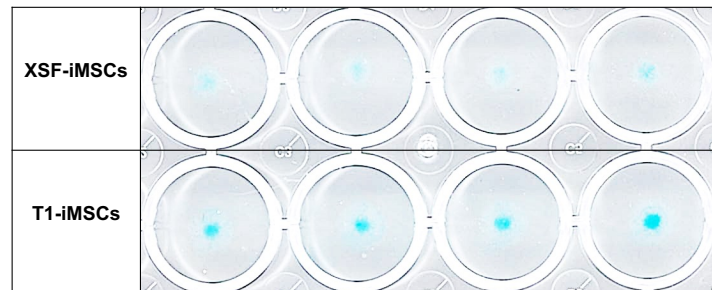


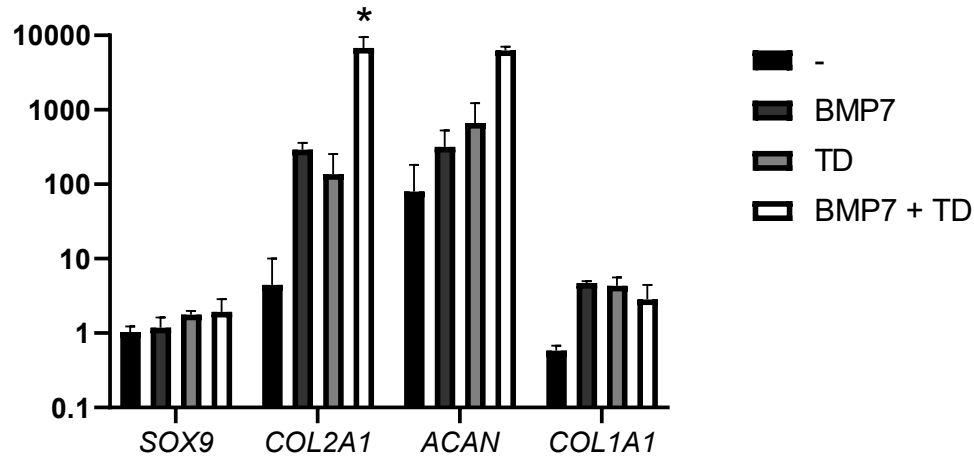
Supplementary Material

Supplementary Figure 1



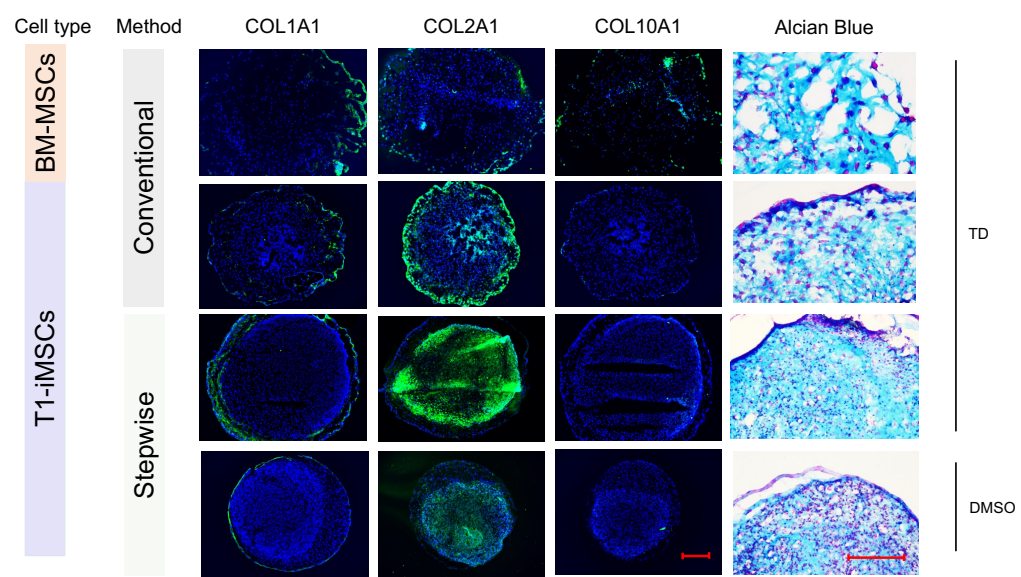
Supplementary Figure 1. Alcian blue staining of T1-iMSCs and XSF-iMSCs differentiated to chondrocytes in the presence of TD (100 nM) in micromass culture for 7 days.

Supplementary Figure 2



Supplementary Figure 2. The mRNA expression of chondrogenic markers determined by RT-qPCR in T1-iMSCs differentiated in micromass culture with basal chondrogenic medium (-) containing BMP7 (100 nM), TD (100 nM) or BMP7 (100 nM) + TD (100 nM). Data are the means $\pm$ SD (n=6) presented as folds relative to iPSCs. *P < 0.05 in BMP7 + TD vs. all others.

Supplementary Figure 3



Supplementary Figure 3. Immunostaining and alcian blue staining of BM-MSC and T1-iMSC chondrogenic spheroids on day 21 generated using different strategies in the presence or absence of TD (100 nM). Scale bars: 100 μm .

Supplementary Table 1**Supplementary Table 1.** Top 10 enriched terms by combined score of the DEGs in T1-iMSCs vs. XSF-iMSCs obtained from two libraries: BioPlanet and Panther.

Source: BioPlanet					
Term	Overlap	P-value	Adjusted P-value	Odds Ratio	Combined Score
TGF-beta regulation of extracellular matrix	76/565	1.94E-26	1.83E-23	5.1	303.5
Adipogenesis	19/133	5.89E-08	2.79E-05	5.1	84.9
BDNF signaling pathway	27/261	1.17E-07	3.68E-05	3.6	56.7
Interleukin-4 regulation of apoptosis	26/267	6.49E-07	1.54E-04	3.3	47.2
FSH regulation of apoptosis	25/263	1.66E-06	3.13E-04	3.2	42.9
Striated muscle contraction	9/38	2.61E-06	4.12E-04	9.4	120.6
Dilated cardiomyopathy	14/100	4.13E-06	5.58E-04	4.9	61.3
Arrhythmogenic right ventricular cardiomyopathy (ARVC)	12/75	4.88E-06	5.77E-04	5.8	70.7
Beta-1 integrin cell surface interactions	11/66	8.06E-06	8.47E-04	6.1	71.1
TAp63 pathway	10/55	9.32E-06	8.82E-04	6.7	77.9
Source: Panther					
5-Hydroxytryptamine degradation P04372	2/5	1.72E-04	0.01	19.968	92.32
Notch signaling pathway P00045	6/38	3.17E-04	0.01	5.6425	37.55
p53 pathway P00059	9/71	4.62E-04	0.01	4.3821	33.66
Plasminogen activating cascade P00050	3/15	1.29E-03	0.02	7.4961	33.46
Integrin signalling pathway P00034	15/156	3.59E-03	0.05	3.2286	27.99
CCKR signaling map ST P06959	15/165	6.36E-03	0.07	3.0335	24.44
Nic acetylcholine signaling pathway P00044	7/68	7.69E-03	0.08	3.4535	17.47
p38 MAPK pathway P05918	4/32	9.82E-03	0.08	4.2866	16.96
Cadherin signaling pathway P00012	12/150	1.15E-03	0.09	2.627	14.79
TGF-beta signaling pathway P00052	8/88	1.91E-03	0.13	3.0113	14.66

Supplementary Table 2

Supplementary Table 2. Primer sequences used for RT-qPCR

GENE NAME	FORWARD	REVERSE
<i>ACTB</i>	AGGTCTTTGCGGATGTCCACGT	CACCATTGGCAATGAGCGGTT C
<i>SOX10</i>	GAGCTGGACCGCACACCTTGG G	AACGCCCACCTCCTCGGACCTC
<i>NGFR</i>	CCGTTGGATTACACGGTCCA	GACAGGGATGAGGTTGTCGG
<i>CD44</i>	GCAATGCTTCTCAGACCACA	GAGGGGAGAGGGTAGACAGG
<i>CD73</i>	CAGTACCAGGGCACTATCTGG	AGTGGCCCCCTTGCTTTAAT
<i>CD90</i>	TGGATTAAGGATGAGGCCCCG	CCTAAGTCACTCGCCATCCC
<i>CD105</i>	CCACTAGCCAGGTCTCGAAG	GATGCAGGAAGACACTGCTG
<i>SOX9</i>	ATTCCTCCTGCCTTTGCTT	CTGGTGTTCTGAGAGGCACA
<i>COL2A1</i>	GGTGGCTTCCATTTAGCTA	TACCGGTATGTTTCGTGCAG
<i>COL1A1</i>	GTGCTAAAGGTGCCAATGGT	CTCCTCGCTTTCCTTCCTCT
<i>COL10A1</i>	ACGCTGAACGATACCAAACG	GCACACCTGGTTTCCCTACA
<i>ACAN</i>	ACAGCTGGGGACATTAGTGG	GTGGAATGCAGAGGTGGTTT
<i>PRG4</i>	GAACGTGCTATAGGACCTTC	CAGACTTTGGATAAGGTCTGC C
<i>RUNX2</i>	CAGACCAGCAGCACTCCATA	CAGCGTCAACACCATCATTC
<i>BGLAP</i>	GGCAGCGAGGTAGTGAAGAG	AGCAGAGCGACACCCTAGAC
<i>SP7</i>	AAGCTGATCTGGTGGTGCAT	GACTCCACAAAGGGCATGAT
<i>CEBPB</i>	TTTGTCCAAACCAACCGCAC	CCCCCAAAGGCTTTGTAACC
<i>CEBPA</i>	TATAGGCTGGGCTTCCCCTT	AGCTTTCTGGTGTGACTCGG
<i>PPARG</i>	CCAGAAGCCTGCATTCTGC	CACGGAGCTGATCCCAAAGT
<i>FABP4</i>	TGGGCCAGGAATTTGACGAA	CACATGTACCAGGACACCCC
<i>FOSL1</i>	CATCAACACCATGAGTGGCA	TCAGTCTCCTGTTCACAAGGC
<i>INHBE</i>	GCTAGCCAAGCAGCAAATCC	CATGGTACAGGTGGTGGGAC
<i>TGFBR2</i>	CTCCTGTGCAGCTTCCCTC	TCCACAGGACGATGTGCAG
<i>MAPK12</i>	CCTTTCCAGTCCGAGCTGTT	GCTTCAGGTCCCTCAGCC
<i>INHBB</i>	GAAGAGGGTGGACCTCAAGC	CCGATGAGGCGGAAGTCAAT
<i>GDF15</i>	GCAAGAACTCAGGACGGTGA	TGGAGTCTTCGGAGTGCAAC
<i>MAP3K7CL</i>	CTCCCCTCTGGATTGTCAGT	CCTGGCTGTGCTGATCATGT
<i>GDF6</i>	CGATCTCTCGCACACTCCTC	CACACGTCGAAGACTTCCCA
<i>GDF5</i>	CGGTCGGCTTTCTCCTTTCA	ACTTTCAAAGCAGCGGCAG
<i>MAPK15</i>	AGATACCTACTCAGGCGGCA	GATGGTCCCCAACTCCTGG

Supplementary Table 3

Supplementary Table 3.

Antibody name	Supplier	Code	Host	Dilution
Anti-Collagen I	Novus Biologicals	NB600-408	Rabbit	1:100
Anti-Collagen II	Invitrogen	MA5-12789	Mouse	1:100
Anti-Collagen X	Invitrogen	14977-1	Mouse	1:100
anti-Rabbit IgG (H+L) Alexa Fluor 488	Thermo Fisher Scientific	A11008	Goat	1:500
anti-Mouse IgG (H+L) Alexa Fluor 488	Thermo Fisher Scientific	A11001	Goat	1:500

Supplementary Movie

Supplementary Movie 1. Four-day recording of the fusion process on chondrogenic spheroids generated with the optimized strategy