GENDER CONSIDERATIONS IN THE TREATMENRT OF NEN PATIENTS

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Overall

 There are gender differences across types of NEN in respect to frequency, and clinical presentation

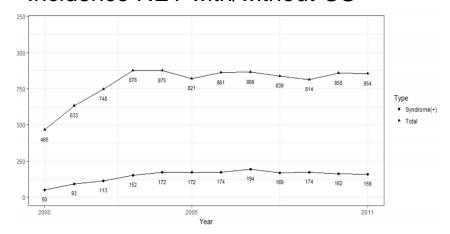
Ex.

- midgut NETs occur more commonly among women
- Small cell lung cancer, an agressive type of NEN, is seen mostly in men (tobacco)
- AC Camargo study (N=456): 55% of overall GEPNET is women

Frequency of carcinoid syndrome

- US database of 9,512 patients (65+yo)
- 18.8% had Carcinoid syndrome
- Risk factors: female gender, mets, well diff, midgut

Incidence NET with/without CS



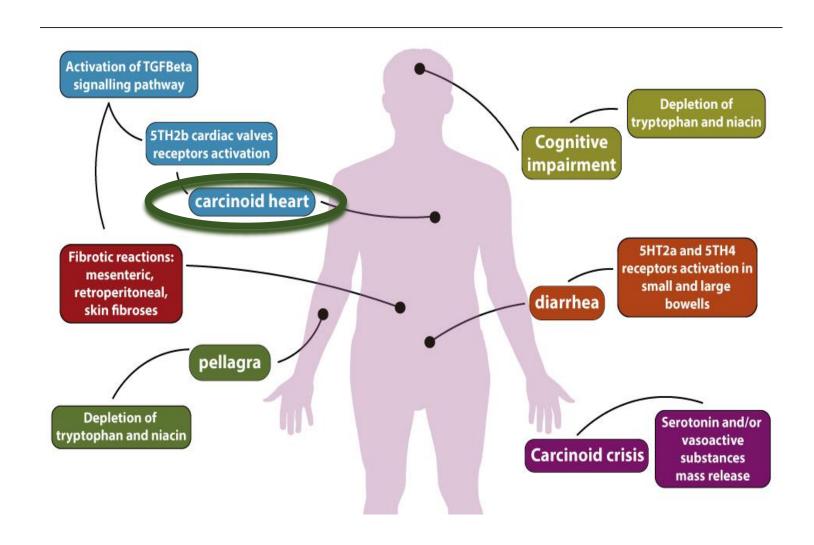
Carcinoid Syndrome





In women of 45 to 55 yo, carcinoid symptoms may be misdiagnosed as menopause

Complications from carcinoid syndrome



Complications from Carcinoid Syndrome: carcinoid heart

 Severe complications that may lead to cardiac insufficiency due to damage of heart valves

 Associated with poor control of carcinoid syndrome (usually longterm)

Not associated with gender

Gender aspects in NEN Treatments

Sexual Hormones in NEN

• In a study conducted by our group, 35% of cases presented positivity for some hormonal receptors (binding proteins) in tumor tissues, either estrogen (ER) or progesterone (PR) receptors.

• Implications for Hormone Replacement Therapy/ Oral contraceptives?

Sexual Hormones in NEN

- Tamoxifen is an oral drug widely used for breast cancer that targets ER.
- Tamoxifen is cheap, safe and easily accessible



HORMONET: Phase II study of hormone therapy with tamoxifen in patients with well differentiated neuroendocrine tumors and hormone receptor positive expression (NCT 03870399)

Particularities in treatments of NEN according to gender

Treatment	Efficacy according to gender is diffent?	Adverse events diferent between genders?
SOMATOSTATIN ANALOGUES	NO	POSSIBLY
EVEROLIMUS	NO	POSSIBLY
SUNITINIB	NO	POSSIBLY
LUTETIUM ¹⁷⁷	NO	NO
CHEMOTHERAPY (Ex. CAPTEM)	NO	NO
HEPATIC EMBOLIZATION	NO	NO

Somatostatin Analogues (SSA) and gender

Women are more prone to gallbladder stones

SSA increase the risk of gallbladder stones

Monitoring is advised

Everolimus

Everolimus-induced stomatitis is more common among women

 Women more commonly have a "sweet tooth" (risk of hyperglicemia)

Monitoring is advised

Sunitinib

 Men tend to have more cardiovascular comorbidities (hypertension, coronary insufficiency)

Sunitinib may induce hypertersion and renal damage

Monitoring is advised

Overall Interpretations about gender as an influencer on NEN treatments

- Overall, the patient <u>clinical scenario</u>, not the gender, <u>dictates the treatment</u>
- The choice of treatment takes into consideration patients' age, comorbid conditions, clinical status, type of NEN, stage, preferences and access.
- Gender is not the main factor
- The role of sex hormones on the biological behaviour of NEN remains to be studied





OBRIGADA!! THANK YOU!! TODA RABA!!

