Figure S1. RNA quality determined by RIN<sup>e</sup> values. RIN<sup>e</sup> values are obtained from Agilent 2200 Tapestation. (A) Electrophoresis and RIN<sup>e</sup> values of samples. (B) The electropherograms of samples. Peaks of 28S,18S and Lower Marker are indicated.



Table SI. Top 20 list of	genes up- or down-reg	ulated in cSR cells com	pared with par	ent H28 cell line.
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Probe ID <sup>a</sup>	Gene symbol	Description	Fold change <sup>b</sup>	Molecular functions <sup>c</sup>
A_23_P88559	LIPC	Lipase, hepatic	934.8	Apolipoprotein binding heparin binding
A_32_P387648	FLG	Filaggrin	326.6	Calcium ion binding structural molecule activity
A_33_P3331746	ST6GAL2	ST6 beta-galactosamide alpha-2,6-sialyltranferase 2	126.7	Beta-galactoside alpha-2,6-sialyltransferase activity
A 23 P157865	TNC	Tenascin C	112.1	Syndecan binding
A_23_P214821	EDNI	Endothelin 1	100.3	Cytokine activity endothelin A receptor binding endothelin B receptor binding
A_23_P218646	TNFRSF6B	Tumor necrosis factor receptor superfamily, member 6b, decoy	91.9	Receptor activity tumor necrosis factor-activated receptor activity
A_24_P190472	SLPI	Leukocyte peptidase inhibitor	76.8	Endopeptidase inhibitor activity enzyme binding
A_23_P372834	AQP1	Aquaporin 1	76.1	Ammonium transmembrane transporter activity carbon dioxide transmembrane transporter activity glycerol channel activity
A_23_P78782	CAII	Carbonic anhydrase XI	69.4	NOT carbonate dehydratase activity
A_23_P121665	SORCS2	Homo sapiens sortilin-related VPS10 domain containing receptor 2	65.9	Neuropeptide receptor activity
A_23_P258612	ATP8A2	Homo sapiens ATPase, aminophospholipid transporter, class I, type 8A, member 2	61.1	ATP binding magnesium ion binding phospholipid-translocating ATPase activity
A_23_P14892	IGFALS	Homo sapiens insulin-like growth factor binding protein, acid labile subunit	58.5	Insulin-like growth factor binding
A 33 P3397865	TNNT1	Homo sapiens troponin T type 1	54.7	Tropomyosin binding troponin T binding
A_24_P406754	LOXL4	Homo sapiens lysyl oxidase-like 4	51.0	Copper ion binding oxidoreductase activity protein binding
A_23_P150609	IGF2	Homo sapiens insulin-like growth factor 2	48.9	Growth factor activity hormone activity insulin receptor binding
A_33_P3285299	GPRIN2	Homo sapiens G protein regulated inducer of neurite outgrowth	48.8	None
A_33_P3377364	ITGB4	Homo sapiens integrin, beta 4	47.3	G-protein coupled receptor binding contributes_to insulin-like growth factor I binding contributes_to neuregulin binding
A_33_P3314401	CLDN16	Homo sapiens claudin 16	46.1	Identical protein binding magnesium ion transmembrane transporter activity structural molecule activity
A_23_P89431	CCL2	Homo sapiens chemokine (C-C motif) ligand 2	43.5	CCR2 chemokine receptor binding chemokine activity heparin binding
A_23_P250629	PSMB8	Proteasome (prosome, macropain) subunit, beta type, 8	42.0	threonine-type endopeptidase activity
A_33_P3280845	THY1	Thy-1 cell surface antigen	-755.7	GPI anchor binding GTPase activator activity integrin binding
A_33_P3380992	AKR1B15	Aldo-keto reductase family 1, member B15	-562.0	Estradiol 17-beta-dehydrogenase activity oxidoreductase activity, acting on the CH-OH group of donors, NAD or NADP as acceptor
A_23_P93641	AKR1B10	Aldo-keto reductase family 1, member B10	-548.5	Aldo-keto reductase (NADP) activity geranylgeranyl reductase activity indanol dehydrogenase activity
A_23_P77908	SLC47A2	Solute carrier family 47 (multidrug and toxin extrusion), member 2	-171.0	Drug:proton antiporter activity drug transmembrane transporter activity

Table SI. Continued.

Probe ID <sup>a</sup>	Gene symbol	Description	Fold change <sup>b</sup>	Molecular functions <sup>c</sup>
A_23_P29046	CBR1	Carbonyl reductase 1	-143.7	15-hydroxyprostaglandin dehydrogenase (NADP+) activity carbonyl reductase (NADPH) activity oxidoreductase activity, acting on NAD(P)H, quinone or similar
A_23_P134237	RARRES2	Retinoic acid receptor responder (tazarotene induced) 2	-130.9	Receptor binding
A_23_P150768	SLCO2B1	Solute carrier organic anion transporter family, member 2B1	-116.6	Bile acid transmembrane transporter activity organic anion transmembrane transporter activity sodium-independent organic anion transmembrane transporter activity
A_24_P166397	KIAA0319	KIAA0319	-105.1	Protein binding
A_24_P190168	TMEM97	Transmembrane protein 97	-80.0	Protein binding
A_23_P13604	PEBP1	Phosphatidylethanolamine binding protein 1	-76.1	ATP binding enzyme binding phosphatidylethanolamine binding
A_23_P15727	FKBP10	FK506 binding protein 10, 65 kDa	-70.9	FK506 binding calcium ion binding peptidyl-prolyl cis-trans isomerase activity
A_23_P251043	SYNDIG1	Synapse differentiation inducing 1	-68.3	Glutamate receptor binding protein binding protein homodimerization activity
A_23_P452	DDR2	Discoidin domain receptor tyrosine kinase 2	-65.0	ATP binding collagen binding protein tyrosine kinase collagen receptor activity
A_23_P409489	DNTT	DNA nucleotidylexotransferase	-60.3	DNA binding DNA nucleotidylexotransferase activity
A_23_P257971	AKR1C1	Aldo-keto reductase family 1, member C1	-54.0	17-alpha,20-alpha-dihydroxypregn-4-en- 3-one dehydrogenase activity alditol: NADP+ 1-oxidoreductase activity aldo-keto reductase (NADP) activity
A 23 P114185	TSPAN7	Tetraspanin 7	-52.0	None
A 23 P252817	SST	Somatostatin	-50.2	Hormone activity
A_23_P129101	HEXA	Hexosaminidase A (alpha polypeptide)	-48.1	Acetylglucosaminyltransferase activity beta-N-acetylhexosaminidase activity
A 23 P163227	CKMT1A	Creatine kinase, mitochondrial	-46.6	ATP binding creatine kinase activity
A_24_P262127	RRAD	Creatine kinase, mitochondrial	-43.7	GTP binding GTPase activity

<sup>a</sup>Data from Agilent Technologies, Inc.; <sup>b</sup>average expression ratio in cSR-A1 and -B1 cell lines were indicated as fold changes; <sup>c</sup>GO terms registered on each gene in molecular function field. Information was taken from the UniProt database (European Bioinformatics, UK, Swiss Institute of Bioinformatics, Switzerland, Protein Information Resource, USA) (http://www.uniprot.org) and the Entrez Gene database (National Center for Biotechnology Information, USA) (http://www.ncbi.nlm.nih. gov/gene).

Table SII. The enriched GO terms of DEGs (Top 20 list of each categories).

GO term	Count <sup>a</sup>	P-value
Biological process		
Oxidation-reduction process	47	2.07x10 <sup>-7</sup>
Angiogenesis	26	2.60x10 <sup>-7</sup>
Negative regulation of fibroblast growth factor receptor signaling pathway	6	$1.04 \times 10^{-5}$
Positive regulation of cell proliferation	36	2.66x10 <sup>-5</sup>
Cell adhesion	34	5.95x10 <sup>-5</sup>
Negative regulation of cell proliferation	31	8.56x10 <sup>-5</sup>
Daunorubicin metabolic process	5	8.72x10 <sup>-5</sup>
Doxorubicin metabolic process	5	8.72x10-5
Response to wounding	11	1.04x10 <sup>-4</sup>
Retinoid metabolic process	10	$2.24 \times 10^{-4}$
Negative regulation of endopeptidase activity	14	$2.73 \times 10^{-4}$
Negative regulation of cell adhesion	8	2.76x10-4
Regulation of cell growth	11	4.08x10 <sup>-4</sup>
Negative regulation of cell migration	12	$4.57 \times 10^{-4}$
Steroid metabolic process	8	$5.23 \times 10^{-4}$
Positive regulation of ERK1 and ERK2 cascade	17	$5.69 \times 10^{-4}$
Signal transduction	62	$6.42 \times 10^{-4}$
Negative regulation of blood coagulation	5	7.76x10 <sup>-4</sup>
Negative regulation of apoptotic process	31	7.95x10-4
Response to drug	25	1.19x10 <sup>-3</sup>
Molecular function		
Integrin binding	17	6.87x10 <sup>-7</sup>
Receptor binding	32	2.42x10 <sup>-6</sup>
Heparin binding	20	3.82x10 <sup>-6</sup>
Insulin-like growth factor binding	7	2.28x10 <sup>-5</sup>
Aldo-keto reductase (NADP) activity	6	$1.14 \times 10^{-4}$
Ferric iron binding	5	2.60x10 <sup>-4</sup>
Insulin-like growth factor I binding	5	5.80x10 <sup>-4</sup>
Syndecan binding	4	7.80x10 <sup>-4</sup>
Transmembrane receptor protein tyrosine kinase activity	7	$1.87 \times 10^{-3}$
Oxidoreductase activity	17	$1.96 \times 10^{-3}$
Oxidoreductase activity, acting on NAD(P)H, quinone or similar compound as acceptor	4	$2.07 \times 10^{-3}$
Calcium ion binding	41	$2.49 \times 10^{-3}$
Indanol dehydrogenase activity	3	$3.56 \times 10^{-3}$
Serine-type endopeptidase inhibitor activity	10	$6.74 \times 10^{-3}$
Iron ion binding	13	$7.03 \times 10^{-3}$
Fibroblast growth factor binding	5	7.65x10 <sup>-3</sup>
Protease binding	10	8.72x10 <sup>-3</sup>
Signal transducer activity	15	$1.24 \times 10^{-2}$
Electron carrier activity	9	$1.32 \times 10^{-2}$
Microtubule binding	15	$1.40 \times 10^{-2}$
Cellular component		
Extracellular space	99	1.13x10 <sup>-12</sup>
Extracellular exosome	154	8.62x10 <sup>-10</sup>
Extracellular matrix	29	1.64x10 <sup>-6</sup>
Extracellular region	90	3.59x10 <sup>-6</sup>
Cell surface	42	4.23x10 <sup>-6</sup>
Basement membrane	11	5.27x10 <sup>-4</sup>
Plasma membrane	175	7.04x10 <sup>-4</sup>
High-density lipoprotein particle	6	7.50x10 <sup>-4</sup>
Growth cone	13	$1.05 \times 10^{-3}$
Proteinaceous extracellular matrix	21	$1.10 \times 10^{-3}$
Focal adhesion	27	$1.17 \times 10^{-3}$
Apical plasma membrane	22	1.69x10 <sup>-3</sup>
Organelle membrane	10	2.47x10 <sup>-3</sup>
Intracellular membrane-bounded organelle	33	2.94x10 <sup>-3</sup>
Integral component of plasma membrane	67	6.37x10 <sup>-3</sup>
Microtubule associated complex	6	6.41x10 <sup>-3</sup>

## Table SII. Continued.

GO term	Count <sup>a</sup>	P-value
Extracellular vesicle	7	6.94x10 <sup>-3</sup>
Endoplasmic reticulum	43	7.58x10 <sup>-3</sup>
External side of plasma membrane	16	8.26x10 <sup>-3</sup>
Membrane raft	16	9.33x10 <sup>-3</sup>

<sup>a</sup>Number of differentially expressed genes that belong to these terms.

Table SIII. KEGG pathway annotation of differentially expressed genes in cSR cells.

KEGG term	Count <sup>a</sup>	P-value
Metabolism of xenobiotics by cytochrome P450	14	1.55x10 <sup>-6</sup>
Chemical carcinogenesis	13	2.13x10 <sup>-5</sup>
Pentose and glucuronate interconversions	8	2.24x10 <sup>-4</sup>
Ascorbate and aldarate metabolism	7	2.95x10 <sup>-4</sup>
Drug metabolism-cytochrome P450	10	6.05x10 <sup>-4</sup>
Transcriptional misregulation in cancer	16	8.66x10 <sup>-4</sup>
Steroid hormone biosynthesis	9	9.20x10 <sup>-4</sup>
Arachidonic acid metabolism	9	1.44x10 <sup>-3</sup>
Glutathione metabolism	8	1.97x10 <sup>-3</sup>
Proteoglycans in cancer	16	4.82x10 <sup>-3</sup>
Drug metabolism-other enzymes	7	5.33x10 <sup>-3</sup>
Retinol metabolism	8	7.80x10 <sup>-3</sup>
Rheumatoid arthritis	9	1.24x10 <sup>-2</sup>
Porphyrin and chlorophyll metabolism	6	1.57x10 <sup>-2</sup>
Glycerolipid metabolism	7	1.62x10 <sup>-2</sup>
Metabolic pathways	57	2.34x10 <sup>-2</sup>
Leukocyte transendothelial migration	10	2.39x10 <sup>-2</sup>
ECM-receptor interaction	8	3.41x10 <sup>-2</sup>
Gastric acid secretion	7	4.41x10 <sup>-2</sup>

<sup>a</sup>Number of differentially expressed genes that belong to these terms.

Table SIV. ABC transporter genes down-regulated in cSR cells compared with parent H28 cell line.

Probe ID <sup>a</sup>	Gene symbol	Description	Fold change <sup>b</sup>	Molecular functions <sup>c</sup>
A_23_P44569	ABCC2	ATP binding cassette subfamily C member 2	-17.3	ATPase-coupled transmembrane transporter activity organic anion transmembrane transporter activity
A_24_P235429	ABCA1	ATP binding cassette subfamily A member 1	-4.7	ATPase-coupled transmembrane transporter activity lipid transporter activity

<sup>a</sup>Data from Agilent Technologies, Inc.; <sup>b</sup>Average expression ratio in cSR-A1 and -B1 cell lines were indicated as fold changes; <sup>c</sup>GO terms registered on each gene in molecular function field. Information was taken from the UniProt database (European Bioinformatics, UK, Swiss Institute of Bioinformatics, Switzerland, Protein Information Resource, USA) (http://www.uniprot.org) and the Entrez Gene database (National Center for Biotechnology Information, USA) (http://www.ncbi.nlm.nih. gov/gene).