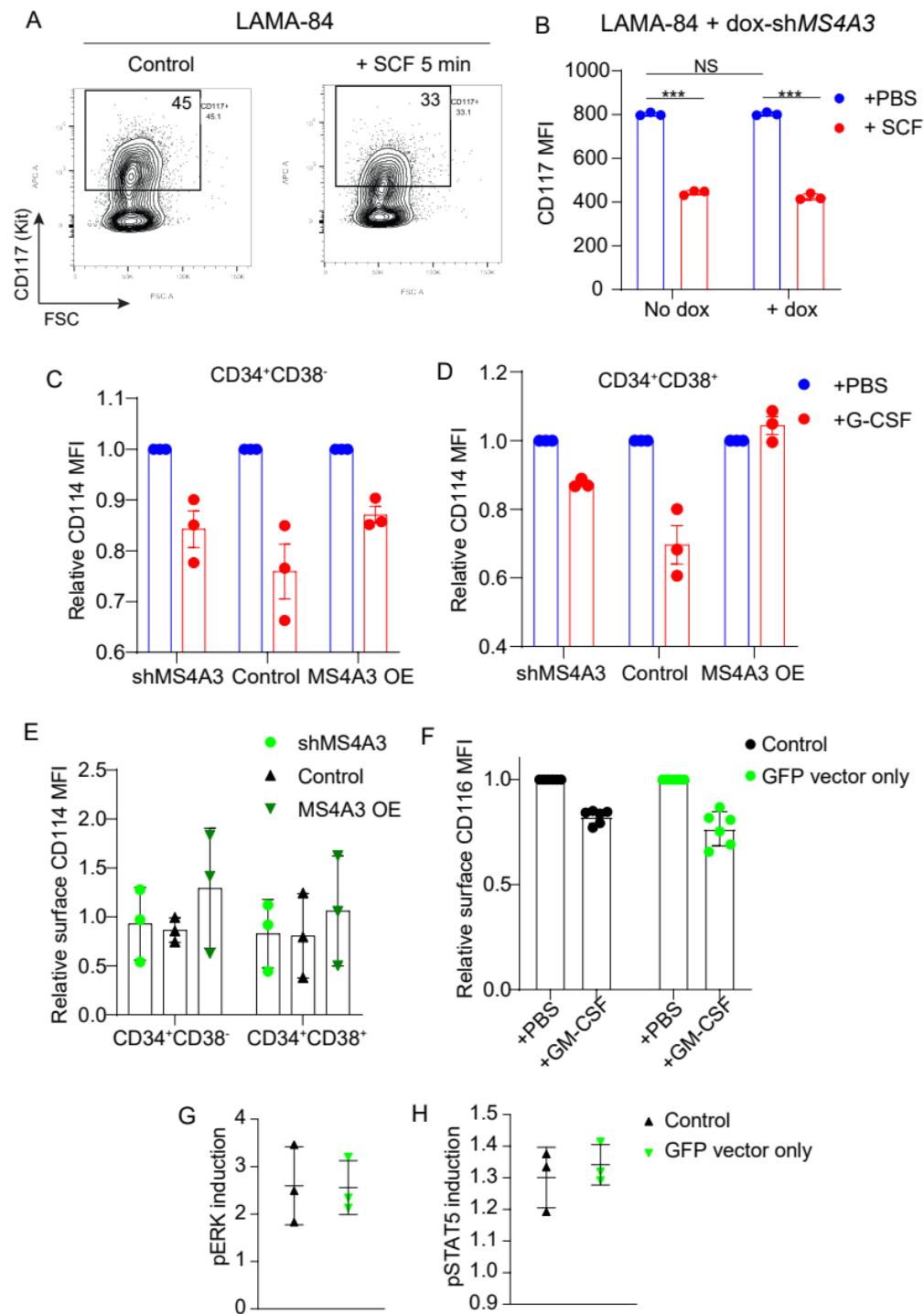
A

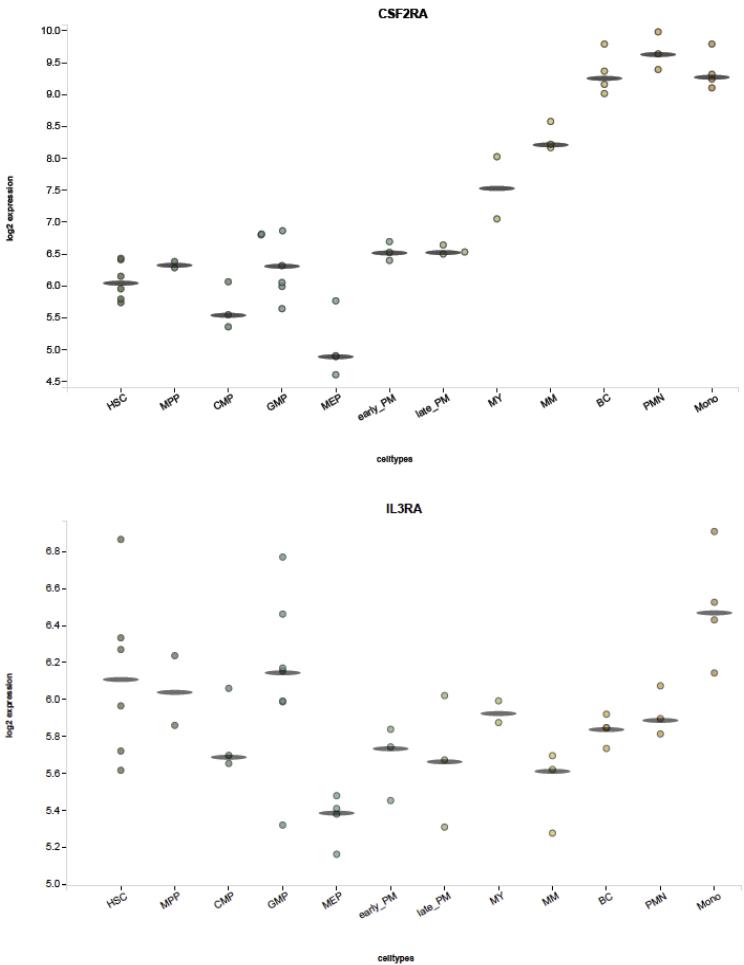
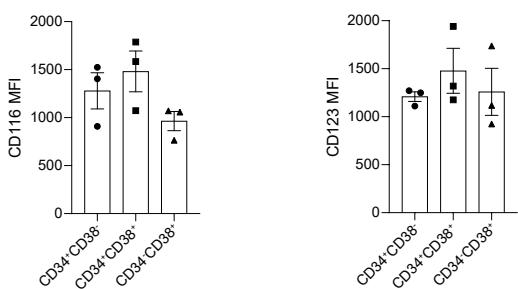
B MS4A3-EGFP pulldown
+ on-bead Ab staining

C



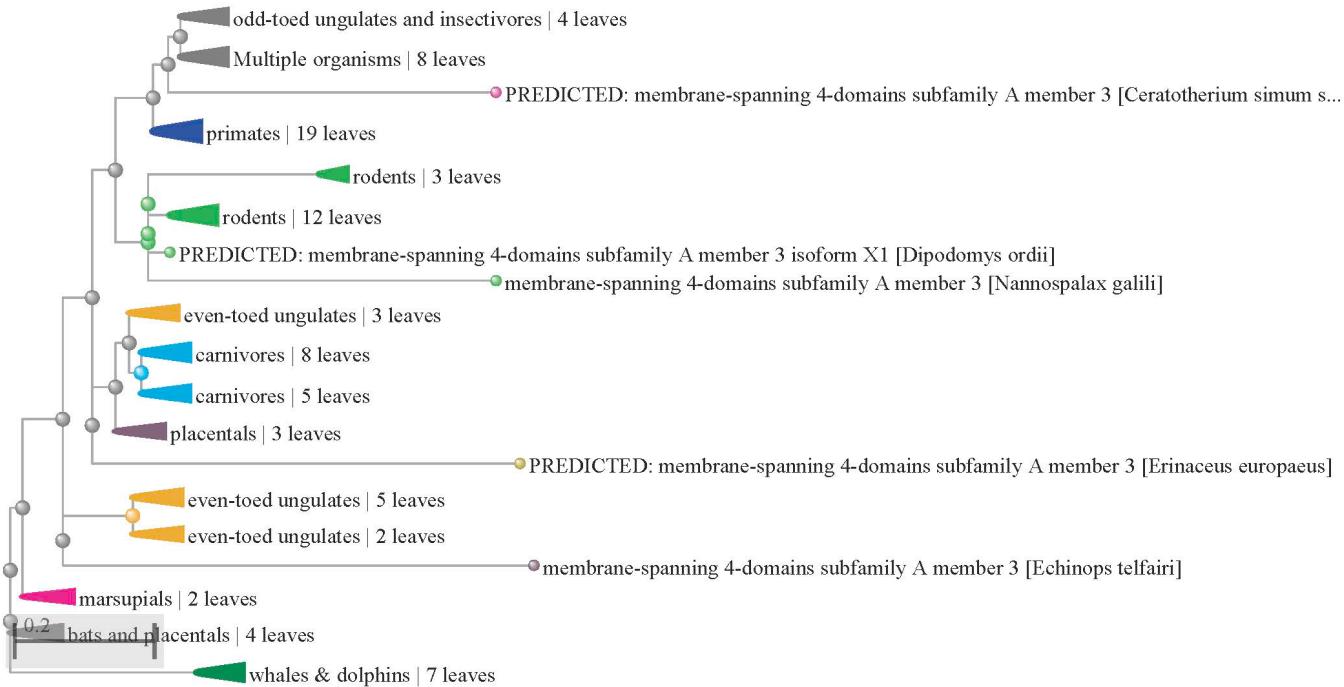
Supplemental Figure 7



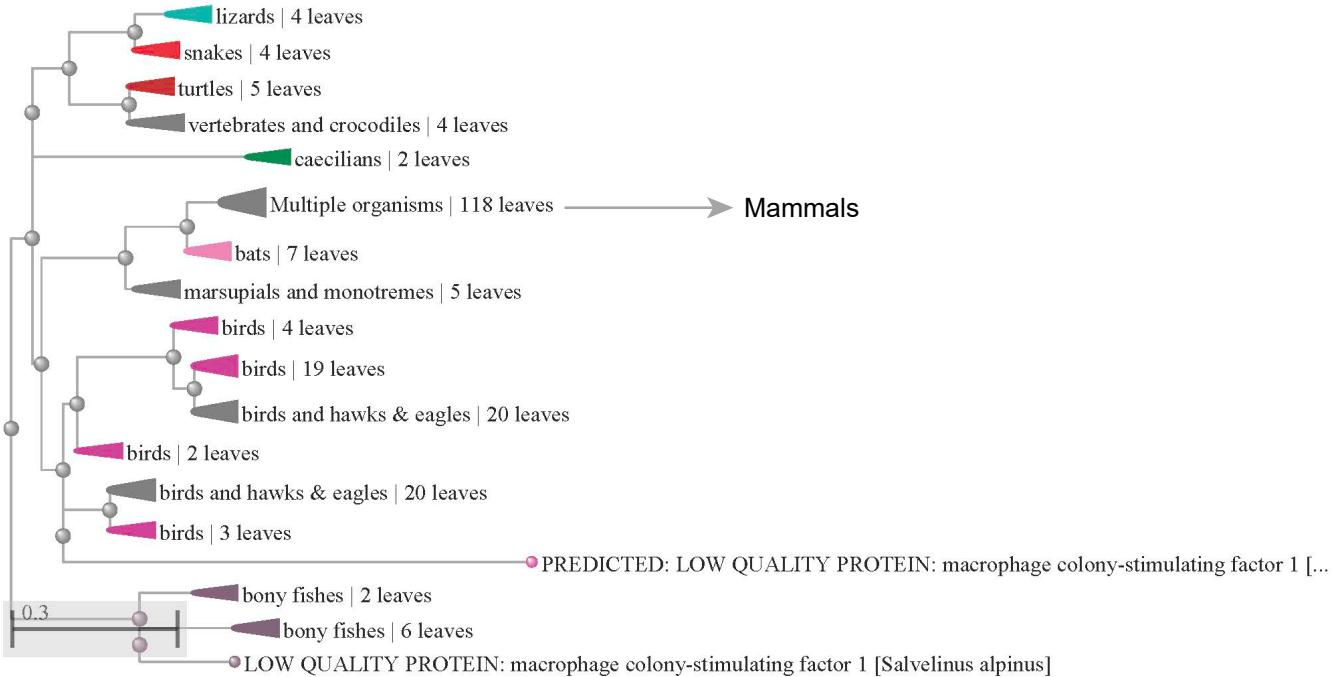
A**B**

Supplemental Figure 9

MS4A3 gene orthologs



M-CSF gene orthologs



Supplementary Table S1. Patient sample summary

Category	Patient ID	CML Disease Phase	BCR-ABL1 Kinase Domain Mutations	Prior TKI Exposure	Type of Resistance	Assays
Newly Diagnosed	M003	CP-CML	None	None	N/A	DNA bisulfite conversion & patch PCR sequencing
	M009	CP-CML	None	None	N/A	qRT-PCR
	M010	CP-CML	None	None	N/A	qRT-PCR
	M011	CP-CML	None	None	N/A	qRT-PCR, shMS4A3, CFSE staining qRT-PCR, shMS4A3, colony assays, immunoblot, LTC-IC, xenografts, CFSE
	M014	CP-CML	None	None	N/A	staining, DNA bisulfite conversion & patch PCR sequencing
	M016	CP-CML	None	None	N/A	qRT-PCR, shMS4A3, colony assays, LTC-IC, xenografts
	M017	CP-CML	None	None	N/A	qRT-PCR, shMS4A3, colony assays
	M018	CP-CML	None	None	N/A	qRT-PCR, shMS4A3, colony assays
	M021	CP-CML	None	None	N/A	Immunoblot, shMS4A3
	M022	CP-CML	None	None	N/A	shMS4A3, colony assays
	M023	CP-CML	None	None	N/A	qRT-PCR, CFSE staining qRT-PCR, shMS4A3, colony assays, DNA bisulfite conversion & patch PCR
	M024	CP-CML	None	None	N/A	sequencing
	M025	CP-CML	None	None	N/A	qRT-PCR, immunoblot, shMS4A3, annexin V, CFSE staining qRT-PCR, immunoblot, shMS4A3, colony assays, annexin V, LTC-IC, xenografts, CFSE staining, CD38 selection, DNA bisulfite conversion & patch PCR
	M028	CP-CML	None	None	N/A	sequencing
	M029	CP-CML	None	None	N/A	Immunoblot
	M030	CP-CML	None	None	N/A	qRT-PCR
	M040	CP-CML	None	None	N/A	qRT-PCR, immunoblot, ectopic MS4A3, shMS4A3, annexin V, CD38 selection
	M041	CP-CML	None	None	N/A	qRT-PCR, immunoblot, shMS4A3
	M043	CP-CML	None	None	N/A	qRT-PCR, shMS4A3, CD38 selection
	M044	CP-CML	None	None	N/A	qRT-PCR, shMS4A3
	M045	CP-CML	None	None	N/A	Immunoblot, CD38 selection, qRT-PCR, shMS4A3
	M046	CP-CML	None	None	N/A	Immunoblot, CD38 selection, qRT-PCR
	M047	CP-CML	None	None	N/A	qRT-PCR, shMS4A3
	M051	CP-CML	None	None	N/A	Immunoblot
	M052	CP-CML	None	None	N/A	Immunoblot
	M055	CP-CML	None	None	N/A	qRT-PCR, differential sorting
	M059	CP-CML	None	None	N/A	qRT-PCR, ectopic MS4A3, CD38 selection, CFSE staining
	M060	CP-CML	None	None	N/A	qRT-PCR, differential sorting
	M066	CP-CML	None	None	N/A	qRT-PCR, CFSE staining
	M071	CP-CML	None	None	N/A	qRT-PCR, DNA bisulfite conversion & patch PCR sequencing
	M073	CP-CML	None	None	N/A	qRT-PCR, shMS4A3
	M074	CP-CML	None	None	N/A	qRT-PCR, differential sorting
	M078	CP-CML	None	None	N/A	qRT-PCR, shMS4A3
AP/BP-CML	M019	BP-CML	None		N/A	qRT-PCR, immunoblot, ectopic MS4A3, shMS4A3, annexin V, colony assays, CFSE staining
	M020	BP-CML	None		N/A	qRT-PCR, immunoblot, ectopic MS4A3, annexin V, colony assays
	M027	AP-CML	None		N/A	qRT-PCR, ectopic MS4A3, annexin V, colony assays
	M033	BP-CML	None		N/A	qRT-PCR, immunoblot, ectopic MS4A3, shMS4A3, annexin V, colony assays, CFSE staining, DNA bisulfite conversion & patch PCR sequencing
	M039	BP-CML	None		N/A	qRT-PCR, immunoblot, ectopic MS4A3, annexin V, colony assays, DNA bisulfite conversion & patch PCR sequencing
	M042	AP-CML	None		N/A	qRT-PCR, ectopic MS4A3, annexin V, DNA bisulfite conversion & patch PCR sequencing
	M049	BP-CML	None		N/A	qRT-PCR, ectopic MS4A3, annexin V, 5-aza treatment
	M050	BP-CML	None		N/A	qRT-PCR, ectopic MS4A3, annexin V, 5-aza treatment
	M065	BP-CML	None		N/A	qRT-PCR, CFSE staining, differential sorting
TKI-R CML	M001	TKI-R (CP-CML)	None	IM, NIL, DAS	Cytogenetic	qRT-PCR
	M002	TKI-R (AP-CML)	None	NIL	Cytogenetic	qRT-PCR, DNA bisulfite conversion & patch PCR sequencing
	M004	TKI-R (CP-CML)	None	IM, NIL, DAS	Cytogenetic	qRT-PCR
	M005	TKI-R (CP-CML)	E255K	IM, NIL, DAS	Cytogenetic	qRT-PCR, DNA bisulfite conversion & patch PCR sequencing
	M006	TKI-R (CP-CML)	None	IM, NIL, DAS	Cytogenetic	qRT-PCR
	M007	TKI-R (AP-CML)	None	IM, DAS	Cytogenetic	qRT-PCR, DNA bisulfite conversion & patch PCR sequencing
	M008	TKI-R	None		Cytogenetic	qRT-PCR
	M012	TKI-R	None		Cytogenetic	qRT-PCR, DNA bisulfite conversion & patch PCR sequencing
	M013	TKI-R (CP-CML)	None	NIL, DAS	Cytogenetic	qRT-PCR, DNA bisulfite conversion & patch PCR sequencing
	M015	TKI-R (CP-CML)	T315I	IM, DAS	Cytogenetic	qRT-PCR, DNA bisulfite conversion & patch PCR sequencing
Healthy Donors	M026	CB	N/A	N/A		qRT-PCR, immunoblot, ectopic MS4A3, shMS4A3, colony assays
	M031	CB	N/A	N/A		qRT-PCR, CFSE staining
	M032	CB	N/A	N/A		qRT-PCR
	M034	CB	N/A	N/A		qRT-PCR, CFSE staining

M035	CB	N/A	N/A	immunoblot
M036	CB	N/A	N/A	immunoblot
M037	CB	N/A	N/A	qRT-PCR, DNA bisulfite conversion & patch PCR sequencing qRT-PCR, immunoblot, ectopic MS4A3, shMS4A3, colony assays, DNA bisulfite
M038	CB	N/A	N/A	conversion & patch PCR sequencing
M048	CB	N/A	N/A	immunoblot
M053	FH	N/A	N/A	qRT-PCR
M054	FH	N/A	N/A	qRT-PCR, CFSE staining
M056	CB	N/A	N/A	qRT-PCR, differential sorting
M057	CB	N/A	N/A	qRT-PCR, CD38 selection
M058	CB	N/A	N/A	qRT-PCR, CD38 selection
M061	CB	N/A	N/A	qRT-PCR, differential sorting
M062	CB	N/A	N/A	qRT-PCR, immunoblot, ectopic MS4A3, colony assays, CFSE staining
M063	CB	N/A	N/A	qRT-PCR, immunoblot, CFSE staining
M067	CB	N/A	N/A	qRT-PCR, CD38 selection
M068	CB	N/A	N/A	qRT-PCR, CD38 selection
M069	CB	N/A	N/A	qRT-PCR, CD38 selection
M070	CB	N/A	N/A	qRT-PCR, DNA bisulfite conversion & patch PCR sequencing
M072	CB	N/A	N/A	qRT-PCR
M075	CB	N/A	N/A	qRT-PCR, CD38 selection
M076	CB	N/A	N/A	qRT-PCR, shMS4A3, CD38 selection
M077	CB	N/A	N/A	qRT-PCR, CD38 selection

Supplementary Table S2. Primer sequences

Gene	Forward Primer	Reverse Primer
Human		
BCR-ABL1	GGT ACC AGG AGT GTT TCT CCA	GAG CGT GCA GAG TGG AGG GA
CEBPB	GACAAGCACAGCGACGAGTA	AGCTGCTCCACCTTCTTCTG
CEBPE	GCAAGAAGGCAGTGAACAAAG	CTCTGCCATGTACTCCAGCA
CSTA	ATACCTGGAGGCTTATCTG	CCAGTAAGTACCAAGTCCTCATT
CTSG	CCTGCTGTGTAACAATGTGG	AAGGCTCTGGCAACACTGTG
CXCR4	GGTGGTCTATGTTGGCGTCT	TGGAGTGTGACAGCTGGAG
EGR1	GGGAGAGGCAGGAAAGACATAA	TCTGAGATCTTCATCTGACCTAAGA
G0S2	CGTGCCACTAAGGTATTCC	GCACGTACAGCTTCACCAC
GAS2	GAAGGTTGGCCTCCACAA	GAAGGAGAAGGGGCAGAAAG
GUS	GAAAATATGTGGTGGAGAGCTATT	CCGAGTGAAGATCCCCTTTTA
MECOM	ACCAGCCCCTGGATCTAAGT	TTCGACGTTGCTTCCTTTT
MS4A3	AGCAGGGATAAAACCCACAA	AATTGCATAGGTCCGGTGAC
PLAUR	GACCCTGAGCTATCGGACTG	CATCCAGGCACTGTTCTCA
PRTN3	ACGCGGAGAACAAACTGAAC	GGGACGAAAGTGCAAATGTT
PTPRe	AGCACCAAGCGACAAGAAGAT	ATGTGTCCAGATGGCAATGA
RNase3	AGGTGAACTGGAACCACAGG	AGATTCCGGGTGCCTTACT
Mouse		
Ms4a3	TGCTGAGGAATTGCACAAAG	ACCAGTCTCCTCTGGCTTCA
Gapdh	GGCATTGCTCTCAATGACAA	TGTGAGGGAGATGCTCAGTG

Supplementary Table S3. Antibodies

Protein	Source	Catalog #
<i>Immunoblot</i>		
Rabbit anti-MS4A3	Aviva Systems Biology	ARP62012-P050
Rabbit anti- α/β -tubulin	Cell Signaling Technology	2148
Rabbit anti-pCDK2 (Thr160)	Cell Signaling Technology	2561S
<i>Immunofluorescent staining</i>		
Rabbit anti-Rab4	Invitrogen	PA3-912
Rabbit anti-Rab5	Cell Signaling Technology	3547
Rabbit anti-Rab7	Cell Signaling Technology	9367
Rabbit anti-Rab11	Cell Signaling Technology	5589
Rabbit anti-LAMP1	Cell Signaling Technology	9091
Rabbit anti-LC3B	Cell Signaling Technology	3868
Rabbit anti-Clathrin	Cell Signaling Technology	4796
Rabbit anti-Caveolin	Cell Signaling Technology	3267
Rabbit anti-58K Golgi protein	Invitrogen	PA5-83166
Rabbit anti-alpha tubulin	Cell Signaling Technology	2125
Alexa Fluor™ 594 Phalloidin	Invitrogen	A12381
Goat anti-Rabbit IgG (H+L) Highly Cross-Adsorbed Secondary Antibody, Alexa Fluor 647	Invitrogen	A-21245
<i>Flow Cytometry</i>		
APC anti-human CD45	BioLegend	368512
FITC anti-human CD34	eBioscience	11-0341-81
APC anti-human CD38	BioLegend	102712
BV421 anti-human CD117	BD Biosciences	562435
APC anti-human CD11b	eBioscience	17-0112-82
<i>ChIP</i>		
Rabbit anti-Histone H3K27me3	Active Motif	39155

Supplementary Table S4. Patch Oligonucleotide Sequences**Patch_L1**

Well Position	Name	Sequence
A1	chr1:151958613-151958784	ggctacttaacaaggaggacctaACTCCCCACCTTCCTCATTCTCT
A2	chr1:151962717-151962873	acatattttcccttgcgtatgaaatgtgACTCCCCACCTTCCTCATTCTCT
A3	chr1:151966081-151966153	gtgggaatccgcgtcaACTCCCCACCTTCCTCATTCTCT
A4	chr1:151966154-151966313	tgcggggaaaaccctgACTCCCCACCTTCCTCATTCTCT
A5	chr1:151966314-151966577	caccggcgcacgtactaaACTCCCCACCTTCCTCATTCTCT
A6	chr1:153329674-153329767	ggcaattctgtatgataccttccttgACTCCCCACCTTCCTCATTCTCT
A7	chr1:153329926-153330006	ccttttagtccaacaaggatggctgaACTCCCCACCTTCCTCATTCTCT
A8	chr1:153330088-153330317	tgggtgttagttcatttcccttatcccaaACTCCCCACCTTCCTCATTCTCT
A9	chr1:153330318-153330500	acagagtgttgcagagctgtgACTCCCCACCTTCCTCATTCTCT
A10	chr1:153330768-153330894	gcgttccagctgcacatttgACTCCCCACCTTCCTCATTCTCT
A11	chr1:153348337-153348415	cctggccctggctcaACTCCCCACCTTCCTCATTCTCT
A12	chr1:153362881-153363077	aattgctagagaccgagtgtctcaACTCCCCACCTTCCTCATTCTCT
B1	chr1:153363139-153363248	tgctgggttgttatttggagatgtgACTCCCCACCTTCCTCATTCTCT
B2	chr1:153363438-153363550	gaagtggagcagcctcctgaACTCCCCACCTTCCTCATTCTCT
B3	chr1:153363557-153363691	cgctataaaaaggagtcgcctctcaACTCCCCACCTTCCTCATTCTCT
B4	chr1:161184287-161184369	ggagacgagcaggaaaaattctaaACTCCCCACCTTCCTCATTCTCT
B5	chr1:161184396-161184474	gttaccagaatgtgggactgtaACTCCCCACCTTCCTCATTCTCT
B6	chr1:161184475-161184585	ctgcccgtcaacatcccctcaACTCCCCACCTTCCTCATTCTCT
B7	chr1:161184586-161184671	gctggctctgaacctcgtatctcaACTCCCCACCTTCCTCATTCTCT
B8	chr1:161184987-161185063	ggcccccctccctcctgACTCCCCACCTTCCTCATTCTCT
B9	chr1:161185076-161185223	gccgtctgacagcactgtgACTCCCCACCTTCCTCATTCTCT
B10	chr1:161186799-161186880	cgaaaaccaaggctggaaaactaaACTCCCCACCTTCCTCATTCTCT
B11	chr1:161188811-161188906	gccagctgggttaatggcatgACTCCCCACCTTCCTCATTCTCT
B12	chr1:186642099-186642206	actctgcctatatttcttacacttgcacttttgACTCCCCACCTTCCTCATTCTCT
C1	chr1:186650355-186650456	gctcacattaactatttacaggtaactgttcaACTCCCCACCTTCCTCATTCTCT
C2	chr1:209847625-209847712	ttagctgtaatcacctgggctgaACTCCCCACCTTCCTCATTCTCT
C3	chr1:209848398-209848506	cttcctactgggtcagcgggtgACTCCCCACCTTCCTCATTCTCT
C4	chr1:209848507-209848581	cgtggaaaggccagtgtgACTCCCCACCTTCCTCATTCTCT
C5	chr1:209848582-209848716	gtggccacgcgtgaACTCCCCACCTTCCTCATTCTCT
C6	chr1:209848717-209848789	ggggccgcttatatctttctctgaACTCCCCACCTTCCTCATTCTCT
C7	chr1:209848891-209848994	gctcatagaaggccgactacctaACTCCCCACCTTCCTCATTCTCT
C8	chr2:64677694-64677801	ccgtggtaatggagataggtaaaatgACTCCCCACCTTCCTCATTCTCT
C9	chr2:64680293-64680366	gtggcccaggctggagtgACTCCCCACCTTCCTCATTCTCT
C10	chr2:64683514-64683698	cctctcccagatatacaagaattctgaACTCCCCACCTTCCTCATTCTCT
C11	chr2:64685929-64686047	gttgtgtctacacaattccactttatgtgACTCCCCACCTTCCTCATTCTCT
C12	chr2:113587642-113587785	aaaccacggccacattgggtctaaACTCCCCACCTTCCTCATTCTCT
D1	chr2:113593790-113593883	cagccatggcagaagtacctgaACTCCCCACCTTCCTCATTCTCT
D2	chr2:113594250-113594412	tgactttaatctcccttacaacttaggtctaaACTCCCCACCTTCCTCATTCTCT
D3	chr2:113594413-113594489	tgaatcaggattcaacagagaaaattctcaACTCCCCACCTTCCTCATTCTCT
D4	chr2:113594618-113594718	aaaaatccagttttatgtggacatcaactgACTCCCCACCTTCCTCATTCTCT
D5	chr2:136872117-136872217	gctagaaaatgtccccagctgtttatgACTCCCCACCTTCCTCATTCTCT
D6	chr2:136873290-136873443	ggtcgtgggttaccagaagaaactgtaACTCCCCACCTTCCTCATTCTCT
D7	chr2:136873749-136873819	ccaagatgtgacttggaaaccctcaACTCCCCACCTTCCTCATTCTCT
D8	chr2:136874486-136874584	cccaagccggaggcccctgACTCCCCACCTTCCTCATTCTCT
D9	chr2:136874585-136874693	ttgctaggagttttgggttcctgACTCCCCACCTTCCTCATTCTCT
D10	chr3:122044160-122044233	tggcgggttggctcaACTCCCCACCTTCCTCATTCTCT
D11	chr3:122058093-122058177	ctgcgaagttagatttacccatgtgACTCCCCACCTTCCTCATTCTCT
D12	chr4:6695222-6695366	tgctgacacgaggagacatctgaACTCCCCACCTTCCTCATTCTCT

E1	chr4:6695367-6695479	gtctcgccatcactgaACTCCCCACCTTCCTCATTCTCT
E2	chr4:6698030-6698255	gcatggagcaagcaccctgaACTCCCCACCTTCCTCATTCTCT
E3	chr4:90815871-90815966	gctgaagggtctgcacaaagctaaACTCCCCACCTTCCTCATTCTCT
E4	chr4:90816031-90816118	gctgcttccacactgaatctgtaaACTCCCCACCTTCCTCATTCTCT
E5	chr4:90816279-90816377	gaccggagtggcgtatgtACTCCCCACCTTCCTCATTCTCT
E6	chr4:90823160-90823235	tgtcaggcttcccagcgtgACTCCCCACCTTCCTCATTCTCT
E7	chr4:90836562-90836635	gctgatcccaatatacaatcgaaactgACTCCCCACCTTCCTCATTCTCT
E8	chr4:156549424-156549498	cttctgactggcccttcttgACTCCCCACCTTCCTCATTCTCT
E9	chr4:156585604-156585709	tgaatcttctaaattcagggttttagaacctgACTCCCCACCTTCCTCATTCTCT
E10	chr4:156587988-156588084	cgcaggaaaagtgttaggaaaactctgACTCCCCACCTTCCTCATTCTCT
E11	chr4:156588085-156588219	cccgctgggtctctaaACTCCCCACCTTCCTCATTCTCT
E12	chr4:156588586-156588666	gaccgcgtgggatgACTCCCCACCTTCCTCATTCTCT
F1	chr4:156589214-156589300	acgaaaacagaaaatggactgtaaACTCCCCACCTTCCTCATTCTCT
F2	chr4:156589354-156589437	agcctgaaatcacatgaccacattgACTCCCCACCTTCCTCATTCTCT
F3	chr5:88017774-88017907	catatthaattaaataaataatctggctgACTCCCCACCTTCCTCATTCTCT
F4	chr5:88017908-88018019	catccaaagaatgctcacaaacctgACTCCCCACCTTCCTCATTCTCT
F5	chr5:88018385-88018459	ttagattattacttagttttttttcttgACTCCCCACCTTCCTCATTCTCT
F6	chr5:88018460-88018715	ggacgaaaggaaagtccctaACTCCCCACCTTCCTCATTCTCT
F7	chr6:130686016-130686122	gggtggggagagccatgACTCCCCACCTTCCTCATTCTCT
F8	chr6:130686123-130686195	gtgatagaaggcaagttatctgttactcaACTCCCCACCTTCCTCATTCTCT
F9	chr6:130690524-130690712	aaccttatgtctctctgttactcaACTCCCCACCTTCCTCATTCTCT
F10	chr6:130747455-130747602	gtctgtttcattttcgtctgACTCCCCACCTTCCTCATTCTCT
F11	chr6:130757713-130757869	tctagaatgagtgaaatctacaggctgACTCCCCACCTTCCTCATTCTCT
F12	chr6:130758118-130758262	atggggatgctaactggcaggctgACTCCCCACCTTCCTCATTCTCT
G1	chr6:132987332-132987408	ggtcttgtcatggattcaaaacactgaACTCCCCACCTTCCTCATTCTCT
G2	chr6:132987409-132987478	ggcaagacaagtcataattgttattgtACTCCCCACCTTCCTCATTCTCT
G3	chr6:132987492-132987573	ggaggaaggtaatctaaaaagagtccctgACTCCCCACCTTCCTCATTCTCT
G4	chr6:133003614-133003784	agaggaaggacaactttacaactgtaaACTCCCCACCTTCCTCATTCTCT
G5	chr6:133004825-133004937	agagacaggagatgacagatgctcaACTCCCCACCTTCCTCATTCTCT
G6	chr6:133035164-133035249	tcattggacttcagcatgactactcaACTCCCCACCTTCCTCATTCTCT
G7	chr6:133035250-133035333	gttttcttaattaaacttcgttagttaaaggtaaACTCCCCACCTTCCTCATTCTCT
G8	chr6:133035334-133035444	acatagagtcttgagttaatctcacaattactgACTCCCCACCTTCCTCATTCTCT
G9	chr7:150211739-150211909	tttcgcttcagggtatagtgatggatgACTCCCCACCTTCCTCATTCTCT
G10	chr7:150323041-150323184	cgatcaattccattgtactgaatcgACTCCCCACCTTCCTCATTCTCT
G11	chr7:150328732-150328802	agggaggtgccttcgtgACTCCCCACCTTCCTCATTCTCT
G12	chr8:30209251-30209362	ccaggctgatctaaactctgaACTCCCCACCTTCCTCATTCTCT
H1	chr8:30209363-30209450	gcaggaagttctgcacacctaACTCCCCACCTTCCTCATTCTCT
H2	chr8:30209451-30209538	cggggcaagaaaataagtaattttttcttgACTCCCCACCTTCCTCATTCTCT
H3	chr8:30209539-30209649	gctgaccagcgttgcgttgACTCCCCACCTTCCTCATTCTCT
H4	chr8:30210276-30210522	gggaccatgttaactgccaacttgACTCCCCACCTTCCTCATTCTCT
H5	chr8:30262205-30262359	cccgagggtggagggttgACTCCCCACCTTCCTCATTCTCT
H6	chr8:30264605-30264707	acgtgactgttattcaggagactgaACTCCCCACCTTCCTCATTCTCT
H7	chr8:30272512-30272647	acaaaggacaccatccatggatgtACTCCCCACCTTCCTCATTCTCT
H8	chr8:30279880-30279964	ccccagcacttggaaaggctgaACTCCCCACCTTCCTCATTCTCT
H9	chr8:30279965-30280034	ctgggttaacatagcaagacacttattactaaACTCCCCACCTTCCTCATTCTCT
H10	chr8:48648672-48648902	gcaaatatccgactgactgctctaACTCCCCACCTTCCTCATTCTCT
H11	chr8:48649023-48649130	cgggactttgccttacttcctcaACTCCCCACCTTCCTCATTCTCT
H12	chr8:48649131-48649216	ggggccctgtgaACTCCCCACCTTCCTCATTCTCT

Patch_L2

Well Position	Name	Sequence
A1	chr8:48649309-48649539	gcaggcccgagctactcaACTCCCCACCTTCCTCATTCTCT
A2	chr8:48649623-48649763	agctttctacatcttactcctgttgatgACTCCCCACCTTCCTCATTCTCT
A3	chr11:59822619-59822847	ggcccagtgcagaaaattctgACTCCCCACCTTCCTCATTCTCT
A4	chr11:59824073-59824143	ccacagacttaacgttacgttgcctatgACTCCCCACCTTCCTCATTCTCT
A5	chr11:59824144-59824297	cttgtcgaagtagggagacactcaACTCCCCACCTTCCTCATTCTCT
A6	chr12:69743601-69743702	ccgtgctccatccctaACTCCCCACCTTCCTCATTCTCT
A7	chr13:31256700-31256846	cttgaagatcccccaaaggctcaACTCCCCACCTTCCTCATTCTCT
A8	chr13:31264669-31264779	ctttccctctgtccctcaACTCCCCACCTTCCTCATTCTCT
A9	chr13:31305958-31306085	cagaatccggagccgcgtgaACTCCCCACCTTCCTCATTCTCT
A10	chr13:31306157-31306256	cgcgcacattaagattctggctgACTCCCCACCTTCCTCATTCTCT
A11	chr13:31308427-31308520	caatgccctcggttcagctgaACTCCCCACCTTCCTCATTCTCT
A12	chr13:31308521-31308612	ggcagcaggaggggctgACTCCCCACCTTCCTCATTCTCT
B1	chr14:25042640-25042723	acaattttttcacaggcttagaggctaaACTCCCCACCTTCCTCATTCTCT
B2	chr14:25043403-25043546	ggagagagggggccgtgACTCCCCACCTTCCTCATTCTCT
B3	chr14:25043773-25043883	ggagggcatggatgtactgACTCCCCACCTTCCTCATTCTCT
B4	chr14:25044917-25045008	gcaaagcattctcctcaataacctgaACTCCCCACCTTCCTCATTCTCT
B5	chr14:25045492-25045582	ccacccttccttcctctcaACTCCCCACCTTCCTCATTCTCT
B6	chr14:25045583-25045665	cagtttgtggcaaactcctgaACTCCCCACCTTCCTCATTCTCT
B7	chr14:94854933-94855052	gacagggccctgtccctaACTCCCCACCTTCCTCATTCTCT
B8	chr14:94855161-94855233	cacaggacgctgtggttctgaACTCCCCACCTTCCTCATTCTCT
B9	chr17:53340951-53341040	ggcaaaaggtaaggcaacattctaaACTCCCCACCTTCCTCATTCTCT
B10	chr17:53341041-53341201	gcaagtgccttctgcctgaACTCCCCACCTTCCTCATTCTCT
B11	chr17:53341202-53341275	ccttgtgacccaggctgaACTCCCCACCTTCCTCATTCTCT
B12	chr17:53341569-53341721	gcgcgtggtaagttatttagttgACTCCCCACCTTCCTCATTCTCT
C1	chr17:53341848-53342046	ggcccccggccaagcttaACTCCCCACCTTCCTCATTCTCT
C2	chr17:53342047-53342128	catagtttcaagctggaaataacctgaACTCCCCACCTTCCTCATTCTCT
C3	chr17:53342679-53342771	agaaaaaaaaattacttatcttataaatacatctgACTCCCCACCTTCCTCATTCTCT
C4	chr17:53342910-53343164	gttccagcaggacacctgaACTCCCCACCTTCCTCATTCTCT
C5	chr17:53345556-53345685	ggagcagaagaaggtaacctcttaACTCCCCACCTTCCTCATTCTCT
C6	chr17:56345755-56345894	gttttgagcacttaatgtgtgtgACTCCCCACCTTCCTCATTCTCT
C7	chr17:56349058-56349181	ggcccactcgcctgACTCCCCACCTTCCTCATTCTCT
C8	chr17:56352856-56353022	cgggatccagatgtccctgACTCCCCACCTTCCTCATTCTCT
C9	chr17:56354467-56354732	accagacattttccagtgtttttttttctaaACTCCCCACCTTCCTCATTCTCT
C10	chr17:56355246-56355465	ctgctcccttgacaacctgACTCCCCACCTTCCTCATTCTCT
C11	chr17:56356893-56357035	gcaccatcacggggatgtgACTCCCCACCTTCCTCATTCTCT
C12	chr18:61553980-61554107	cgtgttaagtcccttgccttaACTCCCCACCTTCCTCATTCTCT
D1	chr18:61557698-61557783	gctgtatgtttgttcatagatctgACTCCCCACCTTCCTCATTCTCT
D2	chr19:826530-826639	cgaaatgtccagaacaaggaaatctgACTCCCCACCTTCCTCATTCTCT
D3	chr19:827348-827467	tttgtgcccaggctggagtgACTCCCCACCTTCCTCATTCTCT
D4	chr19:827468-827539	gcaccggggggctcaACTCCCCACCTTCCTCATTCTCT
D5	chr19:827570-827660	cagcacatggccggcatgACTCCCCACCTTCCTCATTCTCT
D6	chr19:827673-827749	ctgggctcccccatacttctgACTCCCCACCTTCCTCATTCTCT
D7	chr19:827750-827824	cagggacagccccactcaACTCCCCACCTTCCTCATTCTCT
D8	chr19:827825-827967	ggggcaggctgtggctaaACTCCCCACCTTCCTCATTCTCT
D9	chr19:839649-839771	ctccctggcaggccctgACTCCCCACCTTCCTCATTCTCT
D10	chr19:840607-840754	gacgttagcaggggacaggatgACTCCCCACCTTCCTCATTCTCT
D11	chr19:840755-840992	caacatggccacagccctaACTCCCCACCTTCCTCATTCTCT
D12	chr19:840993-841063	catgggggtccagggtgACTCCCCACCTTCCTCATTCTCT

E1	chr19:841064-841176	gtggctcactcaccgctcaACTCCCCACCTTCCTCATTCTCT
E2	chr19:44152949-44153103	ctacagacttgctgtgacctcaACTCCCCACCTTCCTCATTCTCT
E3	chr19:44159620-44159775	acaccttccacttcctgaaatgctgACTCCCCACCTTCCTCATTCTCT
E4	chr19:44173928-44174036	gtcgcccaggctggagtgACTCCCCACCTTCCTCATTCTCT
E5	chr20:48805984-48806131	cctctgctggtaggctcaACTCCCCACCTTCCTCATTCTCT

Patch_R1

Well Position	Name	Sequence
A1	chr1:151958613-151958784	AACACCAACTACCCTCCCACACTcataaagtctgccaaaaaaagttaatgttagac
A2	chr1:151962717-151962873	AACACCAACTACCCTCCCACACTgcagttagatgcagcactagaaa
A3	chr1:151966081-151966153	AACACCAACTACCCTCCCACACTcaccggcttcgcg
A4	chr1:151966154-151966313	AACACCAACTACCCTCCCACACTggaaaggcgacagcc
A5	chr1:151966314-151966577	AACACCAACTACCCTCCCACACTgggtgggatgccctg
A6	chr1:153329674-153329767	AACACCAACTACCCTCCCACACTgcaaggtcacatagctggttg
A7	chr1:153329926-153330006	AACACCAACTACCCTCCCACACTggacctatggggcctga
A8	chr1:153330088-153330317	AACACCAACTACCCTCCCACACTcagggctggcatttataggcag
A9	chr1:153330318-153330500	AACACCAACTACCCTCCCACACTgaaaatgtgaaagaagctttaaaggtaggc
A10	chr1:153330768-153330894	AACACCAACTACCCTCCCACACTcagatcttcgcaccagcttt
A11	chr1:153348337-153348415	AACACCAACTACCCTCCCACACTggctgtaaaagttttgtctcttagtct
A12	chr1:153362881-153363077	AACACCAACTACCCTCCCACACTcaaattccctgcccattggattcc
B1	chr1:153363139-153363248	AACACCAACTACCCTCCCACACTcacatgtctgtgtgaatggacc
B2	chr1:153363438-153363550	AACACCAACTACCCTCCCACACTcatgtcttttcgcaggctgttttc
B3	chr1:153363557-153363691	AACACCAACTACCCTCCCACACTcacagtgattgccacattcacct
B4	chr1:161184287-161184369	AACACCAACTACCCTCCCACACTggcggaaagagggaaatggggatag
B5	chr1:161184396-161184474	AACACCAACTACCCTCCCACACTgacccactatagcccttcc
B6	chr1:161184475-161184585	AACACCAACTACCCTCCCACACTggtgtatccatccgcctcg
B7	chr1:161184586-161184671	AACACCAACTACCCTCCCACACTcaccaccacgcccc
B8	chr1:161184987-161185063	AACACCAACTACCCTCCCACACTgcgcgcggatccctgacc
B9	chr1:161185076-161185223	AACACCAACTACCCTCCCACACTgacccaagctctggacttcc
B10	chr1:161186799-161186880	AACACCAACTACCCTCCCACACTcacaaatctaaaactaccctgtgagg
B11	chr1:161188811-161188906	AACACCAACTACCCTCCCACACTgcattagtctttaacaggggaggagg
B12	chr1:186642099-186642206	AACACCAACTACCCTCCCACACTgagacacagtcttcactactcg
C1	chr1:186650355-186650456	AACACCAACTACCCTCCCACACTcaaattctggccatgcgg
C2	chr1:209847625-209847712	AACACCAACTACCCTCCCACACTggtaagaatccagagtgtgagtaactc
C3	chr1:209848398-209848506	AACACCAACTACCCTCCCACACTcacgcgaaggcccactc
C4	chr1:209848507-209848581	AACACCAACTACCCTCCCACACTgcacgcgtccgactgtgt
C5	chr1:209848582-209848716	AACACCAACTACCCTCCCACACTgacacgcctctgg
C6	chr1:209848717-209848789	AACACCAACTACCCTCCCACACTcagctctccagttggagact
C7	chr1:209848891-209848994	AACACCAACTACCCTCCCACACTcaggaggaggcagcaggaga
C8	chr2:64677694-64677801	AACACCAACTACCCTCCCACACTcacatggaaattctaatttgggagaaag
C9	chr2:64680293-64680366	AACACCAACTACCCTCCCACACTgaggctcttttttttttttttttaa
C10	chr2:64683514-64683698	AACACCAACTACCCTCCCACACTcagaattctggctctcaagtgt
C11	chr2:64685929-64686047	AACACCAACTACCCTCCCACACTcaaaaacaaaacaaaacaaaacaaaacaaa
C12	chr2:113587642-113587785	AACACCAACTACCCTCCCACACTgaggatctctgtccatcagcc
D1	chr2:113593790-113593883	AACACCAACTACCCTCCCACACTgagaaatcacatgcacgttagcc
D2	chr2:113594250-113594412	AACACCAACTACCCTCCCACACTgcctctacttctgttttggaaag
D3	chr2:113594413-113594489	AACACCAACTACCCTCCCACACTgaagctccaccaatactttttcc
D4	chr2:113594618-113594718	AACACCAACTACCCTCCCACACTcatgtataaatctgtgtctccacttt
D5	chr2:136872117-136872217	AACACCAACTACCCTCCCACACTgttgctgtatgtctgtggtagga
D6	chr2:136873290-136873443	AACACCAACTACCCTCCCACACTggggactatgactccatgaaggaa
D7	chr2:136873749-136873819	AACACCAACTACCCTCCCACACTggaaaggttttttctccctcttagtg
D8	chr2:136874486-136874584	AACACCAACTACCCTCCCACACTcagccgcgcattc
D9	chr2:136874585-136874693	AACACCAACTACCCTCCCACACTcaagtggggaaaccgtttgg
D10	chr3:122044160-122044233	AACACCAACTACCCTCCCACACTgactttctgaatggcatcaact
D11	chr3:122058093-122058177	AACACCAACTACCCTCCCACACTcatagtgaattataagtgcctaataattccagc
D12	chr4:6695222-6695366	AACACCAACTACCCTCCCACACTgcgcacagtggcatccattc
E1	chr4:6695367-6695479	AACACCAACTACCCTCCCACACTcatcggtgtatgcacgtttctgga
E2	chr4:6698030-6698255	AACACCAACTACCCTCCCACACTcactgctaacttcattggccctc
E3	chr4:90815871-90815966	AACACCAACTACCCTCCCACACTggaaagtccgggtttctgtttctgt

E4	chr4:90816031-90816118	AACACCAACTACCCTCCCACACTgtagttgaaggcaaggttagctatg
E5	chr4:90816279-90816377	AACACCAACTACCCTCCCACACTggggaggcagtgtgattaaagaag
E6	chr4:90823160-90823235	AACACCAACTACCCTCCCACACTaaaaactgtgacttgagcatattccg
E7	chr4:90836562-90836635	AACACCAACTACCCTCCCACACTgaaccagaaaaccttctgtgtgg
E8	chr4:156549424-156549498	AACACCAACTACCCTCCCACACTcatttatctaggtcaggcctccat
E9	chr4:156585604-156585709	AACACCAACTACCCTCCCACACTcacattaaaggtaaaaaggcactgtc
E10	chr4:156587988-156588084	AACACCAACTACCCTCCCACACTgctcccccaggcgctc
E11	chr4:156588085-156588219	AACACCAACTACCCTCCCACACTcaggcccataaggcattcg
E12	chr4:156588586-156588666	AACACCAACTACCCTCCCACACTggagaggagggacaccagc
F1	chr4:156589214-156589300	AACACCAACTACCCTCCCACACTcagacaggcgcggagac
F2	chr4:156589354-156589437	AACACCAACTACCCTCCCACACTaaaaatgaaataaccatgtgaaaatcgt
F3	chr5:88017774-88017907	AACACCAACTACCCTCCCACACTcaggccatattggcgtatggc
F4	chr5:88017908-88018019	AACACCAACTACCCTCCCACACTcaacatataaaaaatgtatagatgcttgaca
F5	chr5:88018385-88018459	AACACCAACTACCCTCCCACACTgtcaagcgcatgcgactttc
F6	chr5:88018460-88018715	AACACCAACTACCCTCCCACACTgagttcaaatctctccctgccttc
F7	chr6:130686016-130686122	AACACCAACTACCCTCCCACACTcagtgatttgttttcggcgtgt
F8	chr6:130686123-130686195	AACACCAACTACCCTCCCACACTcacaagggtgtgtatgttgc
F9	chr6:130690524-130690712	AACACCAACTACCCTCCCACACTgaggaggagtgggcacagaa
F10	chr6:130747455-130747602	AACACCAACTACCCTCCCACACTcagaaaggattttcccttaacctaatttaca
F11	chr6:130757713-130757869	AACACCAACTACCCTCCCACACTcagtgaggattgtatcatgccactact
F12	chr6:130758118-130758262	AACACCAACTACCCTCCCACACTcaatttatcacatctgcgcggatgt
G1	chr6:132987332-132987408	AACACCAACTACCCTCCCACACTcaggtcaaaatttagaatgtcaaccct
G2	chr6:132987409-132987478	AACACCAACTACCCTCCCACACTaaaaatggcccttacaagtgaa
G3	chr6:132987492-132987573	AACACCAACTACCCTCCCACACTcagttatattatctgtggaaaaactgaag
G4	chr6:133003614-133003784	AACACCAACTACCCTCCCACACTgaatgtgactgtatggagacaggg
G5	chr6:133004825-133004937	AACACCAACTACCCTCCCACACTcaaggctctccactccc
G6	chr6:133035164-133035249	AACACCAACTACCCTCCCACACTgttctggacaccacgtgtcttc
G7	chr6:133035250-133035333	AACACCAACTACCCTCCCACACTcaataaggtagggttttttatgtaa
G8	chr6:133035334-133035444	AACACCAACTACCCTCCCACACTcaaagcaataattttatccccaccaccc
G9	chr7:150211739-150211909	AACACCAACTACCCTCCCACACTggctgggtggagccac
G10	chr7:150323041-150323184	AACACCAACTACCCTCCCACACTgaaagcttcagatttccttcgt
G11	chr7:150328732-150328802	AACACCAACTACCCTCCCACACTcacactgtggccaacatgg
G12	chr8:30209251-30209362	AACACCAACTACCCTCCCACACTgctcttgagtagccgggacta
H1	chr8:30209363-30209450	AACACCAACTACCCTCCCACACTcaggtcgccaggctggag
H2	chr8:30209451-30209538	AACACCAACTACCCTCCCACACTcacattccctatggtaaaatttaatttt
H3	chr8:30209539-30209649	AACACCAACTACCCTCCCACACTcaggtcgctgtccccg
H4	chr8:30210276-30210522	AACACCAACTACCCTCCCACACTcacgttaaacatctgtcatcgac
H5	chr8:30262205-30262359	AACACCAACTACCCTCCCACACTggcgaggatgtcgagatggct
H6	chr8:30264605-30264707	AACACCAACTACCCTCCCACACTggggtcaagaccacgcctgg
H7	chr8:30272512-30272647	AACACCAACTACCCTCCCACACTcagtttagatggggataaggg
H8	chr8:30279880-30279964	AACACCAACTACCCTCCCACACTgaaaatttgttattattatagaaaatttc
H9	chr8:30279965-30280034	AACACCAACTACCCTCCCACACTggcgaggactgttcaagc
H10	chr8:48648672-48648902	AACACCAACTACCCTCCCACACTcagtttaagttaaataaaaaggccgc
H11	chr8:48649023-48649130	AACACCAACTACCCTCCCACACTgaacagtcaagagctccctaaact
H12	chr8:48649131-48649216	AACACCAACTACCCTCCCACACTggtaagcttagcgtttccagc

Patch_R2

Well Position	Name	Sequence
A1	chr8:48649309-48649539	AACACCAACTACCCTCCCACACTgcatgctacttttatattaattttacagtagt
A2	chr8:48649623-48649763	AACACCAACTACCCTCCCACACTcagttctggacataggagcg
A3	chr11:59822619-59822847	AACACCAACTACCCTCCCACACTgagaaagaatgagggcacagcc
A4	chr11:59824073-59824143	AACACCAACTACCCTCCCACACTgaaaagatcactacagcatttccc
A5	chr11:59824144-59824297	AACACCAACTACCCTCCCACACTcagacatccccagtagtccactg
A6	chr12:69743601-69743702	AACACCAACTACCCTCCCACACTcaaatacatctgttaacaccaacgaaca
A7	chr13:31256700-31256846	AACACCAACTACCCTCCCACACTcatcagttagttcattatcacacactc
A8	chr13:31264669-31264779	AACACCAACTACCCTCCCACACTcactccgcggctgc
A9	chr13:31305958-31306085	AACACCAACTACCCTCCCACACTgacagaacaaaaagacaagggca
A10	chr13:31306157-31306256	AACACCAACTACCCTCCCACACTcagaggccccgggatg
A11	chr13:31308427-31308520	AACACCAACTACCCTCCCACACTcatctcggcacggaaagagg
A12	chr13:31308521-31308612	AACACCAACTACCCTCCCACACTgctattggattgttaagaggt
B1	chr14:25042640-25042723	AACACCAACTACCCTCCCACACTgttgtttggattgtacaacaaaatagc
B2	chr14:25043403-25043546	AACACCAACTACCCTCCCACACTcagaggataggcagtgcct
B3	chr14:25043773-25043883	AACACCAACTACCCTCCCACACTcaggttaccacctacgtgg
B4	chr14:25044917-25045008	AACACCAACTACCCTCCCACACTgagggctgtggaaagctc
B5	chr14:25045492-25045582	AACACCAACTACCCTCCCACACTgcagcctgggtatgaaac
B6	chr14:25045583-25045665	AACACCAACTACCCTCCCACACTgcagctgctaagagctcac
B7	chr14:94854933-94855052	AACACCAACTACCCTCCCACACTgccccgtttgtcctcc
B8	chr14:94855161-94855233	AACACCAACTACCCTCCCACACTgtggactctccagagactgtc
B9	chr17:53340951-53341040	AACACCAACTACCCTCCCACACTgtcttaggttccataaaaggacac
B10	chr17:53341041-53341201	AACACCAACTACCCTCCCACACTgtcgctgggaaggatgtg
B11	chr17:53341202-53341275	AACACCAACTACCCTCCCACACTcaagaacgaatgggcacaatgaa
B12	chr17:53341569-53341721	AACACCAACTACCCTCCCACACTcagaggggttaacgggtgg
C1	chr17:53341848-53342046	AACACCAACTACCCTCCCACACTggagtttttttttttttttttttttgctgggtgg
C2	chr17:53342047-53342128	AACACCAACTACCCTCCCACACTgatttttttttttttttttttttgctgggtgg
C3	chr17:53342679-53342771	AACACCAACTACCCTCCCACACTgtttccctccgtcccttg
C4	chr17:53342910-53343164	AACACCAACTACCCTCCCACACTggaggggtccctcccc
C5	chr17:53345556-53345685	AACACCAACTACCCTCCCACACTcaaaatcagaattttagggtagatacaag
C6	chr17:56345755-56345894	AACACCAACTACCCTCCCACACTgtggtaagacatgtatggctgt
C7	chr17:56349058-56349181	AACACCAACTACCCTCCCACACTggAACCTGAAATTGGCGAGGAAA
C8	chr17:56352856-56353022	AACACCAACTACCCTCCCACACTcacacccttacttcgggag
C9	chr17:56354467-56354732	AACACCAACTACCCTCCCACACTcggtgtcaaccatattcaaatg
C10	chr17:56355246-56355465	AACACCAACTACCCTCCCACACTcatccgttctccgctcc
C11	chr17:56356893-56357035	AACACCAACTACCCTCCCACACTgtccccatttttttttcc
C12	chr18:61553980-61554107	AACACCAACTACCCTCCCACACTcactccggctgggtgac
D1	chr18:61557698-61557783	AACACCAACTACCCTCCCACACTcatgctcacaccaagggtgt
D2	chr19:826530-826639	AACACCAACTACCCTCCCACACTcactccggctgggtgac
D3	chr19:827348-827467	AACACCAACTACCCTCCCACACTgggcagtggagctggc
D4	chr19:827468-827539	AACACCAACTACCCTCCCACACTgagtcccgactcccatcc
D5	chr19:827570-827660	AACACCAACTACCCTCCCACACTgtttccctctctagaagatggca
D6	chr19:827673-827749	AACACCAACTACCCTCCCACACTgtgggaacacgggaaattgc
D7	chr19:827750-827824	AACACCAACTACCCTCCCACACTggcggcggctgc
D8	chr19:827825-827967	AACACCAACTACCCTCCCACACTgttctggcctggcagc
D9	chr19:839649-839771	AACACCAACTACCCTCCCACACTgagtctttgggtctgtggaaat
D10	chr19:840607-840754	AACACCAACTACCCTCCCACACTgagcaacacatgcccacgt
D11	chr19:840755-840992	AACACCAACTACCCTCCCACACTcacccacggtaagctcc

D12	chr19:840993-841063	AACACCAACTACCCTCCCACACTgcagcaaggccagcag
E1	chr19:841064-841176	AACACCAACTACCCTCCCACACTgttccccagctgtgcc
E2	chr19:44152949-44153103	AACACCAACTACCCTCCCACACTgcctcaccatcacccctgc
E3	chr19:44159620-44159775	AACACCAACTACCCTCCCACACTgtccctcattcacaatctgacatct
E4	chr19:44173928-44174036	AACACCAACTACCCTCCCACACTggaaatattagtggagcaaaggcat
E5	chr20:48805984-48806131	AACACCAACTACCCTCCCACACTcaccccgaggtcaacttttca