

Association of the rs2234909 variant of the FGFR3 gene with colorectal cancer in a Mexican population





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Background

Colorectal cancer (CRC) ranks as the third most common malignancy worldwide and the leading cause of cancer-related mortality in 2024. While previous studies have explored the role of intracellular genes in CRC development, there is limited evidence regarding the involvement of fibroblast growth factor receptors (FGFR), particularly $FGFR3^{1,2}$.

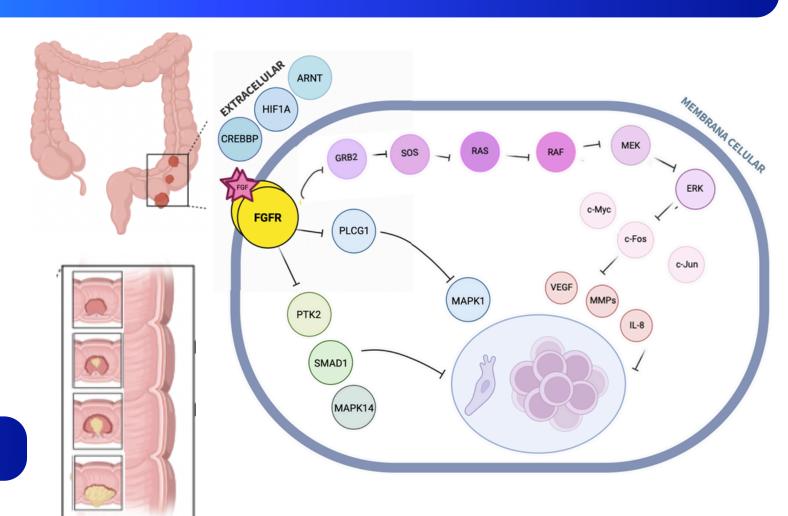
This gene, located on chromosome 4p16.3, consists of 19 exons and plays a key role in cell cycle regulation. Three high-variability regions, or hotspots, have been identified within the gene, with exon 7 being the most significant—accounting for approximately 88% of reported mutations^{3,4}.

Exon 7 encodes the IgIII domain, which is essential for the formation of receptor isoforms via alternative splicing, affecting ligand-binding specificity. The synonymous variant rs2234909, located in exon 7 (codon 294), involves a T>C transition (p.Asn294Asn)^{5,6}.

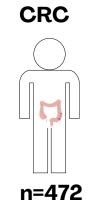
Although it does not change the amino acid sequence, it may promote disulfide bond formation between adjacent monomer receptors or induce structural changes in the tyrosine kinase domain, leading to ligand-independent dimerization^{7,8}.

Objetive

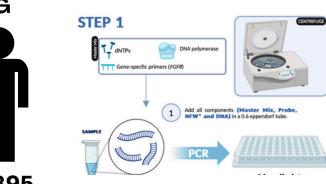
To analyze the association between the FGFR3 rs2234909 variant and CRC in the Mexican population.

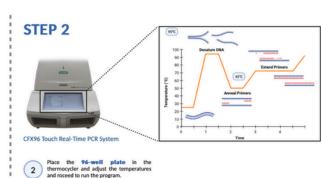


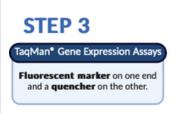
Materials and Methods

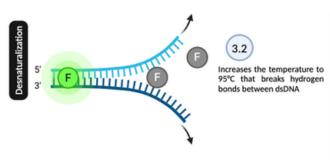


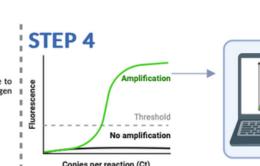


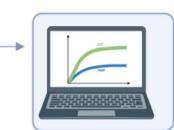










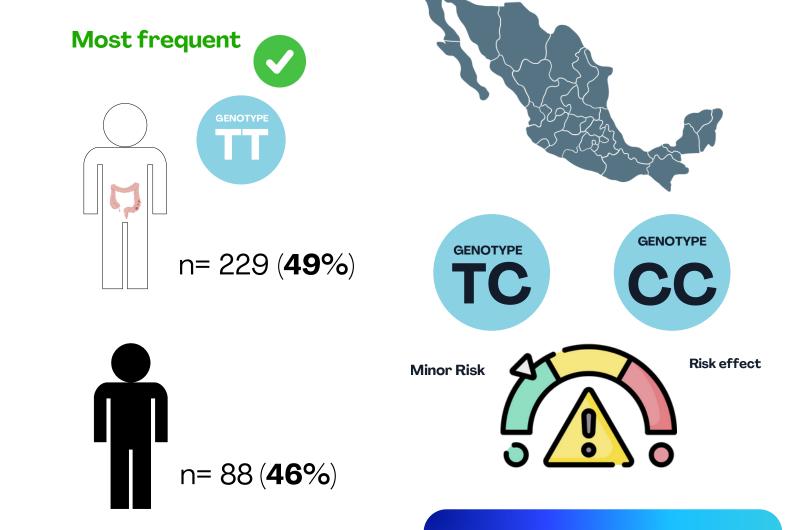


Results

rs2234909 (FGFR3)		Patient group CRC		Reference group		OR	95% (CI)	<i>P</i> -value
Model	Genotype	(n=472)	%	(n=395)	%			
	π	(243)	51	(224)	21	1.0 (reference)		
	TC	(108)	23	(147)	46	0.50	(0.37- 0.67)	0.001
	CC	(121)	26	(24)	6	5.34	(3.36- 8.47)	0.001
Dominant	π	(243)	51	(224)	57	1.0 (reference)		
	TC+CC	(229)	49	(171)	43	1.23	(0.94- 1.61)	0.1242
Recessive	СС	(121)	26	(24)	6	5.34	(3.36- 8.47)	0.001
	TT+TC	(351)	74	(371)	94	1.0 (reference)		

	Patient group CRC		Reference	group	OR	95%(CI)	P-value
Alelle	(2n=944)	%	(2n=790)	%			
A	(594)	63	(595)	73	0.55	(0.45- 0.68)	0.001
ALTERN G	(350)	37	(195)	27	1.79	(1.46- 2.21)	0.001

Conclusion



The rs2234909 variant of FGFR3 may serve as a potential genetic susceptibility marker for CRC in the Mexican population.

References

