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Data Article

Prediction of transcription factor bindings sites affected by SNPs located at the osteopontin promoter



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ABSTRACT

This data contains information related to the research article entitled “Osteopontin splice variants and polymorphisms in Cancer Progression and Prognosis” [1]. Here, we describe an in silico analysis of transcription factors that could have altered binding to their DNA target sequence as a result of SNPs in the osteopontin gene promoter. We concentrated on SNPs associated with cancer risk and development.

The analysis was performed with PROMO v3.0.2 software which incorporates TRANSFACT v6.4 of. We also present a figure depicting the putative transcription factor binding according to genotype.

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Specifications Table

Subject area	<i>Biology, Molecular Biology</i>
More specific subject area	<i>Effect of SNPs in binding of transcription factors for the gene osteopontin</i>
Type of data	<i>Table and figure</i>
How data was acquired	<i>Software PROMO 3.0.2 (using TRANSFAC v.6.4)</i>
Data format	<i>Analyzed</i>
Experimental factors	<i>SNPs sequences were obtained from NCBI Single Nucleotide Polymorphism Database (dbSNP). PROMO parameters were chosen for human sequences and human sites.</i>
Experimental features	<i>SNPs located in OPN promoter with an effect in cancer risk and prognosis were analyzed to compare which transcription factors are binding in the variant sequences.</i>
Data source location	
Data accessibility	<i>The data is available in this article</i>

Value of the data

- These data describe how putative DNA-binding sites for transcriptional factors can be created or interrupted by the changes in sequences generated by SNPs in the promoter of osteopontin.
- Differential binding among SNPs genotypes can potentially explain why these SNPs have been associated with changes in the risk of cancer for a specific population.
- This analysis is an example of how important databases, such as those containing SNP genotypes and the predictive tools for DNA-binding sites for transcriptional factors in a specific sequence, could be used to try to select potential signaling pathways modulating the development of cancer.

1. Data

The table provided in this article is a list of the transcription factors predicted to bind a DNA sequence at the SNPs contained in the osteopontin promoter. We analyzed only those SNPs that statistically in a population have been shown to have an effect on cancer risk and prognosis for the carriers. For each SNP we present both sequences. Each analysis contains the rs ID and the nucleotide position in reference to the osteopontin promoter; a schematic representation of the binding of the transcription factor to their target sequence; and an analysis of how similar the binding site is compared to its canonical binding sequence.

2. Experimental design, materials and methods

Analysis of SNP sequences was performed using software PROMO v3.0.2, (which utilizes TRANSFAC v6.4) [2,3] For each osteopontin gene promoter SNP, the sequences carrying each allele were loaded as the query sequence to search for potential binding sites. The prediction was carried out considering only sites and only human transcription factors. The output of this analysis is presented in Table 1. Each analysis contains the rs that corresponds to each SNP and its position relative to the transcription start site of osteopontin. For each SNP, we present the respective results for both sequences loaded as the query sequences. A schematic representation (boxes in color, also indicated with numbers) of the binding of the transcription factor to the target sequence, and a list of the putative transcription factors binding to the sequence. For each transcription factor site, several predicted parameters are reported. The *transcription*

Table 1 (continued)

rs29001511						rs2853744							
ACAGAGTAAACTACAGTAAATCTCGTGTGGAAATTTGTTGTTTGAATTT						ACAGCCCTCAAGCAGTCTCTGCTCTCAGTCAAGAACTGTTTACT							
-1776 C						-719 G							
ACAGAGTAAACTACAGTAAATCTCGGAAATTTGTTGTTTGAATTT						ACGAGCCCTCAAGCAGTCTCTCTCAGTCAAGAACTGTTTACT							
Factor name	Start position	End position	Dissimilarity	String	RE equally	RE query	Factor name	Start position	End position	Dissimilarity	String	RE equally	RE query
RFX1 [T01673]	1	6	0	CAGAGT	0.03735	0.01628	Pax-5 [T00070]	1	7	8.014558	CGAGCC	0.05493	0.04514
c-Myb [T00137]	7	14	14.265129	AAACTACA	0.00934	0.01497	p53 [T00671]	1	7	6.563521	CGAGCC	0.01221	0.01015
HNF-3alpha [T02512]	14	21	14.000258	AGTAATCT	0.04207	0.1033	AP-1 [T00029]	12	20	10.480716	GAGCAGTCA	0.00389	0.00925
HNF-3alpha [T02512]	29	36	7.000129	AATTTTGT	0.02101	0.12436	AP-1 [T00029]	26	34	6.527374	TCTCAGTCA	0.00458	0.00406
GR-beta [T01920]	18	22	5.042296	AATCC	0.09961	0.1703	c-Jun [T00133]	12	20	12.85308	AGGAGCTCA	0.0061	0.00524
GR-beta [T01920]	28	32	0	AAAT	0.09961	0.39325	c-Jun [T00133]	26	34	8.71589	TCTCAGTCA	0.00191	0.00171
GR-beta [T01920]	29	33	0	AATTT	0.09961	0.39325	c-Myb [T00137]	16	24	10.535275	AGTCAATCG	0.00687	0.00932
GR-beta [T01920]	45	49	1.680765	GAAT	0.09961	0.16207	E2F-1 [T01542]	18	25	10.630964	TCTCCGC	0.01526	0.01801
GR-beta [T01920]	46	50	0	AATTT	0.09961	0.39325	RFX1 [T01673]	33	38	9.512894	CAGAAA	0.18311	0.16592
GR [T05076]	35	41	8.033921	GTTGTTT	0.03735	0.13533	RFX1 [T01673]	36	41	14.26934	AAACTG	0.14648	0.12731
GR [T05076]	37	43	9.33358	GTTGTTT	0.0498	0.20531	STAT6 [T01580]	33	42	0	CAGAAACTCG	0.00019	0.00014
E2F-1 [T01542]	24	31	5.846571	CGGGAAAT	0.00467	0.0009	c-Myb [T00137]	36	43	7.545286	AAACTGCT	0.01068	0.01095
GR-alpha [T00337]	21	25	8.073878	CTGC	0.19922	0.05788	GR-alpha [T00337]	6	10	0.207689	CTCT	0.19531	0.23223
							XBP-1 [T0108]	16	21	0	AGTCAT	0.02441	0.02415
							XBP-1 [T0108]	14	22	13.110131	CGAGTCATC	0.01869	0.02019
rs2853744						rs28537094							
ACGAGCCCTCAAGCAGTCTCTGCTCTCAGTCAAGAACTGTTTACT						CGAGAAAACCTCATGACACAATCTCGCCCTCTGTTGGTGAGGAT							
-719 G						-66 T							
ACGAGCCCTCAAGCAGTCTCTGCTCTCAGTCAAGAACTGTTTACT						CGAGAAAACCTCATGACACAATCTCGCCCTCTGTTGGTGAGGAT							
Factor name	Start position	End position	Dissimilarity	String	RE equally	RE query	Factor name	Start position	End position	Dissimilarity	String	RE equally	RE query
Pax-5 [T00070]	1	7	8.014558	CGAGCC	0.05493	0.04514	RFX1 [T01673]	1	6	9.512894	CAGAAA	0.18677	0.19143
p53 [T00671]	1	7	6.563521	CGAGCC	0.01221	0.01015	RFX1 [T01673]	31	36	9.512894	TCCGTC	0.18677	0.19143
AP-1 [T00029]	12	20	10.480716	GAGCAGTCA	0.00389	0.00925	STAT6 [T01580]	1	14	2.886065	CAGAAACC	0.00389	0.00308
AP-1 [T00029]	26	34	6.527374	TCTCAGTCA	0.00458	0.00406	XBP-1 [T00902]	8	13	7.365101	CTCCAT	0.14941	0.14023
c-Jun [T00133]	12	20	12.85308	AGGAGCTCA	0.0061	0.00524	XBP-1 [T00902]	12	17	1.626297	ATGACA	0.0249	0.02299
c-Jun [T00133]	26	34	8.71589	TCTCAGTCA	0.00191	0.00171	XBP-1 [T0108]	11	19	7.615488	TGACACACA	0.00545	0.00649
PEA3 [T00685]	16	24	10.535275	AGTCAATCG	0.00687	0.00932	ATF3 [T01313]	13	20	6.744803	TGACACAA	0.007	0.00394
E2F-1 [T01542]	18	25	10.630964	TCTCCGC	0.01526	0.01801	C/EBPalpha [T00105]	13	21	3.367013	TGACACAT	0.00961	0.01014
RFX1 [T01673]	33	38	9.512894	CAGAAA	0.18311	0.16592	GR-beta [T01920]	19	23	3.361531	AATCT	0.09961	0.09961
RFX1 [T01673]	36	41	14.26934	AAACTG	0.14648	0.12731	E2F-1 [T01542]	22	29	11.323028	CTCCGGC	0.01245	0.01598
STAT6 [T01580]	33	42	0	CAGAAACTCG	0.00019	0.00014	USF1 [T00874]	16	20	7.629649	CACAA	0.0498	0.04713
c-Myb [T00137]	36	43	7.545286	AAACTGCT	0.01068	0.01095	USF1 [T00874]	34	38	6.294173	CTGTG	0.09961	0.10345
GR-alpha [T00337]	6	10	0.207689	CTCT	0.19531	0.23223	USF1 [T00874]	40	44	8.105784	TGGTGTG	0.09961	0.10144
GR-alpha [T00337]	22	26	0.207689	CTCT	0.19531	0.2548	NF-1 [T00539]	39	46	13.670907	TGGTGGA	0.0249	0.02574
XBP-1 [T00902]	16	21	0	AGTCAT	0.02441	0.02415	GR-alpha [T00337]	8	12	6.263098	CCTCA	0.09961	0.10144
XBP-1 [T0108]	14	22	13.110131	CGAGTCATC	0.01869	0.02019	GR-alpha [T00337]	29	33	8.281568	CTCTC	0.19922	0.22823
							GR-alpha [T00337]	33	37	0	CCTGT	0.19922	0.20028
							GR-alpha [T00337]	44	48	8.281568	CGAGG	0.19922	0.23809
							AP-2alpha [T00305]	28	33	4.22205	GCCTCC	0.0249	0.03119
							AP-2alpha [T02466]	28	33	1.576169	GCCTCC	0.03735	0.04471
							Fra-1 [T01462]	9	19	14.261792	CTCATGACAC	0.0019	0.00175
							JunB [T01977]	9	19	14.261792	CTCATGACAC	0.0019	0.00175

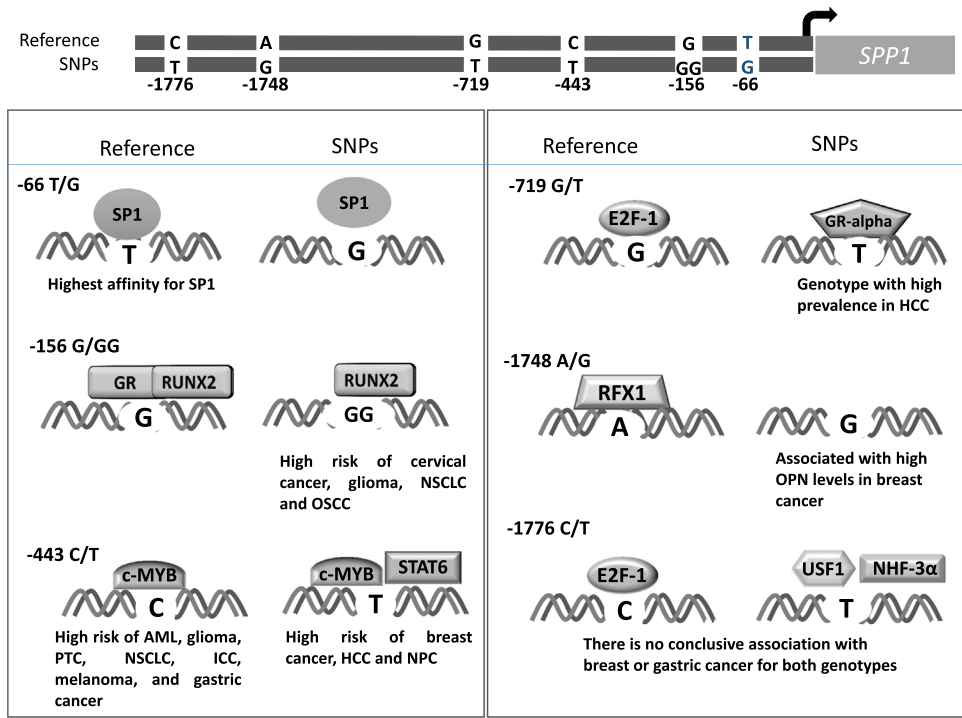


Fig. 1. Schematic representation of changes in transcription factors binding to SNPs located in the promoter of the osteopontin gene. At the top of the image there is a representation of the SNPs located in the osteopontin promoter that have been linked to variation in cancer risk in the carriers. The position of each SNP is given with respect to the transcription starting point. Below, for each SNP, the binding of the transcription factors and changes associated with altered genotype are exemplified.

factor name with the database accession number in brackets; the *start* and *end* positions of the putative binding sequences; *Dissimilarity* (%), which corresponds to the rate of dissimilarity between the putative and consensus sequences for a given transcription factor; *Sequence*, the nucleotide sequence of potential binding site; *Random Expectation (RE)* indicating the expected occurrences of the match in a random sequence of the same length as the query sequence according to the dissimilarity index, presented the *RE equally* (equi-probability for the four nucleotides) and *RE query* (nucleotide frequencies as in the query sequence). Markedly different changes are highlight in grey and the SNP is highlight in red. In Fig. 1 we depict the integration of information obtained from this predictive analysis and data previously reported for transcription factors binding to the osteopontin promoter.

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