









15. Inhibiteurs des Tyrosines Kinases utilisés dans les CBNPC

Le tableau 8 reprend les principales modalités de prescription et de surveillance des ITK qui peuvent être utilisés dans les CBNPC.



TKI	Posologie	Forme	Dosages	Adaptation (/prise)	Cible(s)	Autorisation	Repas	Surveillance biologique	Surveillance Clinique (Appareils)	Autres surveillances
Adagrasib MRTX849 (Mirati)	600mg x 2/j				KRAS G12C			BH, créat	Digestif, général (asthénie, perte appétit), hépatique, hémato, cardiaque	ECG (QT)
Afatinib* GIOTRIF (Boehringer Ingelheim)	40mg x1/j	⊕	20mg 30mg 40mg 50mg	/ 10 à 20mg	EGFR	AMM		BH, iono (K), créat	Digestif – Cutané - Pulmonaire - Oculaire ORL (épistaxis, rhinorrhée) - Général, Déshydratation	FEVG / 3 mois
Alectinib* ALECENSA (Chugai / Roche)	600mg x2/j	⊖	150mg	/ 150mg	ALK	AMM		NFS, BH, CPK	Digestif – Cutané - Myalgie – Bradycardie - Pulmonaire – Oculaire – Œdèmes - Anémie	
Brigatinib# ALUNBRIG (Takeda)	90mg x1/j 7j puis 180mg x1/j	⊕	30mg 90mg 180mg	Voir notice	ALK	AMM		NFS, BH, CPK Lipase, Amylase, Gly, iono, créat	Digestif, pancréatique - Général – Hyperglycémie - Pulmonaire – Oculaire - Cardiaque (HTA, Bradycardie) - Œdème - Céphalées, neuropathies périphériques - Cutané - Musculaire	ECG (QT)
Capmatinib* (Novartis)	400mg x2/j	⊕	200mg	/ 100mg	MET (Mt ex 14)	ATU nominative		BH, Lipase, Amylase, Albumine, iono, créat, NFS	Digestif - Général - Cutané - Œdèmes	ECG (QT)
Ceritinib* ZYKADIA (Novartis)	450mg x1/j	⊖	150mg	/ 150mg	ALK	AMM		NFS, BH, Amylase, Lipase, iono (K), créat, Glycémie	Anémie - Hyperglycémie, hypophosphatémie Pulmonaire – Oculaire - Cardiaque (bradycardie, péricardite, QT) - Digestif, pancréatite - Cutané - Général	ECG (QT)
Crizotinib* XALKORI (Pfizer)	250mg x2/j	⊖	200mg 250mg	A 200mgx2 ou à 250mgx1	ALK, ROS1	AMM		BH, NFS, iono (K), créat	Digestif - Cardiaque (bradycardie, QT) Cutané – hématologique - Pulmonaire - Oculaire	ECG (QT)
Dabrafenib* TAFINLAR (Novartis)	150mg x2/j	⊖	50mg 75mg	/ 50mg	BRAF	AMM		BH, créat	Hypophosphatémie, hyperglycémie - Céphalées - Pulmonaire – Oculaire - Digestif – Cutané - Musculaire - Général	Examen dermato /mois, jusque 6mois après arrêt (Cancers cutanés)
Entrectinib ROZLYTREK (Roche)	600mg x1/j	⊖	100mg 200mg	/ 200mg	ROS1 ALK NTRK	ATU nominative		BH, Uricémie, NFS, créat, Lipase, Amylase, Albumine, iono, Gly	Neurologique – Musculo-squelettique – Oculaire – Général – Digestif – Pulmonaire – Déshydratation – Cutané – Hypotension	FEVG, ECG (QT)
Erlotinib* TARCEVA (Roche)	150mg x1/j	⊕	25mg 100mg 150mg	/ 50mg	EGFR	AMM		BH, iono, créat	Cutané – Digestif – Pulmonaire – Oculaire - Général	
Gefitinib* IRESSA (Astra Zeneca) & génériques	250mg x1/j	⊕	250mg	Aucune	EGFR	AMM		BH, iono (K), créat	Digestif – Cutané – Pulmonaire – Oculaire - Epistaxis	

Larotrectinib* VITRAKVI (Bayer)	100mg x2/j		25mg 100mg Sol. Buv. 20mg/ml	/ 25mg	NTRK	AMM, Post- ATU, en rétrocession Hosp.		NFS, BH	Neurologique (vertige, trouble de la marche, paresthésies) – Digestif - Musculo-squelettique - Général	
Lorlatinib* LORVIQUA (Pfizer)	100mg x1/j		25mg 100mg	/ 25mg	ALK, ROS1	AMM (ALK) Post-ATU, en rétrocession Hosp.		NFS, Lipase, Amylase, Cholesterol, Triglycérides	Général – Psychiatrique – Neurologique – Cutané – Oculaire - Musculo-squelettique - Digestif, pancréatique – Oedèmes	ECG (PR)
Mobocertinib TAK-788 (Takeda)	160mg/j		40mg	NC	EGFR ins 20	ATU nominative	<i>(pauvre en matière grasse) Avec 240mL d'eau.</i>	NFS, créat, BH, amylase, lipase, ionogramme complet (avant, J1C1, J15C1, puis au J1 de chaque cycle)	Digestif, xérose & éruption cutanées, général (fatigue, perte de poids), hématologique (anémie), mucite (buccale), dyspnée & toux, RGO, infection cutanées et onichopathies, déshydratation, augmentation de la créatinine de l'amylase et de la lipase, diminution de la natrémie, kaliémie, magnésémie, céphalée, élévation tension artérielle, douleur dorsale.	ECG avant, J1C1, J1C2, puis au J1 tous les 4 cycles. Echo cœur avant, J1C2, J1C5 et J1C9
Osimertinib* TAGRISSO (Astra-Zeneca)	80mg x1/j		40mg 80mg	/ 40mg	EGFR	AMM		NFS, BH, iono, créat	Digestif – Cutané – Pulmonaire - Hématologique	ECG (QT)
Pozitotinib NOV120101 (Spectrum Pharm.)	16mg/j		2mg 8mg	/ 2mg	EGFR ins 20	ATU nominative		NC	NC	NC
Pralsetinib* BLU-667 (Blueprint)	400mg x1/j		100mg	-	RET	ATU nominative		BH, NFS, CPK	Digestif – Pulmonaire - Cardiaque (HTA)	
Selitrectinib LOXO-195 (Loxo, Bayer)	NC		Sol. Buv. 20mg/ml	NC	NTRK	ATU nominative		NC	NC	NC
Selpercatinib* LOXO-292 (Loxo/Lilly)	160mg x2/j		20mg 80mg Sol. buv. 20mg/mL	-	RET	ATU nominative		BH, créat	Digestif - Cardiaque (HTA) – Général – Œdèmes - Céphalées	
Sotorasib AMG-510 (Amgen)	960mg x 1/j		120mg	/240mg	KRAS G12C	ATU nominative, ATU de cohorte attendue en 2021		iono, créat, NFP, au J1 de chaque cure Coag, TSH, cholesterol, TG au J1C1	Digestif, Hématologique (cytopénies), rénale, hépatique	ECG avant, J1C1,

Trametinib* MEKINIST (Novartis)	2mg x1/j	⊕	0,5mg 2mg	/ 0,5mg	MEK²	AMM		NFS, BH, CPK	Digestif – Oculaire – Musculaire – Cutané – Général – Œdème - Cardiaque (HTA, Altération FEVG) – Pulmonaire- Hématologique	Tension artérielle FEVG /3 mois
Vemurafenib* ZELBORAF (Roche)	960mg x2/j	⊕	240mg	/ 240mg	BRAF	Hors AMM ¹	<i>Eviter le jeun</i>	BH, iono (K), NFS	Digestif – Cutané – Musculaire – Générale - Cardiaque (Trouble du rythme, QT) – Œdème - Oculaire	Examen dermato /mois, jusque 6mois après arrêt (photosensibilité) ECG (QT)

Tableau 8 - Principales modalités d'utilisation des ITK dans les CBNPC.

Sources : * Société Française de Pharmacie Oncologique, Oncolien®, Fiches et vidéos d'aide au bon usage des traitements anticancéreux oraux, <https://oncolien.sfoo.com/> - # Centre National Hospitalier d'Information sur le Médicament, Base de données THERIAQUE, v4.2.4 déployé le 29/08/2017 et mise à jour le 17/12/2020, <http://www.theriaque.org/apps/contenu/acueil.php> - & Site de l'Agence Nationale de Sécurité des Médicaments, Résumé des Caractéristiques du Produit. - † Données de la littérature.

Notes : 1. Le vemurafenib ne dispose pas d'AMM dans le cancer du poumon. 2. Utilisé en association au Dabrafénib pour cibler les BRAF V600E.

Repas : ROUGE : A prendre en dehors des repas (pas de prise entre 1h avant et 3h après le repas) - VERT : A prendre au cours des repas - JAUNE : Pendant ou au cours des repas - GRIS : Inconnu.

Abréviations : BH : Bilan Hépatique, Iono : Ionogramme Sanguin, K : Kaliémie, NFS : Numération Formule Sanguine (+Plaquettes), Créat : Créatinine, CPK : Créatine Kinase, ECG : Electrocardiogramme, FEVG : Fraction d'Ejection du Ventricule Gauche, QT : Espace QT corrigé, PR : Espace PR.

NC : Non connu



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