

# 中山醫學大學附設醫院

# 肺癌診療指引

本臨床指引參考美國NCCN版本

肺癌多專科醫療團隊編修

2023/12/18 Version180 2022/12/26 Version17.0 2021/12/06 Version16.0 2021/01/25 Version15.0 2019/12/23 Version14.0 2019/01/10 Version13.0 2018/01/11 Version12.0 2016/12/15 Version11.0 2015/11/24 Version10.0 2014/12/09 Version 9.0 2013/12/24 Version 8.0 2012/12/11 Version 7.0 2012/01/03 Version 6.0 2010/08/05 Version 5.0 2009/12/15 Version 4.0 2008/05/27 Version 3.0 2008/02/05 Version 2.0

癌症委員會主任委員	癌症委員會執行長	癌症中心主任	團隊負責人

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## 肺癌診療指引

## AJCC 第八版 TNM stage

T	Primary Tumor
TX	Primary tumor cannot be assessed, or tumor proven by the presence of malignant cells in sputum or bronchial washings but not
	visualized by imaging or bronchoscopy
<b>T0</b>	No evidence of primary tumor
Tis	Carcinoma in situ
	Squamous cell carcinoma in situ (SCIS)
	Adenocarcinoma in situ (AIS): adenocarcinoma with pure lepidic pattern, ≤3 cm in greatest dimension
<b>T1</b>	umor ≤3 cm in greatest dimension, surrounded by lung or visceral pleura, without bronchoscopic evidence of invasion more
	proximal than the lobar bronchus (i.e., not in the main bronchus)
	*T1mi Minimally invasive adenocarcinoma: adenocarcinoma (≤3 cm in greatest dimension) with a predominantly lepidic pattern and
	≤5 mm invasion in greatest dimension
	*T1a Tumor ≤1 cm in greatest dimension. A superficial, spreading tumor of any size whose invasive component is limited to the
	bronchial wall and may extend proximal to the main bronchus also is classified as T1a, but these tumors are uncommon.
	*T1b Tumor >1 cm but ≤2 cm in greatest dimension
	*T1c Tumor >2 cm but ≤3 cm in greatest dimension
<b>T2</b>	Tumor >3 cm but ≤5 cm or having any of the following features: (1) Involves the main bronchus, regardless of distance to the
	carina, but without involvement of the carina; (2) Invades visceral pleura (PL1 or PL2); (3) Associated with atelectasis or obstructive
	pneumonitis that extends to the hilar region, involving part or all of the lung
	*T2a Tumor >3 cm but ≤4 cm in greatest dimension
	*T2b Tumor >4 cm but ≤5 cm in greatest dimension
<b>T3</b>	Tumor >5 cm but ≤7 cm in greatest dimension or directly invading any of the following: parietal pleura (PL3), chest wall (including
	superior sulcus tumors), phrenic nerve, parietal pericardium; or separate tumor nodule(s) in the same lobe as the primary
<b>T4</b>	Tumor >7 cm or tumor of any size invading one or more of the following: diaphragm, mediastinum, heart, great vessels, trachea,
	recurrent laryngeal nerve, esophagus, vertebral body, carina; separate tumor nodule(s) in an ipsilateral lobe different from that of the
	primary

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N	Regional Lymph Nodes
NX	Regional lymph nodes cannot be assessed
N0	No regional lymph node metastasis
N1	Metastasis in ipsilateral peribronchial and/or
	ipsilateral
	hilar lymph nodes and intrapulmonary nodes,
	including
	involvement by direct extension
N2	Metastasis in contralateral mediastinal,
	contralateral hilar,
	ipsilateral or contralateral scalene, or
	supraclavicular lymph
	node(s)
N3	Metastasis in contralateral mediastinal,
	contralateral hilar,
	ipsilateral or contralateral scalene, or
	supraclavicular lymph
	node(s)
7.5	
M	Distant Metastasis
M0	No distant metastasis
M1	Distant metastasis
M1a	Separate tumor nodule(s) in a contralateral
	lobe; tumor
	with pleural or pericardial nodules or
	malignant pleural or pericardial effusiona
M1b	Single extrathoracic metastasis in a single
	organ (including involvement of a single
	nonregional node)
M1c	Multiple extrathoracic metastases in a single
	organ or in multiple organs

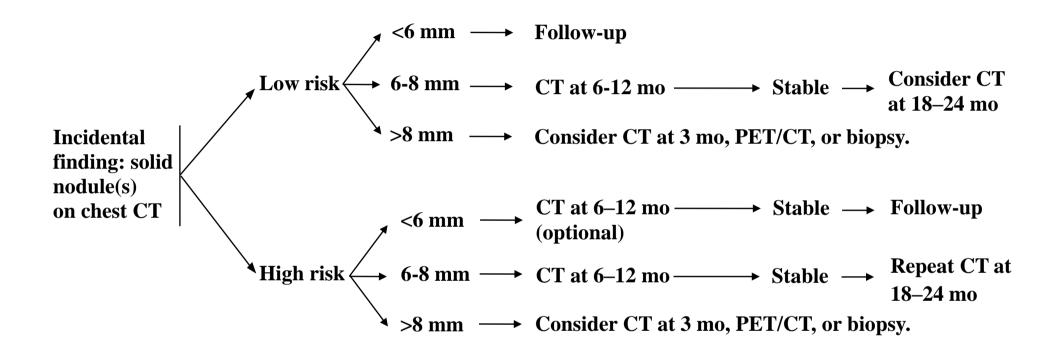
#### T N M TX N0 Occult M0 Carcinoma Stage 0 Tis N0 M0 Stage IA1 T1mi N0 M0 N0 T1a M0Stage IA2 T1b N0 M0 Stage IA3 T1c N0 M0 **Stage IB** T2a N0 M0N0 Stage IIA T2b M0 Stage IIB N1 T1a M0 T1b N1 M0 T2a N1 M0 T2b N1 M0 T3 N0 M0 **Stage IIIA** N2 M0 T1a T1b N2 M0 N2 T2a M0T2b N2 M0 T3 N1 M0T4 N0 M0 T4 N1 M0

肺癌診療指引

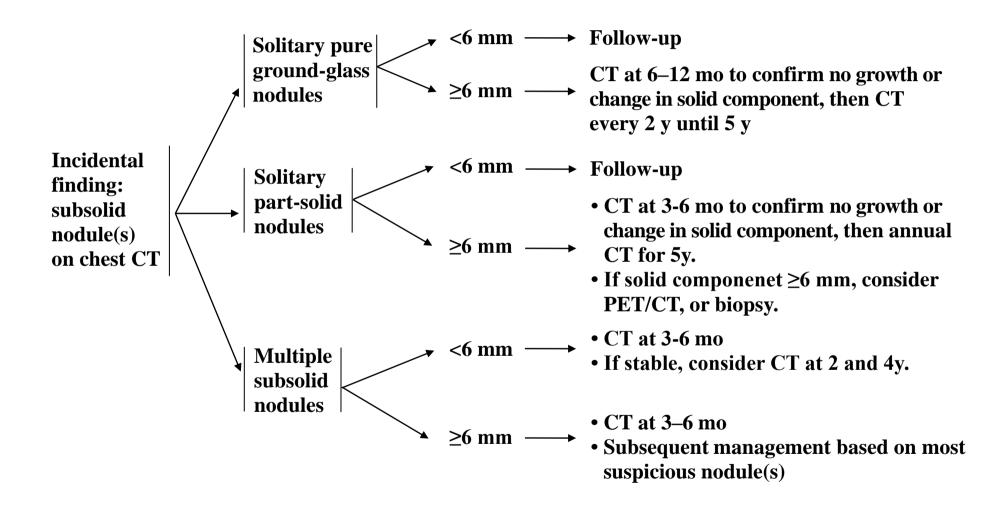
	T	N	M
Stage IIIB	T1a	N3	M0
	T1b	N3	M0
	T1c	N3	M0
	T2a	N3	M0
	T2b	N3	M0
	T3	N2	<b>M</b> 0
	T4	N2	M0
Stage IIIC	T3	N3	M0
	T4	N3	M0
Stage IVA	Any T	Any N	M1a
	Any T	Any N	M1b
Stage IVB	Any T	Any N	M1c

一、檢查

FINDINGS	FOLLOW-UP
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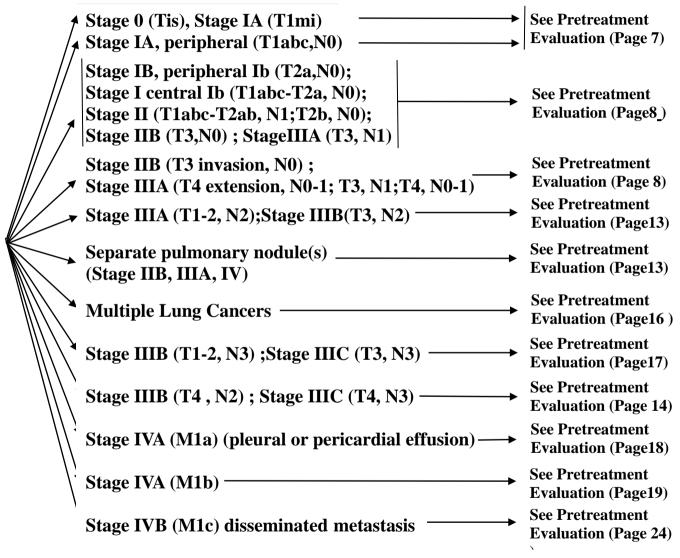
## FINDINGS FOLLOW-UP



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#### **CLINICAL STAGE**

- Pathology review
- H&P (include performance status+weight loss)
- CT chest and upper abdomen with contrast, including adrenals
- CBC, platelets
- Chemistry profile
- Smoking cessation advice, counseling, and pharmacotherapy
- Integrate palliative care
- For tools to aid in the optimal assessment and mamagement of older adults, see the NCCN **Guidelines for Older Adult Oncology**

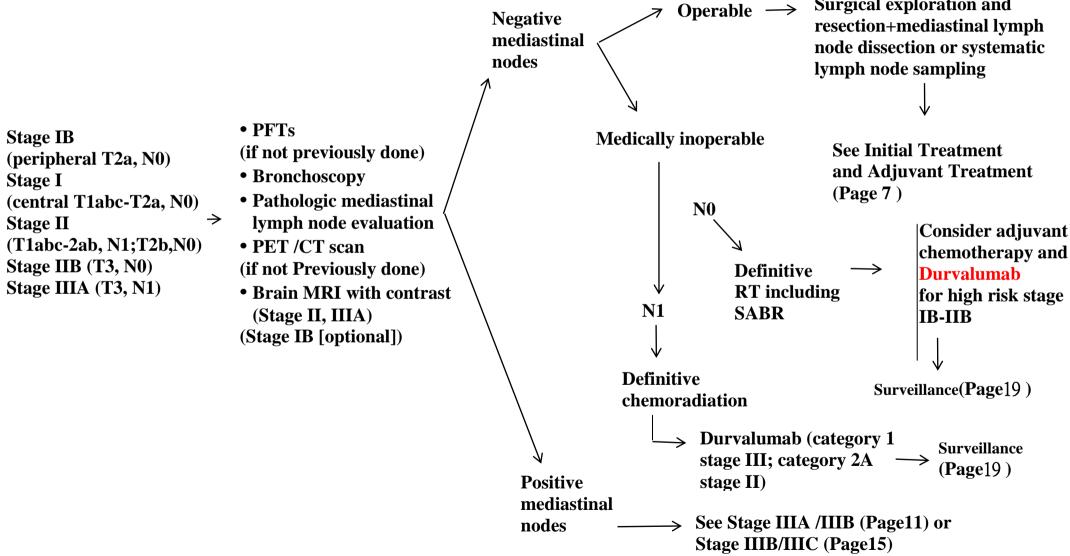


CLINICAL ASSESSMENT	PRETREATMENT EVA	LUATION	INITIAL TREATMENT
Stage 0 (Tis) Stage IA(T1mi)	→ • CT scan (if not Previous	ly done) — C	<b>Operation</b>
Stage IA (peripheral T1abc, N0)	<ul> <li>PFTs (if not previously done)</li> <li>Consider Bronchoscopy (intraoperative preferred)</li> <li>Consider pathologic Mediastinal lymph node Evaluation</li> <li>FDG PET/CT scan (if not Previously done)</li> <li>Observe</li> </ul>	Negative mediastinal nodes  Medically Described in the second sec	Surgical exploration and resection+mediastinal ymph — Treatment node dissection or systematic ymph node  efinitive RT including SABR thers: cryotherapy \ RFA \ MWA
		_	tage IIIA /IIIB(Page11) or IIIB/IIIC (Page15)

註:實際情況及手術與否需與胸腔內科/外科及放射腫瘤科等多專科團隊討論(SDM)

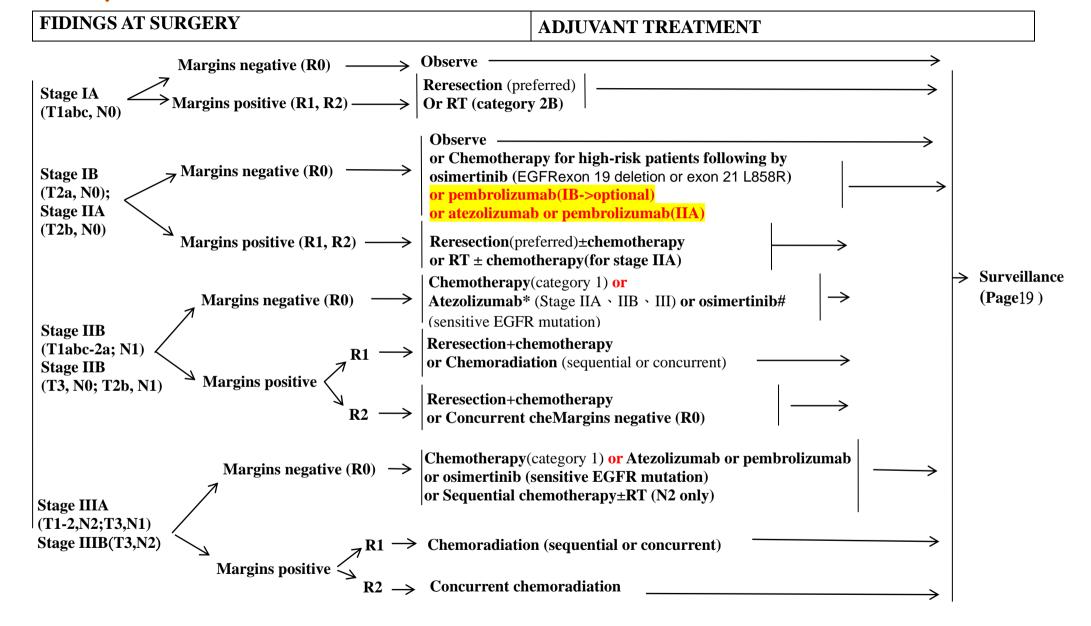
Note: All recommendations are category 2A unless otherwise indicated.

CLINICAL ASSESSMENT	PRETREATMENT EVALUATION	INITIAL TREATMENT
		<b>Z</b> Operable → Surgical exploration and



註:實際情況及手術與否需與胸腔內科/外科及放射腫瘤科等多專科團隊討論(SDM)

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註1:實際情況及手術與否需與胸腔內科/外科及放射腫瘤科等多專科團隊討論(SDM)

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**CLINICAL ASSESSMENT** 

PRETREATMENT EVALUATION

**CLINICAL EVALUATION** 

Stage IIB
(T3 invasion, N0)
Stage IIIA
(T4, extension, N0-1; T3, N1; T4,N0-1)

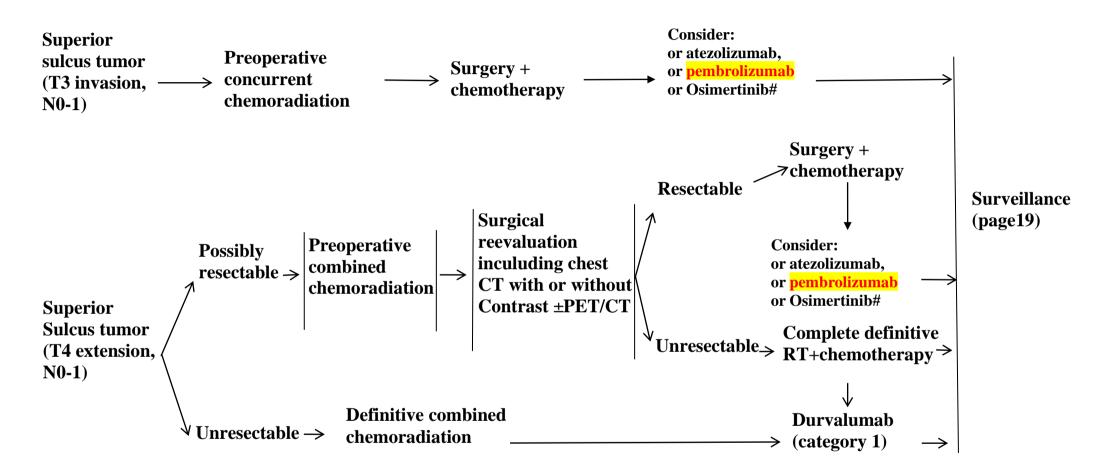
• PFTs (if not previously done)
• Bronchoscopy
• Pathologic mediastinal lymph node
evaluation
• Brain MRI with contrast
• MRI with contrast of spine + thoracic inlet
for superior sulcus lesions abutting the spine
or subclavian vessels.
• FDG PET/CT SCAN(if not previously done)

**Superior sulcus tumor** → Treatment (Page9) (評估有無侵犯血管或其他器官)  $\rightarrow$  Treatment (Page10) Chest wall **Proximal airway**  $\rightarrow$  Treatment (Page10) or mediastinum Stage IIIA(T4, N0-1)  $\longrightarrow$  Treatment (Page 10) **∠** Unresectable disease \_\_\_\_\_\_ Treatment (Page10) See Stage IIIA/IIIB \ Positive mediastinal **→** (Page11) nodes See Treatment for metastasis limited sites Metastatic disease (Page17) or distant disease (Page20-21)

**CLINICAL PRESENTATION** 

#### INITIAL TREATMENT

**ADJUVANT TREATMENT** 

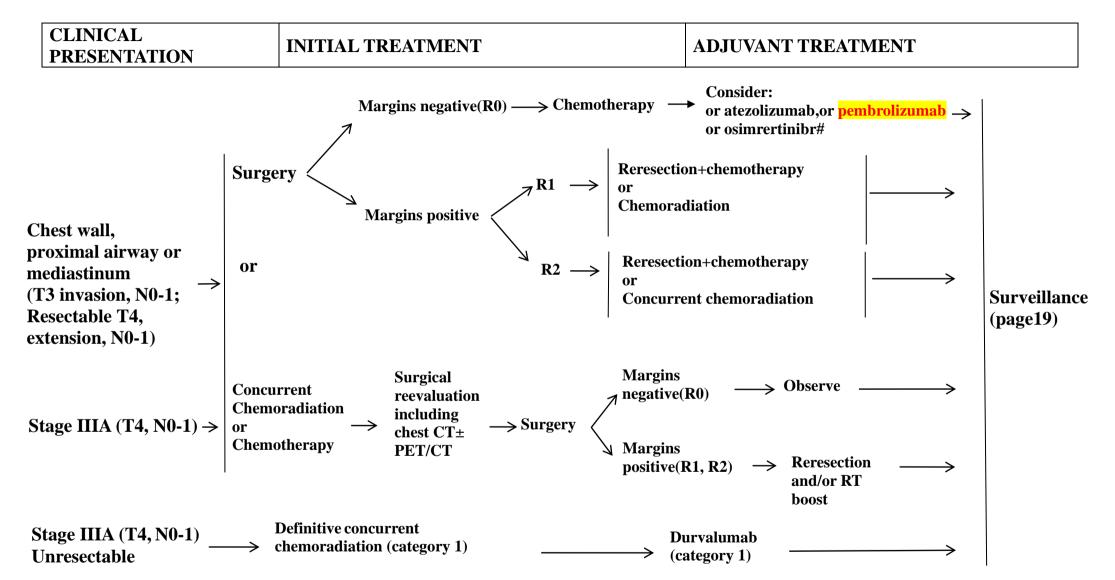


#: osimertinib :建議治療3年以上

註: pembrolizumab (keytruda)適合 II-IIIB(ACJJ8th)或 IB 且 Tumor>4cm,PDL1>1%,可考慮自費使用 from the phase 3 KEYNOTE-091 trial。

註1:實際情況及手術與否需與胸腔內科/外科及放射腫瘤科等多專科團隊討論(SDM)

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Atezolizumab:適用於IB以上,需自費。#: osimertinib:建議治療3年以上

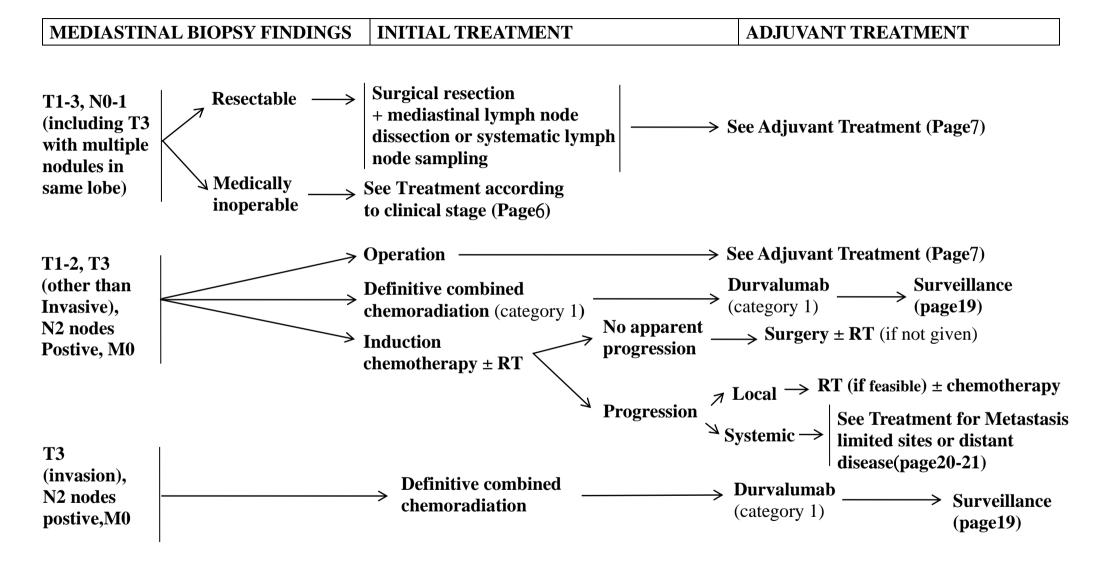
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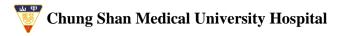
CLINICAL ASSESSMENT	PRETREATMENT EVALUATION	MEDIASTINAL BIOPSY FINDINGS AND RESECTABILITY
Stage IIIA (T1-2, N2) Stage IIIB (T3, N2)	<ul> <li>PFTs (if not previously done)</li> <li>Bronchoscopy</li> <li>Pothologic mediastinal lymph node evaluation</li> <li>Brain CT/MRI</li> <li>PET scan</li> </ul>	N2, N3 nodes negative   See Treatment T1-3.N0-1 (Page12)  N2 nodes positive, M0   See Treatment (Page12)  N3 nodes positive, M0   See Stage IIIB (Page15)  Metastatic disease   Treatment for Metastasis
	• PFTs (if not previously done)	Separate pulmonary   nodule(s), same lobe (T3, N0-1)   or ipsilateral non-primary   lobe (T4, N0-1)
Separate pulmonary nodules (Stage IIB,IIIA, IV)	<ul> <li>Bronchoscopy</li> <li>Pothologic mediastinal lymph node evaluation</li> <li>Brain CT/MRI</li> </ul>	Stage IV (N0, M1a): Contralateral lung (solitary nodule)  See Treatment (Page15)
	• Bone scan • PET scan	Extrathoracic metastatic disease See Treatment for Metastasis (Page19) or distant disease (Page20-21)

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註1:實際情況及手術與否需與胸腔內科/外科及放射腫瘤科等多專科團隊討論(SDM)

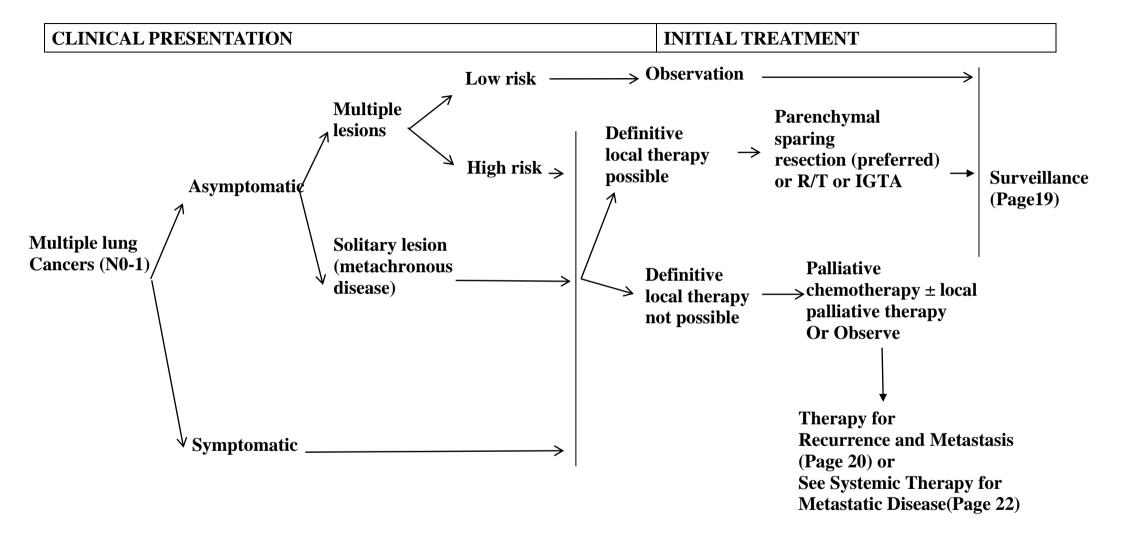
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#### **ADJUVANT TRETMENT CLINICAL PRESENTATION** consider atezolizumab.or $N_{0-1} \longrightarrow Chemotherapy \rightarrow$ oembrolizumab or Osimertinib Separate pulmonary nodule(s), same lobe Chemotherapy (category 1) Margins negative (T3, N0-1), or or $(\mathbf{R0})$ Suvgery Surveillance. ipsilateral non-primary Sequential chemotherapy+ RT (Page19) lobe (T4, N0-1) Chemoradiation (排除micro primary lung cancer) (sequential or concurrent) **Margins** positive Concurrent chemoradiation Stage IVA (N0, M1a) Treat as two primary lung **Contralateral lung** → See Evaluation (Page4) tumors if both curable (solitary nodule) **Disease** → See Systemic Therapy for Metastatic Disease (Page 22) outside Suspected multiple Chest CT with lung cancers (based on of chest contrast **See Initial Treatment** the presence of biopsy-• PET/CT scan (if proven synchronous not previously **Pathologic** lesions or history of done) **√** No disease mediastinal lung cancer) • Brain MRI outside of lymph node **See Systemic Therapy** chest **→** for Metastatic Disease evaluation (Page22)

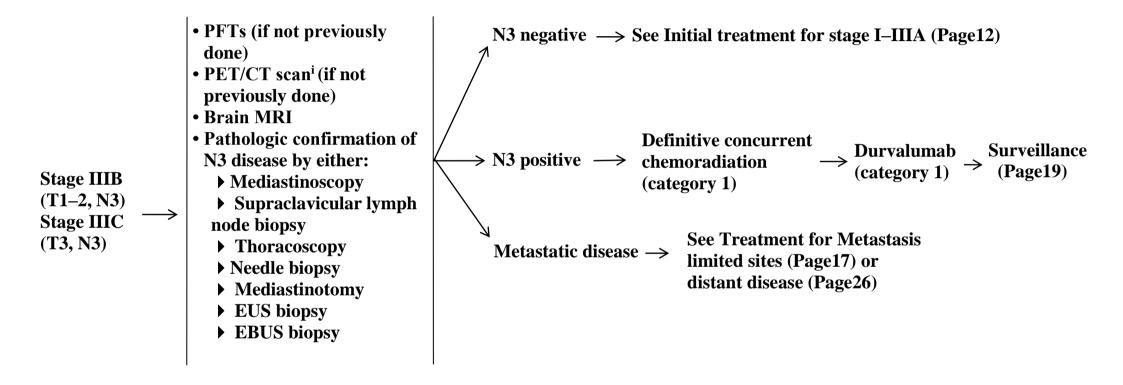
註1:實際情況及手術與否需與胸腔內科/外科及放射腫瘤科等多專科團隊討論(SDM)

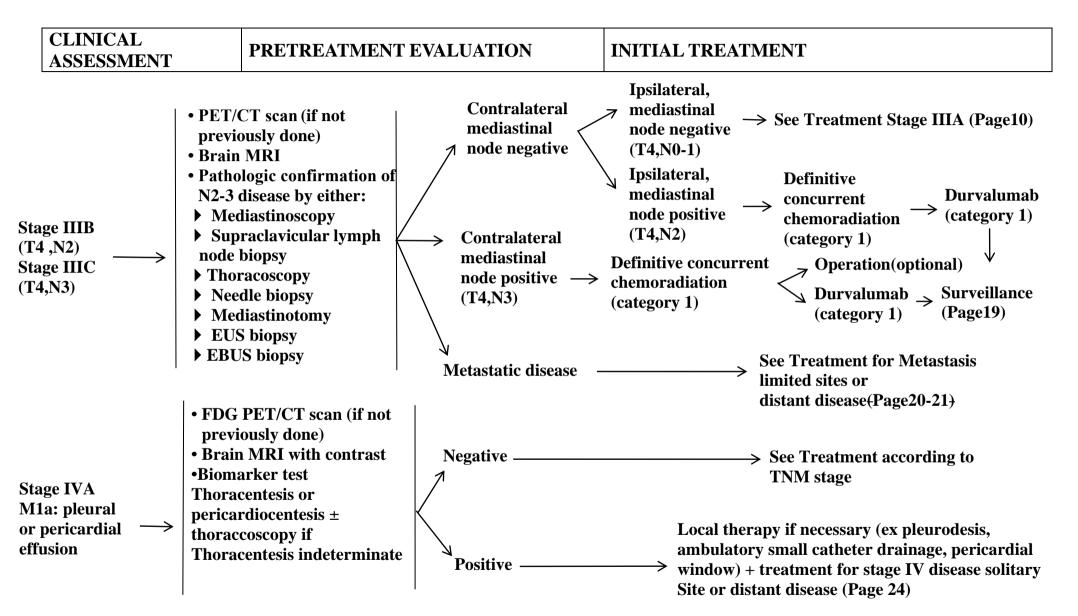
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CLINICAL ASSESSMENT	PRETREATMENT EVALUATION	INITIAL TREATMENT
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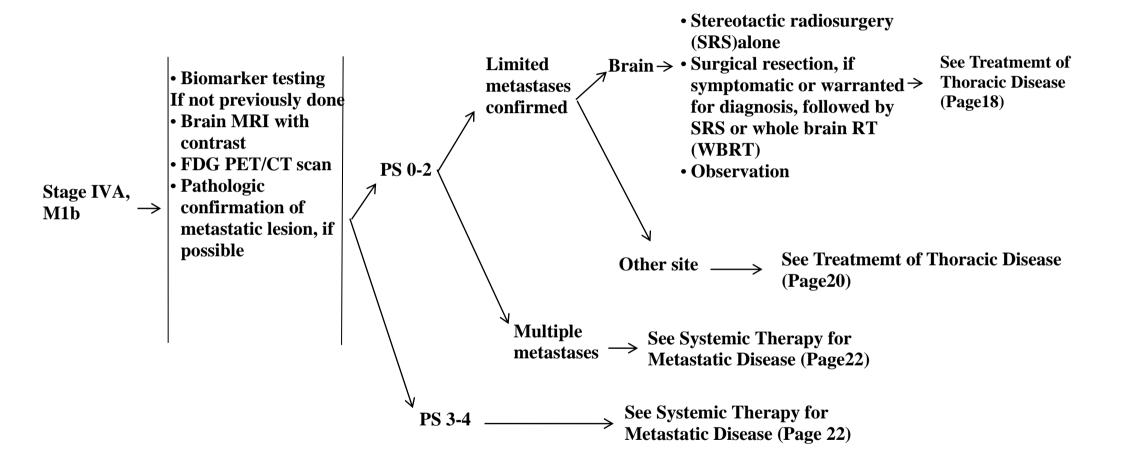
註1:實際情況及手術與否需與胸腔內科/外科及放射腫瘤科等多專科團隊討論(SDM)

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CLINICAL
<b>ASSESSMENT</b>

#### PRETREATMENT EVALUATION

#### INITIAL TREATMENT

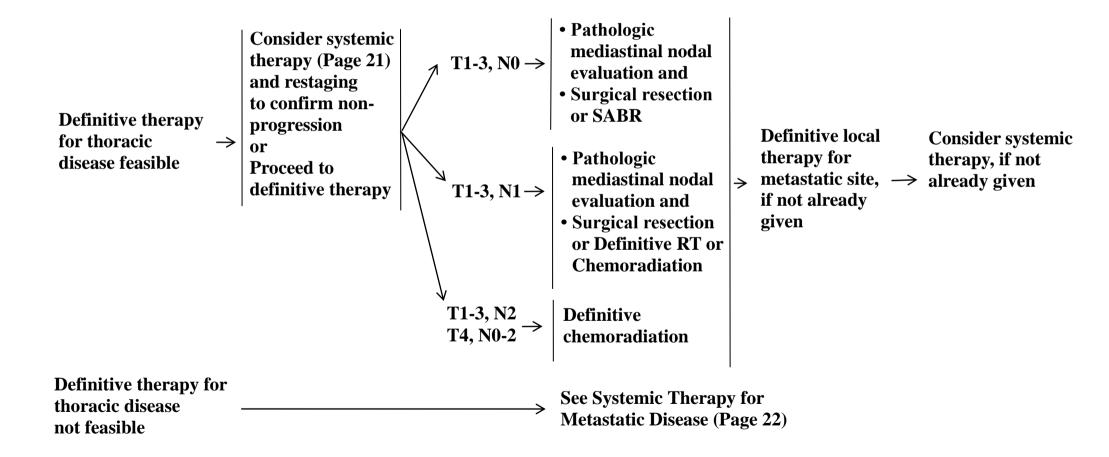


stageIV 若接受 TKI、化療、電療後若可開刀,可與胸腔內/外科等多專科團隊討論

註1:實際情況及手術與否需與胸腔內科/外科及放射腫瘤科等多專科團隊討論(SDM)

Note: All recommendations are category 2A unless otherwise indicated.

#### TREATMENT OF THORACIC DISEASE



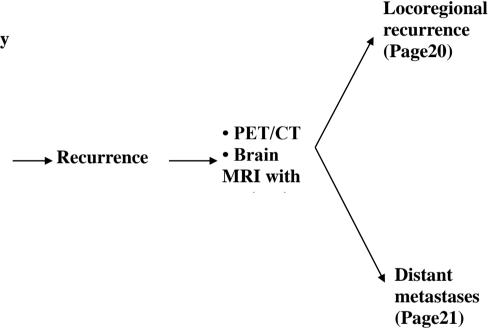
註1:實際情況及手術與否需與胸腔內科/外科及放射腫瘤科等多專科團隊討論(SDM)

Note: All recommendations are category 2A unless otherwise indicated.

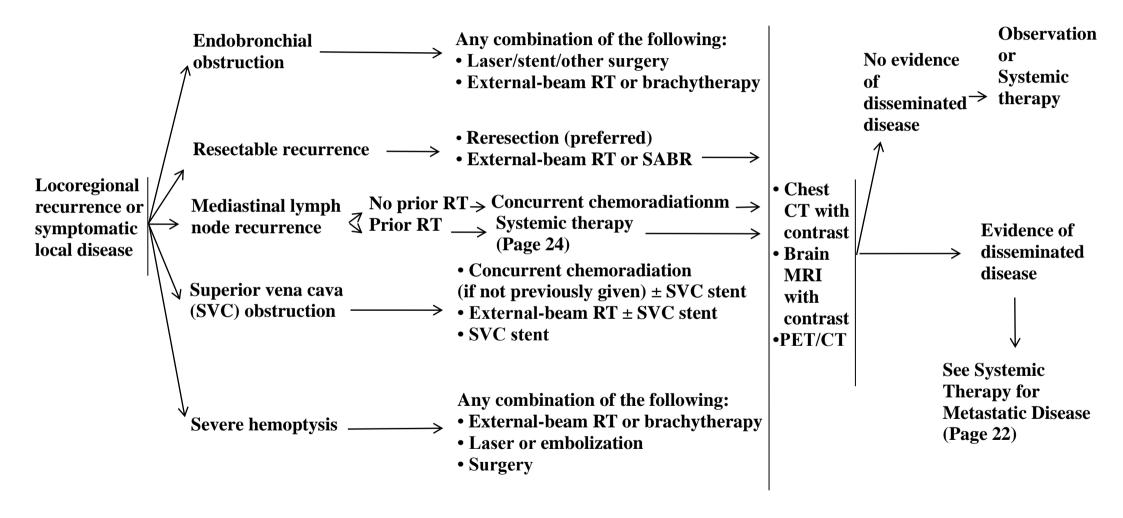
#### SURVEILLANCE AFTER COMPLETIOF DEFINITIVE

No evidence of clinical/radiographic disease

- Stage I–II (primary treatment included surgery ± chemotherapy) H&P and chest CT ± contrast every 3-6 mo for 2–3 y, then H&P and a LDCT
- Stage I–II (primary treatment included RT) or stage III or stage IV (oligometastatic with all sites treated with definitive intent)
   H&P and chest CT ± contrast every 3–6 mo for 3 y, then H&P and chest CT ± contrast every 6 mo for 2 y, then H&P and a LDCT
- Residual or new radiographic abnormalities may require more frequent imaging
- Smoking cessation advice, counseling, and pharmacotherapy
- PET/CT

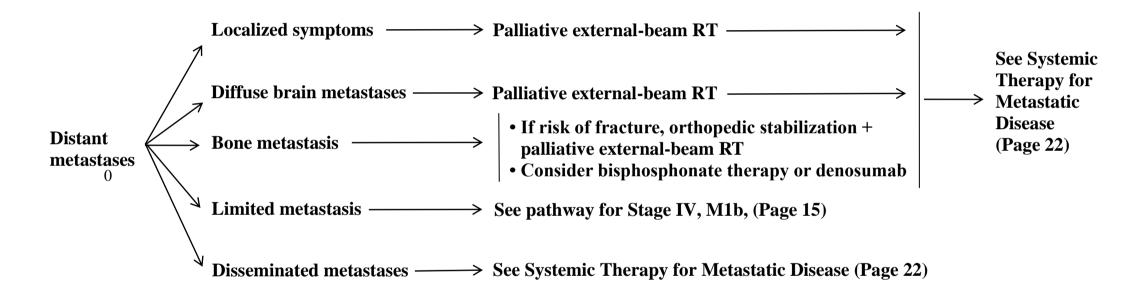


#### THERAPY FOR RECURRENCE AND METASIS



註:實際情況需與醫師討論(SDM)

#### THERAPY FOR RECURRENCE AND METASIS



註:實際情況需與醫師討論(SDM)

**TESTING SUBTYPE** 

• PD-L1 testing

**TESTING RESULTS** 

(Page 24)

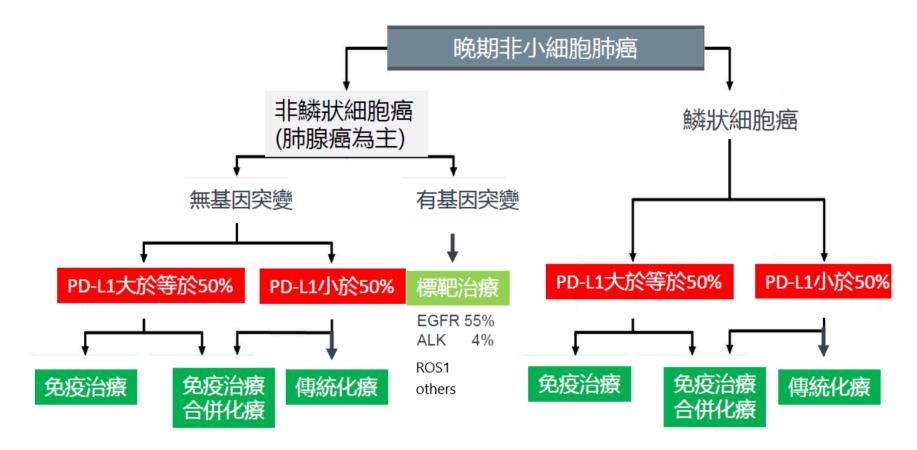
**HISTOLOGIC** 

**CLINICAL PRESENTATION** 

$\begin{array}{c} \textbf{Advanced} \\ \textbf{or} \\ \textbf{metastatic} \longrightarrow \end{array}$	<ul> <li>Establish histologic subtype with adequate tissue for molecular testing</li> <li>NGS test \ IHCtest \ FISH test \ Tumor marker \ HBV/HCV</li> </ul>	• Adenocarcinoma • Large cell • NSCLC not otherwise specified (NOS)	$\rightarrow$	<ul> <li>Molecular testing, ex:</li> <li>*EGFR mutation ,ALK,ROS1,</li> <li>*BRAF, METex14</li> <li>* RET,KRAS NTRK1/2/3,</li> <li>*ERBB2</li> <li>PD-L1 testing</li> </ul>		→ Testing Results (Page 24)
Disease	<ul><li>titer</li><li>Or plasma testing if appropriate)</li><li>Smoking cessation counseling</li><li>Integrate palliative care</li></ul>			<ul> <li>Consider molecular testing ex: (especially non-smoker):</li> <li>*EGFR mutation ,ALK,ROS1,</li> <li>*BRAF, METex14</li> </ul>		
		↓ Squamous cell carcinoma	$\rightarrow$	* RET,KRAS NTRK1/2/3, *ERBB2	$\rightarrow$	<b>Testing Results</b>

註:實際情況需與醫師討論(SDM)

## 晚期非小細胞肺癌藥物治療的選擇

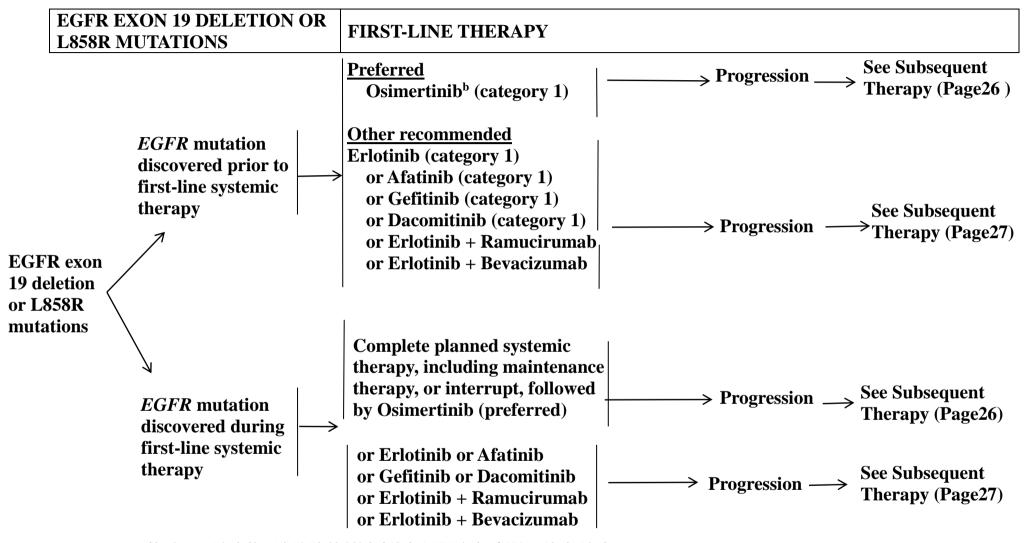


註:實際情況需與醫師討論(SDM)

### **TESTING RESULTS**

EGFR exon 19 deletion or L858R mutation positive	Page 25
EGFR S768I, L861Q, and/or G719X mutation positive	Page 28
EGFR exon 20 insertion mutation positive	Page 29
KRAS G12C mutation positive	Page 30
ALK rearrangement positive	Page31
ROS1 rearrangement positive	Page34
BRAF V600E mutation positive	Page 36
NTRK1/2/3 gene fusion positive	Page 37
METex14 skipping mutation positive	Page 38
RET rearrangement positive	Page 39
ERBB2 (HER2) mutation positive	Page 40
PD-L1 ≥50% and negative for actionable molecular biomarkers above	Page 41
PD-L1 ≥1%-49% and negative for actionable molecular biomarkers above	Page 42
PD-L1 <1% and negative for actionable molecular biomarkers above	Page 43

註:實際情況需與醫師討論(SDM)



Stage:IV 接受 TKI 治療後可與胸腔外科討論進行局部治療(含開刀.放療)治療

Note: All recommendations are category 2A unless otherwise indicated.

<sup>&</sup>lt;sup>a</sup>Clinical Trials: INSIGHT.

b建保條件:限單獨使用於(1)具有 EGFR Exon 19 Del 基因突變且無腦轉移(non-CNS)之轉移性(第IV期)肺腺癌病患之第一線治療。(2)先前已使 用過 EGFR 標靶藥物 Gefitinib、Erlotinib 或 Afatinib 治療失敗,且具有 EGFR T790M 基因突變之局部侵犯性或轉移性之非小細胞肺癌之第二線治療 實際情況雲與醫師討論(SDM)

#### SUBSEQUENT THERAPY EGFR EXON 19 DELETION OR L858R MUTATIONS Consider definitive local therapy **Asymptomatic** (SABR or surgery) for limited lesions • Continue Osimertinib<sup>b</sup> • Consider definitive local therapy **Progression on** Progression, see (eg, SRS) for limited lesions Osimertinib<sup>b</sup> therapy for Brain • Continue Osimertinib<sup>b</sup> multiple lesions, noted below • Consider definitive local therapy Limited (SABR or surgery) metastases **Symptomatic** • Continue Osimertinib<sup>b</sup> or • See subsequent therapy for multiple lesions, noted below **Systemic Multiple** systemic therapy lesions Adenocarcinoma or Squamous cell carcinoma

Stage:IV 接受 TKI 治療後可與胸腔外科討論進行局部治療(含開刀.放療)治療

b建保條件:限單獨使用於(1)具有 EGFR Exon 19 Del 基因突變且無腦轉移 (non-CNS) 之轉移性 (第IV期) 肺腺癌病患之第一線治療。(2)先前已使用過 EGFR 標靶藥物 gefitinib、erlotinib 或 afatinib 治療失敗,且具有 EGFR T790M 基因突變之局部侵犯性或轉移性之非小細胞肺癌之第二線治療用藥。

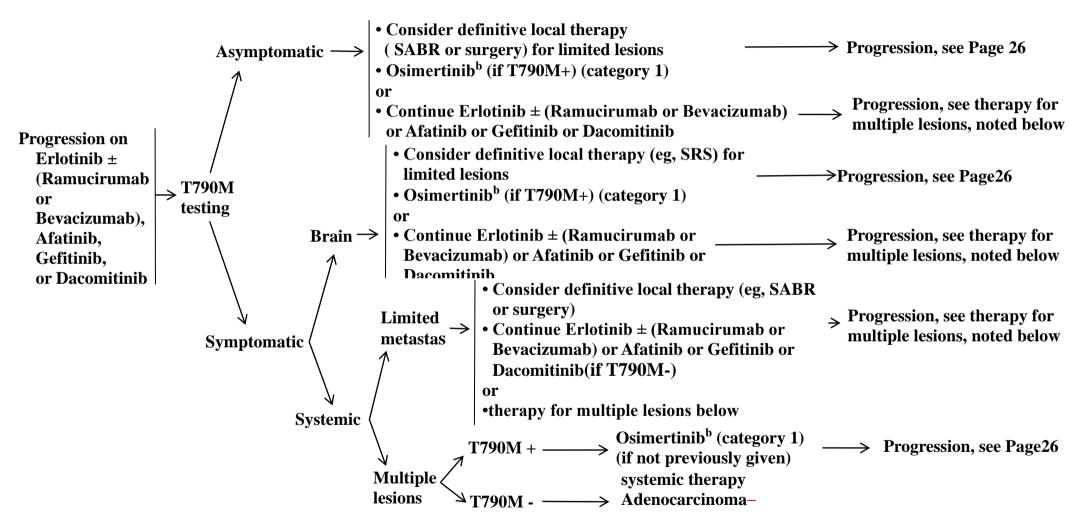
註:實際情況需與醫師討論(SDM)

Note: All recommendations are category 2A unless otherwise indicated.

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EXON 19 DELETION OR L858R MUTATION)

#### SUBSEQUENT THERAPY

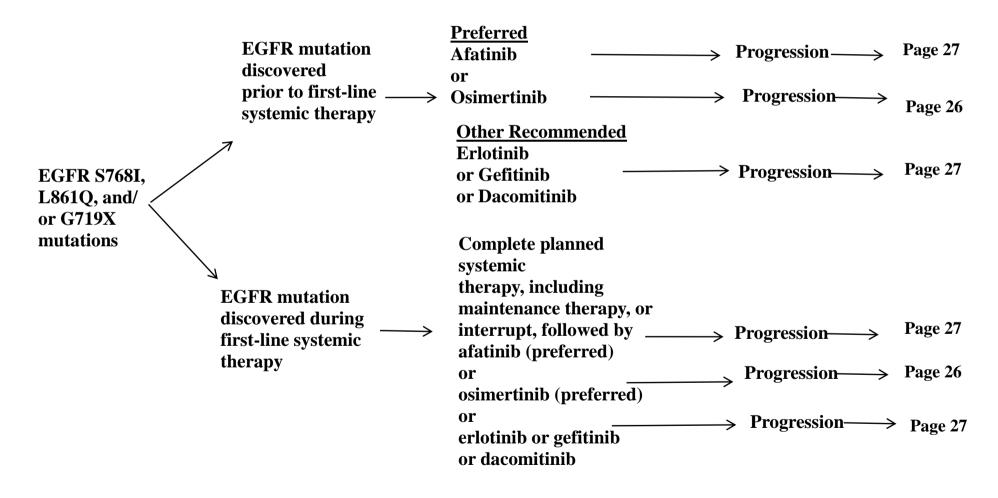


b建保條件:限單獨使用於(1)具有 EGFR Exon 19 Del 基因突變且無腦轉移 (non-CNS) 之轉移性 (第IV期) 肺腺癌病患之第一線治療。(2)先前已使用過 EGFR 標靶藥物 Gefitinib、Erlotinib 或 Afatinib 治療失敗,且具有 EGFR T790M 基因突變之局部侵犯性或轉移性之非小細胞肺癌之第二線治療用藥。

註:實際情況需與醫師討論(SDM)

EGFR S768I, L861Q, and/or G719X MUTATIONS

#### **FIRST-LINE THERAPY**



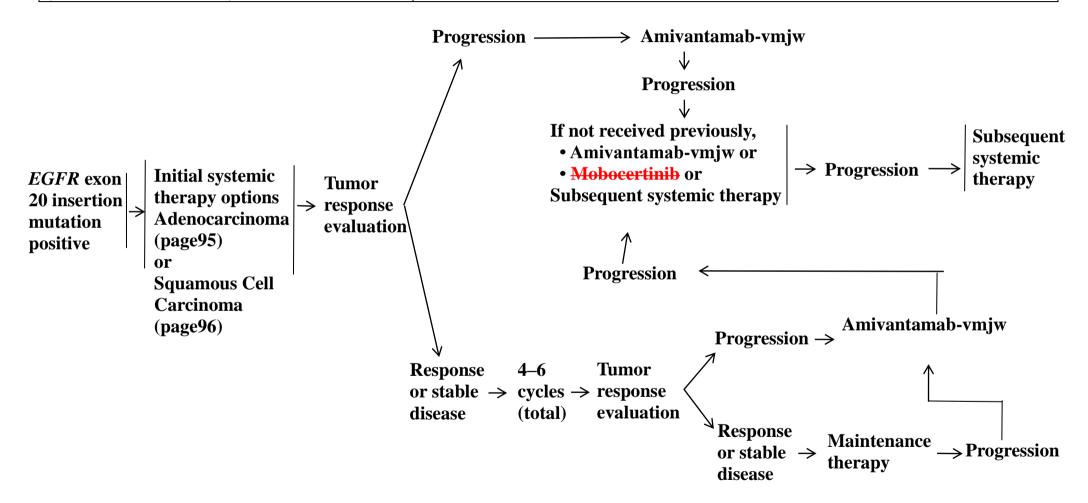
Stage:IV 接受 TKI 治療後可與胸腔外科討論進行局部治療(含開刀.放療)治療

註:實際情況需與醫師討論(SDM)

Note: All recommendations are category 2A unless otherwise indicated.

## EGFR MUTATION POSITIVE (EXON 20 INSERTION)

#### **SUBSEQUENT THERAPY**

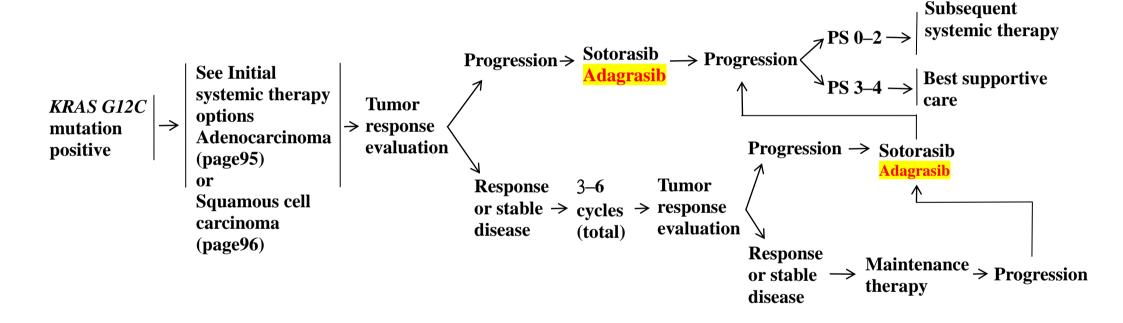


此類個案可建議使用 Afatinib。

註:實際情況需與醫師討論(SDM)

#### KRAS G12C MUTATION POSITIVE FIRST-LINE THERAPY

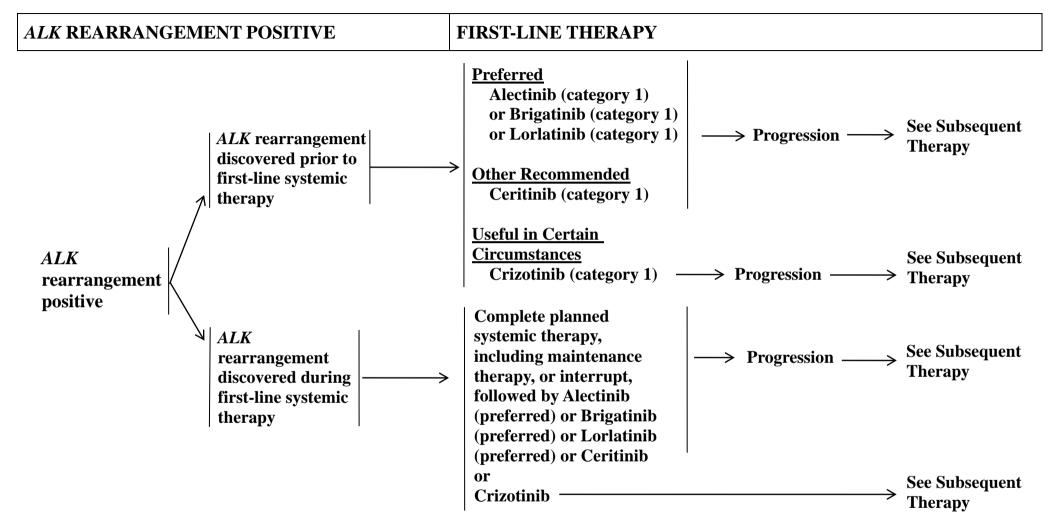
#### SUBSEQUENT THERAPY



Stage:IV 接受 TKI 治療後可與胸腔外科討論進行局部治療(含開刀.放療)治療

註:實際情況需與醫師討論(SDM)

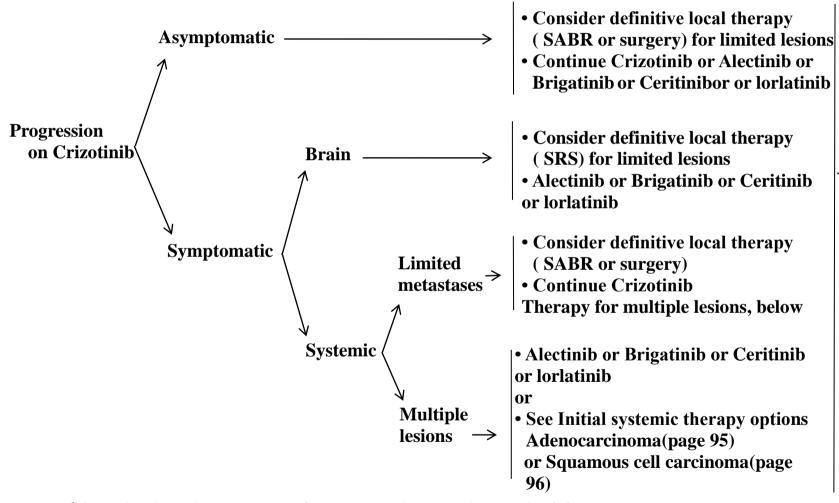
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註:實際情況需與醫師討論(SDM)

#### ALK REARRANGEMENT POSITIVE

#### SUBSEQUENT THERAPY



Progression, Lorlatinib (if not previously given) or **See Initial systemic** therapy options Adenocarcinoma or Squamous cell carcinoma

d建保條件:適用於在 Crizotinib 治療中惡化之 ALK 陽性的晚期非小細胞肺癌患者。

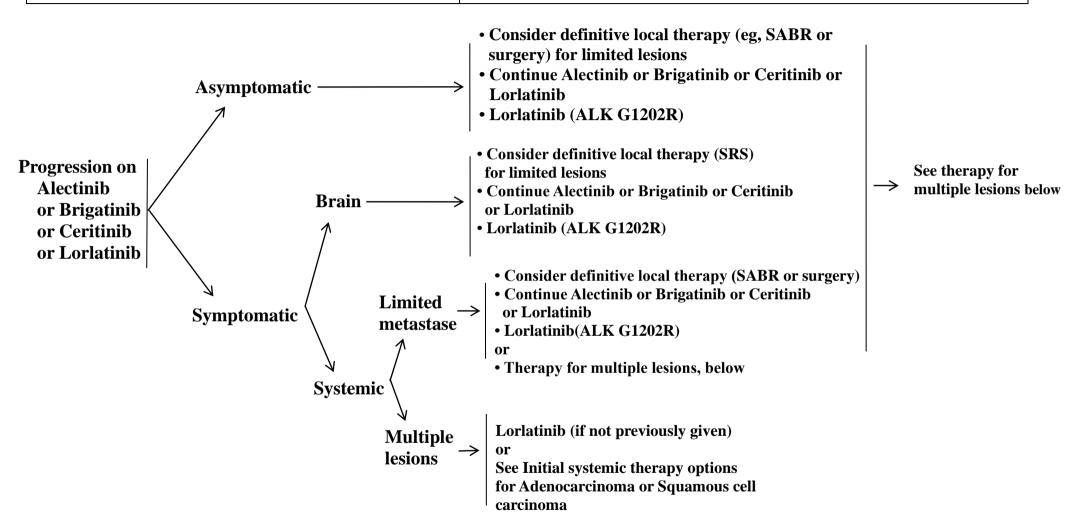
\*健保未給付 註:實際情況需與醫師討論(SDM)

Note: All recommendations are category 2A unless otherwise indicated.

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### ALK REARRANGEMENT POSITIVE

#### SUBSEQUENT THERAPY

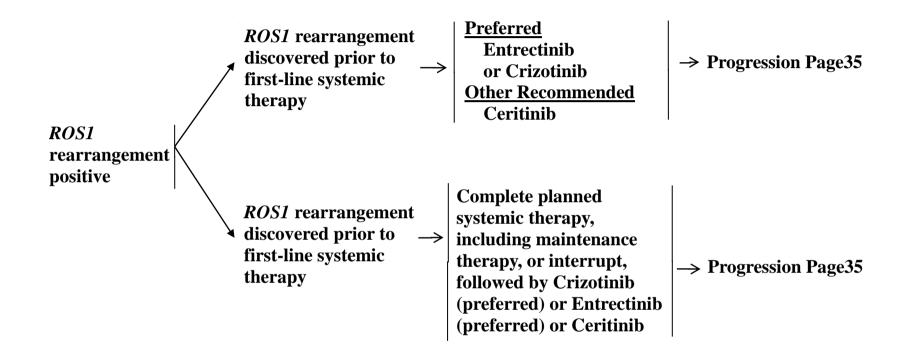


註:實際情況需與醫師討論(SDM)

**ROS1 REARRANGEMENT POSITIVE** 

FIRST-LINE THERAPY

**SUBSEQUENT THERAPY** 



#本院有藥但健保須送申請 ¥健保有給付

註:實際情況需與醫師討論(SDM)

Note: All recommendations are category 2A unless otherwise indicated.

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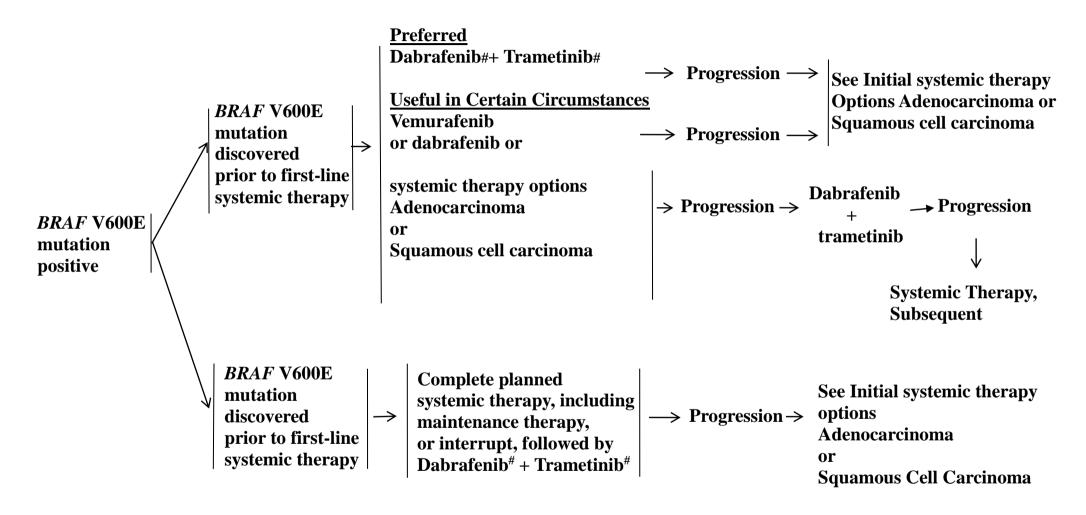
#### **ROS1 REARRANGEMENT POSITIVE** SUBSEQUENT THERAPY • Consider definitive local therapy (eg, SABR or surgery) for limited lesions • Continue entrectinib, crizotinib, or ceritinib or **Asymptomatic** Lorlatinib • Consider definitive local therapy (eg, Progression, Progression, **SRS**) for limited lesions Systemic therapy on • Entrectinib (if previously treated with Adenocarcinoma **Brain** entrectinib, crizotinib or ceritinib) orcrizotinib, **Squamous Cell** ceritinib • Consider definitive local therapy (eg, Carcinoma SABR or surgery) **Symptomatic Symptomatic** Limited → •Continue entrectinib, crizotinib, or metastases ceritinib or • Therapy for multiple lesions, below **Systemic** •Lorlatinib Multiple or • Systemic therapy options lesions Adenocarcinoma or **Squamous Cell Carcinoma**

註:實際情況需與醫師討論(SDM)

**BRAF V600E MUTATION POSITIVE** 

FIRST-LINE THERAPY

**SUBSEQUENT THERAPY** 



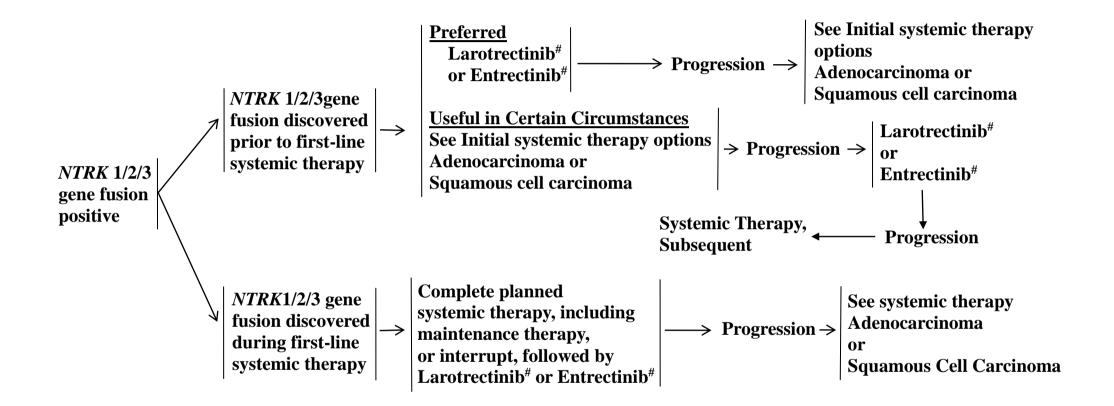
#本院有藥但健保未給付 Stage:IV 接受 TKI 治療後可與胸腔外科討論進行局部治療(含開刀.放療)治療 \$本院無此藥物

註:實際情況需與醫師討論(SDM)

*NTRK* GENE FUSION POSITIVE

FIRST-LINE THERAPY

**SUBSEQUENT THERAPY** 



Stage:IV 接受 TKI 治療後可與胸腔外科討論進行局部治療(含開刀.放療)治療

#本院有樂但健保未給付

註:實際情況需與醫師討論(SDM)

Adenocarcinoma or

Squamous cell carcinoma

SUBSEQUENT THERAPY **METex14 SKIPPING MUTATION** FIRST-LINE THERAPY **Preferred** Capmatinib# or Tepotinib  $\rightarrow$  Progression  $\rightarrow$ See Initial systemic therapy METex14 skipping options mutation discovered **Useful in Certain** Adenocarcinoma or prior to first-line Circumstances Squamous cell carcinoma → Progression – systemic therapy **Crizotinib** or **Preferred** See Initial systemic therapy METex14 Capmatinib# options skipping  $\rightarrow$  Progression  $\rightarrow$ or Tepotinib\$ Adenocarcinoma or mutation **Progression** Squamous cell carcinoma **Useful in Certain Circumstances Crizotinib** Systemic Therapy, Complete planned METex14 skipping **Subsequent** systemic therapy, including mutation discovered See systemic therapy maintenance therapy,  $\rightarrow$  Progression  $\rightarrow$ during first-line options or interrupt, followed by

#本院有藥但健保未給付

Stage:IV 接受 TKI 治療後可與胸腔外科討論進行局部治療(含開刀.放療)治療

Capmatinib<sup>#</sup> or Crizotinib

or Tepotinib<sup>\$</sup>

註:實際情況需與醫師討論(SDM)

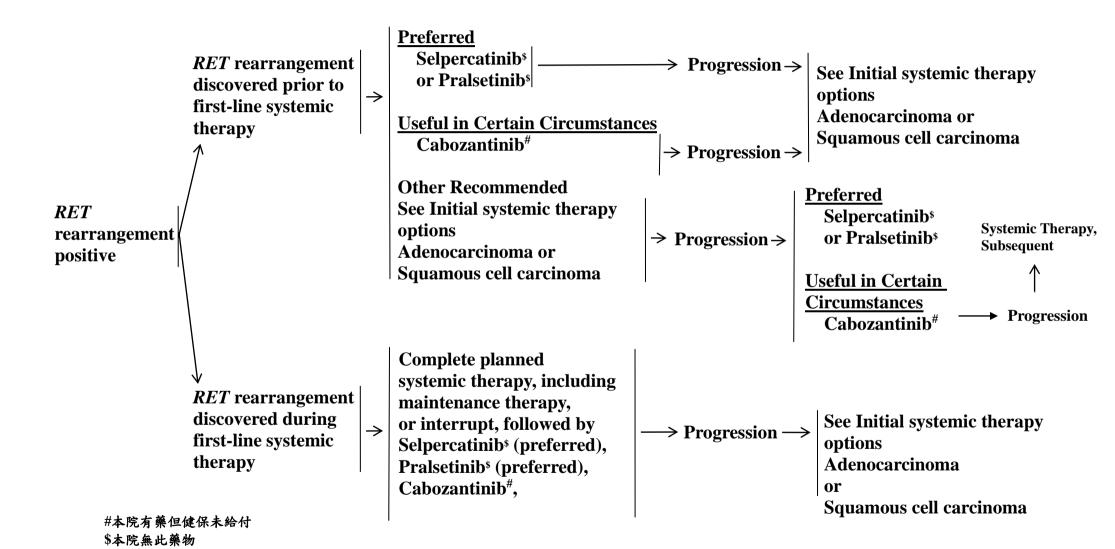
Note: All recommendations are category 2A unless otherwise indicated.

systemic therapy

RET REARRANGEMENT POSITIVI		RET	REARR	ANGEN	<b>JENT</b>	POSITIV	VE
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FIRST-LINE THERAPY

SUBSEQUENT THERAPY



註:實際情況需與醫師討論(SDM)

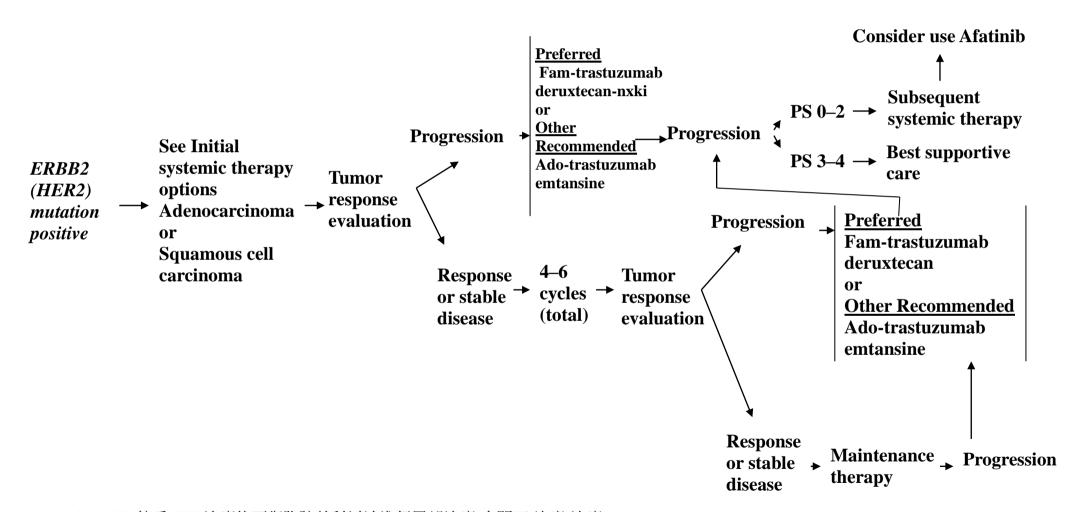
Note: All recommendations are category 2A unless otherwise indicated.

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**ERBB2 (HER2) MUTATION POSITIVE** 

FIRST-LINE THERAPY

SUBSEQUENT THERAPY



Stage:IV 接受 TKI 治療後可與胸腔外科討論進行局部治療(含開刀.放療)治療

註:實際情況需與醫師討論(SDM)

# **PD-L1** expression positive (≥50%) and negative for actionable molecular

**PS 0-2** 

**PS 3-4** 

### Adenocarcinoma, large cell, NSCLC NOS

Squamous cell carcinoma ->

**Best supportive** 

註:實際情況需與醫師討論(SDM)

care

#### FIRST-LINE THERAPY

#### **Preferred**

Pembrolizumab (category 1)

or (Carboplatin or Cisplatin) +

 $Pemetrexed + Pembrolizumab \ (category \ 1)$ 

or Atezolizumab\* (category 1)

or Cemiplimab-rwlc (category 1)

#### **Other Recommended**

Carboplatin + Paclitaxel + Bevacizumab

**±** Atezolizumab\* (category 1)

 $or\ Carboplatin+albumin-bound\ Paclitaxel$ 

**±** Atezolizumab\*

or (Carboplatin or Cisplatin) + Pemetrexed ±I/O(Nivolumab+Ipilimumab)

#### **Useful in Certain Circumstances**

Nivolumab\* + Ipilimumab\* (category 1)

# • Pembrolizumab (category 1)

• Pembrolizumab + Pemetrexed (category 1)

• Atezolizumab and Bevacizumab (category 1)

Atezolizumab

•Nivolumab + Ipilimumab

• Cemiplimab-rwlc (category 1)

or Subsequent Therapy

#### **Preferred**

Pembrolizumab (category 1)

 $or \ Carboplatin + (Paclitaxel \ or \$ 

 $albumin-bound\ Paclitaxel)\ +$ 

Pembrolizumab (category 1)

or Atezolizumab\* (category 1)

or Cemiplimab-rwlc (category 1)

#### **Other Recommended**

Carboplatin+ Paclitaxel ±I/O(Nivolumab +Ipilimumab)

#### **Useful in Certain Circumstances**

Nivolumab\* + Ipilimumab\* (category 1)

Response or stable • Pembro • Atezol

**Continuation maintenance** 

- Pembrolizumab (category 1)
- Atezolizumab
- Nivolumab + ipilimumab
- Cemiplimab-rwlc (category 1)

 $\begin{array}{c|c} \lor & \lor \\ Progression \rightarrow & See Systemic Therapy \\ or Subsequent Therapy \end{array}$ 

4 1 註\*Atezolizumab、Nivolumab、Ipilimumab 未有健保給付

disease

Response

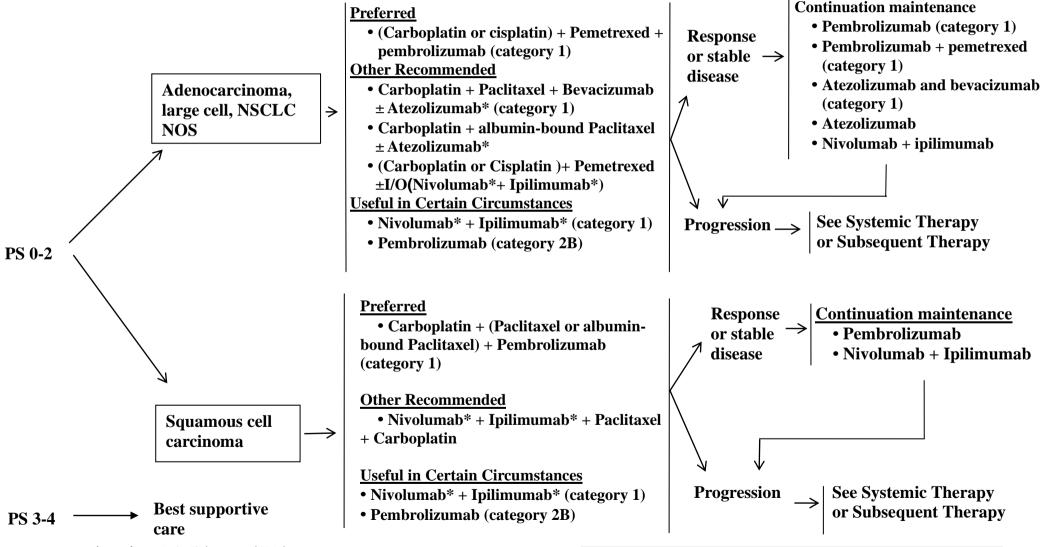
disease

or stable  $\rightarrow$ 

Note: All recommendations are category 2A unless otherwise indicated.

#### PD-L1 EXPRESSION POSITIVE(≥1%--49%)ACTIONABLE MOLECULAR **MARKERS**

#### FIRST-LINE THERAPY

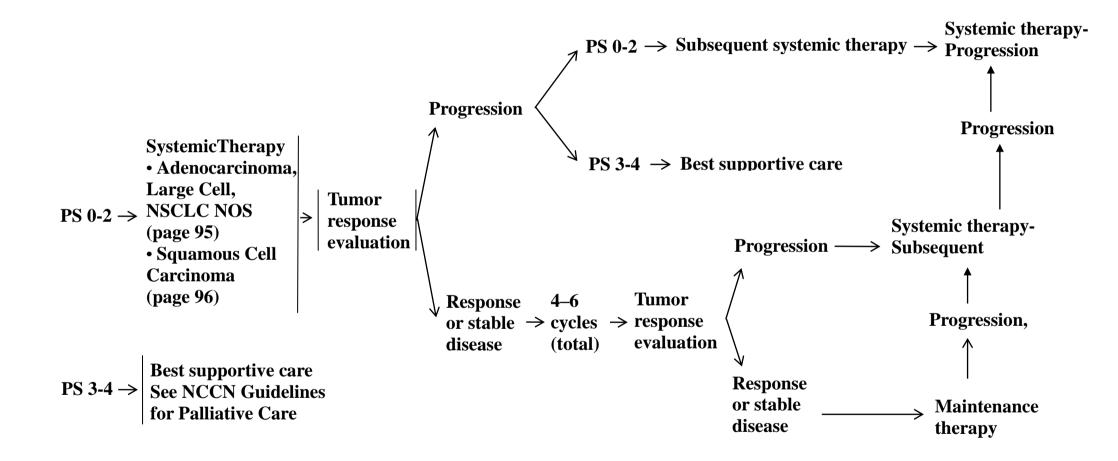


註:實際情況需與醫師討論(SDM)

註\*Atezolizumab、Nivolumab、Ipilimumab 未有健保給付

#### | PD-L1 < 1% AND NEGATIVE FOR ACTIONABLE | MOLECULAR MARKERS INITIAL SYSTEMIC THERAPY

#### SUBSEQUENT THERAPY



註:實際情況需與醫師討論(SDM)

### 三、化療原則

### Neoadjuvant chemotherapy regimens

建議處方:

B A

Platinum + Gemcitabine

Platinum + Paclitaxel

Platinum + Docetaxel

Platinum + Vinorelbine

Platinum + Etoposide

Platinum + Pemetrexed

大化療: Platinum(B) + another C/T agent(A)

小化療:single agent(A)

Platinum (鉑):

Cisplatin (dose adjusted by Ccr)

Carboplatin (dose calculated by Ccr and AUC)

實際施打劑量若因病患肝腎功能有調整,需註記於病歷中

### Adjuvant chemotherapy

建議處方:

B A

Platinum + Gemcitabine

Platinum + Paclitaxel

Platinum + Docetaxel

Platinum + Vinorelbine

Platinum + Etoposide

Platinum + Pemetrexed

Adenocarcinoma (T2 且腫瘤≥3cm) 可考慮使用 Ufur

大化療: Platinum(B) + another C/T agent(A)

小化療: single agent(A)

Platinum (鉑):

Cisplatin (dose adjusted by Ccr)

Carboplatin (dose calculated by Ccr and AUC)

#### Platinum-base doublet

- <u>Cisplatin</u> 50-75mg/m<sup>2</sup> (adjusted by Ccr) or <u>Carboplatin</u> AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable) Day 1 or 15 and **Gemcitabine** 1000 mg/m<sup>2</sup> Day 1, 8, 15 IV
- <u>Cisplatin</u> 50-75mg/m<sup>2</sup> (adjusted by Ccr) or <u>Carboplatin</u> AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable) Day 1 or 15 and <u>Paclitaxel</u> 60-80 mg/m<sup>2</sup> Day 1, 8, 15 or <u>Paclitaxel</u> 160-225mg/m2 D1 IV
- <u>Cisplatin</u> 50-75mg/m<sup>2</sup> (adjusted by Ccr) or <u>Carboplatin</u> AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable ) IV Day 1 or 15 and <u>Etoposide</u> 100mg/m<sup>2</sup> IV Day 1-3 or Day 1, 8, 15(Three weekly cycle)

#### **Platinum-base doublet**

- <u>Cisplatin</u> 50-75mg/m<sup>2</sup> (adjusted by Ccr) or <u>Carboplatin</u> AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable) Day 1 or 8 and <u>Gemcitabine</u> 1000-1250 mg/m<sup>2</sup> Day 1, 8 IV
- <u>Cisplatin</u> 50-75mg/m<sup>2</sup> (adjusted by Ccr) or <u>Carboplatin</u> AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable ) Day 1 or 8,(15) and <u>Vinorelbine</u> 20-30mg/m<sup>2</sup> Day 1, 8, (15) IV
- <u>Cisplatin</u> 50-75mg/m<sup>2</sup> (adjusted by Ccr) or <u>Carboplatin</u> AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable ) Day 1 or 8,(15) and <u>Vinorelbine</u> 60-80mg/m<sup>2</sup> Day 1, 8, (15) PO
- <u>Cisplatin</u> 50-75mg/m<sup>2</sup> (adjusted by Ccr) or <u>Carboplatin</u> AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable) Day 1 or 8 and <u>Docetaxel</u> 25-35mg/m<sup>2</sup> Day 1,8 or <u>Docetaxel</u> 60mg/m<sup>2</sup> Day 1 IV
- <u>Cisplatin</u> 50-75mg/m<sup>2</sup> (adjusted by Ccr) or <u>Carboplatin</u> AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable ) Day 1 and **Pemetrexed** (non-squamous)500mg/m<sup>2</sup> Day 1 IV every 21days
- <u>Ufur</u> 300mg~600mg PO only for Adenocarcinoma (**T2** 且腫瘤≥**3cm**)。
- 輔助化學治療藥物給予時,應依各藥物特性,配合病人狀況,例如:BSA、WBC及特定之血液檢查值等, 調整適當藥物劑量。

#### Systemic therapy regmins for neoadjuvant and adjuvant therapy

Neoadjuvant systemic therapy for patients not candidates for immune checkpoint Inhibitors; Adjuvant systemic therapy

#### Preferred (non-squamous)

• Cisplatin 75 mg/m2 day 1, pemetrexed 500 mg/m2 day 1 every 21 days for 4 cycles

#### Preferred (squamous)

- Cisplatin 75 mg/m2 day 1, gemcitabine 1250 mg/m2 days 1 and 8, every 21 days for 4 cycles
- Cisplatin 75 mg/m2 day 1, docetaxel 75 mg/m2 day 1 every 21 days for 4 cycles

#### Other Recommended

- Cisplatin 50 mg/m2 days 1 and 8; vinorelbine 25 mg/m2 days 1, 8, 15, and 22, every 28 days for 4 cycles
- Cisplatin 100 mg/m2 day 1, vinorelbine 30 mg/m2 days 1, 8, 15, and 22, every 28 days for 4 cycles
- Cisplatin 75–80 mg/m2 day 1, vinorelbine 25–30 mg/m2 days 1 and 8, every 21 days for 4 cycles
- Cisplatin 100 mg/m2 day 1, etoposide 100 mg/m2 days 1–3, every 28 days for 4 cycles

#### Useful in Certain Circumstances

- Chemotherapy Regimens for Patients with Comorbidities or Patients Not Able to Tolerate Cisplatin
- Carboplatin AUC 6 day 1, paclitaxel 200 mg/m2 day 1, every 21 days for 4 cycles
- Carboplatin AUC 5 day 1, gemcitabine 1000 mg/m2 days 1 and 8, every 21 days for 4 cycles
- Carboplatin AUC 5 day 1, pemetrexed 500 mg/m2 day 1 for non-squamous every 21 days for 4 cycles

All chemotherapy regimens listed above can be used for sequential chemotherapy/RT.

#### **Neoadjuvant Systemic Therapy for Patients Candidates for Immune Checkpoint Inhibitors**

• Nivolumab 360 mg and platinum-doublet chemotherapy every 3 weeks for 3 cycles

#### Platinum-doublet chemotherapy options include:

- ♦ Carboplatin AUC 5 or AUC 6 day 1, paclitaxel 175 mg/m² or 200 mg/m² day 1 (any histology)
- ♦ Cisplatin 75 mg/m² day 1, pemetrexed 500 mg/m² day 1 (non-squamous)
- ♦ Cisplatin 75 mg/m² day 1, gemcitabine 1000 mg/m² 1 and 8 and 15 or 1250 mg/m² days 1 and 8 (squamous histology)

\*Cisplatin 75 mg/m² day 1, paclitaxel 175 mg/m² or 200 mg/m² day 1 (any histology)

#### Systemic Therapy Following Previous Neoadjuvant or Adjuvant Systemic Therapy

- Test for PD-L1,EGFR mutations, ALK rearrangements (optional) (stages IB–IIIA, IIIB [T3,N2]). Principles of Molecular and Biomarker Analysis.
- Osimertinib 80 mg daily: stage IB–IIIA or stage IIIB (T3, N2) positive for exon 19 deletion, exon 21 L858R who received previous adjuvant chemotherapy or are ineligible to receive platinum-based chemotherapy.
- Atezolizumab 840 mg every 2 weeks, 1200 mg every 3 weeks, or 1680 mg every 4 weeks for up to 1 year : stage IIB–IIIA, stage IIIB (T3, N2), or high-risk stage IIA NSCLC with PD-L1 ≥1% and negative for EGFR exon 19 deletion or exon 21 L858R mutations or ALK rearrangements who received previous adjuvant chemotherapy and with no contraindications to immune checkpoint inhibitors.\*
- Pembrolizumab 200 mg every 3 weeks or 400 mg every 6 weeks for up to 1 year :stage IIB–IIIA, stage IIIB (T3, N2), or high-risk stage IIA NSCLC and negative for EGFR exon 19 deletion or exon 21 L858R mutations or ALK rearrangements who received previous adjuvant chemotherapy and with no contraindications to immune checkpoint inhibitors.\*,14 The benefit for patients with PD-L1

#### **Concurrent Chemoradiation Regimens**

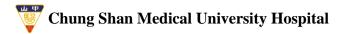
#### Preferred (nonsquamous)

- Carboplatin AUC 5 on day 1, pemetrexed 500 mg/m2 on day 1 every 21 days for 4 cycles; concurrent thoracic RT
- Cisplatin 75 mg/m2 on day 1, pemetrexed 500 mg/m2 on day 1 every 21 days for 3 cycles; concurrent thoracic RT
- ± additional 4 cycles of pemetrexed 500 mg/m2
- Paclitaxel 45–50 mg/m2 weekly; carboplatin AUC 2, concurrent thoracic RT4,\*,†,‡ ± additional 2 cycles every 21 days of paclitaxel 200 mg/m2 and carboplatin AUC 6
- Cisplatin 50 mg/m2 on days 1, 8, 29, and 36; etoposide 50 mg/m2 days 1–5 and 29–33; concurrent thoracic RT

#### Preferred (squamous)

- Paclitaxel 45–50 mg/m2 weekly; carboplatin AUC 2, concurrent thoracic RT ± additional 2 cycles every 21 days of paclitaxel 200 mg/m2 and carboplatin AUC 6
- Cisplatin 50 mg/m2 on days 1, 8, 29, and 36; etoposide 50 mg/m2 days 1–5 and 29–33; concurrent thoracic RT Consolidation Immunotherapy for Patients with Unresectable Stage II/III NSCLC, PS 0–1, and No Disease Progression After Definitive Concurrent Chemoradiation
- Durvalumab 10 mg/kg IV every 2 weeks or 1500 mg every 4 weeks for up to 12 months (patients with a body weight of ≥30 kg)(category 1 for stage III; category 2A for stage II)

註:實際情況需與醫師討論(SDM)



藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60 - 80 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 15	4weekly cycle
Gemcitabine	$1000 \text{ mg/m}^2$	N/S	30mins	Day 1, 8, 15	y y
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 15	4weekly cycle
Gemcitabine	$1000 \text{ mg/m}^2$	N/S	30mins	Day 1, 8, 15	
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60-75mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 8	3weekly cycle
Gemcitabine	1000-1250 mg/m <sup>2</sup>	N/S	30mins	Day 1, 8	3 3

註:實際情況需與醫師討論(SDM)

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60-75mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 8	3 weekly cycle	
Gemcitabine	1000-1250 mg/m <sup>2</sup>	N/S	30mins	Day 1, 8	Ç	
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or Cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 15	4 weekly cycle	
Paclitaxel	60 - 80 mg/m <sup>2</sup>	D5W or N/S	1-2hrs	Day 1, 8, 15		
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60 - 80 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 15	4 weekly cycle	
Paclitaxel	60-80 mg/m <sup>2</sup>	D5W or N/S	1-2hrs	Day 1, 8, 15		

註:實際情況需與醫師討論(SDM)

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60 - 75 mg/m2 (adjusted by Ccr)	N/S	2~12hrs	Day 1		
Paclitaxel	60 - 75 mg/m2 (adjusted by Ccr)	D5W or N/S	or N/S 1-2hrs Day 1		3 weekly cycle	
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Carboplatin	AUC: 3-6 (if Ccr <60ml/min or Cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 15	3 weekly cycle	
Paclitaxel	160-225 mg/m2	D5W or N/S	1-2hrs	Day 1		
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60 - 80mg/m2 (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 15	4 weekly cycle	
Etoposide	100 mg/m2	N/S or D5W	1hrs	Day 1-3 or Day1.8.15		

註:實際情況需與醫師討論(SDM)

Note: All recommendations are category 2A unless otherwise indicated.

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藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60 - 75mg/m2 (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 8	3 weekly cycle
Vinorelbine	60-80mg/m2	PO		Day 1, 8	
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3-6 (if Ccr <60ml/min or Cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 8	3 weekly cycle
Vinorelbine	60-80mg/m2	PO		Day 1, 8	
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60 - 75 mg/m2 (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 8	3 weekly cycle
Docetaxel	25-35mg/m2	D5W or N/S	0.5-1hr	Day 1, 8	<b>J</b> J
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3-6 (if Ccr <60ml/min or Cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 8	3 weekly cycle
Docetaxel	25-35mg/m2	D5W or N/S	0.5-1hr	Day 1, 8	

註:實際情況需與醫師討論(SDM)

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60 - 75 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1	3 weekly cycle	
Docetaxel	$60 \text{ mg/m}^2$	D5W or N/S	0.5-1hr	Day 1	, ,	
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Carboplatin	AUC: 3 – 6 (if Ccr <60ml/min or Cisplatin not suitable)	D5W or N/S	1hr	Day 1	3 weekly cycle	
Docetaxel	$60 \text{ mg/m}^2$	D5W or N/S	0.5-1hr	Day 1		
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60 - 75 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1		
Pemetrexed (non-squamous)	$500 \text{ mg/m}^2$	N/S	10-15mins	Day 1	3 weekly cycle	
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Carboplatin	AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day 1	3 weekly cycle	
Pemetrexed (non-squamous)	500 mg/m <sup>2</sup>	N/S	10-15mins	Day 1		

註:實際情況需與醫師討論(SDM)

Note: All recommendations are category 2A unless otherwise indicated.

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# **Adjuvant chemotherapy**

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60 - 80 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 15	4 11 1
Gemcitabine	$1000 \text{ mg/m}^2$	N/S	30mins	Day 1, 8, 15	4 weekly cycle
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60-75 mg/m2 (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 8	3 weekly cycle
Gemcitabine	1000-1250 mg/m2	N/S	30mins	Day 1, 8	
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or Cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 15	4 weekly cycle
Gemcitabine	1000-1250 mg/m2	N/S	30mins	Day 1, 8,15	<b>J J</b>
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or Cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 8	3 weekly cycle
Gemcitabine	1000-1250 mg/m2	N/S	30mins	Day 1, 8	

註:實際情況需與醫師討論(SDM)

Note: All recommendations are category 2A unless otherwise indicated.

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藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or Cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 15	4 weekly cycle	
Paclitaxel	60 - 80 mg/m2	D5W or N/S	1-2hrs	Day 1, 8, 15		
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60 - 80 mg/m2 (adjusted by Ccr)	N/S	2~12hrs	Day 1or15	4 weekly cycle	
Paclitaxel	60-80mg/m2	D5W or N/S	1-2hrs	Day 1,8,15		
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60 - 75 mg/m2 (adjusted by Ccr)	N/S	2~12hrs	Day 1	3 weekly cycle	
Paclitaxel	160-225 mg/m2	D5W or N/S	1-2hrs	Day 1		
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or Cisplatin not suitable)	D5W or N/S	1hr	Day 1	3 weekly cycle	
Paclitaxel	60 - 80 mg/m2	D5W or N/S	1-2hrs	Day 1		
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Carboplatin	AUC: 3-6 (if Ccr <60ml/min or Cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 15	4 weekly cycle	
Etoposide	100 mg/m2	N/S or D5W	1hrs	Day 1-3 or Day 1,8,15		

註:實際情況需與醫師討論(SDM)

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60-80mg/m2 (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 15	4 weekly cycle	
Etoposide	100 mg/m2	N/S or D5W	1hrs	Day 1-3 or Day1,8,15		
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60 - 75 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 8		
Vinorelbine	60-80mg/m <sup>2</sup>	РО		Day 1, 8	3 weekly cycle	
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Carboplatin	AUC: 3 - 6 (if Ccr < 60ml/min or Cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 8	3 weekly cycle	
Vinorelbine	60-80mg/m2	РО		Day 1, 8		
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 8	3 weekly cycle	
Docetaxel	25-35 mg/m2	D5W or N/S	0.5-1hr	Day 1, 8		

註:實際情況需與醫師討論(SDM)

Note: All recommendations are category 2A unless otherwise indicated.

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藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60 - 75 mg/m2 (adjusted by Ccr)	N/S	2~12hrs	Day 1or8	3 weekly cycle	
Docetaxel	25-35mg/m2	D5W or N/S	0.5-1hr	Day 1or8		
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60 - 75 mg/m2 (adjusted by Ccr)	N/S	2~12hrs	Day 1	3 weekly cycle	
Docetaxel	60mg/m2	D5W or N/S	0.5-1hr	Day 1		
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or Cisplatin not suitable)	D5W or N/S	1hr	Day 1	3 weekly cycle	
Docetaxel	60 mg/m2	D5W or N/S	0.5-1hr	Day 1		
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Cisplatin	60 - 75 mg/m2 (adjusted by Ccr)	N/S	2~12hrs	Day 1		
Pemetrexed (non- squamous)	500 mg/m2	N/S	15mins	Day 1	3 weekly cycle	
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期	
Carboplatin	AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day 1	2 11	
Pemetrexed (non- squamous)	500 mg/m2	N/S	15mins	Day 1	3 weekly cycle	

註:實際情況需與醫師討論(SDM)

Note: All recommendations are category 2A unless otherwise indicated.

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### **Concurrent Chemoradiation Regimen**

R/T (5000-7000 Gy; 25 - 35 次)

1. Paclitaxel 45-60 mg/m<sup>2</sup> IV over 1-2 hour weekly

Carboplatin AUC: 2 (if Ccr < 60ml/min or cisplatin not suitable ) weekly or

AUC: 3-6 (if Ccr < 60ml/min or cisplatin not suitable ) monsly IV

Concurrent thoracic radiotherapy

2. Paclitaxel 45-60 mg/m<sup>2</sup> IV over 1-2 hour weekly

Cisplatin 50-75mg/m<sup>2</sup> (adjusted by Ccr) IV

Concurrent thoracic radiotherapy

3. Etoposide 35-50mg/m<sup>2</sup> IV

Cisplatin 50mg/m<sup>2</sup> (adjusted by Ccr) or

Carboplatin AUC: 2 (if Ccr <60ml/min or cisplatin not suitable ) weekly or

**AUC: 3-6 (if Ccr < 60ml/min or cisplatin not suitable ) monsly IV** 

Concurrent thoracic radiotherapy

4. Pemetrexed (non-squamous)500mg/m<sup>2</sup> IV

Cisplatin 50-75mg/m<sup>2</sup> (adjusted by Ccr) IV

Concurrent thoracic radiotherapy

5. Pemetrexed (non-squamous) 500mg/m<sup>2</sup> IV

Carboplatin AUC: 3-6 (if Ccr < 60ml/min or cisplatin not suitable ) IV

Concurrent thoracic radiotherapy

註:實際情況需與醫師討論(SDM)

# **Concurrent chemoradiation regimen**

◆ Concurrent thoracic radiotherapy (5000 - 7000 Gy; 25 - 35 次)

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60 - 75 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 8, (15)	3 weekly cycle (4 weekly
Paclitaxel	45 - 60 mg/m <sup>2</sup>	D5W or N/S	1-2hrs	Day 1, 8, (15)	`

藥物名稱	藥物劑量	稀釋液	滴注時間	療法週期
Carboplatin	AUC: 2 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	weekly cycle
Paclitaxel	<b>45 - 60 mg/m</b> <sup>2</sup>	D5W or N/S	1-2hrs	weenly cycle

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 8, (15)	3 weekly cycle (4 weekly
Paclitaxel	45 - 60 mg/m <sup>2</sup>	D5W or N/S	1-2hrs	Day 1, 8, (15)	` •

註:實際情況需與醫師討論(SDM)

# **Concurrent chemoradiation regimen**

◆ Concurrent thoracic radiotherapy (5000 -7000Gy; 25 - 35 次)

藥物名稱	藥物劑量	稀釋液	滴注時間	期程
Cisplatin	60 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1, 8, 29, 36
Etoposide	35-50 mg/m <sup>2</sup>	N/S or D5W	1hrs	Day 1-3 or Day1-5,29- 31
藥物名稱	藥物劑量	稀釋液	滴注時間	期程
Carboplatin	AUC: 2 (if Ccr <60ml/min or cisplatin not suitable)	D5W	1hr	weekly
Etoposide	<b>35-50 mg/m</b> <sup>2</sup>	N/S or D5W	1hrs	Day 1-3 or Day1-5,29-31
藥物名稱	藥物劑量	稀釋液	滴注時間	期程
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	4 weekly cycle
Etoposide	35-50 mg/m <sup>2</sup>	N/S or D5W	1hrs	Day 1-3 or Day1-5,29-31

註:實際情況需與醫師討論(SDM)

Note: All recommendations are category 2A unless otherwise indicated.

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# Concurrent chemoradiation regimen

◆ Concurrent thoracic radiotherapy (5000 - 7000Gy; 25 - 35 次)

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60-75 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day1	
Pemetrexed (non-squamous)	<b>500 mg/m</b> <sup>2</sup>	N/S	15mins	Day1	3 weekly cycle

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day1	3 weekly cycle
Pemetrexed (non-squamous)	<b>500 mg/m</b> <sup>2</sup>	N/S	15mins	Day1	3 weekly cycle

註:實際情況需與醫師討論(SDM)

1. **Gemcitabine** 1000 mg/m<sup>2</sup> IV Day 1, 8, 15

<u>Cisplatin</u> 50-75 mg/m<sup>2</sup> (adjusted by Ccr) IV or

**Carboplatin** AUC: 3-6 (if Ccr <60 ml/min or cisplatin not suitable) IV Day 1 or 15

**2. Gemcitabine** 1000-1250 mg/m<sup>2</sup> IV Day 1, 8

<u>Cisplatin</u> 50-75 mg/m<sup>2</sup> (adjusted by Ccr) or

**Carboplatin** AUC: 3-6 (if Ccr <60 ml/min or cisplatin not suitable) IV Day 1 or 8

**3.** <u>Paclitaxel</u> 60-80 mg/m<sup>2</sup> IV Day 1, 8, 15 or <u>Paclitaxel</u> 160-225 mg/m<sup>2</sup> IV Day 1

<u>Cisplatin</u> 50-75 mg/m<sup>2</sup> (adjusted by Ccr) or

**Carboplatin** AUC: 3-6 (if Ccr <60 ml/min or cisplatin not suitable) IV Day 1 or 15

**4. <u>Docetaxel</u>** 25-35 mg/m<sup>2</sup> IV Day 1,8

<u>Cisplatin</u> 50-75 mg/m<sup>2</sup> (adjusted by Ccr) or

**Carboplatin** AUC: 3-6 (if Ccr <60 ml/min or cisplatin not suitable) IV Day 1 or 8

**5. Docetaxel** 60 mg/m<sup>2</sup> IV Day 1

Cisplatin 50-75 mg/m<sup>2</sup> (adjusted by Ccr) or

<u>Carboplatin</u> AUC: 3-6 (if Ccr <60 ml/min or cisplatin not suitable) IV Day 1 (Three week cycle)

**6.** <u>Vinorelbine</u> 60-80 mg/m<sup>2</sup> po Day 1, 8, 15

Cisplatin 50-75 mg/m<sup>2</sup> (adjusted by Ccr) or

**Carboplatin** AUC: 3-6 (if Ccr <60 ml/min or cisplatin not suitable ) IV Day 1 or 15

註:實際情況需與醫師討論(SDM)

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**7. Etoposide** 100 mg/m<sup>2</sup> IV Day 1-3 or Day 1, 8, 15

<u>Cisplatin</u> 50-75 mg/m<sup>2</sup> (adjusted by Ccr) or

**Carboplatin** AUC: 3-6 (if Ccr <60 ml/min or cisplatin not suitable ) IV Day 1 or 15

**8. Pemetrexed** (nonsquamous) 500 mg/m<sup>2</sup> IV on a 21-Day cycle.

<u>Cisplatin</u> 50-75 mg/m<sup>2</sup> (adjusted by Ccr) or

**Carboplatin** AUC: 3-6 (if Ccr <60 ml/min or cisplatin not suitable ) IV on a 21-Day cycle.

**9. TS-1** 80mg~120mg/夭(adjusted by BSA) PO BID Day 1~28,

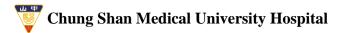
休息 14 天 or Day1~14, 休息7天。

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60-80 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 15	4 wooldy ovolo
Gemcitabine	<b>1000 mg/m</b> <sup>2</sup>	N/S	30mins	Day 1, 8, 15	4 weekly cycle

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 15	4 weekly cycle
Gemcitabine	<b>1000 mg/m</b> <sup>2</sup>	N/S	30mins	Day 1, 8, 15	4 weemy cycle

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60-75 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 8	2 wooldy avala
Gemcitabine	1000-1250 mg/m <sup>2</sup>	N/S	30mins	Day 1, 8	3 weekly cycle

註:實際情況需與醫師討論(SDM)



藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 8	3 weekly cycle
Gemcitabine	1000-1250 mg/m <sup>2</sup>	N/S	30mins	Day 1, 8	5 weekiy cycle

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60 - 80 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 15	4 wookly ovolo
Paclitaxel	<b>60 - 80 mg/m</b> <sup>2</sup>	D5W or N/S	1-2hrs	Day 1, 8, 15	4 weekly cycle

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 15	4 weekly cycle
Paclitaxel	<b>60 - 80 mg/m</b> <sup>2</sup>	D5W or N/S	1-2hrs	Day 1, 8, 15	4 weekly cycle

註:實際情況需與醫師討論(SDM)

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60-75 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1	3 wooldy ovelo
Paclitaxel	160-225 mg/m <sup>2</sup>	D5W or N/S	1-2hrs	Day 1	3 weekly cycle

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60 - 75 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 8	2 wooldy evolu
Docetaxel	<b>25-35mg/m</b> <sup>2</sup>	D5W or N/S	0.5-1hr	Day 1, 8	3 weekly cycle

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day 1	3 weekly cycle
Paclitaxel	<b>160-225 mg/m</b> <sup>2</sup>	D5W or N/S	1-2hrs	Day 1	5 weekly cycle

註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 66

Note: All recommendations are category 2A unless otherwise indicated.

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable)	D5W	1hr	Day 1 or 8	3 weekly cycle
Docetaxel	<b>25-35mg/m</b> <sup>2</sup>	D5W or N/S	0.5-1hr	Day 1, 8	5 weekij cjele

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60 - 80 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day 1 or 15	4 11 1
Etoposide	<b>100 mg/m</b> <sup>2</sup>	N/S or D5W	1hrs	Day 1-3 or Day 1,8,15	4 weekly cycle

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day 1 or 15	4 weekly cycle
Etoposide	<b>100 mg/m</b> <sup>2</sup>	N/S or D5W	1hrs	Day 1-3 or Day 1,8,15	4 weekly cycle

註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 67

Note: All recommendations are category 2A unless otherwise indicated.

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60-75 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day1	
Pemetrexed (nonsquamous)	<b>500 mg/m</b> <sup>2</sup>	N/S	15mins	Day1	3 weekly cycle

藥物名稱	藥物劑量	稀釋液 滴注時間 期程		療法週期	
Carboplatin	AUC: 3 - 6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day1	2 wooldy avolo
Pemetrexed (nonsquamous)	<b>500 mg/m</b> <sup>2</sup>	N/S	15mins	Day1	3 weekly cycle

藥物名稱	藥物劑量	稀釋液	頻次	期程	療法週期
TS-1	80 mg ~ 120 m/夭 (adjusted by BSA)	РО	BID	Day1~28, 休息 14 天 or Day1~14, 休息 7 天	6 weekly cycle or 3 weekly cycle

註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 68

Note: All recommendations are category 2A unless otherwise indicated.

MOLECULAR AND BIOMARKER-DIRECTED THERAPY FOR ADVANCED OR METASTATIC DISEASE						
EGFR Exon 19 Deletion or	ALK Rearrangement	BRAF V600E Mutation	ERBB2 (HER2) Mutation			
Exon 21 L858R:	• First-line therapy:	•First-line therapy : Dabrafenib/trametinib	• Subsequent therapy			
• First-line therapy : Afatinib	Alectinib Brigatinib	Encorafenib/binimetinib Dabrafenib	Fam-trastuzuab deruxtecan-nxki			
Erlotinib, Dacomitinib,	Ceritinib Crizotinib	Vemurafenib	Ado-trastuzumab emtansine			
Gefitinib, Osimertinib	Lorlatinib	•Subsequent therapy Dabrafenib/trametinib				
Erlotinib + ramucirumab7	• Subsequent therapy	Encorafenib/binimetinib				
Erlotinib + bevacizumabc	Alectinib Brigatinib					
(nonsquamous)8	Ceritinib Lorlatinib					
• Subsequent therapy	KRAS G12C Mutation	NTRK1/2/3 Gene Fusion				
Osimertinib	Subsequent therapy	•First-line/Subsequent therapy				
	Sotorasib, Adagrasi	Larotrectinib Entrectinib				
<b>EGFR S768I, L861Q, and/or</b>						
<u>G719X</u>	ROS1 Rearrangement	MET Exon 14 Skipping Mutation				
First-line therapy : Afatinib	•First-line therapy:	•First-line therapy/Subsequent therapy				
Erlotinib Dacomitinib	Ceritinib Crizotinib28	Capmatinib, Crizotinib, Tepotinib				
Gefitinib Osimertinib	Entrectinib					
• Subsequent therapy	•Subsequent therapy:	RET Rearrangeent				
Osimertinib	Lorlatinib30 Entrectinib					
		• First-line therapy/Subsequent therapy Selpercatinib、Pralsetinib、Cabozantinib				
		Seipercatinio Fraisetinio Caoozantinio				
EGFR Exon 20 Insertion						
Mutation College was at the second						
• Subsequent therapy						
Amivantamab-vmjw12						

註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 69

Note: All recommendations are category 2A unless otherwise indicated.

Anti-VEGF 血管新	Anti-VEGF 血管新生抑制劑						
藥物名稱	商品名	中文名	給付	藥物劑量	稀釋液	滴注時間	療法週期
Bevacizumab	Avastin	癌思婷	自費	7.5 mg~15 mg / kg	N/S	30~ 90mins	3 weekly cycle
Bevacizumab	MVASI	艾法施	自費	7.5 mg~15 mg / kg	N/S	30~ 90mins	3 weekly cycle
Ramucirumab	cyramza	欣銳擇	自費	10 mg / kg	N/S	60mins	2 weekly cycle or 3 weekly cycle

註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 70

Note: All recommendations are category 2A unless otherwise indicated.

# **Immunotherapy**

	_					-
學名	商品名	給付	藥物劑量	稀釋液	滴注時間	療法週期
Nivolumab	Opdivo	健保/自費	3 mg/kg / 360mg with Ipilimumab	D5W or N/S	60mins	2 weekly cycle
Durvalumab	Imfinzi	自費	10 mg/kg /1500mg(小細胞 肺癌)	D5W or N/S	60mins	2 weekly cycle
Atezolizumab	Tecentrip	自費	1200 mg	N/S	30~60mins	3 weekly cycle
Pembrolizumab	Keytruda	健保/自費	200 mg or 2mg/kg	D5W or N/S	60mins	3 weekly cycle
	Im	munoth	erapy com	bine thera	<b>py</b>	
藥物名稱	藥物	劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3- <60ml/min not sui	or cisplatin	D5W or N/S	30~60mins	Day1	3 weekly cycle
Pemetrexed	500 m	ng/m <sup>2</sup>	N/S	10-15 mins	Day1	3 weekly cycle
Pembrolizumab	200	mg	D5W or N/S	30~60mins	Day1	3 weekly cycle

註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 71

Note: All recommendations are category 2A unless otherwise indicated.



# 肺癌診療指引

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	50-75mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day1	3 weekly cycle
Pemetrexed	$500 \text{ mg/m}^2$	N/S	15mins	Day1	3 weekly cycle
Pembrolizumab	200 mg	D5W or N/S	60mins	Day1	3 weekly cycle
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60-75mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day1	3 weekly cycle
Pemetrexed	500 mg/m <sup>2</sup>	N/S	15mins	Day1	3 weekly cycle
Pembrolizumab	200 mg	D5W or N/S	60mins	Day1	3 weekly cycle
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day1	3 weekly cycle
Pemetrexed	500 mg/m <sup>2</sup>	N/S	15mins	Day1	3 weekly cycle
Atezolizumab	1200 mg	N/S	30~60mins	Day1	3 weekly cycle
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60-75 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day1	3 weekly cycle
Pemetrexed	500 mg/m <sup>2</sup>	N/S	15 mins	Day1	3 weekly cycle
Atezolizumab	1200 mg	N/S	30~60 mins	Day1	3 weekly cycle

註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM)72

Note: All recommendations are category 2A unless otherwise indicated.

# **Immunotherapy combine therapy**

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Cisplatin	60-75 mg/m <sup>2</sup> (adjusted by Ccr)	N/S	2~12hrs	Day1	3 weekly cycle
Pemetrexed	$500 \text{ mg/m}^2$	N/S	15 mins	Day1	3 weekly cycle
Atezolizumab	1200 mg	N/S	30~60 mins	Day1	3 weekly cycle
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day1	3 weekly cycle
Avastin	15 mg/Kg	N/S	90mins	Day 1	3 weekly cycle
Paclitaxel	160-225 mg/m <sup>2</sup>	D5W or N/S	1-2hrs	Day 1	3 weekly cycle
Atezolizumab	1200 mg	N/S	30~60mins	Day 1	3 weekly cycle
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day1	3 weekly cycle
Avastin	15 mg/Kg	N/S	90mins	Day 1	3 weekly cycle
Pemetrexed	500 mg/m2	N/S	15 mins	Day1	3 weekly cycle
Atezolizumab	1200 mg	N/S	30~60mins	Day1	3 weekly cycle

註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 73

Note: All recommendations are category 2A unless otherwise indicated.

# Immunotherapy combine therapy

藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1 hr	Day1	3 weekly cycle
Pemetrexed	$500 \text{ mg/m}^2$	N/S	15 mins	Day1	3 weekly cycle
Nivolumab	3 mg/kg	D5W or N/S	60 mins	Two weekly cycle	3 weekly cycle
Avastin	15 mg/Kg	N/S	90mins	Day 1	3 weekly cycle
藥物名稱	藥物劑量	稀釋液	滴注時間	期程	療法週期
Carboplatin	AUC:3-6 (if Ccr <60ml/min or cisplatin not suitable)	D5W or N/S	1hr	Day1	3 weekly cycle
Paclitaxel	160-225 mg/m <sup>2</sup>	D5W or N/S	1-2hrs	Day 1	3 weekly cycle
Atezolizumab	1200 mg	N/S	30~60mins	Day 1	3 weekly cycle

### Systemic therapy for advanced or metastatic disease

# ADENOCARCINOMA, LARGE CELL, NSCLC NOS (PS 0-1)

### (No contraindications to PD-1 or PD-L1 inhibitors )

#### **Preferred**

- ▶ Pembrolizumab / Carboplatin / Pemetrexed (category 1)
- ▶ Pembrolizumab / Cisplatin / Pemetrexed (category 1)

#### Other Recommended

- ▶ Atezolizumab / platinum/ Paclitaxel / Bevacizumabe (category 1)
- ▶ Atezolizumab / platinum / albumin-bound Paclitaxel
- ▶ Nivolumab / Ipilimumab
- Nivolumab/ipilimumab/pemetrexed/(carboplatin or cisplatin)

#### ADENOCARCINOMA, LARGE CELL, NSCLC NOS (PS 2)

#### **Preferred**

▶ Carboplatin / Pemetrexed

#### **Other Recommended**

- ▶ Carboplatin / albumin-bound Paclitaxel
- ▶ Carboplatin / Docetaxel
- ▶ Carboplatin / Etoposide
- ▶ Carboplatin / Gemcitabine
- ▶ Carboplatin / Paclitaxel

#### ADENOCARCINOMA, LARGE CELL, NSCLC NOS (PS 3-4)

▶ Best supportive care

### Contraindications to PD-1 or PD-L1 inhibitorsc Useful in Certain Circumstances

- ▶ Bevacizumabe/ Carboplatin / Paclitaxel (category 1)
- ▶ Bevacizumabe / Carboplatin / Pemetrexed
- ▶ Bevacizumabe / Cisplatin / Pemetrexed
- ▶ Carboplatin / albumin-bound Paclitaxel(category 1)
- ▶ Carboplatin / Docetaxel (category 1)
- ▶ Carboplatin / Etoposide (category 1)
- ▶ Carboplatin / Gemcitabine (category 1)
- ▶ Carboplatin / Paclitaxel (category 1)
- ▶ Carboplatin / Pemetrexed (category 1)
- ▶ Cisplatin / Docetaxel (category 1)
- ▶ Cisplatin / Etoposide (category 1)
- ▶ Cisplatin / Gemcitabine (category 1)
- ▶ Cisplatin / Pemetrexed (category 1)
- ▶ Gemcitabine / Docetaxel (category 1)
- ▶ Gemcitabine / Vinorelbine (category 1)
- ▶ Cisplatin/ Paclitaxel

#### SYSTEMIC THERAPY FOR ADVANCED OR METASTATIC DISEASE

#### SQUAMOUS CELL CARCINOMA (PS 0-1)

(No contraindications to PD-1 or PD-L1 inhibitors)

#### **Preferred**

- ▶ Pembrolizumab / Carboplatin / Paclitaxel (category 1)
- ▶ Pembrolizumab / Carboplatin / albumin-bound Paclitaxel (category 1)

#### Other recommended

- ▶ Nivolumab / Ipilimumab
- ▶ Nivolumab / Ipilimumab / Paclitaxel / Carboplatin (category 1)

### **SQUAMOUS CELL CARCINOMA (PS 2)**

#### **Preferred**

- ▶ Carboplatin / albumin-bound Paclitaxel
- ▶ Carboplatin / Gemcitabine
- ▶ Carboplatin / Paclitaxel

#### **Other Recommended**

- ▶ Carboplatin / Docetaxel
- ▶ Carboplatin / Etoposide

#### **Useful in Certain Circumstances**

- ▶ Albumin-bound paclitaxel
- **▶** Docetaxel
- **▶** Gemcitabine
- ▶ Paclitaxel

#### **SQUAMOUS CELL CARCINOMA (PS 3-4)**

▶ Best supportive care

#### Contraindications to PD-1 or PD-L1 inhibitors Useful in Certain Circumstances

- ▶ Carboplatin / albumin-bound Paclitaxel (category 1)
- ▶ Carboplatin / Docetaxel (category 1)
- ▶ Carboplatin / Gemcitabine (category 1)
- ▶ Carboplatin / Paclitaxel (category 1)
- ▶ Cisplatin / Docetaxel (category 1)
- ▶ Cisplatin / Etoposide (category 1)
- ▶ Cisplatin / Gemcitabine (category 1)
- ▶ Cisplatin / Paclitaxel (category 1)
- ▶ Gemcitabine / Docetaxel (category 1)
- ▶ Gemcitabine / Vinorelbine (category 1)

註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 76

### SYSTEMIC THERAPY FOR ADVANCED OR METASTATIC DISEASE – SUBSEQUENT

#### ADENOCARCINOMA, LARGE CELL, NSCLC NOS (PS 0-2)

**Preferred (no previous IO):** 

Systemic immune checkpoint inhibitors

- Nivolumab (category 1)
- Pembrolizumab (category 1)
- Atezolizumab (category 1)

#### Other Recommended (no previous IO):

- Docetaxel
- Pemetrexed
- Gemcitabine
- Ramucirumab/docetaxel
- Albumin-bound paclitaxel

#### **SQUAMOUS CELL CARCINOMA (PS 0-2)**

**Preferred (no previous IO):** 

Systemic immune checkpoint inhibitorse

- Nivolumab (category 1)
- Pembrolizumab (category 1)
- Atezolizumab (category 1)

#### Other Recommended (no previous IO):

- Docetaxel
- Gemcitabine
- Ramucirumab/docetaxel
- Albumin-bound paclitaxel

# ADENOCARCINOMA, LARGE CELL, NSCLC NOS, SQUAMOUS CELL CARCINOMA (PS 3-4): Best supportive care

#### SYSTEMIC THERAPY FOR ADVANCED OR METASTATIC DISEASE – PROGRESSION

#### ADENOCARCINOMA, LARGE CELL, NSCLC NOS

- PS 0–2: nivolumab, pembrolizumab, or atezolizumab, docetaxel (category 2B), pemetrexed (category 2B), gemcitabine (category 2B), ramucirumab/docetaxel (category 2B), or albuminbound paclitaxel (category 2B)
- PS 3–4: Best supportive care Options for further progression are best supportive care or clinical trial

#### SOUAMOUS CELL CARCINOMA

- PS 0–2: nivolumab, pembrolizumab, or atezolizumab, docetaxel (category 2B), gemcitabine (category 2B), ramucirumab/docetaxel (category 2B), or albumin-bound paclitaxel (category 2B)
- PS 3–4: Best supportive care Options for further progression are best supportive care or clinical trial.

註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 77

Note: All recommendations are category 2A unless otherwise indicated.

#### SYSTEMIC THERAPY FOR ADVANCED OR METASTATIC DISEASE – MAINTENANCE

#### **Maintenance Therapy**

- Continuation maintenance refers to the use of at least one of the agents given in first line, beyond 4–6 cycles, in the absence of disease progression. Switch maintenance refers to the initiation of a different agent, not included as part of the first-line regimen, in the absence of disease progression, after 4–6 cycles of initial therapy.
- Patients should receive maintenance therapy for 2 years if they received front-line immunotherapy.
- Patients should receive maintenance therapy until progression if they received second-line immunotherapy.

# ADENOCARCINOMA, LARGE CELL, NSCLC NOS (PS 0-2) Continuation maintenance

#### **Switch maintenance**

Pemetrexed

- Bevacizumab (category 1)
- Pemetrexed (category 1)
- Bevacizumab/pemetrexed
- Pembrolizumab/pemetrexed (category 1)
- Atezolizumab/bevacizumab (category 1)
- Nivolumab/ Ipilimumab
- Atezolizumabn

#### SOUAMOUS CELL CARCINOMA (PS 0-2) Continuation maintenance

- Pembrolizumab
- Nivolumab/ Ipilimumab
- Gemcitabine (category 2B)
- Durvalumab

#### ADENOCARCINOMA, LARGE CELL, NSCLC NOS, SQUAMOUS CELL CARCINOMA (PS 3-4)

Best supportive care

註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 78

Note: All recommendations are category 2A unless otherwise indicated.

# Small Cell Lung Cancer 治療原則

DIAGNOSIS	INITIAL EVALUATION	STAGE
Small cell lung cancer (SCLC) or combined SCLC/ non-small cell lung cancer (NSCLC) on biopsy or cytology of primary or metastatic site	<ul> <li>H&amp;Pb</li> <li>Pathology reviewc</li> <li>CBC</li> <li>Electrolytes, liver function tests (LFTs), BUN, creatinine</li> <li>Chest/abdomen/pelvis CT with contrast</li> <li>Brain MRI (preferred) or CT with contrast</li> <li>Consider PET/CT scan (skull base to mid-thigh), if limited stage is suspected or if needed to clarify stage</li> <li>Smoking cessation counseling and intervention.</li> <li>Molecular profiling (only for never smokers with extensive stage)</li> </ul>	Limited stage ———— See Additional Workup  Extensive stage ———— See Primary Treatment

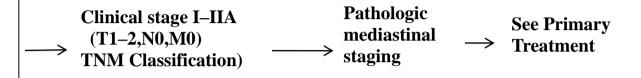


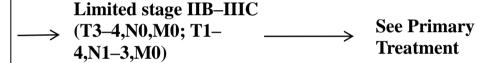
Limited stage	STAGE	TREATMENT
ADDITIONAL WORKUP	STAGE	IREATMENT

• If pleural effusion is present, thoracentesis is recommended; if thoracentesis inconclusive, consider thoracoscopy

**Chung Shan Medical University Hospital** 

- Pulmonary function tests (PFTs) during evaluation for surgery or definitive radiation therapy (RT)
- Bone imaging (radiographs or MRI) as appropriate if PET/CT equivocal (consider biopsy if bone imaging is equivocal)
- Unilateral marrow aspiration/biopsy in select patients





Consider arrange Bone

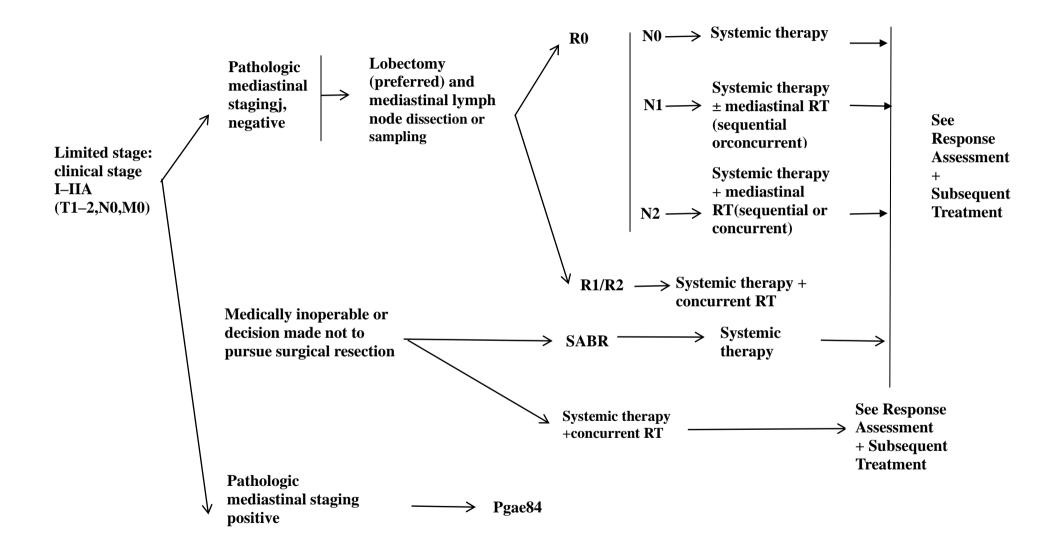
marrow biopsy,
thoracentesis, or bone
studies consistent with
malignancy

See Extensive-Stage Disease

**TESTING RESULTS** 

PRIMARY TREATMENT

ADJUVANT TREATMENT

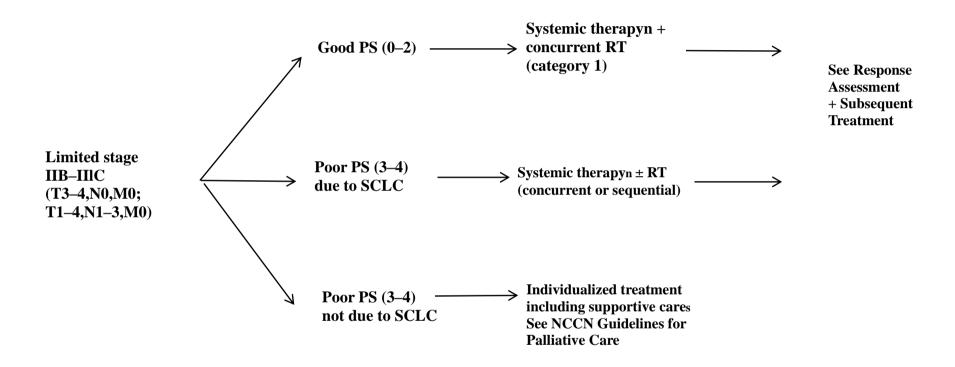


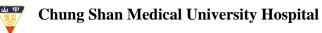
註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 81

Note: All recommendations are category 2A unless otherwise indicated.

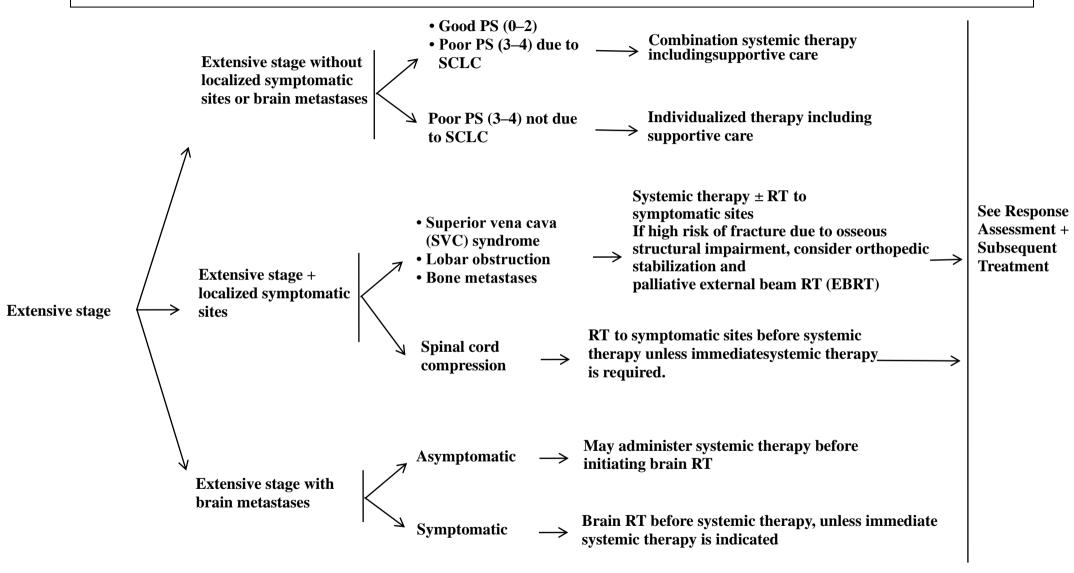


# PRIMARY TREATMENT





#### PRIMARY TREATMENT



註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 83

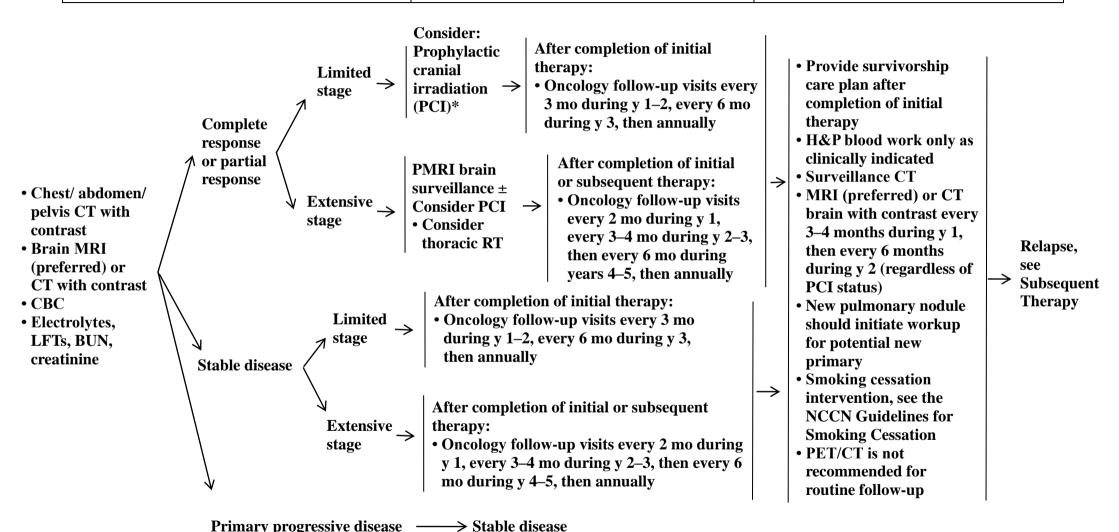
Note: All recommendations are category 2A unless otherwise indicated.



#### RESPONSE ASSESSMENT FOLLOWINGPRIMARY TREATMENT

#### **ADJUVANT RT**

#### **SURVEILLANCE**



註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 84 \*:70 歲以上需特別注意

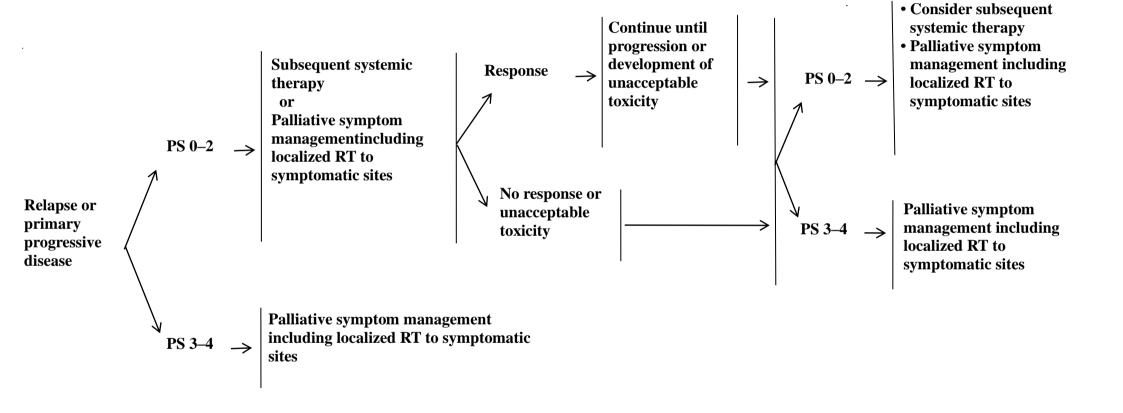
Note: All recommendations are category 2A unless otherwise indicated.



#### PROGRESSIVE DISEASE

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#### SUBSEQUENT THERAPY/PALLIATIVE THERAPY



註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 85

Note: All recommendations are category 2A unless otherwise indicated.

# 山中

# Small Cell Lung Cancer 治療原則-Limited stage (maximum of 4-6 cycles)

- ▶ **Etoposide** 50-100mg/m2 IV Day 1-3 or Day 1-5 or Day 1, 8, 15
- ▶ <u>Cisplatin</u> 50-75mg/m2 (adjusted by Ccr) or
- ▶ <u>Carboplatin</u> AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable ) IV Day 1 or 15

**♦**Good PS (ECOG PS $\leq 1$ ) →

concurrent chemoradiotherapy (5000-7000Gy 25-35 fractions)

**♦**Poor PS (ECOG PS $\geq 2$ ) →

chemotherapy only or sequential chemoradiotherapy (5000-7000Gy 25-35 fractions

註:實際劑量施打劑量若因肝腎功能不佳有調整,需註記於病歷上。

# 山中

# Small Cell Lung Cancer 治療原則-Extensive stage (maximum of 4-6 cycles)

- ▶ **Etoposide** 50-100mg/ M² IV Day 1-3 or Day 1, 8, 15
- ▶ <u>Cisplatin</u> 50-75mg/ M² (adjusted by Ccr) or <u>Carboplatin</u> AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable ) IV Day 1 or 15
- ▶ <u>Irinotecan</u> 50-60 mg/ M² IV Day 1, 8, 15
- ▶ <u>Cisplatin</u> 50-75mg/ M² (adjusted by Ccr) or <u>Carboplatin</u> AUC: 3-6 (if Ccr <60ml/min or cisplatin not suitable) IV Day 1 or 15
- ▶ Atezolizumab\*: 1200 mg Q3W

#### 註:

- 1. 給付期限,自處方用藥起算2年。
- 2. 需事審,每次以12週為期限,每12週需重新評估

註:實際劑量施打劑量若因肝腎功能不佳有調整,需註記於病歷上。

註:實際施打情況及劑量需與醫師討論.並紀錄於病歷上(SDM) 87

Note: All recommendations are category 2A unless otherwise indicated.



# Small Cell Lung Cancer 治療原則-二線治療 Topotecan

### Cycle 1.

1.5mg/m2 IV Day 1, 2, 3, (4,5)or Day 1, 8, 15

### **Subsequent cycle**

2.5mg/m2 IV Day 1, 2, 3, (4,5) or Day 1, 8, 15 (maximum 4mg)

- Lurbinectedin: 3.2mg/ M<sup>2</sup> Q3W
- Irinotecan: 60mg/ M<sup>2</sup> QW

其它藥物如 Ifosfamide, Paclitaxyl, Docetaxel, Gemcitabine,亦可建議使用。 若病人接受過 Platinum、Etoposide 及另一線化學治療後,可考慮使用免疫治療。



# 五、放射線治療指引

# **Non-Small Cell Lung Cancer**

Staging	Treatment	Adjuvant treatment
Operable	Operation (OP)	Postoperative RT if (1) Margin (+)
- cT1~3N0~1		(R1-2) or (2) <u>pN2</u>
Medically inoperable	1) Definitive radiotherapy (RT) or	Chemotherapy (C/T) <u>if high risk</u>
- cT1~2N0	2) Stereotactic ablative radiotherapy (SABR)	
Medically inoperable	1) Definitive chemoradiation (CRT)	Durvalumab (健保尚未給付)
- cT1-3N(+)	2) RT alone if not suitable for C/T	
Resectable	1) <b>OP</b>	Postoperative RT if (1) Margin (+) (R1-2) or
- cT3~4N0~1	2) Neoadjvant C/T OP	(2) <u>pN2</u>
Unresectable	1) Definitive concurrent chemoradiation (CCRT)	Durvalumab (健保尚未給付)
- cT3~4N0~1	2) RT alone if not suitable for C/T	
Superior Sulcus Tumor	1) Neoadjvant CCRT OP	Durvalumab (健保尚未給付) <u>if no OP</u>
- cT3~4N0~1	2) Definitive CCRT	
- cT1~3N2	1) Definitive CCRT	1) Durvalumab (健保尚未給付) <u>if no OP</u>
	2) Neoadjuvant CT OP	2) PORT <u>if not given</u>
	3) Neoadjuvant CCRT OP	<u> </u>
	4) RT alone if not suitable for C/T	
- cT1~3N2	1) Definitive CCRT	Durvalumab (健保尚未給付) <u>if no OP</u>
	2) RT alone if not suitable for C/T	
Stage IVA, IVB	1) Systemic therapy	
	2) Definitive RT to oligometastases	
	3) Palliative RT for symptoms	

註:實際情況需與醫師討論(SDM)

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Note: All recommendations are category 2A unless otherwise indicated.

Staging	Treatment	Adjuvant treatment
Limited stage	1.OP	1. Adjuvant C/T
(cT1~2N0)	2. SBRT	2. Mediastinal RT if pN1/N2
		3. Prophylactic cranial irradiation
		(PCI) for responder
Limited stage	<b>Definitive CCRT</b>	PCI for responder
(cT1~2N0)		
Limited stage	1. Definitive CCRT	PCI for responder
$(cT3\sim4N0, cT1\sim4N+)$	<b>2. Definitive SCRT</b> if poor PS	
Extensive stage	1. Systemic therapy	1.Brain MRI f/u for responder
	2. Palliative RT for symptoms	2.Thoracic RT or ConsiderPCI( optional) for responder
	4) Consolidative RT to primary site	es

#### **NSCLC RT dose**

- 1. **Definitive RT/ CCRT**: 60~70Gy at (1.8~2Gy/ fraction, 5 times per week)
- 2. **Neoadjuvant CCRT:** 45~ **54Gy**at (1.8~2Gy/ fraction, 5 times per week)
- 3. PORT for (margin (-) and pN2): 50~54Gy at (1.8~2Gy/ fraction, 5 times per week)
- 4. **PORT for (ENE or R1):** 54~60Gy at (1.8~2Gy/ fraction, 5 times per week)
- 5. **PORT for (R2):**  $60 \sim 70$ Gy at  $(1.8 \sim 2$ Gy/ fraction, 5 times per week)
- 6. 治療天數應由病況決定,合理範圍: 25~63 天。
- 7. **Palliative RT of metastases :** For patients with oligometastatic disease and good performance status consider higher doses (45–60 Gy) in 1.8–2 Gy daily fractions, or SBRT following principles for treatment of oligometastases

#### **SCLC RT dose**

- 1. **Limited stage Definitive CCRT**: 1) 60~70Gy at (1.8~2Gy/ fraction, 5 times per week), 2)**50**Gy/30fr at (1.5Gy/ fraction, BID)
- 2. Extensive stage: Consolidation thoracic RT w/ 30Gy/10frs ~ 60Gy/ 30fr. 依病人臨床狀況而定。
- 3. **PCI**: 25Gy/ 10fr at (2.5Gy/ fraction, 5 times per week)

#### **SBRT** dose

經胸腔外科醫師評估過後:(1) 無法接受手術。(2)手術風險高之病患(ex. Age≥75, 肺功能差)。

#### **Commonly Used Doses for SABR**

<b>Total Dose</b>	# Fractions	Example Indications
25–34 Gy	1	Peripheral, small (<2 cm) tumors, esp. >1 cm from chest wall
45–60 Gy	3	Peripheral tumors and >1 cm from chest wall
48–50 Gy	4	Central or peripheral tumors <4–5 cm, especially <1 cm from chest wall
50–55 Gy	5	Central or peripheral tumors, especially <1 cm from chest wall
60–70 Gy	8-10	Central tumors

OAR/Regimen	1 Fraction	3 Fractions	4 Fractions	5 Fractions
Spinal cord	14 Gv	18 Gy	26 Gy	30 Gy
Spinar coru	14 Gy	(6 Gy/fx)	(6.5  Gy/fx)	(6 Gy/fx)
Egophogus	15 / Cv	27 Gy	30 Gy	105% of PTV
Esophagus	15.4 Gy	(9 Gy/fx)	(7.5  Gy/fx)	Prescription^
Brachial	17.5 C	24 Gy	27.2 Gy	32 Gy
plexus	17.5 Gy	(8 Gy/fx)	(6.8  Gy/fx)	(6.4 Gy/fx)
Heart / naricandium	22 Cv	30 Gy	34 Gy	105% of PTV
Heart / pericardium	22 Gy	(10  Gy/fx)	(8.5  Gy/fx)	prescription^
Great vessels	27 Cv	NS	49 Gy	105% of PTV
Great vessels	37 Gy	INS	(12.25  Gy/fx)	prescription^
Trachea & proximal	20.2 C	30 Gy	34.8 Gy	105% of PTV
bronchi	20.2 Gy	(10 Gy/fx)	(8.7  Gy/fx)	prescription^
Rib	30 Gy	30 Gy	40 Gy	NS
	<i>ગ</i> ઉપ્ર	(10 Gy/fx)	(10  Gy/fx)	149
Skin	26 Gy	24 Gy	36 Gy	32 Gy

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		(8 Gy/fx)	(9 Gy/fx)	(6.4 Gy/fx)
Stomach	12.4 Gy	NS	27.2 Gy (6.8 Gy/fx)	NS

**Maximum Dose Constraints for SABR** 

Commonly Used Doses for Conventionally Fractionated and Palliative RT

Treatment Type	<b>Total Dose</b>	Fraction Size	<b>Treatment Duration</b>
<b>Definitive RT</b> with or without chemotherapy	60–70 Gy	2 Gy	6–7 weeks
Preoperative RT	45–54 Gy	1.8–2 Gy	5 weeks
Postoperative RT			
<b>▶</b> Negative margins	50–54 Gy	1.8–2 Gy	5–6 weeks
► Extracapsular nodal extension or microscopic	54–60 Gy	1.8–2 Gy	6 weeks
positive margins			
▶ Gross residual tumor	60–70 Gy	2 Gy	6–7 weeks
Palliative RT			
<b>▶</b> Obstructive disease (SVC syndrome or obstructive	30–45 Gy	3 Gy	2–3 weeks
pneumonia)			
<b>▶</b> Bone metastases with soft tissue mass	20–30 Gy	4–3 Gy	1–2 weeks
<b>▶</b> Bone metastases without soft tissue mass	8–30 Gy	8–3 Gy	1 day–2 weeks
▶ Brain metastases	CNS GLs*	CNS GLs*	CNS GLs*
<b>▶</b> Symptomatic chest disease in patients with poor PS	17 Gy	8.5 Gy	1–2 weeks
<b>▶</b> Any metastasis in patients with poor PS	8–20 Gy	8–4 Gy	1 day–1 week

Normal Tissue Dose-Volume Constraints for Conventionally Fractionated RT with Concurrent Chemotherapy

OAR	Constraints in 30–35 fractions
Spinal cord	Max ≤50 Gy
Lung	V20 ≤35%-40%; MLD ≤20 Gy
Heart	V50 ≤25%; Mean ≤20 Gy
Esophagus	Mean ≤34 Gy; Max ≤105% of prescription dose; V60 ≤17%; contralateral sparing is desirable
Brachial plexus	Median dose ≤69 Gy

註:實際情況需與醫師討論(SDM)

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# 六、溫度燒融治療的原則 Thermo-Ablation Therapy (TAT)

- 包括: 射頻消融治療 Radiofrequency Ablation(RFA)、微波消融治療 Microwave Ablation(MWA)、低温消融治療 Cryoablation (CA)
- 1.溫度消融治療(TAT)是局部治療的一種選擇;它可提供原發或轉移性肺部腫瘤的局部消融控制,其治療的併發症與副作用小,費用相對經濟,可適用於心肺功能不良及老年等不宜手術切除之局部控制治療。
- 2.温度消融治療(TAT)中 RFA 的有效消融病灶大小為 2 公分以下; 腫瘤大小 2-5 公分則以 MWA 或 CA 為宜。
- 3.對於早期(Stage 1-2) NSCLC 不適合開刀或是拒絕開刀的, TAT 可作為治療的選項(若適合開刀,仍以開刀作為第一治療選項)。
- 4.對於晚期(Stage 3-4) NSCLC, TAT 可作為局部控制的一個手段, 若病情需要, 可合併藥物及電療。
- 5.對於局部控制,TAT可合併放射治療或免疫治療,可以有加成療效。
- 6.對於 NSCLC 復發的病人, TAT 可作為局部控制的一個手段,對於小於 5 個的多發性肺部轉移腫瘤,可以重複多次 TAT 治療。
- 7.若預期的效果不好(如肋膜積水、縱膈腔腫瘤)則不建議使用 TAT 治療。

### 七、安寧緩和照護原則

- 1.預期存活率小於六個月
- 2.所有第四期病患皆需早期會診安寧緩和醫療照護。

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# 八、肺癌完治率定義

癌別	期別		完治定義
		0 期 1 期	1. 接受根治性手術為完治日 2. 接受 RFA 或 MWA 為完治日
		2 期	1. 接受根治性手術為完治日
	治	3A 期	1. C/T → OP 為完治日 2. OP → C/T 4 cycle 為完治日 3. OP → CCRT 為完治日
肺癌	療期	3B、3C 期	1. CCRT 為完治日 2.標靶藥物持續 3 個月為完治日 3. C/T 4-6 cycle 為完治日
		4 期	1. Palliative 口服或標靶藥物持續 3 個月為完治日 2. Palliative C/T 4 cycle 為完治日 3. 接受 RFA 或 MWA 為完治日 4. 若療程改變,換藥物治療時為完治日 5. 治療中轉安寧算完治日

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