

1 **The prevalence of neonatal near miss and associated factors in Nepal: a cross-sectional**  
2 **study**

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## 26 **Abstract**

27 **Background:** Unlike the infant mortality rate, the rate of neonatal mortality has not declined  
28 and remains a major health challenge in low- and middle-income countries. There is an  
29 urgent need to focus on newborn care, especially during the first 24 hours of birth and the  
30 early neonatal period. Determining which factors contribute to neonatal near miss (NNM) can  
31 be used to assess health care quality and identify factors capable of correction in the  
32 healthcare system to improve neonatal care. Thus, the objective of the current study was to  
33 establish the prevalence of NNM and identify its associated factors.

34 **Methods:** A hospital-based cross-sectional study was conducted at Koshi Hospital, Nepal.  
35 Neonates and their mothers (unspecified maternal age and number of gestational weeks) were  
36 enrolled. The key inclusion criterion was the admission of newborn infants to the neonatal  
37 intensive care unit at Koshi Hospital. Non-Nepali citizens were excluded. Consecutive  
38 sampling was used until the required sample size (i.e., 1,000 newborn infants) was reached.  
39 Simple and multiple logistic regression analysis was performed using SPSS<sup>®</sup> version 24.0.

40 **Results:** One thousand respondents were recruited. The prevalence of NNM was 79 per  
41 1,000 live births. Maternal secondary (adjusted odds ratio (AOR]: 0.46, 95% confidence  
42 interval (CI]: 0.24–0.88) and tertiary education (AOR: 0.18, 95% CI: 0.05–0.56), multiparity  
43 (AOR: 0.52, 95% CI: 0.39–0.86), Caesarean section (AOR: 0.48, 95% CI: 0.19–0.99), and  
44 severe maternal morbidity (AOR: 4.51, 95% CI: 2.07–9.84) were significantly associated  
45 with NNM.

46 **Conclusions:** Parity, severe maternal morbidity, mode of delivery, and maternal education  
47 were significantly associated with NNM. Healthcare workers should be aware of the impact  
48 of obstetric factors so that earlier interventions, especially the Caesarean section, can be  
49 exercised.

50

51 **Keywords:** neonatal near miss, neonatal morbidity, severe maternal morbidity, cross-  
52 sectional study, Nepal

53

#### 54 **Background**

55 The rate of pediatric mortality has long been considered an important indicator of social  
56 development, the level of economic prosperity, and healthcare quality. Globally, a 51%  
57 decline in neonatal mortality was recorded between 1990 and 2017; however, the decline in  
58 neonatal mortality has been slower than that of post-neonatal under-five mortality [1]. At the  
59 country level, annual neonatal mortality rates range from 0.9 to 44.2 deaths per 1,000 live  
60 births [1]. South Asia had 25 neonatal deaths per 1,000 live births in 2018 [2] and is a hub of  
61 the highest number of neonatal deaths along with sub-Saharan Africa [1]; a child born in this  
62 region is typically 10 times more likely to die in the first month of life than a child born in a  
63 high-income country [2]. The objective of Sustainable Development Goal (SDG) 3 and that  
64 of the global Every Newborn Action Plan is to reduce neonatal mortality (i.e., 10 or less per  
65 1,000 live births) by 2030 [3]. The neonatal mortality rate in Nepal was 21 per 1,000 live  
66 births in 2016; 57% of the births were institutional deliveries [4]. More than three-quarters  
67 (79%) of total neonatal deaths were early neonatal deaths (0-6 days) [4]. There are large  
68 variations in neonatal mortality between provinces, i.e. 15 per 1,000 live births in Province 1  
69 and 41 per 1,000 live births in Province 7. The country's flat ecological zones accounted for  
70 60% of neonatal deaths, with 40% of neonatal deaths occurring in hilly and mountainous  
71 zones in 2016 [4]. Nepal needs to reduce the rate of neonatal mortality by half in the next 10  
72 years if it is to achieve SDG 3 Goal. Thus, accelerated efforts are needed to address  
73 interprovincial disparities with regard to neonatal mortality rates.

74 Neonatal near miss (NNM) is a novel concept that has recently emerged and is similar to  
75 maternal near miss (MNM) concept. It provides vital information required for an evaluation

76 of the quality of care provided in hospital and explores opportunities to improve the  
77 performance of healthcare providers [5]. Near-miss events occur three to eight times more  
78 often than neonatal deaths [6, 7]. Thus, NNM evaluations can provide abundant evidence of  
79 the causal pathways responsible for neonatal deaths [8].

80 The conceptualization of the term “NNM” in 2009, similar to “MNM,” was proposed by  
81 Avenant [9]. That same year, Pileggi *et al.* sought to establish pragmatic NNM criteria using  
82 the World Health Organization’s (WHO) Global Survey on Maternal and Perinatal Health  
83 (WHOGS) 2005 data [10]. The initial definition of pragmatic markers included very low birth  
84 weight (i.e., < 1,500 g), < 30 gestational weeks at birth, or an Appearance, Pulse, Grimace,  
85 Activity, and Respiration (Apgar) score of < 7 at five minutes of life in neonates who went on  
86 to survive for seven days [10].

87 Pileggi-Castro *et al.* re-evaluated the NNM definition using WHOGS data and validated their  
88 revised definition using the WHO Multi-Country Survey on Maternal and Newborn Health  
89 data. According to this definition, NNM refers to “an infant who nearly died but survived a  
90 severe complication that occurred during pregnancy, birth, or within seven days of  
91 extrauterine life” [11]. The recommended pragmatic marker cut-off criteria were birthweight  
92 of < 1,750 g, < 33 gestational weeks, or an Apgar score of < 7 at five minutes of life in  
93 newborn infants who go on to survive for seven days. Pileggi-Castro *et al.* [11] also  
94 developed management marker criteria that included the use of therapeutic intravenous  
95 antibiotics, nasal continuous positive airway pressure, intubation, phototherapy within the  
96 first 24 hours, cardiopulmonary resuscitation, vasoactive drugs, anticonvulsants, surfactant  
97 administration, blood products, steroids to treat refractory hypoglycemia, and surgery in early  
98 neonatal life. In one study, the use of a combination of pragmatic and management markers  
99 was demonstrated to be more accurate than the use of either criterion alone in predicting early  
100 neonatal mortality [11]. The pragmatic criteria and management markers developed by

101 Pileggi-Castro *et al.* were shown to have sensitivity and specificity of 93% and 97%,  
102 respectively [11].

103 There is no uniform definition of NNM to this date. Although a vast number of NNM studies  
104 are available, health workers in Ghana were unfamiliar with the concept and interpreted  
105 NNM on their terms [12]. Systematic reviews on NNM, conducted in 2015 and 2017,  
106 recommended that a standard definition for NNM should be developed [13, 14]. A further  
107 two studies validated the NNM definitions but did not establish any statistically significant  
108 differences between them and the sensitivity and specificity values [15, 16].

109 Three categories of distinct NNM pragmatic criteria are commonly used in the literature: (1)  
110 birthweight of < 1,500 g, < 30 gestational weeks, and an Apgar score of < 7 in the first five  
111 minutes of life [6, 10, 17, 18], (2) birth weight of < 1,500 g, < 32 gestational weeks, and an  
112 Apgar score of < 7 in the first five minutes of life, [19-21], and (3) birth weight of < 1,750 g,  
113 < 33 gestational weeks, and an Apgar score of < 7 in the first five minutes of life [7, 11, 22-  
114 25]. Silva *et al.* recommended the addition of mechanical ventilation and congenital  
115 malformation to the pragmatic criteria [21].

116 Literature shows that the management marker criteria are inconsistently applied in most  
117 studies [11, 13, 22, 23, 25], and only a few studies have applied management markers that  
118 differ to those defined according to the initial NNM concept [6, 26-28]. A 2015 systematic  
119 review recommended the addition of antenatal steroids, parenteral nutrition use, congenital  
120 malformation (based on the International Statistical Classification of Diseases and Related  
121 Health Problems, 10<sup>th</sup> revision), and admission to the neonatal intensive care unit (NICU) to  
122 the management criteria proposed by Pileggi-Castro *et al.* [13]. Researchers in South Asia, in  
123 which NNM was assessed, applied pragmatic criteria only [17, 18]. Thus, the current study is  
124 the first to use a combination of pragmatic and management criteria in the South Asia region  
125 to establish the prevalence of NNM and identify associated factors.

126 The worldwide prevalence of NNM ranged from 39.2 to 131 per 1,000 live births in 2014 and  
127 2018 [19, 22]. Exceptionally low and high NNM prevalence was recorded in Brazil, i.e., 17  
128 per 1,000 live births in pregnant mothers and 303 per 1,000 live births in type 1 diabetic  
129 mothers, respectively [20, 25]. A population-based study conducted in Nepal applied a  
130 community-appropriate NNM definition developed by authorities and demonstrated a  
131 prevalence of 22 per 1,000 live births [26]. NNM was shown to be caused by birth asphyxia  
132 (70%), very low birth weight (17%), neonatal sepsis (10%), and prematurity (3%) [26].  
133 The NNM concept was developed to assess the quality of care provided to pregnant women  
134 [29] and their newborn infants, and its application in hospitals was shown to contribute to  
135 improvements in obstetric practice and perinatal care [30]. NNM can be utilized as a  
136 management tool to assess and identify healthcare limitations, as well as develop public  
137 policies with a focus on neonates, considered a particularly vulnerable population group [15].  
138 The NNM definition proposed by Pileggi-Castro *et al.*, which encompasses both pragmatic  
139 and management marker criteria, will be used in the current study. In this study, NNM  
140 referred to “an infant who nearly died but survived a severe complication that occurred  
141 during pregnancy, birth, or within seven days of extrauterine life” [11]. The first week in the  
142 life of a newborn infant is critical. The objective of the current study was to identify the  
143 prevalence of NNM and its associated factors in Nepal. Shifting the focus towards factors  
144 that associate newborns with life-threatening conditions within the first seven days of the  
145 crucial period can be information for policymakers for service improvement.

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## 147 **Methods**

148 This cross-sectional study was conducted on 1000 newborn infants, and their mothers  
149 admitted to the postnatal ward in Koshi Hospital, Morang, Nepal. It is a referral hospital for  
150 Province 1 and offers a NICU service. Koshi Hospital is located in Biratnagar, which is the

151 interim capital city of Province 1 and is the second-most densely populated city after  
152 Kathmandu Valley (total population of 1,058,985, with an average of 27,833 annual  
153 pregnancies) [31]. The hospital has 35 beds in total in postnatal ward and manages  
154 approximately 9,000 annual deliveries. The NICU contains six beds for neonates and admits  
155 approximately 45 neonates per month. Only newborn infants from Morang District were  
156 enrolled in the current study. Those survived for seven days were included. Consecutive  
157 sampling was applied. The sample size was calculated based on the prevalence of NNM  
158 using a single proportion formula. With the NNM prevalence of 2.2% [26], the precision of  
159 0.01, and a 20% non-response rate, the calculated sample size was determined to be 1,000  
160 newborn infants.

161 The research tool comprised of maternal and neonatal medical hospital records. Two research  
162 assistants, who were recent nursing undergraduates, collected the data daily. The hospital's  
163 medical records on current obstetric history, pregnancy complications, and the data on the  
164 newborn infants were reviewed and extracted into a case report form on the day of discharge.  
165 Socio-demographic information was obtained from the mothers.

166 The data were entered and analyzed using IBM SPSS<sup>®</sup> Statistics 24.0. Frequency and  
167 percentages were calculated for the categorical variables; mean, median, standard deviation,  
168 and interquartile range were determined for the numerical variables. Simple and multiple  
169 logistic regression analyses were used to assess the associated factors. Clinically significant  
170 variables in simple logistic regression analysis and those  $< 0.3$  were included in the multiple  
171 logistic regression analysis. Adjusted odds ratios (AORs) and corresponding 95% confidence  
172 interval (CIs) were calculated.

173 The outcome variable was the NNM status [11]. The independent variables were ethnicity,  
174 religion, wealth index, place of residence, maternal education, paternal education, maternal  
175 occupation, paternal occupation, paternal smoking habit, maternal age, age of marriage,

176 duration of the marriage, parity, number of antenatal care visits (ANC), self-reported pre-  
177 pregnancy body mass index, maternal haemorrhagic disorders, maternal hypertensive  
178 disorders, other maternal systemic disorders, maternal severe management indicators, severe  
179 maternal morbidity (SMM) and the sex of the newborn infants.

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## 181 **Results**

182 One thousand newborns and their mothers were recruited for the study between November  
183 2019 and March 2020. There were 18 deaths (12 stillbirths, five intrauterine fetal deaths, and  
184 one early neonatal death) during the study period. There were 10 multiple births, and these  
185 were treated as a single birth. Of these, four were NNM cases.

186 The prevalence of NNM was 79 per 1,000 live births in Koshi Hospital (Table 1). Table 1  
187 shows the pragmatic ( $n = 65$ ) and management markers ( $n = 44$ ) used to evaluate NNM. The  
188 most frequently encountered pragmatic criteria were an Apgar score of  $< 7$  in the first five  
189 minutes of life (63%) followed by birth weight of  $< 1,750$  g (31%). All three pragmatic  
190 criteria were applicable to only one newborn infant. Of the neonates assessed using pragmatic  
191 markers, 35 of the 65 (54%) required NICU admission, and of those evaluated using  
192 management markers and admitted to the NICU, 24 of the 44 (68%) fulfilled at least one of  
193 the pragmatic marker criteria.

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201 **Table 1** Description of pragmatic and management criteria of neonatal near miss cases in Koshi  
 202 Hospital  
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Newborn near miss characteristics	n	(%)
<b><i>Pragmatic criteria</i></b>		
APGAR Score* < 7 in 5 minutes	41	(63.1)
Birth weight < 1750 g	20	(30.7)
Gestation age < 33 weeks	11	(16.9)
Any pragmatic marker of severity	65	(100.0)
<b><i>Management criteria</i></b>		
Use of therapeutic antibiotics	25	(92.6)
Nasal continuous positive airway pressure	19	(70.3)
Cardiopulmonary resuscitation	11	(40.7)
Use of phototherapy in the first 24 hours	6	(22.2)
Any intubation (anytime within the first week)	3	(11.1)
Use of anticonvulsants	2	(7.4)
Use of steroids to treat refractory hypoglycemia	1	(3.7)
Surfactant administration	0	(0)
Use of a vasoactive drug	0	(0)
Use of any blood products	0	(0)
Any surgery	0	(0)
Any management based marker of severity	27	(100.0)
Overall criteria	79	(7.9)

204 \*APGAR Score: Appearance, Pulse, Grimace, Activity, and Respiration  
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206 Of the 44 newborns referred for NICU admission, 17 were self-referred to private hospitals;  
 207 therefore, only 27 were included in Table 1. The majority of the newborn infants in NICU  
 208 were treated with therapeutic antibiotics (93%) and nasal continuous positive airway pressure  
 209 (70%).

210 The socio-demographic, maternal characteristics and characteristics of the newborn infants  
 211 (with and without NNM) are depicted in Table 2. The proportion of adolescent mothers (< 19  
 212 years) with and without NNM was 22% and 10%, respectively.

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219 **Table 2** Distribution of socio-demographic and maternal characteristics of participants based  
 220 on neonatal near miss status

Variables	Neonatal Near Miss (n=79)		non-Neonatal Near Miss (n=921)					
	Mean	SD	n	(%)	Mean	SD	n	(%)
<b><i>Socio-demographic</i></b>								
Mother's age (year) <sup>a</sup>	21	(20, 24)			22	(20, 25)		
Age of marriage (year)	18.91	2.43			19.27	2.59		
Duration of marriage (year) <sup>a</sup>	2	(1, 4)			2	(1, 6)		
Ethnicity								
Muslim			16	(20.3)			86	(9.3)
Terai/Madhese			28	(35.4)			381	(41.4)
Dalits			16	(20.3)			163	(17.7)
Janajati			15	(19.0)			200	(21.7)
Brahmin/Chettri/Newar			4	(5.1)			91	(9.9)
Place of residence								
Urban municipality			48	(60.8)			596	(64.7)
Rural municipality			31	(39.2)			325	(35.3)
Wealth quintile								
Lowest			7	(8.9)			46	(5.0)
Second			14	(17.7)			82	(8.9)
Middle			25	(31.6)			337	(36.6)
Fourth			20	(25.3)			237	(25.7)
Higher			13	(16.5)			219	(23.8)
Mother's education								
None			15	(19.0)			106	(11.5)
Primary			21	(26.6)			187	(20.3)
Secondary			39	(49.4)			494	(53.6)
Tertiary			4	(5.1)			134	(14.5)
Father's education								
None			14	(17.7)			103	(11.2)
Primary			19	(24.1)			133	(14.4)
Secondary			38	(48.1)			512	(55.6)
Tertiary			8	(10.1)			173	(18.8)
Mother's occupation								
Housewife			75	(94.9)			866	(94.0)
Others			4	(5.1)			55	(6.0)
Father's occupation								
Unskilled manual			53	(67.1)			528	(57.3)
Sales and services			18	(22.8)			276	(30.0)
Others			8	(10.1)			117	(12.7)
Father's smoking status								
Yes			16	(20.5)			238	(25.8)
No			63	(79.7)			683	(74.2)
Sex of newborn								
Girl			37	(46.8)			459	(49.8)
Boy			42	(53.2)			462	(53.2)

***Maternal***

Variables	Neonatal Near Miss (n=79)		non-Neonatal Near Miss (n=921)					
	Mean	SD	n	(%)	Mean	SD	n	(%)
Parity								
Nulliparous			51	(64.6)			484	(52.6)
Multiparous			28	(35.4)			437	(47.4)
Mode of delivery								
Vaginal			72	(91.1)			762	(82.7)
Caesarean section			7	(8.9)			159	(17.3)
Pre-pregnancy BMI <sup>b</sup>								
Normal			59	(74.7)			690	(74.9)
Underweight			15	(19.0)			160	(17.4)
Overweight and obese			5	(6.3)			71	(7.7)
Number of ANC <sup>c</sup>								
4 visits			35	(44.3)			465	(50.5)
≤3 visits			35	(44.3)			338	(36.7)
≥5 visits			9	(11.4)			118	(12.8)
SMM								
Present			10	(12.7)			35	(3.8)
Absent			69	(87.3)			886	(96.2)
Maternal haemorrhagic disorders								
Present			2	(2.5)			16	(1.7)
Absent			77	(97.5)			905	(98.3)
Maternal hypertensive disorders								
Present			6	(7.6)			12	(1.3)
Absent			73	(92.4)			909	(98.7)
Maternal severe management indicators								
Present			2	(2.5)			16	(1.7)
Absent			77	(97.5)			905	(98.3)

221 Note: BMI =body mass index, ANC = antenatal care, SMM = severe maternal morbidity

222 <sup>a</sup> Expressed as median (interquartile range). Skewed to the right

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224 The factors associated with NNM (i.e., 20 independent variables) were evaluated using

225 simple logistic regression analysis (Table 3).

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232 **Table 3** Associated factors for neonatal near miss using simple logistic regression analysis

Variables	Crude OR <sup>a</sup>	(95% CI <sup>b</sup> )	Wald stat <sup>c</sup> (df)	P value
Mother's age	0.93	(0.87, 0.99)	4.23 (1)	0.040
Age of marriage	0.94	(0.86, 1.04)	1.42 (1)	0.232
Duration of marriage	0.95	(0.89, 1.02)	1.75 (1)	0.186
Ethnicity				
Muslim	1			
Terai/Madhesi	0.23	(0.07, 0.73)	6.21	0.013
Dalits	0.39	(0.20, 0.76)	7.67	0.006
Janajati	0.52	(0.25, 1.10)	2.86	0.091
Brahmin/Chettri/Newar	0.40	(0.19, 0.85)	5.66	0.017
Place of residence				
Urban municipality	1			
Rural municipality	0.84	(0.52, 1.35)	0.49 (1)	0.482
Wealth quintile				
Highest	1			
Fourth	1.42	(0.69, 2.92)	0.91 (1)	0.340
Middle	1.25	(0.62, 2.49)	0.40 (1)	0.527
Second	2.87	(1.29, 6.38)	6.76 (1)	0.009
Lowest	2.56	(0.97, 6.79)	3.61 (1)	0.058
Mother's education				
None	1			
Primary	0.79	(0.39, 1.60)	0.41 (1)	0.520
Secondary	0.56	(0.29, 1.05)	3.28 (1)	0.070
Tertiary	0.21	(0.07, 0.65)	7.26 (1)	0.007
Father's education				
None	1			
Primary	0.89	(0.50, 2.19)	0.02 (1)	0.895
Secondary	0.54	(0.28, 1.04)	3.34 (1)	0.067
Tertiary	0.34	(0.14, 0.84)	5.48 (1)	0.019
Mother's occupation				
Housewife	1			
Others	0.84	(0.29, 2.38)	0.11 (1)	0.743
Father's occupation				
Unskilled manual	1			
Sales and services	0.65	(0.37, 1.13)	2.32 (1)	0.127
Others	0.68	(0.31, 1.47)	0.95 (1)	0.328
Father's smoking status				
No	1			
Yes	0.73	(0.41, 1.28)	1.19 (1)	0.275
Sex of newborn				
Boy	1			
Girl	0.89	(0.56, 1.40)	0.26 (1)	0.609
Mode of delivery				
Vaginal	1			
Caesarean section	0.47	(0.21, 1.03)	3.55 (1)	0.060

Pre-pregnancy BMI				
Normal	1			
Underweight	1.09	(0.60, 1.98)	0.09 (1)	0.761
Overweight	0.82	(0.32, 2.12)	0.16 (1)	0.687
Parity				
Nulliparous	1			
Multiparous	0.61	(0.38, 0.98)	4.14	0.042
Number of ANC				
4	1			
≤3	1.37	(0.84, 2.24)	1.63 (1)	0.201
≥5	1.01	(0.47, 2.12)	0.00 (1)	0.973
SMM				
Absent	1			
Present	3.67	(1.74, 7.72)	11.72 (1)	0.001
Maternal haemorrhagic disorders				
Absent	1			
Present	1.47	(0.33, 6.51)	0.26 (1)	0.612
Maternal hypertensive disorders				
Absent	1			
Present	6.23	(2.27, 17.07)	12.63 (1)	0.000
Maternal severe management indicators				
Absent	1			
Present	1.47	(0.33, 6.51)	0.26 (1)	0.612

233 Note: BMI =body mass index, ANC = antenatal care, SMM = severe maternal morbidity

234 <sup>a</sup> Crude odds ratio

235 <sup>b</sup> Confidence interval

236 <sup>c</sup> Wald statistics

237 <sup>d</sup> Degree of freedom

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239 Among 20 independent variables, 14 variables, i.e., ethnicity, wealth quintile, maternal

240 education, paternal education, paternal occupation, paternal smoking habit, maternal age, age

241 of marriage, duration of the marriage, parity, mode of delivery, number of ANC visits,

242 maternal hypertensive disorders, and severe maternal morbidity were identified as associated

243 variables using simple logistic regression analysis with  $P < 0.3$ . Thus, they were included in

244 multiple logistic regression analysis.

245 In the multiple logistic regression analysis, mothers' education, parity, mode of delivery, and

246 severe maternal morbidity were found to be significantly associated with NNM. Mothers with

247 secondary (AOR: 0.46, 95% CI: 0.24–0.88) and tertiary education (AOR: 0.18, 95% CI:

248 0.05–0.56) were at lower odds of experiencing NNM than those with no education.

249 Multiparous mothers (AOR: 0.52, 95% CI: 0.39–0.86) were less likely to encounter NNM  
 250 than nulliparous mothers. It was less probable that newborn infants born to mothers who gave  
 251 birth via Caesarean section would be NNM cases (AOR: 0.48, 95% CI: 0.19–0.99) compared  
 252 to neonates born to mothers via normal delivery. Similarly, mothers with severe maternal  
 253 morbidity were at higher odds of giving birth to an infant who would become an NNM case  
 254 (AOR: 4.5, 95% CI: 2.07–9.84) than those without it (Table 4).

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256 **Table 4** Associated factors for neonatal near miss using multiple logistic regression analysis  
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Variables	Adj OR <sup>a</sup>	(95 % CI <sup>b</sup> )	Wald stat <sup>c</sup> (df) <sup>d</sup>	P value
Education of women				
None	1			
Primary	0.66	(0.32, 1.37)	1.22 (1)	0.268
Secondary	0.46	(0.24, 0.88)	5.40 (1)	0.020
Tertiary	0.18	(0.05, 0.56)	8.72 (1)	0.003
Parity				
Nulliparous	1			
Multiparous	0.52	(0.39, 0.86)	6.53 (1)	0.011
Mode of delivery				
Vaginal	1			
Caesarean section	0.48	(0.19, 0.99)	3.89 (1)	0.048
SMM status				
Absent	1			
Present	4.52	(2.07, 9.84)	14.43 (1)	<0.001

258 Note: SMM = severe maternal morbidity

259 <sup>a</sup> Adjusted odds ratio

260 <sup>b</sup> Confidence interval

261 <sup>c</sup> Wald statistics

262 <sup>d</sup> Degree of freedom

263 Note. No significant interaction; no multicollinearity problem; model assumptions met; no influential outliers)

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## 265 Discussion

266 The prevalence of NNM was determined to be 79 per 1,000 live births in Koshi Hospital,

267 Nepal, using a combination of pragmatic and management criteria in the current study.

268 Factors significantly associated with NNM were maternal secondary and tertiary education,

269 multiparity, severe maternal morbidity, and Caesarean section. Secondary and tertiary

270 education, multiparity, and Caesarean section were demonstrated to be protective against and  
271 lowered the risk of NNM.

272 The consensus is lacking regarding a standardized period in which NNM is said to occur  
273 across countries, which makes it difficult to compare NNM between studies. Some studies  
274 have used a near-miss period of 0–6 days [7, 10, 11, 24, 32], while others have utilized 0–27  
275 days [14, 17, 18, 20, 22, 30]. Kale *et al.* recommend extending extrauterine life from seven to  
276 28 days to increase the sensitivity of the definition of near miss by extending the survival  
277 period and a decrease in sensitivity when it was applied from 0–364 days [16]. In the current  
278 study, a period of seven days was used because three-quarters of neonatal deaths occur within  
279 the first week of life, with one quarter taking place in the first 24 hours [33]. Besides, the  
280 chance of information bias increases if NNM information is obtained from parents in the  
281 community.

282 The prevalence of NNM in the current study was compared with that reported in other studies  
283 that used a definition proposed by Pileggi-Castro *et al.* [11]. It was demonstrated in previous  
284 studies that applied the same definition that the prevalence of NNM ranged from 45.1 per  
285 1,000 live births [7] to 72.5 per 1,000 live births [11]. A community study in Nepal defined  
286 NNM using an adaptation that was based on the community based integrated management of  
287 neonatal and childhood illness criteria and reported NNM prevalence of 22 per 1,000 live  
288 births. In another study in Nepal, 37 health workers trained to identify NNM reported 28 such  
289 cases in nine months [27]. The NNM definition used in these studies was adapted for  
290 suitability at the community level. However, the prevalence of NNM in the current hospital-  
291 based study was higher, which could be owing to differences in the definition of NNM and  
292 related criteria, as well as the study settings.

293 Using pragmatic criteria [10], the prevalence of NNM was reported to be 87.6 per 1,000 live  
294 births in India in two studies [17, 18], which is higher than the 65 per 1,000 live births in the

295 present study. A possible explanation for the difference in findings is that a survival period of  
296 28 days was used in India study; hence, the sensitivity increased owing to the more extended  
297 period applied.

298 Multiparity was protective against NNM in the current study, which is contrary to the  
299 findings of prior studies [23, 34]. The findings of a few studies were in agreement with those  
300 of the present study [20, 22]; however, one showed a non-significant association [18]. Both  
301 nulliparous and grand-multiparous mothers are at high risk of complications during delivery  
302 [35, 36], and this places neonates at risk of adverse outcomes [32, 37, 38]. Nulliparity in  
303 older mothers (i.e.,  $\geq 35$  years) was found to be a risk factor for adverse perinatal outcomes  
304 [39, 40], and a higher likelihood of neonates admitted to NICU born among advanced aged  
305 nulliparous women [40-42]. However, in the current study, the proportion of women with  
306 more than four children and aged  $\geq 35$  years was small, and this prevented the ability to draw  
307 further conclusions.

308 Two literature reviews suggest that a significant proportion of nulliparous mothers are in their  
309 teens, are at increased risk of hypertension, and lack experience in childbirth [43, 44]. This  
310 elucidates the likelihood of NNM affecting nulliparous women [32, 38]. Prior studies have  
311 shown that first-born infants are at higher risk of neonatal mortality than the second- or third-  
312 born infants [43, 45]. However, in some studies, parity was not shown to have an association  
313 with neonatal mortality [46].

314 Elsewhere, a high chance of NNM affecting women undergoing Caesarean section has been  
315 demonstrated [20, 21, 23, 47, 48]. In recent studies in India and Ethiopia, although NNM  
316 cases were higher in women who underwent Caesarean delivery, a direct association could  
317 not be established [7, 18]. Contrary to the findings in the literature, the present study showed  
318 that Caesarean delivery was protective against NNM. In support of this, in the United States,  
319 the Caesarean section was observed to reduce neonatal mortality in preterm births [49].

320 The WHO has recommended that Caesarean sections should only be conducted when  
321 medically necessary and recommends an upper limit of 15% concerning the percentage of  
322 deliveries that should be conducted using this method [50]. In the current study, the overall  
323 percentage of births performed using Caesarean delivery was 17%, which is in contrast to  
324 that in public hospitals in Nepal (i.e., 12%) [4]. Elective Caesarean sections are not  
325 performed at Koshi Hospital. The proportion of Caesarean sections performed in mothers  
326 with SMM was two times higher than that performed in mothers without SMM (31% versus  
327 16%) in this study. Previous works of literature show, SMM to be significantly associated  
328 with higher rates of current Caesarean section [51-55] and preterm births born to SMM  
329 mothers are higher than among mothers without SMM [51, 52]. Timely emergency Caesarean  
330 sections performed to manage conditions associated with SMM can prevent an adverse  
331 impact on neonatal health [46].

332 Measures to mitigate medical oversights are in place to manage high-risk pregnancies  
333 effectively [56]. Studies in the literature suggest that very low-birth weight newborns  
334 delivered by Caesarean section have lower neonatal mortality and low five-minute Apgar  
335 scores, which indicates the protective role of Caesarean section [46]. However, overall, there  
336 is a lack of consensus in the literature over a popular opinion that neonatal mortality and  
337 morbidity is higher in infants who are delivered by Caesarean section [55, 57-59].

338 In the current study, maternal secondary and tertiary education decreased the likelihood of  
339 NNM comparative to a lack of maternal education. Prior studies that assessed socio-  
340 demographic factors for NNM have not established a significant association between NNM  
341 and maternal education [7, 18, 21-23, 28]. However, a universal association between  
342 maternal education and neonatal mortality, especially in developing countries, has been  
343 demonstrated [45, 56, 58, 60-63] and supports the current study's findings. Educated mothers  
344 have a better knowledge of healthy behaviours, have a more informed approach to self-care,

345 make better health-related choices, and utilize the healthcare system appropriately [37, 64].  
346 An educated mother is also more likely to have a higher socioeconomic status [58].  
347 The current study found an association between severe maternal morbidity and NNM,  
348 consistent with the finding of a study carried out in Ethiopia [7] but contradictory to the  
349 finding of a study performed in Brazil [25]; therefore, more studies are warranted in this  
350 regard to obtain more conclusive findings. Very few studies have explored the relationship  
351 between MNM and NNM. One study showed a solid association (OR: 17.15, 95% CI: 1.85–  
352 159.12) [25], whereas others have not demonstrated a significant association between MNM  
353 and NNM [22, 48]. In support of the current study, there is an association between MNM and  
354 higher rates of adverse perinatal outcome born to these mothers [65, 66].  
355 Specific severe maternal morbidity condition (i.e., severe hypertension and pre-eclampsia) is  
356 associated with adverse neonatal outcomes [32, 67, 68]. A considerable number of newborn  
357 infants with severe hypoxia and neonatal asphyxia have been born to women with MNM  
358 [67]. Similarly, maternal obstetric complications have been shown to play a role in the  
359 underlying causes of neonatal deaths [45, 69]. Therefore, early screening for poor obstetric  
360 conditions during the antenatal period and the appropriate management of intrapartum  
361 complications is crucial to ensure a reduction in the number of NNM cases.  
362 The WHO guidelines, which are followed in Nepal as part of the national protocol,  
363 recommend a minimum of four antenatal visits for uncomplicated pregnancies. The current  
364 finding did not establish any association between ANC and NNM. However, in other  
365 research, ANC was associated with neonatal mortality in Nepal [62]. Mixed [20, 22] and  
366 converse [7] findings were found. Attending four or more ANC sessions was protective [36,  
367 70], whereas an inadequate number of antenatal visits was associated with neonatal mortality  
368 [69, 71].  
369 There are a few possible explanations for this. Firstly, a quarter (24%) of women in Nepal

370 typically receive all seven components of ANC [72]. The majority of Nepal public  
371 institutions lack basic ultrasonography and laboratory facilities (i.e., blood and urine testing),  
372 which means that a reduced percentage of women undergo blood and urine examinations [72-  
373 74], and most pregnant women only receive health education, iron supplementation, blood  
374 pressure measurements, and tetanus toxoid [72]. Secondly, there is poor compliance by  
375 pregnant mothers with ANC advice [75]. In a qualitative study, it was reported that, in  
376 general, pregnant women do not attend ANC follow-up visits as they hope to avoid  
377 healthcare workers asking for blood and urine samples. They also do not comply with  
378 adequate calcium intake due to poverty. It has been established that women in the wealthiest  
379 quintile [73, 76] who reside in urban areas [73, 76] are highly educated [72, 73, 76],  
380 nulliparous [76], and aged 30–39 years [72], are more likely to utilize adequate ANC [77].  
381 Hence, even mothers who attended less than the recommended four ANC sessions owing to  
382 poor compliance did not show a significant association with NNM in the current study.  
383 Maternal age was significantly associated with NNM when simple logistic regression  
384 analysis was applied, but this was not supported using multiple logistic regression analysis.  
385 Elsewhere, it has been reported that advanced maternal age (> 35 years) was significantly  
386 associated with NNM [22]. Globally, advanced maternal age may increase in pregnancy and  
387 obstetric complications; thus, early intervention is recommended to try to prevent adverse  
388 outcomes [22]. A significant association has been reported between advanced maternal age  
389 [44, 45, 78-82] or younger age (< 20 years) and adverse neonatal outcomes [57, 58, 63, 83,  
390 84]. However, this association was not found in one study [46].  
391 The results of the current study are generalizable to births in government institutions in  
392 Nepal. To the best of our knowledge, this is the only study to have explored factors that  
393 impact NNM in South Asia using both pragmatic and management criteria.

394 The study had several limitations. The cross-sectional nature of the research meant that casual  
395 association could not be proved. Seventeen of the 44 neonates with conditions requiring  
396 admission to the NICU were self-referred to private hospitals; therefore, these data were  
397 unavailable. The date of the last menstrual period was used to calculate gestational age; thus,  
398 incorrect estimations may have been introduced due to recall bias.

399

#### 400 **Recommendations**

401 Nulliparity and severe maternal morbidity should be considered high-risk obstetric  
402 conditions. Therefore, screening should be performed during the antenatal period, and, if  
403 indicated, referral should be made to a tertiary hospital with adequate facilities. Future studies  
404 should explore contributory factors to NNM in illiterate women and those with  
405 communication barriers, as well as the impact of content and quality of ANC. An evaluation  
406 is recommended of the risk of NNM in specific maternal populations, such as older women  
407 and those with multiple pregnancies.

408

#### 409 **Conclusion**

410 Maternal determinants and socioeconomic factors could result in a NNM. The neonates of  
411 mothers with severe maternal morbidity are at increased risk of NNM; conversely, Caesarean  
412 section, multiparity, and secondary and tertiary education were shown to be protective against  
413 NNM in the current study. Caesarean section is encouraged to be exercised as an earlier  
414 intervention, especially among women of high-risk obstetric factors.

415

416

417

418

419 **Abbreviations**

420 SDG: Sustainable Development Goal; NNM: neonatal near miss; WHO: World Health  
421 Organization, WHOGS: WHO Global Survey on Maternal and Perinatal Health; NICU:  
422 neonatal intensive care unit; MNM: maternal near miss; AOR: adjusted odds ratio; CI:  
423 confidence interval; BMI: body mass index; APGAR: Appearance, Pulse, Grimace, Activity,  
424 and Respiration; ANC: antenatal care; SMM: severe maternal morbidity

425

426 **Declarations**

427 *Ethics approval and consent to participate*

428 The ethical approval was obtained from the Human Research Ethics Committee Universiti  
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430 336/2019). Written consent of the participants was taken before the interview. Parental  
431 consent was taken for women less than 18 years of age.

432

433 *Consent for publication*

434 Not applicable.

435

436 *Availability of data and materials*

437 The authors are happy to share anonymized data related to this paper upon receiving a  
438 specific request, along with the purpose of that request. Interested parties may contact

439 [hayatikk@usm.my](mailto:hayatikk@usm.my)

440

441 *Competing interest*

442 The authors declare that they have no competing interests.

443

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448

449 ***Authors' contributions***

450 SR designed the study, involved in data collection, analyze data and prepared the manuscript.  
451 NMN and NHH designed the study, involved in data analysis and critically revised  
452 subsequent drafts of the manuscript for valuable intellectual content. All authors read and  
453 approved the final manuscript.

454

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459

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