

#### **ESMO** Preceptorship Programme

European Society for Medical Oncology

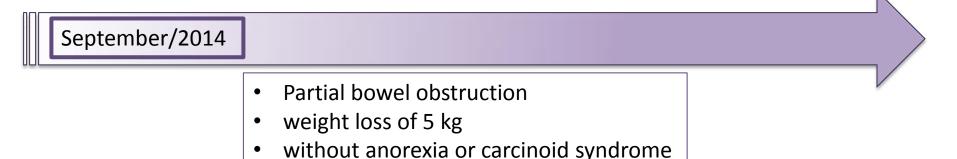
Neuroendocrine neoplasm – Prague, Czech Republic – 28/29 Apr 2017

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## Neuroendocrine tumor – the change of the tumor



- 73 year-old woman, ECOG=1, body weight=55Kg
- Personal History: arterial hypertension, ex-smoker



Colonoscopy:

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- Transverse colon: sessile polyp with 3 mm  $\rightarrow$  Histology: Tubular microadenoma with low grade dysplasia.

- Cecum: neoformation with 5 mm ightarrow Histology: Adenocarcinoma G2, invading submucosa
- TAP-CT + MRI: 2 hepatic nodules in right lobe with 3 and 2 cm in segment VI and V

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October/2014

Right hemicolectomy + right adnexectomy + hepatic segmentectomy V + metastasectomy of VI and VIII segments

- Histology:
  - Neuroendocrine tumor of ileocecal valve, invading subserosa, 0/23 lymph nodes
  - Chromogranin A and synaptophysin positive
  - G2 Ki-67 15%, 10 mitosis/10 HPF
  - Right ovarian and liver (segments V, VI, VII) infiltration
  - pT3N0M1 R0 stage IV WHO classification 2



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November/2014

MRI: two new hepatic lesions with 9 and 10 mm in segment II

- Proposed:
  - Hepato-bilio-pancreatic MDT of the Curry Cabral Hospital (HCC), in Lisbon, for evaluating the possibility of loco-regional therapy



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January/2015

Hepatic metastasectomy

Excised 8 lesions (0,8-1,5 cm) in segment II and 2 diaphragmatic implants.

- Histology:
  - Hepatic and diaphragmatic infiltration due to neuroendocrine tumor G2
  - Ki-67 10%, 13 mitosis/10 HPF



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April/2015

#### DOTA-NOC-PET

Bilobar hepatic lesions with hyperexpression of somatostatin receptors. No evidence of extrahepatic secondary disease with increased expression of receptors for somatostatin.

- Chromogranin A 4,9 mg/dL
- Radiofrequency ablation



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<u>TAP-CT</u>: New hepatic lesions – disease progression

Started octreotid 20mg intramuscular injection, monthly



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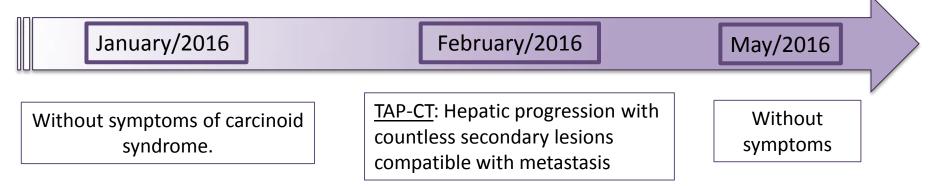
September/2015	November/2015
TAP-CT: Hepatic disease progression	Without symptoms of carcinoid syndrome
<ul> <li>Chromogranin A 9,6 mg/dL</li> <li>Maintain somatostatin analogs</li> </ul>	- Chromogranin A 13,5 mg/dL

- dose increase: octreotid 30 mg

Maintain <u>octreotid 30mg</u>



- 73 year-old woman, ECOG=1
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- Chromogranin A 15,8 mg/dL
- Dose increase: octreotid 40 mg

Change the therapy

- Patient refused chemotherapy / everolimus (QOL)
- Lanreotid 120mg SC, monthly



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Chromogranin A 50 mg/dL

- 73 year-old woman, ECOG=1
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<u>TAP-CT</u>: Hepatic disease progression – Increasing lesion size <u>PET</u>: multiple hepatic lesions and a probable pelvic implant

- Asthenia G1/2
- Chromogranin A 92 mg/dL

- refused intravenous chemotherapy / everolimus (QOL)

Capecitabine (850mg/m<sup>2</sup> 14D)

Temozolamide (150mg/m<sup>2</sup> D10-D14) 28/28D



- 73 year-old woman, ECOG=1
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- Chromogranin A 175 mg/dL
- Toxicity:
  - asthenia G2
  - hand-foot-syndrome G3
- Change to <u>everolimus 10mg/day</u>

- Everolimus 10mg/day
- Toxicity:
  - Thrombocytopenia
- Awaiting imaging evaluation



#### **Discussion**

- The treatment of neuroendocrine tumor is a challenge.
- Patients tend to present after a long history of vague nonspecific symptoms with disseminated metastasis.
- The liver is the predominant site of metastatic disease in gastrointestinal neuroendocrine tumors. Surgery is often advocated in patients with limited liver metastasis.



#### Discussion

- In this case:
  - Re-biopsy liver metastasis to re-assess the profilerative activity?
  - How to treat patients with Ki-67 10-15%? With somatostatin analogs? (Promid/Clarinet trial Ki-67 <10%)</li>
- If the disease continues to progress what to do?
  - Keep chemotherapy and switch line?
  - Peptide receptor-targeted radiotherapy (Yttrium-90 and Lutetium-177) in metastatic carcinoid tumors refractory to octreotid was associated with improvement in symptoms and radiografic regression
- The patient must be respected
  - despite the therapeutic indications and guidelines, we must adjust the therapy according to his will









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