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# ABSTRACT

**Objective**: Postoperative infections of the central nervous system are common complications of neurological surgeries, which can lead to bad consequences for the patient and increase the cost of treatment. Incorrect diagnosis or treatment of meningitis after craniotomy can lead to irrecoverable disabilities. Therefore, this study tends to investigate the prevalence and risk factors of bacterial meningitis after craniotomy in Shahid Bahonar Hospital of Kerman.

Meningitis after Craniotomy in Shahid Bahonar

Prevalence and Risk Factors of Bacterial

Hospital of Kerman in 2016

**Methods**: This was a cross-sectional, analytical, descriptive study. The statistical population was 146 patients who underwent craniotomy in Shahid Bahonar Hospital of Kerman during 2016. To find out the relationship between each risk factor and meningitis, multivariate logistic regression was used. Data was analysed by SPSS software version 16. For comparison of percentages between two groups, Chi-square test or Fisher's exact test was used. In order to compare quantitative variables between the two groups, independent t-test was used; if the data was nonparametric, U-Mann-Whitney test was used.

**Results**: Of 77 patients who underwent elective surgery, 5 cases (6.5%) developed bacterial meningitis; of patents who underwent emergency surgery, 8 cases (11.6%) developed bacterial meningitis. Two of 43 female patients (4.7%) and 11 of 103 male patients (10.7%) developed bacterial meningitis of patients with underlying disease, 4 (5.6%) patients developed bacterial meningitis and 9 (12%) of other patients developed bacterial meningitis. There were 4 cases of cerebrospinal fluid leakage. Of 146 patients, 13 cases (8.9%) developed bacterial meningitis, of which 5 had positive bacterial CSF culture; one case of acinetobacter, one case of S. epidermidis, one case of streptococcus, one case of S.aureus and one case of pseudomonas were reported.

**Conclusion**: it can be concluded that prevalence of bacterial meningitis increases with the increase in the number of received blood units, the presence of associated infection, the increase in duration of stay in hospital and ICU, the increase in duration of mechanical ventilation and repeated craniotomy and lower GCS.

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11 Keywords: Bacterial meningitis, craniotomy, multivariate logistic regression, cerebrospinal 12 fluid leakage.

## 13 1. INTRODUCTION

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15 Postoperative infections of the central nervous system are uncommon; but when they occur, 16 they are followed by serious and bad consequences [1]. Postoperative meningitis is a 17 serious complication which occurs after surgical interventions in the nervous system [2]. Although post-craniotomy meningitis is a rare complication of neurological surgeries, it can 18 cause significant disabilities if diagnosis or treatment is incorrect. Meningitis increases the 19 20 length of hospitalization and is more dangerous than most common complications 21 ofneurological surgeries [3]. Meningitis is inflammation of protective meningethat covers the 22 brain and spinal cord. This inflammation can be bacterial-viral or caused by other 23 microorganisms [4]. In practice, meningitis following surgical procedures for the central

24 nervous system can be considered as bacterial infection, until the opposite is proved. 25 Development of bacterial meningitis following neurological surgeries is different from 26 meningitis acquired from community; its dominant pathogens are gram negative bacteria 27 (Klebsiella, Pseudomonas and Acinetobacter). Meningitis symptoms usually begin in the first 28 week after surgery and caution is required in the case of high fever, focal neurological 29 symptoms, cerebrospinal fluid leakage, and increasedleukocyte and decreased cerebrospinal 30 fluid glucose. The only definitive diagnosis of bacterial meningitis is positive culture of the 31 cerebrospinal fluid (gram staining is negative in 70% of cases) [5]. Strong clinical suspicion 32 is required for meningitis in patients with clinical fever, neck stiffness, and reduced postoperative alertness. Delay in diagnosis or treatment can cause disability or death [6]. For this 33 34 reason, the cerebrospinal fluid of patients suspected of meningitis should be sent to gram 35 staining and culturing, and broad-spectrum antibiotics should be initiated; if the culture is negative, the steroid dose should be increased [5]. Considering the above, this studytends to 36 37 examine the prevalence of bacterial meningitis following craniotomy and its associated risk 38 factors in patients undergoing craniotomy in the Shahid BahonarHospital in 2016.Age, 39 gender, drug addiction, number of blood units received during hospital stay, conditions of 40 surgery (emergency or elective), cultured bacteria type (gram positive or gram negative), the 41 underlying conditions, the score that the patient takes in the APACHE scoring system, 42 associated infection, mechanical ventilation time, time of admission to ICU, spinal cord fluid 43 leakage, repeated surgery, presence of surgical attendant, GCS and hospitalization time will 44 be investigated separately. Finally, their relationship with prevalence of bacterial meningitis 45 after craniotomy is examined, the results of which can help to select appropriate antibiotics and use more effective therapeutic strategies and better management of operating room 46 47 conditions in order to reduce meningitis.

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#### 49 2. MATERIAL AND METHODS

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This was a descriptive-analytic cross-sectional study on records of patients who underwent 51 emergency or elective craniotomy in Shahid Bahonar Hospital of Kerman in 2016. All 52 53 patients undergoing brain surgery within one year (2016) were included through census. In these patients, meningitis was diagnosed based on the presence of fever 48 hours after 54 55 craniotomy, associated with implications of bacterial meningitis in the cerebrospinal fluid. 56 These findings included: white blood cell count > 1000 /microL with a percentage of 57 neutrophils greater than 80 percent, increased protein concentrations>50 mg/dL, decreased concentration of glucose <40 mg/dl(with a CSF:serum glucose ratio of <0.4); lactic acid>3.5 58 59 mmol/lit and smear and culture of bacteria would be considered, if positive.Age, gender, 60 drug addiction, number of blood units received during hospital stay, conditions of surgery 61 (emergency or elective), cultured bacteria type (gram positive or gram negative), the 62 underlying conditions, the score that the patient takes in the APACHE scoring system, 63 associated infection, mechanical ventilation time, time of admission to ICU, CSF leakage, 64 repeated surgery, presence of surgical attendant, GCS and hospitalization time were 65 investigated separately.Data registration form was designed based on the variables and was 66 completed based on the patient records. All records of the patients who underwent brain 67 surgery in Shahid BahonarHospital in Kerman withina year (2016) were enrolled by census 68 method. Prevalence of each of the risk factors was determined with a confidence interval of 69 95%. Multivariate logistic regression was used to find out the relationship between each risk 70 factor with meningitis risk. Data wasanalyzed by SPSS software version 16. For comparison, 71 Chi-square test or Fisher's exact test was used. In order to compare the quantitative 72 variables between the two groups, independent T-test was used; Mann-Whitney-U test was 73 used if data was non-parametric.

### 74 3. RESULTS

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Of 146 patients who underwent craniotomy in Shahid Bahonar Hospital Kerman in 2016, 13
(8.9%) cases developed bacterial meningitis. Comparison of nominal and quantitative variables is presented in Tables1-16 separately.

## Table 1. Comparison of prevalence of bacterial meningitis among patients undergoing craniotomy in Shahid Bahonar Hospital of Kerman in 2016 in terms of surgery type

Meningitis Surgery type	Yes	Νο	Total
Elective	5 (6.5%)	72 (93.5%)	77
Emergency	8 (11.6%)	61 (88.4%)	69
Total	13	133	146

#### 81 P=0.28

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According to Table 1, of 146 patients undergoing craniotomy in Shahid Bahonar hospital in 2016, 77 cases had elective surgery and 69 patients underwent emergency surgery. Of 77 cases undergoing elective surgery, 5 (6.5%) developed bacterial meningitis. Of patients who underwent emergency surgery, 8 cases (11.6%) developed bacterial meningitis; there was no significant difference in prevalence of bacterial meningitis between the two groups of patients undergoing emergency and elective craniotomy.

## Table 2. Comparison of prevalence of bacterial meningitis among patients undergoing craniotomy in Shahid Bahonar Hospital of Kerman in 2016 in terms of gender

Meningitis Gender	Yes	No	Total
Female	2 (4.7%)	41 (95.3%)	43
Male	11 (10.7%)	92 (89.3%)	103
Total	13 (8.9%)	133 (91.1%)	146

#### 91 P=0.346

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According to Table 2, of 146 patients undergoing craniotomy in Shahid Bahonar hospital in
 2016, 43 cases were female and 103 patients were male. Two of43female patients(4.7%)
 and 11 of 103 male patients (10.7%) developed bacterial meningitis. There was no
 significant difference in prevalence of bacterial meningitis betweenmale and female patients.

## Table 3. Comparison of mean of age between the group with bacterial meningitis and other patients undergoing craniotomy in Shahid Bahonar Hospital of Kerman in 2016

Meningitis	Age	
wennights	Mean	Std.
Yes	50.76	5.2
No	45.27	1.9

99 P=0.384

According to Table 3, of 146 patients undergoing craniotomy in Shahid Bahonar hospital in 2016, 13 cases developing bacterial meningitis aged 50.76±5.2 and other patients aged 45.27±1.9, which indicated no significant difference.

## 103 Table 4. Comparison of prevalence of bacterial meningitis among patients undergoing

104 craniotomy in Shahid Bahonar Hospital of Kerman in 2016 in terms of drug addiction

Meningitis Addiction	Yes	No	Total
Yes	54 (14.3%)	30 (85.7%)	35
No	8 (7.2%)	103 (92.8%)	111

Total	13	133	146

- 105 P=0.302%
- 106

107 According to Table 4, of 146 patients undergoing craniotomy in Shahid Bahonar hospital in 108 2016, 35 cases were addicted and 111 patients were not addicted. Five of addicted 109 patients(14.3%) developed bacterial meningitis and 111 of non-addicted patients (7.2%) 110 developed bacterial meningitis. There was nosignificant difference.

#### 111 Table 5. Comparison of prevalence of bacterial meningitis among patients undergoing 112 craniotomy in Shahid Bahonar Hospital of Kerman in 2016 in terms of underlying 113 diseases

Meningitis Underlying disease	Yes	No	Total
Yes	4 (5.6%)	67 (94.2%)	71
No	9 (12%)	66 (88%)	75
Total	13	133	146

#### 114 P=0.177

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116 According to Table 5, of 146 patients undergoing craniotomy in Shahid Bahonar hospital in 2016, 71 cases had underlying disease and 75 patients had no underlying disease. Four 117 ofpatientswith underlying disease (5.6%) developed bacterial meningitis and 9 of other 118 119 patients (12%) developed bacterial meningitis. There was nosignificant difference.

120 Table 6. Comparing the mean of the number of blood units received between the group with bacterial meningitis and other patients undergoing craniotomy in Shahid

- 121
- 122 Bahonar Hospital of Kerman in 2016

Meningitis	Number of blood units received		
Meningius	Mean	Std.	
Yes	2.85	0.64	
No	1.21	0.22	

#### 123 P=0.025 124

125 According to Table 6, the number of the blood units received was 2.85±0.64 in the group with bacterial meningitis and 1.21±0.22 in other groups; there was a significant difference 126 between the two groups in terms of the number of blood units received. 127

#### Table 7. Comparison of prevalence of positive culture ofcerebrospinal fluid in terms of 128 129 bacterium among patients undergoing craniotomy in Shahid Bahonar Hospital of

#### 130 Kerman in 2016

Meningitis Positive culture	Yes	No	Total
Yes	5	0	5
No	8 (5.6%)	133 (94.4%)	141
Total	13	133	146

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132 According to Table 7, of 146 patients undergoing craniotomy in Shahid Bahonar hospital in 133 2016, 13 cases developed bacterial meningitis; 5 of them had positive culture of 134 cerebrospinal fluid.

#### 135 Table 8. Comparison of prevalence of bacterial meningitis among patients undergoing

craniotomy in Shahid Bahonar Hospital of Kerman in 2016 in terms of associated 136 137 infection

Meningitis Associated infection	Yes	No	Total
Yes	9 (52.9%)	8 (47.1%)	17
No	4 (3.1%)	125 (96.9%)	129
Total	13	133	146

138 *P<0.01* 139

According to Table 8, of 146 patients undergoing craniotomy in Shahid Bahonar hospital in
2016, 17 cases had associated infection; 9 of them (52.9%) developed bacterial meningitis.
Of 125 cases who had no associated infection, 4 (3.1%) developed meningitis; bacterial
meningitis was significantly higher in patients with associated infection.

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#### Table 9. Comparing the mean of APACHE score between the group with bacterial meningitis and other patients undergoing craniotomy in Shahid Bahonar Hospital of Kerman in 2016

Meningitis	APACHE score	
weningitis	Mean	Std.
Yes	17	1.62
No	11.36	0.53

148 *P=0.099* 

According to Table 9, the mean of APACHE score was 17±1.62 in the group with bacterial

151 meningitis and 11.36±0.53 in other patients, and this difference was not significant.

# 152Table 10. comparing the mean of mechanical ventilation between the group with153bacterial meningitis and other patients undergoing craniotomy in Shahid Bahonar

154 Hospital of Kerman in 2016

Meningitis	Mechanica	I ventilation time
wennigius	Mean	Std.
Yes	37.23	8.50
No	5.38	0.86

### 155 *P=0.01*

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According to Table 10, mechanical ventilation time was 37.23±8.50 days in the group with
 bacterial meningitis and 5.38±0.86 days in other patients. Mechanical ventilation time was
 significantly higher in the group with bacterial meningitis.

# Table 11. Comparing the mean of ICU admission time between the group with bacterial meningitis and other patients undergoing craniotomy in Shahid Bahonar Hospital of Kerman in 2016

Moningitic	ICU admission time		
Meningitis	Mean	Std.	
Yes	37.92	8.55	
No	6.87	0.88	

163 P<0.001

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According to Table 11, ICU admission time was 37.92±8.55 days in the group with bacterial meningitis and 6.87±0.88 days in other patients; ICU admission time was significantly higher

167 in the group with bacterial meningitis.

# Table 12. comparison of prevalence of bacterial meningitis among patients undergoing craniotomy in Shahid Bahonar Hospital of Kerman in 2016 in terms of cerebrospinal fluid leakage

Meningitis Cerebrospinal fluid leakage	Yes	No	Total
Yes	0 (0%)	4 (100%)	4
No	13 (9.2%)	129 (90.8%)	142
Total	13	133	146

### 171 *P*=1.00

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According to Table 12, 4 of 146 patients undergoing craniotomy had cerebrospinal fluid leakage and none of them developed bacterial meningitis; there was no significant difference

in prevalence of bacterial meningitis in terms of cerebrospinal fluid leakage.

# Table 13. Comparison of prevalence of bacterial meningitis among patients undergoing craniotomy in Shahid Bahonar Hospital of Kerman in 2016 in terms of repeated surgery

Meningitis Repeated surgery	Yes	No	Total
Yes	9 (23.1%)	30 (76.9%)	39
No	4 (3.7%)	103 (96.3%)	107
Total	13	133	146

### 179 P<0.01

According to Table 13, of 146 patients undergoing craniotomy in Shahid Bahonar hospital in
2016, 39 cases had repeated surgery; 9 of them (23.1%) developed bacterial meningitis. Of
cases who had not repeated surgery, 4 developed meningitis; bacterial meningitis was
significantly higher in patients with repeated surgery.

#### 185 **Table 14. Comparison of prevalence of bacterial meningitis among patients** 186 **undergoing craniotomy in Shahid Bahonar Hospital of Kerman in 2016 in terms of** 187 presence of pourseurgoon attendant

187	presence of	neurosurgeon attendant	
		Maninaitia	

Meningitis Presence of attendant	Yes	No	Total
Yes	8 (8.6%)	85 (91.4%)	93
No	5 (9.4%)	48 (90.6%)	53
Total	13 (8.9%)	133 (91.1%)	146

#### 188 *P*=1.00 189

According to Table 14, of 146 patients undergoing craniotomy in Shahid Bahonar hospital in 2016, 93 cases were operated on in the presence of a neurosurgeon attendant; 8 of them (8.6%) developed bacterial meningitis. Of other patients, 5 (9.4%) developed meningitis; there was no significant difference in prevalence of bacterial meningitis in patients undergoing craniotomy in terms of the presence of neurosurgeon attendant.

# Table 15. Comparing the mean of CGS between the group with bacterial meningitis and other patients undergoing craniotomy in Shahid Bahonar Hospital of Kerman in 2016

Meningitis	GSC		
wenngins	Mean	Std.	
Yes	9.85	1.21	
No	12.65	0.32	

198 *P=0.012* 

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According to Table 15, GCS was 9.85±1.21 in the group with bacterial meningitis and 12.65±0.32 in other patients; mean of GCS was significantly lower in the group with bacterial meningitis than other patients.

#### Table 16. comparing the prevalence of bacterial meningitis in patients undergoing craniotomy in Shahid Bahonar Hospital of Kerman in 2016 in terms of hospitalization time

Moningitio	Hospitalization time		
Meningitis	Mean	Std.	
Yes	45.61	8.39	
No	13.45	1.28	

206 P<0.001

According to Table 16, hospitalization time was 45.61±8.39in the group with bacterial
 meningitis and 13.45±1.28 in other patients; hospitalization time was significantly higher in
 the group with bacterial meningitis than other patients.

#### 211

### 212 4. DISCUSSION AND CONCLUSION

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214 In general, 146 patients who underwent craniotomy in Shahid Bahonar Hospital of Kerman in 215 2016were enrolled in this study; 13 cases (8.9%) developed bacterial meningitis, 5 of them 216 had positive CSF culture for bacteria; one case of acinetobacter Bomani ,S. epidermidis, one 217 case of streptococcus, one case of S.aureus and one case of pseudomonas aeroginosa 218 were reported. Various statistics have been reported for the prevalence of meningitis 219 following craniotomy. In a study conducted in Brazil, the incidence of meningitis was 8.9% 220 and gram negative bacilli were the most common causes of infection. Among the risk factors 221 for infection, only repeated surgical procedures were significant [7]. In another study in Italy, 222 the incidence of meningitis after head and neck surgery was 1.4% [8]. In another study, the 223 incidence of meningitis was 5.5%, most of which were due to gram-positive cocci [9]. In a 224 study in India, the incidence of infection was 2.1% with a mortality rate of 5%; the most 225 common organism in this study was gram-negative bacilli [10]. In another study, the 226 incidence of meningitis was 2.7%; the most common organisms were Staphylococcus 227 aureus and Acinetobacter [11]. In a study in Iran, this prevalencewas 4.7%, which is higher than that of the developed countries [5]. According to the results obtained in this study, there 228 229 was a significant relationship between the prevalence of bacterial meningitis after 230 craniotomyand the amount of received blood, associated infection, hospitalization time, ICU 231 admission time, duration of mechanical ventilation, repeated craniotomy and GCS. According 232 to results of this study, the prevalence of bacterial meningitis increased with the increase in 233 the number of received blood units, the presence of associated infection, the increase in 234 duration of stay in hospital and ICU, increased duration of mechanical ventilation, repeated 235 craniotomy and lower GCS. The results showed that patients withdiabetes and those who 236 have cerebrospinal shunts have a higher risk for meningitis [12]. The study, which was 237 conducted in 2015, reported the risk factors for post-operative neurosurgery infections as 238 monitoring after intracranial surgeries, ventricular drainage, cerebrospinal fluid leakage, long 239 procedures, foreign objects, multiple surgeries, and shunt infections and emergency 240 procedures [1]. Another study showed that postoperative cerebrospinal fluid fistula increases 241 the risk of meningitis [13]. Another study done in 2015 found that patients with unhealthy 242 body mass index were at greater risk for cerebrospinal fluid leakage and meningitis. 243 Moreover, older patients had a higher cerebrospinal fluid leakage. The length of lumbar 244 drainage was associated with infection. In this study, BMI was the most important predictor

of cerebrospinal fluid leakage and infection. Other risk factors include age, intraoperative cerebrospinal fluid leakage,duration of lumbar drain, and combined brain surgeries [14].

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