

## Gene Section

### Short Communication

# GUCY2C (guanylate cyclase 2C (heat stable enterotoxin receptor))

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## Identity

**Other names:** GUC2C, STAR

**HGNC (Hugo):** GUCY2C

**Location:** 12p13.1

**Local order:** ATF7IP - PLBD1 - GUCY2C - H2AFJ - HIST4H4.

## DNA/RNA

### Description

The GUCY2C gene is approximately 84 kb in length and has 27 exons.

### Transcription

An approximately 3.8 mRNA is transcribed from the gene.

### Pseudogene

None known.

## Protein

### Note

GUCY2C encodes a guanylyl cyclase.

### Description

1073 amino acid protein with guanylyl cyclase catalytic activity (4.6.1.2).

### Expression

Primarily intestinal epithelial cells.

### Localisation

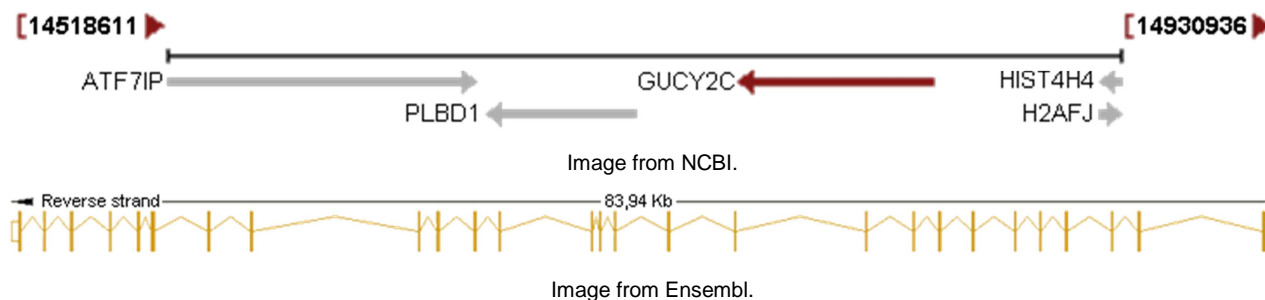
Apical membrane.

### Function

In response to binding endogenous hormones guanylin and uroguanylin, or the exogenous ligand E. coli heat-stable enterotoxin, GUCY2C synthesizes cyclic GMP. Cyclic GMP activates downstream signaling pathways via cGMP-dependent protein kinases, phosphodiesterases and cGMP-gated ion channels.

### Homology

Adenylyl cyclase.





SP: signal peptide; ECD: extracellular ligand binding domain; TM: transmembrane domain; KHD: regulatory kinase-homology domain; CAT: guanylyl cyclase catalytic domain; TAIL: C-terminal tail, interacts with scaffolding proteins.

## Implicated in

### Colorectal cancer

#### Note

The endogenous GUCY2C ligands, guanylin and uroguanylin, are lost early in the neoplastic process. Targeted deletion of Gucy2c in mice results in a phenotype of intestinal cancer susceptibility in the context of predisposing genetic mutations ( $apc^{min}$ ) or exposure to carcinogen (azoxymethane).

## References

Li P, Lin JE, Chervoneva I, Schulz S, Waldman SA, Pitari GM. Homeostatic control of the crypt-villus axis by the bacterial enterotoxin receptor guanylyl cyclase C restricts the

proliferating compartment in intestine. *Am J Pathol.* 2007 Dec;171(6):1847-58

Li P, Schulz S, Bombonati A, Palazzo JP, Hyslop TM, Xu Y, Baran AA, Siracusa LD, Pitari GM, Waldman SA. Guanylyl cyclase C suppresses intestinal tumorigenesis by restricting proliferation and maintaining genomic integrity. *Gastroenterology.* 2007 Aug;133(2):599-607

Lin JE, Li P, Snook AE, Schulz S, Dasgupta A, Hyslop TM, Gibbons AV, Marszlowicz G, Pitari GM, Waldman SA. The hormone receptor GUCY2C suppresses intestinal tumor formation by inhibiting AKT signaling. *Gastroenterology.* 2010 Jan;138(1):241-54

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