



Original Article

Validation of LRINEC (Laboratory Risk Indicators for Diagnosis of Necrotizing Fasciitis) Scoring System for the Diagnosis of Necrotizing Fasciitis

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Abstract

Background : Necrotizing fasciitis (NF) is a rare but potentially fatal infection involving the subcutaneous tissue and fascia. Different classifications and terminology has been used in NF based on affected anatomy, microbial cause and depth of infection.

Clinical scores like the laboratory risk indicators for NF (LRINEC) scores are available to help diagnose NF and differentiate it from other skin and soft tissue infections.

Methods : A total of 100 patients were included in the study between January 2014 to June 2014. This prospective study was done on patients admitted in surgery department of DMCH with symptoms suggestive of soft tissue infections during the study period.

Results : Mean age was found 46.5(±13.8) years in severe cellulitis and 50.9(±9.8) years in necrotizing fasciitis ($p>0.05$). Male were predominant (60%). 80% of NF had DM. The validation test for LRINEC score ≥ 6 vs necrotizing fasciitis had sensitivity of 66%, specificity 84%, accuracy 75%, positive and negative predictive values were 80.49% and 71.19% respectively. Mortality of NF was 6%.

Conclusion : The LRINEC score is an impressive diagnostic tool to distinguish necrotizing fasciitis from other severe soft tissue infections, but it is not useful for early recognition of necrotizing fasciitis.

Keywords: Necrotizing fasciitis; LRINEC score; Cellulitis; Severe cellulitis

Introduction

Necrotizing fasciitis (NF) is a rare but potentially fatal infection involving the subcutaneous tissue and fascia. It is commonly known as flesh-eating disease. Different terms and classifications have been used to describe necrotizing fasciitis of skin and subcutaneous tissue.

These include necrotizing fasciitis, synergistic necrotizing cellulites, streptococcal myonecrosis, and gas gangrene. This variety of classifications and terminology has been based on affected anatomy, microbial cause and depth of infection.

Hippocrates in the fifth century BC gave the first description of this dreaded disease¹. The first report of this disease in the United States was by a Confederate Army surgeon, Joseph Jones in 1871 and he named the entity "hospital gangrene"². Meleney³ in 1924 reported an outbreak of hospital gangrene in Beijing and coined the term hemolytic streptococcal gangrene. The term NF was first introduced by Wilson⁴ in 1952 and is the preferred term today.

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Received: 10-04-2019 **Accepted:** 25-06-2019

Historically, necrotizing infections were classified according to anatomical sites. Fournier's gangrene (involving the perineum) and Meleney's gangrene (involving abdominal wall) are examples. NF can be divided into two categories depending on the causative agents. Type-I disease is polymicrobial, caused by a mixture of aerobic, facultative anaerobic bacteria (*Klebsiella* sp., *Vibrio* sp., *Aeromonas*, *Staphylococcus*, *Clostridia*). Type-II disease is usually monomicrobial, caused by Group A streptococci. In general, Group-II NF is a rapid and progressive disease with a poorer prognosis than the type-I⁵.

Immunocompromised, advanced age, peripheral vascular disease and obesity are some predisposing factors. In a Singapore study, 70.3% of patients with NF had diabetes mellitus⁶. Patients with NF may present in later stages with septic shock, toxic shock syndrome and multi-organ failure. These patients may carry a high rate of mortality. Diagnosis of NF is essentially clinical. The gold standard is surgical exploration and tissue biopsy.

To help decide which patients required surgical exploration, particularly in those with equivocal clinical signs, laboratory and radiological tests might sometimes be useful. Clinical scores like the laboratory risk indicators for NF (LRINEC)⁷ scores are available to help diagnose NF and differentiate it from other skin and soft tissue infections. The LRINEC score is based around the routinely performed laboratory tests: C-reactive protein (CRP), white cell count, hemoglobin, serum sodium, serum creatinine and glucose levels. In Wong's retrospective study, the scoring system was applied to 89 consecutive patients admitted with biopsy-proven necrotizing fasciitis. Control patients were randomly selected patients admitted with severe cellulitis during the same period. A cut-off value of ≥ 6 was then reapplied to the patient cohort. They found a positive predictive value for the diagnosis of necrotizing fasciitis of 92% and a negative predictive value of 96%. When this scoring system was applied to a cohort of patients in a different hospital (56 patients with necrotizing fasciitis), 92.9% had a LRINEC score of ≥ 6 . These findings suggest that this scoring system may be useful in deciding

which patients with severe soft tissue infections should undergo a frozen section biopsy with a view to urgent debridement.

This study was conceived to determine if the LRINEC scoring system was useful and practical tool in the diagnosis of necrotizing skin infections in the local patient population. The LRINEC score was applied in our hospital and its usefulness was assessed by determining its predictive performance against the gold standard surgical biopsy.

Materials and Methods

The study was conducted in Dhaka Medical College Hospital from January, 2014 to June, 2014. It was a prospective observational cross-sectional study. A total of 100 patients were included in the study (50 were necrotizing fasciitis and 50 were severe cellulitis). Study population was based on all patients admitted in surgery department of DMCH with symptoms suggestive of soft tissue infections and purposive sampling technique was used. Statistical analyses were carried out by using the Statistical Package for Social Sciences version 16.0 for Windows (SPSS Inc., Chicago, Illinois, USA).

Inclusion criteria:

All patients with symptoms suggestive of soft tissue infections during the study period

Exclusion criteria:

1. Patients who have received antibiotic treatment in the last 48 hours or a minimum of 3 doses of antibiotic prior to presentation.
2. Patients who have undergone surgical debridement for present episode of soft tissue infection.
3. Patients with localized soft tissue infection.

Severe Cellulitis

The criteria for severe soft tissue infections were as follows based on documentation in the patients' charts: use of parenteral antibiotics for more than 48 hours in a patient with a soft tissue infection; and abscess requiring surgical debridement.

The LRINEC (laboratory risk indicator for necrotizing fasciitis) score

Variable	Score
C-reactive protein (mg/l)	
<150	0
150 or more	4
Total white cell count (per mm ³)	
<15	0
15-25	1
>25	2
Hemoglobin (g/dl)	
13.5	0
11-13.5	1
<11	2
Sodium (mmol/l)	
135 or more	0
<135	2
Creatinine (umol/l)	
141 or less	0
>41	2
Glucose (mmol/l)	
10 or less	0
>10	1

LRINEC score of 6 or greater is considered positive for necrotizing fasciitis.

Results

Mean age was found 46.5(±13.8) years in severe cellulitis and 50.9(±9.8) years in necrotizing fasciitis (p>0.05) (Table 1). Male were predominant, majority (60%) were male and 40% were female. Majority patients had DM in both groups, which was 35(70.0%) in severe cellulitis group and 40(80.0%) in necrotizing fasciitis group as shown in table 2. Among 100 cases 20 patients had no complication in both groups. Rest of 80 complicated cases: septicemia 14(28%) were severe cellulitis and 38(76%) were necrotizing fasciitis, anaemia 08(16%) were severe cellulitis and 05(10%) were necrotizing fasciitis. The validation test for LRINEC score ≥ 6 vs necrotizing fasciitis had sensitivity of 66%, specificity 84%, accuracy 75%, positive and negative predictive values were 80.49% and 71.19% respectively (Table 5). 58% of patients cured in severe cellulitis group and 48% in Necrotizing fasciitis group, 06% mortality were found in necrotizing fasciitis group.

Table 1. Distribution of age of the study population

Age in year	Severe Cellulitis	Necrotizing fasciitis	Total	P value
21-30	7	2	09	
31-40	12	4	16	
41-50	14	23	37	
51-60	8	15	23	
> 60	9	6	15	
Total	50	50	100	
Mean±(SD)	46.5(±13.8)	50.9(±9.8)	48.7(±12.1)	0.069

P value reached from unpaired t-test

Table 2. Co-morbidities of the study population

Co-morbidities	Severe Cellulitis	NF	Total	P value
DM	35	40	75	0.248
Liver cirrhosis	9	7	40	0.585
Cancer	7	5	12	0.527
Peripheral vascular disease	8	15	23	0.096
Others	6	4	10	0.505

P value reached from chi square test

Table 3. Distribution of site involvement of the study population

Site	Severe cellulitis	NF	Total	P value
Lower extremities	37	30	67	0.20
Upper extremities	07	06	13	0.76
Perineum	00	12	12	<0.001
Trunk	03	02	05	1.0
Head & neck	03	02	05	1.0

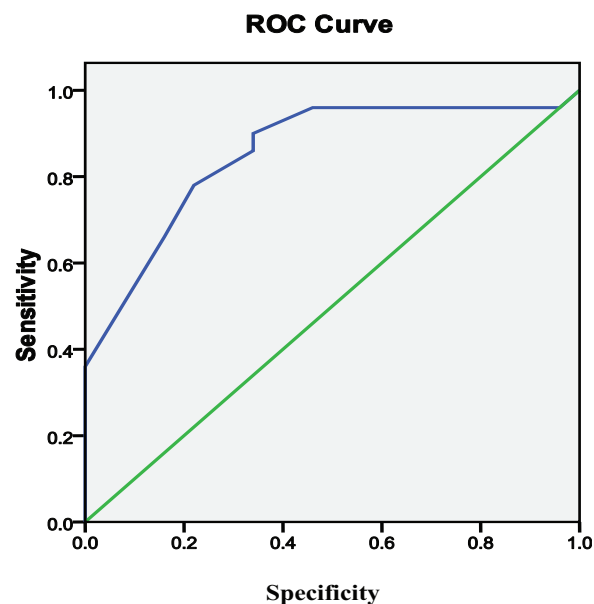
P value reached from chi square test

Table 4. Distribution of complications according to study patients

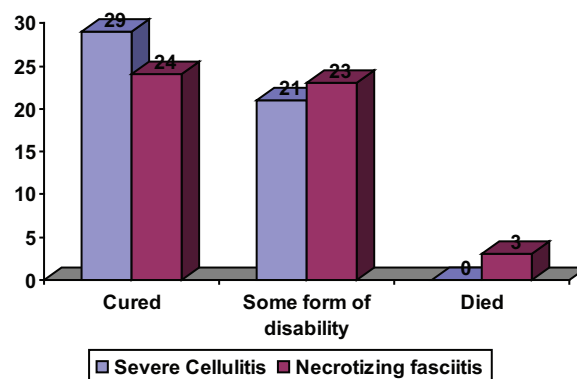
Complications	Severe cellulitis	NF	Total
No complication	18(36%)	02(04%)	20
Septicemia	14(28%)	38(76%)	52
Anaemia	08(16%)	05(10%)	13
Join stiffness	07(14%)	02(4%)	09
Chronic ostemeyelitis	03(6%)	00	03
Multiorgan failure	00	03(6%)	03
Total	50(100%)	50(100%)	100

Table 5. LRINEC score of necrotizing fasciitis and severe cellulitis

LRINEC	Severe Cellulitis	NF	Total	p value
< 6	42(84%) (TN)	17(34%) (FN)	59	< 0.001
≥6	8(16%) (FP)	33(66%) (TP)	41	
Total	50(100)	50(100)	100	

**Figure 1.** ROC of LRINEC score of necrotizing fasciitis and severe cellulitis

P value obtained from chi-square test. TP: True positive; FP: False The test result variable(s): LRINEC score (0.854), necrotizing fasciitis (0.39) has at least one tie between the positive actual state group and the negative actual state group, 95% Confidence Interval of the difference lower -0.77 and upper -0.93.

**Figure 2.** Outcome of the study population

Discussion

The present study was prospective observational cross sectional study which was done in Department of Surgery of Dhaka Medical College Hospital, Dhaka. Inclusion criteria were the patients with soft tissue infections during the study period those given informed written consent for the research.

In this study mean age was found 46.5 (\pm 13.8) years in severe cellulitis and 50.9(\pm 9.8) years in necrotizing fasciitis. Male were predominant, majority 60% were male and 40% were female. More common were in males aged 50-60. In different studies⁸⁻¹⁰, the average age of patients was 55.6; our study was found to be consistent with the literature. Female patient dominance was observed in the series of Tilkorn and colleagues¹¹.

Current study showed majority patients had DM in both groups, which was 35 (70.0%) in severe cellulitis group and 40 (80.0%) in necrotizing fasciitis group. Liao *et al.*¹² found DM in 82.0% of cases, liver cirrhosis in 21.9%, Cancer in 5.5% and others in 7.8% of cases. The most common co-morbid diseases observed in NF are DM, immunosuppression, chronic renal failure, the underlying malignancy, atherosclerosis, chronic obstructive pulmonary disease, and obesity¹³⁻¹⁵. In their study, the most common co-morbid disease was DM (52%).

This study shows that sensitivity LRINEC score \geq 6 against necrotizing fasciitis was 66%, specificity 84%, accuracy 75%, positive and negative predictive values were 80.49% and 71.19% respectively. The probability of having NF in patients with a LRINEC score of 6 or higher was calculated as 92% in the study of Su *et al.*¹³ Wong *et al.*¹⁴ reported that mortality also significantly increases in patients with LRINEC score of 6 or higher. Corbin¹⁵ also showed in his study that the complication risk is higher in patients with LRINEC score of 6 or higher. Mortality is reported in the range of 20-30% in various series^{8,9,16,17}.

Clayton *et al.*¹⁸ presented that mortality is significantly lower in young patients, in patients with BUN of 50 mg/dl or below, and in patients without ongoing sepsis. Faucher *et al.*¹⁹ proposed that co-morbid diseases do not affect mortality. On the other hand, Francis *et al.*²⁰ proposed that mortality is 50% in patients with 3 or more risk factors (being 50 years old or older, diabetes, malnutrition, hypertension, or intravenous drug abuse). As a result of this study, we propose that increased number of debridement due to severity

of disease, factor grown in the deep tissue culture (*Pseudomonas aeruginosa*), and LRINEC scores might be relative to mortality.

The developmental study by Wong *et al* reported that a LRINEC score of more than 6 had a sensitivity of 89.9%, specificity of 96.9%, positive predictive value of 92.0% and negative predictive value of 96%⁹. In 2009, Holland studied a group of 28 patients who had received surgery because of suspected NF. Ten patients were diagnosed with NF postoperatively. The results showed a sensitivity of 80%, specificity of 67%, positive predictive value of 57% and negative predictive value of 86%²¹. Liao *et al.*¹² results showed that the LRINEC score had impressive ability to discriminate NF from severe soft tissue infection, but was not a good diagnostic tool for NF. When clinical data did not indicate a diagnosis of NF, 43.3% (95% CI; 36.9-49.7%) of patients still had a LRINEC score $<$ 6, so a LRINEC score cannot help to decrease the misdiagnosis of patients with NF. Our ROC curve examining the diagnostic ability of a LRINEC score showed moderate value.

Conclusion

This study showed that male was predominant; common co-morbidities were DM, liver cirrhosis, cancer and peripheral vascular disease. Lower extremities were major site of involvement. The LRINEC score is an impressive diagnostic tool to distinguish necrotizing fasciitis from other severe soft tissue infections, but it is not useful for early recognition of necrotizing fasciitis.

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