



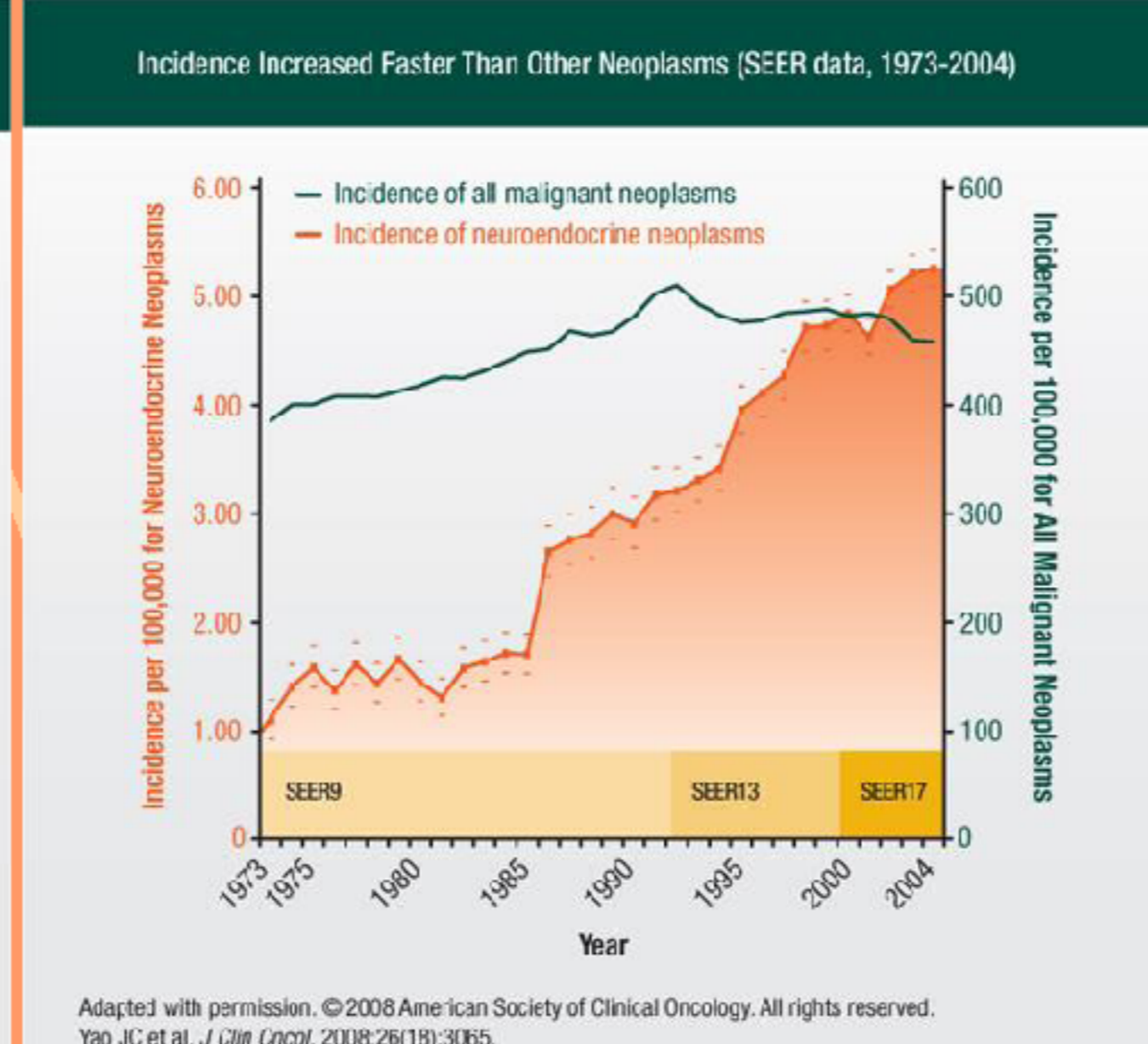
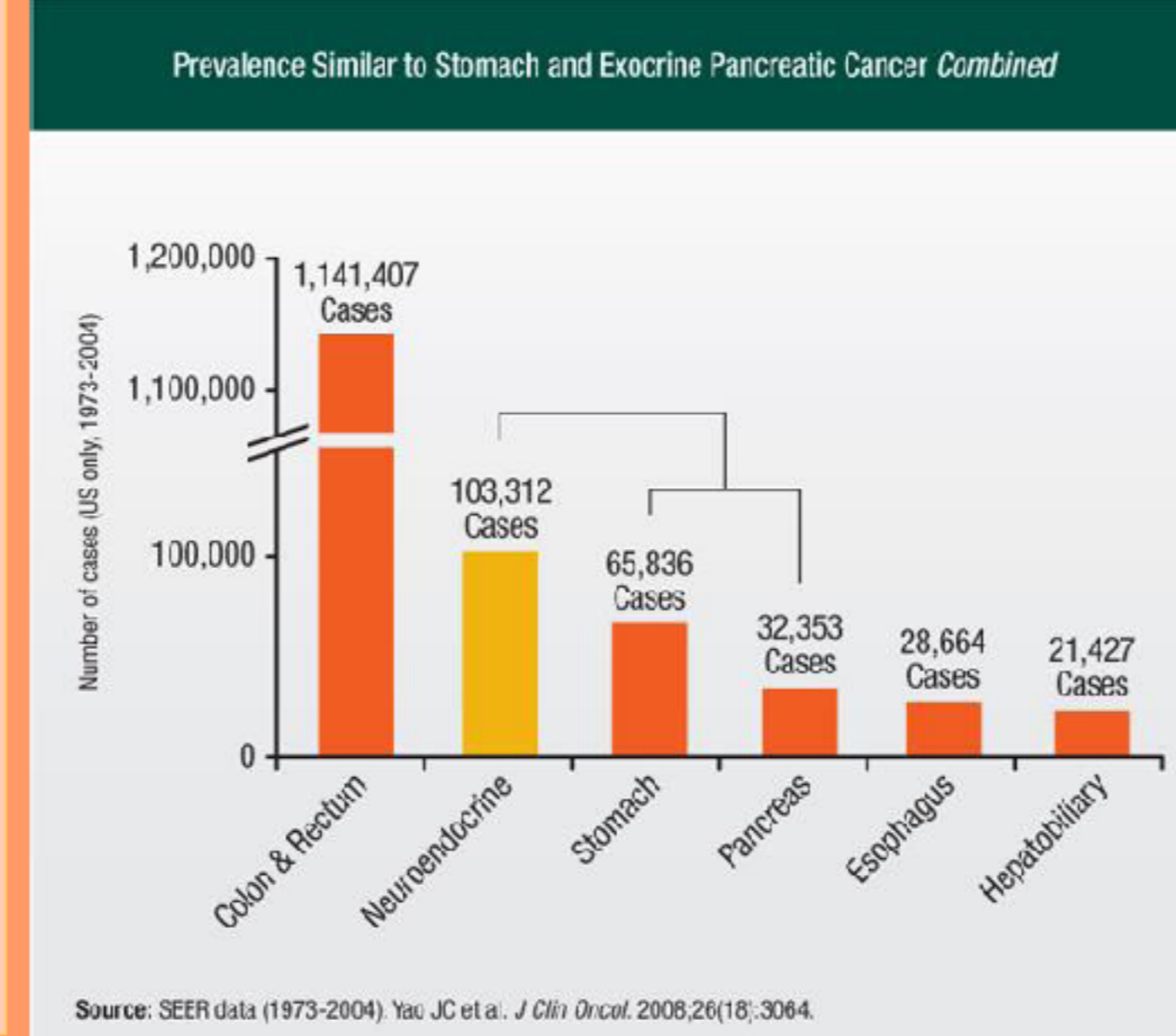
Concomitant long evolving neuroendocrine breast carcinoma and pancreatic tumor- a random association?

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Introduction

➤ Breast neuroendocrine carcinoma is a rare aggressive neuroendocrine tumor (NET).
➤ Its incidence increased in the last period, from less than 0.4% to 3.4% of all breast carcinomas.
➤ According to the World Health Organization (WHO) classification of tumors neuroendocrine tumor in the breast is a category that includes solid neuroendocrine carcinoma, small cell/oat cell carcinoma and large cell neuroendocrine carcinoma.



➤ Neuroendocrine gastroenteropancreatic tumors (GEP-NETs) are a heterogeneous group of tumors with their origin in neuroendocrine cells of the embryological gut.
➤ The primary lesion is located in the gastric mucosa, the small and large intestine, the rectum and pancreas.
➤ The incidence has significantly increased over the last years and is now estimated to be 5.25/100 000/year.
➤ The prevalence has recently been calculated to 35/100 000/year.

Case report

Female patient, aged 73

➤ Presented at the Endocrinology Department with the following **symptoms**:
▪ intense lumbar pain that radiates to the right anterior subcostal region;
▪ loss of appetite; weight loss (15 kg in 2 months);
▪ persistent dry cough

➤ Medical history:

- **1970**: nodular goiter (operated in 1977 - subtotal thyroidectomy)
- **1999**: diabetes mellitus type 2 (October 2011 it requires insulin administration)
- **1999**: hypertension (max systolic BP= 230 mmHg)
- **2008**: angina pectoris
- **2011**:
❖ **Thoracic CT**: suspicion of **sarcoidosis**: mediastinal lymph nodes in the hilum and right lung, nodule in the left hilum of 27/25/38 mm, multiple pulmonary parenchymal

- nodules (<1 cm) in the upper segment of the left lung.
- ❖ **Bronchoscopy with biopsy**: Epithelioid gigantic-cellular granuloma with central ischemic necrosis, + for CD-68
- **2012**: Thoracic control CT – stationary aspect
- **2013**:
❖ **Abdominal CT**
→ an area in the pancreas at the isthmus, measuring 21/20 mm
→ dilated Wirsung duct (5-6 mm).
→ CA-50 = 14 U/mL (N<25 UI/mL)
▪ without further other investigations
- ❖ **Mammography**: Mammary gland fibroadenoma
- January 2014** (due to intense lumbar pain):
• **Lumbar-sacral spinal MRI**: secondary disseminations in the dorsal and lumbar-sacral vertebrae (fig 1)
→ carcinoma / lymphoma / myeloma ?
- January 2014**:
• **Abdominal CT**: intraductal pancreatic lesion with secondary disseminations in the liver, spleen and bone (figure 2, 3, 4, 5)

March 2014:

- **Abdominal ultrasound**: slightly enlarged liver with metastases; in the pancreas a **hypoechoic nodule in the isthmus and body**, measuring **57/29 mm**; in the splenic hilum 2-3 hypoechoic nodules (fig 6).
- **Thoracic X-ray**: right hilar lymphadenopathy 1.5 mm, old posterolateral rib fractures C5-C8, osteolytic lesions of 0.5 cm - left posterior C4, anterior ligament calcification, dorsal osteoarthritis (fig 7).

Laboratory findings		
	Results	Normal range
Chromogranin A (ng/mL)	141.8	0 - 100
Serotonin (ug/L)	990	50 - 200
5-HIAA (mg/24h)	0.2	2-10
CA15-3 (UI/ml)	160	0 - 38.4
CEA (ng/ml)	31.5	0 - 1.5



Figure 1

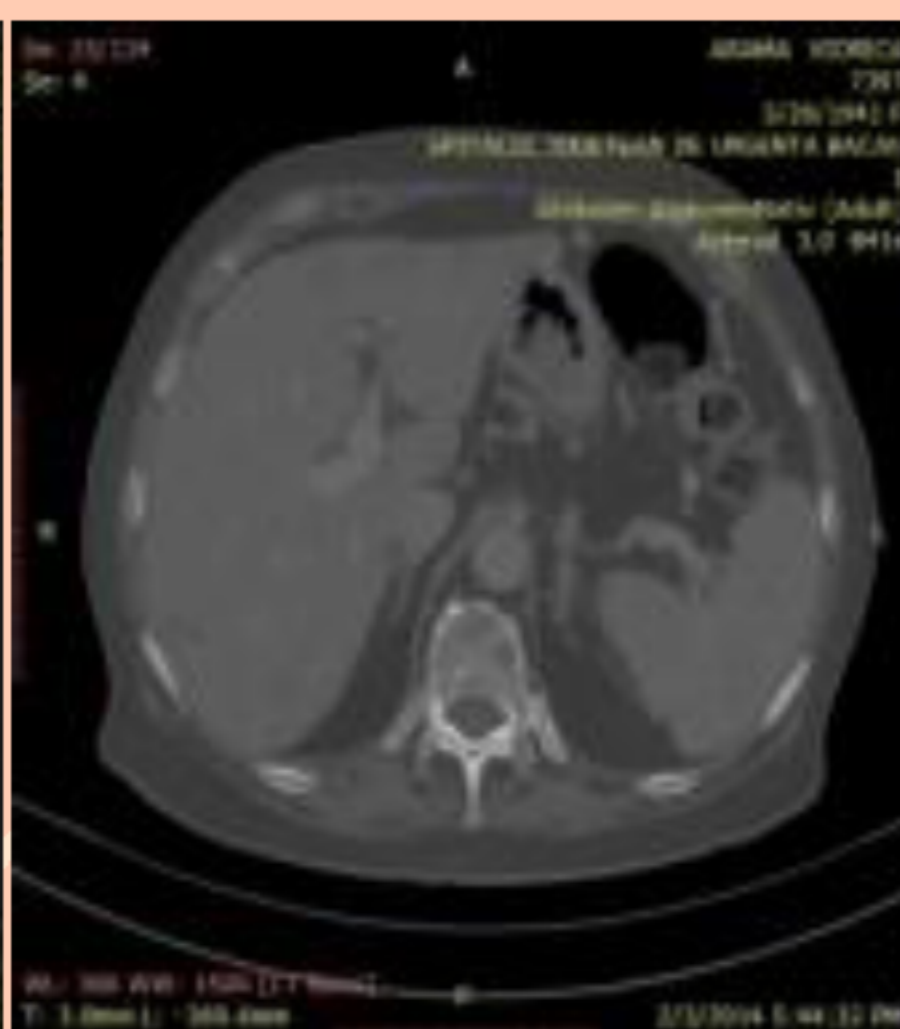


Figure 2



Figure 3

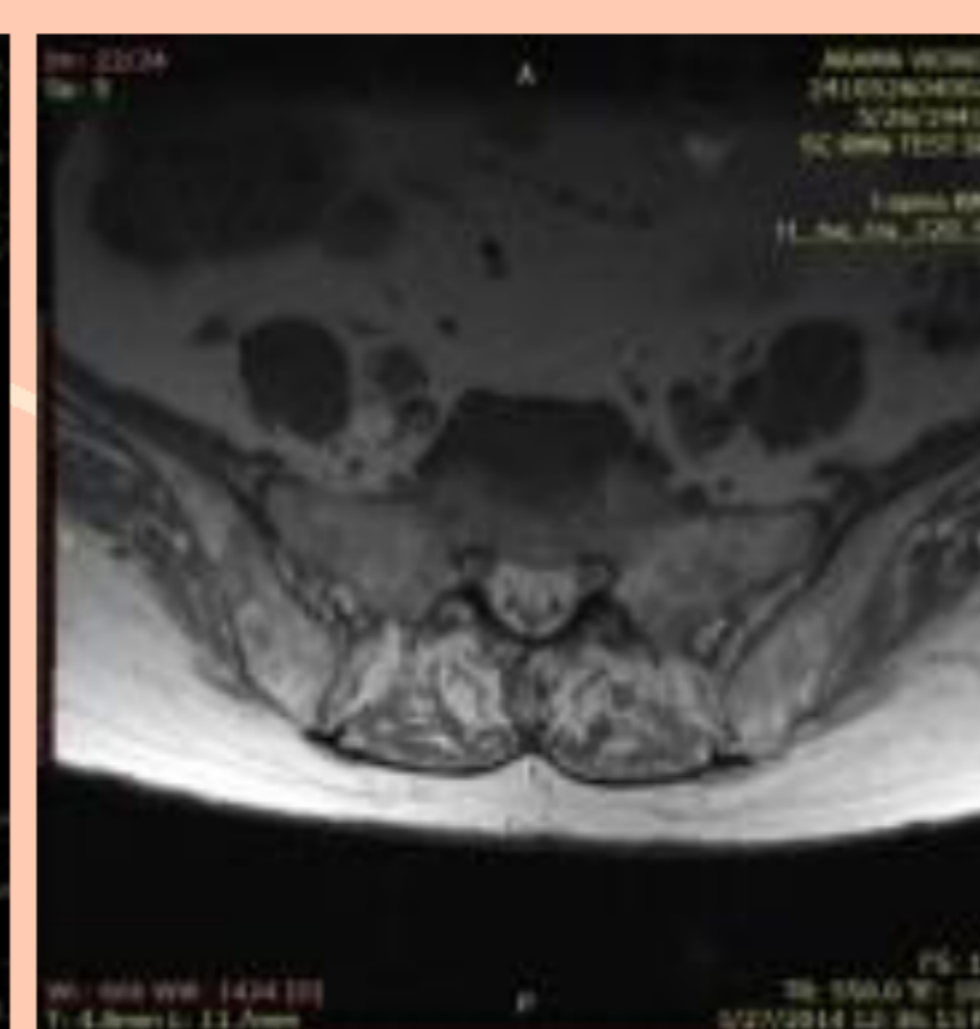


Figure 4

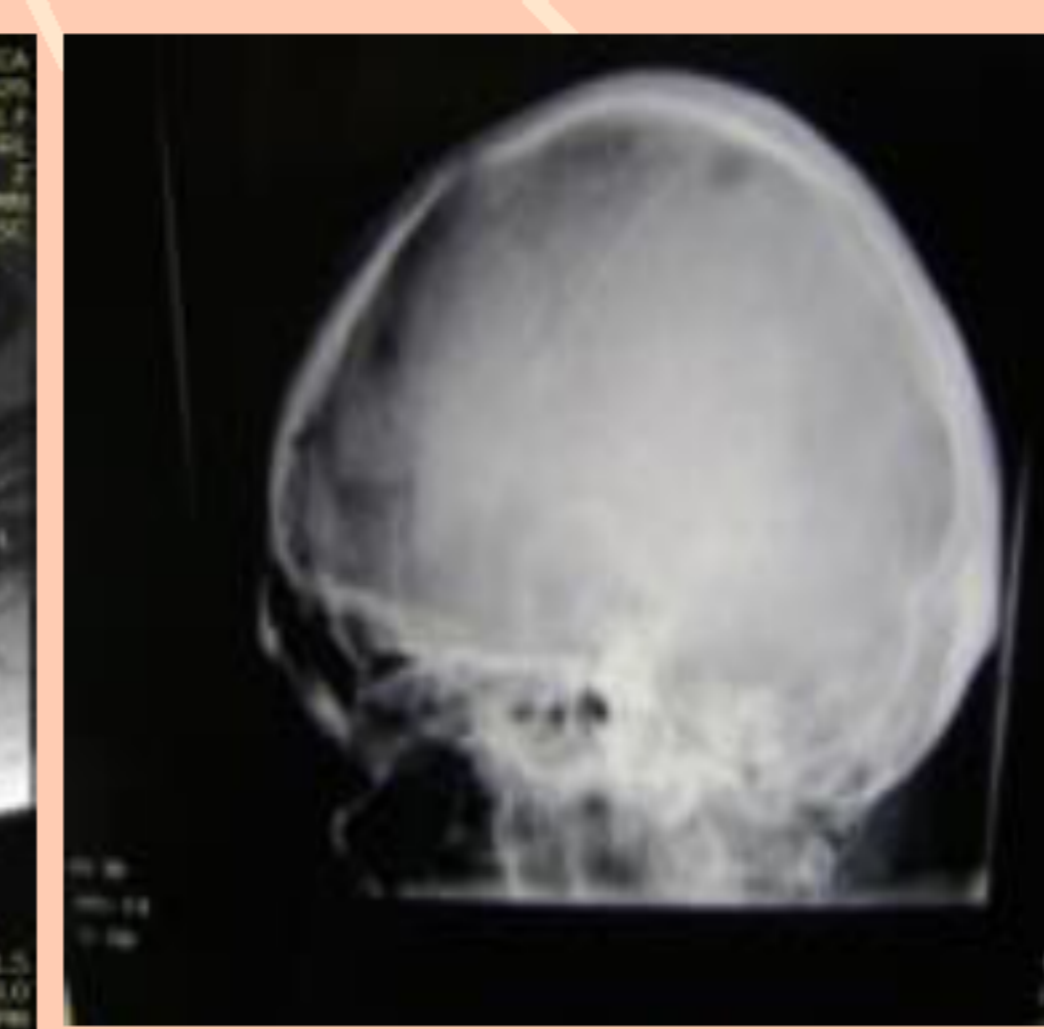


Figure 5

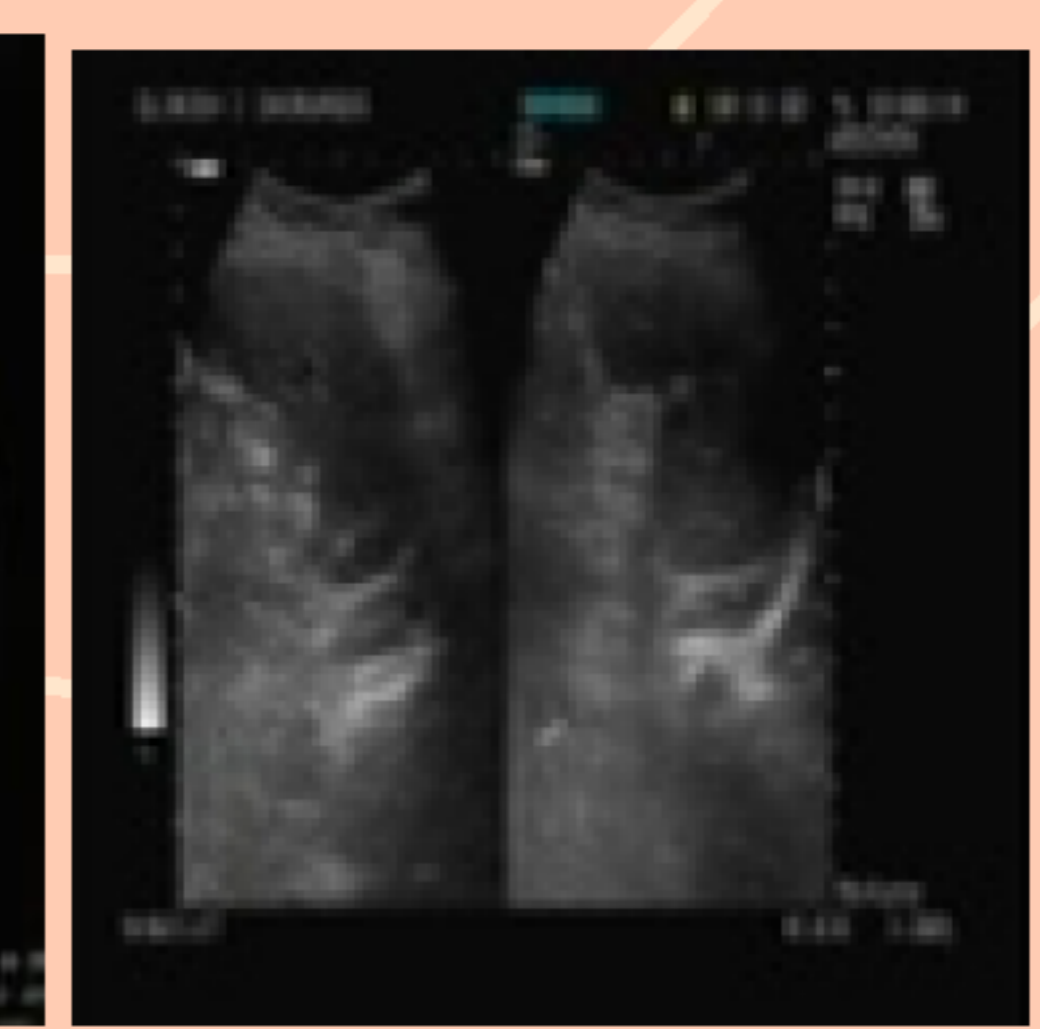


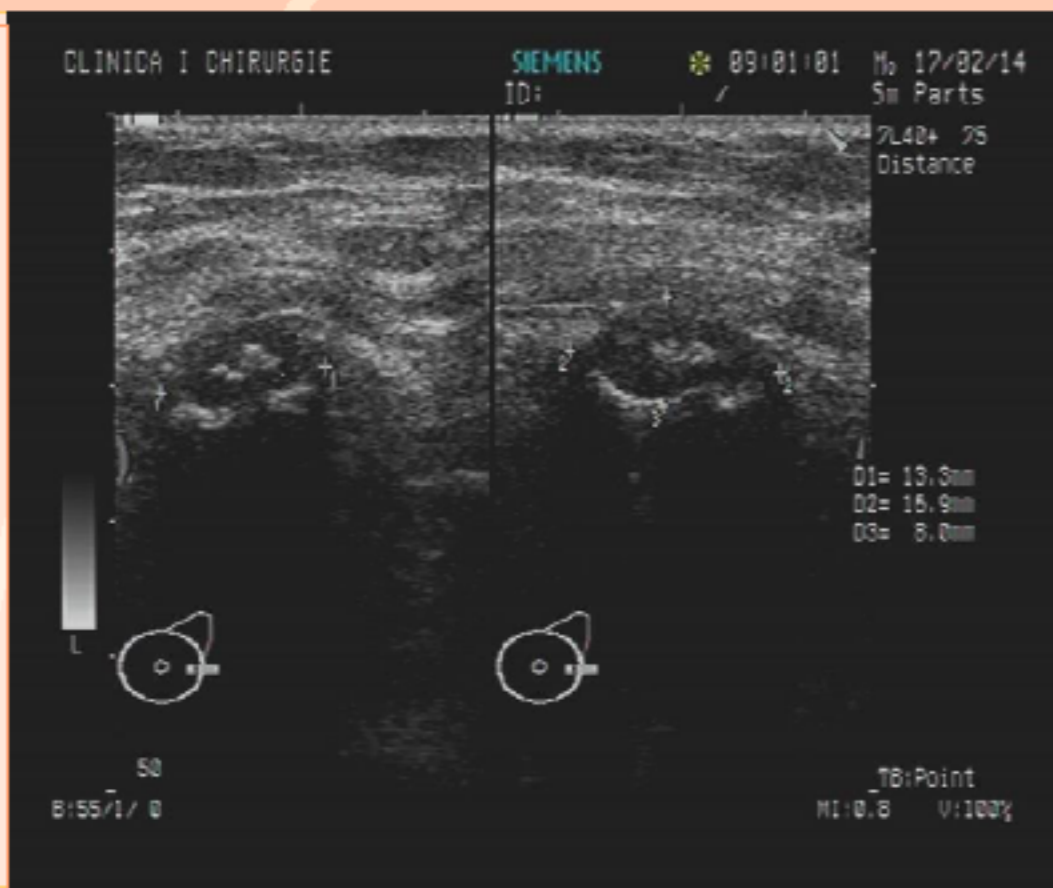
Figure 6



Figure 7

Results

Breast ultrasound:
In the left breast (5 cm from the nipple) multiple solid nodules with a maximum diameter of 10 mm, with coarse calcifications.
+↑ CA15-3, CEA => Breast cancer



Morphopathology investigations
❖ biopsy from one lesion in the liver :**metastasis of poorly differentiated carcinoma**
❖ **immunohistochemistry**: intense + for cytokeratin-7, weak/moderate + for chromogranin, - for synaptophysin, cytokeratin-20, weak/moderate +for estrogen receptor in 50-60% cells. : **CARCINOMA OF THE MAMMARY GLAND WITH NEUROENDOCRINE FEATURES**

Diagnosis
❖ **carcinoma of the mammary gland with neuroendocrine features**
❖ probably **pancreatic neuroendocrine tumour**
❖ **metastasis in the liver, lung and bone**

Oncology evaluation
The oncology evaluation staged the tumour: **cT2N0M1** (liver, lung, bone), ECOG performance status 4.
TREATMENT: somatostatin analogues (sandostatin lar®), zoledronic acid, aromatase inhibitors.

In March 2014, due to pulmonary complications, the patient died.

Discussions

- Long asymptomatic evolution or misdiagnosis?
• bone pain
• difficult balancing diabetes
• chronic respiratory insufficiency
- The period of 3 years of treatment for sarcoidosis
• but with the enhancement of respiratory symptoms and bone pain.
- Are there correlations between breast neuroendocrine carcinoma and pancreatic neuroendocrine tumors?
• There are no similar cases reported in the literature.

Conclusions

- The peculiarity of the case:
- the association of:
❖ breast carcinoma with neuroendocrine features
❖ probably pancreatic neuroendocrine tumour
 - evolving for a long period of time in the past
 - which was initially considered and treated as *sarcoidosis*.

References:

1. Handbook of Gastroenteropancreatic and Thoracic Neuroendocrine Tumours, M. Caplin, J.C. Yao, BioScientifica 2011; Paik WH et al, J Korean Med Sci, 2013; Thomas M. O'Dorisio, Neuroendocrine Cancer Regional Conference, Ohio, 2011; 2. Neuroendocrine gastro-entero-pancreatic tumors:ESMO Clinical Practice Guidelines for diagnosis, treatment and follow-up-K. Öberg1, U. Knigge2, D. Kwakkeboom3 & A. Perren4 on behalf of the ESMO Guidelines Working Group
3. Handbook of gastroenteropancreatic and Thoracic Neuroendocrine Tumours- Martyn Caplin, James Yao 4. Managementul diagnosticului si tratamentul chirurgical in tumorile neuroendocrine gastrointestinale-V. Tomulescu1, O. Stănculea, S. Dima1, V. Herlea2, E. Stoica Mustafa2, T. Dumitrascu1, C. Pechianu2, I. Popescu1-Chirurgia (2011) 106: 151-161 Nr. 2, Martie - Aprilie