# Impact of statin use in the treatment of patients with cancer of the esophagus

MCJ Anderegg¹, SM Lagarde¹, AE Slaman¹, SS Gisbertz¹, ORC Busch¹, SL Meijer², MCCM Hulshof³,

JJGHM Bergman⁴, HWM van Laarhoven⁵, MI van Berge Henegouwen¹
¹Department of Surgery, ²Pathology, ³Radiation Oncology, ⁴Gastroenterology and ⁵Medical Oncology; Academic

Medical Center, Amsterdam, the Netherlands

## **Background**

- o For the vast majority of patients with resectable esophageal malignancies the preferred treatment consists of neoadjuvant chemo(radio)therapy, nC(R)T, followed by esophagectomy.
- o Statins are increasingly used to treat hypercholesterolemia and are currently the most commonly prescribed drug in the world
- Preclinical studies have shown statins to have anti neoplastic effects like arrest of cell-cycle progression, induction of apoptosis, and inhibition of angiogenesis and tumor growth
- o In rectal cancer it has been stated that statin therapy is associated with an improved response to neoadjuvant chemoradiation

### Aim

To determine the effect of statins on pathologic complete response (pCR) and survival after nC(R)T and surgery for locally advanced esophageal cancer

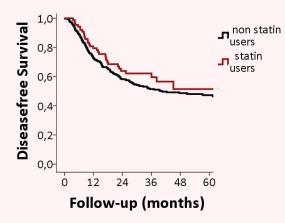
### **Patients & methods**

- o Study period: March 1994 September 2013
- o Inclusion of all consecutive patients treated with nC(R)T, followed by esophagectomy
- o Division: Non-statin users vs. Lipophilic- vs. Hydrophilic statin users
- o Surgical exploration: transthoracic / transhiatal, conventional / minimally invasive
- o Standardized pathologic examination & follow-up
- Multivariate Cox regression analysis to identify independent prognostic factors

Multivariate Cox Model	HR (95% CI)	P-value
Disease free survival		
Statin use	0.787 (0.541-1.144)	0.209
Lipophilic statin use	0.642 (0.385-1.072)	0.090
Overall survival		
Lipophilic statin use	0.664 (0.403-1.094)	0.108

#### Results

- o Number of included patients: 463
- o Neoadjuvant chemo- vs. chemoradiotherapy: 88 (19.0%) vs. 375 (81.0%)
- o Non-statin users vs. statin users: 373 (80.6%) vs. 90 (19.4%)
- o Lipophilic vs. hydrophilic statin users: 49 (54.4%) vs. 41 (45.6%)
- Baseline characteristics statin users: higher age, higher BMI, higher ASA score, similar clinical tumor stage
- o Larger part of statin users received neoadjuvant chemoradiotherapy compared to nonusers (88.9% vs. 79.1%, P 0.033)
- o pCR in non-statin users vs. statin users: 17.4% vs. 22.2% (P 0.291)
- o Median disease free survival in non-statin users vs. statin users: 40.4 vs. 45.0 months (P 0.245)
- o Median overall survival in non-statin users vs. statin users: 41.9 vs. 50.5 months (P 0.284)
- o Subgroup analysis (adenocarcinoma only): median disease free survival in lipophilic vs. non/hydrophilic statin users: 50.9 vs. 39.5 months (P.0.066)



#### Conclusion

Statin users did not have different outcomes compared with non-users. This study does not support modification or discontinuation of statin therapy for patients with esophageal cancer

