

Supplementary Table S1. Cytokines quantifications in the patients series by Knee Osteoarthritis Inflammatory Phenotypes (**KOIP**). Cells show medians and median absolute deviations (continuous) and absolute frequencies and percentages (categorical) for the 13 cytokines evaluated in plasma and synovial fluid samples of the patients' cohort. Statistical significance was assessed using a Kruskal-Wallis (continuous) or a Fisher's test for contingency tables (categorical variables). **KOIP:** Knee Osteoarthritis Inflammatory Phenotype.

	All 168 (100%)	KOIP-1 55 (32.7%)	KOIP-2 51 (30.4%)	KOIP-3 27 (16.1%)	KOIP-4 35 (20.8%)	P-value	
SYNOVIAL FLUID	C-reactive protein (mg/L)	1.23 [0.03, 9.64]	1.52 [0.27, 6.02]	0.78 [0.03, 4.10]	1.61 [0.27, 9.64]	0.98 [0.15, 4.76]	<0.0001
	Tumor Necrosis Factor alpha (pg/mL)	9.02 [1.19, 127.49]	9.09 [1.19, 127.49]	8.14 [2.00, 18.17]	9.90 [2.19, 22.16]	8.90 [1.78, 17.98]	0.1239
	Interleukin 6 (pg/mL)	116.33 [0.11, 15683.26]	208.03 [7.55, 15683.26]	91.67 [0.11, 1857.83]	222.45 [22.77, 5820.23]	71.56 [4.01, 652.84]	0.0012
	Interleukin 8 (pg/mL)	6.21 [0.04, 881.40]	8.00 [0.44, 881.40]	6.21 [0.11, 278.44]	8.21 [0.04, 145.52]	3.92 [0.44, 29.02]	0.0012
	Nerve Growth Factor (pg/mL)	2.26 [0.01, 21.63]	2.26 [0.01, 21.63]	2.26 [0.46, 8.25]	2.20 [0.46, 6.90]	2.20 [0.72, 12.17]	0.038
	Calprotectin (ng/mL)	576.06 [36.79, 5957.01]	763.04 [60.31, 5376.84]	599.99 [36.79, 5957.01]	759.60 [197.72, 4129.89]	420.07 [36.79, 2447.75]	0.0006
	Leptin (pg/mL)	36160.71 [1928.12, 522278.22]	54016.11 [14014.43, 522278.22]	23126.79 [5335.19, 58808.60]	39506.85 [1928.12, 79207.03]	34888.07 [1928.12, 101330.36]	<0.0001
	Irisin (ng/mL)	695.95 [61.56, 3099.45]	938.23 [315.26, 3099.45]	422.07 [61.56, 983.08]	701.88 [161.47, 1270.77]	686.28 [242.12, 1527.24]	<0.0001
	Adiponectin (ng/mL)	2420.01 [300.19, 70423.60]	1937.73 [300.19, 6721.39]	3047.42 [689.12, 14901.06]	3372.91 [1194.76, 70423.60]	1501.12 [370.24, 3730.72]	<0.0001
	Omentin (pg/mL)	4292.68 [0.45, 28995.97]	3073.42 [0.45, 9401.45]	6648.21 [496.86, 28995.97]	6943.02 [1081.05, 25688.53]	2787.84 [130.78, 11575.06]	<0.0001
	Osteopontin (ng/mL)	47.95 [0.78, 3253.98]	48.55 [6.88, 3253.98]	46.75 [0.78, 441.11]	103.82 [15.18, 1400.55]	36.02 [6.88, 180.41]	0.0013
	Visfatin (ng/mL)	2.07 [0.13, 553.81]	2.28 [0.49, 70.61]	1.88 [0.13, 553.81]	2.11 [0.42, 7.53]	1.90 [0.53, 12.96]	0.5075
	Resistin (pg/mL)	1480.20 [32.46, 53485.59]	1692.08 [83.70, 53485.59]	1390.64 [52.92, 10386.82]	1242.23 [83.70, 18547.89]	1329.84 [32.46, 14574.82]	0.421
PLASMA	C-reactive protein (mg/L)	4.26 [0.42, 66.38]	6.87 [1.44, 66.38]	2.59 [0.42, 20.38]	6.00 [0.42, 27.13]	2.90 [0.42, 19.50]	<0.0001
	Interleukin 6 (pg/mL)	2.01 [0.08, 51.18]	2.28 [0.08, 42.02]	1.55 [0.12, 51.18]	2.45 [0.16, 8.28]	1.27 [0.08, 11.19]	0.0149
	Interleukin 8 (pg/mL)	3.19 [0.56, 402.21]	3.21 [0.93, 24.91]	3.79 [0.91, 13.13]	3.52 [0.56, 402.21]	2.30 [0.83, 8.22]	0.0152
	Tumor Necrosis Factor alpha (pg/mL)	6.37 [1.77, 22.13]	7.18 [1.77, 22.13]	6.18 [2.94, 14.66]	6.70 [2.59, 10.74]	6.08 [2.63, 8.86]	0.0997
	Nerve Growth Factor (pg/mL)	1.52 [0.22, 14.81]	1.52 [0.40, 14.81]	1.52 [0.22, 5.19]	1.69 [0.40, 8.31]	1.52 [0.44, 6.52]	0.2699
	Calprotectin (ng/mL)	694.27 [205.01, 12307.99]	851.11 [287.17, 6607.95]	613.63 [205.01, 1837.92]	800.12 [365.98, 12307.99]	588.54 [342.92, 1291.50]	<0.0001
	Leptin (pg/mL)	36688.11 [4039.08, 173177.01]	54392.71 [23739.58, 173177.01]	23566.59 [4039.08, 54817.52]	39809.19 [5017.56, 68130.76]	35530.72 [6972.63, 78971.55]	<0.0001
	Irisin (ng/mL)	707.78 [15.67, 1181.91]	859.25 [421.34, 1181.91]	437.22 [15.67, 884.41]	727.20 [234.19, 1059.97]	726.02 [116.43, 1105.78]	<0.0001
	Adiponectin (ng/mL)	16406.09 [1450.48, 59716.25]	11544.55 [1450.48, 52203.90]	22357.78 [10162.63, 59716.25]	21707.37 [7771.73, 40371.01]	11562.39 [4997.58, 42515.20]	<0.0001
	Omentin (pg/mL)	26859.10 [40.42, 114068.21]	22317.93 [40.42, 55037.25]	44069.15 [2444.37, 114068.21]	41837.49 [2444.37, 114068.21]	22449.96 [2444.37, 68850.68]	<0.0001
	Osteopontin (ng/mL)	13.71 [3.40, 64.53]	12.52 [3.40, 64.53]	14.12 [4.78, 55.83]	16.40 [4.91, 43.44]	12.10 [5.55, 29.52]	0.1872
	Visfatin (ng/mL)	4.01 [2.20, 12.58]	4.02 [2.20, 12.58]	4.23 [2.42, 8.37]	4.01 [2.59, 6.04]	3.74 [2.26, 6.93]	0.281
	Resistin (pg/mL)	2115.23 [791.02, 10079.62]	2389.10 [791.02, 8829.96]	1891.16 [958.55, 4435.30]	2657.56 [1045.95, 10079.62]	1874.89 [1194.29, 3812.08]	0.0001

Supplementary Table S2. Cytokines and Knee Osteoarthritis (KOA) outcomes evaluated in the patients series. g: parameter value for a Tukey ladder of powers transformation applied to continuous variables (when necessary) to symmetrize their distribution and make them more suitable for graphical representation and linear models for ELISA batch correction.

Set	Variable type	Variable (units / values)	Type (transformation parameter)
Cytokines	Inflammation markers - synovial fluid	C-reactive protein - synovial fluid (mg/L)	Continuous (g = 0.25)
		Interleukin 6 - synovial fluid (pg/mL)	Continuous (g = 0)
		Interleukin 8 - synovial fluid (pg/mL)	Continuous (g = 0)
		Tumor Necrosis Factor alpha - synovial fluid (pg/mL)	Continuous (g = 0)
		Nerve Growth Factor - synovial fluid (pg/mL)	Continuous (g = 0.25)
		Calprotectin - synovial fluid (ng/mL)	Continuous (g = 0.25)
	Adipocytokines / Myokines - synovial fluid	Leptin - synovial fluid (pg/mL)	Continuous (g = 0.25)
		Irisin - synovial fluid (ng/mL)	Continuous (g = 0)
		Adiponectin - synovial fluid (ng/mL)	Continuous (g = -0.25)
		Omentin - synovial fluid (pg/mL)	Continuous (g = 0.25)
		Osteopontin - synovial fluid (ng/mL)	Continuous (g = 0)
		Visfatin - synovial fluid (ng/mL)	Continuous (g = 0)
	Inflammation markers - plasma	Resistin - synovial fluid (pg/mL)	Continuous (g = 0)
		C-reactive protein - plasma (mg/L)	Continuous (g = 0)
		Interleukin 6 - plasma (pg/mL)	Continuous (g = 0)
		Interleukin 8 - plasma (pg/mL)	Continuous (g = -0.25)
		Tumor Necrosis Factor alpha - plasma (pg/mL)	Continuous (g = 0)
		Nerve Growth Factor - plasma (pg/mL)	Continuous (g = 0.25)
	Adipocytokines / Myokines - plasma	Calprotectin - plasma (ng/mL)	Continuous (g = -0.50)
		Leptin - plasma (pg/mL)	Continuous (g = 0.25)
Irisin - plasma (ng/mL)		Continuous (g = 0.75)	
Adiponectin - plasma (ng/mL)		Continuous (g = 0)	
Omentin - plasma (pg/mL)		Continuous (g = 0.50)	
Osteopontin - plasma (ng/mL)		Continuous (g = -0.25)	
KOA severity and progression	Radiography - baseline	Visfatin - plasma (ng/mL)	Continuous (g = -0.25)
		Resistin - plasma (pg/mL)	Continuous (g = -0.50)
		Kellgren-Lawrence radiographic grade (1-4)	Categorical - four categories
	Clinical severity - baseline	Osteophytes score (0-10)	Continuous
		Joint space narrowing (0-4)	Categorical - five categories
		KOOS - pain (reversed, 0-100)	Continuous
	Ultrasound - baseline	KOOS - symptoms (reversed, 0-100)	Continuous
		KOOS - functional disability (reversed, 0-100)	Continuous
	Radiographic progression (2 years follow-up)	Joint effusion (mm)	Continuous
		Synovial tissue thickness (mm)	Continuous
		Joint space narrowing radiographic progression (Yes / No)	Binary
		Kellgren-Lawrence radiographic progression (Yes / No)	Binary
		Osteophytes radiographic progression (Yes / No)	Binary
		Joint space narrowing radiographic progression (Yes / No)	Binary

Supplementary Table S4. Association between cytokines and Knee Osteoarthritis (KOA) outcomes in the KOIP-1 group. Table cells show non-parametric correlation-like measures and 95% Confidence Intervals (95%CI) between cytokines and clinical, radiographic and ultrasound severity at baseline and radiographic progression. Correlation-like measures are: Spearman correlation (continuous or ordinal outcomes) and Glass rank biserial correlation (binary outcomes). Radiographic progression according to Kellgren-Lawrence staging and joint space narrowing were treated as ordinal in these analyses. Red color indicates positive correlation, blue represents negative correlation, and color intensity expresses more extreme values of the correlation coefficients. Color intensities were saturated to 0.5 and -0.5 for positive and negative correlation, respectively. **IL-6:** Interleukin 6; **IL-8:** Interleukin 8; **TNF-alpha:** Tumor Necrosis Factor alpha; **NGF:** Nerve Growth Factor; **CRP:** C-Reactive Protein; **KOA:** Knee Osteoarthritis. **KOIP:** Knee Osteoarthritis Inflammatory Phenotype.

	KOOS pain	KOOS symptoms	KOOS functional disability	Joint effusion	Synovial tissue thickness	Kellgren-Lawrence Baseline	Osteophytes Baseline	Joint space narrowing Baseline	Kellgren-Lawrence Progression	Osteophytes Progression	Joint space narrowing Progression
CRP - synovial fluid	0.057 [-0.212, 0.338] 0.6808475	0.169 [-0.099, 0.453] 0.217488	0.037 [-0.265, 0.324] 0.7880936	0.088 [-0.192, 0.374] 0.523407	0.008 [-0.287, 0.289] 0.953269	0.056 [-0.201, 0.316] 0.685440	0.245 [-0.037, 0.482] 0.071720	-0.057 [-0.312, 0.225] 0.6782933	0.322 [-0.024, 0.620] 0.0623661	-0.033 [-0.519, 0.439] 0.8768724	0.013 [-0.318, 0.364] 0.936124
CRP - plasma	0.018 [-0.254, 0.283] 0.8941563	0.084 [-0.205, 0.368] 0.541826	0.092 [-0.198, 0.374] 0.760471	-0.152 [-0.440, 0.135] 0.268017	-0.096 [-0.350, 0.169] 0.919254	-0.014 [-0.271, 0.234] 0.919254	-0.070 [-0.193, 0.325] 0.565797	-0.025 [-0.399, 0.071] 0.2181278	0.025 [-0.318, 0.360] 0.8860196	-0.169 [-0.311, 0.508] 0.6795200	-0.110 [-0.449, 0.226] 0.508546
IL-6 - synovial fluid	-0.122 [-0.364, 0.153] 0.3763797	0.060 [-0.194, 0.328] 0.661141	-0.041 [-0.304, 0.222] 0.7656372	0.031 [-0.232, 0.289] 0.821838	-0.139 [-0.378, 0.135] 0.316026	0.117 [-0.153, 0.368] 0.391270	-0.165 [-0.429, 0.095] 0.8453379	0.027 [-0.373, 0.373] 0.8686328	0.029 [-0.319, 0.373] 0.8768849	0.033 [-0.378, 0.467] 0.8768849	0.304 [0.020, 0.603] 0.068282
IL-6 - plasma	0.007 [-0.293, 0.277] 0.9621484	0.088 [-0.190, 0.339] 0.521256	0.113 [-0.164, 0.376] 0.4132549	0.078 [-0.215, 0.333] 0.573820	0.072 [-0.225, 0.328] 0.604131	0.108 [-0.167, 0.369] 0.434161	-0.083 [-0.340, 0.206] 0.547407	-0.052 [-0.335, 0.225] 0.7068630	0.055 [-0.292, 0.436] 0.7491395	-0.058 [-0.436, 0.333] 0.7863020	0.130 [-0.229, 0.433] 0.434612
IL-8 - synovial fluid	0.030 [-0.266, 0.345] 0.8320551	0.314 [0.058, 0.556] 0.020651	0.199 [-0.110, 0.466] 0.1488220	0.082 [-0.234, 0.372] 0.553326	0.145 [-0.142, 0.404] 0.301267	0.103 [-0.166, 0.345] 0.459411	-0.081 [-0.353, 0.179] 0.558649	-0.093 [-0.167, 0.344] 0.5039524	-0.101 [-0.429, 0.255] 0.5616879	-0.046 [-0.442, 0.469] 0.9789212	0.049 [-0.270, 0.389] 0.771999
IL-8 - plasma	0.038 [-0.235, 0.309] 0.7821161	0.192 [-0.064, 0.444] 0.160820	0.164 [-0.102, 0.424] 0.2326517	0.058 [-0.215, 0.346] 0.675558	-0.121 [-0.434, 0.170] 0.383941	0.165 [-0.109, 0.434] 0.228820	0.028 [-0.242, 0.296] 0.840704	0.065 [-0.200, 0.303] 0.6360916	-0.112 [-0.459, 0.268] 0.5153208	0.225 [-0.186, 0.544] 0.2956847	0.164 [-0.167, 0.465] 0.326266
TNF-alpha - synovial fluid	0.028 [-0.272, 0.286] 0.8418532	0.124 [-0.164, 0.388] 0.370218	0.107 [-0.139, 0.421] 0.2646216	0.107 [-0.151, 0.356] 0.2646216	-0.067 [-0.339, 0.213] 0.634660	0.103 [-0.173, 0.350] 0.406664	-0.053 [-0.340, 0.229] 0.9470219	0.009 [-0.273, 0.303] 0.1777694	-0.236 [-0.560, 0.119] 0.1777694	0.009 [-0.342, 0.400] 0.9683949	0.154 [-0.180, 0.485] 0.362595
TNF-alpha - plasma	0.019 [-0.251, 0.305] 0.8908295	0.070 [-0.216, 0.339] 0.609631	0.028 [-0.285, 0.291] 0.8368486	0.208 [-0.072, 0.480] 0.126986	0.032 [-0.257, 0.309] 0.30908193534	0.038 [-0.226, 0.288] 0.780635	-0.019 [-0.321, 0.269] 0.888887	0.002 [-0.293, 0.279] 0.9884097	0.086 [-0.260, 0.435] 0.6197631	0.178 [-0.300, 0.615] 0.4086700	0.254 [-0.044, 0.575] 0.127874
NGF - synovial fluid	0.028 [-0.274, 0.309] 0.8436937	0.012 [-0.287, 0.304] 0.929738	0.072 [-0.227, 0.369] 0.6101472	0.033 [-0.201, 0.295] 0.814862	0.145 [-0.248, 0.306] 0.816196	-0.001 [-0.303, 0.279] 0.994785	0.060 [-0.194, 0.323] 0.670694	-0.121 [-0.385, 0.170] 0.3861629	-0.139 [-0.471, 0.219] 0.4261336	0.462 [0.036, 0.843] 0.0311171	0.040 [-0.304, 0.375] 0.812906
NGF - plasma	-0.211 [-0.445, 0.022] 0.1257097	-0.158 [-0.417, 0.145] 0.253666	-0.181 [-0.469, 0.099] 0.1898111	0.188 [-0.087, 0.414] 0.173928	0.129 [-0.136, 0.403] 0.357335	0.148 [-0.128, 0.413] 0.286665	-0.125 [-0.376, 0.150] 0.366560	0.141 [-0.135, 0.376] 0.3107590	-0.250 [-0.567, 0.084] 0.1492430	-0.262 [-0.683, 0.197] 0.2218059	-0.033 [-0.370, 0.310] 0.843722
Calprotectin - synovial fluid	0.167 [-0.104, 0.427] 0.2374552	0.222 [-0.061, 0.463] 0.113881	0.136 [-0.160, 0.404] 0.3347281	0.222 [-0.298, 0.222] 0.814926	0.196 [-0.090, 0.462] 0.166977	-0.008 [-0.286, 0.268] 0.951929	-0.047 [-0.310, 0.204] 0.741342	-0.021 [-0.295, 0.242] 0.8831276	-0.039 [-0.430, 0.339] 0.8265385	-0.147 [-0.569, 0.297] 0.9685183	-0.002 [-0.365, 0.325] 0.991202
Calprotectin - plasma	-0.013 [-0.293, 0.283] 0.9252163	0.197 [-0.083, 0.469] 0.148790	0.133 [-0.160, 0.428] 0.3312908	-0.048 [-0.317, 0.215] 0.725346	-0.075 [-0.336, 0.213] 0.588316	0.070 [-0.177, 0.300] 0.609572	0.090 [-0.182, 0.334] 0.515135	0.005 [-0.244, 0.278] 0.9729602	0.181 [-0.204, 0.529] 0.2948672	0.150 [-0.211, 0.512] 0.4857171	0.130 [-0.190, 0.452] 0.434624
Leptin - synovial fluid	-0.076 [-0.386, 0.223] 0.5800336	-0.102 [-0.393, 0.231] 0.457513	-0.024 [-0.325, 0.263] 0.8604284	-0.159 [-0.426, 0.116] 0.245836	-0.152 [-0.422, 0.119] 0.272658	0.154 [-0.120, 0.419] 0.894325	-0.018 [-0.318, 0.236] 0.849325	0.165 [-0.106, 0.426] 0.9290999	0.017 [-0.313, 0.347] 0.9290999	0.483 [0.077, 0.841] 0.0246829	-0.043 [-0.396, 0.297] 0.794529
Leptin - plasma	-0.110 [-0.404, 0.180] 0.4294761	-0.077 [-0.374, 0.219] 0.579981	0.015 [-0.252, 0.288] 0.9156144	-0.158 [-0.413, 0.125] 0.255339	-0.258 [-0.462, 0.222] 0.061894	0.072 [-0.186, 0.326] 0.604795	-0.025 [-0.303, 0.244] 0.859740	0.100 [-0.166, 0.262] 0.8023873	-0.044 [-0.364, 0.341] 0.8023873	0.402 [0.031, 0.762] 0.0625738	-0.061 [-0.385, 0.294] 0.717982
Irisin - synovial fluid	-0.086 [-0.335, 0.188] 0.5414083	-0.032 [-0.318, 0.259] 0.822072	-0.011 [-0.263, 0.236] 0.9387161	-0.077 [-0.365, 0.207] 0.582651	0.103 [-0.178, 0.387] 0.468699	0.268 [-0.007, 0.532] 0.052470	0.182 [-0.124, 0.435] 0.192753	0.231 [-0.074, 0.515] 0.0953657	-0.433 [-0.712, -0.112] 0.0141964	0.158 [-0.328, 0.578] 0.4652117	-0.364 [-0.676, -0.011] 0.023935
Irisin - plasma	-0.164 [-0.440, 0.115] 0.2365206	-0.090 [-0.354, 0.182] 0.517385	-0.098 [-0.375, 0.191] 0.4823262	-0.016 [-0.299, 0.275] 0.907568	0.055 [-0.194, 0.307] 0.697348	0.198 [-0.078, 0.462] 0.151702	0.094 [-0.189, 0.355] 0.500657	0.242 [-0.030, 0.515] 0.077908	-0.038 [-0.424, 0.320] 0.8287400	0.236 [-0.205, 0.743] 0.1145426	-0.238 [-0.546, 0.112] 0.157131
Adiponectin - synovial fluid	-0.115 [-0.390, 0.148] 0.4031430	-0.007 [-0.295, 0.287] 0.961767	-0.055 [-0.320, 0.223] 0.6902647	-0.022 [-0.293, 0.272] 0.871413	0.009 [-0.267, 0.267] 0.947381	0.080 [-0.196, 0.349] 0.560681	-0.024 [-0.331, 0.250] 0.860097	0.059 [-0.217, 0.325] 0.6667851	-0.139 [-0.484, 0.188] 0.4208492	0.608 [0.326, 0.844] 0.0046935	0.070 [-0.268, 0.405] 0.673965
Adiponectin - plasma	-0.251 [-0.482, -0.004] 0.0641210	-0.163 [-0.430, 0.110] 0.234760	-0.197 [-0.439, 0.074] 0.1496734	-0.163 [-0.424, 0.111] 0.234479	-0.173 [-0.430, 0.116] 0.211267	0.022 [-0.249, 0.287] 0.874643	0.021 [-0.243, 0.280] 0.879067	0.140 [-0.132, 0.404] 0.3070675	-0.116 [-0.427, 0.231] 0.5011969	-0.117 [-0.101, 0.705] 0.1482014	0.167 [-0.158, 0.498] 0.316503
Omentin - synovial fluid	-0.265 [-0.501, -0.001] 0.0503973	-0.064 [-0.341, 0.218] 0.643342	-0.218 [-0.462, 0.037] 0.1100286	0.145 [-0.145, 0.406] 0.289727	-0.030 [-0.325, 0.242] 0.827211	0.034 [-0.257, 0.296] 0.805896	-0.003 [-0.279, 0.280] 0.984379	0.049 [-0.267, 0.317] 0.7237695	0.208 [-0.149, 0.531] 0.2294128	0.433 [0.089, 0.735] 0.0440146	0.435 [0.144, 0.723] 0.009205
Omentin - plasma	-0.233 [-0.484, 0.053] 0.0866440	-0.060 [-0.327, 0.224] 0.665615	-0.106 [-0.371, 0.172] 0.4405165	-0.072 [-0.337, 0.169] 0.601943	-0.095 [-0.353, 0.165] 0.492641	0.228 [0.061, 0.555] 0.014846	0.327 [-0.051, 0.455] 0.093683	0.365 [0.097, 0.598] 0.0061057	-0.105 [-0.460, 0.216] 0.542162	0.506 [NA, NA] 0.0187931	-0.043 [-0.400, 0.310] 0.794529
Osteopontin - synovial fluid	0.097 [-0.183, 0.352] 0.4815616	0.116 [-0.149, 0.364] 0.398835	0.109 [-0.146, 0.350] 0.4303205	-0.013 [-0.309, 0.266] 0.922385	-0.092 [-0.348, 0.166] 0.506222	-0.149 [-0.411, 0.118] 0.278336	-0.120 [-0.379, 0.161] 0.381110	-0.168 [-0.435, 0.133] 0.2193841	0.042 [-0.314, 0.377] 0.8082729	0.211 [-0.209, 0.599] 0.3262690	0.244 [-0.092, 0.582] 0.143416
Osteopontin - plasma	0.134 [-0.176, 0.392] 0.3281943	-0.067 [-0.318, 0.212] 0.628722	0.078 [-0.216, 0.367] 0.5728687	-0.181 [-0.424, 0.085] 0.187217	-0.130 [-0.377, 0.130] 0.348827	-0.047 [-0.341, 0.226] 0.735328	0.036 [-0.219, 0.294] 0.248905	0.088 [-0.146, 0.322] 0.5216160	-0.149 [-0.489, 0.189] 0.3897390	-0.058 [-0.456, 0.378] 0.7862967	0.065 [-0.303, 0.407] 0.696040
Visfatin - synovial fluid	0.084 [-0.197, 0.379] 0.5431717	0.017 [-0.283, 0.318] 0.900667	-0.038 [-0.312, 0.284] 0.7831750	-0.150 [-0.375, 0.126] 0.274789	0.036 [-0.256, 0.309] 0.797978	0.060 [-0.219, 0.334] 0.662137	0.018 [-0.221, 0.278] 0.894325	-0.020 [-0.299, 0.231] 0.8820434	-0.055 [-0.396, 0.294] 0.7491395	-0.117 [-0.523, 0.274] 0.5876590	-0.013 [-0.339, 0.309] 0.936129
Resistin - synovial fluid	-0.035 [-0.283, 0.225] 0.7972716	-0.069 [-0.326, 0.188] 0.615252	-0.036 [-0.220, 0.316] 0.6297848	0.066 [-0.341, 0.243] 0.662958	0.066 [-0.225, 0.328] 0.637103	0.023 [-0.333, 0.249] 0.791860	-0.023 [-0.243, 0.295] 0.869784	-0.066 [-0.353, 0.229] 0.6337830	-0.455 [-0.750, -0.142] 0.0054063	0.092 [-0.364, 0.521] 0.6070794	-0.038 [-0.368, 0.308] 0.817789
Resistin - plasma	-0.079 [-0.314, 0.175] 0.5646775	-0.124 [-0.379, 0.133] 0.366273	-0.220 [-0.464, 0.055] 0.1071002	0.012 [-0.273, 0.302] 0.930413	-0.121 [-0.419, 0.191] 0.382289	0.071 [-0.173, 0.317] 0.605001	-0.095 [-0.365, 0.148] 0.488683	-0.015 [-0.277, 0.233] 0.9136636	-0.063 [-0.421, 0.262] 0.7159553	-0.172 [-0.554, 0.250] 0.4234695	-0.013 [-0.337, 0.338] 0.936131

Supplementary Table S5. Association between cytokines and Knee Osteoarthritis (KOA) outcomes in the **KOIP-2 group**. Table cells show non-parametric correlation-like measures and 95% Confidence Intervals (95%CI) between cytokines and clinical, radiographic and ultrasound severity at baseline and radiographic progression. Correlation-like measures are: Spearman correlation (continuous or ordinal outcomes) and Glass rank biserial correlation (binary outcomes). Radiographic progression according to Kellgren-Lawrence staging and joint space narrowing were treated as ordinal in these analyses. Red color indicates positive correlation, blue represents negative correlation, and color intensity expresses more extreme values of the correlation coefficients. Color intensities were saturated to 0.5 and -0.5 for positive and negative correlation, respectively. **IL-6:** Interleukin 6; **IL-8:** Interleukin 8; **TNF-alpha:** Tumor Necrosis Factor alpha; **NGF:** Nerve Growth Factor; **CRP:** C-Reactive Protein; **KOA:** Knee Osteoarthritis. **KOIP:** Knee Osteoarthritis Inflammatory Phenotype.

	KOOS pain	KOOS symptoms	KOOS functional disability	Joint effusion	Synovial tissue thickness	Kellgren-Lawrence Baseline	Osteophytes Baseline	Joint space narrowing Baseline	Kellgren-Lawrence Progression	Osteophytes Progression	Joint space narrowing Progression
CRP - synovial fluid	-0.192 [-0.472, 0.122] 0.181226	0.006 [-0.320, 0.331] 0.968923	-0.145 [-0.422, 0.148] 0.315037	0.022 [-0.286, 0.329] 0.878007	0.175 [-0.123, 0.454] 0.233185	0.068 [-0.261, 0.370] 0.636945	-0.065 [-0.337, 0.217] 0.651815	0.109 [-0.219, 0.412] 0.449363	0.046 [-0.457, 0.503] 0.834243	0.025 [-0.326, 0.405] 0.8967512	0.268 [-0.243, 0.731] 0.2203176
CRP - plasma	-0.005 [-0.268, 0.258] 0.972142	0.028 [-0.273, 0.308] 0.845896	0.005 [-0.273, 0.274] 0.973821	-0.107 [-0.399, 0.172] 0.456404	0.011 [-0.399, 0.300] 0.941143	-0.083 [-0.347, 0.198] 0.563137	0.020 [-0.304, 0.322] 0.891247	-0.020 [-0.347, 0.283] 0.886972	0.187 [-0.346, 0.641] 0.390714	-0.195 [-0.543, 0.150] 0.3015557	0.082 [-0.404, 0.523] 0.6950003
IL-6 - synovial fluid	0.068 [-0.230, 0.373] 0.637689	0.176 [-0.108, 0.467] 0.222709	0.083 [-0.188, 0.437] 0.335177	0.101 [-0.198, 0.394] 0.484166	0.190 [-0.083, 0.437] 0.194681	-0.080 [-0.367, 0.216] 0.581477	-0.089 [-0.394, 0.218] 0.547004	-0.102 [-0.390, 0.211] 0.480385	-0.248 [-0.683, 0.171] 0.256604	0.138 [-0.333, 0.150] 0.4752297	-0.061 [-0.468, 0.364] 0.773450
IL-6 - plasma	0.033 [-0.288, 0.330] 0.820779	0.153 [-0.171, 0.426] 0.283983	0.083 [-0.196, 0.360] 0.564384	0.108 [-0.193, 0.377] 0.448654	-0.118 [-0.396, 0.148] 0.418774	-0.103 [-0.398, 0.198] 0.471828	-0.076 [-0.347, 0.197] 0.597763	-0.116 [-0.414, 0.182] 0.417632	0.098 [-0.335, 0.528] 0.651887	-0.505 [-0.790, -0.130] 0.0075425	-0.138 [-0.590, 0.333] 0.5103045
IL-8 - synovial fluid	0.084 [-0.213, 0.360] 0.564403	0.068 [-0.235, 0.348] 0.640122	0.096 [-0.170, 0.361] 0.512996	-0.292 [-0.549, -0.020] 0.042035	-0.114 [-0.374, 0.142] 0.445407	0.035 [-0.250, 0.339] 0.812890	0.160 [-0.153, 0.454] 0.273189	0.069 [-0.212, 0.330] 0.637051	0.347 [-0.150, 0.743] 0.114358	0.062 [-0.655, 0.017] 0.0876540	0.306 [-0.126, 0.683] 0.1478229
IL-8 - plasma	0.062 [-0.183, 0.319] 0.667046	0.084 [-0.212, 0.353] 0.558917	0.001 [-0.286, 0.280] 0.993202	0.157 [-0.129, 0.431] 0.270712	0.105 [-0.164, 0.377] 0.473627	-0.109 [-0.356, 0.140] 0.447476	-0.176 [-0.439, 0.117] 0.217639	-0.145 [-0.403, 0.143] 0.310341	-0.308 [-0.613, 0.014] 0.158203	0.362 [-0.018, 0.710] 0.0554900	-0.106 [-0.488, 0.304] 0.6141905
TNF-alpha - synovial fluid	0.186 [-0.110, 0.426] 0.216900	0.149 [-0.140, 0.396] 0.322893	0.234 [-0.057, 0.504] 0.116843	-0.095 [-0.393, 0.243] 0.528795	0.053 [-0.225, 0.316] 0.732315	0.019 [-0.297, 0.312] 0.902523	-0.003 [-0.305, 0.301] 0.983231	-0.003 [-0.429, 0.108] 0.215943	0.153 [NA, NA] 0.080723	-0.159 [-0.548, 0.252] 0.4954471	0.062 [-0.419, 0.518] 0.7722214
TNF-alpha - plasma	-0.207 [-0.480, 0.070] 0.144965	-0.140 [-0.413, 0.151] 0.327216	-0.167 [-0.451, 0.111] 0.241859	-0.141 [-0.373, 0.118] 0.323836	0.068 [-0.369, 0.234] 0.642354	-0.082 [-0.371, 0.214] 0.567212	-0.108 [-0.369, 0.151] 0.451310	-0.093 [-0.412, 0.229] 0.516493	-0.130 [-0.598, 0.347] 0.558066	-0.190 [-0.560, 0.181] 0.3134999	-0.271 [-0.724, 0.219] 0.1976563
NGF - synovial fluid	0.113 [-0.174, 0.409] 0.446092	0.057 [-0.162, 0.396] 0.390305	-0.043 [-0.248, 0.328] 0.698342	-0.029 [-0.377, 0.330] 0.844729	-0.329 [-0.566, -0.033] 0.025810	-0.038 [-0.316, 0.256] 0.798773	-0.167 [-0.433, 0.136] 0.256902	-0.201 [-0.301, 0.231] 0.871516	-0.562 [-0.150, 0.185] 0.359345	-0.613 [-0.613, 0.141] 0.2093673	-0.467 [-0.467, 0.283] 0.7044738
NGF - plasma	0.036 [-0.238, 0.304] 0.800945	0.095 [-0.190, 0.364] 0.507137	-0.043 [-0.322, 0.275] 0.763257	-0.063 [-0.362, 0.240] 0.661225	0.042 [-0.253, 0.327] 0.772301	-0.244 [-0.499, 0.020] 0.084875	0.035 [-0.250, 0.319] 0.806272	-0.197 [-0.439, 0.085] 0.165288	0.102 [-0.305, 0.494] 0.640619	-0.100 [-0.455, 0.282] 0.9596576	0.124 [-0.295, 0.533] 0.5553505
Calprotectin - synovial fluid	0.089 [-0.198, 0.372] 0.574663	0.132 [-0.175, 0.414] 0.632246	0.083 [-0.227, 0.376] 0.568887	0.009 [-0.266, 0.309] 0.952726	0.103 [-0.210, 0.403] 0.486637	-0.206 [-0.472, 0.090] 0.150516	-0.003 [-0.332, 0.282] 0.844953	-0.141 [-0.431, 0.134] 0.329142	0.359 [-0.145, 0.767] 0.100583	0.113 [-0.242, 0.469] 0.510842	0.321 [-0.118, 0.707] 0.1275836
Calprotectin - plasma	-0.054 [-0.359, 0.232] 0.704307	-0.048 [-0.333, 0.230] 0.736922	-0.134 [-0.376, 0.165] 0.348779	0.101 [-0.164, 0.359] 0.479215	-0.112 [-0.397, 0.182] 0.443180	-0.255 [-0.544, 0.050] 0.070478	-0.092 [-0.357, 0.212] 0.522538	-0.279 [-0.541, 0.005] 0.047015	-0.073 [-0.623, 0.511] 0.737926	-0.314 [-0.638, 0.037] 0.0963307	-0.341 [-0.790, 0.193] 0.1043065
Leptin - synovial fluid	-0.112 [-0.377, 0.161] 0.432544	-0.068 [-0.326, 0.214] 0.607623	-0.074 [-0.350, 0.187] 0.637623	-0.184 [-0.443, 0.084] 0.196678	-0.054 [-0.363, 0.234] 0.711209	0.042 [-0.243, 0.322] 0.768883	-0.125 [-0.379, 0.138] 0.383268	-0.010 [-0.280, 0.239] 0.944671	-0.333 [-0.727, 0.125] 0.126617	-0.187 [-0.534, 0.174] 0.3382929	-0.571 [-0.861, -0.200] 0.0065948
Leptin - plasma	-0.076 [-0.322, 0.227] 0.595469	0.024 [-0.270, 0.318] 0.868906	0.023 [-0.270, 0.288] 0.874493	-0.201 [-0.472, 0.078] 0.157256	-0.103 [-0.348, 0.181] 0.480895	-0.047 [-0.333, 0.246] 0.743424	-0.088 [-0.380, 0.216] 0.540099	-0.073 [-0.358, 0.199] 0.611502	-0.187 [-0.660, 0.313] 0.390680	-0.145 [-0.536, 0.220] 0.4421415	-0.429 [-0.835, 0.059] 0.0408989
Irisin - synovial fluid	0.105 [-0.158, 0.374] 0.469843	0.196 [-0.095, 0.446] 0.172175	0.142 [-0.146, 0.406] 0.324799	-0.110 [-0.417, 0.200] 0.446436	0.030 [-0.210, 0.267] 0.814649	-0.098 [-0.364, 0.162] 0.496944	-0.139 [-0.412, 0.167] 0.367196	-0.196 [-0.452, 0.129] 0.204229	-0.196 [-0.627, 0.248] 0.370455	-0.177 [-0.519, 0.222] 0.3057954	-0.224 [-0.651, 0.229] 0.2874759
Irisin - plasma	-0.089 [-0.369, 0.196] 0.533790	0.021 [-0.281, 0.290] 0.883898	0.067 [-0.261, 0.369] 0.638315	-0.238 [-0.532, 0.054] 0.092753	-0.091 [-0.369, 0.210] 0.534022	-0.147 [-0.413, 0.144] 0.302119	-0.186 [-0.458, 0.096] 0.190628	-0.109 [-0.378, 0.155] 0.444781	-0.137 [-0.544, 0.300] 0.531605	-0.062 [-0.432, 0.301] 0.7432374	-0.435 [-0.829, 0.016] 0.0382268
Adiponectin - synovial fluid	0.113 [-0.131, 0.374] 0.429112	0.210 [-0.095, 0.450] 0.138468	0.079 [-0.128, 0.390] 0.355233	0.231 [-0.082, 0.508] 0.102952	0.314 [0.018, 0.573] 0.028268	0.011 [-0.258, 0.289] 0.941083	0.086 [-0.216, 0.369] 0.546396	-0.046 [-0.321, 0.247] 0.749713	-0.003 [-0.494, 0.456] 0.749713	-0.305 [-0.629, 0.028] 0.1068200	0.188 [-0.265, 0.600] 0.3701570
Adiponectin - plasma	0.111 [-0.156, 0.378] 0.439477	0.000 [-0.315, 0.294] 0.998736	0.079 [-0.185, 0.344] 0.582468	0.149 [-0.126, 0.433] 0.296014	-0.026 [-0.319, 0.291] 0.859540	-0.055 [-0.310, 0.240] 0.700631	0.186 [-0.096, 0.453] 0.190198	-0.099 [-0.384, 0.210] 0.488163	0.333 [-0.014, 0.645] 0.126604	-0.093 [-0.506, 0.307] 0.6231517	0.424 [0.124, 0.701] 0.0437430
Omentin - synovial fluid	0.277 [-0.013, 0.550] 0.048755	0.265 [-0.033, 0.515] 0.059832	0.228 [-0.093, 0.527] 0.107389	0.152 [-0.166, 0.461] 0.287251	0.106 [-0.177, 0.414] 0.470166	-0.095 [-0.383, 0.178] 0.506556	-0.222 [-0.325, 0.268] 0.880286	-0.114 [-0.388, 0.195] 0.426190	0.162 [-0.371, 0.682] 0.0587817	0.357 [-0.687, 0.024] 0.0587817	0.147 [-0.340, 0.583] 0.4838399
Omentin - plasma	0.171 [-0.113, 0.462] 0.230413	0.150 [-0.141, 0.416] 0.293784	0.136 [-0.145, 0.436] 0.340476	0.096 [-0.168, 0.363] 0.501140	0.083 [-0.228, 0.395] 0.570988	-0.086 [-0.348, 0.198] 0.546971	0.196 [-0.089, 0.468] 0.168473	-0.134 [-0.412, 0.163] 0.348884	0.133 [-0.260, 0.528] 0.541179	-0.112 [-0.535, 0.319] 0.5537400	0.103 [-0.283, 0.482] 0.6240523
Osteopontin - synovial fluid	0.183 [-0.102, 0.471] 0.198656	0.064 [-0.199, 0.346] 0.654943	0.228 [-0.041, 0.469] 0.107057	0.014 [-0.256, 0.302] 0.923357	-0.236 [-0.476, 0.041] 0.103114	-0.210 [-0.475, 0.076] 0.138604	-0.014 [-0.305, 0.293] 0.922460	-0.104 [-0.417, 0.198] 0.465656	0.121 [-0.344, 0.613] 0.580254	-0.371 [-0.691, -0.021] 0.092861	0.029 [-0.403, 0.509] 0.8885989
Osteopontin - plasma	0.098 [-0.207, 0.379] 0.494813	0.167 [-0.110, 0.416] 0.240444	0.169 [-0.097, 0.410] 0.237094	0.171 [-0.083, 0.439] 0.229920	-0.085 [-0.382, 0.228] 0.1061076	0.178 [-0.114, 0.451] 0.210418	0.249 [-0.048, 0.501] 0.078544	0.197 [-0.091, 0.438] 0.166936	-0.137 [-0.518, 0.289] 0.531605	-0.395 [-0.738, -0.019] 0.0364919	-0.224 [-0.590, 0.143] 0.2873233
Visfatin - synovial fluid	0.089 [-0.179, 0.359] 0.534199	0.135 [-0.139, 0.395] 0.385418	0.124 [-0.152, 0.386] 0.385418	0.197 [-0.104, 0.513] 0.166031	0.137 [-0.143, 0.424] 0.346373	0.126 [-0.151, 0.392] 0.376532	0.172 [-0.129, 0.445] 0.227533	0.162 [-0.077, 0.454] 0.167084	0.162 [-0.245, 0.537] 0.458107	0.021 [-0.338, 0.374] 0.9097187	-0.035 [-0.510, 0.460] 0.8665522
Visfatin - plasma	-0.252 [-0.511, -0.004] 0.074890	-0.253 [-0.492, 0.017] 0.073414	-0.156 [-0.425, 0.121] 0.275054	0.154 [-0.140, 0.405] 0.280186	0.208 [-0.085, 0.464] 0.152450	0.078 [-0.232, 0.347] 0.588722	0.007 [-0.302, 0.289] 0.961830	-0.065 [-0.329, 0.212] 0.651418	0.537 [-0.829, -0.177] 0.013948	-0.143 [-0.561, 0.302] 0.4496918	-0.582 [-0.973, -0.211] 0.0055618
Resistin - synovial fluid	0.152 [-0.149, 0.425] 0.285498	-0.037 [-0.313, 0.210] 0.795390	0.127 [-0.143, 0.403] 0.375626	-0.077 [-0.371, 0.222] 0.590927	-0.249 [-0.528, 0.057] 0.084067	-0.175 [-0.438, 0.118] 0.218957	-0.034 [-0.299, 0.236] 0.811473	-0.095 [-0.383, 0.174] 0.506133	0.162 [-0.292, 0.579] 0.457790	0.169 [-0.197, 0.522] 0.3706916	0.241 [-0.225, 0.660] 0.2505229
Resistin - plasma	-0.064 [-0.320, 0.222] 0.653606	-0.181 [-0.461, 0.145] 0.202636	-0.100 [-0.355, 0.171] 0.486370	0.113 [-0.160, 0.350] 0.429731	0.099 [-0.174, 0.363] 0.496551	-0.112 [-0.392, 0.191] 0.435977	-0.199 [-0.482, 0.068] 0.161904	-0.194 [-0.455, 0.087] 0.172896	-0.295 [-0.715, 0.123] 0.176056	0.236 [-0.557, 0.115] 0.2122777	-0.185 [-0.596, 0.342] 0.3776646

Supplementary Table S6. Association between cytokines and Knee Osteoarthritis (KOA) outcomes in the KOIP-3 group. Table cells show non-parametric correlation-like measures and 95% Confidence Intervals (95%CI) between cytokines and clinical, radiographic and ultrasound severity at baseline and radiographic progression. Correlation-like measures are: Spearman correlation (continuous or ordinal outcomes) and Glass rank biserial correlation (binary outcomes). Radiographic progression according to Kellgren-Lawrence staging and joint space narrowing were treated as ordinal in these analyses. Red color indicates positive correlation, blue represents negative correlation, and color intensity expresses more extreme values of the correlation coefficients. Color intensities were saturated to 0.5 and -0.5 for positive and negative correlation, respectively. **IL-6:** Interleukin 6; **IL-8:** Interleukin 8; **TNF-alpha:** Tumor Necrosis Factor alpha; **NGF:** Nerve Growth Factor; **CRP:** C-Reactive Protein; **KOA:** Knee Osteoarthritis. **KOIP:** Knee Osteoarthritis Inflammatory Phenotype.

	KOOS pain	KOOS symptoms	KOOS functional disability	Joint effusion	Synovial tissue thickness	Kellgren-Lawrence Baseline	Osteophytes Baseline	Joint space narrowing Baseline	Kellgren-Lawrence Progression	Osteophytes Progression	Joint space narrowing Progression
CRP - synovial fluid	0.230 [-0.194, 0.591] 0.248541	0.012 [-0.402, 0.396] 0.952875	0.117 [-0.285, 0.504] 0.56014662	0.213 [-0.194, 0.534] 0.286561	0.078 [-0.345, 0.456] 0.698219	0.197 [-0.194, 0.531] 0.324519	0.006 [-0.410, 0.480] 0.9743342	-0.015 [-0.415, 0.392] 0.940895	-0.582 [NA, NA] 0.035734	0.182 [-0.400, 0.722] 0.4813214	0.061 [-0.618, 0.706] 0.8229019
CRP - plasma	0.261 [-0.229, 0.596] 0.188318	0.157 [-0.266, 0.552] 0.435394	0.196 [-0.271, 0.595] 0.32842050	0.321 [-0.071, 0.599] 0.069830	0.354 [-0.002, 0.618] 0.249464	0.229 [-0.145, 0.565] 0.249803	0.095 [-0.320, 0.511] 0.6378534	-0.058 [-0.487, 0.365] 0.773816	-0.429 [-0.919, 0.131] 0.122289	-0.109 [-0.636, 0.461] 0.6726548	0.204 [NA, NA] 0.4556390
IL-6 - synovial fluid	0.070 [-0.374, 0.513] 0.730124	0.099 [-0.294, 0.473] 0.621531	-0.138 [-0.529, 0.286] 0.49322508	0.541 [0.255, 0.731] 0.003598	0.125 [-0.318, 0.522] 0.533631	0.250 [-0.162, 0.562] 0.208900	0.002 [-0.309, 0.375] 0.9938882	0.102 [-0.294, 0.483] 0.612776	-0.231 [-0.750, 0.333] 0.405381	0.273 [-0.245, 0.769] 0.2908457	0.184 [-0.408, 0.778] 0.5019387
IL-6 - plasma	0.098 [-0.367, 0.544] 0.634323	0.108 [-0.330, 0.487] 0.599393	-0.213 [-0.565, 0.202] 0.29583152	0.067 [-0.345, 0.424] 0.746133	-0.237 [-0.615, 0.168] 0.243464	0.070 [-0.300, 0.393] 0.7374701	-0.069 [-0.483, 0.376] 0.995840	0.001 [-0.400, 0.397] 0.932647	-0.024 [-0.514, 0.595] 0.932647	-0.273 [-0.758, 0.109] 0.3050589	0.381 [NA, NA] 0.1869493
IL-8 - synovial fluid	0.222 [-0.229, 0.596] 0.265699	0.329 [-0.110, 0.657] 0.093377	0.062 [-0.429, 0.464] 0.75747769	0.372 [0.039, 0.637] 0.169035	0.273 [-0.143, 0.613] 0.169035	0.115 [-0.203, 0.699] 0.281185	0.286 [-0.049, 0.584] 0.1483444	0.146 [-0.254, 0.518] 0.468515	0.363 [-0.187, 0.820] 0.191040	0.109 [-0.418, 0.655] 0.6726548	0.551 [0.067, 0.918] 0.4379752
IL-8 - plasma	0.104 [-0.329, 0.494] 0.604240	0.045 [-0.314, 0.401] 0.821852	-0.049 [-0.415, 0.353] 0.80987602	-0.287 [-0.584, 0.068] 0.146647	-0.144 [-0.618, 0.308] 0.474399	-0.158 [-0.520, 0.229] 0.432082	0.200 [-0.192, 0.584] 0.3173169	0.084 [-0.285, 0.462] 0.675329	0.538 [0.000, 0.938] 0.052204	0.055 [-0.469, 0.578] 0.8326894	0.531 [-0.022, 1.000] 0.0521434
TNF-alpha - synovial fluid	-0.030 [-0.407, 0.374] 0.811729	-0.018 [-0.415, 0.350] 0.929964	-0.163 [-0.566, 0.286] 0.41540770	-0.051 [-0.459, 0.376] 0.800525	-0.093 [-0.481, 0.368] 0.646096	-0.251 [-0.591, 0.093] 0.206947	0.020 [-0.329, 0.346] 0.9218854	-0.295 [-0.619, 0.057] 0.135553	0.275 [-0.238, 0.700] 0.321912	0.291 [-0.204, 0.763] 0.5298752	0.531 [NA, NA] 0.0521434
TNF-alpha - plasma	-0.123 [-0.491, 0.267] 0.549624	0.095 [-0.291, 0.490] 0.643169	-0.224 [-0.580, 0.213] 0.27036970	0.124 [-0.301, 0.492] 0.545621	-0.019 [-0.364, 0.335] 0.925976	-0.264 [-0.595, 0.105] 0.192837	-0.408 [-0.694, -0.057] 0.0386679	-0.274 [-0.660, 0.121] 0.175237	-0.190 [-0.762, 0.428] 0.498962	-0.434 [-0.858, 0.125] 0.1023776	0.071 [-0.524, 0.667] 0.8045709
NGF - synovial fluid	-0.148 [-0.520, 0.257] 0.461117	0.010 [-0.420, 0.461] 0.960321	0.172 [-0.272, 0.576] 0.39219769	-0.014 [-0.434, 0.359] 0.946414	0.033 [-0.317, 0.370] 0.869517	0.144 [-0.295, 0.539] 0.475105	0.194 [-0.175, 0.539] 0.496702	-0.137 [-0.255, 0.486] 0.747662	-0.088 [-0.625, 0.475] 0.372652	-0.227 [-0.718, 0.275] 0.4498428	-0.204 [-0.704, 0.275] 0.4498428
NGF - plasma	-0.339 [-0.631, 0.037] 0.084025	-0.478 [-0.784, -0.062] 0.011621	-0.621 [-0.800, -0.352] 0.00054013	0.284 [-0.139, 0.676] 0.510438	0.194 [-0.148, 0.513] 0.332852	0.111 [-0.273, 0.504] 0.582444	0.100 [-0.311, 0.463] 0.6207338	0.141 [-0.256, 0.463] 0.482475	0.132 [-0.677, 0.437] 0.631365	-0.145 [-0.654, 0.364] 0.5701812	-0.276 [-0.731, 0.231] 0.3100560
Calprotectin - synovial fluid	-0.070 [-0.362, 0.371] 0.727834	0.104 [-0.327, 0.509] 0.604377	0.047 [-0.419, 0.479] 0.81577987	0.255 [-0.119, 0.560] 0.199743	0.150 [-0.271, 0.518] 0.456147	0.069 [-0.331, 0.435] 0.733511	0.008 [-0.381, 0.376] 0.9682261	-0.159 [-0.542, 0.261] 0.427295	0.011 [-0.531, 0.548] 0.966395	0.091 [-0.491, 0.618] 0.7247712	0.245 [NA, NA] 0.3706500
Calprotectin - plasma	0.014 [-0.370, 0.406] 0.944335	-0.109 [-0.497, 0.267] 0.589533	-0.143 [-0.533, 0.294] 0.47691063	-0.032 [-0.435, 0.386] 0.875019	0.096 [-0.309, 0.488] 0.634117	0.278 [-0.207, 0.665] 0.159925	0.450 [0.096, 0.738] 0.0184255	0.364 [-0.001, 0.711] 0.0661790	0.143 [-0.455, 0.750] 0.606498	0.473 [0.000, 0.852] 0.0671207	0.265 [-0.293, 0.782] 0.3321157
Leptin - synovial fluid	-0.013 [-0.409, 0.409] 0.946752	0.008 [-0.374, 0.397] 0.969782	0.114 [-0.312, 0.527] 0.57047679	0.057 [-0.280, 0.400] 0.778359	-0.034 [-0.434, 0.354] 0.865467	0.070 [-0.334, 0.449] 0.727033	-0.437 [-0.714, -0.036] 0.0226561	-0.071 [-0.487, 0.322] 0.723484	-0.209 [-0.714, 0.319] 0.451566	-0.236 [-0.740, 0.259] 0.3599647	0.082 [NA, NA] 0.7653835
Leptin - plasma	-0.007 [-0.387, 0.379] 0.973362	0.068 [-0.305, 0.413] 0.735142	-0.049 [-0.358, 0.451] 0.75866749	-0.049 [-0.452, 0.367] 0.808731	0.126 [-0.258, 0.506] 0.532626	0.122 [-0.287, 0.565] 0.543971	-0.448 [-0.694, -0.081] 0.0191592	-0.407 [-0.497, 0.302] 0.602765	-0.105 [-0.850, 0.000] 0.027650	-0.143 [-0.904, 0.000] 0.6015081	
Irisin - synovial fluid	0.013 [-0.410, 0.381] 0.947960	0.075 [-0.322, 0.472] 0.947960	0.097 [-0.450, 0.294] 0.63091845	0.224 [-0.170, 0.550] 0.262268	0.116 [-0.341, 0.496] 0.563177	0.144 [-0.325, 0.583] 0.473691	-0.353 [-0.700, 0.069] 0.0713304	-0.319 [-0.401, 0.409] 0.948429	-0.132 [0.025, 0.450] 0.250547	-0.582 [0.027, 0.450] 0.0242357	0.122 [-0.425, 0.615] 0.6544215
Irisin - plasma	-0.112 [-0.491, 0.331] 0.579430	0.067 [-0.341, 0.463] 0.740877	0.049 [-0.337, 0.432] 0.80990452	0.124 [-0.277, 0.502] 0.537725	-0.026 [-0.445, 0.425] 0.898903	0.206 [-0.212, 0.588] 0.302407	-0.306 [-0.651, 0.076] 0.1204924	0.062 [-0.352, 0.460] 0.756950	-0.429 [-0.868, 0.151] 0.122289	-0.436 [-0.855, 0.133] 0.0910224	-0.388 [-0.857, 0.206] 0.1563939
Adiponectin - synovial fluid	0.024 [-0.363, 0.358] 0.906950	0.055 [-0.290, 0.424] 0.76791642	-0.060 [-0.483, 0.346] 0.76791642	0.265 [-0.145, 0.608] 0.181916	-0.129 [-0.492, 0.302] 0.522620	0.234 [-0.193, 0.641] 0.239710	0.206 [-0.480, 0.297] 0.6978681	-0.078 [-0.183, 0.598] 0.178132	-0.407 [-0.875, 0.140] 0.142650	0.182 [-0.353, 0.666] 0.4813214	-0.020 [NA, NA] 0.9405292
Adiponectin - plasma	-0.033 [-0.417, 0.362] 0.869757	0.112 [-0.243, 0.437] 0.579028	0.156 [-0.297, 0.559] 0.43782418	-0.096 [-0.481, 0.278] 0.635255	-0.355 [-0.652, 0.021] 0.069233	0.280 [-0.201, 0.699] 0.157229	-0.161 [-0.527, 0.256] 0.4217065	0.240 [-0.218, 0.604] 0.227355	-0.495 [-1.000, 0.142] 0.074892	-0.127 [-0.596, 0.407] 0.6220655	-0.551 [-1.000, 0.050] 0.0439752
Omentin - synovial fluid	0.165 [-0.198, 0.486] 0.410943	0.237 [-0.115, 0.553] 0.179903	0.093 [-0.172, 0.579] 0.23390323	0.093 [-0.305, 0.462] 0.643962	-0.226 [-0.574, 0.206] 0.256878	0.160 [-0.217, 0.537] 0.426532	-0.219 [-0.578, 0.198] 0.2721665	0.169 [-0.176, 0.520] 0.398809	0.209 [NA, NA] 0.451566	0.182 [-0.363, 0.694] 0.4813214	-0.020 [-0.667, 0.646] 0.9405292
Omentin - plasma	-0.190 [-0.550, 0.214] 0.341452	0.099 [-0.268, 0.449] 0.622610	0.066 [-0.337, 0.474] 0.74256610	-0.041 [-0.422, 0.342] 0.840559	-0.009 [-0.396, 0.392] 0.962609	0.235 [-0.178, 0.581] 0.238994	-0.283 [-0.638, 0.137] 0.1523455	0.052 [-0.364, 0.439] 0.795726	-0.604 [-1.000, 0.020] 0.0203418	-0.291 [-0.799, 0.241] 0.2598752	-0.408 [NA, NA] 0.1356745
Osteopontin - synovial fluid	-0.078 [-0.478, 0.342] 0.699415	-0.140 [-0.532, 0.332] 0.487456	-0.129 [-0.524, 0.308] 0.52268434	0.184 [-0.220, 0.564] 0.357763	0.168 [-0.264, 0.545] 0.401329	0.143 [-0.342, 0.543] 0.479928	0.035 [-0.387, 0.431] 0.8625164	-0.045 [-0.485, 0.376] 0.822667	0.033 [NA, NA] 0.905383	-0.236 [-0.714, 0.327] 0.3599647	0.714 [0.345, 0.978] 0.0090234
Osteopontin - plasma	0.253 [-0.195, 0.619] 0.202447	0.079 [-0.344, 0.522] 0.694270	0.205 [-0.166, 0.516] 0.30587029	0.012 [-0.368, 0.412] 0.954181	0.119 [-0.291, 0.468] 0.553929	0.484 [0.088, 0.759] 0.010467	0.294 [-0.111, 0.609] 0.1366053	0.450 [0.071, 0.745] 0.018620	-0.509 [-0.824, 0.187] 0.191040	-0.306 [-0.625, 0.442] 0.7247712	0.306 [-0.300, 0.827] 0.2631100
Visfatin - synovial fluid	-0.112 [-0.467, 0.263] 0.577334	-0.204 [-0.556, 0.251] 0.308513	-0.156 [-0.552, 0.312] 0.43691518	0.010 [-0.353, 0.390] 0.961409	-0.108 [-0.433, 0.267] 0.592386	0.236 [-0.127, 0.568] 0.236853	0.144 [-0.245, 0.540] 0.4748702	0.233 [-0.132, 0.585] 0.242943	-0.231 [NA, NA] 0.405381	0.309 [-0.218, 0.745] 0.2312660	0.204 [-0.375, 0.789] 0.4556390
Visfatin - plasma	0.110 [-0.281, 0.463] 0.584682	0.107 [-0.275, 0.470] 0.594816	0.207 [-0.164, 0.511] 0.30000485	-0.057 [-0.481, 0.396] 0.778359	-0.043 [-0.460, 0.357] 0.831075	-0.024 [-0.395, 0.335] 0.905655	0.043 [-0.374, 0.443] 0.8312879	0.117 [-0.251, 0.489] 0.562360	0.165 [-0.438, 0.736] 0.452297	0.218 [-0.350, 0.709] 0.3981027	-0.082 [NA, NA] 0.7653835
Resistin - synovial fluid	-0.130 [-0.482, 0.261] 0.517024	0.132 [-0.294, 0.513] 0.517024	0.241 [-0.172, 0.593] 0.22615963	-0.146 [-0.549, 0.295] 0.466635	-0.018 [-0.392, 0.429] 0.930092	-0.085 [-0.411, 0.246] 0.672054	0.070 [-0.425, 0.554] 0.7276493	-0.201 [-0.559, 0.246] 0.306347	0.033 [NA, NA] 0.905383	-0.209 [NA, NA] 0.4179907	0.184 [-0.367, 0.725] 0.5017999
Resistin - plasma	0.068 [-0.357, 0.478] 0.734709	0.188 [-0.203, 0.549] 0.347853	0.206 [-0.167, 0.537] 0.30292871	-0.265 [-0.604, 0.135] 0.180873	-0.347 [-0.649, 0.051] 0.076426	-0.128 [-0.502, 0.297] 0.525673	0.269 [-0.126, 0.596] 0.1749467	-0.033 [-0.399, 0.343] 0.872137	0.319 [NA, NA] 0.250547	0.182 [-0.333, 0.709] 0.4813214	-0.082 [-0.684, 0.537] 0.7653835

Supplementary Table S8. Association of synovial and plasma cytokines with clinical severity in Knee Osteoarthritis Inflammatory Phenotypes (KOIP), adjusted for age, time of disease's evolution and Body Mass Index (BMI). The table presents Spearman correlation coefficients, 95% confidence intervals (95% CI) and p-values, illustrating the association between Knee Osteoarthritis (KOA) severity measures (KOOS-pain, KOOS-functional disability, and ultrasound joint effusion) and selected cytokines analyzed in the study. The analyses are adjusted for age, disease duration, and BMI. Results are presented for the entire patient cohort as well as for individual KOIP groups. **KOA:** Knee Osteoarthritis; **KOIP:** Knee Osteoarthritis Inflammatory Phenotypes (KOIP); **KOOS:** Knee injury and Osteoarthritis Outcome Scores (reversed scores); **95%CI:** 95% confidence interval.

KOA Severity	Cytokine	All		KOIP-1		KOIP-2		KOIP-3		KOIP-4	
		Spearman correlation [95%CI]	P-value	Spearman correlation [95%CI]	P-value	Spearman correlation [95%CI]	P-value	Spearman correlation [95%CI]	P-value	Spearman correlation [95%CI]	P-value
KOOS - Pain (reversed, 0-100)	Omentin - synovial fluid (pg/mL)	0.099 [-0.057, 0.249]	0.21221	-0.241 [-0.487, 0.040]	0.092329	0.300 [-0.025, 0.567]	0.070156	0.146 [-0.209, 0.467]	0.42272	0.231 [-0.199, 0.586]	0.29104
	Osteopontin - synovial fluid (ng/mL)	0.206 [0.055, 0.348]	0.007798	0.118 [-0.176, 0.393]	0.43198	0.114 [-0.207, 0.413]	0.48995	0.098 [-0.349, 0.508]	0.67805	0.461 [0.153, 0.687]	0.0044899
	Interleukin 6 - plasma (pg/mL)	0.007 [-0.147, 0.160]	0.93076	0.047 [-0.240, 0.327]	0.75032	0.027 [-0.261, 0.310]	0.85728	0.130 [-0.366, 0.568]	0.61906	-0.328 [-0.634, 0.067]	0.10145
KOOS - Functional Disability (reversed, 0-100)	Omentin - synovial fluid (pg/mL)	0.060 [-0.095, 0.213]	0.44518	-0.202 [-0.446, 0.069]	0.14207	0.247 [-0.079, 0.524]	0.13584	0.194 [-0.207, 0.539]	0.34369	0.206 [-0.241, 0.581]	0.36769
	Interleukin 6 - plasma (pg/mL)	0.029 [-0.124, 0.182]	0.70824	0.164 [-0.122, 0.425]	0.26152	0.067 [-0.211, 0.335]	0.64094	-0.215 [-0.587, 0.233]	0.34812	-0.299 [-0.565, 0.023]	0.068168
Ultrasound Joint Effusion (mm)	C-reactive protein - synovial fluid (mg/L)	0.007 [-0.146, 0.160]	0.926	0.059 [-0.238, 0.346]	0.70114	0.041 [-0.251, 0.326]	0.78781	0.324 [-0.062, 0.626]	0.097677	-0.570 [-0.763, -0.284]	0.00035628
	C-reactive protein - plasma (mg/L)	-0.029 [-0.181, 0.124]	0.70813	-0.146 [-0.403, 0.133]	0.30445	-0.041 [-0.329, 0.254]	0.78949	0.412 [0.089, 0.657]	0.013912	-0.538 [-0.748, -0.231]	0.0012901
	Leptin - synovial fluid (pg/mL)	0.058 [-0.095, 0.209]	0.45711	-0.123 [-0.410, 0.187]	0.43981	-0.184 [-0.440, 0.099]	0.2007	0.030 [-0.410, 0.458]	0.89965	0.429 [0.091, 0.678]	0.014337
	Leptin - plasma (pg/mL)	-0.062 [-0.213, 0.093]	0.4338	-0.166 [-0.418, 0.110]	0.23865	-0.194 [-0.458, 0.101]	0.19584	-0.034 [-0.420, 0.363]	0.87195	0.253 [-0.099, 0.548]	0.15632
	Irisin - synovial fluid (ng/mL)	0.019 [-0.136, 0.173]	0.81176	-0.183 [-0.434, 0.095]	0.19624	-0.107 [-0.420, 0.228]	0.53419	0.259 [-0.151, 0.593]	0.2124	0.326 [-0.021, 0.603]	0.064766
	Interleukin 6 - synovial fluid (pg/mL)	0.236 [0.086, 0.376]	0.0023444	0.103 [-0.169, 0.361]	0.45915	0.157 [-0.144, 0.430]	0.30618	0.634 [0.390, 0.795]	0.000012813	0.355 [0.035, 0.608]	0.030222
	Interleukin 6 - plasma (pg/mL)	0.055 [-0.099, 0.206]	0.48514	0.118 [-0.169, 0.387]	0.42041	0.142 [-0.149, 0.409]	0.33893	0.051 [-0.351, 0.437]	0.80907	-0.321 [-0.598, 0.025]	0.068546
	Calprotectin - plasma (ng/mL)	-0.003 [-0.156, 0.150]	0.96899	-0.044 [-0.306, 0.224]	0.75206	0.141 [-0.132, 0.394]	0.3115	-0.052 [-0.438, 0.349]	0.8052	-0.308 [-0.554, -0.013]	0.040914
	Omentin - synovial fluid (pg/mL)	0.162 [0.008, 0.308]	0.040004	0.098 [-0.190, 0.370]	0.50697	0.129 [-0.184, 0.418]	0.42145	0.123 [-0.270, 0.480]	0.54479	0.362 [0.021, 0.628]	0.038155

Supplementary Table S9. Association of synovial and plasma cytokines with radiographic progression in Knee Osteoarthritis Inflammatory Phenotypes (KOIP), adjusted for age, time of disease's evolution and Body Mass Index (BMI). The table presents fold-changes, 95% confidence intervals (95% CI) and p-values, illustrating the association between Knee Osteoarthritis (KOA) progression according to different radiographic criteria (Kellgren-Lawrence, Osteophytes and Joint Space Narrowing) and selected cytokines analyzed in the study. Positive fold-changes indicate a higher level of the cytokine in progressors, while negative fold-changes represents a higher level of the cytokine in non-progressors. The analyses are adjusted for age, disease duration, and BMI. Results are presented for the entire patient cohort as well as for individual KOIP groups. **KOA:** Knee Osteoarthritis; **KOIP:** Knee Osteoarthritis Inflammatory Phenotypes (KOIP); **95%CI:** 95% confidence interval.

KOA Radiographic Progression	Cytokines	All		KOIP-1		KOIP-2		KOIP-3		KOIP-4	
		Fold-Change [95%CI]	P-value	Fold-Change [95%CI]	P-value	Fold-Change [95%CI]	P-value	Fold-Change [95%CI]	P-value	Fold-Change [95%CI]	P-value
Kellgren-Lawrence Radiographic Progression	Resistin - synovial fluid (pg/mL)	-1.24 [-1.74, 1.18]	0.24462	-2.08 [-3.65, -1.20]	0.012175	1.51 [-1.41, 2.89]	0.27554	1.31 [-2.27, 3.79]	0.58863	-1.24 [-2.91, 1.88]	0.63494
	C-reactive protein - synovial fluid (mg/L)	1.12 [-1.16, 1.49]	0.39035	1.51 [1.05, 2.19]	0.037626	1.06 [-1.70, 1.81]	0.85564	-1.57 [-2.92, 1.13]	0.11665	1.11 [-1.82, 2.06]	0.75714
	Omentin - plasma (pg/mL)	-1.26 [-1.64, 1.04]	0.076489	-1.40 [-2.20, 1.07]	0.12058	1.06 [-1.45, 1.56]	0.81773	-2.70 [-7.00, -1.31]	0.009627	1.05 [-1.66, 1.70]	0.86185
	Irisin - synovial fluid (ng/mL)	-1.04 [-1.18, 1.10]	0.60394	-1.25 [-1.47, -1.06]	0.008212	-1.07 [-1.49, 1.30]	0.68488	-1.16 [-1.74, 1.31]	0.47166	1.39 [1.14, 1.66]	0.0012974
Osteophytes Radiographic Progression	Osteopontin - synovial fluid (ng/mL)	-1.21 [-1.63, 1.10]	0.19176	1.37 [-1.36, 2.54]	0.33122	-1.45 [-2.30, 1.09]	0.12429	-1.10 [-2.64, 2.15]	0.78894	-1.10 [-1.78, 1.41]	0.64559
	Osteopontin - plasma (ng/mL)	-1.14 [-1.37, 1.04]	0.14962	-1.05 [-1.58, 1.40]	0.78026	-1.42 [-1.98, -1.04]	0.031466	-1.38 [-2.66, 1.42]	0.33039	1.11 [-1.27, 1.60]	0.55478
	Omentin - synovial fluid (pg/mL)	1.02 [-1.32, 1.42]	0.88754	1.85 [1.08, 3.37]	0.028051	-1.72 [-2.92, -1.06]	0.03584	1.05 [-3.20, 3.59]	0.90198	1.41 [-1.36, 2.79]	0.28672
	Omentin - plasma (pg/mL)	1.02 [-1.25, 1.31]	0.86418	1.86 [1.10, 3.64]	0.030952	-1.15 [-1.64, 1.26]	0.46302	-1.76 [-5.02, 1.61]	0.24345	1.36 [-1.17, 2.15]	0.19517
	Leptin - synovial fluid (pg/mL)	1.05 [-1.19, 1.30]	0.68843	1.74 [1.03, 2.94]	0.034066	-1.15 [-1.53, 1.19]	0.38946	-1.08 [-2.10, 1.75]	0.80688	-1.41 [-2.35, 1.16]	0.19797
	Leptin - plasma (pg/mL)	1.05 [-1.14, 1.26]	0.61333	1.43 [1.02, 2.01]	0.044663	-1.05 [-1.50, 1.39]	0.84667	-1.69 [-3.47, 1.37]	0.1892	1.02 [-1.37, 1.43]	0.92611
	Irisin - synovial fluid (ng/mL)	-1.08 [-1.22, 1.05]	0.27223	1.09 [-1.13, 1.34]	0.40431	-1.19 [-1.61, 1.12]	0.23591	-1.68 [-2.64, -1.09]	0.031111	1.19 [-1.00, 1.43]	0.066103
	Irisin - plasma (ng/mL)	-1.03 [-1.17, 1.11]	0.66774	1.17 [-1.02, 1.41]	0.087542	-1.15 [-1.59, 1.25]	0.41349	-1.34 [-2.34, 1.24]	0.22328	1.02 [-1.23, 1.28]	0.84938
	Interleukin 8 - plasma (pg/mL)	1.21 [-1.09, 1.55]	0.1495	1.25 [-1.40, 2.05]	0.35809	1.53 [1.06, 2.23]	0.026868	1.49 [-2.98, 6.09]	0.57305	-1.55 [-2.44, 1.01]	0.045448
	Calprotectin - synovial fluid (ng/mL)	1.33 [-1.06, 1.91]	0.12387	-1.30 [-2.31, 1.46]	0.41521	1.45 [-1.50, 3.34]	0.3442	2.30 [-1.03, 5.71]	0.066658	2.09 [1.05, 4.29]	0.041223
	Adiponectin - synovial fluid (ng/mL)	1.14 [-1.16, 1.48]	0.36748	1.66 [1.07, 2.51]	0.032595	-1.47 [-2.35, 1.06]	0.099879	-1.05 [-2.86, 2.52]	0.90321	1.16 [-1.47, 2.06]	0.54318
	Joint Space Narrowing Radiographic Progression	Osteopontin - synovial fluid (ng/mL)	1.26 [-1.04, 1.68]	0.108	1.31 [-1.28, 2.14]	0.25957	1.02 [-1.62, 1.79]	0.87745	2.59 [1.68, 3.99]	0.00034539	-1.32 [-2.04, 1.17]
JoOmentin - synovial fluid (pg/mL)		1.24 [-1.08, 1.65]	0.16296	1.67 [1.12, 2.50]	0.010921	1.22 [-1.57, 2.07]	0.50537	-1.04 [-2.67, 2.33]	0.94272	1.05 [-1.90, 1.98]	0.89549
Leptin - synovial fluid (pg/mL)		-1.04 [-1.30, 1.22]	0.74315	-1.06 [-1.56, 1.34]	0.73046	-1.67 [-2.30, -1.20]	0.0013274	1.15 [-1.43, 1.84]	0.59876	1.16 [-1.59, 1.97]	0.54063
Leptin - plasma (pg/mL)		-1.14 [-1.42, 1.05]	0.15722	-1.07 [-1.44, 1.22]	0.55993	-1.60 [-2.32, -1.09]	0.016944	-1.26 [-2.31, 1.41]	0.46167	-1.07 [-1.54, 1.32]	0.66752
Irisin - synovial fluid (ng/mL)		1.01 [-1.11, 1.14]	0.84859	-1.12 [-1.30, 1.06]	0.15696	-1.14 [-1.55, 1.21]	0.40734	1.20 [-1.23, 1.76]	0.34398	1.26 [1.05, 1.49]	0.012932
Irisin - plasma (ng/mL)		-1.16 [-1.33, -1.02]	0.021346	-1.10 [-1.24, 1.03]	0.14487	-1.54 [-2.67, -1.01]	0.045653	-1.22 [-1.92, 1.20]	0.28199	-1.15 [-1.51, 1.11]	0.26456