

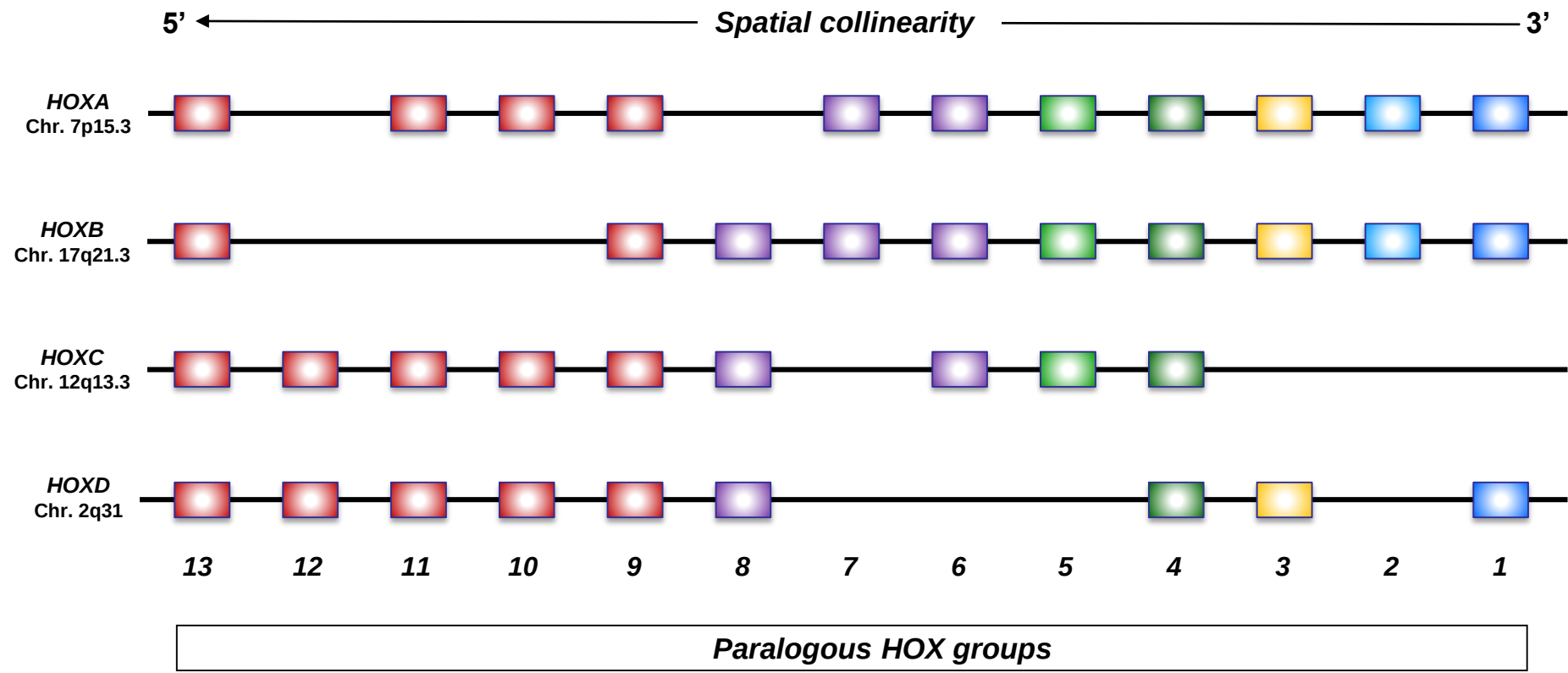
Supplementary file Table S1

Grouping of Homeobox genes according to their main known function.

Anatomical Structure Morphogenesis	<i>EN1, HOXC10, HOXC13, HOXD3, LBX1, SIX2, SIX4</i>
Organ Morphogenesis	<i>CDX1, CDX2, HOXA11, HOXA13, ISL1, LHX1, PAX3, PDHX, PITX2, PITX3, PROX1, SIX6</i>
Body Pattern Formation	<i>ALX3, EMX2, HHEX, HOXA11, HOXA2, HOXA4, HOXA5, HOXA6, HOXB1, HOXB5, HOXB6, HOXC5, HOXD10, HOXD8, LMX1B, PITX2</i>
Ectoderm Development	<i>PROX1, VAX2</i>
Endoderm Development	<i>HOXC11</i>
Brain & Nervous System Development	
Brain Development	<i>ALX1, DLX2, EMX2</i>
Nervous System Development:	<i>ARX, DLX5, DLX6, HOXD10, LBX1, LHX1, OTP, PAX3, PHOX2A, PHOX2B</i>
Skeletal Development:	<i>ALX3, ALX4, DLX3, DLX5, DLX6, EN1, HOXA11, HOXA13, HOXA2, HOXB6, HOXD10, HOXD13, MSX2</i>
Muscle Development:	<i>BARX2, MKX, SIRT1, SIRT2, SIX1</i>
Other Homeobox Genes Involved In Multicellular Organismal Development:	<i>BARX1, CDX4, CUX1, DLX1, EMX1, EN2, HOXA1, HOXA7, HOXA9, HOXB13, HOXB2, HOXB3, HOXB4, HOXB7, HOXB8, HOXB9, HOXC12, HOXC8, HOXC9, HOXD1, HOXD11, HOXD12, HOXD9, ISL2, LBX2, LMX1A, MEIS1, NKX3-1, OTX1, TLX1, VAX1, VSX1, VSX2</i>
Homeobox Genes Involved In Cell Differentiation:	<i>ARX, EMX2, HHEX, HLX, HOPX, LBX1, LHX1, LMX1B, MIXL1, OTP, PHOX2A, SIRT1, VSX2</i>
Other Genes:	<i>PHTF1, SIRT3, SIRT6, SIRT7, ZHX1, ZHX2</i>

Homeobox genes include two subsets of genes coding for transcription factors involved in multiple functions. The clustered HOX genes are indicated in bold.

Supplementary file Figure S2



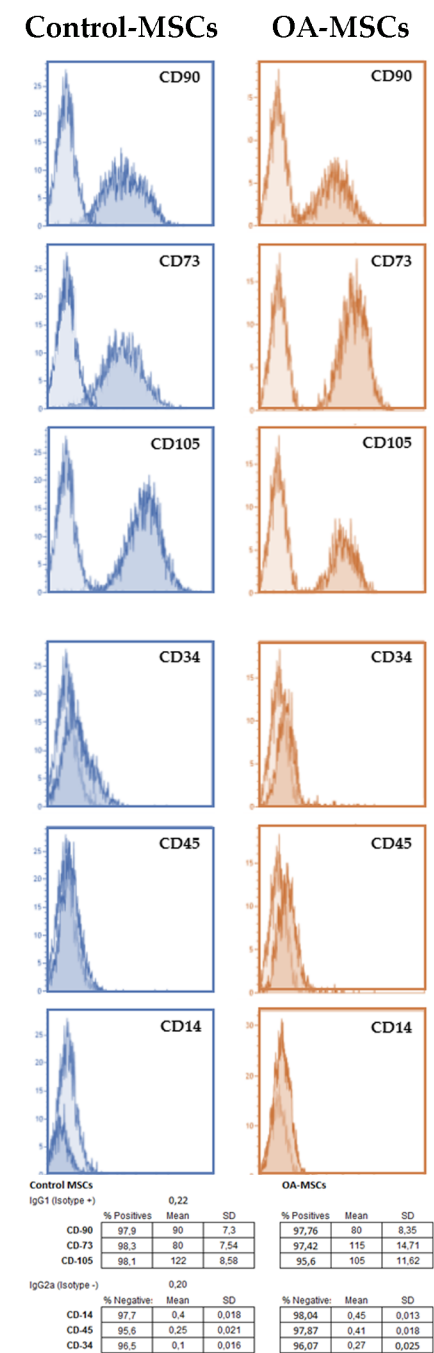
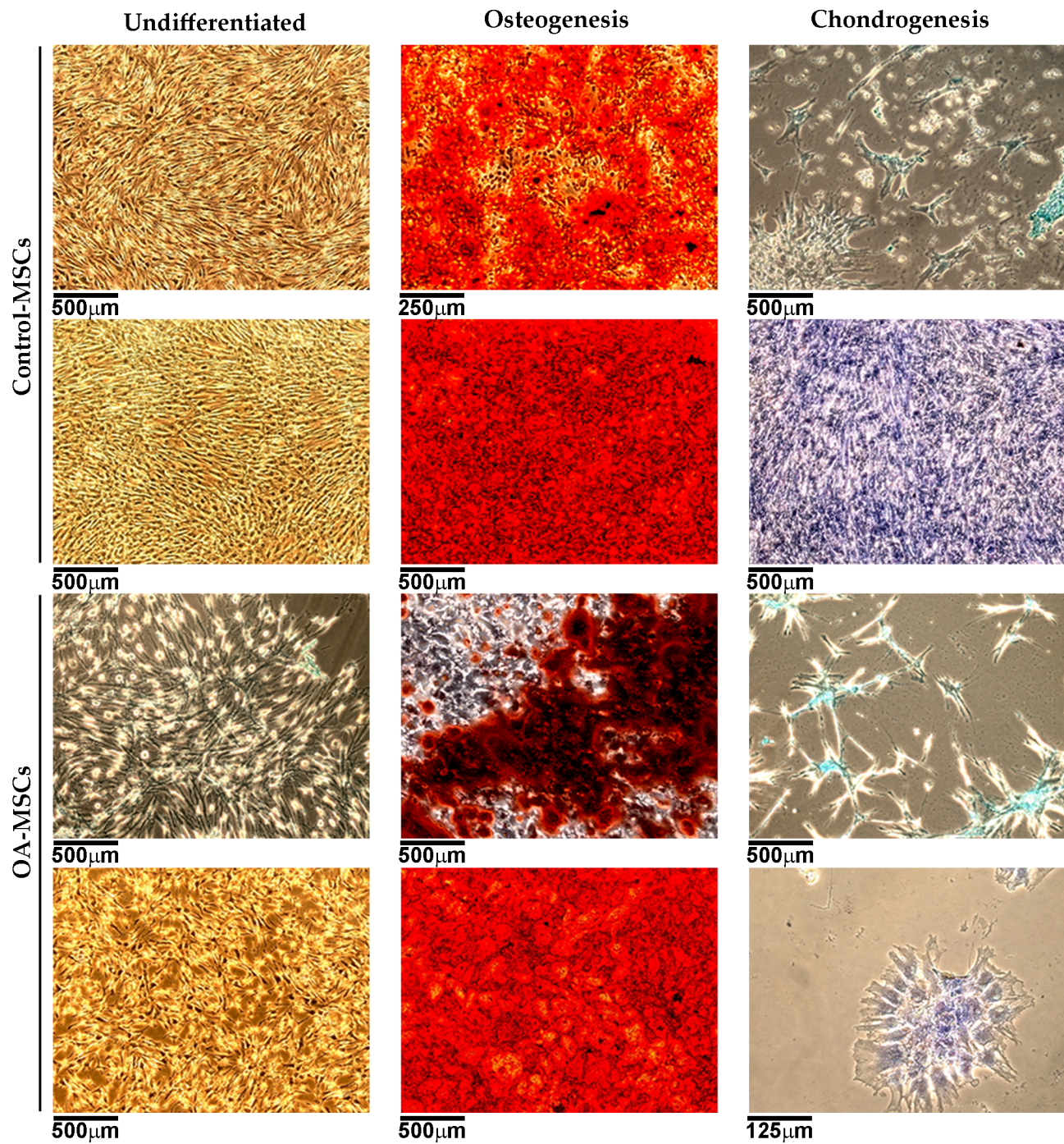
Distribution of the 39 human HOX genes in four clusters located in different chromosomal regions*. Blue indicates anterior HOX genes. Yellow, paralogy group 3 Hox genes, green and purple indicate central HOX genes and Red the posterior HOX genes.

* Apiou F, Flagiello D, Cillo C, Malfoy B, Poupon MF, Dutrillaux B. Fine mapping of human HOX gene clusters. Cytogenet Cell Genet. 1996;73(1-2):114-5.

Supplementary file Table S3

Involvement of Homeobox genes in skeletal and other diseases.

GENE	OMIM	DISEASE ANNOTATION
Homeobox A1	601536	Athabaskan Brainstem Dysgenesis Syndrome; ABDS
Homeobox D4	142981	Homeobox D4; HOXD4
Homeobox D13	610713	Brachydactyly-Syndactyly Syndrome
Homeobox D13	192350	Vater Association
Homeobox D13	186300	Syndactyly, Type V
Homeobox D13	186000	Synpolydactyly 1; SPD1
Homeobox D13	113300	Brachydactyly, Type E1; BDE1
Homeobox D13	113200	Brachydactyly, Type D; BDD
Homeobox A13	176305	Preaxial Deficiency, Postaxial Polydactyly, And Hypospadias
Homeobox A13	140000	Hand-Foot-Uterus Syndrome
Homeobox A11	605432	Radioulnar Synostosis with Amegakaryocytic Thrombocytopenia
Homeobox D10	192950	Vertical Talus, Congenital; CVT
Homeobox D10	142984	Homeobox D10; HOXD10
Homeobox A2	612290	Microtia, Hearing Impairment, and Cleft Palate



Supplementary file Table S5

List of TaqMan Gene Expression Assays used in the study.

Applied Biosystems™ TaqMan® Gene Expression Assays consist of a pair of unlabeled PCR primers and a TaqMan® probe with Applied Biosystems™ FAM™ dye label on the 5′ end and minor groove binder (MGB) and nonfluorescent quencher (NFQ) on the 3′ end. **Note about HOXC12** TaqMan® Gene Expression Assays. According to Applied Biosystems and based on re-evaluation, this assay may detect genomic DNA. Housekeeping genes are indicated in red.

Gene	Catalog #	Assay ID
<i>HOXA1</i>	4453320	Hs00939046_m1
<i>HOXA2</i>	4453320	Hs00534579_m1
<i>HOXA3</i>	4448892	Hs00601076_m1
<i>HOXA4</i>	4448892	Hs01573270_m1
<i>HOXA5</i>	4453320	Hs00430330_m1
<i>HOXA6</i>	4448892	Hs00430615_m1
<i>HOXA7</i>	4448892	Hs00600844_m1
<i>HOXA9</i>	4448892	Hs04931836_s1
<i>HOXA10</i>	4448892	Hs00172012_m1
<i>HOXA11</i>	4448892	Hs00194149_m1
<i>HOXA13</i>	4448892	Hs00426284_m1
<i>HOXB1</i>	4453320	Hs00157973_m1
<i>HOXB2</i>	4448892	Hs01911167_s1
<i>HOXB3</i>	4448892	Hs01587922_m1
<i>HOXB4</i>	4453320	Hs00256884_m1
<i>HOXB5</i>	4448892	Hs00357820_m1
<i>HOXB6</i>	4448892	Hs00980016_m1
<i>HOXB7</i>	4448892	Hs04187556_m1
<i>HOXB8</i>	4448892	Hs00256885_m1
<i>HOXB9</i>	4448892	Hs00256886_m1
<i>HOXB13</i>	4448892	Hs00197189_m1
<i>HOXC4</i>	4448892	Hs00538088_m1
<i>HOXC5</i>	4448892	Hs00232747_m1
<i>HOXC6</i>	4448892	Hs00171690_m1
<i>HOXC8</i>	4448892	Hs00224073_m1
<i>HOXC9</i>	4448892	Hs00396786_m1
<i>HOXC10</i>	4448892	Hs00213579_m1
<i>HOXC11</i>	4448892	Hs00204415_m1
<i>HOXC12</i>	4426964	Hs00329652_s1
<i>HOXC13</i>	4448892	Hs00600868_m1
<i>HOXD1</i>	4448892	Hs00707081_s1
<i>HOXD3</i>	4448892	Hs00232506_m1
<i>HOXD4</i>	4448892	Hs00429605_m1
<i>HOXD8</i>	4448892	Hs00251905_m1
<i>HOXD9</i>	4448892	Hs00610725_g1
<i>HOXD10</i>	4453320	Hs00157974_m1
<i>HOXD11</i>	4448892	Hs00360798_m1
<i>HOXD12</i>	4448892	Hs00706957_s1
<i>HOXD13</i>	4448892	Hs00171253_m1
<i>ACTB</i>	4448892	Hs01060665_g1
<i>18S</i>	4453320	Hs03003631_g1
<i>GAPDH</i>	4331182	Hs02786624_g1

Supplementary file Table S6

TOP 100 Upregulated genes in OA-MSCs

NAME	DESCRIPTION
<i>OGN</i>	Osteoglycin
<i>CHI3L2</i>	Chitinase 3-like 2
<i>CCRL1</i>	Chemokine (C-C motif) receptor-like 1
<i>CHI3L1</i>	Chitinase 3-like 1 (cartilage glycoprotein-39)
<i>MYOC</i>	Myocilin, trabecular meshwork inducible glucocorticoid response
<i>ANGPTL5</i>	Angiopoietin-like 5
<i>CD70</i>	CD70 molecule
<i>RSPO2</i>	R-spondin 2 homolog (<i>Xenopus laevis</i>)
<i>PDPN</i>	Podoplanin
<i>FGF18</i>	Fibroblast growth factor 18
<i>BARX1</i>	BARX homeobox 1
<i>EPB41L3</i>	Erythrocyte membrane protein band 4.1-like 3
<i>LNK1</i>	Ligand of numb-protein X 1
<i>NTN1</i>	Netrin 1
<i>ADH1A</i>	Alcohol dehydrogenase 1A (class I), alpha polypeptide
<i>MEOX2</i>	Mesenchyme homeobox 2
<i>MYRIP</i>	Myosin VIIA and Rab interacting protein
<i>MIA</i>	Melanoma inhibitory activity
<i>HAPLN1</i>	Hyaluronan and proteoglycan link protein 1
<i>PRG4</i>	Proteoglycan 4
<i>FGL2</i>	Fibrinogen-like 2
<i>SCARA5</i>	Scavenger receptor class A, member 5 (putative)
<i>MMP1</i>	Matrix metalloproteinase 1 (interstitial collagenase)
<i>FLJ14167</i>	Hypothetical protein FLJ14167
<i>CRTAC1</i>	Cartilage acidic protein 1
<i>SEMA3E</i>	Sema domain, immunoglobulin domain (Ig), short basic domain, secreted, (semaphorin) 3E
<i>PLA2G2A</i>	Phospholipase A2, group IIA (platelets, synovial fluid)
<i>RGMA</i>	RGM domain family, member A
<i>ADAMTSL3</i>	ADAMTS-like 3
<i>VEPH1</i>	Ventricular zone expressed PH domain homolog 1 (zebrafish)
<i>IL13RA2</i>	Interleukin 13 receptor, alpha 2
<i>ADH1C</i>	Alcohol dehydrogenase 1C (class I), gamma polypeptide
<i>FGD4</i>	FYVE, rhogef and PH domain containing 4
<i>ERG</i>	V-ets erythroblastosis virus E26 oncogene homolog (avian)
<i>COLEC12</i>	Collectin sub-family member 12
<i>CFB</i>	Complement factor B
<i>ZNF521</i>	Zinc finger protein 521
<i>TCEAL7</i>	Transcription elongation factor A (SII)-like 7
<i>CP</i>	Ceruloplasmin (ferroxidase)
<i>ADORA1</i>	Adenosine A1 receptor
<i>LAMA2</i>	Laminin, alpha 2 (merosin, congenital muscular dystrophy)
<i>ZNF659</i>	Zinc finger protein 659
<i>ZNF533</i>	Zinc finger protein 533
<i>CRYM</i>	Crystallin, mu
<i>ABCA6</i>	ATP-binding cassette, sub-family A (ABC1), member 6
<i>HYAL1</i>	Hyaluronoglucosaminidase 1
<i>FGF9</i>	Fibroblast growth factor 9 (glia-activating factor)
<i>KIAA1199</i>	Kiaa1199
<i>CDC47L</i>	Cell division cycle associated 7-like
<i>CIT</i>	Citron (rho-interacting, serine/threonine kinase 21)
<i>OAF</i>	OAF homolog (<i>Drosophila</i>)
<i>DLX4</i>	Distal-less homeobox 4
<i>OMD</i>	Osteomodulin

<i>CPXM2</i>	Carboxypeptidase X (M14 family), member 2
<i>WISP3</i>	WNT1 inducible signaling pathway protein 3
<i>C2orf40</i>	Chromosome 2 open reading frame 40
<i>DLK1</i>	Delta-like 1 homolog (Drosophila)
<i>ADRA2A</i>	Adrenergic, alpha-2A-, receptor
<i>ASPN</i>	Asporin
<i>SCARA3</i>	Scavenger receptor class A, member 3
<i>DNM1</i>	Dynamin 1
<i>CFI</i>	Complement factor I
<i>EGR3</i>	Early growth response 3
<i>BEGAIN</i>	Brain-enriched guanylate kinase-associated homolog (rat)
<i>BDKRB1</i>	Bradykinin receptor B1
<i>DENND2A</i>	DENN/MADD domain containing 2A
<i>FLJ30901</i>	Hypothetical protein FLJ30901
<i>PCOLCE2</i>	Procollagen C-endopeptidase enhancer 2
<i>RIPK3</i>	Receptor-interacting serine-threonine kinase 3
<i>GALNT12</i>	UDP-N-acetyl-alpha-D-galactosamine:polypeptide N-acetylgalactosaminyltransferase 12 (galnac-T12)
<i>TEK</i>	TEK tyrosine kinase, endothelial (venous malformations, multiple cutaneous and mucosal)
<i>CPT1C</i>	Carnitine palmitoyltransferase 1C
<i>LHX2</i>	LIM homeobox 2
<i>LOC91461</i>	Hypothetical protein BC007901
<i>NNAT</i>	Neuronatin
<i>SNED1</i>	Sushi, nidogen and EGF-like domains 1
<i>DPP4</i>	Dipeptidyl-peptidase 4 (CD26, adenosine deaminase complexing protein 2)
<i>RBPM52</i>	RNA binding protein with multiple splicing 2
<i>SECTM1</i>	Secreted and transmembrane 1
<i>KIAA1324L</i>	KIAA1324-like
<i>C10orf90</i>	Chromosome 10 open reading frame 90
<i>SERPINA1</i>	Serpin peptidase inhibitor, clade A (alpha-1 antiproteinase, antitrypsin), member 1
<i>AEBP1</i>	AE binding protein 1
<i>CDKN3</i>	Cyclin-dependent kinase inhibitor 3 (CDK2-associated dual specificity phosphatase)
<i>GPX3</i>	Glutathione peroxidase 3 (plasma)
<i>RARRES1</i>	Retinoic acid receptor responder (tazarotene induced) 1
<i>SLC47A1</i>	Solute carrier family 47, member 1
<i>CSDC2</i>	Cold shock domain containing C2, RNA binding
<i>CYP3A5</i>	Cytochrome P450, family 3, subfamily A, polypeptide 5
<i>CLIC2</i>	Chloride intracellular channel 2
<i>PAQR9</i>	Progestin and adipoq receptor family member IX
<i>NTF3</i>	Neurotrophin 3
<i>SOX5</i>	SRY (sex determining region Y)-box 5
<i>OSAP</i>	Ovary-specific acidic protein
<i>C3</i>	Complement component 3
<i>ITGB8</i>	Integrin, beta 8
<i>TNFRSF1B</i>	Tumor necrosis factor receptor superfamily, member 1B
<i>ASPM</i>	Asp (abnormal spindle) homolog, microcephaly associated (Drosophila)
<i>PITX1</i>	Paired-like homeodomain 1

TOP 100 Downregulated genes in OA-MSCs

NAME	DESCRIPTION
<i>IGHA2</i>	Immunoglobulin heavy constant alpha 2 (A2m marker)
<i>LPL</i>	Lipoprotein lipase
<i>IGL@</i>	Immunoglobulin lambda locus
<i>IGHG4</i>	Immunoglobulin heavy constant gamma 4 (G4m marker)
<i>IGHM</i>	Immunoglobulin heavy constant mu
<i>HEYL</i>	Hairy/enhancer-of-split related with YRPW motif-like
<i>SPP1</i>	Secreted phosphoprotein 1 (osteopontin, bone sialoprotein I, early T-lymphocyte activation 1)
<i>CD163</i>	CD163 molecule
<i>IGLL1</i>	Immunoglobulin lambda-like polypeptide 1
<i>CCL3</i>	Chemokine (C-C motif) ligand 3
<i>NTN2L</i>	Netrin 2-like (chicken)
<i>IGH@</i>	Immunoglobulin heavy locus
<i>CTA-246H3.1</i>	Similar to omega protein
<i>SFRP2</i>	Secreted frizzled-related protein 2
<i>IGLV2-14</i>	Immunoglobulin lambda variable 2-14
<i>TYROBP</i>	TYRO protein tyrosine kinase binding protein
<i>HOXC13</i>	Homeobox C13
<i>C1QB</i>	Complement component 1, q subcomponent, B chain
<i>PRL</i>	Prolactin
<i>HOXB6</i>	Homeobox B6
<i>PLA2G7</i>	Phospholipase A2, group VII (platelet-activating factor acetylhydrolase, plasma)
<i>CHRD12</i>	Chordin-like 2
<i>WFDC1</i>	WAP four-disulfide core domain 1
<i>ACP5</i>	Acid phosphatase 5, tartrate resistant
<i>MYH11</i>	Myosin, heavy chain 11, smooth muscle
<i>MCAM</i>	Melanoma cell adhesion molecule
<i>FCGR3A</i>	Fc fragment of igg, low affinity iiiia, receptor (CD16a)
<i>IGKV1D-13</i>	Immunoglobulin kappa variable 1D-13
<i>AIF1</i>	Allograft inflammatory factor 1
<i>FRAS1</i>	Fraser syndrome 1
<i>LCP1</i>	Lymphocyte cytosolic protein 1 (L-plastin)
<i>MS4A7</i>	Membrane-spanning 4-domains, subfamily A, member 7
<i>C1QA</i>	Complement component 1, q subcomponent, A chain
<i>IGHG1</i>	Immunoglobulin heavy constant gamma 1 (G1m marker)
<i>TREM2</i>	Triggering receptor expressed on myeloid cells 2
<i>ZMAT4</i>	Zinc finger, matrin type 4
<i>TM4SF20</i>	Transmembrane 4 L six family member 20
<i>MMP9</i>	Matrix metalloproteinase 9 (gelatinase B, 92kda gelatinase, 92kda type IV collagenase)
<i>SLAMF7</i>	SLAM family member 7
<i>COL10A1</i>	Collagen, type X, alpha 1(Schmid metaphyseal chondrodysplasia)
<i>IGKV1-5</i>	Immunoglobulin kappa variable 1-5
<i>CD52</i>	CD52 molecule
<i>CSF1R</i>	Colony stimulating factor 1 receptor
<i>FPR1</i>	Formyl peptide receptor 1
<i>CDH15</i>	Cadherin 15, M-cadherin (myotubule)
<i>ALPL</i>	Alkaline phosphatase, liver/bone/kidney
<i>CCL18</i>	Chemokine (C-C motif) ligand 18 (pulmonary and activation-regulated)
<i>PPP1R14A</i>	Protein phosphatase 1, regulatory (inhibitor) subunit 14A
<i>CCL3L3</i>	Chemokine (C-C motif) ligand 3-like 3
<i>NEFM</i>	Neurofilament, medium polypeptide 150kda
<i>CD53</i>	CD53 molecule
<i>TSPAN7</i>	Tetraspanin 7
<i>HEY2</i>	Hairy/enhancer-of-split related with YRPW motif 2
<i>FMO3</i>	Flavin containing monooxygenase 3
<i>DYSF</i>	Dysferlin, limb girdle muscular dystrophy 2B (autosomal recessive)
<i>LAIR1</i>	Leukocyte-associated immunoglobulin-like receptor 1
<i>TSPAN18</i>	Tetraspanin 18

<i>UNC5C</i>	Unc-5 homolog C (C. Elegans)
<i>HAS1</i>	Hyaluronan synthase 1
<i>HOXB4</i>	Homeobox B4
<i>LAPTM5</i>	Lysosomal associated multispinning membrane protein 5
<i>VMO1</i>	Vitelline membrane outer layer 1 homolog (chicken)
<i>CXCR4</i>	Chemokine (C-X-C motif) receptor 4
<i>CNN1</i>	Calponin 1, basic, smooth muscle
<i>LOC401233</i>	Similar to HIV TAT specific factor 1; cofactor required for Tat activation of HIV-1 transcription
<i>HOXB3</i>	Homeobox B3
<i>HCLS1</i>	Hematopoietic cell-specific Lyn substrate 1
<i>JPH2</i>	Junctophilin 2
<i>EPC1</i>	Enhancer of polycomb homolog 1 (Drosophila)
<i>COBL</i>	Cordon-bleu homolog (mouse)
<i>IGKC</i>	Immunoglobulin kappa constant
<i>COL11A1</i>	Collagen, type XI, alpha 1
<i>SLC16A6</i>	Solute carrier family 16, member 6 (monocarboxylic acid transporter 7)
<i>BEX1</i>	Brain expressed, X-linked 1
<i>IGHV4-31</i>	Immunoglobulin heavy variable 4-31
<i>MGAT4A</i>	Mannosyl (alpha-1,3-)-glycoprotein beta-1,4-N-acetylglucosaminyltransferase, isozyme A
<i>FLRT3</i>	Fibronectin leucine rich transmembrane protein 3
<i>IGJ</i>	Immunoglobulin J polypeptide, linker protein for immunoglobulin alpha and mu polypeptides
<i>KIF26B</i>	Kinesin family member 26B
<i>PLCG2</i>	Phospholipase C, gamma 2 (phosphatidylinositol-specific)
<i>MS4A4A</i>	Membrane-spanning 4-domains, subfamily A, member 4
<i>C15orf48</i>	Chromosome 15 open reading frame 48
<i>NR4A3</i>	Nuclear receptor subfamily 4, group A, member 3
<i>ASB2</i>	Ankyrin repeat and SOCS box-containing 2
<i>RNF128</i>	Ring finger protein 128
<i>KCNMB4</i>	Potassium large conductance calcium-activated channel, subfamily M, beta member 4
<i>KRT7</i>	Keratin 7
<i>NGFR</i>	Nerve growth factor receptor (TNFR superfamily, member 16)
<i>MGC23985</i>	Similar to AVLV472
<i>CCL4</i>	Chemokine (C-C motif) ligand 4
<i>CES1</i>	Carboxylesterase 1 (monocyte/macrophage serine esterase 1)
<i>LILRB3</i>	Leukocyte immunoglobulin-like receptor, subfamily B (with TM and ITIM domains), member 3
<i>PLN</i>	Phospholamban
<i>ADAMTS9</i>	ADAM metalloproteinase with thrombospondin type 1 motif, 9
<i>HS3ST2</i>	Heparan sulfate (glucosamine) 3-O-sulfotransferase 2
<i>OASL</i>	2'-5'-oligoadenylate synthetase-like
<i>CDH3</i>	Cadherin 3, type 1, P-cadherin (placental)
<i>HLA-DMB</i>	Major histocompatibility complex, class II, DM beta
<i>ANKRD1</i>	Ankyrin repeat domain 1 (cardiac muscle)
<i>SIX6</i>	SIX homeobox 6

Top 100 upregulated and downregulated genes in OA-MSCs from the DNA-Array. In red are indicated all the homeobox genes and **Hox clustered** genes are in bold.

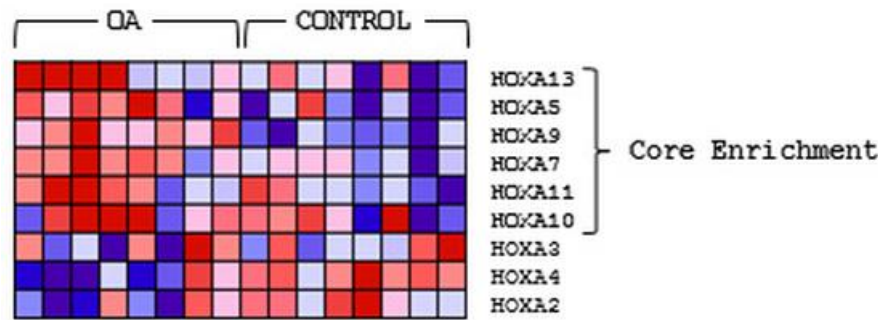
Supplementary file Table S7

Enrichment in OA phenotype (8 samples)

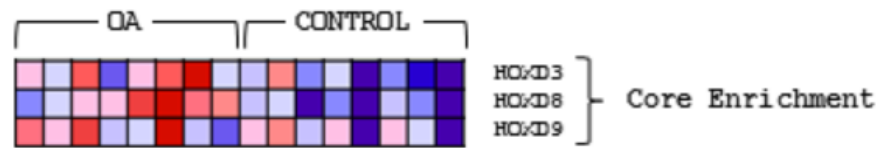
2 / 4 gene sets are **upregulated** in phenotype OA: HOXA and HOXD
Only 1 gene set (HOMEBOXA) is significant at FDR < 25%

NAME	SIZE	NOM p-val	FDR q-val	FWER p-val
HOMEBOX A	6	0.00193	0.00303	0.003

PROBE	DESCRIPTION (from dataset)	GENE SYMBOL	RANK IN GENE LIST	CORE ENRICHMENT
HOXA13	homeobox A13	HOXA13	380	Yes
HOXA5	homeobox A5	HOXA5	1080	Yes
HOXA9	homeobox A9	HOXA9	1382	Yes
HOXA7	homeobox A7	HOXA7	1668	Yes
HOXA11	homeobox A11	HOXA11	1987	Yes
HOXA10	homeobox A10	HOXA10	3486	Yes
HOXA3	homeobox A3	HOXA3	7346	No
HOXA4	homeobox A4	HOXA4	13707	No
HOXA2	homeobox A2	HOXA2	13778	No



PROBE	DESCRIPTION (from dataset)	GENE SYMBOL	RANK IN GENE LIST	CORE ENRICHMENT
HOXD3	homeobox D3	HOXD3	380	Yes
HOXD8	homeobox D8	HOXD8	1080	Yes
HOXD9	homeobox D9	HOXD9	1382	Yes



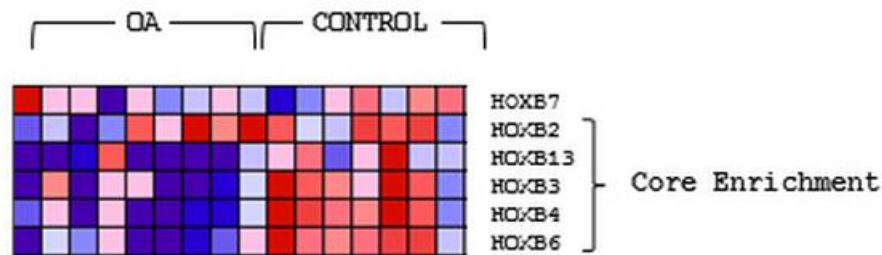
Enrichment in phenotype: CONTROL (8 samples)

- 2 / 4 gene sets are upregulated in phenotype CONTROL
- 2 gene sets are significantly enriched at FDR < 25%
- 1 gene sets are significantly enriched at nominal pvalue < 1%
- 1 gene sets are significantly enriched at nominal pvalue < 5%

NAME	SIZE	NOM p-val	FDR q-val	FWER p-val
HOMEBOX B	6	0.00193	0.00303	0.003
HOMEBOX C	7	0.11666	0.11344	0.172

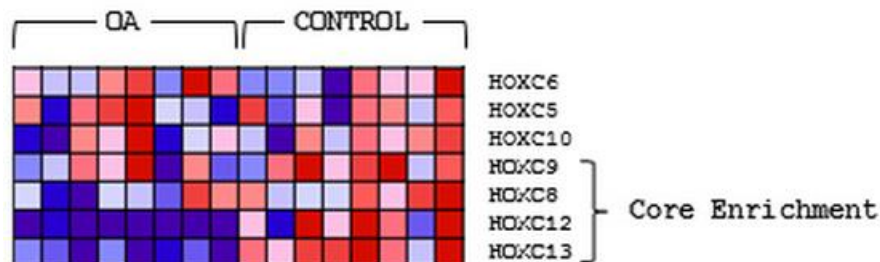
1 gene set (HOMEBOX B) is significantly enriched at nominal pvalue < 1%

PROBE	DESCRIPTION (from dataset)	GENE SYMBOL	RANK IN GENE LIST	CORE ENRICHMENT
HOXB7	homeobox B7	HOXB7	6427	No
HOXB2	homeobox B2	HOXB2	13273	Yes
HOXB13	homeobox B13	HOXB13	13836	Yes
HOXB3	homeobox B3	HOXB3	14288	Yes
HOXB4	homeobox B4	HOXB4	14294	Yes
HOXB6	homeobox B6	HOXB6	14334	Yes



1 gene set (HOMEBOX C) is significantly enriched at nominal pvalue < 25%

PROBE	DESCRIPTION (from dataset)	GENE SYMBOL	RANK IN GENE LIST	CORE ENRICHMENT
HOXC6	homeobox C6	HOXC6	3919	No
HOXC5	homeobox C5	HOXC5	6455	No
HOXC10	homeobox C10	HOXC10	8916	No
HOXC9	homeobox C9	HOXC9	12460	Yes
HOXC8	homeobox C8	HOXC8	13511	Yes
HOXC12	homeobox C12	HOXC12	14209	Yes
HOXC13	homeobox C13	HOXC13	14337	Yes



Supplementary file Table S8

HOX gene functional signatures are enriched in bone marrow Mesenchymal Stem Cells

Enrichment in OA subjects (8 samples)

5 / 12 gene sets are upregulated in OA phenotype

2 gene sets are significant at FDR < 25%

0 gene sets are significantly enriched at nominal pvalue < 1%

0 gene sets are significantly enriched at nominal pvalue < 5%

GENE SET NAME	SIZE	NOM p-val	FDR q-val
<i>HOXA</i>	9	0.0613	0.2324
Other HOX Genes	5	0.1232	0.2183
Body Pattern Formation	10	0.1080	0.2963
Anatomical Structure Morphogenesis	5	0.2202	0.2788
Hox Genes Involved In Cell Differentiation	7	0.4541	0.4401

Enrichment in control subjects (8 samples)

7 / 12 gene sets are upregulated in phenotype CONTROL

6 gene sets are significantly enriched at FDR < 25%

1 gene sets are significantly enriched at nominal pvalue < 1%

3 gene sets are significantly enriched at nominal pvalue < 5%

GENE SET NAME	SIZE	NOM p-val	FDR q-val
Other Hox Involved Multicellular Organismal Dev.	17	0.004	0.029
<i>HOXC</i>	6	0.035	0.128
Multicellular Organismal development (Hox)	7	0.035	0.094
Body Pattern Formation (Hox)	5	0.066	0.115
Skeletal Development	10	0.151	0.206
Nervous System Development	5	0.232	0.247
Organ Morphogenesis	7	0.979	0.988

HOXA

HOXA13	homeobox A13
HOXA5	homeobox A5
HOXA9	homeobox A9
HOXA7	homeobox A7
HOXA11	homeobox A11
HOXA10	homeobox A10
HOXA3	homeobox A3
HOXA4	homeobox A4
HOXA2	homeobox A2

BODY PATTERN FORMATION

EMX2	empty spiracles homeobox 2
HHEX	hematopoietically expressed homeobox
HOXA5	homeobox A5
HOXD8	homeobox D8
HOXA11	homeobox A11
HOXC5	homeobox C5
PITX2	paired-like homeodomain 2
HOXA4	homeobox A4
HOXA2	homeobox A2
HOXB6	homeobox B6

OTHER GENES

SIRT3	sirtuin (silent mating type information regulation 2 homolog) 3 (S. cerevisiae)
SIRT7	sirtuin (silent mating type information regulation 2 homolog) 7 (S. cerevisiae)
ZHX2	zinc fingers and homeoboxes 2
SIRT6	sirtuin (silent mating type information regulation 2 homolog) 6 (S. cerevisiae)
ZHX1	zinc fingers and homeoboxes 1

ANATOMICAL STRUCTURE MORPHOGENESIS

HOXD3	homeobox D3
SIX4	SIX homeobox 4
SIX2	SIX homeobox 2
LBX1	ladybird homeobox 1
HOXC13	homeobox C13

HOX GENES INVOLVED IN CELL DIFFERENTIATION

EMX2	empty spiracles homeobox 2
HHEX	hematopoietically expressed homeobox
LHX1	LIM homeobox 1
LBX1	ladybird homeobox 1
SIRT1	sirtuin (silent mating type information regulation 2 homolog) 1 (S. cerevisiae)
PHOX2A	paired-like homeobox 2a
HLX	H2.0-like homeobox

OTHER HOMEBOX GENES INVOLVED IN MULTICELLULAR ORGANISMAL DEVELOPMENT

HOXA9	homeobox A9
HOXD9	homeobox D9
HOXA7	homeobox A7
HOXB7	homeobox B7
EN2	engrailed homeobox 2

EMX1	empty spiracles homeobox 1
LBX2	ladybird homeobox 2
DLX1	distal-less homeobox 1
HOXC9	homeobox C9
MEIS1	Meis homeobox 1
HOXB2	homeobox B2
HOXC8	homeobox C8
NKX3-1	NK3 homeobox 1
HOXB13	homeobox B13
HOXC12	homeobox C12
HOXB3	homeobox B3
HOXB4	homeobox B4

MULTICELLULAR ORGANISMAL DEVELOPMENT(HOX)

HOXA9	homeobox A9
HOXA7	homeobox A7
HOXB7	homeobox B7
HOXB2	homeobox B2
HOXC8	homeobox C8
HOXB13	homeobox B13
HOXB4	homeobox B4

HOXC

HOXC6	homeobox C6
HOXC5	homeobox C5
HOXC9	homeobox C9
HOXC8	homeobox C8
HOXC12	homeobox C12
HOXC13	homeobox C13

BODY PATTERN FORMATION (HOX)

HOXD3	homeobox D3
HOXA5	homeobox A5
HOXA4	homeobox A4
HOXA2	homeobox A2
HOXB6	homeobox B6

SKELETAL DEVELOPMENT

HOXA13	homeobox A13
DLX3	distal-less homeobox 3
MSX2	msh homeobox 2
HOXA11	homeobox A11
EN1	engrailed homeobox 1
ALX4	aristaless-like homeobox 4
DLX6	distal-less homeobox 6
HOXA2	homeobox A2
DLX5	distal-less homeobox 5
HOXB6	homeobox B6

NERVOUS SYSTEM DEVELOPMENT

LHX1	LIM homeobox 1
LBX1	ladybird homeobox 1
PHOX2A	paired-like homeobox 2a

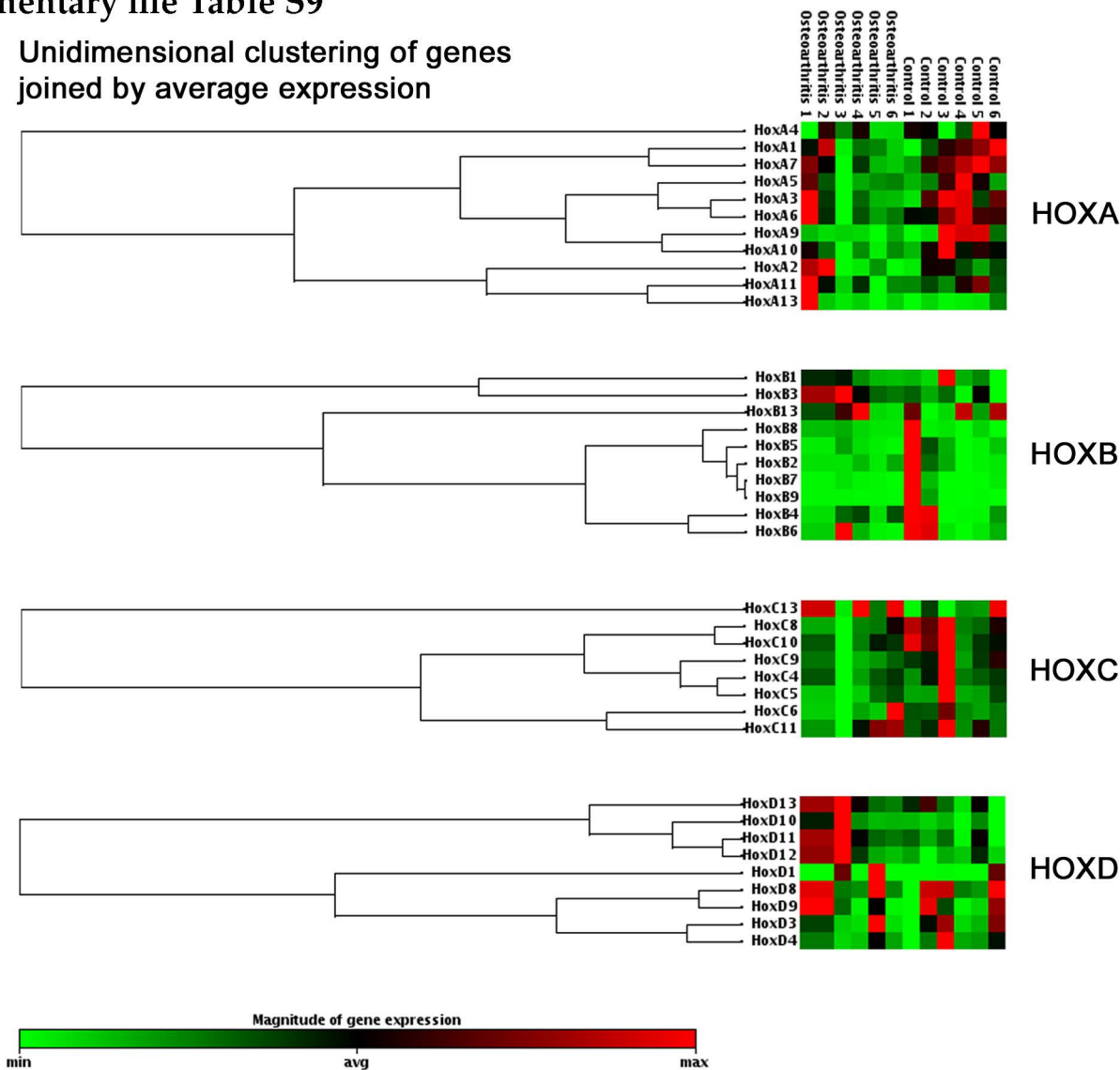
DLX6	distal-less homeobox 6
DLX5	distal-less homeobox 5

ORGAN MORPHOGENESIS

HOXA13	homeobox A13
HOXA11	homeobox A11
LHX1	LIM homeobox 1
PITX3	paired-like homeodomain 3
PITX2	paired-like homeodomain 2
PDHX	pyruvate dehydrogenase complex, component X
SIX6	SIX homeobox 6

Supplementary file Table S9

Unidimensional clustering of genes
joined by average expression



Bidimensional clustering of genes
joined by average expression

