

APC (adenomatous polyposis coli – tumor suppressor gene) → long arm Cr 5
gatekeeper gene

- cloned in 1991 through
 - identification of interstitial deletion on Cr5q in FAP pts (loss of heterozygosity – LOH)
 - linkage analysis
- later studies demonstrated that APC plays a pivotal role also in sporadic CRC (>70% – peculiar feature of CRC)
- > 90% APC gene mutation → premature STOP signal → truncated protein
this abnormal protein typical of APC mutations can be used for genetic screening

why is the role of the “non-mutated” APC gene product (i.e. APC protein) in physiological conditions?
what has β -catenin to do with?

- β -catenin: protein necessary to keep a monolayer cylindric type of epithelium together
- it belongs to a protein complex that contains GSK, axin, APC (destruction complex)

APC is the transporter protein, the other components are “skeleton” components, necessary for cell survival → because it is the transporter, it does NOT directly causes cancer

APC is responsible for β -catenin proteolysis

by 2 hit mutation or deletion, APC is downregulated

if β -catenin in the cytoplasm goes over the threshold level, it translocate into the nucleus going over the nuclear membrane (protein overflow)

gut (main) role: absorption

two types of absorption

1. transcellular
2. intracellular (sort of phagocytosis)

cells next to each other open and close → necessity to have “muscles”

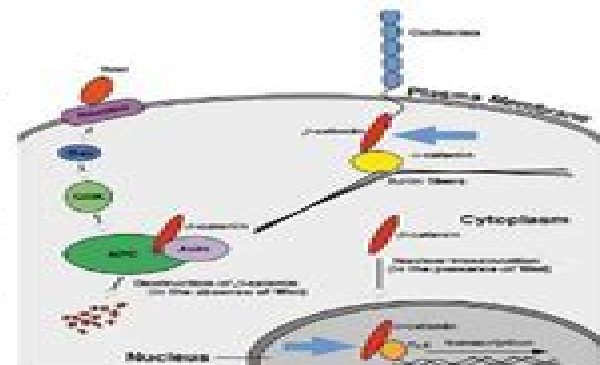
in physiological condition, APC is linked to GSK/axin/ β -catenin leading them to destruction

mutated APC → β -catenin accumulation with consequent translocation into the nucleus, linked to the TCF (= tumor cell factor) → this complex leads to oncogenes activation (transcription) in the nucleus

are APC mutation the only cause of β -catenin accumulation? what happens in patients with normal APC and high levels of β -catenin?

Wnt = wingless/intern gene whose recessive mutation is characterized by wingless *Drosophila Melanogaster*

- ➔ human genes coding for Wnt proteins: in the absence of Wnt ligands, β -catenin is destroyed
- ➔ Wnt → frizzled → dishevelled (Dsh) activation → high affinity to the destruction complex
Wnt is the ligand for frizzled receptors: when activated, this receptor activates an internal cytoplasmatic protein called dishevelled that, once activated, gets to the destruction complex (high affinity)
 - Wnt overexpressed leads to tumor onset due to β -catenin accumulation
 - absence of Wnt ligands, β -catenin is destroyed



Appunti Medicina Interna

S Marginson



Appunti Medicina Interna:

RUGARLI Medicina interna sistematica Claudio Rugarli, 2015-10-19T00:00:00+02:00 Il Rugarli medicina interna sistematica arrivato alla sua settima edizione lo strumento fondamentale che accompagna lo studente di medicina nel corso del triennio clinico Anche in questa edizione ha mantenuto la struttura per apparati con una prima parte introduttiva con nozioni di base per categorie nosologiche definizione classificazione fisiopatologia diagnostica laboratorio e strumentale caratteristiche cliniche del paziente tipo seguita da una rassegna delle singole malattie presentate sempre con lo stesso schema epidemiologia eziopatogenesi fisiopatologia clinica diagnosi cenni di terapia decorso e prognosi Nella revisione e aggiornamento dei contenuti gli Autori hanno prestato particolare attenzione alla parte relativa alla diagnosi Infatti con l'intento di abituare gli studenti al ragionamento clinico oltre a un cappello introduttivo sul metodo stato dato spazio per ciascuna patologia ai dati relativi alla prevalenza e o elementi predisponenti alla sensibilità e alla specificità E' stato inoltre dedicato ampio spazio alle principali comorbidità senza per arrivare alla trattazione del paziente complesso **Dizionario classico di medicina interna ed esterna**, 1831 *Lavori dei congressi di medicina interna*, 1902 *Il Policlinico*, 1903 *Bibliotheca medico-chirurgica, pharmaceutico-chemica et veterinaria*, 1891 **Il policlinico. Sezione pratica periodico di medicina, chirurgia e igiene**, 1930 **Annuario della R. Università degli studi di Padova** Università di Padova, 1893 *La Formazione nelle Medicine Tradizionali e Non Convenzionali in Italia. Attualità, esigenze, criticità e prospettive* AA. VV., Gian Franco Gensini, Paolo Roberti di Sarsina, Mara Tognetti Bordogna, 2015-03-05T00:00:00+01:00 1370 1 31 *Minerva medica*, 1959 *Catalogo generale della libreria Italiana*, 1900 *Una dottoressa ai Caraibi* Julie Danvers, 2024-03-15T00:00:00+01:00 Quando la dottoressa Danielle vince una borsa di studio ai Caraibi viene letteralmente travolta in tutti i sensi da un cardiologo che assomiglia a un semidio alto abbronzato bellissimo Ma Danielle non solo un medico anche una principessa e il suo titolo deve rimanere a tutti i costi un segreto Per questo una relazione con Cade Logan fuori discussione Cade ha chiuso definitivamente le porte al romanticismo ma l'attrazione che prova per Dani innegabile Dopo essersi concesso il lusso di cedere alla tentazione teme che lei possa insinuarsi nel suo cuore che da troppi anni ormai chiuso a chiave Soprattutto quando gli rivela che la loro passione ha avuto una scioccante conseguenza reale *Annuario della R. Università degli studi di Padova per l'anno accademico ...*, 1894 **Current Catalog** National Library of Medicine (U.S.), 1982 First multi year cumulation covers six years 1965 70 **Annuario della R. Università di Pisa** Università di Pisa, 1912 **Il primato degli Italiani nella Botanica** Saccardo, 1894 **LA SALUTE AUTORITARIA Dal mito dell'efficacia al rito dell'efficienza e altri scritti** **AUTHORITARIAN HEALTH From the myth of effectiveness to a ritual efficiency** Guglielmo M. Trovato, 2015-12-07 Tutti dovrebbero avere una grande superbia un grande orgoglio un alto senso di sé di sé come individui e di sé come comunità questo sempre il cammino della civiltà e della storia Vincenzo Consolo Questo un libro per la medicina e per la libertà Tina Anselmi Finalmente qualcosa scritto con sensibilità etica e di ricerca

Graham Wilkie Guglielmo Trovato un medico ma ha anche imparato la medicina e ha imparato a insegnarla Medicina come professionalit si prende cura di esseri umani e come ricerca diffonde nuove conoscenze con metodo scientifico Trovato insegna da quarant anni in una facolt di medicina di una universit italiana Avendo lavorato anche altrove dagli Stati Uniti all Inghilterra alla Francia Dal mito della efficacia al rito della efficienza il titolo del libro che inizi a scrivere nel 1992 raccogliendo gli appunti di sue conversazioni con gli studenti di medicina Ora riproposto con una nuova breve premessa e due articoli in appendice e in nota **Supplemento al policlinico periodico di medicina, chirurgia ed igiene** ,1895

Bibliografia d'Italia ,1878 **Bibliography of the History of Medicine** , **Bibliografia d'Italia compilata sui documenti comunicati dal ministero dell'istruzione pubblica** Silvio Bocca,1878

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In a fast-paced world fueled by information and interconnectivity, the spellbinding force of linguistics has acquired newfound prominence. Its capacity to evoke emotions, stimulate contemplation, and stimulate metamorphosis is really astonishing. Within the pages of "**Appunti Medicina Interna**," an enthralling opus penned by a highly acclaimed wordsmith, readers set about an immersive expedition to unravel the intricate significance of language and its indelible imprint on our lives. Throughout this assessment, we shall delve in to the book is central motifs, appraise its distinctive narrative style, and gauge its overarching influence on the minds of its readers.

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