

# **Systemic Therapy Update**

Volume 23 Issue 6 June 2020

## For Health Professionals Who Care for Cancer Patients

#### **Inside This Issue:**

#### **Editor's Choice**

**New Pembrolizumab Programs:** Locally Advanced or Metastatic Urothelial Carcinoma | Platinum-Pemetrexed with Pembrolizumab for Advanced Non-Squamous NSCLC | Paclitaxel-Carboplatin with Pembrolizumab for Advanced Squamous NSCLC | Platinum-Gemcitabine with Pembrolizumab for Advanced Squamous NSCLC | Relapsed or Refractory Hodgkin Lymphoma | Adjuvant Treatment of Resected Stage III-IV NED Melanoma

#### **Drug Update**

Immunotherapy Dosing Schedules | Biosimilar Rituximab Coming Soon | Discontinuation of Brand-Name Warfarin | Patient Assistance Programs

#### **Drug Shortages**

New: Bromocriptine, Nystatin | Resolved: Cabergoline

#### **ST Update Editorial Board**

Membership Update

#### Cancer Drug Manual<sup>®</sup>

New: Cemiplimab | Revised: Cabazitaxel, Pembrolizumab Benefit Drug List

New: UGUAVPEM, UGUAVPEM6, ULUAVATZ4, LUAVPMBM, LUAVPMBM6, ULUAVPCPMB, ULUAVPGPMB, LUAVPPMBM, ULUAVPPPMB, ULYPEM, ULYPEM6, USMAJPEM, USMAVPEM6 Revised: Asparaginase (LKNOS, LYNOS) Deleted: GIGAVCCT

#### **NEW Protocols, PPPOs and Patient Handouts**

GU: UGUAVPEM, UGUAVPEM6 LU: ULUAVATZ4, LUAVPMBM, LUAVPMBM6, ULUAVPCPMB, ULUAVPGPMB, LUAVPPMBM, ULUAVPPPMB LY: ULYPEM, ULYPEM6 SM: USMAJPEM, USMAVPEM6

#### **REVISED Protocols, PPPOs and Patient Handouts**

BR: BRAVPTRAD, BRAVTCAP GI: GIGAVFFOXT GO: GOCXCRT,
GOOVGEM, GOOVTOP GU: GUBEP HN: HNAVPE, HNNAVPE,
HNOTVAN LK: ULKAMLAS, ULKMDSA LU: ULUAVATZ
MY: UMYCARLD, UMYLDF, UMYLDREL, UMYLENMTN,
UMYPOMDEX SC: SCDRUGRX, SCNAUSEA SM: USMAJDT,
USMAJNIV, USMAJNIV4, USMAVFIPI, USMAVIPI, USMAVIPNI,
USMAVNIV, USMANIV4, USMAVPEM, SMAVTMZ, USMMCCAVE,
SMMCCPE

Antiemetics Updated: BRAJAC, BRAJACT, BRAJACTG, BRAJACTT, BRAJACTTG, BRAJACTW, BRAJDAC, BRAJFEC, BRAJFECD, BRAJFECDT, BRAVAC, BRLAACD, BRLAACDT, BRLACTWAC, BRLATACG, BRLATWAC, GIENDO2, GIFIRINOX, GIGAJCPRT, GIGAVCC, GIGAVCFT, GIGECC, GIPAJFIROX, GOBEP, GOCISP, GOEP, GOOVCIS, GUAJPG, GUAVPG, GUBEP, GUEDPM, GUEP, GUMVAC, GUNAJPG, HNAVFUP, HNAVPC, HNAVPD, HNAVPE, HNLAALTPRT, UHNLADCF, HNLAPRT, HNNAVFUP, HNNAVPE, HNLAALTPRT, UHNLAPG, HNSAVFAC, HNSAVFUP, HNSAVNP, HNSAVPAC, LUAJNP, LUAVDC, LUAVPG, LUAVPP, LUMMPG, LUMMPP, LUOTCAV, LUOTPAC, LUSCCAV, LUSCPI, SAAI, SAAI3, SAAJADIC, SAAJAP, SAAVADIC, SAAVAP, SAAVI, SAAVI3, SAAVIME3, SADTIC, SAIME, SAVAC, SAVDC, SAVDCM, SMDTIC

#### **Resources and Contact Information**

### Editor's Choice

#### **New Programs**

Effective 01 June 2020, the BC Cancer Provincial Systemic Therapy Program has approved the following new pembrolizumab-containing treatment programs. Recommended pembrolizumab dosing schedules are dependent on clinical and pharmacokinetic data, as well as on the administration schedule of concomitant agents; therefore, protocols may include 3-weekly, 6-weekly, or both 3- and 6-weekly dosing schedules. In general, 6-weekly dosing allows for added flexibility in scheduling patient appointments. Full details of these programs can be found on the BC Cancer website in the <u>Chemotherapy Protocols</u> section.

For patients approved for 3-weekly pembrolizumab treatment by the BC Cancer Compassionate Access Program (CAP), new CAP approval is <u>not</u> required when switching to a 6-weekly dosing schedule. Note that because of the significant workload associated with the large number of eligible patients, <u>new</u> patients should be prioritized by BC Cancer centres and Community Oncology Network (CON) sites. Patients currently being treated with pembrolizumab via the manufacturer patient assistance program can be transitioned to a BC Cancer treatment protocol from 01 June through 15 October 2020.

#### Genitourinary

**Pembrolizumab for Locally Advanced or Metastatic Urothelial Carcinoma (UGUAVPEM, UGUAVPEM6)** — The Genitourinary Tumour Group is introducing pembrolizumab as the first immunotherapy agent to be approved in patients with locally advanced or metastatic urothelial carcinoma who have disease progression during or following platinum-containing chemotherapy. Until now, there has been no established standard of care for patients progressing after platinum-containing chemotherapy; thus, pembrolizumab fills an unmet need in this patient population.<sup>1</sup> A BC Cancer CAP approval is required. As per below, pembrolizumab can be given every 3 or 6 weeks.

Protocols Dosing Schedules			
UGUAVPEM	Pembrolizumab 2 mg/kg IV every 3 weeks (maximum 200 mg)		
UGUAVPEM6	Pembrolizumab 4 mg/kg IV every 6 weeks (maximum 400 mg)		

Approval of this new treatment is based on the phase III KEYNOTE-045 trial, which compared pembrolizumab with investigator's choice of chemotherapy (paclitaxel, docetaxel or vinflunine) in the second-line setting.<sup>2</sup> After a median follow-up of 22.5 months, the median overall survival (mOS) was significantly improved in the pembrolizumab group (10.3 months vs. 7.4 months, HR 0.70, 95% CI 0.57-0.86). Pembrolizumab was associated with a lower rate of any-grade (60.9% vs. 90.2%) and grades 3 to 5 (15.0% vs. 49.4%) treatment-related adverse events (TRAEs). Immune-mediated adverse events occurred more frequently with pembrolizumab, including hypothyroidism (6.4% vs. 1.2%), pneumonitis (4.1% vs. 0.4%), hyperthyroidism (3.8% vs. 0.4%) and colitis (2.3% vs. 0.4%).

#### Lung

**First-Line Platinum-Pemetrexed with Pembrolizumab for Advanced Non-Squamous Non-Small Cell Lung Cancer (NSCLC)** — The Lung Tumour Group is implementing this first-line combination therapy for patients whose disease demonstrates any level of programmed death-ligand 1 (PD-L1) expression, but is negative for EGFR, ALK or ROS1 mutations. Four cycles of 3-weekly platinum-pemetrexed and pembrolizumab (ULUAVPPPMB) are followed by 3-weekly pemetrexed and pembrolizumab maintenance therapy for up to two years of total treatment (LUAVPPMBM). If pemetrexed maintenance is discontinued due to toxicity, patients may continue maintenance pembrolizumab alone (LUAVPMBM or LUAVPMBM6). A BC Cancer CAP approval is required for the initial 4-cycle treatment protocol. Note that the use of pembrolizumab in the first-line setting precludes the use of atezolizumab or nivolumab in subsequent lines of therapy.

Initial Protocol		Maintenance Protocol
		LUAVPPMBM
ULUAVPPPMB		Pemetrexed + Pembrolizumab every 3 weeks
atinum + Pemetrexed + Pembrolizumab	THEN	If pemetrexed-intolerant:
		LUAVPMBM Pembrolizumab every 3 weeks
every 3 weeks x 4 cycles		OR
		LUAVPMBM6 Pembrolizumab every 6 weeks

Approval of these programs is based on the phase III KEYNOTE-189 trial, which compared platinumpemetrexed chemotherapy plus either pembrolizumab or placebo.<sup>3,4</sup> The pembrolizumab-combination group demonstrated significantly longer mOS (not reached vs. 11.3 months, HR 0.49, 95% CI 0.38-0.64); the overall survival rate at 12 months was also significantly improved (69.2% vs. 49.4%, HR 0.49, 95% CI 42.1-56.2). Rates of grade 3 or higher adverse events were comparable (67.2% vs. 65.8%), although adverse events that led to treatment discontinuation were reported more frequently in the pembrolizumab-combination group (13.8% vs. 7.9%).

**First-Line Paclitaxel-Carboplatin with Pembrolizumab for Advanced Squamous NSCLC** — The Lung Tumour Group is implementing this first-line combination therapy with paclitaxel-carboplatin and pembrolizumab for patients with advanced, squamous NSCLC with any level of PD-L1 expression. Four cycles of 3-weekly paclitaxel-carboplatin and pembrolizumab (**ULUAVPCPMB**) are followed by pembrolizumab maintenance therapy given every 3 weeks (**LUAVPMBM**) or 6 weeks (**LUAVPMBM6**) for up to two years of total treatment. Platinum-gemcitabine and pembrolizumab may be used for patients intolerant to taxanes (**ULUAVPGPMB**). A BC Cancer CAP approval is required for the initial 4-cycle treatment protocol. Note that the use of pembrolizumab in the first-line setting precludes the use of atezolizumab or nivolumab in subsequent lines of therapy.

Initial Protocol		Maintenance Protocol
ULUAVPCPMB Paclitaxel + Carboplatin + Pembrolizumab every 3 weeks x 4 cycles If taxane-intolerant: ULUAVPGPMB Platinum + Gemcitabine + Pembrolizumab every 3 weeks x 4 cycles	THEN	<b>LUAVPMBM</b> Pembrolizumab <i>every 3 weeks</i> OR <b>LUAVPMBM6</b> Pembrolizumab <i>every 6 weeks</i>

Approval of this new treatment is based on the phase III KEYNOTE-407 trial, which compared paclitaxelcarboplatin chemotherapy plus either pembrolizumab or placebo.<sup>5,6</sup> The pembrolizumab-combination group demonstrated significantly longer mOS (15.9 months vs. 11.3 months, HR 0.64, 95% CI 0.49-0.85). Rates of grade 3 or higher adverse events were comparable (69.8% vs. 68.2%), although treatment discontinuation due to adverse events was reported more frequently in the pembrolizumab-combination group (13.3% vs. 6.4%).

#### Lymphoma

**Pembrolizumab for Relapsed or Refractory Hodgkin Lymphoma (ULYPEM, ULYPEM6)** — The BC Cancer Lymphoma Tumour Group is introducing pembrolizumab for patients with classical Hodgkin lymphoma (cHL) that has relapsed or progressed after autologous stem cell transplantation and brentuximab vedotin (BV). A BC Cancer CAP approval is required. Pembrolizumab joins nivolumab (ULYNIV, ULYNIV4) as a treatment option in this small, heavily pretreated patient population.

Protocols	Dosing Schedules				
ULYPEM	Pembrolizumab 2 mg/kg IV every 3 weeks (maximum 200 mg)				
ULYPEM6	Pembrolizumab 4 mg/kg IV every 6 weeks (maximum 400 mg)				

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Approval of this new treatment is based on the phase II KEYNOTE-087 trial, which evaluated pembrolizumab in patients with relapsed or refractory cHL. Applicable cohorts included patients whose disease had progressed after BV; an objective response rate was achieved in 73.9% (95% CI 61.9% to 83.7%) and 64.2% (95% CI 52.8% to 74.6%) of these patients.<sup>7,8</sup> The most common grade 1 or 2 TRAEs were hypothyroidism (12.4%) and pyrexia (10.5%), while the most common grade 3 TRAEs were neutropenia (2.4%), diarrhea (1%) and dyspnea (1%); no grade 4 TRAEs were reported in the trial.

#### Melanoma

**Pembrolizumab for Adjuvant Treatment of Resected Stage III-IV NED Melanoma (USMAJPEM)** — The BC Cancer Skin and Melanoma Tumour Group is introducing pembrolizumab as a new adjuvant treatment option for fully resected stage III or IV melanoma, regardless of BRAF mutation status. A BC Cancer CAP approval is required. Note that patients are eligible to receive one of pembrolizumab, nivolumab (USMAJNIV, USMAJNIV4) or combination dabrafenib/trametinib (USMAJDT), but not the sequential use of these agents.

Protocol	Dosing Schedule			
USMAJPEM	Pembrolizumab 2 mg/kg IV every 3 weeks (maximum 200 mg)			

Approval of this new program is based on the phase III, placebo-controlled KEYNOTE-054 trial, in which pembrolizumab demonstrated an improved 12-month recurrence-free survival (75.4% vs. 61.0%, HR 0.57, 98.4% CI 0.43-0.74).<sup>9,10</sup> More grades 3 to 5 TRAEs were reported in the pembrolizumab group (14.7% vs. 3.4%). Most grade 3 or 4 immune-related adverse events resolved within 2 months after the last dose of pembrolizumab.

#### References

- 1. Pan-Canadian Oncology Drug Review (pCODR) Expert Review Committee (pERC). Final recommendation for pembrolizumab (Keytruda®) for metastatic urothelial carcinoma. 15 February 2018.
- 2. Bellmunt J, de Wit R, Vaughn DJ, et al. Pembrolizumab as second-line therapy for advanced urothelial carcinoma. *New Engl J Med* 2017;376(11):1015-1026. <u>https://doi.org/10.1056/NEJMoa1613683</u>
- 3. Pan-Canadian Oncology Drug Review (pCODR) Expert Review Committee (pERC). Final recommendation for pembrolizumab (Keytruda®) for non-squamous non-small cell lung cancer (NSCLC). 16 May 2019.
- 4. Gandhi L, Rodríguez-Abreu D, Gadgeel S, et al. Pembrolizumab plus chemotherapy in metastatic non-small cell lung cancer. *New Engl J Med* 2018;378(22):2078-2092. https://doi.org/10.1056/NEJMoa1801005
- 5. Pan-Canadian Oncology Drug Review (pCODR) Expert Review Committee (pERC). Final recommendation for pembrolizumab (Keytruda®) for squamous non-small cell lung cancer (NSCLC). 12 December 2019.
- 6. Paz-Ares L, Luft A, Vicente D, et al. Pembrolizumab plus chemotherapy for squamous non-small cell lung cancer. *New Engl J Med* 2018;379(21):2040-2051. <u>https://doi.org/10.1056/NEJMoa1810865</u>
- 7. Pan-Canadian Oncology Drug Review (pCODR) Expert Review Committee (pERC). Final recommendation for pembrolizumab (Keytruda®) for classical Hodgkin lymphoma. 19 October 2017.
- 8. Chen R, Zinzani PL, Fanale MA, et al. Phase II study of the efficacy and safety of pembrolizumab for relapsed/refractory classic Hodgkin lymphoma. *J Clin Oncol* 2017;35:2125-2132. <u>https://doi.org/10.1200/JCO.2016.72.1316</u>
- 9. Pan-Canadian Oncology Drug Review (pCODR) Expert Review Committee (pERC). Final recommendation for pembrolizumab (Keytruda®) for melanoma adjuvant therapy. 18 July 2019.
- 10. Eggermont AMM, Blank CU, Mandala M, et al. Adjuvant pembrolizumab versus placebo in resected stage III melanoma. *N Engl J Med* 2018;378(19):1789-1801. https://doi.org/10.1056/NEJMoa1802357

### Drug Update

#### **New Immunotherapy Dosing Schedule Options**

Effective 01 June 2020, BC Cancer has approved new dosing schedule options for atezolizumab and pembrolizumab. These changes allow for additional flexibility in scheduling patient clinic appointments. For patients previously approved for these treatments by the BC Cancer Compassionate Access Program (CAP), new CAP approval is <u>not</u> required when switching to a new dosing schedule.

#### Atezolizumab

**4-Weekly Atezolizumab Dosing for Advanced NSCLC (ULUAVATZ4)** — Atezolizumab may now be administered as a 3-weekly or 4-weekly fixed-dose regimen.<sup>1</sup> Note that patients switching from the existing 3-weekly dosing schedule (ULUAVATZ) to the 4-weekly dosing schedule should receive the first 4-weekly dose on the day they are due for their next 3-weekly dose.

Atezolizumab dosing schedules are summarized below:

Protocols	Dosing Schedules
ULUAVATZ	Atezolizumab 1200 mg IV every 3 weeks
ULUAVATZ4 (new)	Atezolizumab 1680 mg IV every 4 weeks

#### Pembrolizumab

**6-Weekly Pembrolizumab Dosing for Unresectable or Metastatic Melanoma (USMAVPEM6)** — The BC Cancer Skin and Melanoma Tumour Group is adding a 6-weekly dosing schedule option for pembrolizumab in the metastatic setting, based on clinical data from the KEYNOTE-555 trial.<sup>2</sup> Note that patients switching from the existing 3-weekly dosing schedule (USMAVPEM) to the 6-weekly dosing schedule should receive the first 6-weekly dose on the day they are due for their next 3-weekly dose.

Pembrolizumab dosing schedules are summarized below:

Protocols	Dosing Schedules
USMAVPEM	Pembrolizumab 2 mg/kg IV every 3 weeks (maximum 200 mg)
USMAVPEM6 (new)	Pembrolizumab 4 mg/kg IV every 6 weeks (maximum 400 mg)

#### References

- 1. Morrissey KM, Marchand M, Patel H, et al. Alternative dosing regimens for atezolizumab: An example of model-informed drug development in the postmarketing setting. *Cancer Chemother Pharmacol* 2019;84:1257-1267. <u>https://doi.org/10.1007/s00280-019-03954-8</u>
- 2. Lala M, Akala O, Chartash E, et al. Pembrolizumab 400 mg Q6W dosing: First clinical outcomes data from KEYNOTE-555 cohort B in patients with metastatic melanoma. American Association for Cancer Research Annual Virtual Meeting 2020. Abstract #CT042.

### Drug Update

#### **Biosimilar Rituximab Coming Soon**

Effective 01 July 2020, the BC Cancer Provincial Systemic Therapy Program will implement the use of biosimilar rituximab for rituximab given intravenously. Note that the reference biologic product will continue to be used when rituximab is administered subcutaneously. Funding details and updates to applicable rituximab-containing documents will be outlined in the July 2020 Systemic Therapy Update.

More information and resource materials on biosimilar drugs are available on the BC Cancer website, located in the <u>Biosimilar Drugs</u> section.

#### **Discontinuation of Brand-Name Warfarin**

Warfarin is used for anticoagulation in select oncology patients. In BC, the majority of patients use a generic warfarin product. Effective 01 June 2020, the sale and distribution of all strengths of brand-name warfarin tablets (Coumadin<sup>®</sup>, Bristol Myers Squibb) will be discontinued in Canada. Full discontinuation is expected by 30 August 2020.

Clinicians and patients should be aware that warfarin generic alternatives remain available and covered under PharmaCare. If switching from brand-name warfarin to a generic warfarin product, additional laboratory (INR) testing is <u>not</u> required. Full details are summarized in the 21 May 2020 issue of the <u>BC PharmaCare Newsletter</u>.

#### **Manufacturer Patient Assistance Programs**

The listing of patient assistance programs offered by pharmaceutical manufacturers has been updated and can be accessed on the BC Cancer website under Health Professionals > Systemic Therapy > <u>Reimbursement & Forms</u>.

## **Drug Shortages**

The following are updates of drug supply shortages in BC. Full details about new, updated or resolved drug shortages, including recommended treatment alternatives, can be found in the *Briefing Notes* and email communications previously circulated to BC Cancer and the Community Oncology Network (CON).

#### New

#### Bromocriptine

(Adapted from BC Cancer Briefing Note 08May2020)

There is a current shortage of bromocriptine in Canada. The estimated release date for additional supply is July 2020. BC Cancer centres and CON sites may have limited quantities on hand and may run out prior to the release date. As a mitigation strategy, BC Cancer centres and CON sites may reduce the total quantity dispensed to enable all active patients to continue on treatment.

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### **Drug Shortages**

Bromocriptine is used at BC Cancer for the treatment of pituitary adenomas producing prolactin or growth hormone (CNB). If necessary, patients can be switched to cabergoline (CNCAB) or quinagolide (CNQUIN) as alternative treatment options. Existing bromocriptine supplies should be prioritized for patients who are pregnant or are trying to conceive, and for patients who are unable to tolerate cabergoline or quinagolide.

#### **Nystatin Oral Suspension**

(Adapted from BC Cancer Briefing Note 28May2020)

Commercially available nystatin oral suspension is an antifungal ingredient used in the preparation of BC Cancer Magic Mouthwash.

There is a current backorder on all brands of oral nystatin suspension in Canada. The estimated release date for additional supply is mid-June 2020. Although there is no anticipated supply interruption of oral nystatin suspension or BC Cancer Magic Mouthwash at BC Cancer centres, community pharmacies may have variable supplies of nystatin oral suspension on hand.

If a community pharmacy does not have nystatin oral suspension on hand, BC Cancer Magic Mouthwash may be compounded by substituting nystatin powder for nystatin oral suspension at an equivalent dose. Prescriptions compounded with nystatin powder will be covered by BC PharmaCare Special Authority with the existing PIN for BC Cancer Magic Mouthwash (22123334).

#### Resolved

#### Cabergoline

(Adapted from BC Cancer email communication 08May2020)

Cabergoline supplies are now readily available.

## Systemic Therapy Update Editorial Board

#### **Membership Update**

The Systemic Therapy Update Editorial Board sincerely thanks **Dr. Caroline Lohrisch** (Medical Oncologist, BC Cancer Breast Tumour Group and Department Head, Medical Oncology, BC Cancer – Vancouver Centre) for her invaluable contributions over the years. She is stepping down from the Editorial Board as a long-term member after serving as the Medical Oncology representative since November 2005. The ST Update has benefited from Dr. Lohrisch's tremendous expertise in systemic therapy, as well as from her aptitude for clarity in the written language.

At this time, the Editorial Board would like to welcome **Dr. Alina Gerrie** (Hematologist, BC Cancer Lymphoma Tumour Group) to the Board. Welcome Dr. Gerrie!

## Cancer Drug Manual<sup>©</sup>

All BC Cancer Drug Manual<sup>©</sup> documents can be accessed from the <u>Cancer Drug Manual<sup>©</sup></u> home page on the BC Cancer website.

#### **New Documents**

Note that the following drug is <u>not</u> a BC Cancer Benefit Drug and requires application to the BC Cancer Compassionate Access Program (CAP). The corresponding Monograph and Patient Handout are made available for reference only.

The **Cemiplimab Interim Monograph** has been updated to the full **Monograph**, and a **Patient Handout** has been developed. Expert review was provided by Dr. Vanessa Bernstein (Medical Oncologist) and Robert Tillmanns (Pharmacist) of the Skin and Melanoma Tumour Group. The updated monograph includes expanded sections, including *Pharmacokinetics, Special Precautions*, and *Dosage Guidelines*. Cemiplimab is a programmed death receptor-1 (PD-1) immune checkpoint inhibitor used in the treatment of squamous cell skin cancer. Usual cemiplimab dosing follows either a weight-based regimen (3 mg/kg IV every 2 weeks) or a flat-dose regimen (350 mg IV every 3 weeks).

Highlights from these documents include:

- potentially life-threatening immune-related reactions such as pneumonitis, hepatitis, endocrinopathies, encephalitis, meningitis and nephritis
- other common side effects include nausea, musculoskeletal pain, infection, diarrhea, constipation, fatigue and rash

Cemiplimab was previously added to the **Chemotherapy Preparation and Stability Chart** and evaluated for the **BC Cancer Hazardous Drug List**.

#### **Revised Documents**

Highlights of key changes are listed below:

#### Cabazitaxel Monograph and Chemotherapy Preparation and Stability Chart

*Cautions:* deleted brand-specific information about alcohol

Supply and Storage: added Sandoz as new brand; updated Sanofi-Aventis brand information Solution Preparation and Stability: deleted brand-specific information from Additional Information Chemotherapy Preparation and Stability Chart: added Sandoz formulation as new brand

#### Pembrolizumab Monograph

Uses: updated Health Canada-approved indications

## Benefit Drug List

### **New Programs**

Effective 01 June 2020, the following new treatment programs have been added to the BC Cancer <u>Benefit Drug List</u>:

Protocol Title	Protocol Code	Benefit Status
Treatment of Locally Advanced or Metastatic Urothelial Carcinoma using <b>Pembrolizumab</b>	UGUAVPEM	Restricted
Treatment of Locally Advanced or Metastatic Urothelial Carcinoma using 6-Weekly <b>Pembrolizumab</b>	UGUAVPEM6	Restricted
Treatment of Advanced Non-Small Cell Lung Cancer using 4-Weekly Atezolizumab	ULUAVATZ4	Restricted
Maintenance Therapy of Advanced Non-Small Cell Lung Cancer with Pembrolizumab	LUAVPMBM	Class I
Maintenance Therapy of Advanced Non-Small Cell Lung Cancer with 6-Weekly Pembrolizumab	LUAVPMBM6	Class I
First-Line Treatment of Advanced Squamous Non-Small Cell Lung Cancer with <b>Paclitaxel, Carboplatin</b> and <b>Pembrolizumab</b>	ULUAVPCPMB	Restricted
First-Line Treatment of Advanced Squamous Non-Small Cell Lung Cancer with <b>Platinum, Gemcitabine</b> and <b>Pembrolizumab</b>	ULUAVPGPMB	Restricted
Maintenance Therapy of Advanced Non-Squamous Non-Small Cell Lung Cancer with <b>Pemetrexed</b> and <b>Pembrolizumab</b>	LUAVPPMBM	Class I
First-Line Treatment of Advanced Non-Squamous Non-Small Cell Lung Cancer with <b>Platinum, Pemetrexed</b> and <b>Pembrolizumab</b>	ULUAVPPPMB	Restricted
Treatment of Relapsed or Refractory Hodgkin Lymphoma using Pembrolizumab	ULYPEM	Restricted
Treatment of Relapsed or Refractory Hodgkin Lymphoma using 6-Weekly Pembrolizumab	ULYPEM6	Restricted
Adjuvant Treatment of Resected Stage III – IV NED Melanoma using Pembrolizumab	USMAJPEM	Restricted
Treatment of Unresectable or Metastatic Melanoma using 6-Weekly Pembrolizumab	USMAVPEM6	Restricted

## Benefit Drug List

### **Revised Programs**

Effective 01 June 2020, the following treatment program has been revised on the BC Cancer <u>Benefit Drug</u> <u>List</u>:

Protocol Title	Protocol Code	Benefit Status
Treatment of Acute Lymphoblastic Leukemia and Lymphoblastic Lymphoma using <b>Asparaginase-Erwinia (Erwinase®)</b> in Patients Allergic to Asparaginase (Kidrolase®) or Pegaspargase (Oncaspar®)	LKNOS LYNOS	Class I (expanded eligibility)

### **Deleted Programs**

Effective 01 June 2020, the following treatment programs have been deleted from the BC Cancer <u>Benefit Drug List</u>:

Protocol Title	Protocol Code	Benefit Status
Palliative Treatment of Metastatic or Locally Advanced Gastric, Gastroesophageal Junction or Esophageal Adenocarcinoma using <b>Cisplatin</b> , Capecitabine and Trastuzumab	GIGAVCCT	Deleted (replaced by GIGAVCOXT)

## Highlights of New & Revised Protocols, PPPOs and Patient Handouts

**BC Cancer Protocol Summaries, Provincial Pre-Printed Orders (PPPOs) and Patient Handouts** are revised periodically. New, revised or deleted protocols, PPPOs and patient handouts for this month are listed below, with document revisions indicated in the respective columns. Protocol codes for treatment requiring BC Cancer Compassionate Access Program (CAP) approval are prefixed with the letter **U**.

<b>NEW Protocols, PPPOs and Patient Handouts</b> (new documents checked <b>1</b> )				
Code	Protocol Title	Protocol	РРРО	Handout
UGUAVPEM	Treatment of Locally Advanced or Metastatic Urothelial Carcinoma using Pembrolizumab	$\checkmark$	V	
UGUAVPEM6	Treatment of Locally Advanced or Metastatic Urothelial Carcinoma using 6-Weekly Pembrolizumab	$\checkmark$	V	
ULUAVATZ4	Treatment of Advanced Non-Small Cell Lung Cancer using 4-Weekly Atezolizumab	$\checkmark$	V	$\checkmark$
LUAVPMBM	Maintenance Therapy of Advanced Non-Small Cell Lung Cancer with Pembrolizumab	V	V	
LUAVPMBM6	Maintenance Therapy of Advanced Non-Small Cell Lung Cancer with 6-Weekly Pembrolizumab	$\checkmark$	V	
ULUAVPCPMB	First-Line Treatment of Advanced Squamous Non-Small Cell Lung Cancer with Paclitaxel, Carboplatin and Pembrolizumab		V	
ULUAVPGPMB	First-Line Treatment of Advanced Squamous Non-Small Cell Lung Cancer with Platinum, Gemcitabine and Pembrolizumab		Cisplatin and carboplatin PPPOs	
LUAVPPMBM	Maintenance Therapy of Advanced Non-Squamous Non-Small Cell Lung Cancer with Pemetrexed and Pembrolizumab		Ŋ	
ULUAVPPPMB	First-Line Treatment of Advanced Non-Squamous Non- Small Cell Lung Cancer with Platinum, Pemetrexed and Pembrolizumab	V	Cisplatin and carboplatin PPPOs	
ULYPEM	Treatment of Relapsed or Refractory Hodgkin Lymphoma using Pembrolizumab	$\checkmark$	V	
ULYPEM6	Treatment of Relapsed or Refractory Hodgkin Lymphoma using 6-Weekly Pembrolizumab	$\checkmark$	V	
USMAJPEM	Adjuvant Treatment of Resected Stage III – IV NED Melanoma using Pembrolizumab	$\checkmark$	V	$\checkmark$
USMAVPEM6	Treatment of Unresectable or Metastatic Melanoma using 6-Weekly Pembrolizumab	$\checkmark$	V	$\checkmark$

<b>REVISED Protocols, PPPOs and Patient Handouts</b> (revisions in respective columns)				
Code	Protocol Title	Protocol	РРРО	Handout
BR   Breast				
BRAVPTRAD	Palliative Therapy for Metastatic Breast Cancer using Pertuzumab, Trastuzumab (Herceptin®), and Docetaxel as First-Line Treatment for Advanced Breast Cancer		Minor typo corrected	
BRAVTCAP	Palliative Therapy for Metastatic Breast Cancer using Trastuzumab and Capecitabine		Herceptin® brand name deleted	
GI   Gastroint	testinal			
GIGAVFFOXT	Palliative Treatment of Metastatic or Locally Advanced Gastric, Gastroesophageal Junction or Esophageal Adenocarcinoma using Oxaliplatin, Fluorouracil, Leucovorin and Trastuzumab		Minor typo corrected	
GO   Gynecol	ogic			
GOCXCRT	Treatment of High-Risk Squamous Carcinoma, Adenocarcinoma, or Adenosquamous Carcinoma of the Cervix with Concurrent Cisplatin and Radiation	Eligibility, Tests and Dose Modifications (renal) revised	Pre-treatment tests revised	
GOOVGEM	Treatment of Relapsed/Progressing Epithelial Ovarian, Primary Peritoneal or Fallopian Tube Carcinoma using Gemcitabine		Tests revised and institutional logo updated	
GOOVTOP	Treatment of Relapsed/Progressing Epithelial Ovarian, Primary Peritoneal or Fallopian Tube Carcinoma using Topotecan	Tests clarified	Tests clarified and institutional logo updated	
GU   Genitou	rinary			
GUBEP	Curative Therapy for Germ Cell Cancer using Bleomycin, Etoposide and Cisplatin		Minor typo corrected	
HN   Head an	d Neck			
HNAVPE	Treatment of Recurrent and Metastatic Squamous Cell Cancer with Platinum and Etoposide	Institutional name and Dose Modifications revised		
HNNAVPE	Treatment of Recurrent and/or Metastatic Nasopharyngeal Cancer with Platinum and Etoposide	Dose Modifications revised		
HNOTVAN	Treatment for Locally Advanced or Metastatic Medullary Thyroid Cancer using Vandetanib	Tests and Precautions updated	Tests, Treatment and Return Appointment Orders updated	

<b>REVISED Protocols, PPPOs and Patient Handouts</b> (revisions in respective columns)				
Code	Protocol Title	Protocol	РРРО	Handout
LK   Leukemia				
ULKAMLAS	Therapy of Acute Myeloid Leukemia using Azacitidine and Sorafenib	Tests updated (serum bicarbonate and chloride added)	Tests updated (serum bicarbonate and chloride added)	
ULKMDSA	Therapy of Myelodysplastic Syndrome and Acute Myeloid Leukemia using Azacitidine	Tests updated (serum bicarbonate and chloride added)	Tests updated (serum bicarbonate and chloride added)	
LU   Lung				
ULUAVATZ	Treatment of Advanced Non-Small Cell Lung Cancer using Atezolizumab	Eligibility clarified	Hypersensitivity tray added	
MY   Myelom	a			
UMYCARLD	Therapy of Multiple Myeloma using Carfilzomib, Lenalidomide with Dexamethasone		Dispensing quantity revised	
UMYLDF	Treatment of Previously Untreated Multiple Myeloma and Not-Eligible for Stem Cell Transplant using Lenalidomide with Low-Dose Dexamethasone		Dispensing quantity revised	
UMYLDREL	Therapy of Relapsed Multiple Myeloma using Lenalidomide with Dexamethasone		Dispensing quantity revised	
UMYLENMTN	Maintenance Therapy of Multiple Myeloma using Lenalidomide		Dispensing quantity revised	
UMYPOMDEX	Therapy of Multiple Myeloma Using Pomalidomide with Dexamethasone		Dispensing quantity revised	
SA   Sarcoma				
SAAI	Therapy for Advanced Soft Tissue Sarcoma using Doxorubicin, Ifosfamide-Mesna	Mesna diluent updated		
SC   Supportiv	ve Care	·		
SCDRUGRX	Management of Infusion-Related Reactions to Systemic Therapy Agents	Bronchospasm management and references updated	Inhalers added	
SCNAUSEA	Prevention and Treatment of Chemotherapy-Induced Nausea and Vomiting in Adults	Netupitant- palonosetron and antiemetics for multi-day chemotherapy clarified		

<b>REVISED Protocols, PPPOs and Patient Handouts</b> (revisions in respective columns)					
Code	Protocol Title	Protocol	РРРО	Handout	
SM   Skin and	l Melanoma				
USMAJDT	Adjuvant Treatment of Stage III and IV, BRAF-Mutated, Fully Resected Melanoma using Dabrafenib and Trametinib	Eligibility updated			
USMAJNIV	Adjuvant Treatment of Resected Stage III – IV NED Melanoma using Nivolumab	Eligibility updated and treatment duration clarified	Repeat treatment checkbox added		
USMAJNIV4	Adjuvant Treatment of Resected Stage III – IV NED Melanoma using 4-Weekly Nivolumab	Eligibility updated and treatment duration clarified		Description of immunotherapy updated and corticosteroid examples clarified (dexamethasone added)	
USMAVFIPI	First-Line Treatment of Unresectable or Metastatic Melanoma using Ipilimumab				
USMAVIPI	Treatment of Unresectable or Metastatic Melanoma using Ipilimumab				
USMAVIPNI	Treatment of Unresectable or Metastatic Melanoma using Ipilimumab and Nivolumab				
USMAVNIV	Treatment of Unresectable or Metastatic Melanoma using Nivolumab		Repeat treatment checkbox added		
USMAVNIV4	Treatment of Unresectable or Metastatic Melanoma using 4-Weekly Nivolumab			_	
USMAVPEM	Treatment of Unresectable or Metastatic Melanoma using Pembrolizumab				
SMAVTMZ	Palliative Therapy for Malignant Melanoma with Brain Metastases using Temozolomide	Contact physician updated and protocol reformatted			
USMMCCAVE	Second-Line Treatment of Recurrent or Metastatic Merkel Cell Carcinoma using Avelumab			Immunotherapy and corticosteroids updated	
SMMCCPE	Treatment of Recurrent or Metastatic Merkel Cell Carcinoma (MCC) with Cisplatin and Etoposide	Contact physician and institutional name updated; Protocol reformatted			

Antiemetic regimens in the Premedications section have been revised (based on updated recommendations for the use of NK<sub>1</sub> receptor antagonists) in the following BC Cancer protocols and/or pre-printed orders:

CODE	Protocol Title	
BRAJAC	Adjuvant Therapy for Breast Cancer using Doxorubicin and Cyclophosphamide	
BRAJACT	Neoadjuvant or Adjuvant Therapy for Breast Cancer using Doxorubicin and Cyclophosphamide Followed by Paclitaxel	
BRAJACTG	Neoadjuvant or Adjuvant Therapy for Breast Cancer using Dose-Dense Therapy: Doxorubicin and Cyclophosphamide Followed by Paclitaxel	
BRAJACTT	Neoadjuvant or Adjuvant Therapy for Breast Cancer using Doxorubicin and Cyclophosphamide Followed by Paclitaxel and Trastuzumab	
BRAJACTTG	Neoadjuvant or Adjuvant Therapy for Breast Cancer using Dose-Dense Therapy: Doxorubicin and Cyclophosphamide Followed by Paclitaxel and Trastuzumab	
BRAJACTW	Neoadjuvant or Adjuvant Therapy for Early Breast Cancer using Doxorubicin and Cyclophosphamide Followed by Weekly Paclitaxel	
BRAJDAC	Neoadjuvant or Adjuvant Therapy for Breast Cancer using Cyclophosphamide, Doxorubicin and Docetaxel	
BRAJFEC	Adjuvant Therapy for Breast Cancer using Fluorouracil, Epirubicin and Cyclophosphamide	
BRAJFECD	Neoadjuvant or Adjuvant Therapy for Breast Cancer using Fluorouracil, Epirubicin, Cyclophosphamide and Docetaxel	
BRAJFECDT	Neoadjuvant or Adjuvant Therapy for Breast Cancer using Fluorouracil, Epirubicin and Cyclophosphamide Followed by Docetaxel and Trastuzumab	
BRAVAC	Palliative Therapy for Metastatic Breast Cancer using Doxorubicin and Cyclophosphamide	
BRLAACD	Treatment of Locally Advanced Breast Cancer using Doxorubicin and Cyclophosphamide Followed by Docetaxel	
BRLAACDT	Treatment of Locally Advanced Breast Cancer using Doxorubicin and Cyclophosphamide Followed by Docetaxel and Trastuzumab	
BRLACTWAC	Neoadjuvant Therapy for Triple-Negative Breast Cancer using Carboplatin and Weekly Paclitaxel Followed by Doxorubicin and Cyclophosphamide	
BRLATACG	Neoadjuvant Therapy for Breast Cancer using Dose-Dense Therapy: Paclitaxel Followed by Doxorubicin and Cyclophosphamide	
BRLATWAC	Neoadjuvant Therapy for Locally Advanced Breast Cancer using Weekly Paclitaxel Followed by Doxorubicin and Cyclophosphamide	
GIENDO2	Palliative Therapy for Pancreatic Endocrine Tumours using Streptozocin and Doxorubicin	
GIFIRINOX	Palliative Combination Chemotherapy for Advanced Pancreatic Adenocarcinoma using Irinotecan, Oxaliplatin, Fluorouracil and Leucovorin	
GIGAJCPRT	Adjuvant Chemotherapy of Gastric Cancer Patients with Completely Resected Gastric Cancer using Cisplatin and Capecitabine and Radiation Therapy	
GIGAVCC	Palliative Therapy of Metastatic or Locally Advanced Anal Squamous Cell Carcinoma using Cisplatin and Capecitabine	
GIGAVCFT	Palliative Treatment of Metastatic or Inoperable, Locally Advanced Gastric or Gastroesophageal Junction Adenocarcinoma using Cisplatin, Infusional Fluorouracil and Trastuzumab	
GIGECC	Perioperative Treatment of Resectable Adenocarcinoma of the Stomach, Gastroesophageal Junction or Lower 1/3 Esophagus using Epirubicin, Cisplatin and Capecitabine	
GIPAJFIROX	Adjuvant Chemotherapy for Resected Pancreatic Adenocarcinoma using Irinotecan, Oxaliplatin, Fluorouracil and Leucovorin	
GOBEP	Therapy of Non-Dysgerminomatous Ovarian Germ Cell Cancer using Bleomycin, Etoposide and Cisplatin	
GOCISP	Alternative Treatment of Gynecological Malignancies using Cisplatin and Paclitaxel	
GOEP	Therapy of Non-Dysgerminomatous Ovarian Germ Cell Cancer using Etoposide and Cisplatin	
GOOVCIS	Therapy for Invasive Epithelial Ovarian Cancer using Cisplatin	
GUAJPG	Adjuvant Therapy for Urothelial Carcinoma using Cisplatin and Gemcitabine	
GUAVPG	Palliative Therapy for Urothelial Carcinoma using Cisplatin and Gemcitabine	

CODE	Protocol Title	
GUBEP	Curative Therapy for Germ Cell Cancer using Bleomycin, Etoposide and Cisplatin	
GUEDPM	Treatment of Metastatic Adrenocortical Cancer with Etoposide, Doxorubicin, Cisplatin and Mitotane	
GUEP	Therapy for Nonseminoma Germ Cell Cancer using Etoposide-Cisplatin	
GUMVAC	Therapy for Transitional Cell Cancers of the Urothelium using Methotrexate, Vinblastine, Doxorubicin and Cisplatin	
GUNAJPG	Neoadjuvant Therapy for Urothelial Carcinoma using Cisplatin and Gemcitabine	
HNAVFUP	Treatment of Advanced Squamous Cell Carcinoma of the Head and Neck Cancer using Fluorouracil and Platinum	
HNAVPC	Treatment for Unresectable, Locoregionally Recurrent or Metastatic Squamous Cell Carcinoma of the Head and Neck using Paclitaxel and Cisplatin or Carboplatin	
HNAVPD	Treatment of Recurrent or Metastatic Squamous Cell Carcinoma of the Head and Neck with Platinum and Docetaxel	
HNAVPE	Treatment of Recurrent and Metastatic Squamous Cell Cancer with Platinum and Etoposide	
HNLAALTPRT	Treatment of Locally Advanced (Alternate) Head and Neck Cancer using Cisplatin During Radiation Therapy	
UHNLADCF	Treatment of Locally Advanced Squamous Cell Carcinoma of the Head and Neck with Docetaxel, Cisplatin and Infusional Fluorouracil	
HNLAPRT	Combined Chemotherapy (Cisplatin) and Radiation Treatment for Locally Advanced Squamous Cell Carcinoma of the Head and Neck	
HNNAVFUP	Treatment for Advanced Nasopharyngeal Cancer of the Head and Neck using Platinum and Fluorouracil	
HNNAVPE	Treatment of Recurrent and/or Metastatic Nasopharyngeal Cancer with Platinum and Etoposide	
HNNAVPG	Treatment of Locoregionally Recurrent and/or Metastatic Nasopharyngeal Cancer with Platinum and Gemcitabine	
HNNLAPG	Induction Treatment of Locally Advanced Nasopharyngeal Cancer with Cisplatin and Gemcitabine	
HNSAVFAC	Palliative Therapy for Advanced Salivary Gland Cancers using Cyclophosphamide, Doxorubicin and Fluorouracil	
HNSAVFUP	Treatment of Advanced Head and Neck Cancer using Cisplatin and Fluorouracil	
HNSAVNP	Treatment of Advanced Salivary Gland Cancers with Cisplatin and Vinorelbine	
HNSAVPAC	Treatment of Advanced Salivary Gland Cancers with Platinum, Doxorubicin and Cyclophosphamide	
LUAJNP	Adjuvant Cisplatin and Vinorelbine Following Resection of Non-Small Cell Lung Cancer	
LUAVDC	First-Line Treatment of Advanced Non-Small Cell Lung Cancer with Cisplatin and Docetaxel	
LUAVPG	Treatment of Advanced Non-Small Cell Lung Cancer with Platinum and Gemcitabine	
LUAVPP	First-Line Treatment of Advanced Non-Small Cell Lung Cancer with Platinum and Pemetrexed	
LUMMPG	Treatment of Malignant Mesothelioma with Platinum and Gemcitabine	
LUMMPP	Treatment of Malignant Mesothelioma with Platinum and Pemetrexed	
LUOTCAV	Treatment of Thymoma/Thymic Carcinoma with Cyclophosphamide, Doxorubicin and Vincristine	
LUOTPAC	Treatment of Thymoma with Platinum, Doxorubicin and Cyclophosphamide	
LUSCCAV	Treatment of Extensive Small Cell Lung Cancer with Cyclophosphamide, Doxorubicin and Vincristine	
LUSCPI	Second-Line Treatment of Extensive Stage Small Cell Lung Cancer with Irinotecan with or without Platinum	
SAAI	Therapy for Advanced Soft Tissue Sarcoma using Doxorubicin, Ifosfamide-Mesna	
SAAI3	3-Day Doxorubicin-Ifosfamide-Mesna for Use in Patients with Advanced Soft Tissue Sarcoma	
SAAJADIC	Adjuvant Treatment of Patients with Soft Tissue Sarcoma using Doxorubicin and Dacarbazine	
SAAJAP	3-Day Doxorubicin-Ifosfamide-Mesna for Use in Patients with Advanced Soft Tissue Sarcoma	
SAAVADIC	Treatment of Patients with Soft Tissue Sarcoma using Doxorubicin and Dacarbazine	
SAAVAP	Therapy of Advanced Osteosarcoma using Doxorubicin and Cisplatin	
SAAVI	Therapy for Advanced Soft Tissue Sarcoma using Ifosfamide	
SAAVI3	3-Day Ifosfamide for Use in Patients with Advanced Soft Tissue Sarcoma	
SAAVIME3	3-Day Etoposide and Ifosfamide-Mesna for Patients with Advanced Soft Tissue or Bony Sarcomas	
SADTIC	High-Dose Single-Agent Dacarbazine for Metastatic Soft Tissue Sarcoma	

CODE	Protocol Title	
SAIME	Etoposide and Ifosfamide-Mesna for Use in Sarcomas	
SAVAC	Treatment of Sarcomas with Vincristine, Doxorubicin and Cyclophosphamide	
SAVDC	Adjuvant Therapy for Rhabdomyosarcoma using Vincristine, Dactinomycin, and Cyclophosphamide	
SAVDCM	Adjuvant Therapy for Rhabdomyosarcoma using Vincristine, Dactinomycin, Cyclophosphamide and Mesna	
SMDTIC	Therapy for Metastatic Malignant Melanoma using High-Dose Single-Agent Dacarbazine	

## **Resources and Contact Information**

Resource	Phone	Email / Toll Free / Fax		
Systemic Therapy Update: www.bccancer.bc.ca/health-professionals/clinical-resources/systemic-therapy/systemic-therapy-update				
Systemic Therapy Update Editor	604-877-6000 x 672649	bulletin@bccancer.bc.ca		
Oncology Drug Information	604-877-6275	druginfo@bccancer.bc.ca		
Cancer Drug Manual Editor	250-519-5500 x 693742	nbadry@bccancer.bc.ca		
Pharmacy Oncology Certification	250-712-3900 x 686820	rxchemocert@bccancer.bc.ca		
Nurse Educators	604-877-6000 x 672638	nursinged@bccancer.bc.ca		
Compassionate Access Program (CAP)	604-877-6277	cap bcca@bccancer.bc.ca fax 604-708-2026		
OSCAR – Online System for Cancer		oscar@bccancer.bc.ca		
Drugs Adjudication and Reimbursement	888-355-0355	fax 604-708-2051		
Library/Cancer Information	604-675-8003	toll free 888-675-8001 x 8003		
		requests@bccancer.bc.ca		
Library Document Delivery	604-675-8002	requests@bccancer.bc.ca		
Pharmacy Professional Practice	604-877-6000 x 672247	mlin@bccancer.bc.ca		
Professional Practice, Nursing	604-877-6000 x 672623	BCCancerPPNAdmin@ehcnet.phsa.ca		
Provincial Systemic Therapy Program	604-877-6000 x 672247	mlin@bccancer.bc.ca		
BC Cancer – Abbotsford	604-851-4710	toll free 877-547-3777		
BC Cancer – Kelowna	250-712-3900	toll free 888-563-7773		
BC Cancer – Prince George	250-645-7300	toll free 855-775-7300		
BC Cancer – Surrey	604-930-2098	toll free 800-523-2885		
BC Cancer – Vancouver	604-877-6000	toll free 800-663-3333		
BC Cancer – Victoria	250-519-5500	toll free 800-670-3322		

Community Oncology Network (CON) sites: To update your contact information, please contact: <u>bulletin@bccancer.bc.ca</u>

## Editorial Review Board

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