LYMPH NODE METASTASIS IN ORAL CANCER

Thesis submitted in accordance with
the requirements of the University of Liverpool
for the degree of Doctor in Philosophy

bу

JULIA ANNE WOOLGAR

November, 1993.

VOLUME 1. - TEXT.

VOLUME 2. - TABLES, FIGURES, APPENDICES, and BIBLIOGRAPHY.

CONTENTS - VOLUME 2

The Tables, Figures and Appendices are arranged in the sequence in which they are cited in the text.

The Bibliography is arranged in the alphabetical sequence of the surname of the (first) author.

		<u>Page No.</u>
Section 1.	Tables.	1
Section 2.	Figures.	61
Section 3.	Appendices.	119
Section 4.	Bibliography.	140

Section 1.

TABLES.

- 1. Tables 4.1-4.15.
- 2. Tables 5.1-5.8.
- 3. Tables 6.1-6.33.
- 4. Tables 7.1-7.3.

Table 4.1

CLINICAL FEATURES, METASTATIC STATUS AND TYPE OF NECK DISSECTION IN UNILATERAL PROCEDURE PATIENTS.

ND No.	Sex	Age	e Site	Stage	Metasta	tic Status	Extent of Dissection
					Clinical	Histo- logical	
56	F	69	Buccal Mucosa	T1	Negative	Negative	Functional
2	F	67	Buccal Mucosa	T2	Negative	Positive	Radical
5	M	62	Buccal Mucosa	T2	Negative	Positive	SOH
99	\mathbf{F}	33	Buccal Mucosa	T2	Negative	Negative	Functional
55	M	49	Buccal Mucosa	Т4	Positive	Negative	Functional
125	F	43	Buccal Mucosa	Т4	Positive	Positive	Functional
86	F	64	Lower Alveolar Ridge		Negative	Negative	Functional
46	F	68	Lower Alveolar Ridge		Negative	Negative	Functional
65	F	81	Lower Alveolar Ridge		Negative	Negative	SOH
90	F	72	Lower Alveolar Ridge		Positive	Negative	Functional
98	M	50	Lower Alveolar Ridge		Positive	Negative	Radical
131	F	75	Lower Alveolar Ridge		Negative	Negative	Functional
128	M	79	Retromolar Trigone	T1	Negative	Negative	SOH
10	M	69	Retromolar Trigone	T3	Negative	Negative	Functional
9	M	61	Oral Tongue	T1	Positive	Positive	Radical
22	F	55	Oral Tongue	T1	Negative	Negative	SOH
47	M	46	Oral Tongue	T1	Negative	Negative	Functional
85 122	M	45 72	Oral Tongue	T1	Negative	Negative Positive	SOH
13	M F	72 67	Oral Tongue	T1 T2	Positive		Functional Radical
16	r M	50	Oral Tongue Oral Tongue	T2	Negative Positive	Negative Negative	Radical
18	M	46	Oral Tongue	T2	Positive	Negative Negative	Radical
31	M	65	Oral Tongue	T2	Positive	Positive	Radical
41	F	64	Oral Tongue	T2	Positive	Negative	Functional
120	M	66	Oral Tongue	T2	Negative	Positive	Functional
129	M	63	Oral Tongue	T2	Negative	Positive	Functional
144	M	81	Oral Tongue	T2	Negative	Positive	SOH
39	F	68	Oral Tongue	T3	Positive	Positive	Radical
52	M	67	Oral Tongue	T3	Negative	Positive	Functional
134	М	37	Oral Tongue	Т3	Positive	Negative	Functional
110	F	65	Oral Tongue	Т4	Positive	Positive	Radical
132	M	75	Floor of Mouth	T1	Positive	Negative	SOH
111	F	77	Floor of Mouth	T2	Negative	Positive	Functional
135	M	71	Floor of Mouth	T2	Positive	Positive	Functional
51	M	72	Floor of Mouth	Т4	Negative	Positive	SOH
96	\mathbf{F}	63	Floor of Mouth	Т4	Positive	Positive	Radical
109	M	53	Floor of Mouth	Т4	Negative	Positive	Functional
116	\mathbf{F}	61	Oropharynx	T1	Positive	Negative	SOH
124	M	49	Oropharynx	T2	Negative	Negative	Functional
108	M	51	Oropharynx	T4	Positive	Positive	Radical

a SOH = Supra-omohyoid.

Table 4.2 THE SITE AND STAGE OF THE PRIMARY TUMOUR IN THE SIMULTANEOUS UNILATERAL PROCEDURE PATIENTS a .

Git C Dui u			-		
Site of Primary	T1	Stage o T2	f Primary T3	Т4	All T Stages
Buccal Mucosa	1(0)	3(2)	0(0)	2(1)	6(3)
Lower Alveolar Ridge	1(0)	0(0)	1(0)	4(0)	6(0)
Retromolar Trigone	1(0)	0(0)	1(0)	0(0)	2(0)
Oral Tongue	5(2)	8(4)	3(2)	1(1)	17(9)
Floor of Mouth	1(0)	2(2)	0(0)	3(3)	6(5)
Oropharynx	1(0)	1(0)	0(0)	1(1)	3(1)
All Sites	10(2)	14(8)	5(2)	11(6)	40(18)

 $^{^{\}rm a}$ The number of patients with histologically diagnosed nodal metastasis is shown in brackets.

Table 4.3

CLINICAL FEATURES, METASTATIC STATUS AND TYPE OF NECK DISSECTION IN BILATERAL PROCEDURE PATIENTS.

ND No.	Sex	Ag	e Site	Stag	ge Metast	atic Statu	ıs Extent of Dissection ^a
					Clinical	Histo- logical	
104 105	M	74	Lower Alveolar Ridge	Т4	Negative Negative	Negative Negative	Functional Functional
63 64	M	64	Oral Tongue	Т2	Negative Positive	Negative Positive	SOH Functional
112 113	F	61	Oral Tongue	Т2	Positive Negative	Positive Negative	Functional SOH
152 153	M	60	Oral Tongue	Т2	Negative Negative	Positive Negative	Functional Functional
92 93	F	79	Oral Tongue	Т3	Negative Positive	Negative Negative	SOH Radical
83 84	M	51	Floor of Mouth	Т1	Negative Positive	Negative Positive	SOH Functional
106 107	F	63	Floor of Mouth	T1	Negative Negative	Positive Positive	SOH SOH
117 118	М	43	Floor of Mouth	T 1	Negative Positive	Negative Positive	SOH SOH Functional
138	F	57	Floor of Mouth	T1	Negative	Negative	SOH SOH
139 154	F	58	Floor of Mouth	т1	Negative Negative	Negative Negative	SOH
155 42	M	63	Floor of Mouth	Т2	Negative Positive	Negative Negative	SOH Functional
43 75	М	64	Floor of Mouth	Т2	Negative Negative	Negative Negative	SOH Functional
76 114	F	63	Floor of Mouth	Т2	Negative Positive	Negative Positive	Functional Radical
115 136	М	56	Floor of Mouth	Т2	Negative Negative	Positive Positive	SOH Functional
137 58	M	65	Floor of Mouth	Т4	Negative Positive	Negative Negative	SOH Functional
59 72	M	46	Floor of Mouth	Т4	Positive Positive	Negative Positive	Functional Radical
73 94	M	68	Floor of Mouth	Т4	Negative Positive	Negative Positive	Functional Radical
95 141	M	63	Oropharynx	Т1	Negative Positive	Negative Positive	SOH Radical
142 158	M	43	Oropharynx	Т3	Positive Positive	Positive Positive	Functional Radical
159 156	M	50	Oropharynx	Т4	Negative Negative	Negative Negative	Functional SOH
157					Positive	Positive	Functional

a SOH = Supra-omohyoid.

Table 4.4 THE SITE AND STAGE OF THE PRIMARY TUMOUR IN THE SIMULTANEOUS BILATERAL PROCEDURE PATIENTS $^{\mathbf{a}}$.

Site of Primary		Stage of 3	Primarv		
	T1	T2	T3	Т4	All T Stages
Lower Alveolar Ridge	0(0)	0(0)	0(0)	1(0)	1(0)
Oral Tongue	0(0)	3(3)	1(0)	0(0)	4(3)
Floor of Mouth	5(3)	4(2)	0(0)	3(2)	12(7)
Oropharynx	1(1)	0(0)	1(1)	1(1)	3(3)
All Sites	6(4)	7(5)	2(1)	5(3)	20(13)

^a The number of patients with histologically diagnosed nodal metastasis is shown in brackets.

Table 4.5

THE SITE AND STAGE OF THE PRIMARY TUMOUR IN THE SIMULTANEOUS PROCEDURE PATIENTS^a.

Site of Primary		Stage o	'Y		
	T1	т2	т3	T4	All T Stages
Buccal Mucosa	1(0)	3(2)	0(0)	2(1)	6(3)
Lower Alveolar Ridge	1(0)	0(0)	1(0)	5(0)	7(0)
Retromolar Trigone	1(0)	0(0)	1(0)	0(0)	2(0)
Oral Tongue	5(2)	11(7)	4(2)	1(1)	21(12)
Floor of Mouth	6(3)	6(4)	0(0)	6(5)	18(12)
Oropharynx	2(1)	1(0)	1(1)	2(2)	6(4)
All Sites	16(6)	21(13)	7(3)	16(9)	60(31)

^a The number of patients with histologically diagnosed nodal metastasis is shown in brackets.

Table 4.6

PATHOLOGICAL FINDINGS IN SERIES I PATIENTS.

	Unilateral Procedures (40)	Bilateral Procedures (40)	
With No Evidence of Metastasis	22(55%)	7(35%)	29(48%)
With Metastasis	18(45%)	13(65%)	31(52%)
-with discrete nodes only	14	9	23
-with fused nodal masses	4	4	8
Positive Patients with ECS ^a	11(61%)	11(85%)	22(71%)
-with macroscopic ECS	6	7	13
-with microscopic ECS	5	4	9

a ECS = Extracapsular spread.

Table 4.7

PATHOLOGICAL FINDINGS IN EIGHTY SIDES OF NECK DISSECTION (SERIES 1).

	Jnilateral Dissections	Bilateral Dissections	Simultaneous Dissections	
Sides of Neck Dissection	40	40	80	
With No Evidence of Metastasis	s 22(55%)	24(60%)	46(58%)	
With Metastasis	18(45%)	16(40%)	34(43%)	
-with discrete nodes only	14	11	25	
-with fused nodal masses	4	5	9	
-with ECS ^a	11	13	24	
-with macroscopic ECS	6	8	14	
-with microscopic ECS-with permeation of perinodal	5 L	4	9	
lymphatics only -percentage of positive	0	1	1	
dissections with ECS	61%	81%	71%	
Number of Metastatic Deposits	54	26	80	
-fused nodal masses	10	6	16	
-discrete positive nodes	44	20	64	
<pre>-with no ECS -with permeation of per:</pre>	24 i-	8	32(50%)	
nodal lymphatics only	1	2	3(5%)	
-with microscopic ECS	11	6	17(27%)	
-with macroscopic ECS	8	4	12(19%)	

a ECS = Extracapsular spread.

Table 4.8

SIZE AND DISTRIBUTION OF METASTATIC LYMPH NODES AND FUSED NODAL MASSES IN THIRTY-FOUR SIDES OF NECK DISSECTION.

Size (cm.)	Yield of	Metastati	c Deposi	ts ^a at Ana	atomic	al Level
of Node/Mass	I	II	III	IA	V	All Levels
Less than 0.5cm	. 1	0	0	0	2	3
0.5-0.9cm.	9	2	1	2	3	17
1.0-1.4cm.	6	4	5	2	2	19
1.5-1.9cm.	3	2	1	1	0	7
2.0-2.4cm.	4	7(2	0	2	0	13(2)
2.5-2.9cm.	1	5(3)	1	0	0	7(3)
3.0-3.4cm.	0	5(2)	1(1)	2(2)	0	8(5)
3.5-3.9cm.	0	0	0	1(1)	0	1(1)
More than 3.9cm	. 2(2)	3(3)	0	0	0	5(5)
All Sizes	26(2)	28(10)	9(1)	10(3)	7	80(16)

^a The total number of metastatic deposits (discrete and/or fused nodes) is shown. The number of fused nodal masses is shown in brackets. Fused nodal masses involving several anatomical levels are assigned to the first (lowest numerical) level of involvement.

Table 4.9

DISTRIBUTION OF CERVICAL METASTATIC DEPOSITS ACCORDING TO THEIR ANATOMICAL LEVEL.

Anatomical Level	Number of Discrete Positive Nodes	Number of Fused Nodal Masses ^a	Total Number of Metastatic Deposits	Percentage of All Metastatic Deposits
Level I	24	2	26	33%
Level II	18	10	28	35%
Level III	8	1	9	11%
Level IV	7	3	10	13%
Level V	7	0	7	9%
All Levels	64	16	80	100%

^a Fused nodal masses involving several anatomical levels are assigned to the first (lowest numerical) level of involvement.

Table 4.10 SIZE AND DISTRIBUTION OF LYMPH NODES IN EIGHTY SIDES OF NECK DISSECTION.

Nodal Size		Yiel	d of Nodes	at Anatom	ical Leve	ı1 ^a
	I	II	III	IV	V	All Levels
< 0.5cm.	28(1)	122(0)	57(0)	68(0)	277(2)	552 (3)
0.5-0.9cm.	173(9)	239(2)	158(1)	186(2)	226(3)	982(17)
1.0-1.4cm.	97(6)	126(4)	99(5)	82(2)	42(2)	446(19)
1.5-1.9cm.	34(3)	71(2)	40(1)	29(1)	8(0)	182 (7)
2.0-2.4cm.	13(4)	31(5)	11(0)	11(2)	0(0)	66(11)
2.5-2.9cm.	1(1)	13(2)	2(1)	2(0)	0(0)	18 (4)
3.0-3.4cm.	0(0)	10(3)	0(0)	0(0)	0(0)	10 (3)
3.5-3.9cm.	0(0)	1(0)	0(0)	0(0)	0(0)	1 (0)
> 3.9cm.	0(0)	1(0)	0(0)	0(0)	0(0)	1 (0)
All Sizes	346(24)	614(18)	367(8)	378(7)	553(7)	2258(64)

 $^{^{\}mathbf{a}}$ The number of discrete nodes containing metastatic carcinoma is shown in brackets.

Table 4.11 $\mbox{FREQUENCY OF HISTOLOGICAL METASTASIS IN RELATION TO LYMPH NODE SIZE}^{\bf a}.$

Size of Lymph Node (cm.)	Number of Nodes	Number of Metastatic Nodes	Relative %age of Metastatic Nodes	
All Sizes	2,258	64	2.8%	
= or > 0.5cm.	1,706	61	3.6%	
= or > 1.0 cm.	724	44	6.1%	
- or > 1.5cm.	278	25	9.0%	
= or > 2.0cm.	96	18	18.8%	
- or > 2.5cm.	30	7	23.3%	
= or > 3.0cm.	12	3	25.0%	
- or > 3.5cm.	2	0	0.0%	
= or > 4.0cm.	1	0	0.0%	

^a Only discrete lymph nodes are considered.

Lymph Node Size (cm.)	Extent Minimal	of Nodal F Partial	_	All Positive Discrete Nodes
Less than 0.5cm.	0	2	1	3
0.5-0.9cm.	2	13	2	17
1.0-1.4cm.	2	11	6	19
1.5-1.9cm.	2	3	2	7
2.0-2.4cm.	3	5	3	11
2.5-2.9cm.	0	1	3	4
3.0-3.4cm.	1	0	2	3
All Sizes	10	35	19	64

The extent of nodal replacement was subjectively graded as 'Minimal','Partial' or 'Total' as defined in the text (Chapter 4, Histological Methods).

Table 4.13

RELATIONSHIP BETWEEN ANATOMICAL LEVEL OF DISCRETE POSITIVE LYMPH NODES AND INCIDENCE AND EXTENT OF EXTRACAPSULAR SPREAD (ECS).

Extent of	Anatomical Level					
ECS	I	II	III	IV	V	All Levels
Positive, No ECS	10	12	4	3	3	32(50%)
Positive, With ECS -permeation of peri-	14	6	4	4	4	32(50%)
nodal lymphatics only	1	2	0	0	0	3 (5%)
-microscopic ECS	8	3	3	1	2	17(27%)
-macroscopic ECS	5	1	1	3	2	12(19%)
Total Number of						
Positive Nodes	24	18	8	7	7	64(100%)

Table 4.14

RELATIONSHIP BETWEEN SIZE OF METASTATIC LYMPH NODES AND INCIDENCE AND EXTENT OF EXTRACAPSULAR SPREAD (ECS).

Lymph Node		nt of Extracap	-		411 D:+:
Size (cm.)	Absent	Permeation of Perinodal Lymphatics Only	Micro- scopic ECS		All Positive Discrete Nodes
< 0.5cm.	2	0	2	0	4
0.5-0.9cm.	6	0	7	3	16
1.0-1.4cm.	11	1	3	4	19
1.5-1.9cm.	4	1	2	0	7
2.0-2.4cm.	5	1	2	3	11
2.5-2.9cm.	1	0	1	2	4
3.0-3.4cm.	3	0	0	0	3
All Sizes	32	3	17	12	64

Table 4.15

RELATIONSHIP BETWEEN INCIDENCE AND EXTENT OF EXTRACAPSULAR SPREAD (ECS) AND EXTENT OF NODAL REPLACEMENT.

Incidence/		Nodal Replace	ement a	
Extent of ECS	Minimal	Partial	Total	All Positive Discrete Nodes
ECS, Absent	9	18	5	32
Permeation of Perin Lymphatics Only	odal 1	2	0	3
ECS, Microscopic	0	14	3	17
ECS, Macroscopic	0	1	11	12
All Positive Discrete Nodes	10	35	19	64

The extent of nodal replacement was subjectively graded as 'Minimal', 'Partial', or 'Total' as defined in the text (Chapter 4, Histological Methods).

Table 5.1 ACCURACY OF CLINICAL ASSESSMENT OF METASTATIC STATUS IN RELATION TO SITE AND T STAGE OF PRIMARY TUMOUR.

T Site/ T Stage	Clin. N+ Hist. N+	Clin. N+ Hist. N-			Accuracy
Low-risk Sites ^a (n = 16)	1	3	10	2	11/16 (69%)
High-risk Sites ^b (n = 64)	19	10	23	12	42/64 (66%)
T1 (n = 22) T2 (n = 28) T3 (n = 9) T4 (n = 21)	5 2	2 4 2 5	12 10 4 7	2 9 1 2	18/22 (82%) 15/28 (54%) 6/9 (67%) 14/21 (67%)
All T Sites All T Stage (n = 80)	•	13	33	14	53/80 (66%)

 $[\]begin{array}{c} a \\ b \\ \end{array} \label{lower_alveolar_ridge/retromolar_trigone/buccal mucosa. \\ \\ b \\ \end{array}$

Table 5.2 CLINICAL AND PATHOLOGICAL FEATURES IN CLINICALLY FALSE-POSITIVE NECK DISSECTIONS.

Case	e T Site/ T Stage		urgical ocedure ^a I	Cli	
134 93	Oral Tongue Oral Tongue	T3 T3	Uni/F Bi/R	I	
_	FOM ^C FOM FOM	T2 T4 T1	Bi/F Bi/F Uni/SOH	I I	Chronic sialadenitis/fibrosis of SMG
90 98	Lower Ridge Lower Ridge	T4 T4	Uni/F Uni/R	I	Direct spread of primary tumour Direct spread of primary tumour
41	Oral Tongue	Т2	Uni/F	I	Reactive hyperplasia: four nodes (1.2-1.5cm.)
58§	FOM	Т4	Bi/F	Ι	Reactive hyperplasia: three nodes (2.0-2.2cm.)
55	Buccal Mucosa	Т4	Uni/F	II	Calcified node (4.5cm.). Reactive hyperplasia: five nodes (1.0-2.7cm.)
16	Oral Tongue	Т2	Uni/R	II	Reactive hyperplasia: twelve nodes (0.5-3.5cm.)
18	Oral Tongue	Т2	Uni/R	II	Reactive hyperplasia (granulomatous): thirteen nodes (0.5-2.4cm.)
116	Oropharynx	T1	Uni/SOH	II	Reactive hyperplasia: three nodes (2.0-2.2cm).

a Uni = Unilateral procedure, Bi = Bilateral procedure, R = Radical,
 F = Functional, SOH = Supra-omohyoid.
b SMG = Submandibular salivary gland.

c FOM = Floor of mouth.

[§] One patient had bilateral false-positive clinical assessments.

Table 5.3 CLINICAL AND PATHOLOGICAL FEATURES IN CLINICALLY FALSE-NEGATIVE NECK DISSECTIONS.

Case ND	T Site/ T Stage	Su Pro	rgical cedure ^a L	His evel	Size	ally Posi Degree o Replaceme	
2 5	Buccal Mucosa Buccal Mucosa		Uni/R Uni/SOH	I I I		Total Minimal Mimimal	In bony depression
152 129	Oral Tongue Oral Tongue	T2 T2	Bi/F Uni/F	I I II	0.3cm. 0.7cm. 1.7cm.	Partial	Microscopic ECS
144	Oral Tongue	T2	Uni/SOH	I II	0.5cm. 0.6cm.	Partial Minimal	Microscopic ECS
120	Oral Tongue	Т2	Uni/F	I II II	1.0cm. 1.2cm.	Partial Minimal Partial Partial	Microscopic ECS Emboli in perinodal
52	Oral Tongue	Т3	Uni/F	IV III III			lymphatics Microscopic ECS Microscopic ECS
107§ 106§	FOM ^b FOM	T1 T1	Bi/SOH Bi/SOH	I I I	0.7cm.	Partial Partial Partial	Microscopic ECS Microscopic ECS Microscopic ECS
115 136	FOM FOM	T2 T2	Bi/SOH Bi/F	Î II	0.8cm.	Minimal Partial	Emboli in perinodal lymphatics
111	FOM	Т2	Uni/F	II II	2.1cm.		Microscopic ECS
109 51	FOM FOM	T4 T4	Uni/F Uni/SOH	II II	1.6cm. 2.5cm.	Partial Total	Microscopic ECS

a Uni = Unilateral procedure, Bi = Bilateral procedure, R = Radical,
 F = Functional, SOH = Supra-omohyoid.
§ One patient had bilateral false-positive clinical assessments.
c FOM = Floor of mouth.

Table 5.4

ANATOMICAL DISTRIBUTION OF LYMPH NODES IN SEVENTY-ONE SIDES OF NECK DISSECTION GROUPED ACCORDING TO TYPE OF SURGICAL PROCEDURE.

Surgical Procedure	I	Nodal II	l Yield a	at Anatom IV	ical Leve V	II-IV	All Levels
RADICAL (1	n = 11)			_			
Mean	5.0	11.1	8.7	6.4	13.8	26.2	45.0
SD	2.0	4.3	3.7	3.4	5.5	9.4	11.1
Median	5.0	11.0	10.0	6.0	12.0	26.0	41.0
Range	2-8	4-17	3-15	1-14	7-23	13-43	31-68
Sum	55	122	96	70	152	288	495
FUNCTIONA	L(n = 3)	35)					
Mean	4.6	6.9	4.6	5.3	9.0	16.7	30.3
SD	1.7	2.8	2.5	2.9	5.5	5.8	9.2
Median	4.0	6.0	4.0	5.0	9.0	16.0	28.0
Range	3-11	3-12	1-12	1-13	3-32	7-31	18-54
Sum	162	241	160	184	315	585	1062
SUPRA-OMO	HYOID (r	n = 25					
Mean	4.0	5.8	3.1	3.4		12.4	16.4
SD	1.3	3.6	1.9	2.7		6.8	6.9
Median	4.0	4.5	2.0	2.0		14.0	14.0
Range	2-7	2-19	1-7	1-10		5-35	7-38
Sum	99	146	78	86		310	409
ALL DISSE	CTIONS ($(n = 71, \epsilon)$	except L	evel V; r	n = 46		
Mean	4.5	7.2	4.7	4.8	10.2	16.7	27.7
SD	1.7	3.7	3.1	3.1	5.8	8.1	13.1
Median	4.0	6.5	4.0	4.0	9.0	15.0	25
Range	2-11	2-19	1-15	1-14	3-32	5-43	7-68
Sum	316	509	334	340	467	1183	1966

Table 5.5 COMPARISON OF NODAL YIELD IN RADICAL, FUNCTIONAL AND SUPRA-OMOHYOID NECK DISSECTIONS $^{\mathbf{a}}$.

Type of Neck Dissection and Anatomical Level	t Value	d.f. ^b	Probability Value (P)
RADICAL versus FUNCTIONAL			
(n = 11) $(n = 35)$			
Level I	0.602	44	0.56
Level II	3.082 ^c	10	0.01*
Level III	3.444	10	0.006*
Level IV	1.071	44	0.29
Level V	2.550	44	0.01*
Levels II-IV	3.157 ^c	10	0.01*
All Levels	4.394	44	<0.0001*
RADICAL versus SUPRA-OMOHYC	OID		
(n = 11) $(n = 25)$			
Level I	1.863	34	0.07
Level II	3.822	34	0.0006*
Level III	4.713 ^c	10	0.0009*
Level IV	2.773	34	0.009*
Levels II-IV	4.972	34	<0.0001*
All Levels	7.924 ^c	10	<0.0001*
FUNCTIONAL versus SUPRA-OMO	OHYOID		
(n = 35) $(n = 25)$	5)		
Level I	1.614	58	0.11
Level II	1.277	58	0.20
Level III	2.432	58	0.02*
Level IV	2.490	58	0.01*
Levels II-IV	2.641	58	0.01*
All Levels	6.422	58	<0.0001*

a Groups of neck dissections are compared by the Two Sample t Test.
b Degrees of freedom.
c The t value is corrected for unequal variance.
* Significant at the 5% level.

Table 5.6 COMPARISON OF NODAL YIELD IN UNILATERAL AND BILATERAL RADICAL NECK DISSECTIONS $^{\mathfrak{a}}$.

Anate Leve	omical 1	Unilateral Procedures n = 8	Bilateral to Procedures n = 3	: Value	d.f. ^b	Probability Value
I	Mean (SD) Median Range Sum	5.3 (2.1) 5.5 2-8 42	4.3 (1.5) 4.0 3-6 13	0.675	9	0.52
II	Mean (SD) Median Range Sum	12.0 (4.5) 13.5 4-17 96	8.7 (2.5) 9.0 6-11 26	1.180	9	0.27
III	Mean (SD) Median Range Sum	9.6 (3.4) 10.0 5-15 77	6.3 (4.2) 5.0 3-11 19	1.351	9	0.21
IV	Mean (SD) Median Range Sum	7.4 (3.3) 6.5 3-14 59	3.7 (2.3) 5.0 1-5 11	1.766	9	0.11
V	Mean (SD) Median Range Sum	13.8 (6.0) 12.5 7-23 110	14.0 (5.3) 12.0 10-20 42	0.063	9	0.95
Leve	els II-IV Mean (SD) Median Range Sum	29.0 (8.8) 27.5 14-43 232	18.7 (7.4) 16.0 13-27 56	1.795	9	0.10
A11	Levels Mean (SD) Median Range Sum	48.0 (11.4) 46.0 35-68 384	37.0 (5.3) 39.0 31-41 111	1.570	9	0.15

 $^{^{\}mathbf{a}}$ Unilateral and bilateral neck dissections are compared by the Two Sample t Test.
b Degrees of freedom.

Table 5.7 COMPARISON OF NODAL YIELD IN UNILATERAL AND BILATERAL FUNCTIONAL NECK DISSECTIONS $^{\mathtt{a}}$.

Anate Leve	omical 1	Unilateral Procedures n = 19	Bilateral Procedures n = 16	t Value	d.f. ^b	Probability Value
I	Mean (SD) Median Range Sum	4.8 (1.9) 4.0 3-11 91	4.4 (1.5) 4.0 3-8 71	0.593	33	0.56
II	Mean (SD) Median Range Sum	7.9 (2.8) 9.0 3-12 150	5.7 (2.3) 5.0 3-12 91	2.547	33	0.02*
III	Mean (SD) Median Range Sum	5.9 (2.4) 6.0 2-12 112	3.0 (1.6) 2.5 1-6 48	4.087	33	0.0003*
IV	Mean (SD) Median Range Sum	6.3 (3.3) 5.0 2-13 120	4.0 (1.6) 4.0 1-6 64	2.724 ^c	15	0.02*
V	Mean (SD) Median Range Sum	8.6 (3.7) 9.0 3-16 163	9.5 (7.1) 8.0 3-32 152	0.470 ^c	15	0.65
Leve	els II-IV Mean (SD) Median Range Sum	20.1 (5.1) 20.0 11-31 382	12.7 (3.7) 13.0 7-22 203	4.862	33	<0.0001*
A11	Levels Mean (SD) Median Range Sum	33.5 (7.5) 33.0 21-47 636	26.6 (9.8) 24.5 18-54 426	2.337	33	0.02*

Unilateral and bilateral neck dissections are compared by the Two Sample t Test.
Degrees of freedom.
The t value is corrected for unequal variance.
* Significant at the 5% level.

Table 5.8 COMPARISON OF NODAL YIELD IN UNILATERAL AND BILATERAL SUPRA-OMOHYOID NECK DISSECTIONS $^{\mathtt{a}}$.

Anat Leve	omical 1	Unilateral Procedures n = 9	Bilateral Procedures n = 16	t Value	d.f. ^b	Probability Value
I	Mean (SD) Median Range Sum	3.8 (1.6) 3.0 2-7 34	4.1 (1.2) 4.0 2-7 65	0.503	23	0.63
II	Mean (SD) Median Range Sum	7.3 (5.0) 7.0 2-19 66	5.0 (2.3) 4.0 2-10 80	1.329 ^c	8	0.22
III	Mean (SD) Median Range Sum	3.7 (2.1) 3.0 1-7 33	2.8 (1.7) 2.0 1-7 45	1.096	23	0.28
IV	Mean (SD) Median Range Sum	4.4 (3.6) 4.0 1-10 40	2.9 (1.9) 2.0 1-9 46	1.207 ^c	8	0.26
Leve	els II-IV Mean (SD) Median Range Sum	15.4 (9.5) 15.0 5-35 139	10.7 (4.2) 10.0 5-19 171	1.432 ^c	8	0.19
All	Levels Mean (SD) Median Range Sum	19.2 (9.1) 18.0 7-38 173	14.8 (4.9) 14.0 9-24 236	1.363	8	0.21

 $^{^{\}mathrm{a}}$ Unilateral and bilateral neck dissections are compared by the Two Sample t Test.

b Degrees of freedom.

c The t value is corrected for unequal variance.

Table 6.1

THE HISTOLOGICAL MALIGNANCY GRADING SYSTEM AND POINT SCORING CRITERIA.

Histological Feature	1	Point Scor 2	ce 3	4
Degree of Keratin- isation	High, well-formed pearls	Moderate, attempts at pearl formation	Poor, single cells	None identified
Nuclear Polymor- phism	Slight	Moderate	Abundant	Extreme
Mitoses, per 10 HPF	0-15	16-35	36-55	>55
Pattern of Invasion	Pushing	Bands	Cords or islands, >15 cells	Single cells small cords or islands, <15 cells
Stage of Invasion	Borderline	Invasion into lamina propria	Invasion into submucosa	Invasion into muscle
Lympho- plasma- cytic Infiltrate	Continuous rim	Moderate, many large patches	Slight, either diffuse or a few small patches	None

Table 6.2 COMPARISON OF HISTOLOGICAL MALIGNANCY GRADING AND IDENTIFICATION OF PERINEURAL AND VASCULAR INVASION IN THIRTY PILOT STUDY CASES ASSESSED ON TWO SEPARATE OCCASIONS.

Pilot Test Case		K	N	P	Histo	logi M	cal	Feat P	ure ^a S		L	ı	N	Ъ	V	b
1	1	1	3	3	3	3	3	4	3	3	2	3	0	0	0	0
2	1	1	2	2	2	2	3	3	4	4	1	1	1	1	0	0
3	1	1	2	2	2	2	1	1	4	4	3	3	1	0	0	0
4	2	2	2	2	2	2	2	2	3	3	1	1	0	0	0	0
5	2	2	2	3	3	3	2	2	4	4	2	2	0	0	0	0
6	1	1	1	1	2	3	3	4	4	4	3	3	0	1	0	0
7	1	2	2	2	1	2	3	3	4	4	2	2	0	0	0	0
8	1	1	3	3	3	3	3	3	4	4	3	3	1	1	1	1
9	3	3	1	1	1	1	3	3	4	4	3	3	0	0	0	0
10	1	1	2	2	2	2	2	2	4	4	2	2	0	0	0	0
11	4	4	2	2	2	2	4	3	4	4	4	3	1	1	0	1
12	3	3	2	2	3	3	3	3	4	4	3	3	1	1	1	1
13	2	1	3	3	1	1	2	2	4	4	2	2	0	0	0	0
14	1	1	3	3	2	2	3	3	4	4	3	3	0	0	0	0
15	3	3	3	3	4	3	3	3	4	4	4	4	1	1	1	1
16	1	1	3	3	2	2	3	3	4	4	2	2	0	0	0	0
17	3	3	3	3	1	1	3	3	3	3	1	1	0	0	0	0
18	3	3	1	1	3	3	3	3	4	4	4	4	1	0	0	0
19	2	2	2	3	2	2	3	3	3	3	2	2	0	0	0	0
20	2	2	2	2	3	3	3	3	4	4	3	3	1	1	0	0
21	2	2	2	2	2	2	1	1	4	4	3	3	0	0	0	0
22	1	1	2	2	2	2	3	3	4	4	1	1	0	0	0	0
23	3	3	2	2	2	2	3	3	3	3	2	3	0	0	0	0
24	1	1	2	2	2	2	3	3	4	4	3	3	0	0	1	0
25	1	1	2	2	2	2	2	2	4	4	2	3	0	0	0	0
26	3	3	2	2	4	4	3	3	4	4	4	4	1	1	1	1
27	1	1	1	1	2	2	2	2	4	4	3	3	0	0	0	0
28	4	3	2	2	2	2	4	3	4	4	3	3	1	1	1	0
29	3	3	1	1	2	2	3	3	4	4	3	3	0	0	0	0
30	3	3	3	3	3	3	4	4	4	4	1	1	0	0	0	0

 $^{^{\}rm a}$ K = Degree of Keratinisation, NP = Nuclear Polymorphism, M = Frequency of Mitosis, P = Pattern of Invasion, S = Stage of Invasion, L = Lymphoplasmacytic Infiltrate, N = Perineural Invasion, V = Vascular Invasion.

b Not Identified = 0, Identified = 1.

Table 6.3

CONSISTENCY OF MALIGNANCY GRADING AND IDENTIFICATION OF PERINEURAL AND VASCULAR INVASION IN PILOT STUDY CASES.

Histological Feature	Consistency (%)		
Degree of Keratinisation	90%		
Nuclear Polymorphism	93%		
Frequency of Mitosis	90%		
Pattern of Invasion	87%		
tage of Invasion	100%		
ymphoplasmacytic Infiltrate	87%		
Perineural Invasion	90%		
Jascular Invasion	90%		

Table 6.4
CLINICAL FEATURES IN SERIES II PATIENTS.

Case Number	Sex	Age	T Site	T Stage
CASES WITH LYN	1PH NODE META	STASIS		
ND9	Male	61	Tongue	Т1
ND31	Male	65	Tongue	T2
ND39	Female	68	Tongue	T3
ND40	Female	62	Tongue	T1
ND51	Male	72	Floor of Mouth	T4
ND52	Male	67	Tongue	Т3
ND63/64	Male	64	Tongue	T2
ND72/73	Male	46	Floor of Mouth	T4
ND74/89	Male	51	Floor of Mouth	Tl
ND94	Male	68	Floor of Mouth	T4
ND96	Female	63	Floor of Mouth	T4
ND106/107	Female	61	Floor of Mouth	T1
ND109	Male	53	Floor of Mouth	T4
ND111	Female	77	Floor of Mouth	T2
ND112	Female	61	Tongue	T2
ND114/115	Female	62	Floor of Mouth	T2
ND114/113 ND118	Male	43	Floor of Mouth	T1
ND120	Male	66		T2
ND120 ND129	Male	63	Tongue Tongue	T2
ND129	Male	71	Floor of Mouth	T2
ND135	Male	56	Floor of Mouth	T2
ND136 ND144	Male Male	81		T2
ND144 ND145	Male	52	Tongue Floor of Mouth	T1
ND143 ND152	Male	60	Tongue	T2
ND132 ND170	Male Male	56	Floor of Mouth	T1
ND170 ND175	Male Male	42		T2
ND175 ND176	Female	72	Tongue Tongue	T2
ND170	remare	12	Totigue	12
CASES WITHOUT	LYMPH NODE : Female		Тот тис	Т2
ND13	remare Male	67 50	Tongue	T2
ND16		46	Tongue	T2
ND18	Male Female	55	Tongue	T1
ND22		64	Tongue	T2
ND41	Female	63	Tongue Floor of Mouth	T2
ND42/43	Male	46		T1
ND47	Male	46 47	Tongue	T1
ND48	Female		Floor of Mouth	T4
ND58/59	Male	65	Floor of Mouth	T2
ND75/76	Male	64	Floor of Mouth	T1
ND85	Male	45 70	Tongue	
ND92/93	Female	79 74	Tongue	T3
ND132	Male	74 27	Floor of Mouth	T1
ND134	Male	3 7	Tongue	T3
ND138/139	Female	57 50	Floor of Mouth	T1
ND154/155	Female	58	Floor of Mouth	T1
ND166/167	Male	76	Floor of Mouth	T1
ND169	Male	64	Floor of Mouth	T1

Table 6.5 EXTENT OF NODAL METASTASIS IN SERIES II PATIENTS.

Case No.	Tumour Site ^a	Tumour Stage	Laterality of Nodal Involvement	No. of Positive Nodes/No Masses	Level of Involvement dal	ecs ^b
ND9	Tongue	т1	Unilateral	Two	III, IV	Macro
ND31	Tongue Tongue	T2	Unilateral	One	III, IV I	No ECS
ND31	Tongue	T3	Unilateral	Four	II, III, IV	Macro
ND40	Tongue	T1	Unilateral	One	II, III, IV	Macro
ND40 ND51	FOM	T4	Unilateral	One	I	Micro
ND51		T3	Unilateral	Two		No ECS
ND63/64	Tongue Tongue	T2	Unilateral	Three	II, IV	Macro
ND72/73	FOM	T4	Unilateral	Two	I, II	Macro
ND72/73 ND74/89	FOM	T1	Bilateral	Three	II, IV	Macro
ND74/09 ND94	FOM	T4	Unilateral	One	I,II I	Macro
ND94 ND96	FOM	14 T4		One Two	_	Macro
		14 T1	Unilateral		I, II	
ND106/107 ND109	FOM	11 T4	Bilateral	Three	I	Micro No ECS
ND109 ND111	FOM	T2	Unilateral	One	II	
			Unilateral	Two	II	Micro
ND112	Tongue	T2 T2	Unilateral	Two	I	Macro
ND114/115			Bilateral	Five	I, II, III	Macro
ND118	FOM	T1	Unilateral	One	I	Micro
ND120	Tongue		Unilateral	Five	I, II, III	Micro
ND129	Tongue		Unilateral	0ne	II	No ECS
ND135	FOM	T2	Unilateral	Five	II, III, IV, V	
ND136	FOM	T2	Unilateral	0ne	II	No ECS
ND144	Tongue		Unilateral	Three	I, II, III	Micro
ND145	FOM	T1	Unilateral	One	III	Macro
ND152	Tongue		Unilateral	Two	I	Micro
ND170	FOM	T1	Unilateral	Nine	I, II, III, IV	
ND175	Tongue		Unilateral	Two	II, III	Micro
ND176	Tongue	Т2	Unilateral	Six	II, III, IV	Macro

a FOM = Floor of mouth.
b Extracapsular spread (ECS) is graded as 'No ECS', 'Macroscopic' or 'Microscopic' as defined in the text (Chapter 4, ' Gross Dissection and Sampling' and 'Histological Methods'.

Table 6.6

TOBACCO USAGE, ALCOHOL CONSUMPTION AND EPITHELIAL DYSPLASIA IN SERIES II PATIENTS.

Case Number	Tumour Site		Cigarettes, number per day	Alcohol units per day	, Epithelial Dysplasia
CASES WITH	LYMPH NODE METAS	TASIS			
ND9	Tongue	<u>T</u> 1	0	2	Not identified
ND31	Tongue	T2	20	2	Not identified
ND39	Tongue	Т3	15	2	Not identified
ND40	Tongue	T1	0	0	Not identified
ND51	Floor of Mouth	Т4	30	8	Identified
ND52	Tongue	Т3	20	4	Not Identified
ND63/64	Tongue	Т2	20	6	Identified
ND72/73	Floor of Mouth	Т4	25	4	Not identified
ND74/89	Floor of Mouth	T1	15	3	Not identified
ND94	Floor of Mouth	Т4	25	5	Not identified
ND96	Floor of Mouth	Т4	10	0	Not identified
ND106/107	Floor of Mouth	T1	0	0	Identified
ND109	Floor of Mouth	T4	20	18	Identified
ND111	Floor of Mouth	T2	5	0	Not identified
ND112	Tongue	T2	6	0	Not identified
ND114/115	Floor of Mouth	T2	10	Ō	Not identified
ND118	Floor of Mouth	T1	15	Ō	Identified
ND120	Tongue	T2	25	8	Identified
ND129	Tongue	T2	10	2	Not identified
ND135	Floor of Mouth	T2	12	6	Not identified
ND136	Floor of Mouth	T2	40	12	Identified
ND144	Tongue	T2	50	5	Identified
ND145	Floor of Mouth	T1	30	12	Identified
ND152	Tongue	T2	30	10	Identified
ND170	Floor of Mouth	T1	20	16	Identified
ND175	Tongue	T2	10	6	Identified
ND176	Tongue	T2	0	Ö	Identified
CASES WITH	OUT LYMPH NODE ME	TASTASIS	<u>.</u>		
ND13	Tongue	Т2	15	4	Not identified
ND16	Tongue	Т2	50	16	Identified
ND18	Tongue	Т2	30	20	Not identified
ND22	Tongue	T 1	0	1	Identified
ND41	Tongue	Т2	0	0	Not identified
ND42/43	Floor of Mouth	T2	25	20	Identified
ND47	Tongue	T1	30	28	Identified
ND48	Floor of Mouth	T1	25	15	Identified
ND58/59	Floor of Mouth	T 4	40	16	Identified
ND75/76	Floor of Mouth	Т2	20	8	Not identified
ND85	Tongue	т1	0	0	Not identified
ND92/93	Tongue	Т3	50	0	Not identified
ND132	Floor of Mouth	T1	25	1	Not identified
ND134	Tongue	Т3	40	12	Identified
ND138/139	Floor of Mouth	T1	25	4	Identified
ND154/155	Floor of Mouth	T1	20	6	Identified
ND166/167	Floor of Mouth	T1	30	2	Identified
ND169	Floor of Mouth	T1	15	8	Identified

Table 6.7

TUMOUR SURFACE-DIMENSION AND THICKNESS IN SERIES II PATIENTS.

Case Number	Site T	Stage	Surface Dimension, D (mm.)	Actual Thickness, T _A (mm.)	
CASES WITH	LYMPH NODE ME	TASTASIS	}		
ND9	Tongue	T1	24	18	18
ND31	Tongue	T2	20	9	11
ND39	Tongue	Т3	60	25	25
ND40	Tongue	T1	9	4	3
ND51	Floor of Mouth		15	10	10
ND52	Tongue	Т3	45	15	20
ND63/64	Tongue	T2	22	11	9
ND72/73	Floor of Mouth		40	17	15
ND74/89	Floor of Mouth		12	4	4
ND94	Floor of Mouth		30	15	11
ND96	Floor of Mouth		65	10	12
	Floor of Mouth		18	5	6
ND109	Floor of Mouth		28	10	10
ND111	Floor of Mouth		22	6	8
ND112	Tongue	T2	35	18	18
	Floor of Mouth		30	10	10
ND118	Floor of Mouth		8	3	3
ND120	Tongue	T2	30	15	15
ND129	Tongue	T2	30	21	21
ND135	Floor of Mouth		20	7	11
ND136	Floor of Mouth		28	6	12
ND144	Tongue	T2	35	17	17
ND145	Floor of Mouth		5	2	2
ND152	Tongue	T2	30	14	16
ND170	Floor of Mouth		16	2	2
ND175	Tongue	T2	35	15	17
ND176	Tongue	T2	40	14	16
CASES WIT	HOUT LYMPH NODE	E METAST	ASIS		
ND13	Tongue	T2	20	13	13
ND16	Tongue	T2	33	11	11
ND18	Tongue	T2	20	8	10
ND22	Tongue	T1	21	13	16
ND41	Tongue	T2	13	18	18
ND42/43	Floor of Mouth		8	6	6
ND47	Tongue	T1	17	2	1
ND48	Floor of Mouth		4	2	2
ND58/59	Floor of Mouth		43	4	2
ND75/76	Floor of Mouth		22	3	3
ND85	Tongue	T1	7	2	3 2
ND92/93	Tongue	T3	44	7	7
ND132	Floor of Mout		8	4	3
ND134	Tongue	T3	42	12	14
	Floor of Mout		5	2	2
	Floor of Mout		15	5	5
	Floor of Mout		5	5	5
11D T O O / T O /	THOUL OF HOUL			•	•

Table 6.8 TUMOUR SURFACE-DIMENSION AND THICKNESS IN SERIES II PATIENTS WITH AND WITHOUT METASTASIS.

Pathological Feature ^a	Cases With Metastasis ^b	Cases Without Metastasis ^b	t value	P value
Dimension, D	27.9 (14.3)	18.9 (13.3)	2.13	0.04*
Thickness, T _A ^c	11.2 (6.1)	6.7 (4.8)	2.65	0.008*
Thickness, T_R^c	11.9 (6.2)	6.7 (5.5)	2.88	0.005*

 $^{^{\}rm a}$ The mean measurement (mm.) and standard deviation are given. $^{\rm b}$ Cases with and without metastasis are compared by the Two Sample t

Test.

C 'Actual Tumour Thickness, T_A ', and 'Reconstructed Tumour Thickness, T_R ', are described in the text (Chapter 6, 'Histological Assessment').

Table 6.9

FREQUENCY OF EPITHELIAL DYSPLASIA IN RELATION TO TOBACCO AND ALCOHOL USAGE.

Tobacco and Alcohol Usage ^a	Epithelial Dysplasia Identified	Epithelial Dysplasia Not Identified
Both Heavy Smoker and Heavy Drinker	11	1
Either Heavy Smoker or Heavy Drinker	5	5
Moderate/Non-Smoker and/or Moderate/Non-Drinker	8	15

A 'Heavy Smoker' is defined as more than 24 cigarettes per day. A 'Heavy Drinker' is defined as more than six units of alcohol per day.

Table 6.10
HISTOLOGICAL MALIGNANCY GRADE IN SERIES II PATIENTS WITH LYMPH NODE METASTASIS.

Case No.	Malignar						Total mancy Grade
•	Degree of Kerat- inisation	Nuclear Polymor- phism	Frequency of Mitosis	Pattern of Invasion	Stage of Invasion	Lympho- plasma- cytic Infiltrate	
ND9	4	2	2	4	4	4	20
ND31	1	2	2	3	4	1	13
ND39	4	3	4	3	4	4	22
ND40	3	3	3	4	4	1	18
ND51	1	4	3	3	3	2	16
ND52	1	3	3	3	4	3	17
ND63/64	4 4	1	4	3	4	4	20
ND72/7	3 2	2	3	3	4	3	17
ND74/89		1	2	4	4	3	17
ND94	1	2	2	3	4	3	15
ND96	3	2	4	3	4	4	20
ND106/	107 3	3	2	4	3	2	17
ND109	1	3	4	4	4	4	20
ND111	3	3	4	4	4	4	22
ND112	2	2	3	3	4	3	17
ND114/		3	4	4	4	3	21
ND118	2	4	2	3	4	3	18
ND120	3	4	3	4	4	4	22
ND129	4	2	3	4	4	2	19
ND135	3	2	4	3	4	3	19
ND136	4	3	4	4	4	4	23
ND144	4	3	4	4	4	3	22
ND145	3	1	2	3	3	2	14
ND152	3	3	3	4	4	2	19
ND170	3	4	3	4	4	3	21
ND175	2	2	3	4	4	3	18
ND176	3	2	3	4	4	4	20

Table 6.11
HISTOLOGICAL MALIGNANCY GRADE IN SERIES II PATIENTS WITHOUT LYMPH NODE METASTASIS.

Case No.	Histological Feature Point Score Total Malignancy Grade						
C	Degree of Kerat- inisation	Nuclear Polymor- phism	Frequency of Mitosis	Pattern of Invasion	Stage of Invasion	Lympho- plasma- cytic Infiltra	
ND13	1	2	2	2	4	1	12
ND16	1	3	2	3	4	3	16
ND18	2	3	1	1	4	2	13
ND22	1	1	2	2	4	3	13
ND41	3	2	4	3	4	3	19
ND42/43	3 3	2	2	3	3	4	17
ND47	2	2	2	1	4	3	14
ND48	1	2	2	2	3	1	11
ND58/59		2	1	2	4	2	12
ND75/76	6 2	3	2	3	3	2	15
ND85	2	3	1	3	3	1	13
ND92/93		2	2	1	4	3	13
ND132	2	1	3	3	3	3	15
ND134	1	3	3	3	4	1	15
ND138/		1	2	1	3	1	10
ND154/		3	2 3	3	3 3	2 3	15 15
ND166/1 ND169	167 2 2	3 3	3 4	1 1	3	2	15 15
NDIOS	2	3	4	L	3	Z	13

Table 6.12

COMPARISON OF HISTOLOGICAL MALIGNANCY GRADE IN SERIES II PATIENTS WITH AND WITHOUT METASTASIS.

Histological Feature	Chi Squared Test Statisti	Degrees of c Freedom	Probability Value
Degree of Keratinisation	14.17	3	0.003*
Nuclear Polymorphism	3.07	3	0.38
Frequency of Mitosis	11.20	3	0.01*
Pattern of Invasion	25.00	3	0.00002*
Stage of Invasion	8.35	1	0.004*
Lymphoplasmacytic Infiltrate	7.06	3	0.07

^{*} Significant at the 5% level.

Table 6.13

FREQUENCY OF DEGREE OF KERATINISATION POINT SCORES IN PRIMARY TUMOURS WITH AND WITHOUT METASTASIS.

Point Score ^a	Cases With Metastasis	Cases Without Metastasis
1	5	7
2	4	9
3	12	2
4	6	0

^a The criteria for grading and the four point score categories are defined in the text (Chapter 6, Histological Assessment).

Table 6.14

FREQUENCY OF NUCLEAR POLYMORPHISM POINT SCORES IN PRIMARY TUMOURS WITH AND WITHOUT METASTASIS.

Point Score ^a	Cases With Metastasis	Cases Without Metastasis
1	3	3
2	10	7
3	10	8
4	4	0

^a The criteria for grading and the four point score categories are defined in the text (Chapter 6, Histological Assessment).

Table 6.15

FREQUENCY OF MITOSIS POINT SCORES IN PRIMARY TUMOURS WITH AND WITHOUT METASTASIS.

Point Score ^a	Cases With Metastasis	Cases Without Metastasis
1	0	3
2	7	10
3	11	3
4	9	2

 $^{^{\}mathbf{a}}$ The four point score categories are defined in the text (Chapter 6, Histological Assessment).

Table 6.16

FREQUENCY OF PATTERN OF INVASION POINT SCORES IN PRIMARY TUMOURS WITH AND WITHOUT METASTASIS.

Point Score ^a	Cases With Metastasis	Cases Without Metastasis
1	0	6
2	0	4
3	12	8
4	15	0

^a The criteria for grading and the four point score categories are defined in the text (Chapter 6, Histological Assessment).

Table 6.17

• FREQUENCY OF STAGE OF INVASION POINT SCORES IN PRIMARY TUMOURS WITH AND WITHOUT METASTASIS.

Point Score ^a	Cases With Metastasis	Cases Without Metastasis
1	0	0
2	0	0
3	3	9
4	24	9

^a The criteria for grading and the four point score categories are defined in the text (Chapter 6, Histological Assessment).

Table 6.18

FREQUENCY OF LYMHOPLASMACYTIC INFILTRATE POINT SCORES IN PRIMARY TUMOURS WITH AND WITHOUT METASTASIS.

Point Score ^a	Cases With Metastasis	Cases Without Metastasis
1	2	5
2	5	5
3	11	7
4	9	1

^a The criteria for grading and the four point score categories are defined in the text (Chapter 6, Histological Assessment).

Table 6.19 CORRELATION BETWEEN COMPONENT FEATURES OF THE HISTOLOGICAL MALIGNANCY GRADING SYSTEMa.

	_			-		
			istological		-	_
_	K	N 	M 	P	S 	L
N	r _s =0.01 P=0.94					
M	$r_s=0.45$ P=0.002*	$r_s = 0.19$ P = 0.21				
P	r _s =0.57 P=<0.001*	r _s =0.27 P=0.07	r _s =0.39 P=0.007*			
S	$r_s = 0.13$ P = 0.39	r _s =-0.01 P=0.94	r _s =0.28 P=0.06	r _s =0.31 P=0.04*		
L	r _s =0.43 P=0.003*	r _s =-0.02 P=0.88	r _s =0.48 P=0.001*	r _s =0.34 P=0.02*	r _s =0.39 P=0.009*	
G	r _s =0.73 P=<0.001*	r _s = 0.35 P=0.02*	r _s =0.75 P=<0.001*	r _s =0.78 P=<0.001*	r _s =0.44 P=0.002*	r _s =0.70 P=<0.001

 $^{^{\}rm a}$ Spearman's Rank Correlation Test. The test statistic, $\rm r_{\rm s},$ and

associated probability, P, are shown.

b K = Degree of Keratinisation, N = Nuclear Polymorphism, M = Frequency of Mitosis, P = Pattern of Invasion, S = Stage of Invasion, L = Lymphoplasmacytic infiltrate, G = Histological Malignancy Grade.

^{*} Significant at the 5% level.

Table 6.20
PERINEURAL AND VASCULAR INVASION IN SERIES II TUMOURS.

Tumour	Tumour	Perineural	Vascular
Site 	Stage 	Invasion	Invasion
LYMPH NODE METAS	<u> </u>		
Tongue	T1	Identified	Identified
Tongue	T2	Identified	Identified
Tongue	Т3	Identified	Identified
Tongue	T1	Not identified	Not identified
Floor of Mouth	Т4	Not identified	Not identified
Tongue	Т3	Identified	Identified
Tongue	T2	Identified	Identified
Floor of Mouth	Т4	Identified	Identified
Floor of Mouth	T1	Identified	Not identified
Floor of Mouth	Т4	Identified	Identified
Floor of Mouth	Т4	Identified	Identified
Floor of Mouth	T1	Identified	Identified
Floor of Mouth	Т4	Identified	Not identified
		Identified	Identified
Tongue		Identified	Identified
Floor of Mouth		Not identified	Identified
Floor of Mouth			Not identified
			Identified
_			Identified
			Identified
			Identified
			Identified
•			Not identified
			Identified
_			Identified
			Identified
Tongue	T2	Identified	Identified
OUT LYMPH NODE ME	<u> </u>		
Tongue	T2	Identified	Not identified
	T2	Not identified	Not identified
	T2	Not identified	Not identified
	T1	Not identified	Not identified
-	T2	Identified	Identified
Floor of Mouth	Т2	Identified	Not identified
Tongue	T1	Not identified	Not identified
•	T1	Identified	Not identified
		Not identified	Not identified
		Not identified	Not identified
			Not identified
_			Not identified
_			Not identified
			Identified
_			Not identified
			Not identified
			Not identified
Floor of Mouth	T1	Not identified	Not identified
	LYMPH NODE METAST Tongue Tongue Tongue Tongue Tongue Floor of Mouth Tongue Floor of Mouth Tongue Tongue Floor of Mouth Tongue Floor of Mouth Tongue Floor of Mouth	LYMPH NODE METASTASIS Tongue T1 Tongue T2 Tongue T3 Tongue T1 Floor of Mouth T4 Tongue T2 Floor of Mouth T4 Floor of Mouth T1 Floor of Mouth T2 Tongue T2 Floor of Mouth T2 Tongue T2 Floor of Mouth T2 Tongue T2 Floor of Mouth T1 Tongue T2 Floor of Mouth T2 Tongue T2 Floor of Mouth T2 Tongue T2 Floor of Mouth T1 Tongue T2 Tongue T1 Tongue T2 Tongue T1 Tongue T3 Floor of Mouth T1 Tongue T3 Floor of Mouth T1 Tongue T3 Floor of Mouth T1 Floor of Mouth T1 Floor of Mouth T1 Tongue T3 Floor of Mouth T1 Floor of Mouth T1	LYMPH NODE METASTASIS Tongue T1 Identified Tongue T2 Identified Tongue T3 Identified Tongue T4 Identified Tongue T5 Identified Tongue T6 Identified Tongue T7 Identified Tongue T8 Identified Tongue T9 Identified Floor of Mouth T4 Identified Floor of Mouth T5 Identified Floor of Mouth T6 Identified Floor of Mouth T7 Identified Floor of Mouth T2 Identified Floor of Mouth T2 Identified Floor of Mouth T3 Not identified Floor of Mouth T4 Identified Floor of Mouth T5 Identified Floor of Mouth T6 Identified Floor of Mouth T7 Identified Floor of Mouth T8 Identified Floor of Mouth T9 Identified Floor of Mouth T1 Identified Floor of Mouth T1 Identified Floor of Mouth T1 Identified Tongue T2 Identified Floor of Mouth T1 Identified Tongue T2 Identified Tongue T1 Inot identified Tongue T1 Inot identified Tongue T1 Inot identified Floor of Mouth T1 Inot identified

Table 6.21

CORRELATION BETWEEN COMPONENT FEATURES OF HISTOLOGICAL MALIGNANCY GRADE AND PERINEURAL AND VASCULAR INVASION^a.

	Histological Malignancy Grade	Perineural Invasion	Vascular Invasion
Degree of Keratin- isation	$r_s = 0.73$ $P = <0.001*$	r _s = 0.20 P = 0.19	r _s = 0.48 P = 0.001*
Nuclear Poly- morphism	$r_s = 0.35$ P = 0.02*	$r_s = -0.17$ $P = 0.26$	$r_s = 0.07$ P = 0.65
Frequency of Mitosis	$r_s = 0.75$ P = <0.001*	$r_s = 0.26$ P = 0.09	$r_s = 0.56$ $P^s = <0.001*$
Pattern of Invasion	$r_s = 0.78$ $P = <0.001*$	$r_s = 0.40$ P = 0.007*	$r_s = 0.56$ P = <0.001*
Stage of Invasion	$r_S = 0.44$ P = 0.002*	$r_s = 0.34$ $P = 0.02*$	$r_s = 0.52$ P = <0.001*
Lympho- plasma cytic Infiltrate	$r_s = 0.70$ $P = <0.001*$	$r_s = 0.28$ $P = 0.06*$	$r_s = 0.38$ P = 0.01*
Histo- logical Malignancy Grade		$r_s = 0.36$ P = 0.02*	
Perineural Invasion	$r_s = 0.35$ P = 0.02*		$r_s = 0.60$ P = <0.001*
Vascular Invasion	$r_s = 0.38$ P = 0.01*		

 $^{^{\}rm a}$ Spearman's Rank Correlation Test. The test statistic, $\rm r_{\rm s},$ and associated probability, P, are shown.

^{*} Significant at the 5% level.

Table 6.22

POTENTIAL PREDICTOR VARIABLES AND THEIR ASSOCIATED PROBABILITY VALUE.

Predictor Variable	Probability Valu	ue Statistical Method
Tumour Site	1.00	Yates' Chi Squared Test
Tumour T Stage	0.28	Chi Squared Test
Sex	0.75	Yates' Chi Squared Test
Age	0.39	Two Sample t Test
Tobacco Usage	0.08	Mann-Whitney U Test
Alcohol Consumption	0.13	Mann-Whitney U Test
Tumour Dimension	0.04*	Two Sample t Test
Actual Tumour Thickness	0.008*	Two Sample t Test
Reconstructed Tumour Thickr	ness 0.005*	Two Sample t Test
Epithelial Dysplasia	0.58	Yates' Chi Squared Test
Histological Malignancy Gra	ide 0.0001*	Mann-Whitney U Test
Degree of Keratinisation	0.003*	Chi Squared Test
Nuclear Polymorphism	0.38	Chi Squared Test
Frequency of Mitosis	0.01*	Chi squared Test
Pattern of Invasion	0.00002*	Chi Squared Test
Stage of Invasion	0.004*	Chi Squared Test
Lymphoplasmacytic Infiltrat	e 0.07	Chi Squared Test
Perineural Invasion	0.01*	Yates' Chi Squared Test
Vascular Invasion	0.00005*	Yates' Chi Squared Test

 $[\]star$ Significant at the 5% level.

Table 6.23
TRANSFORMATION OF MULTICATEGORICAL DATA INTO BINARY FORM.

Histological	Binary	
Feature	0	1
Degree of Keratinisation	Point scores 1 and 2 Formation of keratin pearls	Point scores 3 and 4 Individual cell or no keratinisation
Frequency of Mitosis	Point scores 1 and 2 0-35 per 10 HPF	Point scores 3 and 4 >35 per 10 HPF
Pattern of Invasion	Point scores 1 and 2 Pushing or broad strands and cords	Point scores 3 and 4 Thin cords or small islands or single cells
Stage of Invasion	Point score 3 Invasion into submucosa	Point score 4 Invasion into muscle
Lymphoplasmacytic Infiltrate	Point scores 1 and 2 Continuous rim or many large patches	Point scores 3 and 4 Slight or none

Table 6.24 LOGISTIC REGRESSION MODEL WITH ELEVEN INDEPENDENT VARIABLES.

Variable	Regression Coefficient (Beta)	Standard Error	Chi Squared (X ²)	Probab- ility
Intercept ^a	-17.55	30.34	0.33	0.56
Tumour Dimension	0.00	0.08	0.01	0.92
Actual Tumour Thickness	0.01	0.34	0.00	0.96
Reconstructed Tumour Thickness	-0.22	0.36	0.36	0.55
Histological Malignancy Grade	0.63	0.51	1.54	0.22
Degree of Keratinisation	0.17	1.80	0.01	0.92
Frequency of Mitosis	0.13	1.63	0.01	0.94
Pattern of Invasion	8.29	§		
Stage of Invasion	1.86	1.90	0.95	0.33
Lymphoplasmacytic Infiltrate	-1.21	1.62	0.56	0.46
Perineural Invasion	-0.19	1.97	0.01	0.93
Vascular Invasion	2.38	2.54	0.88	0.35

 $[\]mbox{\colored}$ Denotes infinity. $\mbox{\colored}$ The intercept is the logit of the probability when all factors have a value of zero.

Table 6.25

CLASSIFICATION TABLE FOR THE LOGISTIC REGRESSION MODEL WITH ELEVEN INDEPENDENT VARIABLES.

	Predicted Negative	Predicted Positive	Total
rue Negative	16	2	18
True Positive	2	25	27
Total	18	27	45

Sensitivity: 92.6% Specificity: 88.9% Correct: 91.1% False Positive Rate: 7.4% False Negative Rate: 11.1%

Table 6.26 LOGISTIC REGRESSION MODEL WITH NINE INDEPENDENT VARIABLES.

Variable	Regression Coefficient (Beta)	Standard Error	Chi Squared (X ²)	Probab- ility
Intercept ^a	-17.57	30.53	0.33	0.56
Reconstructed Tumour Thickness	-0.20	0.20	0.93	0.34
Histological Malignancy Grade	0.64	0.50	1.67	0.20
Degree of Keratinisation	0.10	1.63	0.00	0.95
Frequency of Mitosis	0.10	1.60	0.00	0.95
Pattern of Invasion	8.20	§		
Stage of Invasion	1.88	1.88	1.00	0.32
Lymphoplasmacytic Infiltrate	-1.20	1.56	0.59	0.44
Perineural Invasion	-0.20	1.98	0.01	0.92
Vascular Invasion	2.43	2.49	0.95	0.33

 $[\]mbox{\$}$ Denotes infinity. \mbox{a} The intercept is the logit of the probability when all factors have a value of zero.

Table 6.27

LOGISTIC REGRESSION MODEL WITH FOUR INDEPENDENT VARIABLES.

Variable	Regression Coefficient (Beta)	Standard Error	Chi Squared (X ²)	Probab- ility
•				
Intercept ^a	-11.42	4.32	7.00	0.008*
Reconstructed Tumour Thickness	-0.17	0.16	1.04	0.31
Histological Malignancy Grade	0.74	0.30	6.15	0.01*
Perineural Invasion	-0.01	1.47	0.00	0.99
Vascular Invasion	3.47	2.17	2.57	0.11

 $^{^{\}mathrm{a}}$ The intercept is the logit of the probability when all factors have a value of zero.

^{*} Significant at the 5% level.

Table 6.28

CLASSIFICATION TABLE FOR THE LOGISTIC REGRESSION MODEL WITH FOUR INDEPENDENT VARIABLES.

Predicted Negative	Predicted Positive	Total
15	3	18
3	24	27
18	27	45
	Negative 15	Negative Positive 15 3 3 24

Sensitivity: 88.9% Specificity: 83.3% Correct: 86.7% False Positive Rate: 11.1% False Negative Rate: 16.7%

Table 6.29 LOGISTIC REGRESSION MODEL WITH THREE INDEPENDENT VARIABLES.

Variable	Regression Coefficient (Beta)	Standard Error	Chi Squared (X ²)	Probab- ility
Intercept ^a	-11.41	4.18	7.44	0.006*
Reconstructed Tumour Thickness	-0.17	0.16	1.05	0.31
Histological Malignancy Grade	0.73	0.28	6.72	0.009*
Vascular Invasion	3.47	1.93	3.21	0.07

^a The intercept is the logit of the probability when all factors have a value of zero.
* Significant at the 5% level.

Table 6.30 'BEST' LOGISTIC REGRESSION MODEL WITH TWO INDEPENDENT VARIABLES.

Variable	Regression Coefficient (Beta)	Standard Error	Chi Squared (X ²)	Probab- ility
Intercept ^a	-10.73	3.81	7.93	0.005*
Histological Malignancy Grade	0.64	0.24	7.04	0.008*
Vascular Invasion	1.95	1.04	3.52	0.05*

 $^{^{\}mathbf{a}}$ The intercept is the logit of the probability when all factors have a value of zero.

* Significant at the 5% level.

Table 6.31 'ALTERNATIVE' LOGISTIC REGRESSION MODEL WITH TWO INDEPENDENT VARIABLES.

Variable	Regression Coefficient (Beta)	Standard Error	Chi Squared (X ²)	Probab- ility
Intercept ^a	-8.28	19.86	0.17	0.68
Pattern of Invasion	8.28	19.86	0.17	0.68
Vascular Invasion	2.35	0.94	6.28	0.01*

 $^{^{\}mathbf{a}}$ The intercept is the logit of the probability when all factors have a value of zero.
* Significant at the 5% level.

Table 6.32

CLASSIFICATION TABLE FOR 'ALTERNATIVE' TWO-VARIABLE LOGISTIC REGRESSION MODEL.

	Predicted Negative	Predicted Positive	Total	
True Negative	16	2	18	
True Positive	6	21	27	
Total	18	27	45	

Sensitivity: 77.8%
Specificity: 88.9%
Correct: 82.2%
False Positive Rate: 8.7%
False Negative Rate: 27.3%

Table 6.33
ESTIMATED PROBABILITY OF NODAL METASTASIS IN SERIES II PATIENTS.

Case No. Histological Malignancy Grade Vascular Invasion Probability

CACEC LITTU IND	NI NODE METACTACI		
ND 9	<u>PH NODE METASTASI</u> 20	<u>s</u> Identified	0.98
ND 31	13	Identified	0.38§
ND 39	22	Identified	0.99
ND 40	18	Not Identified	0.68
ND 51	16	Not Identified	0.37§
ND 52	17	Identified	0.89
ND 63/64	20	Identified	0.98
ND 72/73	17	Identified	0.89
ND 74/89	17	Not Identified	0.53
ND 94	15	Identified	0.70
ND 96	20	Identified	0.98
ND 106/107	17	Identified	0.89
ND 109	20	Not Identified	0.89
ND 111	22	Identified	0.99
ND 112	17	Identified	0.89
ND 114/115	21	Identified	0.99
ND 118	18	Not Identified	0.68
ND 120	22	Identified	0.99
ND 129	19	Identified	0.97
ND 135	19	Identified	0.97
ND 136	23	Identified	0.99
ND 144	22	Identified	0.99
ND 145	14	Not Identified	0.14§
ND 152	19	Identified	0.97
ND 170	21	Identified	0.99
ND 175	18	Identified	0.94
ND 176	20	Identified	0.98
CASES WITHOUT I	LYMPH NODE METAST	CASIS	
ND 13	12	Not Identified	0.04
ND 16	16	Not Identified	0.37
ND 18	13	Not Identified	0.08
ND 22	13	Not Identified	0.08
ND 41	19	Identified	0.97§
ND 42/43	17	Not Identified	0.53§
ND 47	14	Not Identified	0.14
ND 48	11	Not Identified	0.02
ND 58/59	12	Not Identified	0.04
ND 75/76	15	Not Identified	0.24
ND 85	13	Not Identified	0.08
ND 92/93	13	Not Identified	0.08
ND 132	15	Not Identified	0.24
ND 134	15	Identified	0.69§
ND 138/139	10	Not Identified	0.01
ND 154/155	15	Not Identified	0.24
ND 166/167	15	Not Identified	0.24
ND 169	15	Not Identified	0.24
103 TO3	1.)	Not identified	0.24

[§] Denotes a misclassification.

Table 7.1

COMPARISON OF STEREOLOGICAL ESTIMATION OF MEAN NUCLEAR VOLUME ON TWO SEPARATE OCCASIONS.

ilot est ase	Mean Nuclear Volume (µm³)	Coefficient of Error
	745.2 736.2	1.2 1.5
	492.4 514.5	1.3 1.3
	1305.0 1259.0	1.6 1.5
	224.8 235.0	1.3 1.3
; ;	380.0 407.3	1.5 1.5

Table 7.2

NUCLEAR FEATURES IN TUMOURS IN SERIES II PATIENTS WITH METASTASIS.

Case Number	Mean Nuclear Volume	Coefficient of Error	Nuclear Polymorphism Point Score ^a
ND9	880.7	0.14	2
ND31	597.8	0.13	2
ND39	1305.0	0.16	3
ND40	498.2	0.16	3
ND51	981.9	0.13	4
ND52	490.5	0.14	3
ND63/64	224.8	0.16	
ND72/73	630.0	0.13	1 2
ND74/89	360.6	0.17	
ND94	792.0	0.14	1 2 2 3 3
ND96	460.1	0.14	2
ND106/107	766.1	0.14	3
ND109	1124.0	0.13	3
ND111	522.9	0.17	3
ND112	638.1	0.14	2
ND114/115	398.0	0.13	3
ND118	952.2	0.13	4
ND120	858.2	0.13	4
ND129	632.3	0.13	2
ND135	947.0	0.13	2
ND136	559.7	0.14	3 3
ND144	405.6	0.14	
ND145	380.0	0.15	1
ND152	1089.0	0.13	3
ND170	603.3	0.15	4
ND175	954.3	0.16	2
ND176	1290.0	0.13	2

^a Criteria for point scoring 'Nuclear Polymorphism' are detailed in the text (Chapter 6, Histological Assessment).

Table 7.3

NUCLEAR FEATURES IN TUMOURS IN SERIES II PATIENTS WITHOUT METASTASIS.

Case Number	Mean Nuclear Volume	Coefficient of Error	Nuclear Polymorphism Point Score ^a
ND13	523.6	0.17	2
ND16	745.2	0.12	3
ND18	492.4	0.13	3
ND22	528.6	0.17	1
ND41	1249.0	0.18	2
ND42/43	406.4	0.15	2
ND47	239.5	0.15	2
ND48	488.0	0.19	2
ND58/59	353.0	0.16	2
ND75/76	1079.0	0.17	3
ND85	530.9	0.15	3
ND92/93	971.6	0.13	2
ND132	566.8	0.14	1
ND134	1340.0	0.17	3
ND138/139	524.8	0.13	1
ND154/15	768.9	0.15	3
ND166/16	7 719.6	0.15	3 3
ND169	576.0	0.14	3

^a Criteria for point scoring 'Nuclear Polymorphism' are detailed in the text (Chapter 6, Histological Assessment).

Section 2.

FIGURES.

- 1. Figures 4.1-4.13.
- 2. Figures 5.1-5.3.
- 3. Figures 6.1-6.15.
- 4. Figures 7.1-7.4.

Figure 4.1a

A UNILATERAL SURGICAL NECK DISSECTION SPECIMEN FOLLOWING PINNING OUT AND FIXATION.



The neck dissection is a radical procedure. The sternomastoid and omohyoid muscles have been pinned out in their extended positions prior to fixation. A bulky metastatic mass is present in the submandibular triangle.

Figure 4.1b

A BILATERAL SURGICAL NECK DISSECTION FOLLOWING PINNING OUT AND FIXATION.



Both neck dissections are functional procedures, and are continuous with the primary tumour, a T4 squamous cell carcinoma of floor of mouth, necessitating resection of the body of mandible and the oral tongue.

Figure 4.2a

EXTENT OF NODAL REPLACEMENT: MINIMAL.



Metastatic carcinoma is seen as small islands of tumour cells within the peripheral sinuses of the node, with minimal replacement of normal nodal architecture. Haematoxylin and eosin. Original magnification, $\times 100$.

Figure 4.2b

EXTENT OF NODAL REPLACEMENT: PARTIAL.



Metastatic carcinoma is seen replacing approximately 15-20% of the normal nodal architecture. Haematoxylin and eosin. Original magnification, x40.

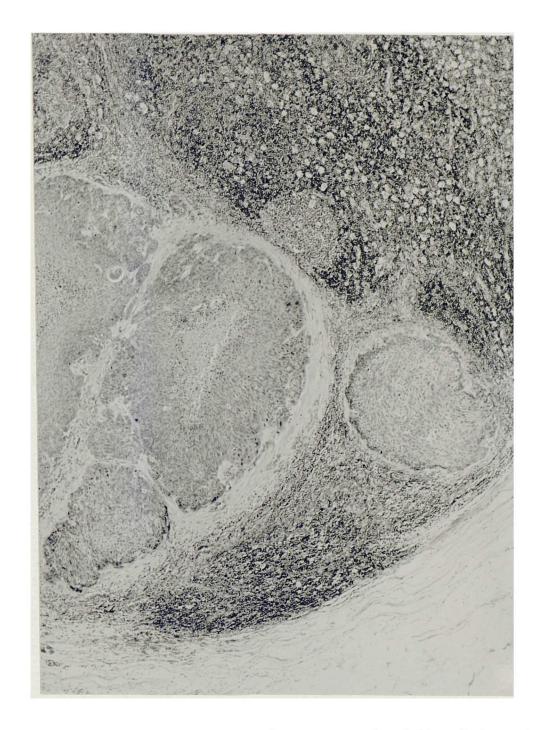
Figure 4.2c

EXTENT OF NODAL REPLACEMENT: TOTAL.



The lymph node is almost completely replaced by metastatic carcinoma. Haematoxylin and eosin. Original magnification, $\varkappa 40\,.$

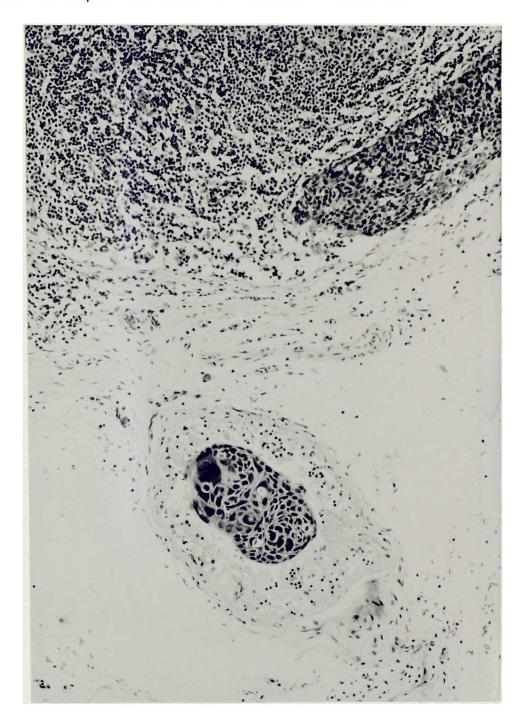
Figure 4.3a
HISTOLOGICAL ASSESSMENT OF EXTRACAPSULAR SPREAD: NO EXTRACAPSULAR SPREAD.



Metastatic carcinoma is confined to the cortex and medulla of the node and there is no involvement of the capsule. Haematoxylin and eosin. Original magnification, x40.

Figure 4.3b

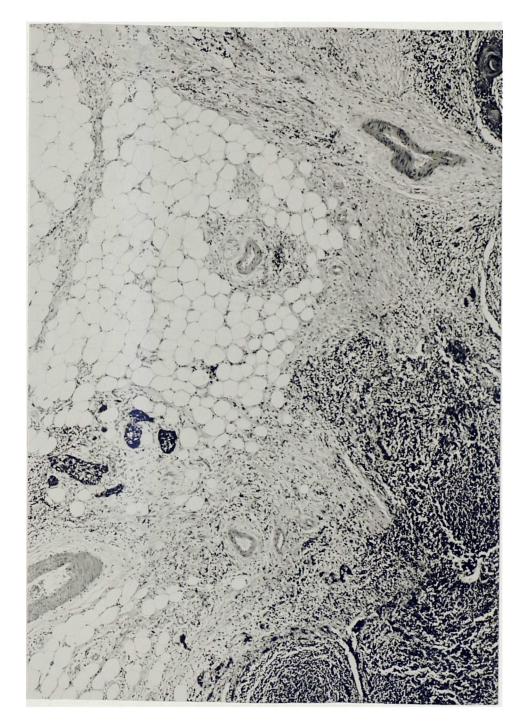
HISTOLOGICAL ASSESSMENT OF EXTRACAPSULAR SPREAD: MICROSCOPIC EMBOLISATION AND/OR PERMEATION OF PERINODAL LYMPHATICS.



Tumour cells are seen within a lymphatic vessel in the fibro-adipose tissue surrounding the node. Haematoxylin and eosin. Original magnification, x100.

Figure 4.3c

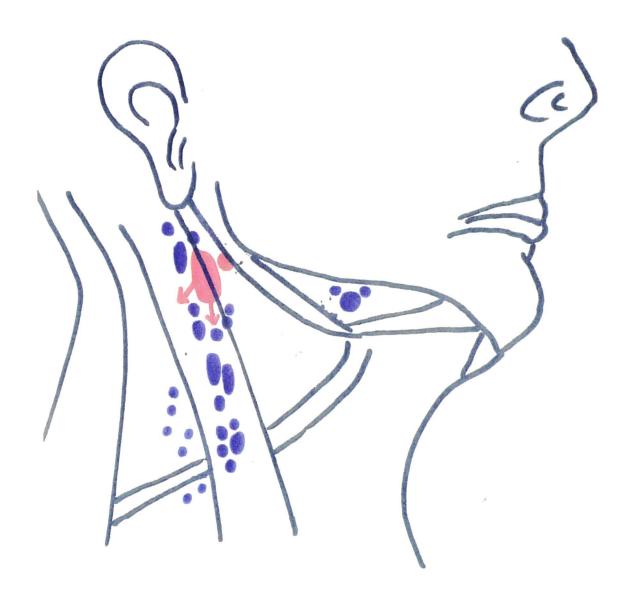
HISTOLOGICAL ASSESSMENT OF EXTRACAPSULAR SPREAD: MICROSCOPIC EXTRACAPSULAR SPREAD TO PERINODAL TISSUES AND/OR ANATOMICAL STRUCTURES.



Small islands of tumour are seen infiltrating perinodal fibro-adipose tissue. Islands of tumour are seen, also, within the capsule and periphery of the node. Haematoxylin and eosin. Original magnification, x40.

Figure 4.4a

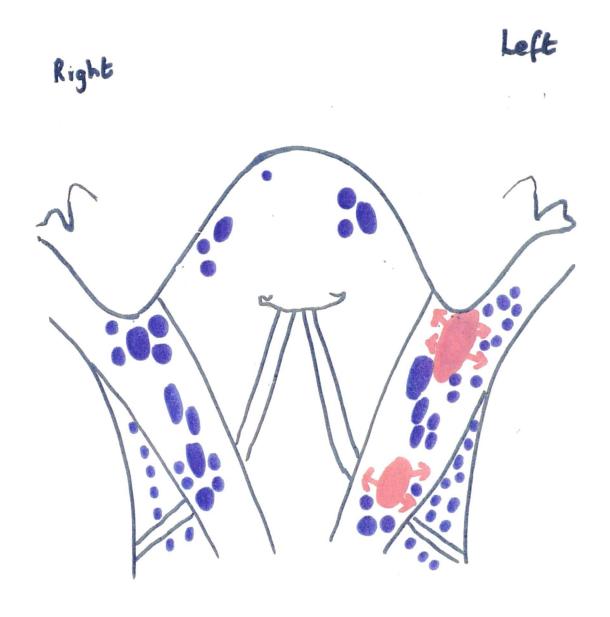
TOPOGRAPHICAL DEPICTION OF THE PATHOLOGICAL FINDINGS IN A UNILATERAL NECK DISSECTION SPECIMEN.



Metastatic carcinoma is present in two nodes (indicated in red) at level II (superior cervical). Extracapsular spread (indicated by red arrows) is present in relation to the larger positive node. The size and position of 26 negative nodes is indicated in blue.

Figure 4.4b

TOPOGRAPHICAL DEPICTION OF THE PATHOLOGICAL FINDINGS IN A BILATERAL NECK DISSECTION SPECIMEN.



On the left, metastatic masses (indicated in red) are present at levels II and IV (superior and inferior cervical, respectively). The arrows indicate extracapsular spread. The size and position of 34 negative nodes is indicated in blue.

On the right, there was no evidence of metastasis. The size and position of 24 negative nodes is indicated in blue.

Figure 4.5

NUMBER AND ANATOMICAL LEVEL OF METASTATIC DEPOSITS^a IN EIGHTEEN UNILATERAL PROCEDURE PATIENTS.

Case ND	T Site	T Stage	Anatomical Level I II III IV V
2	Buccal Mucosa	Т2	+, + + + + +
5	Buccal Mucosa	T2	
125	Buccal Mucosa	T 4	
			+ + + + + + +
9	Oral Tongue	T1	
122	Oral Tongue	T1	
31	Oral Tongue	T2	-
120	Oral Tongue	T2	SECURIOR STATE OF SECURIOR STA
129	Oral Tongue	T2	
144	Oral Tongue	T2	
39	Oral Tongue	т3	Replacement Replac
52	Oral Tongue	Т3	
110	Oral Tongue	Т4	+ + + + +
111	Floor of Mouth	T2	
135	Floor of Mouth	T2	
51	Floor of Mouth	Т4	-
96	Floor of Mouth	T4	
109	Floor of Mouth	T4	-
108	Oropharynx	Т4	+ + + + + + + + + + + + + + + + + + +

Positive discrete lymph nodes are indicated in red. Metastatic masses are indicated in black.

Figure 4.6 NUMBER AND ANATOMICAL LEVEL OF METASTATIC DEPOSITS^a IN THIRTEEN BILATERAL PROCEDURE PATIENTS.

Case :	Γ Site	e ^b ∕S	tage	v		t Neck A		nical I	evel	eft N		v v	,
			+	+	+	+	+	++	+	+	+	+	+
64/63	To	T2				-	_ =						
113/112	To	T2						=	-				
153/152	To	T2						_	Name of the last o				
			+	+	+	+	+	++	+	+	+	+	+
84/83	FOM	T1					-	E/Minor					
107/106	FOM	T1					-	-	hilliane.				
118/117	FOM	T1					-	-					
115/114	FOM	T2					_			-			
137/136	FOM	T2							-	-			
73/72	FOM	T4							- Minor	name to the same of the same o	-		
95/94	FOM	T4						_					
			+	+	+	4	+	++	+	+	+	+	+
142/141	Orop	T1				-			*****	entate un			
159/158	Orop	Т3							_	_			
157/156	Orop	T 4				-							
			+	+	+	+	+	++	+	+ .	+	+	+

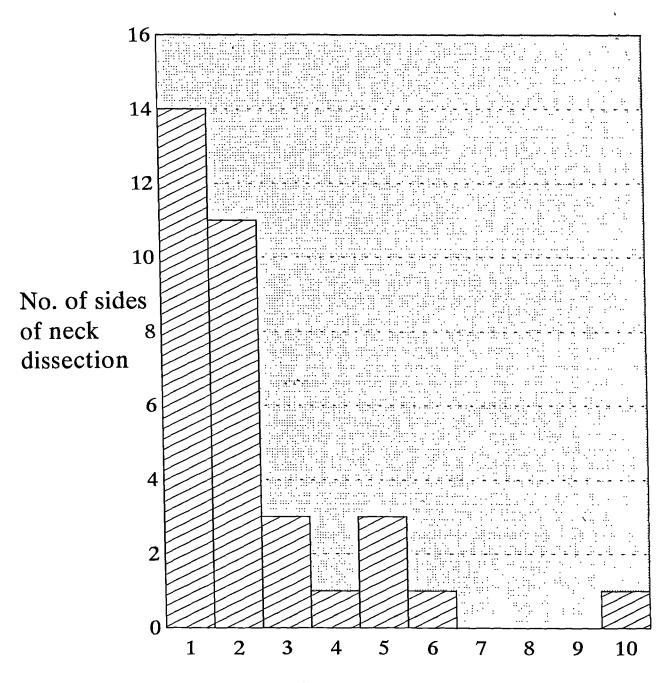
 $^{^{\}rm a}$ Positive discrete lymph nodes are indicated in red. Metastatic masses of fused nodes are indicated in black. b To = Oral tongue, FOM = Floor of mouth, Orop = Oropharynx.

Figure 4.7

PREVALENCE OF METASTASIS AT DIFFERENT ANATOMICAL LEVELS IN THIRTY-FOUR SIDES OF NECK DISSECTION.

Dissect		+==+ + + + +
12	(35%)	
9	(26%)	
2	(6%)	
3	(9%)	
1	(3%)	
1	(3%)	
2	(6%)	
1	(3%)	
2	(6%)	
1	(3%)	

NUMBER OF METASTATIC DEPOSITS PER Fig 4.8 DISSECTION IN THIRTY-FOUR SIDES OF NECK DISSECTION.



Number of metastatic deposits

Figure 4.9a
METASTATIC MASS SHOWING MACROSCOPIC CYSTIC CHANGE.



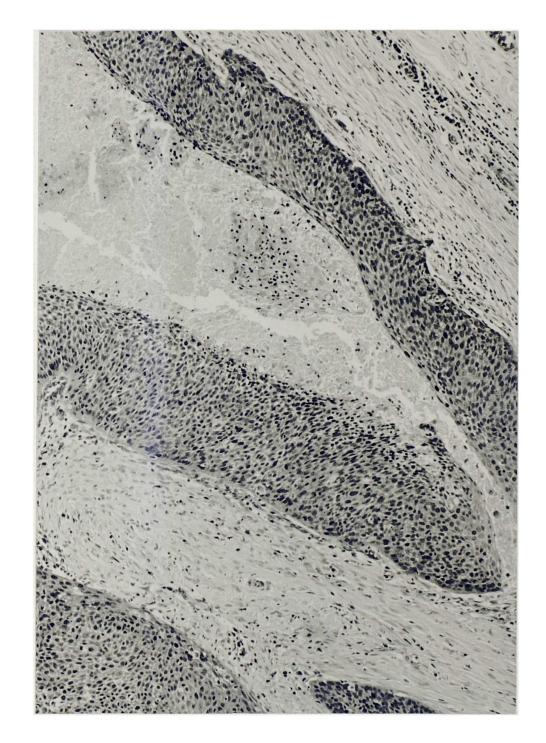
A metastatic mass (measuring 4.0cm.), formed by fusion of three nodes. The central lumen, which occupied one half of the mass, contained gelatinous material. Solid parts of the mass are projecting into the lumen.



The wall of the cyst is lined, in part, by a thin layer of epithelium. The lumen contained, mainly, necrotic debris. Haematoxylin and eosin. Original magnification, x40.

Figure 4.10

CYSTIC METASTATIC MASS: TRANSITIONAL-TYPE EPITHELIAL LINING.



The papillary processes are lined by multiple layers of non-keratinising cells. These are orientated towards the surface, thus creating the appearances of transitional-type epithelium. Haematoxylin and eosin. Original magnification, x40.

Figure 4.11

CYSTIC METASTATIC LYMPH NODE: MICROSCOPIC APPEARANCE.



The lumen is full of keratin lamellae. A peripheral rim of lymphoid tissue is preserved and there is no extracapsular spread. Haematoxylin and eosin. Original magnification, x100.

Figure 4.12

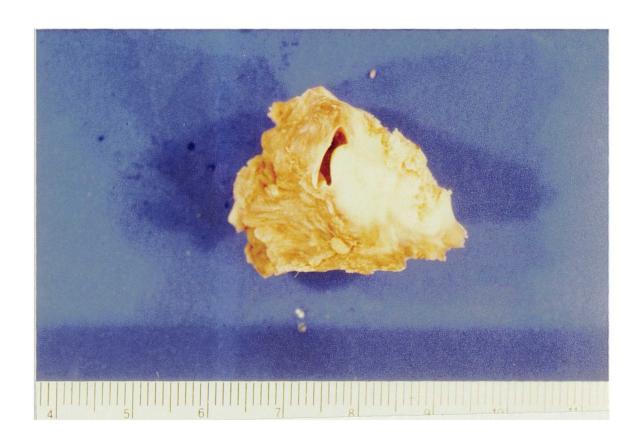
SPREAD OF METASTATIC CARCINOMA TO SUBCUTANEOUS TISSUES AND SKIN.



The epidermis is ulcerated with a fibrinous slough. Squamous cell carcinoma is extensively replacing the subcutaneous tissues. Haematoxylin and eosin. Original magnification, x40.

Figure 4.13a

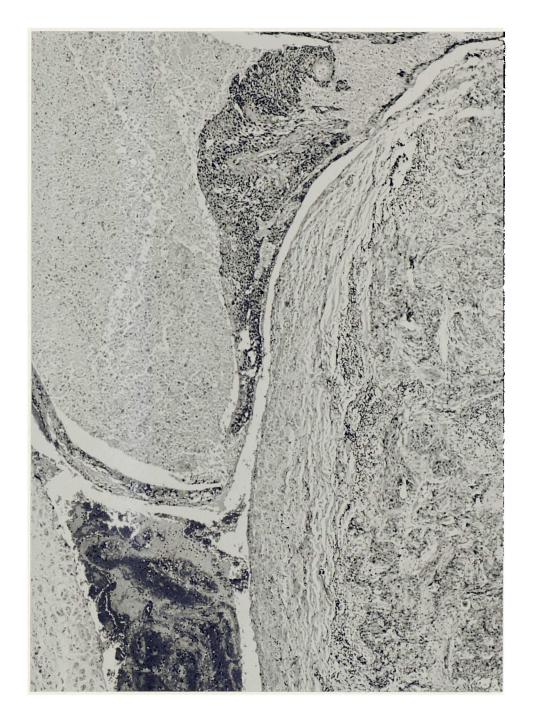
INVASION OF INTERNAL JUGULAR VEIN BY METASTATIC CARCINOMA: MACROSCOPIC APPEARANCE.



Tumour is spreading from a metastatic mass into the wall of the internal jugular vein.

Figure 4.13b

INVASION OF INTERNAL JUGULAR VEIN BY METASTATIC CARCINOMA: MICROSCOPIC APPEARANCE.



The lumen of the vein is occupied by a mass of tumour showing central necrosis. Squamous cell carcinoma has extensively involved the adventitia of the vein, and, in areas, breached the muscular media. Haematoxylin and eosin. Original magnification, x40.

Figure 5.1

ACCURACY OF THE CLINICAL ASSESSMENT OF THE METASTATIC STATUS OF CERVICAL NODES IN UNILATERAL NECK DISSECTION PATIENTS.

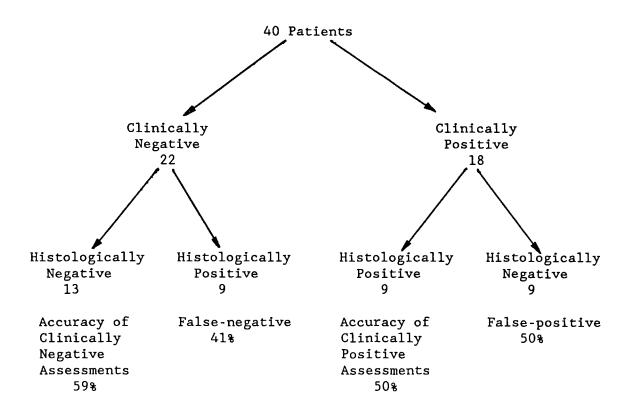


Figure 5.2

ACCURACY OF THE CLINICAL ASSESSMENT OF THE METASTATIC STATUS OF CERVICAL NODES IN BILATERAL NECK DISSECTION PATIENTS.

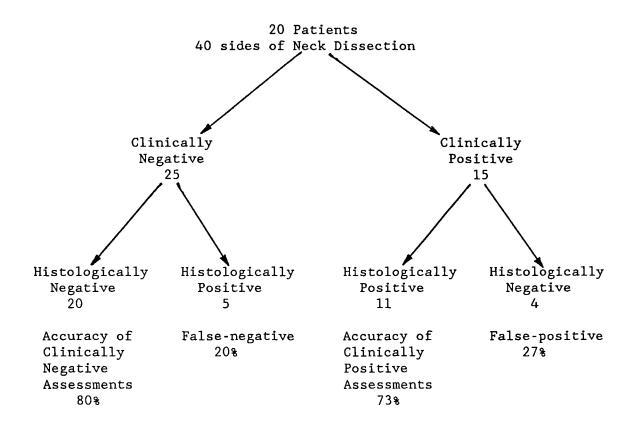


Figure 5.3

ACCURACY OF THE CLINICAL ASSESSMENT OF THE METASTATIC STATUS OF CERVICAL NODES IN SIMULTANEOUS PROCEDURE NECK DISSECTION PATIENTS.

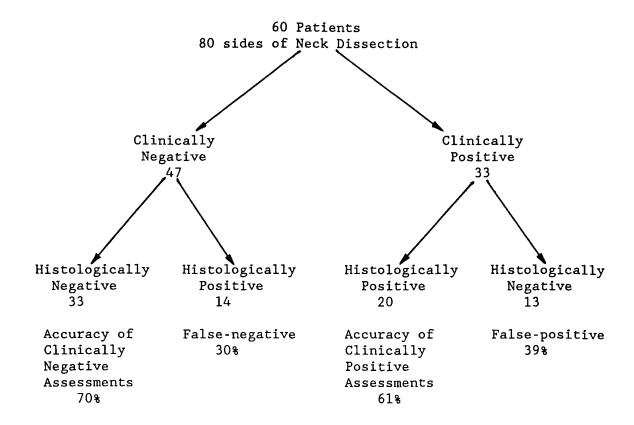
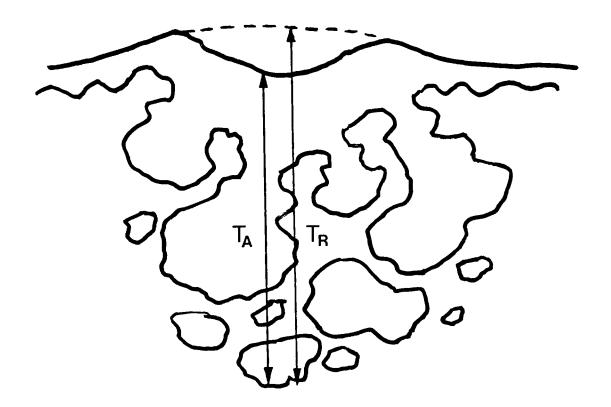


Figure 6.1a

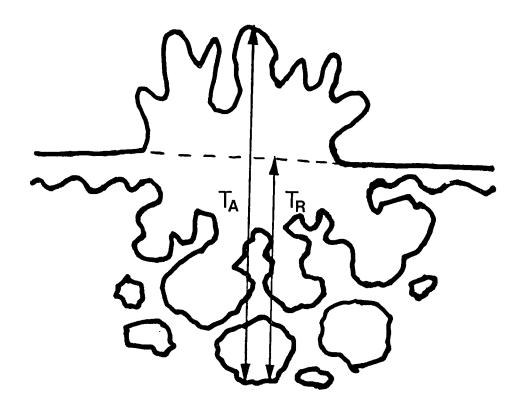
ACTUAL TUMOUR THICKNESS (T_A) AND RECONSTRUCTED TUMOUR THICKNESS (T_R) IN AN ULCERATED, ENDOPHYTIC CARCINOMA.



The actual tumour thickness (\mathtt{T}_A) was measured from the floor of the ulcer to the deepest extent of growth at the advancing tumour front. The reconstructed tumour thickness (\mathtt{T}_R) was measured from a theoretically reconstructed normal mucosal line to the deepest extent of growth at the advancing tumour front.

Figure 6.1b

ACTUAL TUMOUR THICKNESS (\mathbf{T}_{A}) AND RECONSTRUCTED TUMOUR THICKNESS (\mathbf{T}_{R}) IN AN EXOPHYTIC CARCINOMA.



The actual tumour thickness (T_A) was measured from the papillary surface (including the most superfical layer of keratinocytes with open nuclei) to the deepest extent of growth at the advancing tumour front. The reconstructed tumour thickness (T_R) was measured from a theoretically reconstructed normal mucosal line to the deepest extent of growth at the advancing tumour front.

Figure 6.2a

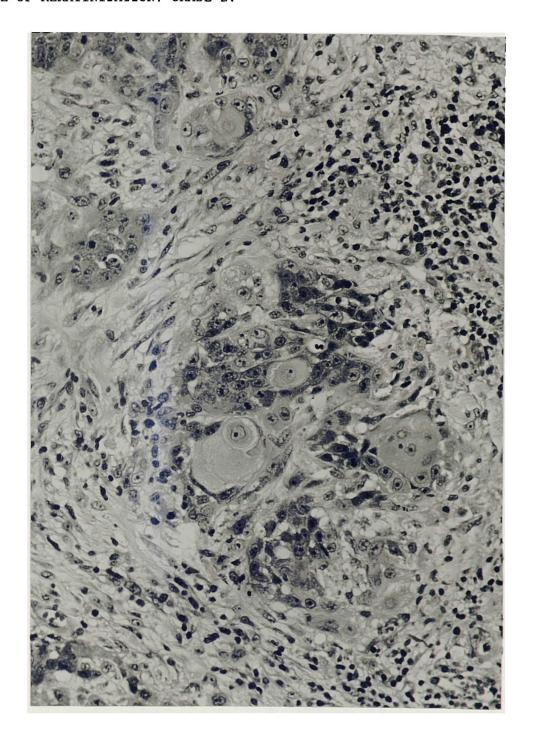
DEGREE OF KERATINISATION: GRADE 1.



Abundant well-formed keratin pearls are seen at the advancing front. Haematoxylin and eosin. Original magnification, x100.

Figure 6.2b

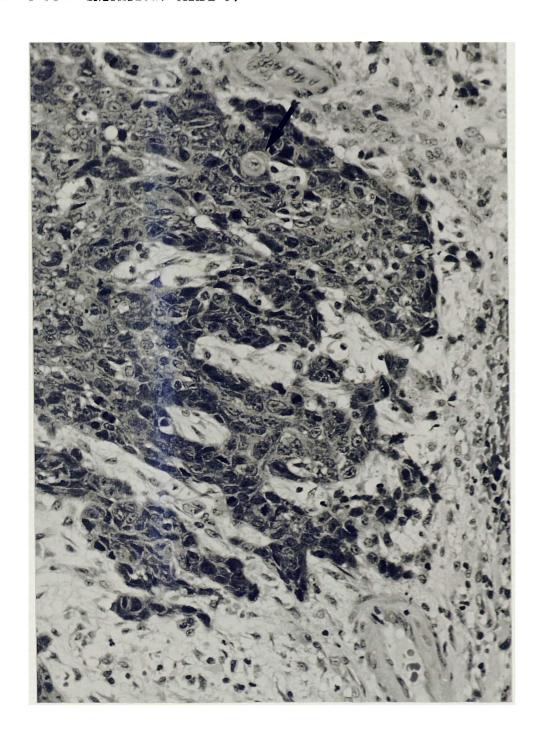
DEGREE OF KERATINISATION: GRADE 2.



Poorly-formed keratin pearls are seen at the advancing front. Haematoxylin and eosin. Original magnification, x200.

Figure 6.2c

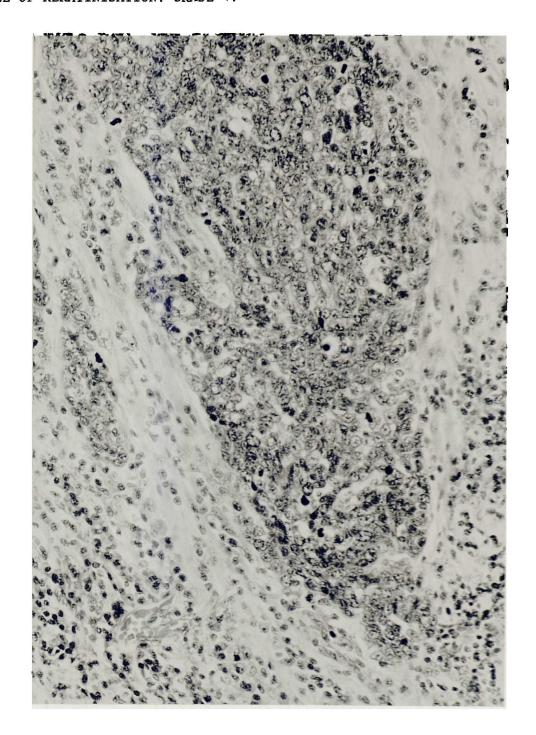
DEGREE OF KERATINISATION: GRADE 3.



At the advancing front, no keratin pearls are present, but keratin is seen within individual tumour cells (indicated by arrow). Haematoxylin and eosin. Original magnification, x200.

Figure 6.2d

DEGREE OF KERATINISATION: GRADE 4.



Tumour cells at the advancing front show no evidence of keratinisation. Haematoxylin and eosin. Original magnification, x200.

Figure 6.3a
NUCLEAR POLYMORPHISM: GRADE 1.



The tumour cell nuclei at the advancing front are small, almost uniform in size, with minimal aberrations. Haematoxylin and eosin. Original magnification, x200.

Figure 6.3b

NUCLEAR POLYMORPHISM: GRADE 2.



Tumour cell nuclei at the advancing front are large, with distinct nucleoli. Multiple/large anaplastic nuclei are not detected. Haematoxylin and eosin. Original magnification, x400.

Figure 6.3c

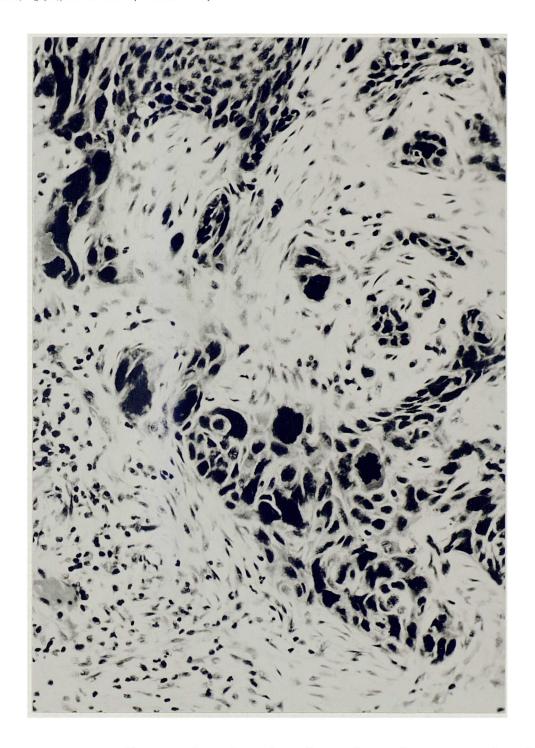
NUCLEAR POLYMORPHISM: GRADE 3.



Some tumour cells at the advancing front have large, anaplastic nuclei. Haematoxylin and eosin. Original magnification, x200.

Figure 6.3d

NUCLEAR POLYMORPHISM: GRADE 4.



Numerous tumour cells at the advancing front have large, anaplastic nuclei or atypical mitotic figures. Haematoxylin and eosin. Original magnification, x200.

Figure 6.4a

PATTERN OF INVASION: GRADE 1.



The tumour has a well-delineated, pushing border. Haematoxylin and eosin. Original magnification, x40.

Figure 6.4b

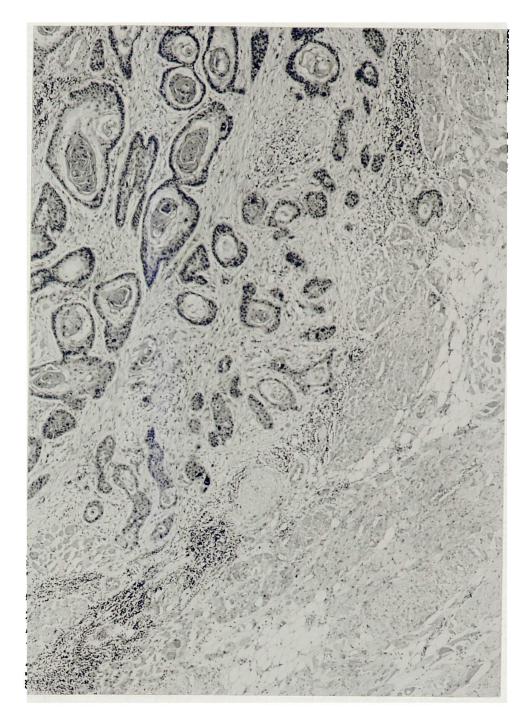
PATTERN OF INVASION: GRADE 2.



Broad bands and solid cords of tumour cells are seen at the advancing front. Haematoxylin and eosin. Original magnification, x40.

Figure 6.4c

PATTERN OF INVASION: GRADE 3.



Small islands of tumour cells (each containing more than 15 cells) are seen at the advancing front. Haematoxylin and eosin. Original magnification, x40.

Figure 6.4d

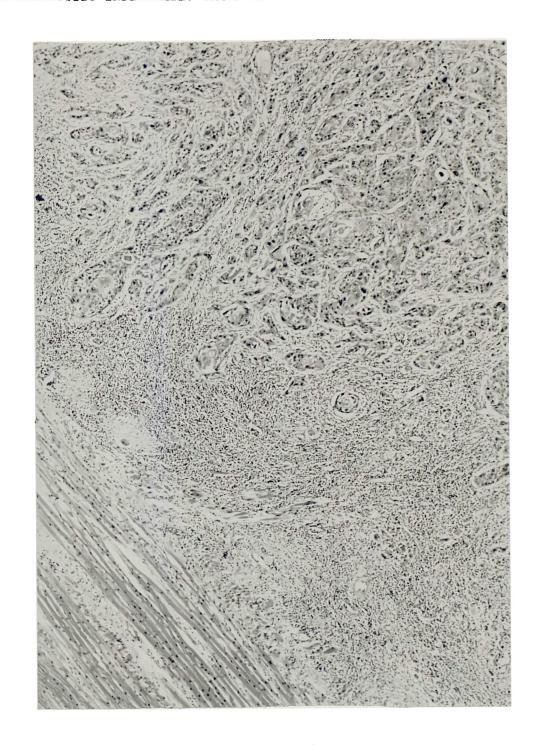
PATTERN OF INVASION: GRADE 4.



The advancing front is poorly delineated with tiny islands and individual tumour cells widely dispersed. Haematoxylin and eosin. Original magnification, x40.

Figure 6.5a

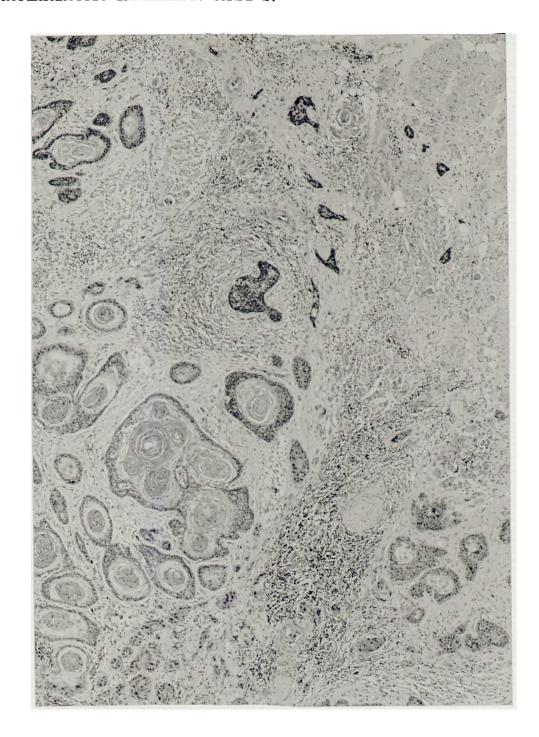
LYMPHOPLASMACYTIC INFILTRATE: GRADE 1.



A dense, continuous band of lymphocytes and plasma cells surrounds the advancing tumour front. Haematoxylin and eosin. Original magnification, x40.

Figure 6.5b

LYMPHOPLASMACYTIC INFILTRATE: GRADE 2.



Large patches of lymphocytes and plasma cells are seen at the advancing tumour front. Haematoxylin and eosin. Original magnification, x40.

Figure 6.5c
LYMPHOPLASMACYTIC INFILTRATE: GRADE 3.



A sparse and diffuse infiltrate of lymphocytes and plasma cells is seen at the advancing tumour front. Haematoxylin and eosin. Original magnification, x40.

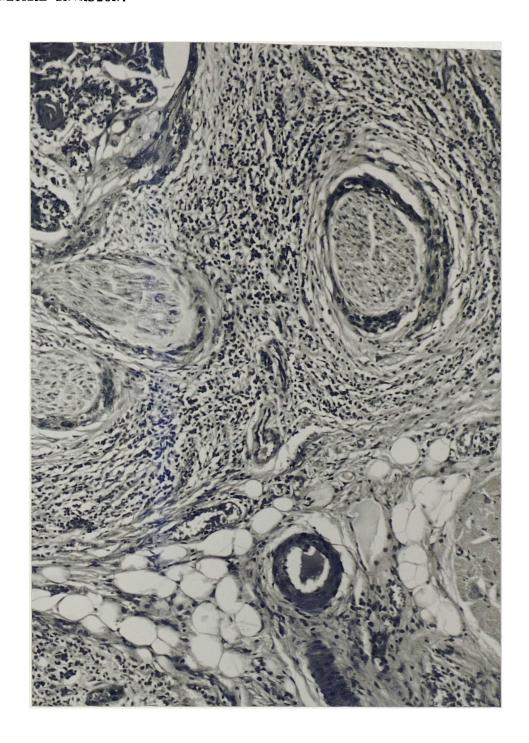
Figure 6.5d

LYMPHOPLASMACYTIC INFILTRATE: GRADE 4.



No lymphocytes or plasma cells are seen at the advancing tumour front. Haematoxylin and eosin. Original magnification, ${\bf x40}.$

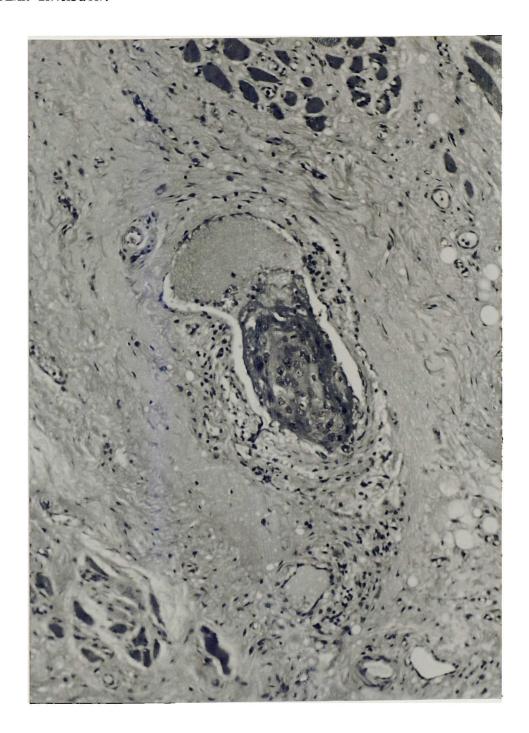
Figure 6.6
PERINEURAL INVASION.



Tumour cells are seen infiltrating perineural spaces. Haematoxylin and eosin. Original magnification, ${\tt x100}\,.$

Figure 6.7a

VASCULAR INVASION.



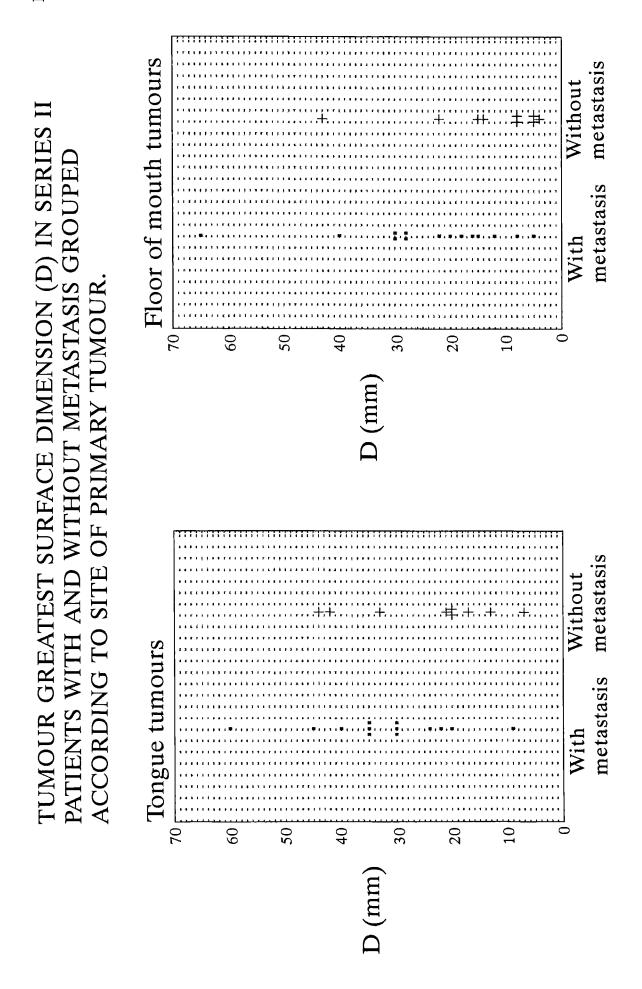
An embolus of tumour cells is seen within an endothelial-lined space. Haematoxylin and eosin. Original magnification, x100.

Figure 6.7b

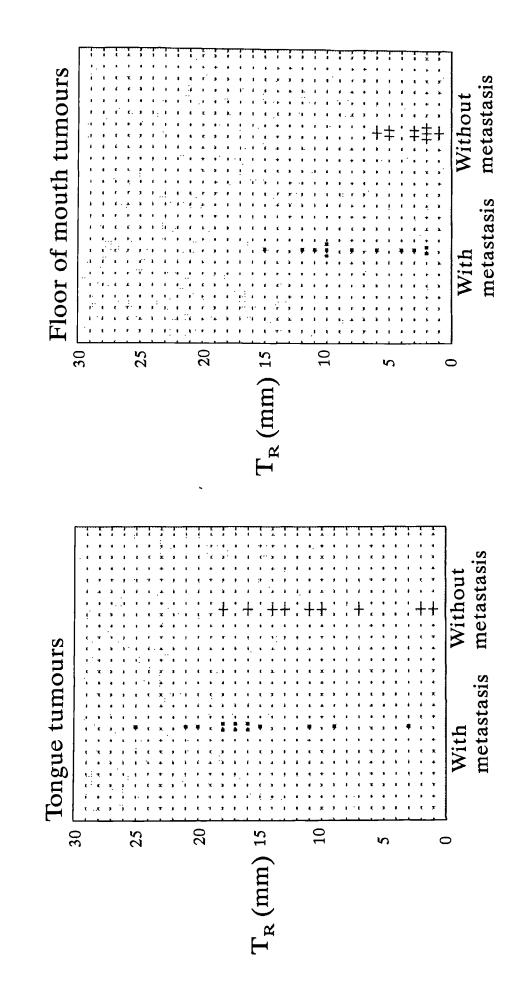
VASCULAR INVASION.



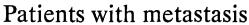
Tumour is seen invading a vessel (with a muscular media), with fibrin partially occluding the lumen. Haematoxylin and eosin. Original magnification, x100.

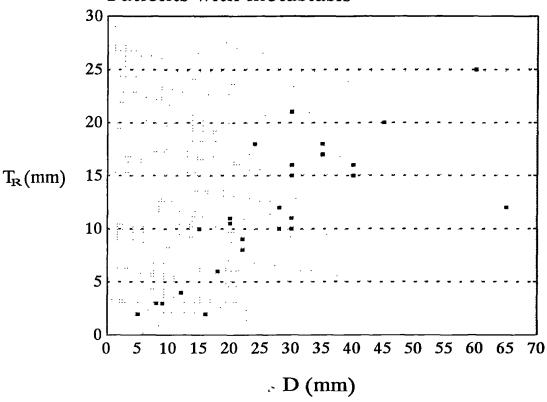


RECONSTRUCTED TUMOUR THICKNESS IN SERIES II PATIENTS WITH AND WITHOUT METASTASIS GROUPED ACCORDING TO SITE OF PRIMARY TUMOUR.

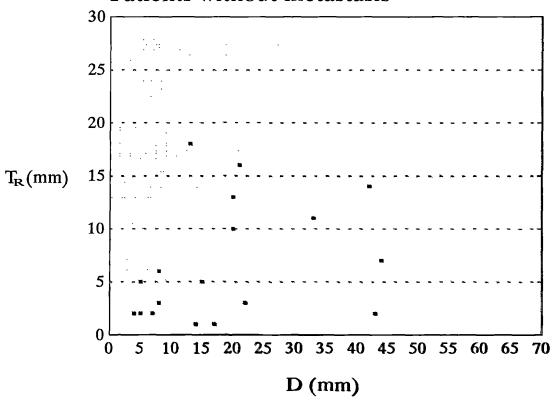


RELATIONSHIP BETWEEN RECONSTRUCTED TUMOUR THICKNESS (T_R) AND GREATEST SURFACE DIMENSION (D) IN SERIES II PATIENTS

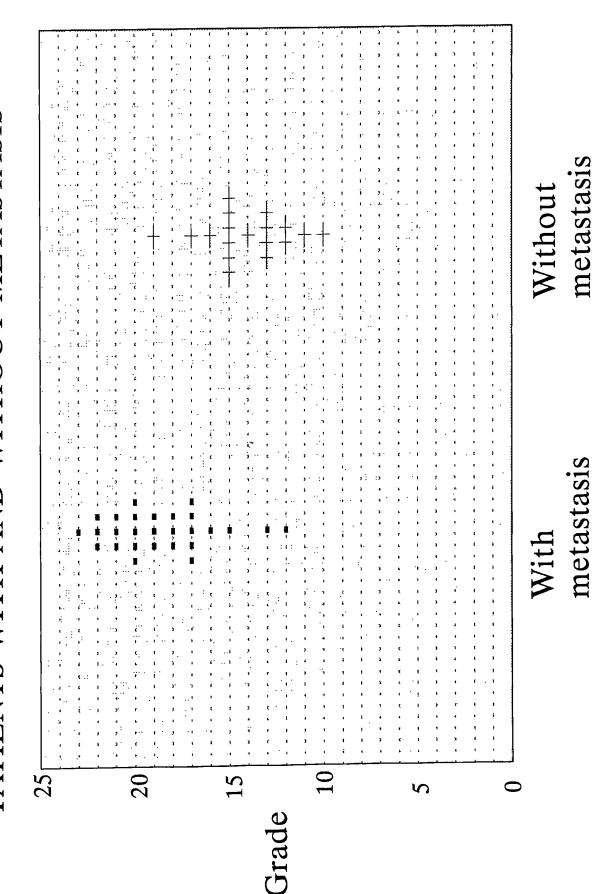


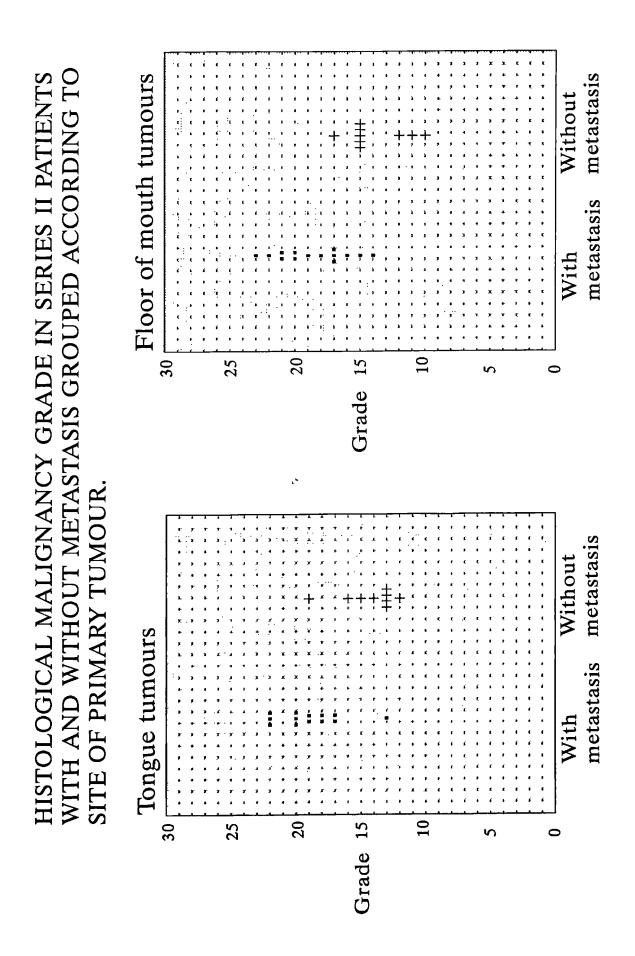


Patients without metastasis

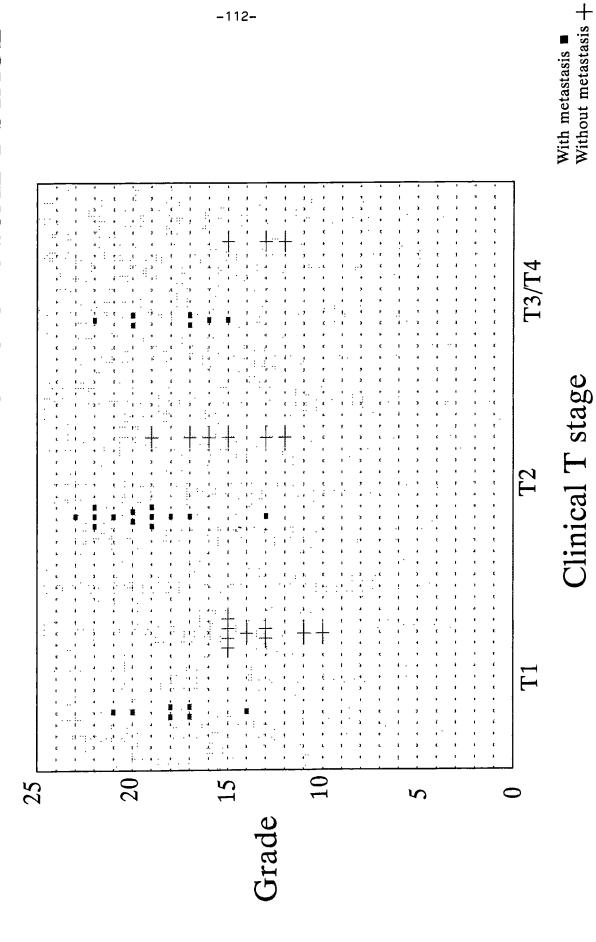


HISTOLOGICAL MALIGNANCY GRADE IN SERIES II PATIENTS WITH AND WITHOUT METASTASIS

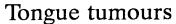


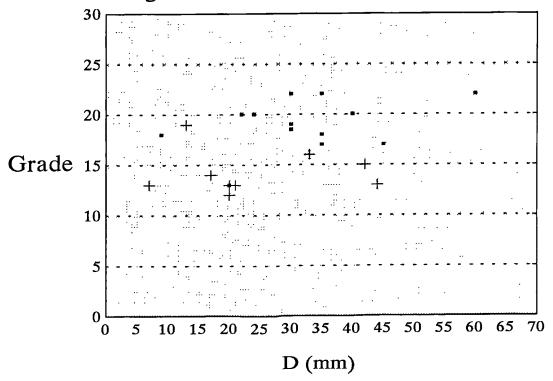


TUMOURS GROUPED ACCORDING TO CLINICAL T STAGE HISTOLOGICAL MALIGNANCY GRADE IN SERIES II

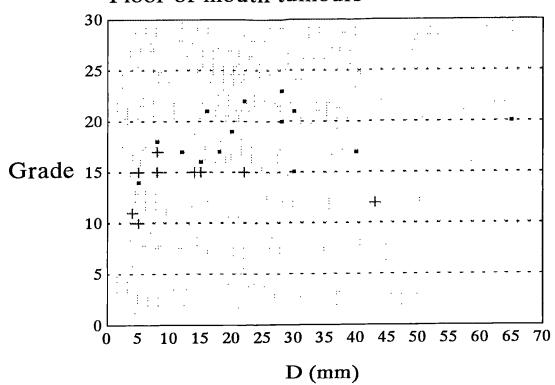


RELATIONSHIP BETWEEN HISTOLOGICAL MALIGNANCY GRADE AND GREATEST SURFACE DIMENSION (D) IN SERIES II PATIENTS.



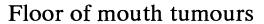


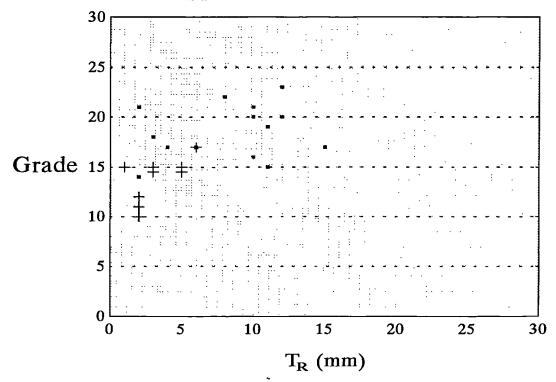
Floor-of-mouth tumours



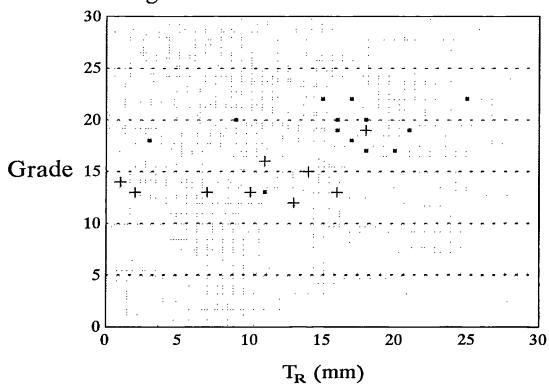
With metasasis ■
Without metastasis +

RELATIONSHIP BETWEEN HISTOLOGICAL MALIGNANCY GRADE AND RECONSTRUCTED TUMOUR THICKNESS IN SERIES II PATIENTS.





Tongue tumours



With metastasis ■
Without metastasis +

Figure 7.1

SCHEMATIC DIAGRAM OF THE INTERACTIVE IMAGE ANALYSIS SYSTEM FOR STEREOLOGICAL ESTIMATION OF MEAN NUCLEAR VOLUME BY POINT-SAMPLING OF LINEAR INTERCEPTS.

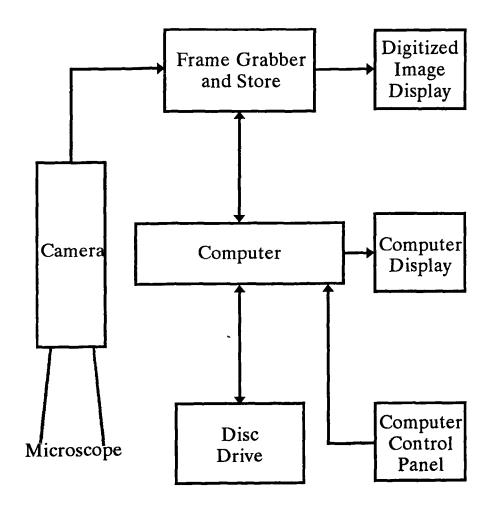
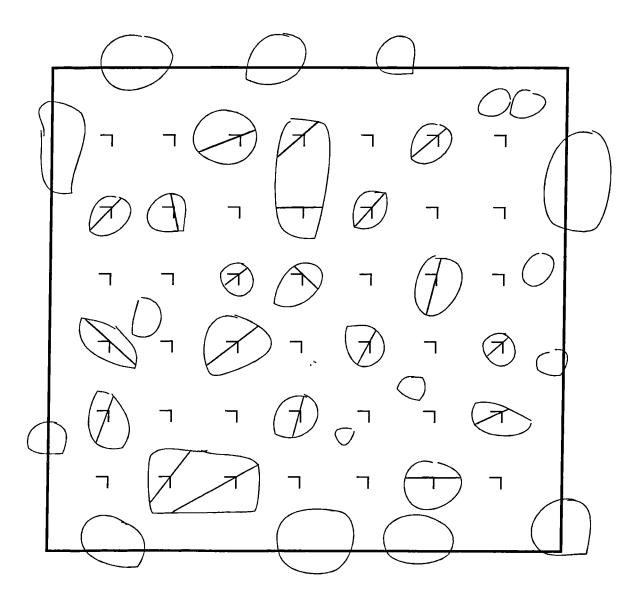
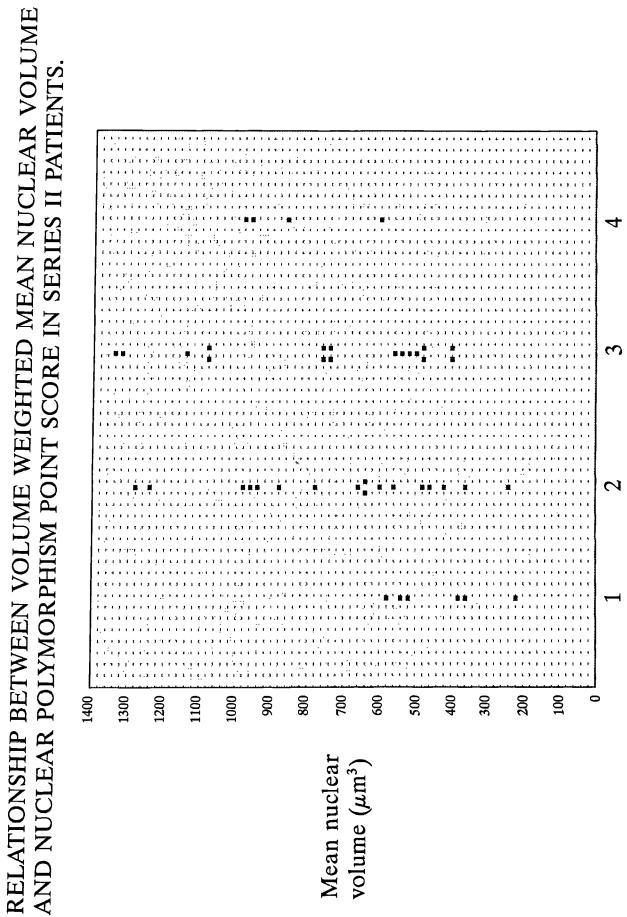


Figure 7.2

VERTICAL SECTION OF A SQUAMOUS CELL CARCINOMA SHOWING KERATINOCYTE NUCLEI WITH TEST GRID IN POSITION FOR MEASUREMENT OF OBSERVED POINT-SAMPLED INTERCEPT LENGTHS (1_0) .

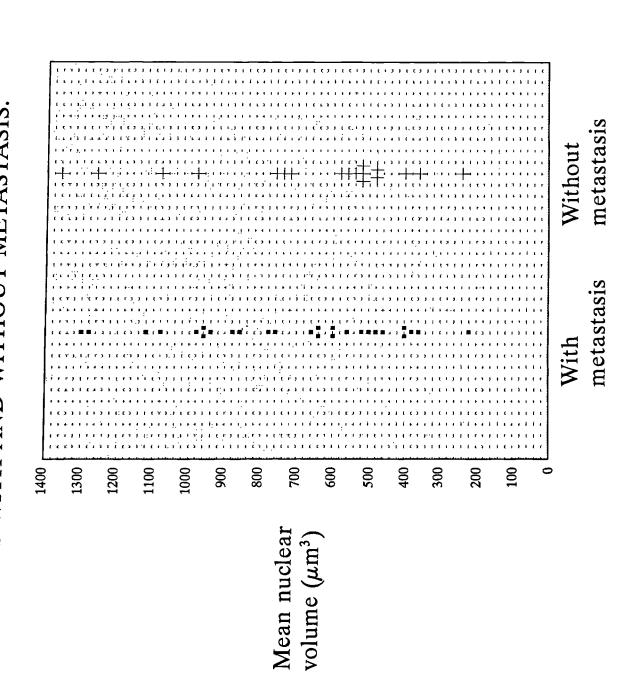


When a point hits a nuclear profile, a signal to generate a test-line is given, via the computer control panel. The test-line is generated in a three-dimensionally isotropic direction. Using the mouse part of the control panel, the cursors are then positioned at the intercept of the test-line and the nuclear profile. A second signal is given, and the observed intercept length, \mathbf{l}_0 (indicated by the red line), is then measured and recorded automatically. The cursor is moved to another nucleus, and the process is repeated until the intercept lengths of all point-sampled nuclei have been measured.



Nuclear polymorphism point score

VOLUME-WEIGHTED MEAN NUCLEAR VOLUME IN SERIES II PATIENTS WITH AND WITHOUT METASTASIS.



Section 3.

APPENDICES.

- 1. Appendix 1.
- 2. Appendix 2.
- 3. Appendix 3.

Appendix 1.

AMERICAN JOINT COMMITTEE ON CANCER (1988) THM CLASSIFICATION DEFINITIONS FOR STAGING CANCERS OF THE LIP AND ORAL CAVITY

Primary Tumour (T)

- T1 Tumour 2cm. or less in greatest dimension.
- T2 Tumour more than 2cm. but not more than 4cm. in greatest dimension.
- T3 Tumour more than 4cm. in greatest dimension.
- T4 Tumour invades adjacent structures, e.g., through cortical bone, into deep (extrinsic) muscle of tongue, maxillary sinus, skin.

Lymph Node (N)

- NO No regional lymph node metastasis.
- N1 Metastasis in a single ipsilateral lymph node, 3cm. or less in greatest dimension.
- N2a Metastasis in a single ipsilateral lymph node more than 3cm. but not more than 6cm. in greatest dimension.
- N2b Metastasis in multiple ipsilateral lymph nodes, none more than 6cm. in greatest dimension.
- N2c Metastasis in bilateral or contralateral lymph nodes, none more than 6cm. in greatest dimension.
- N3 Metastasis in a lymph node more than 6cm. in greatest dimension.

Distant Metastasis (M)

- MO No distant metastasis.
- M1 Distant metastasis.

Stage	Groupin	<u>ıg</u>			
Stage	I	T1		NO	MO
Stage	II	T2		NO	MO
Stage	III	Т3		NO	MO
		T1		N1	MO
		T2		N1	MO
		T3		N1	ΜO
Stage	IV	T 4		И0	MO
_		T 4		N1	MO
	Any	T		N2	MO
	Any			N3	MΟ
	Any		Any	N	M1

Appendix 2.

NECK DISSECTIONS (OCTOBER 1989 - OCTOBER 1992)

For each neck dissection (ND), the laboratory reference number (P), the sex and age of the patient, the type of surgical procedure and the histological diagnosis are given.

ND1 (P27746), male, 64 years.

Radical dissection of right neck (unilateral procedure) in continuity with resection of residual T3 SCC, right oral tongue (pre-operative chemotherapy).

One metastatic mass: level II.

Four positive nodes: level I(1), level II(1), level III(1), level IV(1).

ND2 (P27791), female, 67 years.

Radical dissection of right neck (unilateral procedure) simultaneous with resection of T2 SCC, right buccal mucosa. One positive node: level I.

ND3 (P27864), female, 45 years. Suprahyoid dissection of left neck (bilateral procedure) in continuity with resection of T1 SCC, anterior floor of mouth. No evidence of metastasis.

ND4 (P27864), female, 45 years. Suprahyoid dissection of right neck (bilateral procedure) in continuity with resection of T1 SCC, anterior floor of mouth. No evidence of metastasis.

ND5 (P27893), male, 62 years. Supra-omohyoid dissection of left neck (unilateral procedure) simultaneous with resection of T2 SCC, left buccal mucosa. Two positive nodes: level I.

ND6 (P27940), male, 61 years.
Radical dissection of left neck (unilateral procedure). Previous resection of T4 SCC, left maxillary antrum.
One metastatic mass: level I.

ND7 (P28063), female, 51 years. Radical dissection of left neck (unilateral procedure). Previous resection of medullary carcinoma of thyroid. Metastatic thyroid medullary carcinoma.

ND8 (P28065), female, 72 years.
Functional dissection of left neck (unilateral procedure) in continuity with resection of recurrence of T1 SCC, left buccal mucosa (previous surgery).
No evidence of metastasis.

ND9 (P28125), male, 61 years. Radical dissection of left neck (unilateral procedure) in continuity with resection of T1 SCC, left oral tongue. Two positive nodes: level III (1), level IV (1). ND10 (P28158), male, 69 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T3 SCC, right retromolar trigone. No evidence of metastasis.

ND11 (P28180), female, 38 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of carcinoma ex pleomorphic adenoma, left parotid. Metastatic adenocarcinoma.

ND12 (P28329), female, 74 years.

Radical dissection of left neck (unilateral procedure). Previous resection of synchronous T1 SCCs, dorsum and undersurface of oral tongue and lower lip.

One positive node: level II.

ND13 (P28337), female, 67 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of T2 SCC, left oral tongue.
No evidence of metastasis.

ND14 (P28361), female, 67 years.

Suprahyoid dissection of right neck (unilateral procedure). Previous resection of T1 SCC, right buccal mucosa. One positive node: level I.

ND15 (P28373), male, 60 years.

Radical dissection of left neck (unilateral procedure). Clinically covert primary SCC.

One metastatic mass: level II.

ND16 (P28379), male, 50 years.

Radical dissection of right neck (unilateral procedure) in continuity with resection of T2 SCC, right oral tongue.

No evidence of metastasis.

ND17 (P28455), female, 83 years.

Radical dissection of left neck (unilateral procedure). Previous resection of SCC, skin of forehead.

One metastatic mass: level II.

Two positive nodes: parotid gland.

ND18 (P28510), male, 46 years.

Radical dissection of right neck (unilateral procedure) in continuity with resection of T2 SCC, right oral tongue. No evidence of metastasis.

ND19 (P28511), male, 68 years.

Functional dissection of left neck (bilateral procedure) simultaneous with resection of T2 SCC, lower lip.

No evidence of metastasis.

ND20 (P28511), male, 68 years.

Functional dissection of right neck (bilateral procedure) simultaneous with resection of T2 SCC, lower lip.
No evidence of metastasis.

ND21 (P28543), female, 68 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of residual T2 SCC, left oral tongue (pre-operative radiotherapy).

One positive node: level II.

ND22 (P28593), female, 55 years.

Supra-omohyoid dissection of right neck (unilateral procedure) in continuity with resection of T1 SCC, right oral tongue. No evidence of metastasis.

ND23 (P28641), male, 61 years.

Suprahyoid dissection of left neck (unilateral procedure) in continuity with resection of T1 SCC, left floor of mouth.

No evidence of metastasis.

ND24 (P28655), male, 52 years.

Suprahyoid dissection of right neck (unilateral procedure) in continuity with resection of T1 SCC, right floor of mouth.

No evidence of metastasis.

ND25 (P28745), female, 78 years.

Radical dissection of right neck (unilateral procedure). Previous resection of T2 SCC, right oral tongue.

One positive node: level II.

ND26 (P28774), male, 72 years.

Radical dissection of right neck (unilateral procedure). Previous resection of metachronous T1 SCC, left oral tongue and T2 SCC, right oropharynx and suprahyoid dissection of right neck.

One metastatic mass: level II-III.

ND27 (P28880), male, 58 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of primary intraosseous carcinoma, right mandible.

Two positive nodes: level II.

ND28 (P28892), male, 42 years.

Radical dissection of right neck (unilateral procedure). Previous resection of T1 SCC, right oral tongue. One positive node: level II.

ND29 (P28914), female, 49 years.

Radical dissection of left neck (bilateral procedure) in continuity with resection of recurrent adenocarcinoma of left submandibular gland.

Metastatic salivary adenocarcinoma.

ND30 (P28914), female, 49 years.

Functional dissection of right neck (bilateral procedure) simultaneous with resection of recurrent adenocarcinoma of left submandibular gland.

Metastatic salivary adenocarcinoma.

ND31 (P28956), male, 65 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of T2 SCC, left oral tongue.

One positive node: level I.

ND32 (P29037), male, 79 years.

Radical dissection of left neck (unilateral procedure) simultaneous with resection of malignant melanoma, skin of forehead.

Metastatic malignant melanoma.

ND33 (P29038), male, 64 years.

Radical dissection of left neck (unilateral procedure). Malignant lymphoma.

ND34 (P29072), female, 27 years.

Functional dissection of left neck (unilateral procedure). Previous resection of malignant melanoma, skin of forehead. Metastatic malignant melanoma.

ND35 (P29077), male, 45 years.

Radical dissection of left neck (bilateral procedure) in continuity with resection of residual T3 SCC, oropharynx (pre-operative chemotherapy and radiotherapy).

One metastatic mass: level II.

ND36 (P29077), male, 45 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of residual T3 SCC, oropharynx (preoperative chemotherapy and radiotherapy).

No evidence of metastasis.

ND37 (P29127), male, 51 years.

Suprahyoid dissection of left neck (bilateral procedure) in continuity with resection of T1 SCC, anterior floor of mouth.

One positive node: level I.

ND38 (P29127), male, 51 yeras.

Suprahyoid dissection of right neck (bilateral procedure) in continuity with resection of T1 SCC, anterior floor of mouth.

No evidence of metastasis.

ND39 (P29285), female, 68 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of T3 SCC, left oral tongue.

Four metastatic masses: level II(2), level III(1), level IV(1). One positive node: level III.

ND40 (P29316), female, 62 years.

Radical dissection of left neck (unilateral procedure). Previous resection of T1 SCC, left oral tongue.

One positive node: level II.

ND41 (P29317), female, 64 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of T2 SCC, left oral tongue.

No evidence of metastasis.

ND42 (P29362), male, 63 years.

Functional dissection of left neck (bilateral procedure) in continuity with resection of T2 SCC, anterior floor of mouth.

No evidence of metastasis.

ND43 (P29362), male, 63 years.

Supra-omohyoid dissection of right neck (bilateral procedure) simultaneous with resection of T2 SCC, anterior floor of mouth. No evidence of metastasis.

ND44 (P29435), male, 77 years.

Functional dissection of left neck (bilateral procedure) in continuity with resection of residual T4 SCC, floor of mouth (pre-operative chemotherapy).

One metastatic mass: level I.

Two positive nodes: level I(1), level II(1).

ND45 (P29435), male, 77 years.

Radical dissection of right neck (bilateral procedure) in continuity with resection of residual T4 SCC, floor of mouth (pre-operative chemotherapy).

Two metastatic masses: level I.

Three positive nodes: level II(2), level III(1).

ND46 (P29555), female, 68 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of T3 SCC, left lower alveolar ridge. No evidence of metastasis.

ND47 (P29756), male, 46 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T1 SCC, right oral tongue.

No evidence of metastasis.

ND48 (P29785), female, 47 years.

Functional dissection of right neck (unilateral procedure). Previous resection of T1 SCC, floor of mouth and bilateral suprahyoid dissections (ND3 and ND4).

No evidence of metastasis.

ND49 (P29786), male, 74 years.

Supra-omohyoid dissection of left neck (unilateral procedure) in continuity with resection of recurrence of T2 SCC, left retromolar trigone.

No evidence of metastasis.

ND50 (P29945), female, 78 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of recurrence of T1 SCC, right oral tongue (pre-operative radiotherapy).

No evidence of metastasis.

ND51 (P29866), male, 72 years.

Supra-omohyoid dissection of right neck (unilateral procedure) in continuity with resection of T4 SCC, right floor of mouth. One positive node: level II.

ND52 (P29902), male, 67 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of T3 SCC, left oral tongue. Two positive nodes: level II(1), level IV(1).

ND53 (P29975), female, 79 years.

Radical dissection of right neck (unilateral procedure) in continuity with resection of malignant melanoma, skin of cheek.

Metastatic malignant melanoma.

ND54 (P30010), male, 42 years.

Radical dissection of right neck (unilateral procedure). Previous resection of T2 SCC, lower lip.
One metastatic mass: level I.

ND55 (P30048), male, 49 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T4 SCC, right buccal mucosa. No evidence of metastasis.

ND56 (P30144), female, 69 years.

Functional dissection of left neck (unilateral procedure) simutaneous with resection of T1 SCC, left buccal mucosa. No evidence of metastasis.

ND57 (P30194), female, 70 years.

Radical dissection of right neck (unilateral procedure). Previous resection of SCC, skin of cheek.

Two metastatic masses: level I(1), parotid-level II(1).

Eight positive nodes: level I(1), level II(4), level III(2), level IV(1).

ND58 (P30298), male, 65 years.

Functional dissection of left neck (bilateral procedure) in continuity with resection of T4 SCC, anterior floor of mouth.

No evidence of metastasis.

ND59 (P30298), male, 65 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of T4 SCC, anterior floor of mouth. No evidence of metastasis.

ND60 (P30333), male, 47 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of residual T4 SCC, right oropharynx (preoperative chemotherapy and radiotherapy).

Intranodal keratin granulomata: level II.

ND61 (P30484), female, 64 years.

Functional dissection of left neck (unilateral procedure) simultaneous with resection of recurrence of T2 SCC, left oropharynx (pre-operative radiotherapy).

No evidence of metastasis.

ND62 (P30528), male, 63 years.

Radical dissection of left neck (unilateral procedure). Previous resection of T3 SCC, right oropharynx and radical dissection of right neck, followed by chemotherapy and radiotherapy.

Three metastatic masses: level II-IV(1), level IV(2).

One positive node: level V.

ND63 (P30639), male, 64 years.

Supra-omohyoid dissection of left neck (bilateral procedure) in continuity with resection of T2 SCC, right oral tongue. No evidence of metastasis.

ND64 (P30639), male, 64 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of T2 SCC, right oral tongue. Three positive nodes: level I(2), level II(1).

ND65 (P30676), female, 81 years.

Supra-omohyoid dissection of left neck (unilateral procedure) in continuity with resection of T4 SCC, left lower alveolar ridge. No evidence of metastasis.

ND66 (P30851), male, 52 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of residual T4 SCC, left retromolar trigone (pre-operative chemotherapy).

One metastatic mass: level II-V.

Seven positive nodes: level I(4), level II(2), level IV(1).

ND67 (P30938), male, 57 years.

Functional dissection of left neck (bilateral procedure) in continuity with resection of residual T4 SCC, oropharynx (pre-operative radiotherapy).

Extranodal keratin granulomata: level II.

ND68 (P30938), male, 57 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of residual T4 SCC, oropharynx (preoperative radiotherapy).

Intranodal keratin granulomata: level II.

ND69 (P30968), female, 68 years.

Functional dissection of left neck (unilateral procedure). Clinically covert primary SCC.

One metastatic mass: level I.

Two positive nodes: level I.

ND70 (P30983), male, 55 years.

Functional dissection of left neck (bilateral procedure) in continuity with resection of residual T4 SCC, left oropharynx (pre-operative chemotherapy and radiotherapy).

One positive node: level IV.

Intranodal keratin granulomata: level II.

ND71 (P30983), male, 55 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of residual T4 SCC, left oropharynx (preoperative chemotherapy and radiotherapy).

Two positive nodes: level IV.

ND72 (P31036), male, 46 years.

Radical dissection of left neck (bilateral procedure) in continuity with resection of T4 SCC, left floor of mouth.

Two metastatic masses: level II(1), level IV(1).

ND73 (P31036), male, 46 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of T4 SCC, left floor of mouth.

No evidence of metastasis.

ND74 (P31094), male, 52 years.

Radical dissection of right neck (unilateral procedure). Previous resection of T1 SCC, anterior floor of mouth and bilateral suprahyoid dissections (ND37 and ND38), followed by radiotherapy.

One metastatic mass: level I-II.

ND75 (P31146), male, 64 years.

Functional dissection of left neck (bilateral procedure) in continuity with resection of T2 SCC, anterior floor of mouth.

No evidence of metastasis.

ND76 (P31146), male, 64 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of T2 SCC, anterior floor of mouth.

No evidence of metastasis.

ND77 (P31161), female, 75 years.

Radical dissection of right neck (unilateral procedure). Previous resection of synchronous T1 SCCs, dorsum and undersurface of left oral tongue and lower lip and radical dissection of left neck (ND12), followed by radiotherapy.

One metastatic mass: level III-IV.

ND78 (P31188), male, 67 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of malignant melanoma, infra-auricular skim. Metastatic malignant melanoma.

ND79 (P31402), male, 58 years.

Functional dissection of left neck (bilateral procedure) in continuity with resection of residual T4 SCC, left lower alveolar ridge (preoperative chemotherapy).

No evidence of metastasis.

MD80 (P31402), male, 58 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of residual T4 SCC, left lower alweolar ridge (pre-operative chemotherapy). No evidence of metastasis.

ND81 (P31436), male, 53 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of residual T4 SCC, left oropharynx (pre-operative chemotherapy).

Two metastatic masses: level I(1), level II-III(1).

Two positive nodes: level III(2).

ND82 (P31471), male, 63 years.

Radical dissection of right neck in continuity with resection of Tl SCC, right oropharynx (unilateral procedure). Previous surgery and radiotherapy for two metachronous SCCs, supraglottic region and larynx.

One metastatic mass: level I.

ND83 (P31513), male, 51 years.

Supra-omohyoid dissection of left neck (bilateral procedure) in continuity with resection of T1 SCC, anterior floor of mouth.

No evidence of metastasis.

ND84 (P31513), male, 51 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of T1 SCC, anterior floor of mouth.

One metastatic mass: level I.

ND85 (P31539), male, 45 years.

Supra-omohyoid dissection of right neck (unilateral procedure) in continuity with resection of T1 SCC, right oral tongue.

No evidence of metastasis.

ND86 (P31584), female, 64 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of T1 SCC, left lower alveolar ridge.

No evidence of metastasis.

ND87 (P31624), female, 77 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of recurrent mucoepidermoid carcinoma, left oropharynx. Metastatic mucoepidermoid carcinoma.

ND88 (P31655), male, 58 years.

Functional dissection of right neck (unilateral procedure) simultaneous with resection of recurrence of SCC, right floor of mouth. Previous resection of Tl SCC, right floor of mouth and suprahyoid dissection of right neck.

No evidence of metastasis.

ND89 (P31730), male, 52 years.

Radical dissection of left neck (unilateral procedure). Previous resection of T1 SCC, anterior floor of mouth and bilateral suprahyoid dissections (ND37 and ND38), followed by radiotherapy and radical dissection of right neck (ND74).

One metastatic mass: level I-II.

One positive node: level II.

ND90 (P31794), female, 72 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T4 SCC, right lower alveolar ridge. No evidence of metastasis.

ND91 (P31900), female, 53 years.

neck (unilateral procedure) Supra-omohyoid dissection of left simultaneous with resection of adenoid cystic carcinoma, left buccal mucosa.

Metastatic adenoid cystic carcinoma.

ND92 (P31921), female, 79 years.
Supra-omohyoid dissection of left neck (bilateral procedure) in continuity with resection of T3 SCC, right oral tongue. No evidence of metastasis.

ND93 (P31921), female, 79 years.

Radical dissection of right neck (bilateral procedure) in continuity with resection of T3 SCC, right oral tongue. No evidence of metastasis.

ND94 (P31940), male, 68 years.

Radical dissection of left neck (bilateral procedure) in continuity with resection of T4 SCC, anterior floor of mouth. One positive node: level I(1).

ND95 (P31940), male, 68 years.

Supra-omohyoid dissection of right neck (bilateral procedure) in continuity with resection of T4 SCC, anterior floor of mouth. No evidence of metastasis.

ND96 (P32000), female, 63 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of T4 SCC, left floor of mouth. Two positive nodes: level I(1), level II(1).

ND97 (P32001), male, 53 years.

Functional dissection of right neck (unilateral procedure). Previous resection of adenoid cystic carcinoma of left parotid and radical dissection of left neck.

Metastatic papillary adenocarcinoma of thyroid.

ND98 (P32046), male, 50 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of T4 SCC, left lower alveolar ridge. No evidence of metastasis.

ND99 (P32088), female, 33 years.

Functional dissection of right neck (unilateral procedure) continuity with resection of T2 SCC, right buccal mucosa. No evidence of metastasis.

ND100 (P32124), female, 85 years.

Radical dissection of left neck (unilateral procedure). Previous resection of T2 SCC, left oral tongue and suprahyoid dissection of left neck, followed by radiotherapy.

One metastatic mass: level II-IV.

One positive node: level V.

ND101 (P32158), female, 76 years.

Functional dissection of left neck (unilateral procedure) simultaneous with resection of T4 SCC, left maxillary antrum.

One metastatic mass: level I.

ND102 (P32195), male, 78 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T4 SCC, right lower alveolar ridge. One positive node: level I.

ND103 (P32301), male, 39 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of malignant melanoma, infra-auricular skin.

Metastatic malignant melanoma.

ND104 (P32372), male, 74 years.

Functional dissection of left neck (bilateral procedure) in continuity with resection of T4 SCC, anterior lower alveolar ridge. No evidence of metastasis.

ND105 (P32372), male, 74 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of T4 SCC, anterior lower alveolar ridge. No evidence of metastasis.

ND106 (P32420), female, 63 years.

Supra-omohyoid dissection of left neck (bilateral procedure) in continuity with resection of T1 SCC, anterior floor of mouth. Two positive nodes: level I.

ND107 (P32420), female, 63 years.

Supra-omohyoid dissection of right neck (bilateral procedure) in continuity with resection of T1 SCC, anterior floor of mouth. One positive node: level I.

ND108 (P32449), male, 51 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of T4 SCC, left oropharynx.

Four metastatic masses: level II-V(1), II-III(2), level IV-V(1). Six positive nodes: level III(1), level IV(3), level V(2).

ND109 (P32450), male, 53 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of T4 SCC, left floor of mouth.

One positive node: level II.

ND110 (P32457), female, 65 years.

Radical dissection of right neck (unilateral procedure) in continuity with resection of T4 SCC, right oral tongue.

One metastatic mass: level I-IV.

Five positive nodes: level IV(1), V(4).

ND111 (P32483), female, 77 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T2 SCC, right floor of mouth. Two positive nodes: level II.

ND112 (P32531), female, 61 years.

Functional dissection of left neck (bilateral procedure) in continuity with resection of T2 SCC, left oral tongue.

Two positive nodes: level I.

ND113 (P32531), female, 61 years.

Supra-omohyoid dissection of right neck (bilateral procedure) in continuity with resection of T2 SCC, left oral tongue. No evidence of metastasis.

ND114 (P32541), female, 63 years.

Radical dissection of left neck (bilateral procedure) in continuity with resection of T2 SCC, left floor of mouth.

One metastatic mass: level II.

Three positive nodes: level I(2), level III(1).

ND115 (P32541), female, 63 years.

Supra-omohyoid dissection of right neck (bilateral procedure) in continuity with resection T2 SCC, left floor of mouth.

One positive node: level I.

ND116 (P32570), female, 61 years.

Supra-omohyoid dissection of left neck (unilateral procedure) simultaneous with resection of T1 SCC, left oropharynx. No evidence of metastasis.

ND117 (P32618), male, 43 years.

Supra-omohyoid dissection of left neck (bilateral procedure) in continuity with resection of T1 SCC, anterior floor of mouth.

No evidence of metastasis.

ND118 (P32618), male, 43 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of Tl SCC, anterior floor of mouth. One positive node: level I.

ND119 (P32689), female, 45 years.

Supra-omohyoid dissection of left neck (unilateral procedure) in continuity with resection of polymorphous low-grade adenocarcinoma, left retromolar trigone.

No evidence of metastasis.

ND120 (P32722), male, 66 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T2 SCC, right oral tongue. Five positive nodes: level I(1), level II(3), level III(1).

ND121 (P32723), male, 85 years.

Radical dissection of right neck (unilateral procedure). Previous resection of malignant melanoma of hard palate.
Metastatic malignant melanoma.

ND122 (P32738), male, 72 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of two synchronous T1 SCCs, lateral and undersurface of left oral tongue.

Two positive nodes: level II(1), level III(1).

ND123 (P32781), female, 39 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of recurrence of adenoid cystic carcinoma of right parotid.

No evidence of metastasis.

ND124 (P32812), male, 49 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T2 SCC, right oropharynx.

No evidence of metastasis.

ND125 (P32855), female, 43 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T4 SCC, right buccal mucosa. Three positive nodes: level I.

ND126 (P32856), male, 71 years.

Radical dissection of right neck (unilateral procedure). Previous resection of Tl SCC, right retromolar trigone.

One metastatic mass: level II.

One positive node: level II.

ND127 (P32889), female, 34 years.

Functional dissection of right neck (unilateral procedure). Previous resection of carcinoid tumour, right lung.
Metastatic carcinoid tumour.

ND128 (P32890), male, 79 years.

Supra-omohyoid dissection of left neck (unilateral procedure) in continuity with resection of T1 SCC, left retromolar trigone.

No evidence of metastasis.

ND129 (P32891), male, 63 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T2 SCC, right oral tongue. One positive node: level II.

ND130 (P32918), female, 59 years.

Radical dissection of left neck (unilateral procedure) in continuity with resection of carcinoma ex pleomorphic adenoma, left parotid.

Metastastic adenocarcinoma.

ND131 (P32956), female, 75 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T4 SCC, right lower alveolar ridge.

No evidence of metastasis.

ND132 (P32976), male, 74 years.

Supra-omohyoid dissection of right neck (unilateral procedure) in continuity with resection of Tl SCC, right floor of mouth.

No evidence of metastasis.

ND133 (P32978), male, 72 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of residual T2 SCC, left oral tongue (preoperative radiotherapy).

One positive node: level II.

ND134 (P33033), male, 37 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of T3 SCC, left oral tongue.

No evidence of metastasis.

ND135 (P33071), male, 71 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T2 SCC, right floor of mouth.

One metastatic mass: level II.

Four positive nodes: level II(1), level III(1), level IV(1), level V(1).

ND136 (P33091), male, 56 years.

Functional dissection of left neck (bilateral procedure) in continuity with resection of T2 SCC, left floor of mouth.

One positive node: level II.

ND137 (P33091), male, 56 years.

Supra-omohyoid dissection of right neck (bilateral procedure) in continuity with resection of T2 SCC, left floor of mouth.

No evidence of metastasis.

ND138 (P33129), female, 57 years.

Supra-omohyoid dissection of left neck (bilateral procedure) in continuity with resection of T1 SCC, anterior floor of mouth.

No evidence of metastasis.

ND139 (P33129), female, 57 years.

Supra-omohyoid dissection of right neck (bilateral procedure) in continuity with resection of Tl SCC, anterior floor of mouth.

No evidence of metastasis.

ND140 (P33169), female, 76 years.

Supra-omohyoid dissection of right neck (unilateral procedure) simultaneous with resection of recurrence of T1 SCC, right buccal mucosa (previous radiotherapy).

No evidence of metastasis.

ND141 (P33231), male, 63 years.

Radical dissection of left neck (bilateral procedure) simultaneous with resection of two synchronous T1 SCCs, left oropharynx.

One metastatic mass: level II.

ND142 (P33231), male, 63 years.

Functional dissection of right neck (bilateral procedure) simultaneous with resection of two synchronous T1 SCCs, left oropharynx. One metastatic mass: level II.

ND143 (P33243), female, 56 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of malignant melanoma, infra-auricular skin. No evidence of metastasis.

ND144 (P32273), male, 81 years.

Supra-omohyoid dissection of left neck (unilateral procedure) in continuity with resection of T2 SCC, left oral tongue. Three positive nodes: level I(1), level II(1), level III(1).

ND145 (P33285), male, 55 years.

Radical dissection of right neck (unilateral procedure). Previous resection of T1 SCC, right floor of mouth and suprahyoid dissection of right neck (ND24).

One metastatic mass: level III-IV.

ND146 (P33342), male, 54 years.

Radical dissection of left neck (bilateral procedure) in continuity with resection of residual T4 SCC, left retromolar trigone (preoperative chemotherapy and radiotherapy).

One metastatic mass: level I-V.

Five positive nodes: level IV(4), level V(1).

ND147 (P33342), male, 54 years.

Supra-omohyoid dissection of right neck (bilateral procedure) in continuity with resection of residual T4 SCC, left retromolar trigone (pre-operative chemotherapy and radiotherapy). No evidence of metastasis.

ND148 (P33353), male, 78 years.

Functional dissection of left neck (unilateral procedure). Previous resection of T4 SCC, right lower alveolar ridge and functional dissection of right neck (ND102).

One metastatic mass: level II-III.

ND149 (P33381), male, 63 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of T4 SCC, left maxillary antrum. No evidence of metastasis.

ND150 (P33463), female, 51 years.

Functional dissection of left neck (bilateral procedure) in continuity with resection of recurrence of carcinoma ex pleomorphic adenoma, right submandibular gland.

No evidence of metastasis.

ND151 (P33463), female, 51 years.

Radical dissection of right neck (bilateral procedure) in continuity with resection of recurrence of carcinoma ex pleomorphic adenoma, right submandibular gland.

No evidence of metastasis.

ND152 (P33485), male, 60 years.

Functional dissection of left neck (bilateral procedure) in continuity with resection of T2 SCC, left oral tongue.

Two positive nodes: level I.

ND153 (P33485), male, 60 years.

Functional dissection of right neck (bilateral procedure) simultaneous with resection of T2 SCC, left oral tongue.

No evidence of metastasis.

ND154 (P33527), female, 58 years.

Supra-omohyoid dissection of left neck (bilateral procedure) in continuity with resection of T1 SCC, anterior floor of mouth.

No evidence of metastasis.

ND155 (P33527), female, 58 years.

Supra-omohyoid dissection of right neck (bilateral procedure) in continuity with resection of T1 SCC, anterior floor of mouth.

No evidence of metastasis.

ND156 (P33566), male, 50 years.

Supra-omohyoid dissection of left neck (bilateral procedure) simultaneous with resection of T4 SCC, right oropharynx. No evidence of metastasis.

ND157 (P33566), male, 50 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of T4 SCC, right oropharynx.

Two positive nodes: level II.

ND158 (P33575), male, 43 years.

Radical dissection of left neck (bilateral procedure) in continuity with resection of T3 SCC, left oropharynx.

One positive node: level II.

ND159 (P33575), male, 43 years.

Functional dissection of right neck (bilateral procedure) simultaneous with resection of T3 SCC, left oropharynx.

No evidence of metastasis.

ND160 (P33617), male, 77 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of recurrence of T4 SCC, right lower alveolar ridge (previous radiotherapy).

No evidence of metastasis.

ND161 (P33671), male, 50 years.

Supra-omohyoid dissection of left neck (bilateral procedure) simultaneous with resection of T4 SCC, right lower alveolar ridge. No evidence of metastasis.

ND162 (P33671), male, 50 years.

Functional dissection of right neck (bilateral procedure) in continuity with resection of T4 SCC, right lower alveolar ridge. No evidence of metastasis.

ND163 (P33687), female, 74 years.

Functional dissection of right neck (unilateral procedure) simultaneous with resection of recurrence of T1 SCC, left buccal mucosa. Previous functional dissection of left neck (ND8), followed by radiotherapy.

Six positive nodes: level I(3), level II(1), level III(1), level IV(1).

ND164 (P33711), female, 61 years.

Radical dissection of right neck (unilateral procedure). Previous resection of T2 SCC, left oral tongue, functional dissection of left neck (ND112) and supra-omohyoid dissection of right neck (ND113). One metastatic mass: level IV-V.

ND165 (P33712), male, 55 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of T1 SCC, left buccal mucosa. No evidence of metastasis.

ND166 (P33713), male, 76 years.

Supra-omohyoid dissection of left neck (bilateral procedure) in continuity with resection of Tl SCC, anterior floor of mouth. No evidence of metastasis.

ND167 (P33713), male, 76 years.

Supra-omohyoid dissection of right neck (bilateral procedure) simultaneous with resection of T1 SCC, anterior floor of mouth. No evidence of metastasis.

ND168 (P33738), male, 77 years.

Radical dissection of left neck (unilateral procedure). Previous resection of SCC, skin of left temple. Three positive nodes: parotid gland.

ND169 (P33739), male, 64 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of T1 SCC, left floor of mouth. No evidence of metastasis.

ND170 (P33744), male, 56 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T1 SCC, right floor of mouth.

Nine positive nodes: level I(1), level II(2), level III(2), level IV(4).

ND171 (P00012), female, 49 years.

Supra-omohyoid dissection of left neck (unilateral procedure) in continuity with resection of T1 SCC, left floor of mouth. Three positive nodes: level I(2), level II(1).

ND172 (P00051), male, 48 years.

Supra-omohyoid dissection of left neck (bilateral procedure). Previous resection of malignant melanoma, lower lip.
No evidence of metastasis.

ND173 (P00051), male, 48 years.

Supraomohyoid dissection of right neck (bilateral procedure). Previous resection of malignant melanoma, lower lip.
No evidence of metastasis.

ND174 (P00074), male, 52 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T4.SCC, right retromolar trigone. Two positive nodes: level II.

ND175 (P00090), male, 42 years.

Functional dissection of left neck (unilateral procedure) in continuity with resection of T2 SCC, right oral tongue.
Two positive nodes: level II(1), level III(1).

ND176 (P00127), male, 72 years.

Functional dissection of right neck (unilateral procedure) in continuity with resection of T2 SCC, right oral tongue. Six positive nodes: level II(3), level III(2), level IV(1).

Appendix 3.

AMERICAN JOINT COMMITTEE ON CANCER (1988) THM CLASSIFICATION DEFINITIONS OF ANATOMICAL SITES

Oral Cavity

<u>Buccal Mucosa</u>. This includes all the mucous membrane lining the inner surface of the cheeks and lips from the line of contact of the opposing lips to the line of attachment of mucosa of the alveolar ridge (upper and lower) and pterygomandibular raphe.

Lower Alveolar Ridge. The ridge includes the alveolar process of the mandible and its covering mucosa, which extends from the line of attachment of mucosa in the buccal gutter to the line of free mucosa of the floor of the mouth. Posteriorly it extends to the ascending ramus of the mandible.

<u>Upper Alveolar Ridge</u>. The upper ridge is the alveolar process of the maxilla and its covering mucosa, which extends from the line of attachment of mucosa in the upper gingival buccal gutter to the junction of the hard palate. Its posterior margin is the upper end of the pterygopalatine arch.

Retromolar Trigone. This is the attached mucosa overlying the ascending ramus of the mandible from the level of the posterior surface of the last molar tooth to the apex superiorly, adjacent to the tuberosity of the maxilla.

Floor of Mouth. This is a semilunar space over the mylohyoid and hyoglossus muscles, extending from the inner surface of the lower alveolar ridge to the undersurface of the tongue. Its posterior boundary is the base of the anterior pillar of the tonsil.

Hard Palate. This is the semilunar area between the upper alveolar ridge and the mucous membrane covering the palatine process of the maxillary palatine bones. It extends from the inner surface of the upper alveolar ridge to the posterior edge of the palatine bone.

Oral Tongue. This is the freely mobile portion of the tongue that extends anteriorly from the line of circumvallate papillae to the undersurface of the tongue at the junction of the floor of the mouth. It is composed of four areas: the tip, the lateral borders, the dorsum and the undersurface.

Oropharynx

The boundaries of the oropharynx are:

Anterior wall (glosso-epiglottic area), formed by the tongue posterior to the vallate papillae (base of tongue or posterior third) and the vallecula.

Lateral wall, formed by the tonsil, the tonsillar fossa and faucial pillars, and the glosso-tonsillar sulci.

Posterior wall.

Superior wall, formed by the inferior surface of soft palate and the uvula.

Section 4.

BIBLIOGRAPHY.

Ackerman LV. Verrucous carcinoma of the oral cavity. Surgery 1948; 23: 670-8.

Ackerman LV. Oral cavity and oropharynx. In: Surgical Pathology. 4th edn. St. Louis: C.V.Mosby, 1968: 173-99.

Ali S, Tiwari RM, Snow GB. False-positive and false-negative neck nodes. Head Neck Surg 1985; 8: 78-82.

Altman DG. Comparing groups - Continuous data. In: Practical Statistics for Medical Research. London: Chapman and Hall, 1991a: 179-218.

Altman DG. Comparing groups - Categorical data. In: Practical Statistics for Medical Research. London: Chapman and Hall, 1991b: 229-76.

Altman DG. Relation between two continuous variables. In: Practical Statistics for Medical Research. London: Chapman and Hall, 1991c: 277-324.

Altman DG. Relation between several variables. In: *Practical Statistics for Medical Research*. London: Chapman and Hall, 1991d: 325-64.

Ambros RA, Trost RC, Campbell AY, Lambert WC. Prognostic value of morphometry in papillary thyroid carcinoma. *Hum Pathol* 1989; 20: 215-8.

American Cancer Society. Cancer Facts and Figures. New York: American Cancer Society, 1983.

American Joint Committee on Cancer. Purposes and principles of staging. In: Beahrs OH, Henson DE, Hutter RVP, Myers MH, eds. Manual for Staging of Cancer. 3rd edn. Philadelphia: J.B.Lippincott, 1988a: 3-5.

American Joint Committee on Cancer. Head and neck sites. In: Beahrs OH, Henson DE, Hutter RVP, Myers MH, eds. Manual for Staging of Cancer. 3rd edn. Philadelphia: J.B.Lippincott, 1988b: 27-38.

Anneroth G, Hansen LS. A methodologic study of histologic classification and grading of malignancy in oral squamous cell carcinoma. Scand J Dent Res 1984; 92: 448-68.

Anneroth G, Batsakis J, Luna M. Review of the literature and a recommended system of malignancy grading in oral squamous cell carcinomas. Scand J Dent Res 1987; 95: 229-49.

Arendorf TM, Walker DM, Kingdom RJ, Roll JRS, Newcombe RG. Tobacco smoking and denture wearing in oral candidal leukoplakia. Br Dent J 1983; 155: 340-3.

Arthur K, Farr HW. Prognostic significance of histologic grade in epidermoid carcinoma of the mouth and pharynx. *Am J Surg* 1972; 124: 489-92.

Arthur JF, Fenner ML. The influence of histological grading on prognosis in carcinoma of the tongue (a computer analysis of 299 cases). Clin Radiol 1966; 17: 384-96.

Aznavoorian S, Murphy AN, Stetler-Stevenson WG, Liotta LA. Molecular aspects of tumour cell invasion and metastasis. *Cancer* 1993; 71: 1368-83.

Baak JPA, Kurver PHJ, Snoo-Niewlaat, AJE de, Graef S de, Makkink B, Boon ME. Prognostic indicators in breast cancer - Morphometric methods. *Histopathology* 1982; 6: 327-39.

Baak JPA, Oort J. Obtaining quantitative data. In: A Manual of Morphometry in Diagnostic Pathology. Berlin: Springer-Verlag, 1983a: 15-26.

Baak JPA, Oort J. The case for morphometry in diagnostic pathology. In: A Manual of Morphometry in Diagnostic Pathology. Berlin: Springer-Verlag, 1983b: 2-5.

Baak JPA, Oort J. Introduction to the applications. In: A Manual of Morphometry in Diagnostic Pathology. Berlin: Springer-Verlag, 1983c: 48-51.

Baak JPA, Dop H van, Kurver PHJ, Hermans J. The value of morphometry to classic prognosticators in breast cancer. *Cancer* 1985; 56: 374-82.

Baddeley AJ, Gundersen HJG, Cruz-Orive LM. Estimation of surface area from vertical sections. J Microsc 1986; 142(3): 259-76.

Ballantyne AJ, McCarten AB, Ibanez ML. The extension of cancer of the head and neck through the peripheral nerves. *Am J Surg* 1963; 106: 651-67.

Ballantyne AJ. Significance of retropharyngeal nodes in cancer of the head and neck. Am J Surg 1964; 108: 500-4.

Barnes L, Johnson JT. Clinical and pathological considerations in the evaluation of major head and neck specimens resected for oral cancer. Part I. *Pathol Annu* 1986a; 21(1): 173-250.

Barnes L, Johnson JT. Clinical and pathological considerations in the evaluation of major head and neck specimens resected for oral cancer. Part II. Pathol Annu 1986b; 21(2): 83-110.

Bataini JP, Bernier J, Brugere J, Jaulerry C, Picco C, Brunin F. Natural history of neck disease in patients with squamous cell carcinoma of oropharynx and pharyngolarynx. *Radiother Oncol* 1985; 3: 245-55.

Batsakis JG. Squamous cell carcinomas of the oral cavity and the oropharynx. In: Tumors of the Head and Neck. Clinical and Pathological Considerations. 2nd edn. Baltimore: Williams and Wilkins, 1979: 144-76.

Bearhs OH, Barber KW. The value of radical dissection of structures of the neck in the management of carcinoma of the lip, mouth, and larynx. Arch Surg 1962; 85: 49-54.

Bennett SH, Futrell JW, Roth JA, Hove RC, Ketcham AS. Prognostic significance of histologic host response in cancer of the larynx or hypopharynx. Cancer 1971; 28: 1255-65.

Berg JW, Schottenfeld D, Ritier F. Incidence of multiple primary cancer, 3: Cancers of the respiratory and upper digestive system as multiple primary cancers. *J Natl Cancer Inst* 1970; 44: 263-74.

Berlinger NT, Tsakraklides V, Pollak K, Adams GL, Yang M, Good RA. Prognostic significance of lymph node histology in patients with squamous cell carcinoma of the larynx, pharynx or oral cavity. Laryngoscope 1976; 86: 792-803.

Binnie WH, Rankin KV. Epidemiology of oral cancer. In: Wright BA, Wright JM, Binnie WH, eds. Oral Cancer: Clinical and Pathological Considerations. Florida: CRC Press, 1988: 2-11.

Black RJ, Gluckman JL, Shumrick DA. Multiple primary tumours of the upper aerodigestive tract. Clin Otolaryngol 1983; 8: 277-81.

Bloom HJG, Richardson WW. Histological grading and prognosis in breast cancer. A study of 1409 cases of which 359 have been followed for 15 years. Br J Cancer 1957; 11: 359-77.

Blot WJ, McLaughlin JK, Winn DM, Austin DF, Greenberg RS, Preston-Martin SM, Beinstein L, Schoenberg JB, Stemhagen A, Fraumeni JF. Smoking and drinking in relation to oral and pharyngeal cancer. Cancer Res 1988; 48: 3282-97.

Bocca E, Pignataro O. A conservation technique in radical neck dissection. Ann Otol Rhinol Laryngol 1967; 76: 975-87.

Bocca E. Conservation neck dissection. Laryngoscope 1975; 85: 1511-5.

Bocca E, Pignataro O, Oldini C. Supraglottic laryngectomy: 30 years of experience. Ann Otol Rhinol Laryngol 1983; 92: 14-8.

Bocca E, Pignataro O, Oldini C, Cappa C. Functional neck dissection: An evaluation and review of 843 cases. Laryngoscope 1984; 94: 942-5.

Bouquot JE, Weiland LH, Kurland LT. Leukoplakia and carcinoma in situ synchronously associated with invasive oral/oropharyngeal carcinoma in Rochester, Minn., 1975-1984. *Oral Surg Oral Med Oral Pathol* 1988; 65: 199-204.

- Breslow A. Thickness, cross-sectional areas and depth of invasion in the prognosis of cutaneous melanoma. *Ann Surg* 1970; 172: 902-8.
- Breslow A. Tumour thickness, level of invasion and node dissection in stage I cutaneous melanoma. Ann Surg 1975; 182: 572-5.
- Briggs RJS, Pienta KJ, Hruban RH, Richtsmeier WJ. Nuclear morphometry for prediction of metastatic potential in early squamous cell carcinoma of the floor of mouth. Arch Otolaryngol Head Neck Surg 1992; 118: 531-3.
- Broders AC. Squamous cell epithelioma of the lip; a study of five hundred and thirty seven cases. JAMA 1920; 74: 656-64.
- Broders AC. Carcinoma. Grading and practical application. Arch Pathol 1926; 2: 376-81.
- Broders AC. The microscopic grading of cancer. Surg Clin North Am 1941; 21: 947-62.
- Browne MA, Gaydecki PA. High speed splice fitting with application to boundary analysis in low contrast digital image computers. *Biol Med* 1987; 17: 109-16.
- Bryne M, Koppang HS, Lilleng R, Stene T, Bang G, Dabelsteen E. New malignancy grading is a better prognostic indicator than Broders' grading in oral squamous cell carcinomas. *J Oral Pathol Med* 1989; 18: 432-7.
- Bryne M. Prognostic value of various molecular and cellular features in oral squamous cell carcinomas: A review. *J Oral Pathol Med* 1991; 20: 413-20.
- Bryne M, Neilsen K, Koppang HS, Dablesteen E. Reproducibility of two malignancy grading systems with reportedly prognostic value for oral cancer patients. *J Oral Pathol Med* 1991a; 20: 369-72.
- Bryne M, Neilsen K, Koppang HS, Dabelsteen E. Grading of nuclear polymorphism; higher prognostic value than stereologically assessed nuclear volume in oral carcinomas. *APMIS* 1991b; 99: 333-9.
- Bryne M, Koppang H, Lilleng R, Kjaerheim A. Malignancy grading of the deep invasive margins of oral squamous cell carcinomas has high prognostic value. *J Pathol* 1992; 166: 375-81.
- Bullough W. Mitotic activity in the tissues of dead mice, and in tissues kept in physiological salt solutions. Exp Cell Res 1950; 1: 410-20.
- Burman JF, Carter RL. Lysis of type-I collagen by squamous carcinomas of the head and neck. Int J Cancer 1985; 36: 109-16.
- Butler TP, Gullino PM. Quantitation of cell shedding into efferent blood of mammary adenocarcinoma. Cancer Res 1975; 35: 512-6.

Byers RM. Squamous cell carcinoma of the oral tongue in patients less than thirty years of age. Am J Surg 1975; 130: 475-8.

Byers RM, O'Brien J, Waxler J. The therapeutic and prognostic implications of nerve invasion in cancer of the lower lip. Int J Radiat Oncol Biol Phys 1978; 4: 215-7.

Byers RM. Modified neck dissection. A study of 967 cases from 1970 to 1980. Am J Surg 1985; 150: 414-21.

Byers RM, Wolf PF, Ballantyne AJ. Rationale for elective modified neck dissection. Head Neck Surg 1988; 10: 160-7.

Cachin Y, Gerard-Marchant R, Flamant R, Lazar P, Lacour J, Richard J. Preferential metastatic invasion of subdigastric lymph node group. In: Ruttiman A, ed. Progress in Lymphology, Proceedings of the International Symposium on Lymphology, Zurich, Switzerland, 1966. Stuggart: Georg Thieme, 1967: 287-90.

Cachin Y, Sancho-Garnier H, Micheau C, Marandas P. Nodal metastases from carcinomas of the oropharynx. Otolaryngol Clin North Am 1979; 12: 145-54.

Cady B, Catlin D. Epidermoid carcinoma of the gum. Cancer 1969; 23: 551-69.

Cahan WG, Castro EB, Rosen PP, Strong EW. Separate primary carcinomas of the esophagus and head and neck region in the same patient. Cancer 1976; 37: 85-9.

Callery CD, Spiro RH, Strong EW. Changing trends in the management of squamous carcinoma of the tongue. *Am J Surg* 1984; 148: 449-54.

Cancer Research Campaign. Atlas of Cancer Incidence in England and Wales. London: Cancer Research Campaign, 1991.

Carenfelt C, Eliasson K. Cervical metastases following radical neck dissection that preserved the spinal accessory nerve. *Head Neck Surg* 1980; 2: 181-4.

Carr RJ, Langdon JD. Multiple primaries in mouth cancer - The price of success. Br J Oral Maxillofac Surg 1989; 27: 394-9.

Carter RL, Pittam MR, Tanner NSB. Pain and dysphagia in patients with squamous carcinomas of the head and neck: The role of perineural spread. *J R Soc Med* 1982; 75: 598-606.

Carter RL, Clifford P. Spread of squamous carcinomas of the oral cavity. In: Henk JM, Langdon JD, eds. *Malignant Tumours of the Oral Cavity*. London: Edward Arnold, 1985: 32-42.

Carter RL, Barr LC, O'Brien CJ, Soo KC, Shaw HJ. Transcapsular spread of metastatic squamous cell carcinoma from cervical lymph nodes. Am J Surg 1985; 150: 495-9.

- Carter RL, Bliss JM, Soo KC, O'Brien CJ. Radical neck dissections for squamous carcinomas: Pathological findings and their clinical implications with particular reference to transcapsular spread. Int J Radiat Oncol Biol Phys 1987; 13: 828-32.
- Chu W, Strawitz JG. Results in suprahyoid, modified radical, and standard radical neck dissections for metastatic squamous cell carcinoma: Recurrence and survival. *Am J Surg* 1978; 136: 512-5.
- Clark WH, From L, Bernardino E, Mihm M. Histogenesis and biologic behaviour of primary human malignant melanoma of the skin. Cancer Res 1969; 29: 705-27.
- Close LG, Merkel M, Vuitch MF, Reisch J, Schaeffer SD. Computed tomographic evaluation of regional lymph node involvement in cancer of the oral cavity and oropharynx. *Head Neck* 1989; 11: 309-17.
- Collan Y, Torkbeli T, Pesonen E. Application of morphometry in tumour pathology. Anal Quant Cytol Histol 1987; 9: 79-88.
- Conley J. The significance of the cervical lymphatics in cancer of the head and neck. In: Ruttiman A, ed. Progress in Lymphology. Proceedings of the International Symposium on Lymphology, Zurich, Switzerland, 1966. Stuggart: Georg Thieme, 1967a: 274-5.
- Conley J. Biopsy in the head and neck. In: Ruttiman A, ed. Progress in Lymphology. Proceedings of the International Symposium on Lymphology, Zurich, Switzerland, 1966. Stuggart: Georg Thieme, 1967b: 294-9.
- Cooper JS, Pajek TF, Rubin P, Tupchong L. Second malignancies in patients who have head and neck cancers; incidence, effect on survival and implications for chemoprevention based on the RTOG experience. *Int J Radiat Oncol Biol Phys* 1989; 17: 449-56.
- Cox DR. Regression models and life-tables. Journal of the Royal Statistical Society 1972; 34: 187-220.
- Crile G. Excision of cancer of the head and neck. With special reference to the plan of dissection based on one hundred and thirty-two operations. *JAMA* 1906; 47: 1780-6.
- Crissman JD, Gluckman J, Whiteley J, Quenelle D. Squamous cell carcinoma of the floor of the mouth. Head Neck Surg 1980; 3: 2-7.
- Crissman JD, Liu WY, Gluckman JL, Cummings G. Prognostic value of histopathologic parameters in squamous cell carcinoma of the oropharynx. Cancer 1984; 54: 2995-3001.
- Crissman JD. Tumor-host interactions as prognostic factors in the histologic assessment of carcinomas. *Pathol Annu* 1986; 21(1): 29-52.

Cunningham MJ, Johnson JT, Myers EN, Schramm VL, Thearle PB. Cervical lymph node metastasis after local excision of early squamous cell carcinoma of the oral cavity. *Am J Surg* 1986; 152: 361-5.

Daftary DK, Murti PR, Bhonsle RB, Gupta PC, Mehta FS, Pindborg JJ. Risk factors and risk markers for oral cancer in high incidence areas of the world. In: Johnson NW, ed. Oral Cancer: Detection of Patients and Lesions at risk. England: Cambridge University Press, 1991: 29-63.

Dallenbach F, Komitowski D. Digital picture analysis of borderline papillary serous cystadenomas of the ovary. In: Dallenbach-Hellweg G, ed. Ovarial Tumoren. Berlin: Springer-Verlag, 1982.

De Hoff RT. The determination of the size distribution of ellipsoidal particles from measurements made on random plane sections. Transactions of the American Institute of Mining and Metallurgical Engineers 1962; 224: 474-80.

De La Pava S, Pickren JW. On lymph node clearing as applied to head and neck tumours. In: Ruttiman A, ed. Progress in Lymphology, Proceedings of the International Symposium on Lymphology, Zurich, Switzerland, 1966. Stuggart: Georg Thieme, 1967: 290-2.

Delemarre JFM, van der Waal I. Clinical and histopathologic aspects of median rhomboid glossitis. *Int J Oral Surg* 1973; 2: 203-8.

De Santo LW, Beahrs OH. Modified and complete neck dissection in the treatment of squamous cells carcinoma of the head and neck. Surg Gynecol Obstet 1988; 167: 259-69.

Diamond DA, Berry SJ, Umbricht C, Jewett HJ, Coffey DS. Computerised image analysis of nuclear shape as a prognostic factor for prostatic cancer. *Prostate* 1982a; 3: 321-32.

Diamond DA, Berry SJ, Jewett HJ, Eggleston JC, Coffey DS. A new method to assess metastatic potential of human prostate cancer: Relative nuclear roundness. *J Urol* 1982b; 128: 729-34.

Di Troia JF. Nodal metastases and prognosis in carcinoma of the oral cavity. Otolaryngol Clin North Am 1972; 5: 333-42.

Dodd GD, Dolan PA, Ballantyne AJ, Ibanez ML, Chau P. The dissemination of tumors of the head and neck via the cranial nerves. Radiol Clin North Am 1970; 8: 445-61.

Donegan JO, Gluckman JL, Crissman JD. The role of suprahyoid neck dissection in the management of cancer of the tongue and floor of the mouth. Head Neck Surg 1982; 4: 209-12.

Donhuijsen K, Schmidt U, Hirche H, Beuningen D van, Budach V. Changes in mitotic rate and cell cycle fractions caused by delayed fixation. Hum Pathol 1990; 21: 709-14.

Droulias C, Whitehurst JO. The lymphatics of the tongue in relation to cancer. Am Surg 1976; 42: 670-4.

Easson EC, Russell MH. The Curability of Cancer at Various Sites. Baltimore: Williams and Wilkins, 1968: 25-8.

Elbrond O, Jorgensen K, Andersen AP. Carcinoma of the tongue. Acta Otolaryngol 1973; 75: 310-6.

Eneroth CM, Hjertman L, Moberger B. Squamous cell carcinomas of the palate. Acta Otolaryngol 1972; 73: 418-27.

Eneroth CM. Aspiration biopsy of head and neck tumours. Swedish Dent J 1973; 66: 177-9.

Eneroth CM, Moberger G. Histological malignancy grading in squamous cell carcinoma of the palate. *Acta Otolaryngol* 1973; 75: 293-5.

Ernst P. Uber das wachstum und die verbreitung bostariger geshwulste insbesondere des krebes in den lymphbahnen der nerven. Beitrage zur Pathologischen Anatomie und zer Allgemeinen Pathologie 1905; 7: 29-39.

Evans SJW. Survival of Patients with Oral Cancer. [Dissertation]. London: University of London, 1978.

Evans SJW, Langdon JD, Rapidis AD, Johnson NW. Prognostic significance of STNMP and velocity of tumour growth in oral cancer. Cancer 1982; 49: 773-6.

Farr HW, Arthur K. Epidermoid carcinoma of the mouth and pharynx, 1960-1964. *J Laryngol* 1972; 86: 243-53.

Farr HW, Goldfarb PM, Farr CM. Epidermoid carcinoma of the mouth and pharynx at Memorial Sloan Kettering Cancer Center. Am J Surg 1980; 140: 563-7.

Feind CR. The head and neck. In: Haagensen CD, Feind CR, Herter FP, Slanetz CA, Weinberg JA, eds. *The Lymphatics in Cancer*. Philadelphia: WB Saunders, 1972: 59-227.

Feinmesser R, Freeman JL, Noyek AM, Birt BD. Metastatic neck disease: A clinically, radiographically, pathological, correlative study. Arch Otolaryngol Head Neck Surg 1987; 113: 1307-10.

Fidler IJ, Balch CM. The biology of cancer metastasis and implications for therapy. Curr Probl Surg 1987; 24: 137-209.

Fidler IJ. Origin and biology of cancer metastasis. Cytometry 1989; 10: 673-80.

Fidler IJ. Cancer metastasis. Br Med Bull 1991; 47: 157-77.

Fisch UP, Sigel ME. Cervical lymphatic system as visualized by lymphography. Ann Otol Rhinol Laryngol 1964; 73: 869-82.

Fisher ER, Gregorio RM, Fisher B. The pathology of invasive breast cancer. Cancer 1975; 36: 1-85.

Fisher HR. Grading of biopsies of laryngeal carcinomas by multiple criteria. Can J Otolaryngol 1975; 4: 881-4.

Fletcher GH. Elective irradiation of subclinical disease in cancers of the head and neck. Cancer 1972; 29: 1450-5.

Fletcher GH. Clinical dose-response curves of human malignant epithelial tumours. Br J Radiol 1973; 46: 1-12.

Folberg R, Peter J, Gruman LM, Woolson RF, Jeng G, Montague PR, Moninger TO, Yi H, Moore KC. The morphologic characteristics of tumour blood vessels as a marker of tumour progression in primary uveal melanoma: a matched case control study. *Hum Pathol* 1992; 23: 1298-302.

Foss RD, Warnock GR, Clark WB, Graham SJ, Morton AL, Yunan ES. Malignant cyst of the lateral aspect of the neck: Branchial cleft carcinoma or metastasis? Oral Surg Oral Med Oral Pathol 1991; 71: 214-7.

Frable WJ, Frable MAS. Thin-needle aspiration biopsy. The diagnosis of head and neck tumors revisited. Cancer 1979; 43: 1541-8.

Franks LM. Structure and biological malignancy of tumors. In: Garratini S, Franchi G, eds. *Chemotherapy of Cancer Dissemination and Metastases*. New York: Raven Press, 1973: 71-81.

Friedell GH, Steiner G, Kristner RW. Prognostic value of blood vessel invasion in cervical cancer. Obstet Gynecol 1967; 39: 781-90.

Friedman HI, Cooper PH, Wanebo HJ. Prognostic and therapeutic use of microstaging of cutaneous squamous cell carcinoma of the trunk and extremities. *Cancer* 1985; 56: 1099-105.

Friedman M, Shelton VK, Mafee MM. Metastatic neck disease: Evaluation of computed tomography. Arch Otolaryngol 1984; 110: 443-7.

Friedman M, Mafee MM, Pacella BL, Strorigl TL, Dew LL, Toriumi DM. Rationale for elective neck dissection in 1989. Laryngoscope 1990; 100: 54-9.

Frierson HF, Cooper PH. Prognostic factors in squamous cell carcinoma of the lower lip. Hum Pathol 1986; 17: 346-54.

Fu YS, Hall TL. DNA measurements in tissue sections. Anal Quant Cytol Histol 1985; 7: 90-6.

Futrell JW, Hoye RC, Ketcham AS. Predicting survival in cancer of the larynx or hypopharynx. Am J Surg 1971; 122: 451-7.

Gilmore BB, Repola DA, Batsakis JG. Carcinoma of the larynx: Lymph node reaction patterns. Laryngoscope 1978; 88: 1333-8.

Gluckman JL, Crissman JD. Survival rates in 548 patients with multiple neoplasms of the upper aerodigestive tract. Laryngoscope 1983; 93: 1-4.

Goepfert H, Jesse RH. Squamous cell carcinoma cervical node metastases. In: Gates GA, ed. Current Therapy in Otolaryngology - Head and Neck Surgery. New Jersey: BC Decker, 1982: 224-9.

Goepfert H. Are we making any progress? Arch Otolaryngol 1984; 110: 562-3.

Goepfert H, Dichtel WJ, Medina J, Linberg RD, Luna MD. Perineural invasion in squamous cell skin carcinoma of the head and neck. Am J Surg 1984; 148: 542-7.

Graem N, Helweg-Larson K, Keiding N. Precision of histological grading of malignancy. *Acta Pathol Microbiol Scand* [A] 1980; 88: 307-17.

Graham A. Malignant epithelial tumours of the thyroid with special reference to invasion of blood vessels. Surg Gynecol Obstet 1924; 39: 781-90.

Grandi C, Alloisio M, Moglia D, Podrecca S, Sala L, Salvatori P, Molinari R. Prognostic significance of lymphatic spread in head and neck carcinomas: Therapeutic implications. *Head Neck Surg* 1985; 8: 67-73.

Greenspan D, Schiodt M, Greenspan JS, Pindborg JJ. Neoplasms. In: AIDS and the Mouth. Copenhagen: Munksgaard, 1990: 135-45.

Gundersen HJG, Jensen EB. Particle sizes and their distributions estimated from line- and point-sampled intercepts. Including graphical unfolding. *J Microsc* 1983; 131(3): 291-310.

Gundersen HJG, Jensen EB. Stereological estimation of the volume-weighted mean nuclear volume of arbitrary particles observed on random sections. *J Microsc* 1985; 138: 127-42.

Gundersen HJG. Stereology of arbitrary particles. A review of unbiased number and size estimators and the presentation of some new ones, in memory of William R. Thompson. J Microsc 1986; 143(1): 3-45.

Gundersen HJG, Bendtsen TF, Korbo L, Marcussen N, Moller A, Nielsen K, Nyengaard JR, Pakkenberg B, Sorensen FB, Vesterby A, West MJ. Some new, simple and efficient stereological methods and their use in pathological research and diagnosis. APMIS 1988a; 96: 379-94.

Gundersen HJG, Bagger P, Bendtsen TF, Evans SM, Korbo L, Marcussen N, Moller A, Nielsen K, Nyengaard JR, Pakkenberg B, Sorensen FB, Vesterby A, West MJ. The new stereological tools: Disector, fractionator, nucleator and point sampled intercepts and their use in pathological research and diagnosis. APMIS 1988b; 96: 857-881.

Haagensen CD. General anatomy of the lymphatic system. In: Haagensen CD, Feind CR, Herter FP, Slanetz CA, Weinberg JA, eds. *The Lymphatics in Cancer*. Philadelphia: WB Saunders, 1972: 1-13.

Hajek PC, Salmonowitz E, Turk R, Tscholakoff D, Kumpan W, Czembirek H. Lymph nodes of the neck. Evaluation with US. Radiology 1986; 158: 739-42.

Hansemann D. Ueber asymmetrische zelltheilung in epithelkrebsen und deren biologishe bedeutung. Virchows Arch A Pathol Anat Histopathol 1890; 119: 299-326.

Hansemann D. Ueber pathologische mitosen. Virchows Arch A Pathol Anat Histopathol 1891; 123: 356-70.

Hansemann D. Ueber die anaplasie der geschwulstzellen und die asymmetrische mitose. Virchows Arch A Pathol Anat Histopathol 1892; 129: 436-49.

Hansemann D. Studien uber die spezificitat, den altruismus und die anaplasie der zellen mit besonderer berucksichtigung der geschwulste. Berlin: A. Hirschwald, 1893.

Hansemann D. Die mikroskopische diagnose der bosartigen geschwulste. 2nd. ed. Berlin: A. Hirschwald, 1902.

Health Education Authority. Sensible Drinking. Here's How. London: Health Education Authority, 1989.

Helweg-Larson K, Graem N, Meistrup-Larsen K, Meistrup-Larsen U. Clinical relevance of histological grading of cancer of the larynx. Acta Pathol Microbiol Scand [A] 1978; 86: 499-504.

Henk JM, Langdon JD. Management of the primary tumour. In: Malignant Tumours of the Oral Cavity. London: Edward Arnold, 1985a: 158-85.

Henk JM, Langdon JD. Management of the regional lymph nodes. In: Malignant Tumours of the Oral Cavity. London: Edward Arnold, 1985b: 186-203.

Hepner GH. Tumor heterogeneity. Cancer Res 1984; 44: 2259-65.

Hibbert J, Marks NJ, Winter PJ, Shaheen OH. Prognostic factors in oral squamous cell carcinoma and their relation to clinical staging. Clin Otolaryngol 1983; 8: 197-203.

Hillsamer PJ, Schuller DE, McGhee RB, Chakeres D, Young DC. Improving diagnostic accuracy of cervical metastases with computed tomography and magnetic resonance imaging. Arch Otolaryngol Head Neck Surg 1990; 116: 1297-301.

Hindle I, Nally F. Oral cancer: A comparative study between 1962-67 and 1980-84 in England and Wales. Br Dent J 1991; 170: 15-20.

Hiratsuka H, Imamura M, Ishii Y, Kohama GI, Kikuchi K. Immunohistologic detection of lymphocyte subpopulations infiltrating in human oral cancer with special reference to its clinical significance. Cancer 1984; 53: 2456-66.

Holm LE, Lundquist PG, Silfversward C, Sobin A. Histological grading of malignancy in squamous cell carcinoma of the oral tongue. *Acta Otolaryngol* 1982; 94: 185-92.

Horak ER, Leek R, Klenek N, LeJeune S, Smith K, Stuart N, Greenall M, Stepniewska A, Harris AL. Angiogenesis, assessed by platelet/endothelial cell adhesion molecule antibodies, as indicator of node metastases and survival in breast cancer. *Lancet* 1992; 340: 1120-4.

Hordijk GJ, de Jong JMA. Synchronous and metachronous tumours in patients with head and neck cancer. J Laryngol Otol 1983; 97: 619-21.

Hsairi M, Luce D, Point D, Rodriguez J, Brugere J, Leclerc A. Risk factors for simultaneous carcinoma of the head and neck. *Head Neck* 1989; 11: 426-30.

Ildstad ST, Bigelow ME, Remensnyder JP. Squamous cell carcinoma of the tongue: A comparison of the anterior two-thirds of the tongue with its base. Am J Surg 1983; 146: 456-61.

International Agency For Research On Cancer. Tobacco habits other than smoking, betel quid and areca-nut chewing and some related nitrosamines. Monograph 37. Lyon: IARC, 1985.

International (Ludwig) Breast Cancer Study Group. Prognostic importance of occult axillary lymph node micrometastases from breast cancers. Lancet 1990; 335: 1565-8.

Jakobsson PA. Glottic carcinoma of the larynx. Factors influencing prognosis after radiotherapy. [Dissertation]. Sweden: University of Uppsala, 1973.

Jakobsson PA, Eneroth CM, Killander D, Moberger G, Martensson B. Histologic classification and grading of malignancy in carcinoma of the larynx. Acta Radiol 1973; 12: 1-8.

Jesse RH, Barkley HT, Lindberg RD, Fletcher GH. Cancer of the oral cavity. Is elective neck dissection beneficial? *Am J Surg* 1970; 120: 505-8.

Jesse RH, Lindberg RD. The efficacy of combining radiation therapy with a surgical procedure in patients with cervical metastasis from squamous cancer of the oropharynx and hypopharynx. *Cancer* 1975; 35: 1163-6.

Jesse RH, Ballantyne AJ, Larson D. Radical or modified neck dissection: A therapeutic dilemma. Am J Surg 1978; 136: 516-9.

Johnson JT, Barnes EL, Myers EN, Schramm VL, Borochovitz D, Sigler BA. The extracapsular spread of tumors in cervical node metastasis. Arch Otolaryngol 1981; 107: 725-9.

Johnson NW. The role of histopathology in diagnosis and prognosis of oral squamous cell carcinoma. *Proc Royal Soc Med* 1976; 69: 740-7.

Johnson NW. Histological and histochemical studies of oral cancer. Int Dent J 1977; 27: 25-34.

Johnson NW. Oro-facial Neoplasms: Global Epidemiology, Risk Factors and Recommendations for Research. Final Report of Working Group 2 of the Commission on Oral Health, Research and Epidemiology of the Federation Dentaire Internationale. Technical Report 36. London: FDI, 1990.

Jones JH, Coyle JI. Squamous carcinoma of the lip: a study of the interface between neoplastic epithelium and the underlying mesenchyma. *J Dent Res* 1969; 48(suppl): 702-8.

Kalnins IK, Leonard AG, Sako K, Razack MS, Shedd DP. Correlation between prognosis and degree of lymph node involvement in carcinoma of the oral cavity. Am J Surg 1977; 134: 450-4.

Keim WF, Lowenberg S. Marginal mandibulectomy in the treatment of carcinoma of the floor of mouth. Laryngoscope 1970; 80: 1566-79.

Keszler A, Cabrini RL. Histometric study of leukoplakia, lichen planus and carcinoma in situ of oral mucosa. *J Oral Pathol* 1983; 12: 330-5.

Kloppel G, Lingenthal G, Von Bulow M, Kern HF. Histological and fine structural features of pancreatic ductal adenocarcinomas in relation to growth and prognosis: Studies in xenografted tumours and clinicopathological correlation in a series of 75 cases. Histopathology 1985; 9: 841-56.

Kodama Y, Inokuchi K, Soejima K. Growth patterns and prognosis in early gastric carcinoma. Cancer 1983; 51: 320-9.

Kotwall C, Sako K, Razack MS, Rao U, Bakamjian V, Shedd DP. Metastatic patterns in squamous cell carcinoma of the head and neck. Am J Surg 1987; 154: 439-42.

Kotwall C, Razack MS, Sako K, Rao U. Multiple primary cancers in squamous cell cancer of head and neck. *J Surg Oncol* 1989; 40: 97-9.

Krause CJ, Lee GJ, McCabe BF. Carcinoma of the oral cavity. Arch Otolaryngol 1973; 97: 354-8.

Kreicbergs A, Slezak E, Soederberg G. The prognostic significance of different histomorphologic features in chondrosarcoma. Virchows Arch A Pathol Anat Histopath 1981; 390: 1-10.

Krogh P, Hald B, Holmstrup P. Possible mycological etiology of oral mucosal cancer: Catalytic potential of infecting Candidal albicans and other yeasts in production of N-nitrosobenzyl methylamine. Carcinogenesis 1987; 8: 1543-8.

Langdon JD, Harvey PW, Rapidis AD, Patel MF, Johnson NW. Oral cancer: The behaviour and response to treatment of 194 cases. *J Maxillofac Surg* 1977; 5: 221-37.

Langdon JD, Rapidis AD. Oral cancer and sex - Why do females do better? J Maxillofac Surg 1979; 7: 177-81.

Langley FA. Quality control in histopathology and diagnostic cytology. *Histopathology* 1978; 2: 3-18.

Larson DL, Rodin AE, Roberts DK, O'Steen WK, Rapperport AS, Lewis SR. Perineural lymphatics: Myth or fact. Am J Surg 1966; 112: 488-92.

Larson MP, Steinberg GD, Brendler CB. Use of Ulex europaeus agglutinin I (UEA I) to distinguish vascular and "pseudovascular" invasion in transitional cell carcinoma of bladder with lamina propria invasion. *Mod Pathol* 1990; 3: 83-8.

Lee AKC, Delellis RA, Silverman ML. Lymphatic and blood vessel invasion in breast carcinoma: A useful prognostic indicator. *Hum Pathol* 1986; 17: 984-7.

Lee JG, Krause CJ. Radical neck dissection: Elective, therapeutic, and secondary. Ann Otol Rhinol Laryngol 1975; 101: 656-9.

Lennert K. The significance of the unspecific inflammatory reaction in the cervical lymphatic system. In: Ruttiman A, ed. Progress in Lymphology, Proceedings of the International Symposium on Lymphology, Zurich, Switzerland, 1966. Stuggart: Georg Thieme, 1967: 293-4.

Lever WF, Schaumburg-Lever G. Tumors and cysts of the epidermis. In: *Histopathology of the Skin*, 7th edn. Philadephia: J.B.Lippincott, 1990a: 523-77.

Lever, WF, Schaumburg-Lever G. Benign melanocytic tumors and malignant melanoma. In: *Histopathology of the Skin*, 7th edn. Philadephia: J.B.Lippincott, 1990b: 756-805.

Levin ML, Goldstein H, Gerhardt PR. Cancer and tobacco smoking. JAMA 1950; 143: 336-8.

- Lindberg R. Distribution of cervical lymph node metastases from squamous cell carcinoma of the upper respiratory and digestive tracts. Cancer 1972; 29: 1446-9.
- Lindquist C. Risk factors of lip cancer: A critical evaluation based on epidemiological comparisons. Am J Public Health 1979; 69: 256-60.
- Lore JM. The neck. In: An Atlas of Head and Neck Surgery. 2nd. edn. Philadelphia: W.B.Saunders, 1973: 539-94.
- Lund C, Sogaard H, Elbrond O, Jorgensen K, Andersen AP. Epidermoid carcinoma of the lip. Histologic grading in the clinical evaluation. Acta Radiol Oncol Radiat Phys Biol 1975a; 14: 465-74.
- Lund C, Sogaard H, Elbrond O, Jorgensen K, Andersen AP. Epidermoid carcinoma of the tongue. Histologic grading in the clinical evaluation. Acta Radiol Oncol Radiat Phys Biol 1975b; 14: 513-21.
- Lund C, Sogaard H, Jorgensen K, Elbrond O, Hjelm-Hansen M, Andersen AP. Histological grading of epidermoid carcinomas in the head and neck. Dan Med Bull 1977; 24: 162-6.
- Lyall D, Schetlin CF. Cancer of the tongue. Ann Surg 1952; 135: 489-96.
- McCullough MW, Million RR, Parsons JT, Mendenhall WM, Cassisi NJ. Treatment results for simultaneous primary squamous cell carcinomas of the head and neck. Laryngoscope 1988; 98: 79-82.
- MacFarlane GJ, Boyle P, Scully C. Rising mortality from cancer of the tongue in young Scottish males. Lancet 1987; 2(8564): 912.
- McGavran MH, Bauer WC, Ogura JH. The incidence of cervical lymph node metastases from epidermoid carcinoma of the larynx and their relationship to certain characteristics of the primary tumor. *Cancer* 1961; 14: 55-66.
- McGregor AD, MacDonald DG. Routes of entry of squamous cell carcinoma to the mandible. Head Neck Surg 1988; 10: 294-301.
- McGuirt WF, Matthews B, Koufman JA. Multiple simultaneous tumors in patients with head and neck cancer. A prospective, sequential panendoscopic study. Cancer 1982; 50: 1195-9.
- McKelvie P. Penetration of the cervical lymph node system by cancer. Proc Royal Soc Med 1976; 69: 409-11.
- McKenna DM, Holden H, Farrell RWR. Immunohistochemical detection of micrometastasis in cervical lymph nodes in head and neck cancer. *J Pathol* 1991; 164: 356.
- McQuarrie DG, Mayberg M, Ferguson M, Shons AR. A physiologic approach to the problems of simultaneous bilateral neck dissection. *Am J Surg* 1977; 134: 455-60.

Macchiarini P, Fontanini G, Hardin J, Squartini F, Angeletti CA. Relation of neovascularisation to metastasis of non-small cell lung cancer. Lancet 1992; 340: 145-6.

Maddox WA. Vicissitudes of head and neck cancer. Am J Surg 1984; 148: 428-32.

Marlowe FI, Goodman RS, Mobini J, Dave U. Cystic metastasis from occult tonsillar primary simulating branchiogenic carcinoma: the case for tonsillectomy as a "grand biopsy". Laryngoscope 1984; 94: 833-5.

Maitland NJ, Brombridge B, Cox MF, Crane IJ, Prime SS, Scully C. Detection of Human Papillomavirus genes in human oral tissue biopsies and cultures by polymerase chain reaction. Br J Cancer 1989; 59: 598-603.

Malicka K. Attempt at evaluation of defensive activity of lymph nodes on basis of microscopic and clinical studies in cases of laryngeal cancer. *Pol Med J* 1971; 10: 154-64.

Mancuso AA, Harnsberger HR, Muraki AS, Stevens MH. Computed tomography of cervical and retropharyngeal nodes: Normal anatomy, variants of normal, and applications in staging of cancer. Radiology 1983; 148: 709-23.

Marshall JR, Graham S, Haughey BP, Shedd D, O'Shea R, Brasure J, Wilkinson GS, West D. Smoking, alcohol, dentition and diet in the epidemiology of oral cancer. Eur J Cancer 1992; 28B(1): 9-15.

Martin H, Del Valle B, Ehrlich H, Cahan WG. Neck dissection. Cancer 1951; 4: 441-99.

Martin ME, Munster M, Sugarbaker ED. Cancer of the tongue. Arch Surg 1940; 41: 888-936.

Mashberg A, Garfinkel L, Harris S. Alcohol as a primary risk factor in oral squamous carcinoma. CA Cancer J Clin 1981; 31: 146-55.

Mattfeldt T, Schurmann G, Matthaei-Mauer U, Feichter G, Moller P. Morphometrische aquivalente des differenzierungs grades bei epithelialen tumoren und lymphomen. Verh Dtsch Ges Pathol 1988; 72: 343-48.

Mattson ME, Winn DM. Smokeless tobacco: Association with increased cancer risk. NCI Monogr 1989; 8: 13-6.

Medina JE, Byers RM. Supraomohyoid neck dissection: Rationale, indications, and surgical technique. *Head Neck* 1989; 11: 111-22.

Mendelson BC, Woods JE, Beahrs OH. Neck dissection in the treatment of carcinoma of the anterior two-thirds of the tongue. Surg Gynecol Obstet 1976; 143: 75-80.

Mendenhall WM, Parsons JT, Mendenhall NP, Brant TA, Stringer SP, Cassisi NJ, Million RR. Carcinoma of the skin of the head and neck with perineural invasion. *Head Neck* 1989; 11: 301-8.

Merino OR, Lindberg RD, Fletcher GH. An analysis of distant metastases from squamous cell carcinoma of the upper respiratory and digestive tracts. *Cancer* 1977; 40: 145-51.

Micheau C, Cachin Y, Caillou C. Cystic metastases in the neck revealing occult carcinoma of the tonsil: a report of six cases. Cancer 1974; 33: 228-33.

Mittal KR, Schwartz PE, Barwick KW. Architectural (FIGO) grading, nuclear grading and other prognostic indicators in stage I endometrial adenocarcinoma with identification of high risk and low risk groups. Cancer 1988; 61: 538-45.

Mohit-Tabatabai MA, Sobel HJ, Rush BF, Mashberg A. Relation of thickness of floor of mouth stage I and II cancers to regional metastasis. Am J Surg 1986; 152: 351-3.

Moller H. Changing incidence of cancer of the tongue, oral cavity and pharynx in Denmark. *J Oral Pathol Med* 1989; 18: 224-9.

Moore C, Flynn MB, Greenberg RA. Evaluation of size in prognosis of oral cancer. Cancer 1986a; 58: 158-62.

Moore C, Kuhns JG, Greenberg RA. Thickness as prognostic aid in upper aerodigestive tract cancer. Arch Surg 1986b; 121: 1410-4.

Morgan RF, Hirata RM, Jaques DA, Saunders JR. Value of contralateral supra-omohyoid dissections. Am J Surg 1983; 146: 439-40.

Moss MC, Browne MA, Howard CV, Joyner DJ. An interactive image analysis system for mean particle estimation using stereological principles. *J Microsc* 1983; 156: 79-90.

Mostofi FK. Pathology and spread of renal cell carcinoma. In: King JS, ed. *Renal Neoplasia*. Boston: Little, Brown and Company, 1967: 230-47.

Nahun AM, Bone RC, Davidson TM. The case for elective prophylactic neck dissection. Laryngoscope 1977; 87: 588-99.

Nathanson A, Agren K. Evaluation of some prognostic factors in small squamous cell carcinoma of the mobile tongue: A multicenter study in Sweden. *Head Neck* 1989; 11: 387-92.

Nicolson GL. Tumor cell instability, diversification and progression to the metastatic phenotype: From oncogene to oncofetal expression. Cancer Res 1987; 47: 1473-87.

Nielsen K, Colstrup H, Nilsson T, Gundersen HJG. Stereological estimates of nuclear volume correlated with histopathological grading and prognosis of bladder tumours. *Virchows Archiv B Cell Pathol* 1986; 52: 41-54.

Nielsen K, Orntoft T, Wolf H. Stereological estimates of nuclear volume in non-invasive bladder tumours correlated with recurrence pattern. Cancer 1989; 64: 2269-74.

Noone RB, Bonner H, Raymond S, Brown AS, Graham WP, Lehr HB. Lymph node metastases in oral carcinoma. A correlation of histopathology with survival. *Plast Reconstr Surg* 1974; 53: 158-66.

Norris HJ. Mitosis counting. Hum Pathol 1976; 7: 483-4.

Ogden GR. Second malignant tumours in head and neck cancer. Commoner than elsewhere. Br Med J 1991; 302: 193-4.

Ojo MA, MacDonald DG. Mitotic phase distribution in oral epithelial hyperplasia and neoplasia. *Histopathology* 1988; 13: 213-9.

Okamoto M, Ozeki S, Watanabe T, Iida Y, Tashiro H. Cervical lymph node metastasis in carcinoma of the tongue. Correlation between clinical and histopathological findings and metastasis. *J Craniomaxillofac Surg* 1988; 16: 31-4.

Parker RG, Enstrom JE. Second primary cancers of the head and neck following treatment of initial primary head and neck cancer. *Int J Radiat Oncol Biol Phys* 1988; 14: 561-4.

Parkin DM, Laara E, Muir CS. Estimates of the worldwide frequency of sixteen major cancers in 1980. Int J Cancer 1988; 41: 184-97.

Pernanen K. Validity of survey data on alcohol use. Res Adv Alc Drug Probl 1974; 1: 355-60.

Pindborg JJ, Kiaer J, Gupta PC. Studies in oral leukoplakias. Prevalence of leukoplakia among 10,000 persons in Lucknow, India, with special reference to use of tobacco and betel nut. *Bull WHO* 1967; 37: 109-16.

Pindborg JJ. Palate. In: Oral Cancer and Precancer. Bristol: John Wright & Sons, 1980: 77-84.

Platz H, Fries R, Hudec M, Tjoa AM, Wagner RR. The prognosic relevance of various factors at the time of first admission of the patient. *J Maxillofac Surg* 1983; 11: 3-12.

Ploton D, Menager M, Jeannesson P, Himber G, Pigeon F, Adnet JJ. Improvement in the staining and in the visualisation of the argyrophilic proteins of the nucleolar organiser region at the optical level. Histochem J 1986; 18: 5-14.

Poleksic S, Kalwaic HJ. Prognostic value of vascular invasion in squamous cell carcinoma of the head and neck. *Plast Reconstr Surg* 1978; 61: 234-40.

Probert JC, Thompson RW, Bagshaw MA. Patterns of spread of distant metastases in head and neck cancer. Cancer 1974; 33: 127-33.

Quinn CM, Wright NA. The clinical assessment of proliferation and growth in human tumours: evaluation of methods and applications as prognostic variables. *J Pathol* 1990; 160: 93-102.

Rabin RS, Johnson J, Claassen D. Identification of subsets of lymphocytes infiltrating head and neck tumour tissue. A preliminary report. Laryngoscope 1984; 94: 688-90.

Reddy CRRM, Prahlad D, Ramulu C. Incidence of oral cancer with particular reference to hard palate cancer in 1 million population in the district of Visakhapatnam. *Indian J Cancer* 1975; 12: 72-6.

Reifen E, Noyek AM, Mullen BM. Nuclear morphometry and stereology in nasopharyngeal carcinoma. Laryngoscope 1992; 102: 53-5.

Rhys Evans PH, Morgan D, Smallman LS. Preservation of neck dissection for histological study. *J Laryngol Otol* 1987; 101: 823-7.

Richard JM, Sancho-Garnier H, Micheau C, Saravane D, Cachin Y. Prognostic factors in cervical lymph node metastasis in upper respiratory and digestive tract carcinomas: Study of 1,713 cases during a 15-year period. Laryngoscope 1987; 97: 97-101.

Rigaut JP, Boysen M, Reith A. Karyometry of pseudostratified, metaplastic and dysplastic nasal epithelium by morphometry and stereology. 2. Automated image analysis (I.B.A.S.) of the basal layer of nickel workers. *Pathol Res Pract* 1985; 180: 151-60.

Ring AH, Sako K, Rao U, Razack MS, Reese P. Immunologic patterns of regional lymph nodes in squamous cell carcinoma of the floor of the mouth. Prognostic significance. *Am J Surg* 1985; 150: 461-5.

Rosai J. Lymph node dissection. In: Ackerman's Surgical Pathology. 7th edn. St.Louis: C.V.Mosby, 1989: 1913-5.

Rothman K. Alcohol. In: Fraumeni JF, ed. Persons at High Risk of Cancer: An Approach to Cancer Etiology and Control. New York: Academic Press, 1975: 38-59.

Rouviere H. Anatomy of the Human Lymphatic System. Michigan: Edwards Bros, Inc, 1938.

Safaii H, Azar HA. Keratin granulomas in irradiated squamous cell carcinoma of various sites. Cancer Res 1966; 26: 500-8.

Saltykov SA. The determination of the size distribution of particles in an opaque material from a measurement of the size distribution of their sections. In: *Proceedings of the Second International Congress for Stereology, Chicago*. Berlin: Springer-Verlag, 1967: 163-5.

Sancho H, Hauss G, Saravane D. Metastatic cervical adenopathies: study of the histologic clinical correlations and consequences on the prognosis. *Tumori* 1977; 63: 259-66.

Scholl P, Byers RM, Batsakis JG, Wolf P, Santini H. Microscopic cut-through of cancer in the surgical treatment of squamous carcinoma of the tongue. Prognostic and therapeutic implications. Am J Surg 1986; 152: 354-60.

Schottenfeld D, Ganit RC, Wynder EL. The role of alcohol and tobacco in multiple primary cancers of the upper digestive system, larynx and lung: A prospective study. Prev Med 1974; 3: 277-93.

Schrek R. Ultrastructure of blood lymphocytes from chronic lymphocytic and lymphosarcoma cell leukaemia. *J Natl Cancer Inst* 1972; 48: 51-63.

Schroeder HE, Munzel-Pedrazzoli S. Application of stereologic methods to stratified gingival epithelia. *J Microsc* 1970; 92(3): 179-98.

Schuller DE, McGuirt WF, McCabe BF, Young D. The prognostic significance of metastatic cervical lymph nodes. *Laryngoscope* 1980; 90: 557-70.

Schwab W. Szintigraphic study of the cervical lymphatic system. In: Ruttiman A, ed. Progress in Lymphology, Proceedings of the International Symposium on Lymphology, Zurich, Switzerland, 1966. Stuggart: Georg Thieme, 1967: 280-3.

Scully C. The immunology of cancer of the head and neck with particular reference to oral cancer. Oral Surg Oral Med Oral Pathol 1982; 53: 157-67.

Scully C. Viruses and oral squamous carcinoma. Eur J Cancer 1992; 28B(1): 57-9.

Sessions DG. Surgical pathology of cancer of the larynx and hypopharynx. Laryngoscope 1976; 86: 814-39.

Shabana AHM, El-Labban NG, Lee KW. Morphometric analysis of basal cell layer in oral premalignant white lesions and squamous cell carcinoma. *J Clin Pathol* 1987; 40: 454-8.

Shah JP, Cendon RA, Farr HW, Strong EW. Carcinoma of the oral cavity. Factors affecting treatment failure at the primary site and neck. Am J Surg 1976; 132: 504-7.

- Shah JP, Candela FC, Poddar AK. The patterns of cervical lymph node metastases from squamous carcinoma of the oral cavity. Cancer 1990; 66: 109-13.
- Shaha AR, Spiro RH, Shah JP, Strong EW. Squamous carcinoma of the floor of the mouth. Am J Surg 1984; 148: 455-9.
- Shaha AR, Hoover EL, Mitrani M, Marti JR, Krespi YP. Synchronicity, multicentricity, and metachronicity of head and neck cancer. Head Neck Surg 1988; 10: 225-8.
- Sharpe DT. The pattern of lymph node metastases in intra-oral squamous cell carcinoma. Brit J Plast Surg 1981; 34: 97-101.
- Shear M, Ichilcik E. Morphologic changes in lymph node deposits of oral squamous carcinoma. *Int J Oral Surg* 1973; 2: 1-12.
- Shear M, Hawkins DM, Farr HW. The prediction of lymph node metastases from oral squamous carcinoma. Cancer 1976; 37: 1901-7.
- Shingaki S, Saito R, Kawasaki T, Nakajima T. Recurrence of carcinoma of the oral cavity, oropharynx and maxillary sinus after radical neck dissection. *J Maxillofac Surg* 1985; 13: 231-5.
- Shingaki S, Suzuki I, Wakajima T. Evaluation of histopathologic parameters in predicting cervical lymph node metastasis of oral and oropharyngeal carcinomas. *Oral Surg Oral Med Oral Pathol* 1988; 66: 683-8.
- Shons AR, McQuarrie DG. Multiple primary epidermoid carcinomas of the upper aerodigestive tract. Arch Surg 1985; 120: 1007-9.
- Sicher H. Oral mucous membrane. In: Orban's Oral Histology and Embryology. 5th edn. St. Louis: C.V.Mosby, 1962: 220-71.
- Silverman S, Griffith M. Smoking characteristics of patients with oral carcinomas and the risk for second oral primary carcinoma. *J Am Dent Assoc* 1972; 85: 637-40.
- Silverman S, Migliorati CA, Lozada-Nur F, Greenspan D, Conant MA. Oral findings in people with or at high risk for AIDS: A study of 375 homosexual males. J Am Dent Assoc 1986; 112: 187-92.
- Skinner DB. En bloc resection for neoplasms of the esophagus and cardia. Thorac Cardiovasc Surg 1983; 85: 59-69.
- Skolnik EM, Yee KF, Friedman M, Golden TA. The posterior triangle in radical neck surgery. Arch Otolaryngol 1976; 102: 1-4.
- Slaughter DP, Southwick HW, Smejkal W. Field cancerization in oral stratified squamous epithelium. Cancer 1953; 6: 963-8.
- Snow GB, Annyas AA, Slooten EA Van, Bartelink H, Hart AAM. Prognostic factors of neck node metastasis. Clin Otolaryngol 1982; 7: 185-92.

Som PM. Detection of metastasis in cervical lymph nodes: CT and MR criteria and differential diagnosis. AJR 1992; 158: 961-9.

Son YH, Kapp DS. Oral cavity and oropharyngeal cancer in a younger population. Cancer 1985; 55: 441-4.

Sorensen FB. Objective histopathological grading of cutaneous malignant melanomas by stereological estimation of nuclear volume: prediction of survival and disease-free period. *Cancer* 1989; 63: 1784-98.

Sorensen FB, Bennedbaek O, Pilgaard J, Spaun E. Stereological estimation of nuclear volume and other quantitative histological parameters in the prognostic evaluation of supraglottic laryngeal squamous cell carcinoma. APMIS 1989; 97: 987-95.

Sorensen FB. Stereological estimation of the mean and variance of nuclear volume from vertical sections. *J Microsc* 1991; 162(2): 203-29.

Sorensen FB, Bichel P, Jakobsen A. Stereological estimates of nuclear volume in squamous cell carcinoma of the uterine cervix and its precursors. *Virchows Archiv* [A] 1991a; 418(3): 225-33.

Sorensen FB, Kristensen IB, Grymer F, Jakobsen A. DNA-level and sterological estimates of nuclear volume in stage I cutaneous malignant melanomas: a comparative study with analysis of prognostic impact. Am J Dermatopathol 1991b; 13(1): 1-9.

Spiro JD, Spiro RH, Shah JP, Sessions RB, Strong EW. Supra-omohyoid neck dissection - a critical assessment. *Am J Surg* 1988; 156: 286-9.

Spiro RH, Strong EW. Epidermoid carcinoma of the mobile tongue. Treatment by partial glossectomy alone. Am J Surg 1971; 122: 707-10.

Spiro RH, Strong EW. Epidermoid carcinoma of the oral cavity and oropharynx. Elective vs. therapeutic radical neck dissection as treatment. Arch Surg 1973; 107: 382-4.

Spiro RH, Strong EW. Surgical treatment of cancer of the tongue. Surg Clin North Am 1974; 54: 759-65.

Spiro RH, Alfonso AE, Farr HW, Strong EW. Cervical node metastasis from epidermoid carcinoma of the oral cavity and oropharynx. A critical assessment of current staging. *Am J Surg* 1974; 128: 562-7.

Spiro RW, Huvos AG, Wong GY, Spiro JD, Gnecco CA, Strong EW. Predictive value of tumor thickness in squamous carcinoma confined to the tongue and floor of the mouth. Am J Surg 1986; 152: 345-50.

Springall RJ, Rytina ERC, Millis RR. Incidence and significance of micrometastases in axillary lymph nodes detected by immunohistochemical techniques. *J Pathol* 1990; 160: 174

Srivastava A, Laidler P, Davies RP, Morgan K, Hughes LE. The prognostic significance of tumor vascularity in intermediate thickness (0.76-4.0mm. thick) skin melanoma. A quantitative histologic study. *Am J Pathol* 1988; 133: 419-23.

Stell PM, Green JR. Management of metastases to the lymph glands of the neck. Proc Royal Soc Med 1976; 69: 411-3.

Stell PM, Morton RP, Singh SD. Cervical lymph node metastases: The significance of the level of the lymph node. *Clin Oncol* 1983; 9: 101-7.

Stell PM, McCormick MS. Cancer of the head and neck: Are we doing any better? Lancet 1985; 2(8464): 1127.

Stenkvist B, Bengtsson E, Eriksson O, Jarkrans T, Nordin B, Westman-Naeser S. Correlation between cytometric features and mitotic frequence in human breast carcinoma. *Cytometry* 1981; 1: 287-91.

Stephenson TJ, Griffiths DWR, Mills PM. Comparison of Ulex europaeus I lectin binding and factor VIII-related antigen as markers of vascular endothelium in follicular carcinoma of the thyroid. Histopathology 1986; 10: 251-60.

Stephenson TJ. Quantitation of the nucleus. In: Underwood JCE, ed. Pathology of the Nucleus. Berlin: Springer-Verlag, 1990: 152-213.

Sterio DE. The unbiased estimation of number and sizes of arbitrary particles using the dissector. J Microsc 1984; 134: 127-36.

Stern WBR, Silver CE, Zeifer BA, Persky MS, Heller KS. Computed tomography of the clinically negative neck. *Head Neck* 1990; 12: 109-13.

Stoddart TG. Conference of cancer of the lip. J Can Med Assoc 1966; 90: 666-70.

Sugarbaker EV. Cancer metastasis: A product of tumor-host interactions. Curr Probl Cancer 1979; 3: 1-59.

Syrjanen SM, Syrjanen KJ, Happonen PR. Human Papillomavirus (HPV) DNA sequences in oral precancerous lesions and squamous cell carcinoma demonstrated by in situ hybridization. *J Oral Pathol* 1988; 17: 273-8.

Tanner NSB, Carter RL, Dalley VM, Clifford P, Shaw HJ. The irradiated radical neck dissection in squamous carcinoma: A clinicopathological study. Clin Otolarngol 1980; 5: 259-71.

Tannock IF. Tumour growth and cell kinetics. In: Tannock IF, Hill RP, eds. *The Basic Science of Oncology*. Oxford, England: Pergamon Press, 1987: 57-79.

- Tannock IF. Principles of cell profileration Cell kinetics. In: De Vita VT, ed. *Principles and Practice of Oncology*, 2nd edn. Philadelphia: Lippincott, 1988: 3-13.
- Toker C. Some observations on the deposition of metastatic carcinoma within cervical lymph nodes. Cancer 1963; 16: 364-74.
- Tytor M, Franzen G, Olofsson J, Brunk U, Nordenskjold B. DNA content, malignancy grading and prognosis in Tl and T2 oral cavity carcinomas. Br J Cancer 1987; 56: 647-52.
- Underwood JCE, Giri DD. Nucleolar organizer regions as diagnostic discriminants for malignancy. J Pathol 1988; 155: 95-9.
- Underwood JCE. Nuclear morphometry and grading in tumours. In: Pathology of the Nucleus. Berlin: Springer-Verlag, 1990: 1-15.
- Underwood JCE. Prognostic indices in epithelial neoplasms. In: Anthony PP, MacSween RNM, eds. Recent Advances in Histopathology. Number 15. Edinburgh: Churchill Livingstone, 1992: 17-36.
- Urist MM, O'Brien CJ, Soong SJ, Visscher DW, Maddox WA. Squamous cell carcinoma of the buccal mucosa: Analysis of prognostic factors. Am J Surg 1987; 154: 411-4.
- Valentine JA, Scott J, West CR, Hill CA. A histological analysis of the early effects of alcohol and tobacco usage on human lingual epithelium. *J Oral Pathol* 1985; 14: 654-65.
- Vandenbrouck C, Sancho-Garnier H, Chassagne D, Saravane D, Cachin Y, Micheau C. Elective versus therapeutic radical neck dissection in epidermoid carcinoma of the oral cavity. Results of a randomised clinical trial. Cancer 1980; 46: 386-90.
- Venables CW, Craft IL. Carcinoma of the tongue in early adult life. Br J Cancer 1967; 21: 645-50.
- Vikram B, Strong EW, Shah JP, Spiro R. Failure in the neck following multimodality treatment for advanced head and neck cancer. Head Neck Surg 1984a; 6: 724-9.
- Vikram B, Strong EW, Shah JP, Spiro R. Failure at distant sites following multimodality treatment for advanced head and neck cancer. Head Neck Surg 1984b; 6: 730-3.
- Vikram B, Strong EW, Shah JP, Spiro R. Second malignant neoplasms in patients successfully treated with multimodality treatment for advanced head and neck cancer. *Head Neck Surg* 1984c; 6: 734-7.
- Virchow R. Die Krankhaften Geschwulste. Berlin: Hirschwald, 1863.
- Vries N de, Waal I van der, Snow GB. Multiple primary tumours in oral cancer. Int J Maxillofac Surg 1986; 15: 85-7.

Vries N de, Lange G de, Drexhage HA, Snow GB. Immunoglobulin allotypes in head and neck cancer patients with multiple primary tumours. *Acta Otolaryngol* 1987a; 104: 187-91.

Vries N de, Drexhage HA, Waal LP de, Lange G de, Snow GB. Human leucocyte antigens and immunoglobulin allotypes in head and neck cancer patients with and without multiple primary tumours. *Cancer* 1987b; 60: 957-61.

Vries N de. The magnitude of the problem. In: Vries N de, Gluckman JL, eds. Multiple Primary Tumors of the Head and Neck. Stuttgart: Georg Thieme Verlag, 1990: 1-29.

Walsh A. Logit regression. In: Statistics for the Social Sciences with Computer Applications. New York: Harper and Row, 1990: 322-8.

Walter JB, Israel MS. Structure and effects of some common tumours. In: *General Pathology*. 5th edn. Edinburgh: Churchill Livingstone, 1979: 316-41.

Waterhouse JAH. Cancer Handbook of Epidemiology and Prognosis. Edinburgh: Livingstone, 1974.

Weibel ER. Stereological principles for morphometry in electron microscopic cytology. *Int Rev Cytol* 1969; 26: 235-302.

Weibel ER. Glossary of terms. In: Stereological Methods. Volume I. Practical Methods for Biological Morphometry. London: Academic Press, 1979a: 352-3.

Weibel ER. Introduction. What is stereology? In: Stereological Methods. Volume I. Practical Methods for Biological Morphometry. London: Academic Press, 1979b: 1-3.

Weidner N, Semple JP, Welch WR, Folkman J. Tumor angiogenesis and metastasis correlation in invasive breast cancer. N Engl J Med 1991; 324: 1-8.

Weiss L. Studies on cellular adhesion in tissue culture. VII. Surface activity and cell detatchment. Exp Cell Res 1964; 33: 277-99.

Weiss L, Mayhew E, Glaves Rapp D, Holmes JC. Metastatic inefficiency in mice bearing B16 melanomas. Br J Cancer 1982; 45: 44-50

Weymuller EA, Kiviat NB, Duckert LG. Aspiration cytology: An efficient and cost-effective modality. Laryngoscope 1983; 93: 561-4.

Wilkinson EJ, Hause L. Probability in lymph node sectioning. Cancer 1974; 33: 1269-74.

Willen R, Nathanson A, Moberger G, Anneroth G. Squamous cell carcinoma of the gingiva. Histological classification and grading of malignancy. Acta Otolaryngol 1975; 79: 146-54.

- Willen R, Willen H, Stendahl U. Invasive squamous cell carcinoma of the uterine cervix. Influence of vehicle osmolarity on biopsy quality. Acta Radiol 1983; 22: 257-61.
- Willis RA. Epidermoid carcinoma of the head and neck, with special reference to metastasis. *J Pathol Bacteriol* 1930; 33: 501-26.
- Willis RA. Modes of origin of tumors Solitary localised squamous cell growth of the skin. Cancer Res 1964; 4: 630-44.
- Willis RA. Metastasis via the lymphatics. In: The Spread of Tumours in the Human Body. 3rd edn. London: Butterworth & Co, 1973a: 18-32.
- Willis RA. Direct spread of tumours. In: The Spread of Tumours in the Human Body. 3rd edn. London: Butterworth & Co, 1973b: 1-17.
- Woolf N. The relationship of neoplastic cells with their environment: Tumour spread. In: *Cell*, *Tissue and Disease*. *The Basis of Pathology*. 2nd edn. England: Bailliere Tindall, 1986: 397-411.
- World Health Organisation. Definition of leukoplakia and related lesions: An aid to studies on oral precancer. Oral Surg Oral Med Oral Pathol 1978; 46: 518-39.
- Wright A, Shear M. Epithelial dysplasia immediately adjacent to oral squamous cell carcinoma. *J Oral Pathol* 1985; 14: 559-64.
- Wright BA, O'Brien JC, Liberman ZH, Bradfielf JS, Mennel RG. Treatment of oral cancer. In: Wright BA, Wright JA, Binnie WH, eds. Oral Cancer: Clinical and Pathological Considerations. Florida: CRC Press, 1988: 89-126.
- Wynder EL, Hultberg S, Jacobbson F. Environmental factors in cancer of the upper alimentary tract: A Swedish study with special reference to Plummer-Vinson (Paterson-Kelly) syndrome. *Cancer* 1957; 10: 470-82.
- Wynder EL, Mushinski MH, Spivak JC. Tobacco and alcohol consumption in relation to the development of multiple primary cancers. Cancer 1977; 40: 1872-8.
- Yamamoto E, Miyakawa A, Kohama G. Mode of invasion and lymph node metastasis in squamous cell carcinoma of the oral cavity. *Head Neck Surg* 1984; 6: 938-47.
- Zoller M, Goodman ML, Cummings CW. Guidelines for prognosis in head and neck cancer with nodal metastasis. Laryngoscope 1978; 88: 135-40.