

BC Cancer Protocol Summary for Palliative Therapy for Unresectable, Platinum-refractory, Recurrent or Metastatic Squamous Cell Cancer of the Head and Neck Using Nivolumab

Protocol Code

UHNAVNIIV

Tumour Group

Head and Neck

Contact Physician

Dr. Cheryl Ho

ELIGIBILITY:

- Histologically confirmed recurrent or metastatic SCCHN (oral cavity, oropharynx, pharynx, larynx, primary unknown), stage III/IV and not amenable to local therapy with curative intent (surgery or radiation therapy with or without chemotherapy)
- Patients have received at least 1 prior line of platinum chemotherapy in the neoadjuvant, adjuvant, concurrent, or metastatic setting
- ECOG 0-2
- Adequate hepatic and renal function
- Patients may be PDL1 positive or negative
- Patients may be p16 positive or negative
- Access to a treatment centre with expertise to manage immune-mediated adverse reactions of nivolumab
- A BC Cancer “Compassionate Access Program” (CAP) approval
- CAP approval is not required to switch between UHNAVNIIV and UHNAVNIIV4

EXCLUSIONS:

- Recurrent or metastatic cancers of the salivary gland, nasopharyngeal carcinoma, or non-squamous histologies
- Active central nervous system metastases (should be asymptomatic and/or stable)
- Active autoimmune disease, active hepatitis B, C or HIV (HCV antibody or negative HCV RNA permitted)
- Use with caution in patients with long term immunosuppressive therapy or systemic corticosteroids (requiring more than 10 mg prednisone/day or equivalent)

TESTS:

- Baseline: CBC/differential, creatinine, alkaline phosphatase, ALT, total bilirubin, LDH, sodium, potassium, random glucose, TSH
- Before each treatment: CBC/differential, creatinine, alkaline phosphatase, ALT, total bilirubin, LDH, sodium, potassium, glucose, TSH
- If clinically indicated: chest x-ray, free T3 and free T4, morning serum cortisol, lipase, serum ACTH levels, FSH, LH, testosterone, estradiol
- Weekly telephone assessment for signs and symptoms of side effects while on treatment (optional but recommended)

PREMEDICATIONS:

- Antiemetic protocol for low emetogenic chemotherapy protocols (see [SCNAUSEA](#)). Antiemetics are not usually required.

TREATMENT:

Drug	Dose	BC Cancer Administration Guideline
nivolumab	3 mg/kg (maximum 240 mg)	IV in 50 to 100 mL NS over 30 minutes using a 0.2 micron in-line filter

Repeat **every 2 weeks** until disease progression or unacceptable toxicity.

DOSE MODIFICATIONS:

No specific dose modifications. Toxicity managed by treatment delay and other measures (see [SCIMMUNE](#) protocol for management of immune-mediated adverse reactions to checkpoint inhibitors immunotherapy).

PRECAUTIONS:

- **Serious immune-mediated reactions:** these can be severe to fatal and usually occur during the treatment course. They may include enterocolitis, intestinal perforation or hemorrhage, hepatitis, dermatitis, neuropathy, endocrinopathy, as well as toxicities in other organ systems. Early diagnosis and appropriate management are essential to minimize life-threatening complications (**see [SCIMMUNE](#) protocol for management of immune-mediated adverse reactions to checkpoint inhibitors immunotherapy**).
- **Infusion-related reactions:** isolated cases of severe reaction have been reported. In case of a severe reaction (Grade 3 or 4), nivolumab infusion should be permanently discontinued and appropriate medical therapy administered. Patients with mild or moderate infusion reaction may receive nivolumab with close monitoring. Premedications with acetaminophen and antihistamine may be considered if there is a history of reaction.

Call Dr. Cheryl Ho or tumour group delegate at (604) 877-6000 or 1-800-663-3333 with any problems or questions regarding this treatment program.

References:

1. Ferris RL, Blumenschein G, Fayette J, et al. Nivolumab for recurrent squamous-cell carcinoma of the head and neck. *N Engl J Med* 2016; 375(19):1856-1867.
2. Gillison ML, Blumenschein G, Fayette J, et.al. Phase III, open-label, randomized study of Nivolumab (nivo) vs investigator's choice (IC) for recurrent or metastatic (R/M) head and neck squamous cell carcinoma (HNSCC): CheckMate-141. ASCO Meeting Abstracts 34:6009, May 2016.
3. Ferris L, Blumenschein G, Fayette J, et.al. Further evaluations of nivolumab (nivo) versus investigators choice (IC) chemotherapy for recurrent or metastatic (R/M) squamous cell carcinoma of the head and neck (HNSCC): CheckMate 141. *J Clin Oncol Asco Meeting Abstracts* 34:6009, May 2016.
4. Lyford-Pike S, Peng S, Young GD, et.al. Evidence for a role of the PD-1:PD-L1 pathway in immune resistance of HPV-associated head and neck squamous cell carcinoma. *Cancer Res* 2013;73 (6):1733-1741.
5. Bristol-Myers Squibb Canada. OPDIVO® product monograph. Montreal, Quebec; 26 August 2016.

6. Bristol-Myers Squibb Canada. OPDIVO® Immune-mediated adverse reaction management guide. Montreal, Quebec. April 2016.
7. Waterhouse D, Horn L, Reynolds C, et al. Safety profile of nivolumab administered as 30-min infusion: analysis of data from CheckMate 153. *Cancer Chemother Pharmacol* 2018;81:679-86.