# Title Head and Neck Cancer in the Elderly

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# INTRODUCTION

- Head and neck cancer is a disease of older adults
- an estimated 61% of patients ages 65 and older by the year 2030
- More than 90% are squamous cell carcinomas that arise from the mucosal surfaces of the oral cavity, oropharynx, and larynx
- early disease (stages I–II), there is an 80% cure rate
- A majority of patients (>60%) with (HNSCC) present with locoregionally advanced disease (stages III and IVA/B) and metastatic tumors (stage IVC)
- more than half of patients with locally advanced HNSSC develop recurrences and 30% are at risk of distant metastases

# Head and Neck Cancer (HNC) A disease of the old?

- Worldwide HNC is still increasing (688.000 in 2012)
- Majority still tobacco and alcohol related
- Increase of viral-associated OPC, less so in elderly
- 47% of SCCHN patients >65 years of age
- The incidence of HNC among older patients is expected to increase 34% over the next 10 years, and 64% over the next 20 years
- Most studies use the age of 70 (or even 75) as a cut-off for being old

# (HNC) The problem of aging

- Aging is associated with:
  - loss of function reserve of multiple organ systems
  - increased prevalence of chronic diseases
  - enhanced susceptibility to stress
- However, age-related modifications occur at different rates in different individuals
- Chronological age ≠ biological age



#### The effect of aging on outcome

- Few studies available specifically focused at older age
- Variable outcomes in studies:
  - similar outcome when similarly treated
  - worse outcome, due to higher comorbidity status
  - independent prognostic variable?
- Multiple studies concluded that geriatric SCCHN patients receive non-standard and less aggressive therapies

### **Decision Making**

- Disease factors (e.g. site, stage, biology [HPV,EGFR], specific risk factors for locoregional or distant relapse)
- Patient factors (age, sex, performance status, nutritional status, comorbid chronic disease, oral health, lifestyle habits, socio-economic status)
- Treatment factors (surgery, radiotherapy, chemotherapy, immunotherapy, targeted therapy)
- What do patients want?

#### Treatment Options

Early-stage SCCHN (stage I-II)

- ERT vs BT vs S (depending on patient/disease factors)
- Locoregionally advanced SCCHN (stages III-IV)
  - Surgery  $\rightarrow$  adjuvant RT or concurrent CRT (CCRT)
  - Definitive CCRT (surgery remains an option)
  - Definitive cetuximab/RT (surgery remains an option)
  - Induction chemotherapy (ICT)  $\rightarrow$  local treatment
- Recurrent/metastatic SCCHN
  - Recurrent resectable: postop. RT or CCRT3
  - R/M-SCCHN: PFE (fit patients); single drug (PS2), BSC2

#### The Importance of Comorbidity

Comorbidity scores may contribue in decision proces:

- Charlson Comobidity Index (CCI)
- Adult Comorbidity Evaluation 27 (ACE27)
- Comorbidity of importance both in the primary disease setting and in the recurrent disease setting
   Inclusion of comorbidity led to a further refinement in the

prognostic model in OPSCC patients

## Chemotherapy in the Elderly Dangers

- Decrease in renal blood flow  $\rightarrow$  GFR  $\downarrow$   $\rightarrow$  cave: cisplatin, carboplatin, MTX
- Serum creatinine does not reflect renal function in the elderly
- Consider replacing cisplatin with carboplatin
- Gastric enzymes & splanchnic blood flow  $\downarrow \rightarrow$  absorption oral drugs  $\downarrow$
- Liver mass and cytochrome 450 \$\sqrthcolor\$ (cave combination with drugs that have exclusive liver metabolism (e.g. opioids)
- Elderly patients with  $GFR \downarrow$  and preexisting neuropathy (Cis, Vinca, Tax)
- Fluorouracil and fluoropyrimidine therapy → diarrhea, mucositis. Earlyrehydration in CT-induced diarrhea recommended (NCCN)
- Elderly have a physiologic decline in intracellular DPD levels (cave 5-FU)
- Risk of myelosuppression ↑ (over 65 yrs of age): consider G-CSF

#### **Better Selection of Patients**

- Geriatric screening tools to estimate whether patients are able to tolerate CCRT or BRT
  - G8 screening tool
  - Flemish version of the Triage Risk Screening Tool (fTRST)
  - Goningen Frailty Indicator
  - Vulnerable Elders Survey-13 (VES-13)
  - Abbreviated Comprehensive Geriatric Assessment

# G8 screening tool

- The total score of the G8 screening tool can range from1-17
- A high-risk
   profile= score ≤14

	Items	Possible answers	Score
	Food intake in the last 3 months	<ul> <li>0: severe reduction in food intake</li> <li>1: moderate reduction in food intake</li> <li>2: normal food intake</li> </ul>	
	Weight loss during the last 3 months	<ul> <li>0: weight loss &gt;3kg</li> <li>1: does not know</li> <li>2: weight loss between 1 and 3 kg</li> <li>3: no weight loss</li> </ul>	
	Mobility	<ul> <li>0: bed/chair bound</li> <li>1: able to get out of bed/chair but does not go out</li> <li>2: goes out</li> </ul>	
	Neuropsychological problems	<ul> <li>0: severe dementia or depression</li> <li>1: mild dementia or depression</li> <li>2: no psychological problems</li> </ul>	
	Body Mass Index (BMI)	<ul> <li>0: BMI &lt;19</li> <li>1: BMI 19 to &lt;21</li> <li>2: BMI 21 to &lt;23</li> <li>3: BMI 23 or greater</li> </ul>	
	Takes more than 3 medications per day	0: yes 1: no	
	Self-rated health status (compared to other people of the same age)	<ul> <li>0: not as good</li> <li>0.5: does not know</li> <li>1: as good</li> <li>2: better</li> </ul>	
	Age	0: >85 1: 80-85 2: <80	
	Total score (0-17) [Cut-off $\leq$ 14 indicating impairment]		

## Flemish version of the Triage Risk Screening Tool (fTRST)

- The total score of the fTRST can range from 0-6
- A high risk profile
   in the oncologic
   population is a
   score ≥1

Items		Score	
		Yes	No
1.	Presence of cognitive impairment (disorientation, diagnosis of dementia, or delirium)	2	0
2.	Lives alone or no caregiver available, willing, or able	1	0
3.	Difficulty with walking or transfers or fall(s) in the past 6 months	1	0
4.	Hospitalized in the last 3 months	1	0
5.	Polypharmacy: $\geq$ 5 medications	1	0
То	tal score	0-6	
	normal if $\geq 2$ within the geriatric population and $\geq 1$ within th pulation	e oncologic	

Results: both screening tools have a high sensitivity and moderate NPV to detect patients with a hig-risk profile. G8 had the strongest prognostic value for OS HR for G8 normal vs abnormal, 0.38 (95%CI: 0.27-0.52

## ORAL CAVITY

Early-stage

- Stage I : any tumor <2 cm with < 5 mm depth of invasion
- Stage II : tumor 2-4 cm with or depth of invasion >5 mm and < 10
- addressed quickly and effectively with surgery without functional loss
- Definitive radiotherapy is often not the primary treatment modality in because of the length of treatment required and the long-term adverse effects, including xerostomia, altered sense of taste, dental issues, and risk of osteoradionecrosis & increased Mortality
- elderly patients who are medically unfit for surgery, radiotherapy remains an effective curative option(external beam radiotherapy or brachytherapy)
- Trial demonstrated an increased survival benefit to elective neck dissection in oral cavity cancer (except for invasion < 3 mm )</li>
- sentinel lymph node biopsy can provide a negative predictive value of 95%

#### OROPHARYNX

- half of all oropharyngeal carcinoma attributed to HPV
- Early-stage carcinoma : lesion < 4 cm in its greatest dimension without invasion into surrounding structures or nodal involvement
- Soft palate tumors are treated with radiotherapy, because surgery often leads to higher morbidity, especially velopharyngeal insufficiency
- tonsil lesions usually can be surgically resected with minimal risk to speech and swallowing
- Radiotherapy generally is favored for base-of-tongue cancers
- Ipsilateral neck treatment is appropriate for lesions localized to the tonsil, whereas bilateral lymphatic treatment is necessary for most other oropharyngeal sites
- data in advanced-stage HPV-related HNSCC show that de-escalated therapy(better prognosis and longer survival)

#### HYPOPHARYNX

- Early-stage cancer is confined to the hypopharynx <4 cm without vocal cord fixation or nodal involvement</p>
- Radiotherapy is the general default choice for early-stage hypopharyngeal cancer because of its adequate survival outcomes and functional preservation
- no differences in the incidence of mucositis, weight loss, osteoradionecrosis, or laryngeal dyspnea between the general population and the elderly
- all patients with clinically node-negative disease should have bilateral elective treatment to the neck



- Early-stage laryngeal cancers are tumors that do not invade the thyroid cartilage or show evidence of nodal involvement
- radiation and surgery have been shown to have similar survival
- IMRT minimize radiotherapy to the carotid arteries with respectable local control
- Minimally invasive approach is the preferred method to preserve function if tumor-free margins can be achieved
- Iow risk that glottic cancers will spread to lymph nodes, observation after treatment of the primary tumor is recommended.32 Supraglottic and subglottic tumors require bilateral elective therapy of the neck

#### NASOPHARYNX

- The standard of care for early nasopharyngeal cancer is radiotherapy, and it has excellent outcomes
- Surgery plays little role in the primary treatment, because this area is not readily accessible for an oncologic resection
- Traditionally, the neck is treated with elective whole-neck radiotherapy
- elective radiotherapy can spare the lower neck in clinically nodenegative disease without sacrifice of oncologic outcomes