

# Morbidity Related to the Groin Lymph Node Dissection for Vulvar Cancer

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## Abstract

**Objective:** To determine the incidence of morbidity following groin lymphadenectomy for vulvar cancer, to explore causal factors, and examine strategies to reduce morbidity.

**Method:** A retrospective analysis of clinical and histopathological data was conducted on patients treated for invasive cancer of the vulva at a tertiary hospital in Sydney, Australia, from 1987 to 2016.

**Results:** Some type of groin dissection was performed on 525 groins in 333 patients. Lymphocysts occurred in 36.6% of groins and was higher in patients having an inguino-femorallymph node dissection compared to those having groin node debulking, or a sentinel node procedure (42.5% versus 14.6% versus 0% respectively;  $p < 0.0001$ ). In multivariable analysis, no significant difference in lymphocyst incidence was observed between patients with or without a groin drain. Wound breakdown occurred in 8.2% and wound infection in 10.7% of groins. Lymphedema occurred in 31.6% of lower limbs. The number of nodes resected was the only factor significantly associated with all complications, but current smoking and increasing age also increased the risk of wound breakdown.

**Conclusion:** A more extensive lymph node dissection is a significant risk factor for lymphocyst formation, groin wound infection, groin wound breakdown, and lower limb lymphedema. Debulking of bulky positive lymph nodes rather than complete inguino-femorallymphadenectomy reduces the risk of all post-operative complications. Our incidence of groin wound breakdown was less than 10% despite resection of the saphenous vein in all cases. Preservation of all subcutaneous fat above Camper's fascia appears to be the most critical factor in wound healing.

## Introduction

The status of the groin lymph nodes is the most important prognostic factor for patients with vulvar cancer. Selected early vulvar cancers may be amenable to sentinel node biopsy, but many patients will require an inguino-femoral lymphadenectomy in order to adequately treat the groin nodes.

The use of a separate incision approach significantly improved wound healing and decreased post-operative hospital stay, but the long-term problem of lower limb lymphedema remained. Several attempts have been made to try to reduce the risk of lymphedema, including elimination of groin dissection in patients with 'micro-invasive' vulvar cancer [1], the performance of a superficial inguinal lymphadenectomy [2] and the use of primary groin irradiation. These approaches were shown to increase the incidence of groin recurrence [3-5].

The purpose of this study was to determine the incidence of short and long-term postoperative morbidity of groin node dissection in a large cohort of patients, to investigate causal factors, and to postulate possible strategies to further reduce this morbidity.

## Materials and Methods

### Study design

A retrospective observational single institutional study.

Following ethics approval obtained from the South Eastern Sydney Local Health District Human Research Ethics Committee (Reference Number 15/151), the medical records of 429 consecutive patients

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treated for primary invasive vulvar cancer at the Royal Hospital for Women in Sydney, between February 1987 and June 2016 were reviewed. Ninety-six patients were excluded as their groins were not surgically treated. The remaining 333 patients underwent either unilateral or bilateral inguino-femoral lymphadenectomy, groin node debulking, or a sentinel node procedure and were included in the analysis. Data retrieved from the medical records included age at diagnosis, body mass index (BMI), smoking status, co-morbidities, disease stage, tumour diameter, histologic type, histologic grade, primary treatment, adjuvant treatment, type of lymph node dissection, number of lymph nodes removed, intra-operative insertion of a groin drain, duration of drain use, post-operative groin wound infection, groin wound dehiscence/breakdown, lymphocyst formation, length of stay and hospital readmission. Follow up data on lymphedema and patient disease status was retrieved from the outpatient clinical files. All patients were staged according to the 2009 International Federation of Gynecology and Obstetrics (FIGO) staging system [6].

Lymphocyst formation was recorded if confirmed by an ultrasonic scan, or if fluid was drained from the groin. Groin wound breakdown

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was defined as opening of the wound requiring either wound packing, or a negative pressure dressing. Groin wound infection was defined as erythema or a purulent exudate necessitating the use of antibiotics. Chronic lower limb lymphedema was recorded if documented as clinically obvious (mild, moderate, severe) during routine follow up, or patient reported as requiring compression garments and lymphatic massage to manage.

Three forms of groin node resection were performed; (1) complete inguino-femoral lymphadenectomy (2) resection of bulky positive nodes and (3) sentinel node biopsy.

The technique for inguino-femoral lymphadenectomy was to make a linear incision down to Camper's fascia, 1 cm above the groin crease, extending from a line perpendicular to the pubic tubercle medially to about 2 cm medial to the anterior superior iliac spine laterally. Camper's fascia was incised, and the fat in the femoral triangle deep to the fascia was removed as inguinal lymph nodes. All subcutaneous fat was preserved. The femoral nodes were obtained by removing the fat beneath the cribriform fascia in the fossa ovalis, medial to the femoral vein. After 1991, the fascia lata was left intact, but previously it was removed, and a sartorius muscle transposition performed to protect the femoral vessels. The saphenous vein was removed routinely.

Patients with palpable groin nodes were treated by resection of bulky nodes and frozen section diagnosis. If metastatic disease was confirmed, only palpably enlarged nodes were removed. When sentinel node biopsy was performed, pre-operative lymphoscintigraphy was combined with intraoperative blue dye injection for nodal identification.

Groin suction drains were routinely used up until 2002, and then variably over subsequent years. They were removed when fluid production was less than 50 millilitres over 24 hours. All patients received one dose of prophylactic antibiotics pre-operatively and thrombotic prophylaxis post-operatively.

### Statistical Analysis

Risk factors for short and long-term complications were assessed with univariate analysis. Descriptive analysis was performed using the Statistical Package for the Social Sciences (SPSS) version 25 software (IBM Corp., Armonk, New York, USA) including frequencies and medians. Cross tabulations were performed to examine associations between two variables using Pearson's  $\chi^2$  test (SPSS), or the Cochran-Armitage trend test to assess linear trends using Stata Statistical Software 15 [7]. A p value of <0.05 was considered statistically significant.

To investigate the factors associated with groin morbidity in multivariable models, the lme4 package [8] in R [9] was used to fit a mixed-effects logistic regression model for each outcome. Patient factors (age, BMI, diabetic and smoking status) and treatment factors (number of nodes removed, groin drain insertion, radiotherapy) were included as fixed effects, with random intercepts to account for within-patient correlation. Odds ratios and their 95% confidence intervals for each fixed effect were calculated by exponentiating the parameter estimates and Wald confidence intervals produced by the model.

### Results

We included 333 eligible patients. Table 1 shows the characteristics of the study group. Among the 333 patients, 525 groins were dissected, 192 patients (57.7%) undergoing a bilateral procedure and 141 (42.3%) a unilateral procedure. Inguino-femoral lymphadenectomy was performed in 278 patients (79.7%) (416 groins), a nodal debulking

in 65 patients (18.6%) (103 groins), and a sentinel node biopsy in 6 patients (1.7%) (6 groins). The median number of nodes removed per groin was 9 for patients having an inguino-femoral lymphadenectomy, 3 for a nodal debulking and 2.3 for a sentinel node procedure.

Sixty-nine patients (20.7%) received adjuvant radiotherapy to the groins and pelvis, while 12 patients (3.6%) received primary radiotherapy to the vulva and both groins. All 12 patients underwent some form of groin node procedure prior to their radiotherapy.

Groin wound drains were used in 211 patients (63.4%) and 348 groins (66.3%), with the drain left in-situ for a median of 6 days (range 2 - 16). Overall median length of post-operative hospital stay was 13

Patient Characteristic	Study Group (n = 333)
<b>Age, years (Range 20 – 96)</b>	
Mean	67
Median	71
<b>Smoking status</b>	
Current	77 (23.1%)
Former	45 (13.5%)
Never	211 (63.2%)
<b>BMI (Range 14.6 – 54.7)</b>	
< 20	13 (3.9%)
20 - < 30	230 (69.1%)
30 – 35	66 (19.8%)
> 35	24 (7.2%)
<b>Diabetic</b>	
Insulin dependent	16 (4.8%)
Non-Insulin dependent	31 (9.3%)
<b>Histopathological sub-type</b>	
Squamous cell carcinoma	302 (90.7%)
Melanoma	10 (3%)
Adenocarcinoma	8 (2.4%)
Sarcoma	6 (1.8%)
Other	7 (2%)
<b>FIGO Stage 2009<sup>†</sup></b>	
1B	182 (54.6%)
1I	12 (3.6%)
111A (1)	39 (11.7%)
111A (11)	15 (4.5%)
111B (1)	5 (1.5%)
111B (11)	11 (3.3%)
111C	42 (12.6%)
1VA (1)	3 (0.9%)
1VA (11)	7 (2.1%)
1VB	5 (1.5%)
<b>Groin Radiotherapy</b>	
Neoadjuvant	12 (3.6%)
Adjuvant	69 (20.7%)

Table 1: Patient Characteristics  
BMI = Body mass index, FIGO = International Federation of Gynecology and Obstetrics.

<sup>†</sup> FIGO Staging not done on Melanoma (n = 10), or Neuro-endocrine tumours (n = 2).

days (range 2 - 65) and was significantly longer when a groin drain was used (14 days versus 10 days respectively,  $p = 0.005$ ). The median follow-up was 49 months (range 6 - 366 months). Twenty-two patients (6.6%) were excluded from the analysis for long term complications (lymphedema and recurrent lower limb cellulitis) due to follow up of less than 6 months. Eleven of these patients died within five months of surgery (4 of progressive disease), and 11 were lost to follow up.

### Short-term complications of the groin dissection

The commonest immediate post-operative complication was lymphocyst formation which occurred in 36.6% of the groins dissected (Table 2). There was no difference in lymphocyst incidence in groins having an inguino-femoral lymphadenectomy before 1991 when the fascia lata was resected compared to after 1991 when the fascia lata was preserved (39.4% vs 42.7% respectively,  $P = 0.4$ ). Lymphocyst formation was most strongly associated with a greater number of nodes removed ( $p = 0.0001$ ) (Table 3). When adjusted for other risk factors, the number of nodes removed remained statistically significant for lymphocyst formation ( $p = 0.0001$ ; OR 1.24 [95% CI 1.12-1.36] per node) (Table 4).

Univariate analysis indicated no difference in the incidence of lymphocyst formation when a groin drain was used. There was a bias in the indication for the use of drains, as they were more commonly used following an inguino-femoral lymphadenectomy (72.4%)

than following nodal debulking, (43.8%) ( $p < 0.001$ ). Use of a drain compared to no drain resulted in no significant difference in the incidence of lymphocyst formation for either an inguino-femoral lymphadenectomy (39.7% vs 48.7% respectively,  $p = 0.121$ ) or nodal debulking (17.4% vs 13.5% respectively,  $p = 0.647$ ) on univariate analysis. After adjusting for the number of nodes removed, patients having more nodes removed had a lower rate of lymphocyst formation with a groin drain, but this failed to reach statistical significance ( $p = 0.06$ ) (Table 4).

The next most common short-term complication was groin wound infection, which occurred in 10.7% of the groins dissected. This was more common in the groins having an inguino-femoral lymphadenectomy (11.3%) than a nodal debulking (7.8%), but the difference was not significant in univariate analysis ( $p = 0.4$ ) (Table 2). However, in multivariable analysis, increasing number of nodes removed was associated with an increased incidence of groin wound infection ( $p = 0.02$ ) (Table 4).

The least common short-term complication was groin wound breakdown, which occurred in 8.2% of groins dissected (Table 2). In univariate analysis, the factors significantly associated with a higher rate of groin wound breakdown were increasing number of nodes removed ( $p = 0.005$ ), current smoking ( $p = 0.02$ ) and obesity ( $p < 0.001$ ) (Table 3). On multivariable analysis, increasing age was also associated with groin wound breakdown ( $p = 0.02$ ; OR 1.74, [95% CI

Total Number of Groins 525 Total No of Patients = 333	No of groins (no of patients)	Complication per groin (per patient)	% per groin (% per patient)	P value <sup>†</sup>
<b>Lymphocyst</b>				
Inguino-femoral LND	416 (278)	177 (150)	42.5% (54.3%)	<.0001
Nodal debulking	103 (65)	15 (14)	14.6% (21.5%)	(<.0001)
Sentinel node	6 (6)	0 (0)		
<i>Incidence per patient</i>	333	164	49.2 %	
<b>Groin wound breakdown</b>				
Inguino-Femoral LND	416 (278)	39 (33)	9.4% (11.9)	0.1570
Nodal debulking	103 (65)	4 (4)	3.9% (3.8%)	(0.3946)
Sentinel node	6 (6)	0 (0)		
<i>Incidence per patient</i>	333	37	13%	
<b>Groin wound infection</b>				
Inguino-Femoral LND	416 (278)	47 (45)	11.3% (16.1%)	0.3713
Nodal debulking	103 (65)	8 (6)	7.8% (9.2%)	(0.3773)
Sentinel node	6 (6)	1 (1)	16.6%	
<i>Incidence per patient</i>	333	52	15.6%	
<b>Lymphedema</b>				
Inguino-Femoral LND	392 (262)	137 (113)	35% (43.1%)	0.0032
Nodal debulking	92 (58)	18 (13)	19.6% (22.4%)	(0.0025)
Sentinel node	6	0	0%	
<i>Incidence per patient</i>	311	126	40.5%	
<b>Recurrent cellulitis per patient<sup>‡</sup></b>				
Inguino-Femoral LND	262	17	6.5%	1.000
Nodal debulking	58	4	6.9%	
Sentinel node	6	0	0%	
<i>Incidence per patient</i>	311	21	6.8%	

Table 2: Incidence of short and long-term complications to the type of groin dissection.

For lymphedema 35 Groins (22 patients) excluded due to follow up < 6 months.

<sup>†</sup> Cochran-Armitage trend test.

<sup>‡</sup> Recurrent cellulitis data only available per patient, 22 patients excluded due to follow up < 6 months.

1.11 – 2.74] per 10 years), along with current smoking (p = 0.02) and number of nodes removed (p = 0.04) (Table 4).

### Long-term complications of groin node dissection

Lymphedema was the major long-term complication occurring in 31.6% of the groins dissected. Lymphedema was more common in groins having an inguino-femoral lymphadenectomy (35%) compared to those having a nodal debulking (19.6%) or a sentinel node procedure (0%) (p = 0.003) (Table 2). An increasing number of nodes removed was significantly associated with an increasing incidence of lymphedema (p = 0.003) (Table 5).

On univariate analysis, there was evidence that obesity was associated with an increased incidence of lymphedema (p=0.01)(Table 5). However, no significant association was found in the multivariable analysis, where BMI was included as a continuous variable (Table 4).

When radiotherapy to the groin was included in our multivariable analysis, the wide 95% confidence interval did not allow a strong conclusion to be drawn about its association with lymphedema (p = 0.4; OR 1.61 [95% CI 0.53 - 4.90]) Table 4.

Recurrent lower limb cellulitis was documented in 6.8% of patients (Table 2), but this was probably substantially under-reported because over 50% of our patient population were referred from regional and rural areas and would have been treated for this complication locally. For this reason, recurrent cellulitis was excluded from further analysis.

### Discussion

This is one of the largest series in the literature reporting on groin morbidity following groin node dissection for vulvar cancer. The principal findings were the relatively high incidence of lymphocyst formation and lymphedema, and the relatively low incidence of groin wound breakdown and infection, despite routine resection of the saphenous vein.

Our lymphocyst incidence of 36.6% per groin falls within the reported range of 13% to 60% [10-15]. The incidence of lymphocysts increased significantly as the number of nodes resected increased.

The issue of drains is controversial. In view of the relatively high incidence of lymphocysts despite drain usage, the senior author began to omit the insertion of a drain in 2002. Instead, Camper's fascia was firmly sutured to the underlying fascia lata. Lymphocysts continued to be a problem but another recent Australian study has also reported no statistically significant difference in lymphocyst formation between patients with and without groin drains [15]. In a prospective Dutch study where drains were routinely used, the incidence of lymphocyst formation was reported to be lower when the drain was left in situ until drainage was < 30mls (range 2-40 days), compared to routine removal on the 5<sup>th</sup> post-operative day (16 % versus 60% respectively) [14].

The post-operative drain management after axillary lymphadenectomy for breast cancer has been studied more extensively. Two

Variable	Total groins 528 (Total patients 333) No of Groins (No patients)	Lymphocyst incidence per groin (per patient)			Groin Wound Breakdown incidence per groin (per patient)			Groin Wound Infection incidence per groin (per patient)		
		Number	% (%)	p value	Number	% (%)	p value	Number	% (%)	p value
<b>Age in years</b>										
≤ 50	76 (51)	27 (23)	35.5% (45%)	0.4773†	6 (5)	7.9% (9.8%)	0.4129†	12 (11)	15.8% (21.5%)	0.2701†
51 – 70	179 (112)	64 (55)	35.6% (49%)	(0.8274)	11 (11)	6.1% (9.8%)	(0.7625)	21 (19)	11.7% (16.9%)	(0.4278)
> 70	270 (170)	101 (86)	37.4% (50.5%)		26 (21)	9.6% (12.3%)		23 (22)	8.5% (12.9%)	
<b>Smokers</b>										
Current	119 (77)	50 (42)	42% (54.5%)	0.2489†	16 (15)	13.4% (19.5%)	0.020‡	18 (17)	15.1% (22%)	0.2731†
Past	74 (45)	23 (20)	31% (44.4%)	(0.4040)	8 (6)	10.8% (13.3%)	(0.023)	7 (7)	9.4% (15.5%)	(0.2930)
Never	332 (211)	119 (102)	35.8% (48.3%)		19 (16)	5.7% (7.6%)		31 (28)	9.3% (13.3%)	
<b>Diabetic</b>										
No	451 (286)	159 (138)	35.3% (48.2%)	0.1766†	38 (32)	8.4% (11.2%)	0.3784†	46 (44)	10.2% (15.4%)	0.4600†
Non-Insulin	51 (31)	20 (16)	39.2% (51.6%)	(0.5202)	2 (2)	3.9% (6.4%)	(0.4657)	8 (6)	15.7% (19.3%)	(0.7920)
Insulin	23 (16)	13 (10)	56.5% (62.5%)		3 (3)	13% (18.7%)		2 (2)	8.7% (12.5%)	
<b>Nodes removed</b>										
≤ 4 nodes	103 (83)	18 (16)	17.5% (19.3%)	0.0001†	2 (2)	1.9% (2.4%)	0.0054†	6 (5)	5.8% (6%)	0.1994†
5 – 8 nodes	168 (148)	56 (57)	33.3% (38.5%)	(< 0.0001)	11 (10)	6.5% (6.7%)	(0.0047)	22 (22)	13% (14.8%)	(0.1645)
9 + nodes	254 (195)	118 (102)	46.4% (52.3%)		30 (27)	11.8% (13.8%)		28 (25)	11% (12.8%)	
<b>BMI</b>										
< 30	388 (243)	136 (115)	35 (47.3%)	0.191‡	20 (17)	5.2% (7%)	< 0.001‡	40 (38)	10.3% (15.6%)	0.725‡
≥ 30	137 (90)	56 (49)	40.9% (54.4%)	(0.248)	23 (20)	16.8% (22.2%)	(< 0.001)	16 (14)	11.7% (15.5%)	(0.985)
<b>Groin Drain</b>										
Yes	348 (211)	123 (99)	35.3% (46.9%)	0.854‡	34 (28)	9.8% (13.3%)	0.064‡	42 (39)	12.1% (18.6%)	0.155‡
No	177 (122)	69 (65)	39% (52.8%)	(0.669)	9 (9)	5.1% (7.4%)	(0.094)	14 (13)	7.9% (10.7%)	(0.152)

Table 3: The incidence of short-term complications of the groin node dissection and their association to study variables.

† Cochran Armitage Trend Test, ‡ Pearson Chi Square Test.



	P value	Odds Ratio (95% CI)
<b>Lymphocyst</b>		
Age (+ 10 years)	0.4510	1.09 (0.87 – 1.36)
BMI (+ 5 points)	0.6103	0.93 (0.71 – 1.22)
Diabetes		
Non-insulin dependant	0.8496	1.10 (0.40 – 3.06)
Insulin dependant	0.2840	2.17 (0.52 – 9.01)
Smoker		
Past	0.4780	0.72 (0.30 – 1.77)
Current	0.1670	1.74 (0.79 – 3.84)
Number of Nodes (+)	0.0001	1.24 (1.12 – 1.36)
Groin drain (Yes)	0.0578	0.53 (0.27 – 1.02)
<b>Groin Wound Breakdown</b>		
Age (+ 10 years)	0.0166	1.74 (1.11 – 2.74)
BMI (+ 5 points)	0.0724	1.47 (0.96 – 2.23)
Diabetes		
Non-insulin dependant	0.3448	0.41 (0.65 – 2.60)
Insulin dependant	0.5352	1.82 (2.74 – 12.1)
Smoker		
Past	0.1730	2.44 (6.76 – 8.80)
Current	0.0237	4.83 (1.23 – 18.9)
Number of Nodes (+)	0.0360	1.12 (1.01 – 1.24)
Groin drain (Yes)	0.1476	2.32 (0.74 – 7.28)
<b>Groin Wound Infection</b>		
Age (+10 years)	0.1767	0.83 (0.64 – 1.08)
BMI (+ 5 points)	0.2302	0.80 (0.60 – 1.15)
Diabetes		
Non-insulin dependant	0.1157	2.51 (0.80 – 7.93)
Insulin dependant	0.7432	1.36 (0.21 – 8.70)
Smoker		
Past	0.7162	1.21 (0.43 – 3.50)
Current	0.4544	1.41 (0.60 – 3.50)
Number of Nodes (+)	0.0164	1.11 (1.02 – 2.80)
Groin drain (Yes)	0.5839	1.25 (0.56 – 2.80)
<b>Lymphedema</b>		
Age (+ 10 years)	0.6315	0.92 (0.67 - 1.28)
BMI (+ 5 points)	0.4210	1.18 (0.80 - 1.80)
Diabetes		
Non-insulin dependant	0.6899	0.74 (0.17 – 3.30)
Insulin dependant	0.9183	1.11 (0.14 – 8.75)
Smoker		
Past	0.8533	1.11 (0.32 – 3.90)
Current	0.9348	1.05 (0.34 – 3.24)
Radiotherapy	0.4013	1.61 (0.53 – 4.90)
Number of Nodes (+)	0.0109	1.16 (1.03 – 1.30)
Groin drain (Yes)	0.4234	0.68 (0.30 – 1.73)
Groin Infection	0.7785	0.85 (0.30 – 2.60)
Groin Breakdown	0.4700	1.70 (0.40 – 7.13)
Lymphocyst	0.0578	2.17 (0.97 – 4.81)

Table 4: Odds ratios with 95% confidence intervals (CI) for short and long-term complications of the groin dissection associated in multivariable models with patient-specific random effects.

recent studies of breast cancer patients have also concluded that the use of an axillary drain did not significantly affect symptomatic seroma rates, or any other wound complication rates [16,17]. In both these studies, post-operative hospital stay was significantly longer in the drainage groups, which was also our experience.

Our wound infection rate of 10.7% per groin is low when compared to other studies, where incidences are reported to range from 21% to 59% [10,12,14,15,18]. The only risk factor we identified was an increasing number of nodes removed. One recent study found that the incidence of post-operative groin cellulitis was lower in patients without a groin drain [15].

Our 8.2% incidence of groin wound breakdown is one of the lowest rates reported [10,11,13,14,18-20]. In addition to increasing number of nodes removed, increasing age was also a significant risk factor. This association has been noted in some studies [11,20], but not in others [10,12].

A Gynecologic Oncology Group study reported that the presence of a drain significantly increased the risk of groin wound breakdown [21]. Our groin breakdown rate was also higher in patients having a drain (9.8% versus 5.1%), but this was not significant on either univariate or multivariable analysis.

As expected, we found that current smokers were at a higher risk for groin wound breakdown. To our knowledge, only one other vulvar cancer study has reported this association [12]. However, two recent studies from the United States have reported significantly increased wound dehiscence rates in smoking cohorts. One study involved plastic and general surgical patients [22], and the other patients undergoing radical cystectomy [23].

The issue of saphenous vein preservation versus resection is controversial [19,20,24]. Two reports have suggested that saphenous vein preservation decreases groin wound breakdown, but the incidence of groin breakdown in these papers (13% and 16% respectively) was higher than our 8% incidence with vein resection [19,20]. Other studies have reported no correlation between saphenous vein ligation and complication rates for the groin dissection [10,13]. We believe that the most important aspect of preventing groin wound breakdown is the preservation of all the subcutaneous fat above Camper's fascia.

The reported incidence of lymphedema ranges from 10.9% [25] to 67% [21]. Our overall incidence of 31.6% per groin is in the mid-range of those reported [13,15,19,20,24,26,27]. The incidence was strongly correlated with the number of nodes resected and most studies concur with this finding [12,18,20,25,28]. Some authors have suggested that preservation of the saphenous vein may decrease the incidence of lymphedema [19,20], but as the problem is related to lymphatic obstruction, not venous congestion, this hypothesis lacks biologic credibility.

We have previously reported that nodal debulking for patients with bulky positive groin nodes followed by post-operative groin and pelvic radiation does not compromise survival [29], and the procedure is applicable to all patients with bulky positive nodes. The safety of the procedure was recently confirmed in a study from Leiden University [30]. In our experience, only 3.9% of groins experienced a wound breakdown after a lymph node debulking, 7.8% a wound infection and 14.6% developed a lymphocyst. These data support the earlier initiation of post-operative groin and pelvic radiation and would suggest that nodal debulking rather than an inguino-femoral lymphadenectomy should be considered the treatment of choice for patients with bulky positive nodes.

Total patients 311 Total groins 490	No of groins (no of patients)	Complication per groin (per patient)	% per groin (% per patient)	P value
Lymphedema				
<i>No of nodes removed</i>				
≤ 4 nodes	94 (76)	16 (13)	17% (17.1%)	0.0026†
5 – 8 nodes	160 (140)	52 (47)	32.5% (33.5%)	(0.0003)
9 + nodes	236 (180)	87 (78)	36.9% (43.3%)	
Lymphedema				
<i>Age groups</i>				
≤ 50 years	72 (48)	19 (18)	26.4%(37.5%)	0.217‡
51 – 70 years	174(109)	64 (48)	36.8% (44%)	(0.707)
> 70 years	244 (154)	72 (60)	29.5% (39.1%)	
Lymphedema				
With drain	323 (193)	100 (82)	31% (42.5%)	0.576‡
Without drain	167 (118)	55 (44)	32.9% (37.3%)	(0.821)
Lymphedema				
Current Smokers	115 (74)	38 (31)	33% (41.2%)	0.799‡
Past smokers	72 (43)	25 (18)	34.7% (41.8%)	(0.955)
Non-smokers	303 (194)	93 (77)	30.7% (39.7%)	
Lymphedema				
BMI < 30	359 (224)	102 (81)	28.4% (36.1%)	0.011‡
BMI ≥ 30	131 (87)	53 (46)	40.5% (52.9%)	(0.015)

Table 5: Incidence of lymphedema and its association to study variables.

35 Groins (22 patients) excluded due to follow up < 6 months.

† Cochran-Armitage trend test, ‡ Pearson Chi Square Test.

In accordance with two previous studies [10,25,28], and in contrast to two other studies [20,28] we found no significant association between the incidence of lymphedema and the addition of groinradiotherapy, although the relatively small numbers having radiation therapy may have not provided sufficient power to detect important differences.

Like some earlier studies [10,11], we found evidence of an association between the development of a lymphocyst and the subsequent development of lymphedema in univariate analysis ( $p = 0.04$ ), although the evidence was weaker after accounting for other factors ( $p = 0.06$ ). Obesity was also found to be a risk factor on univariate but not multivariable analysis, possibly due to the small number of patients in the higher BMI range. To our knowledge, higher BMI as a risk factor for developing lymphedema has only been reported in two other studies [28,31].

The major limitation of this study is the retrospective nature of the review. The incidence of long-term complications, particularly recurrent cellulitis, may have been under-reported because over 50% of the patients came from rural areas, and some were only seen annually. The strengths of the study are its large sample size, its per groin analysis, and the management of all patients in one specialised unit with a common treatment protocol.

## Conclusions

Appropriate groin node dissection is a critical part of the treatment for all patients with vulvar cancer, except those with stage IA disease. Lymphocyst formation in the immediate post-operative period and lymphedema after several months are the major morbidities, and both are associated with the number of lymph nodes removed. In this study, the use of groin drains did not significantly decrease the incidence

of lymphocyst formation. Groin node debulking for all patients with bulky positive nodes, and sentinel node biopsy for patients with small primary tumours are the only legitimate ways to reduce the number of resected groin nodes. Groin wound breakdown should occur in less than 10% of groins if care is taken to preserve the subcutaneous fat above Camper's fascia, regardless of resection of the saphenous vein.

## Author Contribution

The authors declare their responsibility for the content of this publication. EB and NH developed the concept of the article. EB conducted the data collection and analysis and drafted the manuscript. NH edited all drafts. MD provided statistical guidance and assistance with interpretation and reporting of results. All authors approved the final version for submission.

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## Competing Interests

The authors declare no conflicts of interest.

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