高雄榮民總醫院

皮膚癌(SCC、 Keratoacanthoma)

診療原則

修訂日期:2022.04.19

癌委會公告日期:2022.07.18

Reference: NCCN Clinical Practice Guideline in OncologyTM ,Skin cancer, V.1.2019

SCC診療指引審視修訂會議討論日期

- 前次會議：2021/04/29

上一版	新版
NCCN Guidelines 2021年版	更換附件為:NCCN Guidelines 2022年版

鱗狀上皮細胞癌(SCC、Keratoacanthoma)

診斷	初步評估	分期	初始治療	療效評估	輔助治療	追蹤

臨床診斷之可疑病灶，經皮膚切片及病理組織學證實。

1、病史
2、理學檢查
(1)完整皮膚檢查
(2)淋巴結檢查
3、影像學檢查§
(1)CXR
(2)CT
(3)MRI

侷限型#
Local

擴散型

Low risk+

curettage & electrodesiccation

excision

margin positive
Moh's surgery
Wide excision
RT△

margin negative

RT△±C/T or Cryotherapy if not OP

High risk+

wide excision

margin positive
Moh's surgery
RT△

margin negative

Moh's surgery

margin positive
RT△

margin negative

RT△±C/T or Cryotherapy if not OP

Monthly*III
↓
Quarterly*III
↓
6 months*II
↓
Yearly for life

§： Image studies is indicated for extensive disease (deep structural involvement such as bone, deep soft tissue, perineural disease)

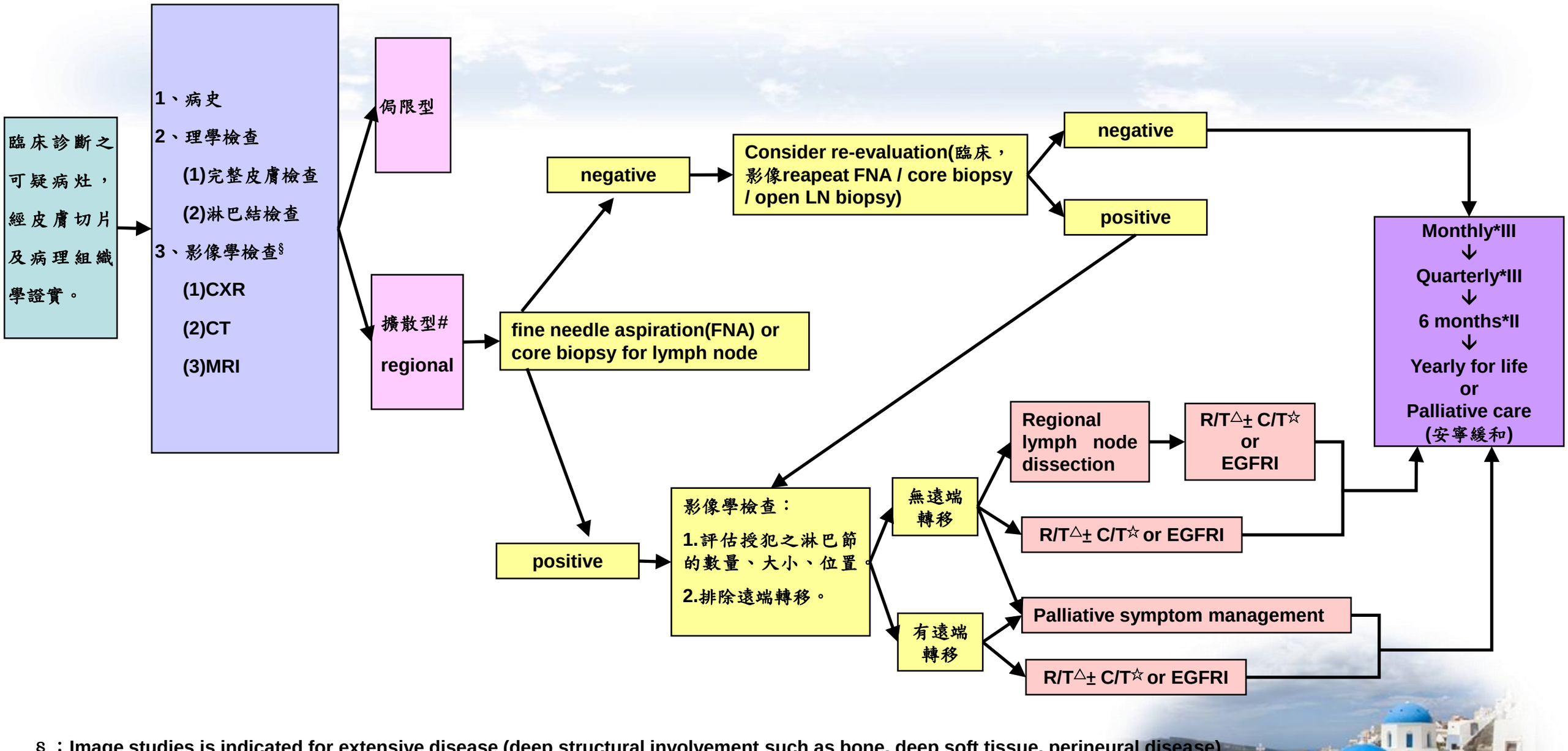
＋：附件一

△： RT主要針對手術不適用之情形, 附件二

#： T any, N0, M0, 附件三

鱗狀上皮細胞癌(SCC、Keratoacanthoma)

診斷	初步評估	分期	再評估(針對淋巴結)	初步治療	輔助治療	追蹤



§：Image studies is indicated for extensive disease (deep structural involvement such as bone, deep soft tissue, perineural disease) if perineural disease is suspected, MRI is preferred.

¥： Palliative symptom management, including salvage C/T

△： RT主要針對手術不適用之情形, 附件二

＃：Palpable regional lymph node(s) or abnormal lymph nodes identified by image studies. (擴散型的“初始皮膚病灶”治療同侷限型中high risk) T any, N1, M0 or M1 (附件三)

☆： chemotherapy regimen & EGFRI, 附件四

鱗狀上皮細胞癌(SCC、Keratoacanthoma)

復發

1、病史
2、理學檢查
(1)完整皮膚檢查
(2)淋巴結檢查
3、影像學檢查
(1)CXR
(2)CT
(3)MRI

侷限型 Local

- curettage & electrodesiccation
- wide excision
  - margin positive
    - Moh's surgery
    - RT
  - margin negative
- Moh's surgery
  - margin positive
    - RT
  - margin negative
- RT if not OP

擴散型 regional

- negative
  - Consider re-evaluation(臨床，影像reapeat FNA / core biopsy / open LN biopsy)
    - negative
    - positive
- fine needle aspiration(FNA) or core biopsy for lymph node
- positive

影像學檢查：
1.評估授犯之淋巴節的數量、大小、位置。
2.排除遠端轉移。

- 無遠端轉移
  - Regional lymph node dissection
    - R/T ± C/T or EGFRI
  - R/T ± C/T or EGFRI
  - Palliative symptom management
- 有遠端轉移
  - Palliative symptom management
  - R/T ± C/T or EGFRI

Monthly*III
↓
Quarterly
or
Palliative care
(安寧緩和)

皮膚癌多專科團隊

鱗狀上皮細胞癌(SCC、Keratoacanthoma)

癌症藥物停藥準則

- 根據CTCAE (Common Terminology Criteria for Adverse Events, Version 4.0 Published: May 28, 2009 【v4.03: June 14, 2010】)，出現Grade 3 ~ Grade 4 adverse event。
- 停藥至adverse event回復至Grade 1或Baseline時可再次用藥，但有些患者必須調整用藥劑量。
- 使用BRAF inhibitor時可能產生cutaneous SCC。此現象雖被CTCAE列為Grade 3 toxic effect, 但此現象不必停藥或調整劑量。
- 特定藥物治療下疾病仍持續進展，根據追蹤及評估顯示疾病對此特定藥物治療無效(考慮停止投藥並選擇其他治療方法)。
- 病患要求 (Hospice care或其他因素)。
- 病患死亡。

鱗狀上皮細胞癌(SCC)_ regional disease

附件一：

National Comprehensive Cancer Network®

NCCN Guidelines Version 1.2022
Squamous Cell Skin Cancer

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STRATIFICATION TO DETERMINE TREATMENT OPTIONS AND FOLLOW-UP FOR LOCAL CSCC BASED ON RISK FACTORS FOR LOCAL RECURRENCE, METASTASES, OR DEATH FROM DISEASE

Risk Group[1]	Low Risk	High Risk	Very High Risk
Treatment options	See SCC-2	See SCC-3	See SCC-3
H&P			
Location/size[2]	Trunk, extremities ≤2 cm	Trunk, extremities >2 cm – ≤4 cm	>4 cm (any location)
		Head, neck, hands, feet, pretibia, and anogenital (any size)[5]	
Borders	Well-defined	Poorly defined	
Primary vs. recurrent	Primary	Recurrent	
Immunosuppression	(-)	(+)	
Site of prior RT or chronic inflammatory process	(-)	(+)	
Rapidly growing tumor	(-)	(+)	
Neurologic symptoms	(-)	(+)	
Pathology (See SCC-A)			
Degree of differentiation	Well or moderately differentiated		Poor differentiation
Histologic features: Acantholytic (adenoid), adenosquamous (showing mucin production), or metaplastic (carcinosarcomatous) subtypes	(-)	(+)	Desmoplastic SCC
Depth[3,4]: Thickness or level of invasion	≤6 mm and no invasion beyond subcutaneous fat		>6 mm or invasion beyond subcutaneous fat
Perineural involvement	(-)	(+)	Tumor cells within the nerve sheath of a nerve lying deeper than the dermis or measuring ≥0.1 mm
Lymphatic or vascular involvement	(-)	(-)	(+)

See footnotes on SCC-B (2 of 2)

Note: All recommendations are category 2A unless otherwise indicated.
Clinical Trials: NCCN believes that the best management of any patient with cancer is in a clinical trial. Participation in clinical trials is especially encouraged.

皮膚癌
多專科團隊

鱗狀上皮細胞癌(SCC)_ regional disease

附件二

National Comprehensive Cancer Network®

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PRINCIPLES OF RADIATION THERAPY

General Principles

- Protracted fractionation is associated with improved cosmetic results and should be utilized for poorly vascularized or cartilaginous areas.
- For extensive perineural invasion, clinically evident perineural involvement, or involvement of named nerves (particularly in the head and neck region), consider including the course of the local nerves proximally.
- RT is contraindicated for genetic conditions predisposing to skin cancer (eg, basal cell nevus syndrome) and relatively contraindicated for patients with connective tissue diseases (eg, scleroderma).
- Given higher complication rates, re-irradiation should not be routinely utilized for recurrent disease within a prior radiation field.
- Isotope-based brachytherapy can be an effective treatment for certain sites of disease, particularly on the head and neck.
- There are insufficient long-term efficacy and safety data to support the routine use of electronic surface brachytherapy.

General Treatment Information

Primary Tumor	Examples of Dose Fractionation and Treatment Duration
Definitive RT	
Tumor diameter <2 cm	60–64 Gy over 6 to 7 weeks 50–55 Gy over 3 to 4 weeks 40 Gy over 2 weeks 30 Gy in 5 fractions over 2 to 3 weeks
Tumor diameter ≥2 cm, T3/T4, or those with invasion of bone or deep tissue	60–70 Gy over 6 to 7 weeks 45–55 Gy over 3 to 4 weeks
Postoperative Adjuvant RT	60–64 Gy over 6 to 7 weeks 50 Gy over 4 weeks
Regional Disease	
• Lymph node regions, after lymph node dissection	
▸ Negative margins, no ECE	50–60 Gy over 5 to 6 weeks
▸ Positive margins or ECE	60–66 Gy over 6 to 7 weeks
• Lymph node regions, without lymph node dissection	
▸ Clinically negative, at risk	50 Gy over 5 weeks
▸ Clinically positive	60–70 Gy over 6 to 7 weeks
• Clinically at-risk nerves	50–60 Gy over 5 to 6 weeks

Note: All recommendations are category 2A unless otherwise indicated.
Clinical Trials: NCCN believes that the best management of any patient with cancer is in a clinical trial. Participation in clinical trials is especially encouraged.

鱗狀上皮細胞癌(SCC)_ regional disease

附件三-1·

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American Joint Committee on Cancer (AJCC)
TNM Staging Classification for Cutaneous Carcinoma of the Head and Neck (8th ed., 2017)[1,2]

Table 1. **Definitions for T, N, M**

T	Primary Tumor
TX	Primary tumor cannot be assessed
Tis	Carcinoma *in situ*
T1	Tumor smaller than or equal to 2 cm in greatest dimension
T2	Tumor larger than 2 cm, but smaller than or equal to 4 cm in greatest dimension
T3	Tumor larger than 4 cm in maximum dimension or minor bone erosion or perineural invasion or deep invasion*
T4	Tumor with gross cortical bone/marrow, skull base invasion and/or skull base foramen invasion
T4a	Tumor with gross cortical bone/marrow invasion
T4b	Tumor with skull base invasion and/or skull base foramen involvement

*Deep invasion is defined as invasion beyond the subcutaneous fat or >6 mm (as measured from the granular layer of adjacent normal epidermis to the base of the tumor); perineural invasion for T3 classification is defined as tumor cells within the nerve sheath of a nerve lying deeper than the dermis or measuring 0.1 mm or larger in caliber, or presenting with clinical or radiographic involvement of named nerves without skull base invasion or transgression.

Clinical N (cN)

cN	Regional Lymph Nodes
NX	Regional lymph nodes cannot be assessed
N0	No regional lymph node metastasis
N1	Metastasis in a single ipsilateral lymph node, 3 cm or smaller in greatest dimension and ENE(−)
N2	Metastasis in a single ipsilateral node larger than 3 cm but not larger than 6 cm in greatest dimension and ENE(−); *or* metastases in multiple ipsilateral lymph nodes, none larger than 6 cm in greatest dimension and ENE(−); *or* in bilateral or contralateral lymph nodes, none larger than 6 cm in greatest dimension and ENE(−)
N2a	Metastasis in a single ipsilateral node larger than 3 cm but not larger than 6 cm in greatest dimension and ENE(−)
N2b	Metastases in multiple ipsilateral nodes, none larger than 6 cm in greatest dimension and ENE(−)
N2c	Metastases in bilateral or contralateral lymph nodes, none larger than 6 cm in greatest dimension and ENE(−)
N3	Metastasis in a lymph node larger than 6 cm in greatest dimension and ENE(−); *or* metastasis in any node(s) and clinically overt ENE [ENE(+)]
N3a	Metastasis in a lymph node larger than 6 cm in greatest dimension and ENE(−)
N3b	Metastasis in any node(s) and ENE (+)

Note: A designation of "U" or "L" may be used for any N category to indicate metastasis above the lower border of the cricoid (U) or below the lower border of the cricoid (L). Similarly, clinical and pathological extranodal extension (ENE) should be recorded as ENE(−) or ENE(+).

[1] These staging tables are for cutaneous squamous cell carcinoma, cutaneous carcinoma, basal cell carcinoma of the head and neck, and all other nonmelanoma skin carcinomas of the head and neck (except Merkel cell carcinoma). Anatomic site of external vermilion lip is included because it has a more similar embryologic origin to skin, and its etiology—which is often based on ultraviolet exposure—is more similar to other nonmelanoma skin cancers. The AJCC Staging Manual, Eighth Edition does not include staging for cutaneous carcinoma outside the head and neck.

[2] Used with the permission of the American College of Surgeons, Chicago, Illinois. The original source for this information is the AJCC Cancer Staging Manual, Eighth Edition (2017) published by Springer International Publishing

Continued

鱗狀上皮細胞癌(SCC)- regional disease

附件三-2:

National Comprehensive Cancer Network®

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Squamous Cell Skin Cancer

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American Joint Committee on Cancer (AJCC)
TNM Staging Classification for Cutaneous Carcinoma of the Head and Neck (8th ed., 2017)[1,2]

Pathological N (pN)

pN	**Regional Lymph Nodes**
NX	Regional lymph nodes cannot be assessed
N0	No regional lymph node metastasis
N1	Metastasis in a single ipsilateral lymph node, 3 cm or smaller in greatest dimension and ENE(−)
N2	Metastasis in a single ipsilateral lymph node, 3 cm or smaller in greatest dimension and ENE(+); *or* larger than 3 cm but not larger than 6 cm in greatest dimension and ENE(−); *or* metastases in multiple ipsilateral lymph nodes, none larger than 6 cm in greatest dimension and ENE(−); *or* in bilateral or contralateral lymph node(s), none larger than 6 cm in greatest dimension, ENE(−)
N2a	Metastasis in single ipsilateral node 3 cm or smaller in greatest dimension and ENE(+); *or* a single ipsilateral node larger than 3 cm but not larger than 6 cm in greatest dimension and ENE(−)
N2b	Metastases in multiple ipsilateral nodes, none larger than 6 cm in greatest dimension and ENE(−)
N2c	Metastases in bilateral or contralateral lymph node(s), none larger than 6 cm in greatest dimension and ENE(−)
N3	Metastasis in a lymph node larger than 6 cm in greatest dimension and ENE(−); *or* in a single ipsilateral node larger than 3 cm in greatest dimension and ENE(+); *or* multiple ipsilateral, contralateral, or bilateral nodes, any with ENE(+); *or* a single contralateral node of any size and ENE(+)
N3a	Metastasis in a lymph node larger than 6 cm in greatest dimension and ENE(−)
N3b	Metastasis in a single ipsilateral node larger than 3 cm in greatest dimension and ENE(+); *or* multiple ipsilateral, contralateral, or bilateral nodes, any with ENE(+); *or* a single contralateral node of any size and ENE(+)

Note: A designation of "U" or "L" may be used for any N category to indicate metastasis above the lower border of the cricoid (U) or below the lower border of the cricoid (L). Similarly, clinical and pathological extranodal extension (ENE) should be recorded as ENE(−) or ENE(+).

M	**Distant Metastasis**
M0	No distant metastasis
M1	Distant metastasis
G	**Histologic Grade**
GX	Grade cannot be assessed
G1	Well differentiated
G2	Moderately differentiated
G3	Poorly differentiated
G4	Undifferentiated

Table 2. **AJCC Prognostic Stage Groups**

	T	N	M
Stage 0	Tis	N0	M0
Stage I	T1	N0	M0
Stage II	T2	N0	M0
Stage III	T3	N0	M0
	T1	N1	M0
	T2	N1	M0
	T3	N1	M0
Stage IV	T1	N2	M0
	T2	N2	M0
	T3	N2	M0
	Any T	N3	M0
	T4	Any N	M0
	Any T	Any N	M1

[1] These staging tables are for cutaneous squamous cell carcinoma, cutaneous carcinoma, basal cell carcinoma of the head and neck, and all other nonmelanoma skin carcinomas of the head and neck (except Merkel cell carcinoma). Anatomic site of external vermilion lip is included because it has a more similar embryologic origin to skin, and its etiology—which is often based on ultraviolet exposure—is more similar to other nonmelanoma skin cancers. The AJCC Staging Manual, Eighth Edition does not include staging for cutaneous carcinoma outside the head and neck.

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鱗狀上皮細胞癌(SCC)- regional disease

附件四-1:chemotherapy regimen or metastasis

化學治療處方	
chemotherapy regimen	
published C/T regimens	**schedule**
Cisplatin, 100 mg/m2 IV D1	**Q 21-28 days x 4 cycles**
5-FU, 1 g/m2 IV D1–3	**Q 21-28 days x 4 cycles**

鱗狀上皮細胞癌(SCC)_ regional disease

附件四-2:chemotherapy regimen & EGFRI or metastasis

化學治療處方

chemotherapy regimen & EGFRI	
published C/T regimens	**schedule**
Cisplatin 100 mg/m2 IV D1	**Q 21 days * 6 cycles**
5-FU 1 g/m2 IV D1-4	**Q 21 days * 6 cycles**
*** Cetuximab 400 mg/m2 ; 250 mg/m2 IV**	**400 mg/m2 * Week 1；then 250 mg/m2 * QW**

* Cetuximab could be continued as long as the response or the stabilization persisted

鱗狀上皮細胞癌(SCC)_ regional disease

附件四-3:EGFRI or metastasis

化學治療處方	
EGFRI	
published C/T regimens	**schedule**
•Cetuximab, 400 mg/m2 IV Week 1, then 250 mg/m2 QW	**Till IV or unacceptable toxicity**

* Cetuximab could be continued as long as the response or the stabilization persisted

PRINCIPLES OF SYSTEMIC THERAPY

Local Disease (Including Multiple Primaries) Amenable to Curative Surgery
- Systemic therapy is not recommended.

Primary and Recurrent Locally Advanced Disease in Non-Surgical Candidates (See SCC-3)
- For patients who have residual disease and further surgery is not feasible, recommend RT, and multidisciplinary teams can consider concurrent systemic therapy in select cases (Table 1).
- For patients who have complicated cases of locally advanced disease in which curative surgery and curative RT are not feasible,[1] recommend multidisciplinary consultation to consider systemic therapy alone (Table 2).

New Regional Disease (See SCC-4 and SCC-5)
- For most cases of fully resected regional disease, adjuvant systemic therapy is not recommended, unless within a clinical trial.
- For patients with resected high-risk regional disease, consider RT ± systemic therapy (Table 1).
- For patients with unresectable, inoperable, or incompletely resected disease, multidisciplinary consultation is needed to consider:
  - RT ± systemic therapy (Table 1)
  - Systemic therapy alone if curative RT not feasible[1] (Table 2)

Regional Recurrence or Distant Metastatic Disease (See SCC-6)
- For regional recurrence or distant metastases, multidisciplinary team can consider systemic therapy alone (Table 2) or in combination with RT (Table 1).

Table 1: Systemic Therapy Options for Use with RT

Preferred Regimens	Other Recommended Regimens	Useful in Certain Circumstances
• Cisplatin[2] • Clinical trial[3,4]	• None	• EGFR inhibitors (eg, cetuximab)[2] • Cisplatin + 5-FU[2] • Carboplatin ± paclitaxel[2,5,6]

Table 2: Options for Systemic Therapy Alone

Preferred Regimens	Other Recommended Regimens	Useful in Certain Circumstances
• Cemiplimab-rwlc[3,4] (if curative RT or surgery is not feasible[1] for locally advanced, recurrent, or metastatic disease) • Pembrolizumab[3,4] (if curative RT or surgery is not feasible[1] for locally advanced, recurrent, or metastatic disease) • Clinical trial[3,4]	• If ineligible for or progressed on immune checkpoint inhibitors and clinical trials, consider: ▸ Carboplatin + paclitaxel	• If ineligible for or progressed on immune checkpoint inhibitors and clinical trials, consider: ▸ EGFR inhibitors (eg, cetuximab)[2] ▸ Capecitabine ▸ Cisplatin[2] ▸ Cisplatin + 5-FU[2] ▸ Carboplatin[2]

See Footnotes and References on SCC-F (2 of 2)

Note: All recommendations are category 2A unless otherwise indicated.
Clinical Trials: NCCN believes that the best management of any patient with cancer is in a clinical trial. Participation in clinical trials is especially encouraged.

Reference

1. NCCN Clinical Practice Guideline in Oncology, Basal and Squamous Cell Skin Cancers, Version 1.2022
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