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Delays in Emergency Obstetrics referrals in Addis Abeba hospitals

Endalkachew Mekonnen Assefa¹, Yemane Berhane²

Abstract

Background: Preventing severe maternal outcomes or deaths requires timely and appropriate referrals to facilities capable of providing adequate and appropriate treatment. In the city of Addis Abeba the geographic access to health facilities that provide obstetric emergency services is universal. However, timely referral to obstetric facilities when needed is often delayed leading to unnecessary maternal complications due to many factors. This study tries to assess where the delays occur in the referral chain.

Methods: A facility based cross sectional study was conducted in two public hospitals, Zewditu and Gandhi Memorial Hospitals, in Addis Abeba, the capital city of Ethiopia.The study was conducted between December 2018 and February 2019. Data were collected using a pretested and structured questionnaire. All mothers referral to the two hospitals during the study period were included in the study.

Results: A total of 403 pregnant women referred for delivery to the study hospitals were included in the study. Three-fourth (301, 74.7%) of referred pregnant mothers had the third delay (delayed care at the receiving hospital); (211, 52.4%) had first delay (delay in making decision to seek care). Overall 366(90.8%) mothers had experienced at least one of the three delays and 71(17.6%) had all the three delays. Twenty-nine (7.2%) referred mothers had severe maternal outcomes (SMO). The most leading causes/diagnosis of SMO were blood transfusion 17 (58.6%) followed bypost partum hemorrhage 15 (52%) then eclampsia 9 (31%). In addition, mothers who experienced severe maternal outcomes were 2.9 times more to have at least one of the three delays.

Conclusion & recommendation: The majority of the mothers in this study had serious delays in both making decision to seek care for birthing and in actually receiving care once at a hospital. Strengthening the guidelines to reduce delays in maternity care is critical to prevent severe maternal outcomes.

Key words: Addis Abeba, Severe maternal outcomes, Three-delay model

Introduction

The majority of maternal deaths are clustered around labour, delivery and the 24 hours postpartum (1),(2). It is estimated that just five conditions (postpartum hemorrhage; puerperal sepsis; preeclampsia and eclampsia; obstructed or prolonged labour, and complications of unsafe abortion), account for at least60% of all maternal mortality (3). Life-threatening situations may develop rapidly and without warning, often in previously uncomplicated pregnancies(4).

Globally, it is recognized that significant inroads in maternal mortality cannot be made without dramatically increasing access to emergency obstetrical care (EmOC). The World Health Organization estimates that at least 88–98% of maternal deaths can be averted with timely access to existing, emergency obstetric interventions (5). This produces a triple return on investment, saving mothers and newborns and preventing stillbirths(6).

A number of factors can influence a woman's ability to access effective interventions to treat complications in the event of an obstetric emergency. Thaddeus and Maine (1994) group these into three broad categories using a classic, pathways-based framework known as the 'three delays model'(7).The 'three delays model' attempts to explain delays in women accessing emergency obstetric care as the result of: 1) decision-making, 2) accessing services and 3) receipt of appropriate care once a health facility is reached.

Referral is often associated with the second delay of the three delays model– associated with reaching the appropriate level of care. But in fact, a referral system can reduce all three delays. If a population knows that a system is reliable and affordable, families may make the decision to seek care more quickly (the first delay(8).

The major obstacles that affect the referral system reported by both the health workers and mothers were: 1) financial barriers (for transportation and service payments at health facilities), 2) lack of means of transportation, 3) distance, and 4) lack of awareness of services and the importance of services (9).

Factors associated with health seeking behavior are multidimensional. Socio-cultural and economic problems, lack of awareness, the quality of health services, and infrastructure such as transport services all affect whether and where a woman will seek care, how long it will take to reach care, and whether she receives the appropriate care in a timely fashion(10).

Studies showed referrals in pregnancy and childbirth can be (1) institutional or self-referral, depending on the involvement of first line services; (2) antenatal, delivery or postnatal referral; and (3) elective or emergency referral. Pregnant mothers may referred due to demographic risks, obstetric historical risks, prenatal complications and delivery and immediate postnatal complications(11). On other hand Studies show that high risk prediction may not necessarily mean that the woman will have complication and many women identified as being at risk go on to have normal deliveries(12). Defining a framework and process for obstetric referrals may lead to reduction of maternal mortality and morbidity. Referral should be broadly defined to include not only transport, but it should be timely referral to minimize or prevent the delay for transportation (called second delay), and ensure prehospital care while transporting a patient to the referral facility(13), (14).

It is widely accepted that substantial reductions in maternal mortality and severe morbidity are impossible to achieve without an effective referral system for complicated cases(9), (15). There is a paucity of evidence from Ethiopia on the proportion of complicated and emergency obstetric cases that are detected and referred to appropriate higher level health institutions.

Objectives

The objectives of this study were to identify indications of emergency obstetrics referral and, types of delay and maternal health outcomes based on the three delay model among mothers who referred for childbirth.

Methods

We used the STROBE cross sectional reporting guidelines (42).

Study design and setting

A facility based cross-sectional study was conducted between December 10/2018-February 28/2019 in two government hospitals Zewditu Memorial Hospital (ZMH) and Gandhi Memorial Hospitals (GMH). Both hospitals are under Addis Abeba Health Bureau and affiliated with Addis Abeba University College of Health Sciences. Gandhi Memorial Hospital is a referral maternity hospital and Zewditu Memorial Hospital is also a comprehensive referral hospital. Both hospitals have 40 (forty) health centers under them allocated by Ministry of Health and other health facilities which can refer to them after contact. Both hospitals are giving Comprehensive emergency obstetrics care (CEmoc) and attending more than 17,000 deliveries per a year.

The referral system for obstetric emergency in Addis Abeba is organized to include Basic Emergency obstetric care (BEmoc) facilities and CEmoc facilities. The referral system is developed to work both ways. Referral between facilities is facilitated by the liason office or Maternal Health Task Force. An ambulance system is organized to transportmothers which should be accompanied by midwives. The midwife provides care during transportation and hand-over the mother to the receiving hospital care provider with a referral paper. In the city, all maternity services including labour/delivery and the ambulance services are provided free of charge in all government health facilities.

Eligibility criteria

All pregnant mothers who were referred for labor and delivery services to the study hospitals and who gave consent were included in the study. The mothers were identified from emergency or labor wardsdaily and interviewed before they were discharged from the hospitals. Residents and interns at ob-gyn department helped collecting the data using a pre-tested structured and structured

questionnaire. The completed questionnaires were reviewed by principal investigator. Incomplete questionnaires were returned to the person who completed the questionnaire by asking the mother for the second time.

The referral papers reviewed and date, time and diagnosis of referral were recorded for each mother. The triage paper and patient chart are also reviewed including mode of transportation, date and time of arrival, sources of referrals, obstetrics performances, the time taken to admit/get the service after arrival, diagnosis at receiving hospital, gestational age, place and mode of delivery, newborn outcomes and severe maternal complications types and managements. Mothers interviewed on demographic and economic characteristics, time interval to seek medical advice and reason of delay to seek the care (if there was).

The three delays time frame were operationally defined through a consultative process involving obstetrician & gynecologists who had working experience of 7-20 years in the selected hospitals. Accordingly, first delay, time elapsed time between the recognition of a maternal complication and the decision to transport the patient to a health facility, was considered if the decision was not made within 60 minutes. The second delay was if the mother did not reach to referral hospitals within 60 minutes of referral. The third delay was if the mother didn't receive care or intervention within 30 minutes. Severe Maternal outcomes were any maternal complication including laboratory evidences of organ damage and maternal death during the process of delivery or before discharge from the hospital.Potentially life threatening maternal conditions (PLTMC) was considered when the mother had at least one of the following; hemorrhagic complications, hypertensive disorders and complications, end organ injury, blood product transfusion, ICU admission, uterine rupture, hysterectomy/laparatomy.

Study size: Single proportion formula used. By assuming 50% of the referred mothers have delay, degree of precision of 5% (d) and confidence interval of 95% (Z=1.96) the sample size was 403.

Analyses

Data were entered in epi-info version 7.2.2.6 and transported to SPSS Version-21 statistics software for cleaning and analysis. Descriptive statistical were used to present mothers by their socio-demographic characteristics, referral diagnosis, diagnosis at receiving hospitals, obstetrics characteristics, mode of delivery, newborn outcomes, the three delays, and severe maternal outcomes. Severe maternal outcomes were analyzed for the three delays. The relationship between the three delays and SMO were examined using the multivariate logistic regression. The goodness of the model was tested by Hosmer-Lemeshow test for goodness-of-fit. Selected variables were included in the model to account for maternal characteristics differences other than delays in seeking and receiving care.

Ethics approval

Ethical clearance was obtained from both Addis Continental Institute of Public Health (ACIPH) and Addis Abeba Health Bureau (AAHB). Support letters were written to both study hospitals from the local health authorities to gain access. Informed consent was obtained from each mother who participated in the study after ensuring their privacy and confidentiality of information.

Results

A total of 403 pregnant mothers referred for labor/delivery services to the study hospitals were included. The majorities weremarried (380, 94.3%), and have completed at least secondary school (54.3%). The median age was 26 years (range 18-43 years) and the median gravidity was 3 (range 1-7). Majority mothers were primigravida (56.1%) and most pregnant mothers (58.8%) were at term pregnancy (37 weeks-41w6d)(Table1).

Table 1.Socio-demographic and obstetrics characteristics of respondents, Addis Abeba, Ethiopia,2019 (n=403)

Characteristics n=403	Number (%)
Study Hospital	
Gandhi Memorial Hospital (GMH)	173 (42.9
Zewditu Memorial Hospital (ZMH)	230 (57.1
Age (years) median: 26 years (range 18-43)	
< 20	32 (7.9
20-25	158 (39.2
26-30	151 (37.5
31-35	46 (11.4
≥36	16 (4.0
Marital status	0,
Married	380 (94.3
Others(unmarried, divorced)	23 (5.
Educational level	
No formal education	49 (12.1
Primary school	135 (33.
Secondary school	117 (29.
Preparatory	35 (8.
Vocational and above	67 (16.0

Gravidity n=403		
1		226 (56.1)
2-4		165 (40.9)
≥5		12 (3.0)
Parity n=146		
1		90 (61.6)
≥2		56 (38.4)
Abortion n=60	0	
1	6	48 (80)
≥ 2		12 (20)
Gestational age		
28-33w6d		9(2.2)
34-36w6d		22 (5.5)
37-41w6d		237 (58.8)
≥ 42		42 (10.4)
Unknown		93 (23.1)

The majority of the pregnant mothers were referred from health centers (387, 96%) and transported by the ambulance provided by the health facility. The three most common referral diagnosis were term premature rupture of membrane (PROM) (25.8%), latent first stage of labour abnormality (16.6%), and post term pregnancy (15.4%)(Table 2).

The most common diagnosis at the receiving hospital following reassessment of the mother were latent first stage of labor (126, 31.3%), term premature rupture of membrane (123, 30.5%), and active first stage of labor (18.9%) (Table 2).

Table 2. Source of referral, transportation, referral and receiving hospital diagnosis, mode & place of delivery, among referred pregnant mothers for delivery, Addis Abeba, Ethiopia, 2019.

Characteristics

Number (%)

Health Center	387 (96.0)
Others	16 (4.0)
Transportation	
Ambulance	290 (72)
Others(Taxi, personal car)	113 (28)
Receiving hospital contacted before the women referred	
Yes	157 (39.0)
No	246 (61.0)
Referral diagnosis ** N=403	
Antepartum hemorrhage (APH)	11
Pregnancy induced hypertension (PIH)	51 (2
Previous c-section scar	19
Latent phase labor abnormality	67 (2
Active phase labor abnormality	52 (2
Second stage labor abnormality	20
Post term pregnancy	62 (2
Malpresentation/malposition	14
Premature rupture of membrane (PROM)	104 (2
Cephalopelvic disproportion (CPD)	6
Non-reassuring fetal heart pattern(NRFHRP)	25
Meconium staining amniotic fluid (MSAF)	25
Others	19

Ante partum hemorrhage (APH)	13 (3.2)
Pregnancy induced hemorrhage (PIH)	54 (13.4)
Previous C-section scar	18 (4.5)
Latent first stage of labor (LFSOL)	126 (31.3)
Active first stage of labor	76 (18.9)
Second stage of labor	29 (7.2)
Post-term	52 (12.9)
Malpresentation/malposition	18 (4.5)
Premature rupture of membrane (PROM)	123 (30.5)
Cephalopelvic disproportion (CPD)	7 (1.7)
Non-reassuring fetal heart pattern (NRFHRP)	13 (3.2)
Meconium staining	21 (5.2)
Others	28 (6.9)

** percentage is > 100 % due to more than one answer is possible

The majority of the mothers delivered through vaginal route (254, 63.3%) and the remaining by Csection (148, 36.7%). The most common indication for C-section were non-reassuring fetal heart rate pattern (27%), cephalo-pelvic disproportion/malpresentation/malposition (24.3%), and meconium staining in latent first stage of labor (22.3%). The majority of the babies were born alive (96.5%). There were eight (2%) intra-partum fetal losses among total deliveries (table 3).

Table 3.Mode of delivery, place of delivery and perinatal outcome of referred mothers, Addis Abeba, Ethiopia 2019.

Node of delivery		
Vaginal delivery	229 (56.8)	
Assisted breech delivery	4 (1.0)	
C-section	148 (36.7)	
Instrumental delivery	21(5.2)	
Vacuum	16 (4.0)	

Forceps		5 (1.2)
Laparatomy		1 (0.2)
C-section indication N=148		
Non-reassuring fetal heart rate	e (NRFHR)	40 (27.0)
Meconium in LFSOL		33 (22.3)
Cephalopelvic disproportion (C	CPD)	21 (14.2)
Malpresentation/malposition		15 (10.1)
Previous C-scar with labor/lab	or abnormality	10 (6.8)
Cord prolapse/presentation		7 (4.7)
Non reassuring biophysical pro	ofile (NRBPP)	5 (3.4)
АРН		4 (2.7)
Others		13(8.8)
Place of delivery N=403	Ó	•
Labor ward/Operation room	Ì	386 (95.8)
Emergency OPD		17 (4.2)
Outcome	N=403	2
Alive		389 (96.5)
Newborn referred to NICU	Yes	137 (34)
	No	252 (62.5)
Stillbirth		14 (3.5)
Fetal heart beat positive on arrival		8(2.0)
Fetal heart beat negative	on arrival	6(1.5)

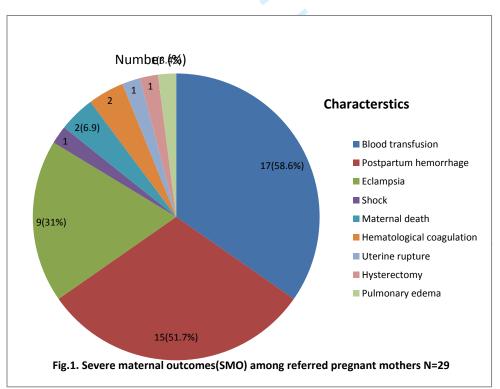
Among 403 referred mothers for childbirth, 71(17.6%) mothers experienced all the three delays. Almost three-fourth referred mothers (74.7%) experienced third delay followed by first delay (52.4%). Majority (366, 90.8%) mothers had at least one of the delays (Table 3).

Characteristics	Number	· (%)
N=403	Yes	Νο
First delay	211 (52.4)	192 (47.6
Second delay	163 (40.4)	240(59.6
Third delay	301(74.7)	102(25.3
At least one delay	366 (90.8)	37 (9.2
All three delay	< 71 (17.6)	332 (82.4

Table 3. The frequency of the three delays among referred mothers, Addis Abeba, Ethiopia 2019.

** multiple answer possible

Twenty-nine (7.2%) mothers had severe maternal outcomes. The most common severe maternal outcome was blood transfusion (58.6%) followed by post partum hemorrhage (51.7%) then eclampsia (31%). Nearly three-fourth mothers with SMO (78.5%) had more than one complication (Figure 1).



The most common delays in mothers with SMO were third delay (58.6%) followed by first delay (51.7%). All most three-fourth mothers experienced at least one of the delays (79.3%) and one-quarter (24.1%) mothers had all the delays (table 4).

Statistically significant association was observed between SMO and third delay, at least one of the delays. Referred mothers with the third delay had 2.2 times (95% CI of 1.025-4.840) more likely to be a risk for severe maternal outcomes. Mothers who experienced at least one of the delays had 2.9 times (95% CI 1.093-7.620) more likely to be a risk for severe maternal outcomes (table 4). After adjusted for age, gravidity, parity, educational level and marital status none of the delays were significant. This may be due to small sample of mothers with SMO.Also the three delays were not significant for age, gravidity, parity, educational level, gestational age and marital status.

Table 4. SMO and types of delays Addis Abeba, Ethiopia 2019.

Characteristic	S	Severe maternal outcomes			
Types of delay	,	No	Yes	P-value	COR 95 % CI
		Number (%)	Number (%)		
First delay	No	178 (47.6%)	14 (48.3)	•	
	Yes	196 (52.4)	15 (51.7)	0.944	1.028(0.483-2.189)
Second delay	No	224 (59.9)	16 (55.2)		
	Yes	150 (40.1)	13 (44.8)	0.618	1.213(0.567-2.596)
Third delay	No	90(24.1)	12 (41.4)		
	Yes	284 (75.9)	15 (58.6)	0.043	2.227(1.025-4.840)
All delays	No	310 (82.9)	22 (75.9)		
	Yes	64 (17.1)	7 (24.1)	0.342	1.541(0.632-3.761)
At least one	No	31 (8.3)	6 (20.7)		
delay	Yes	343 (91.7)	23(79.3)	0.032	2.889(1.093-7.620)

**AOR not significant after adjusted for age, marital status, educational level, gestational age, gravidity, parity.

Eighty (19.9%) of referred mothers had at least one potential life threatening conditions (PLTC). The most common complications was hypertensive disorders 56 (70%) followed by blood transfusion 17 (21.3%) then post partum hemorrhage (table 5).

Table 5.Potentially life threatening conditions among referred mothers Addis Abeba, Ethiopia,2019

Characteristics* N=80	Number (%)
Hemorrhagic complications	
Ante partum hemorrhage (AP,PP)	13 (16.3)
Post partum hemorrhage (PPH)	15 (18.8)
Ruptured uterus	1 (1.25)
Coagulopathy	2 (2.5)
Hypertensive disorders	
Severe hypertension/Preeclampsia	50 (62.5)
Eclampsia	9 (11.3)
HELLP syndrome	2 (2.5)
Others	
Pulmonary edema	1 (1.25)
Shock	1(1.25)
Thrombocytopenia	2 (2.5)
Management indication of severity	1
Transfusion of blood derivatives	17 (21.3)
Major surgical intervention(hysterectomy)	1(1.25)
*Multiple response possible	

*Multiple response possible

Discussion

All three types of delay were common in maternity services in the study hospitals; the most sever being the delay within the receiving hospital. The most common referral diagnoses were premature rupture of membranes (PROM), latent phase labor abnormality and post-term pregnancy. The third delay was significantly associated with severe maternal outcomes.

This study is similar to previous study in Addis Abeba showed that PROM was the most common referral diagnosis contributed 19.7 %(16); this is different from other reports in other sub-Saharan countries (17). There were many discrepancies of cases diagnosis between at referring and at receiving hospitals, the most common diagnosis at receiving hospitals are the latent first stage of labor (31.3%) followed by premature rupture of membrane (29%). Some cases were under-/over-diagnosed. Non-reassuring fetal heart rate pattern/fetal distress were twice over diagnosed; on the other hand latent first stage of labor twice less under-diagnosed at referring facilities. These discrepancies might occur due to differences between the qualification of care providers at the health center and referral hospitals. In the health centers midwives are the main caregivers while in the receiving hospitals midwives, medical interns, general practitioners, residents, and obstetrician and gynecologist involved in maternal care. Variation in providers' competences can be contributing factor to the observedgaps (18).

Studies have found that distance to facilities is a clear barrier to women accessing health facilities (19) (20) but in Addis Abeba, proximity to services does not appear to be a problem, as the median distance to a facility that provides surgical services is 5 kilometers, well below the national average of 45 kilometers (21). Two-fifth of mothers had the second delay. When we compare with other studies it may be low however this proportion of the second delay is not expected because referring facilities/catchment health centers are near to the receiving hospitals and expected to refer by ambulance (21).

Our study showed the rate of occurrence of SMO indicators were higher than the findings of an earlier study done in other parts of the country (22) (23) (24) (25) and other countries (26) (27). This high proportion SMO might be due to the fact that our study selected facilities are referral hospitals which are serving complicated cases and cases referred from other health facilities which were beyond their capacity/needed further interventions. This study showed that direct obstetric causes were the most common leading factors of SMO and the most common diagnosis were postpartum hemorrhage (52%) followed by eclampsia (31%) and the most common intervention was blood transfusion (58.6%). Obstetric hemorrhage and hypertensive disorders (eclampsia, hematological coagulation, and pulmonary edema) were found to be the top underlying complications among cases of SMO; similarly, hypertensive disorders and obstetric hemorrhage were the most common underlying causes of PLTC. This is comparable to the findings from studies in other parts of the country (23), (24) (28) and other countries (27, 29-31) including sub-Saharan countries (32).

Emergency obstetric care use by women is influenced by a complex interaction of factors leading to delay in decision-making, accessing services and receipt of proper care once a health facility is reached (33). Receiving appropriate care after reaching at the health facility (delay three) was the most common

(58.6%) followed by delay in seeking care (delay one) (51.7%) then reaching at the appropriate health facility (delay two) (44.8%) were identified among SMO and more than half of (58.6%) SMO cases had encountered at least one of the delays which were similar to study done elsewhere in the country (23) (34) (35) (36); however delay two and one is seen less frequently than the findings from other developing countries (17) (27) (33) (34) (38) (39). This can be justified by overloaded of cases, limited hospitals capacity, a difference in socio-demographic characteristics of the study population and the proximity of health facilities.

In this study, failure to receiving appropriate care after reaching at the health facility (delay three) was found to have the strongest association with SMO: with a two-fold increase in the risk. This supports the WHO hypothesis relating a high case fatality in the hospital as an indicator for the presence of delay in receiving an adequate and proper treatment (40) and it indicates the poor performance of obstetrics services (41). Seeking care from a facility that is incapacitated health facility and system, poor leadership (mismanagements of hospital resources, poor co-ordinations and lack of understanding of obstetrics emergencies) contribute to significant delay after reaching the health facility. These factors were reported as significant contributors to delay in several studies (23)(34) (36). Studies from Tigray, 88% of all maternal deaths were attributed to health system failure (35). In our study, 59 % of SMO cases and both maternal deaths (MD) had health system related factors as a possible reason for delay three.

The limitation of this study include that it was conducted in two referral hospitals which often receive complicated cases and referred mother with complications; and the results might not be representative of other institutions and the community. In addition, cases were missed because they transferred to health centers after two hours of uncomplicated vaginal delivery.

However, we believe that this study gives the extent of delays and SMO &its indicators experienced by women who were referred for an emergency delivery. In fact, if the delay is so severe in these wellestablished referral centers one may expect it to be worse in some not well staffed and equipped centers. We decided to focus on women with severe maternal outcomes (SMO) rather than less severe forms of obstetric complications because their situations are the closest to maternal deaths.

Conclusion

The burden of severe maternal outcomes (SMO) is high due to preventable and/or treatable direct obstetric. The majority of the mothers in this study had serious delays in both making decision to seek care for birthing and in actually receiving care once at a hospital. We recommend strengthening health referral systems, provision of training to maternal health service providers at the referring facilities, and addressing specific health system bottlenecks during labor and birth in order to ensure no mother will be endanger.

Abbreviations

AFSOL: Active First Stage of Labor

ANC: Antenatal Care

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BEmOC: Basic Emergency Obstetrics Care
CEmOC: Comprehensive Emergency Obstetrics Care
EOPD: Emergency Out-Patient Department
GMH: Gandhi Memorial Hospital
HELLP: Hemolysis, Elevated liver enzyme, Low Platelets
LFSOL: Latent First Stage of Labor
LW: Labor Ward
MD: Maternal Death Ratio
MNM: Maternal Near-Miss
MWH: Maternity Waiting Home
NICU: Neonatal Intensive Care Unit
PIH: Pregnancy Induced Hypertension
PLTC: Potential Life Threatening Conditions
PPH: Post Partum Hemorrhage
PROM: Premature Rupture of Membrane
SMO: Severe Maternal Outcomes
WHO: World Health Organization
ZMH: Zewditu Memorial Hospital
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Author details

¹Endalkachew Mekonnen Assefa (MD, MPH), Addis Abeba University-College of Health Sciences Department of Obstetrics and Gynecology; e-mail-<u>endmekon@gmail.com</u>; phone no. +251-913605674: corresponding author

²Yemane Berhane (MD, PhD), Addis Continental Institute of Public Health; <u>yemaneberhane@gmail.com</u>

Contributors: EMA wrote the draft of the paper, performed the data analysis and wrote the final manuscript. YB advised the paper's scope and the draft of manuscript. Both authors approved the final version.

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Data availability statement: data are available from the corresponding author upon reasonable request.

References

1. Campbell O, Graham WJ. Measuring maternal mortality and morbidity: levels and trends. London: Lond Sch Hyg Trop Med. 1990;

2. Li XF, Fortney JA, Kotelchuck M, Glover LH. The postpartum period: the key to maternal mortality. Int J Gynaecol Obstet.1996;54:1–10.

3. Khan KS, Wojdyla D, Say L, Gulmezoglu AM, Van Look PF. WHO analysis of causes of maternal death: a systematic review.2006;Lancet 367:1066–1074.

4. Fortney J. Emergency obstetric care: the keystone in the arch of safe motherhood. .; IJOG. 2001;(74):95–97.

5. World Health Organization (WHO). Maternal Mortality. Fact Sheet. Media Centre. 2016;

6. World Health Organization (WHO). True magnitude of stillbirths and maternal and neonatal deaths underreported. Media Cent. 2016;

7. Thaddeus S, Maine D. Too far to walk: maternal mortality in context. Soc Sci Med. 1994;38:1091–110.

8. The Effectiveness of Emergency Obstetric Referral Interventions in Developing Country Settings: A Systematic Review [Internet]. [cited 2018 Jul 28]. Available from: http://journals.plos.org/plosmedicine/article?id=10.1371/journal.pmed.1001264

9. Knight HE, Self A, Kennedy SH. Why Are Women Dying When They Reach Hospital on Time? A Systematic Review of the 'Third Delay.' PLoS ONE [Internet]. 2013 May 21 [cited 2018 Sep 7];8(5). Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3660500/

10. JSI Research and Training Institute. The Last 10 Kilometers (L10K). Emergency referral for pregnant women and newborns: A Rapid Community and Health System Assessment. BETA Development Consulting Firm, Addis Abeba. 2012 Jun;

11. U. Onwudiegwu, O. C.Ezechi. Emergency obstetric admissions: late referrals, misdiagnoses and consequences. J J Obstet Gynaecol. 2001;Volume 21(Issue 6).

12. Bossyns P, Abache R, Abdoulaye MS, Miyé H, Depoorter A-M, Van Lerberghe W. Monitoring the referral system through benchmarking in rural Niger: an evaluation of the functional relation between health centres and the district hospital. BMC Health Serv Res. 2006 Apr 12;6(1):51.

13.Giovine A, Ostrowski C. Technical report on Improving Transportation and Referral for Maternal Health: Knowledge Gaps and Recommendations. Wilson Cent. 2010;

14. Ronsmans C, Graham WJ, Lancet. Maternal Survival Series steering group: Maternal mortality: who, when, where, and why. Lancet. 2006;(368(9542)):1189–2000.

15. Qureshi RN, Sikandar R, Hoodbhoy Z, Bano R, Jabeen N. Referral pattern of emergencies in obstetrics: Implications for defining scope of services and policy. JPMA J Pak Med Assoc. 2016;66(12):1606–10.

16. Alemnesh H. Mirkuzie, Mitike Molla Sisay and Mulu Muleta Bedane. High proportions of obstetric referrals in Addis Abeba: the case of term premature rupture of membranes. 2016. 9:40.

17. David M. Goodman, Emmanuel K. Srofenyoh, Adeyemi J. Olufolabi, Sung Min Kim and Medge D. Owen. The third delay: understanding waiting time for obstetric referrals at a large regional hospital in Ghana:BMC Pregnancy and Childbirth (2017) 17:216; DOI 10.1186/s12884-017-1407-4.

18. Alemnesh H Mirkuzie1, Mitike Molla Sisay, Alemnesh Tekelebirhan Reta, et al. Current evidence on basic emergency obstetric and newborn care services in Addis Abeba, Ethiopia; a cross sectional study:2013.

19. McNamee P, Ternent L, Hussein J. Barriers in accessing maternal healthcare: evidence from low-and middle-income countries. Expert Rev Pharmacoecon Outcomes Res.2009;9(1):41–8.

20. Gabrysch SC, Oona MR. Still to far to walk: Literature review of the determinants of derlivery service use. BMC Pregnancy Childbirth. 2009;9(34).http://www.biomedcentral.com/1471-2393/9/34.

21. Federal Ministry of Health, Addis Abeba Regional Health Bureau. Emergency Obstetric and Newborn Care Services: Standard Operation Manual for Activities of Catchement Teams in Addis Abeba. 2012.

22. Dile M, Abate T, Seyum T. Proportion of maternal near misses and associated factors in referral hospitals of Amhara regional state, Northwest Ethiopia: institution based cross sectional study. GynecolObstet (Sunnyvale). 2015;5:308. https://doi.org/10.4172/2161-0932.1000308.

23. Wondimagegnehu Sisay Woldeyes1,2, Dejene Asefa3* and Geremew Muleta. Incidence and determinants of severe maternal outcome in Jimma University teaching hospital, south-West Ethiopia: a prospective cross-sectional study. Woldeyes Al BMC Pregnancy Childbirth 2018. 18:255(18:255).

24. Gebrehiwot Y, Tewolde BT. Improving maternity care in Ethiopia through facility based review of maternal deaths and near misses. Int J GynecolObstet. 2014;127:S29–34.

25. Liyew EF, Yalew AW, Afework MF, Esse. Incidence and causes of maternal near-miss in selected hospitals of Addis Abeba, Ethiopia. ONE. 2017;12(6):e0179013. https://doi.org/10.1371/journal.pone.0179013.

26. Jabir M, et al. Maternal near miss and quality of maternal health care in Baghdad, Iraq. BMC Pregnancy Childbirth. 2013;13:11.

27. Leonardo Antonio Chavane, Patricia Bailey, Osvaldo Loquiha, Martinho Dgedge, Marc Aerts5, and Marleen Temmerman. Maternal death and delays in accessing emergency obstetric care in Mozambique.BMC Pregnancy and Childbirth. (2018) 18:71 https://doi.org/10.1186/s12884-018-1699-z.

28. Özge Tunçalp, Michelle J. Hindin, Kwame Adu-Bonsaffoh , Richard M., Adanu. Assessment of maternal near-miss and quality of care in a hospitalbased study in Accra, Ghana. Int J Gynecol Obstet 123 (2013) 58–63.

29. Say L, Souza JP, Pattinson RC. WHO working group on Maternal Mortality and Morbidity classifications. Maternal near miss-towards a standard tool for monitoring quality of maternal health care. 2009. 23:287–96.

30. Nelissen EJ, et al. Maternal near miss and mortality in a rural referral hospital in northern Tanzania: a cross-sectional study. BMC Pregnancy Childbirth. 2013;13:141.

31. Oladapo et al. When getting there is not enough: a nationwide cross-sectional study of 998 maternal deaths and 1451 near-misses in public tertiary hospitals in a low-income country.BJOG.2015.www.bjog.org.DOI: 10.1111/1471-0528.13450.

32. Dan K Kaye, Othman Kakaire, and Michael O Osinde. Systematic review of the magnitude and case fatality ratio for severe maternal morbidity in sub-Saharan Africa between 1995 and 2010. BMC Pregnancy Childbirth. 2011;11(65).doi: 10.1186/1471-2393-11-65.

33. Ayele Geleto, Catherine Chojenta, Abdulbasit Musaand Deborah Loxton. Barriers to access and utilization of emergency obstetric care at health facilities in sub-Saharan Africa: a systematic review of literature. (2018) 7:183.https://doi.org/10.1186/s13643-018-0842-2.

34. Knight HE, Self A, Kennedy SH. Why are women dying when they reach hospital on time? A systematic review of the 'third delay'. PLoS One. 2013; 8(5):e63846. https://doi.org/10.1371/journal.pone.0063846. 35. Hailu S, Enqueselassie F, Berhane Y. Health facility-based maternal death audit in Tigray, Ethiopia. Ethiop J Health Dev. 2009; 23:115–9.

36. Rodolfo Carvalho Pacagnella, Jose Guilherme Cecatti, Maria Jose Osis & João, Paulo Souza. The role of delays in severe maternal morbidity and mortality: expanding the conceptual framework.Reproductive Health Matters 2012;20(39):155–163. www.rhm-elsevier.com. Doi:0.1016/S0968-8080(12)39601-8.

37. Rodolfo C Pacagnella, José G Cecatti, Mary A Parpinelli, Maria H Sousa, Samira M Haddad, Maria L Costa, João P Souza, Robert C Pattinson and the Brazilian Network for the Surveillance of Severe Maternal Morbidity, study group. Delays in receiving obstetric care and poor maternal outcomes: results from a national multicentre cross-sectional study.BMC Pregnancy and Childbirth 2014, 14:159.

38. Bouchra Assarag, Bruno Dujardin, Alexandre Delamou, Fatima-Zahra Meski,, Vincent De Brouwere. Determinants of Maternal Near-Miss in Morocco: Too Late, Too Far, Too Sloppy?PLOS ONE | January 22, 2015.DOI:10.1371/journal.pone.0116675.

39. Saira Yunus, Shahnaz Kauser, Shazia Ali. Three "Delays" as a framework for critical analysis of maternal near miss and maternal mortality. Journal of South Asian Federation of Obstetrics and Gynecology, May-August 2013;5(2):57-59.

40. World Health Organization. Evaluating the quality of care for severe pregnancy complications: the WHO near-miss approach for maternal health. Geneva: WHO; 2011.

41. Filippi V, Ronsmans C, Gohou V, et al. Maternity wards or emergency obstetric rooms? Incidence of near-miss events in African hospitals. Acta Obstetricia et Gynecologica Scandinavica 2005;84(1):11–16.

42. Von Elm E, Altman DG, Egger M, Pocock SJ, Gotzsche PC, Vandenbroucke JP. The Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) Statement: guidelines for reporting observational studies.

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1 2			what was done and what was found
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8 9 10	rationale		investigation being reported
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18 19 20	Study design	<u>#3</u>	Present key elements of study design early in the paper
21 22	Setting	<u>#3</u>	Describe the setting, locations, and relevant dates, including
23 24 25			periods of recruitment, exposure, follow-up, and data collection
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28 29 30			selection of participants.
31 32		<u>#4</u>	Clearly define all outcomes, exposures, predictors, potential
33 34 35			confounders, and effect modifiers. Give diagnostic criteria, if
36 37 38			applicable
39 40 41	Data sources /	<u>#4</u>	For each variable of interest give sources of data and details of
42 43	measurement		methods of assessment (measurement). Describe comparability
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1 2	Quantitative	<u>#4</u>	Explain how quantitative variables were handled in the
3 4	variables		analyses. If applicable, describe which groupings were chosen,
5 6 7			and why
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11 12 13	methods		for confounding
14 15	Statistical	<u>#n/a</u>	Describe any methods used to examine subgroups and
16 17 18	methods		interactions
19 20 21	Statistical	<u>#4</u>	Explain how missing data were addressed
22 23	methods		
24 25 26	Statistical	<u>#4</u>	If applicable, describe analytical methods taking account of
27 28 29	methods		sampling strategy
30 31 32	Statistical	<u>#n/a</u>	Describe any sensitivity analyses
33 34	methods		
35 36 37	Results		
38 39	Participants	<u>#n/a</u>	Report numbers of individuals at each stage of study—eg
40 41 42			numbers potentially eligible, examined for eligibility, confirmed
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47 48 49			unexposed groups if applicable.
50 51 52			Note it was cross-sectional study
53 54 55	Participants	<u>#n/a</u>	Give reasons for non-participation at each stage
56 57 58			Note: all participants participated
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1 2 3	Participants	<u>#n/a</u>	Consider use of a flow diagram
4 5 6			Note: it was just at one study.
7 8	Descriptive data	<u>#5</u>	Give characteristics of study participants (eg demographic,
9 10 11			clinical, social) and information on exposures and potential
12 13			confounders. Give information separately for exposed and
14 15 16			unexposed groups if applicable.
17 18	Descriptive data	<u>#n/a</u>	Indicate number of participants with missing data for each
19 20 21			variable of interest
22 23 24			Note: there was no missed data due to different interests.
25 26	Outcome data	<u>#6</u>	Report numbers of outcome events or summary measures.
27 28 29			Give information separately for exposed and unexposed groups
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33 34 35 36 37 38 39	Main results	<u>#11</u>	Give unadjusted estimates and, if applicable, confounder- adjusted estimates and their precision (eg, 95% confidence
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2 3			potential bias or imprecision. Discuss both direction and
4 5 6			magnitude of any potential bias.
7 8	Interpretation	<u>#13</u>	Give a cautious overall interpretation considering objectives,
9 20			limitations, multiplicity of analyses, results from similar studies,
21 22 23			and other relevant evidence.
24 25 26	Generalisability	<u>#13</u>	Discuss the generalisability (external validity) of the study
27 28			results
29 80 81 82	Other Information		
83 84	Funding	<u>#14</u>	Give the source of funding and the role of the funders for the
85 86 87			present study and, if applicable, for the original study on which
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Delays in Emergency Obstetrics referrals in Addis Ababa hospitals: a facility-based, crosssectional study

Endalkachew Mekonnen Assefa¹, Yemane Berhane²

Abstract

Objectives: To assess where the delays occur in the referral chain at most and maternal health outcomes based on the three delay model in Addis Ababa.

Design: This study was a facility-based cross sectional study

Setting: Two public and tertiary hospitals in Addis Ababa

Participants: All pregnant women who were referred for only labor and delivery services after 28 weeks of gestation between December 2018 and February 2019 in Zewditu and Gandhi Memorial hospitals.

Primary and secondary outcome measures: The primary outcome was the type of delays from the three-delay model which met operationally defined time. Maternal health outcomes based on the three-delay model was secondary outcome.

Results: A total of 403 pregnant women referred for delivery to the study hospitals were included in the study. Three-fourth (301, 74.7%) of referred pregnant women had the third delay (delayed receiving appropriate care); (211, 52.4%) had first delay (delay in making decision to seek care). Overall 366(90.8%) pregnant women had experienced at least one of the three delays and 71(17.6%) had all the three delays. Twenty-nine (7.2%) referred women had severe maternal outcomes (SMO). The most leading causes/diagnosis of SMO were blood transfusion 17 (58.6%) followed by post partum hemorrhage 15 (52%) then eclampsia 9 (31%). In addition, women who experienced severe maternal outcomes were 2.9 times more likely to have at least one of the three delays.

Conclusion & recommendation: This study highlights the persistence of delays at all levels and especially delay three and its contribution to the severe maternal outcomes. Strengthening health referral systems and addressing specific health system bottlenecks during labor and birth in order to ensure no mother will be endanger. We also recommend a qualitative method of study (focus group discussion and in-depth interview) and observing the tertiary hospitals set-up and readiness to manage obstetrics emergencies.

Key words: Addis Ababa, Emergency obstetrics referral, severe maternal outcomes, Three-delay model

Article summary

Strengths and limitations

- This study gives the extent of delays and severe maternal outcomes experienced by women who were referred for labor and delivery
- > This study focused on women with severe maternal outcomes (SMO) rather than less severe forms of obstetric complications because their situations are the closest to maternal deaths.
- Women after delivery might be missed if they transferred to health centers and to other hospitals after delivery for different reasons.
- The results might not be representative of other institutions and the community because it was conducted in two referral hospitals which often receive and treat complicated cases.

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Introduction

The majority of maternal deaths are clustered around labour, delivery and the 24 hours postpartum (1),(2). It is estimated that just five conditions (postpartum hemorrhage; puerperal sepsis; preeclampsia and eclampsia; obstructed or prolonged labour, and complications of unsafe abortion), account for at least60% of all maternal mortality (3). Life-threatening situations may develop rapidly and without warning, often in previously uncomplicated pregnancies(4).

Globally, it is recognized that significant inroads in maternal mortality cannot be made without dramatically increasing access to emergency obstetrical care (EmOC). The World Health Organization estimates that at least 88–98% of maternal deaths can be averted with timely access to existing, emergency obstetric interventions (5). This produces a triple return on investment, saving women and newborns and preventing stillbirths(6).

A number of factors can influence a woman's ability to access effective interventions to treat complications in the event of an obstetric emergency. Thaddeus and Maine (1994) group these into three broad categories using a classic, pathways-based framework known as the 'three delays model' (7). The 'three delays model' attempts to explain delays in women accessing emergency obstetric care as the result of: 1) decision-making, 2) accessing services and 3) receipt of appropriate care once a health facility is reached.

Referral is often associated with the second delay of the three delays model– associated with reaching the appropriate level of care. But in fact, a referral system can reduce all three delays. If a population knows that a system is reliable and affordable, families may make the decision to seek care more quickly (the first delay(8).

The major obstacles that affect the referral system reported by both the health workers and women were: 1) financial barriers (for transportation and service payments at health facilities), 2) lack of means of transportation, 3) distance, and 4) lack of awareness of services and the importance of services (9).

Factors associated with health seeking behavior are multidimensional. Socio-cultural and economic problems, lack of awareness, the quality of health services, and infrastructure such as transport services all affect whether and where a woman will seek care, how long it will take to reach care, and whether she receives the appropriate care in a timely fashion(10).

Studies showed referrals in pregnancy and childbirth can be (1) institutional or self-referral, depending on the involvement of first line services; (2) antenatal, delivery or postnatal referral; and (3) elective or emergency referral. Pregnant women may referred due to demographic risks, obstetric historical risks, prenatal complications and delivery and immediate postnatal complications(11). On other hand Studies show that high risk prediction may not necessarily mean that the woman will have complication and many women identified as being at risk go on to have normal deliveries(12).

Defining a framework and process for obstetric referrals may lead to reduction of maternal mortality and morbidity. Referral should be broadly defined to include not only transport, but it should be timely referral to minimize or prevent the delay for transportation (called second delay), and ensure prehospital care while transporting a patient to the referral facility(13), (14).

It is widely accepted that substantial reductions in maternal mortality and severe morbidity are impossible to achieve without early decision-making to seek care, an effective referral system for complicated cases and receiving timely and appropriate care (9), (15).

The objective of this study was to determine the types of delay and maternal health outcomes based on the three delay model among women who referred for labor and delivery. Results from this study may be an input for the hospitals, health bureau, policy-makers and other stakeholders to act on bottle-necks of emergency obstetrics services by identifying the most common types of delay.

Methods

We used the STROBE cross sectional reporting guidelines (16).

Study design

A facility based cross-sectional study was conducted between December 10/2018-February 28/2019 in two government hospitals Zewditu Memorial Hospital (ZMH) and Gandhi Memorial Hospitals (GMH).

Study setting

This study conducted in tertiary hospitals located in capital city of Ethiopia, Addis Ababa. Both hospitals are under Addis Ababa Health Bureau and affiliated with Addis Ababa University College of Health Sciences. Gandhi Memorial Hospital is a referral maternity hospital and Zewditu Memorial Hospital is also a comprehensive referral hospital. Both hospitals have 40 (forty) health centers under them allocated by Ministry of Health and other health facilities which can refer to them. Both hospitals are giving Comprehensive emergency obstetrics care (CEmOC) and attending more than 17,000 deliveries per a year.

The referral system for obstetric emergency in Addis Ababa is organized to include Basic Emergency obstetric care (BEmOC) facilities and CEmOC facilities. The referral system is developed to work both ways. Referral between facilities is facilitated by the liason office or Maternal Health Task Force. An ambulance system is organized to transport women which should be accompanied by midwives. The midwife provides care during transportation and hand-over the mother to the receiving hospital care provider with a referral paper. In the city, all maternity services including labour/delivery and the ambulance services are provided free of charge in all government health facilities.

Eligibility criteria

All pregnant women who were referred for only labor and delivery services after 28 weeks of gestation or baby outcome \geq 1 kg and delivered in the selected hospitals, and also who gave consent were included.

Data Collection Tools

After they gave birth, women were identified and interviewed from emergency OPD, labor ward and inpatient wards every day before they discharged from the hospital by data collectors using pretested and structured questionnaires.

Data collection Procedures

The referral papers reviewed and date, time and diagnosis of referral were recorded for each mother. The triage paper and patient chart are also reviewed including mode of transportation, date and time of arrival, sources of referrals, obstetrics performances, the time taken to admit/get the service after arrival, diagnosis at receiving hospital, gestational age, place and mode of delivery, newborn outcomes and severe maternal complications types and managements. Women interviewed on socio-demographic characteristics, time interval to seek medical advice and reason of delay to seek the care (if there was).

The completed questionnaires were reviewed by principal investigator and supervisors. Incomplete questionnaires were filled if the women were not discharged; otherwise incomplete questionnaires were discarded.

Main outcomes and measures of the study

The three delays time frame were operationally defined through a consultative process involving six obstetrician & gynecologists (three from each hospitals) who had working experience of 7-20 years in the selected hospitals. Accordingly, first delay defined time elapsed between the recognition of complication/s, and the decision to transport and reach to a health facility, was considered if it took more than 60 minutes. The second delay was defined if the mother did not reach to referral hospitals within 60 minutes of referral. The third delay was if the mother didn't receive care or admitted within 30 minutes. Severe maternal outcomes were any maternal complications including blood transfusion (any type & units), Post-partum hemorrhage, shock, eclampsia, uterine rupture, pulmonary edema, laparatomy, laboratory evidences of organ damage and maternal death during the process of delivery and/or before discharge from the hospital. Potentially life threatening maternal conditions (PLTMC) was considered when the mother had at least one of the following; hemorrhagic complications, hypertensive disorders and complications, end organ injury, blood product transfusion, ICU admission, uterine rupture, hysterectomy/laparatomy.

Sample size: Single proportion formula used by assuming 50% of the referred women have delay, degree of precision of 5% (d), confidence interval of 95% (Z=1.96), assuming 5% non-response rate and the final sample size was 403.

Statistical analysis

Data were entered in epi-info version 7.2.2.6 and transported to SPSS Version-21 statistics software for cleaning and analysis. Descriptive statistical were used to present women by their socio-demographic characteristics, referral diagnosis, diagnosis at receiving hospitals, obstetrics characteristics, mode of delivery, newborn outcomes, the three delays, and severe maternal outcomes. Severe maternal outcomes were analyzed for the three delays. The relationship between the three delays and SMO were examined using the multivariate logistic regression. The goodness of the model was tested by Hosmer-Lemeshow test for goodness-of-fit. Selected variables were included in the model to account for maternal characteristics differences other than delays in seeking and receiving care.

Ethical issues

Ethical clearance was obtained from both Addis Continental Institute of Public Health (ACIPH) and Addis Ababa Health Bureau (AAHB) Institution Review Board (IRB). Support letters were written to both study hospitals from AAHB-IRB to gain access.

Patient and public involvement:

Women asked for some questions after obtained informed consent on voluntary basis. All the responses given by the participants and the results obtained kept anonymous and confidential. There was no public involvement in the design, conduct and interpretation of the study. Patients were not asked to advise on interpretation or writing up of results. We did not include patient involvement in the design of this study .We have presented a summary of the findings at medical and public health schools and among health providers in Addis Ababa and plan to continue presenting the results at professional society's conferences. Results were shared with administrative of both selected hospitals and Addis Ababa health Bureau to facilitate improved obstetrics services. There are no plans to disseminate the results of this research to study participants.

Results

Table 1 shows the descriptive information for socio-demographic and obstetrics characteristics of referred pregnant women. The mean age of 403 pregnant women referred for labor/delivery services was 26.47 ± 4.5 years and ranged from 18 to 43. The majorities were married (380,94.3%), and have completed at least secondary school (54.3%). Majority women were primigravida (56.1%) , the mean gravidity was 1.77 ± 1.1 and ranged between 1 to 7. Most pregnant women (58.8%) were at term pregnancy (37 weeks- 41w6d) (Table1).

Table 1.Socio-demographic and obstetrics characteristics of respondents, Addis Ababa, Ethiopia, 2019 (n=403)

Characteristics n=403

Number (%)

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Gandhi Memorial Hospital (GMH)	173 (42.9)
Zewditu Memorial Hospital (ZMH)	230 (57.1)
Age (years) median: 26 years (range 18-43)	
< 20	32 (7.9)
20-25	158 (39.2)
26-30	151 (37.5)
31-35	46 (11.4)
≥36	16 (4.0)
Marital status	
Married	380 (94.3)
Others(unmarried, divorced)	23 (5.7)
Educational level	
No formal education	49 (12.2)
Primary school	135 (33.5)
Secondary school	117 (29.0)
Preparatory	35 (8.7)
Vocational and above	67 (16.6)
Gravidity n=403	27.
1	226 (56.1)
2-4	165 (40.9)
≥5	12 (3.0)
Parity n=146	
1	90 (61.6)
≥2	56 (38.4)
Abortion n=60	

1	48 (80)
≥ 2	12 (20)
Gestational age	
28-33w6d	9(2.2)
34-36w6d	22 (5.5)
37-41w6d	237 (58.8)
≥ 42	42 (10.4)
Unknown	93 (23.1)

The majority of the pregnant women were referred from health centers (387, 96%) and transported by the ambulance (72%) (table 2).

The majority of the women delivered through vaginal route (254, 63.3%) followed by Cesarean-section (148, 36.7%). The most common indication for C-section were non-reassuring fetal heart rate pattern (27%) followed by cephalo-pelvic disproportion/malpresentation/malposition (24.3%) then meconium staining in latent first stage of labor (22.3%). The majority of the babies were born alive (389, 96.5%). There were eight (2%) intra-partum fetal losses (Table 2).

Table 2. Source of referral, transportation, mode & place of delivery and perinatal outcome, among referred pregnant women for delivery, Addis Ababa, Ethiopia, 2019.

Characteristics		Number (%)
Source of referral N=403	21	
Health Center	1	387 (96.0)
Others		16 (4.0)
Transportation		
Ambulance		290 (72)
Others(Taxi, personal car)		113 (28)
Receiving hospital contacted before the women referred		
Yes		157 (39.0)

No	246 (61.0)	
Mode of delivery		
Vaginal delivery	229 (56.8)	
Assisted breech delivery	4 (1.0)	
C-section	148 (36.7)	
Instrumental delivery	21(5.2) 16 (4.0)	
Vacuum		
Forceps	5 (1.2)	
Laparatomy	1 (0.2)	
C-section indication N=148		
Non-reassuring fetal heart rate (NRFHR)	40 (27.0)	
Meconium in LFSOL	33 (22.3)	
Cephalopelvic disproportion (CPD)	21 (14.2)	
Malpresentation/malposition	15 (10.1)	
Previous C-scar with labor/labor abnormality	10 (6.8)	
Cord prolapse/presentation	7 (4.7)	
Non reassuring biophysical profile (NRBPP)	5 (3.4)	
АРН	4 (2.7)	
Others	13(8.8)	
Place of delivery N=403		
Labor ward/Operation room	386 (95.8)	
Emergency OPD	17 (4.2)	
Outcome N=403		
Alive	389 (96.5)	
Newborn referred to NICU Yes	137 (34)	

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		No	252 (62.5)
Still	birth		14 (3.5)
	Fetal heart beat positive	on arrival	8(2.0)
	Fetal heart beat negative	on arrival	6(1.5)

Among 403 referred women for childbirth, 71(17.6%) women experienced all the three delays. Almost three-fourth referred women (74.7%) experienced third delay followed by first delay (52.4%). Majority (366, 90.8%) women had at least one of the delays (Table 3).

Table 3. The frequency of the three delays among referred women, Addis Ababa, Ethiopia 2019.

Characteristics	Numbe	er (%)
N=403	Yes	Νο
First delay	211 (52.4)	192 (47.6)
Second delay	163 (40.4)	240(59.6)
Third delay	301(74.7)	102(25.3)
At least one delay	366 (90.8)	37 (9.2)
All three delay	71 (17.6)	332 (82.4)

** multiple answer possible

Twenty-nine (7.2%) women had severe maternal outcomes. The most common severe maternal outcome was blood transfusion (58.6%) followed by post partum hemorrhage (51.7%) then eclampsia (31%). Nearly three-fourth women with SMO (78.5%) had more than one complication (Figure 1).

The most common delays in women with SMO were third delay (58.6%) followed by first delay (51.7%). All most three-fourth women experienced at least one of the delays (79.3%) and one-quarter (24.1%) women had all the delays (table 4).

Statistically significant association was observed between SMO and third delay, at least one of the delays. Referred women with the third delay had 2.2 times (95% CI of 1.025-4.840) more likely to be a risk for severe maternal outcomes. Women who experienced at least one of the delays had 2.9 times (95% CI 1.093-7.620) more likely to be a risk for severe maternal outcomes (table 4).

After adjusted for age, gravidity, parity, educational level and marital status none of the delays were significant. This may be due to small sample of women with SMO. Also the three delays were not significant for age, gravidity, parity, educational level, gestational age and marital status.

Table 4. SMO and types of delays Addis Ababa, Ethiopia 2019.

Characteristic	S	Severe materna	l outcomes		
Types of delay		No	Yes	P-value	COR 95 % CI
		Number (%)	Number (%)		
First delay	No	178 (47.6%)	14 (48.3)		
	Yes	196 (52.4)	15 (51.7)	0.944	1.028(0.483-2.189)
Second delay	No	224 (59.9)	16 (55.2)		
	Yes	150 (40.1)	13 (44.8)	0.618	1.213(0.567-2.596)
Third delay	No	90(24.1)	12 (41.4)		
	Yes	284 (75.9)	15 (58.6)	0.043	2.227(1.025-4.840)
All delays	No	310 (82.9)	22 (75.9)		
	Yes	64 (17.1)	7 (24.1)	0.342	1.541(0.632-3.761)
At least one	No	31 (8.3)	6 (20.7)		
delay	Yes	343 (91.7)	23(79.3)	0.032	2.889(1.093-7.620)

**AOR not significant after adjusted for age, marital status, educational level, gestational age, gravidity, parity.

Eighty (19.9%) of referred women had at least one potential life threatening conditions (PLTC). The most common complications was hypertensive disorders 56 (70%) followed by blood transfusion 17 (21.3%) then post partum hemorrhage (table 5).

Table 5. Potentially life threatening conditions among referred women Addis Ababa, Ethiopia, 2019

Characteristics* N=80	Number (%)

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Ante partum hemorrhage (AP,PP)	13 (16.3)
Post partum hemorrhage (PPH)	15 (18.8)
Ruptured uterus	1 (1.25)
Coagulopathy	2 (2.5)
Hypertensive disorders	
Severe hypertension/Preeclampsia	50 (62.5)
Eclampsia	9 (11.3)
HELLP syndrome	2 (2.5)
Others	
Pulmonary edema	1 (1.25)
Shock	1(1.25)
Thrombocytopenia	2 (2.5)
Management indication of severity	
Transfusion of blood derivatives	17 (21.3)
Major surgical intervention(hysterectomy)	1(1.25)
*Multiple response possible	0,

All three types of delay were common in maternity services in the study hospitals; the most severe being the delay within the receiving hospital. The third delay was significantly associated with severe maternal outcomes.

Studies have found that distance to facilities is a clear barrier to women accessing health facilities (17) (18) but in Addis Ababa, proximity to services does not appear to be a problem, as the median distance to a facility that provides surgical services is 5 kilometers, well below the national average of 45 kilometers (19). Two-fifth of women had the second delay. When we compare with other studies it may be low however this proportion of the second delay is not expected because referring

facilities/catchment health centers are near to the receiving hospitals and expected to refer by ambulance (20).

This study showed the rate of occurrence of SMO indicators were higher than the findings of an earlier study done in other parts of the country (20) (21) (22) (23) and other countries (24-26). This high proportion SMO might be due to the fact that our study selected facilities are referral hospitals which are serving complicated cases and cases referred from other health facilities which were beyond their capacity/needed further interventions. This study showed that direct obstetric causes were the most common leading factors of SMO and the most common diagnosis were postpartum hemorrhage (52%) followed by eclampsia (31%) and the most common intervention was blood transfusion (58.6%). Obstetric hemorrhage and hypertensive disorders (eclampsia, hematological coagulation, and pulmonary edema) were found to be the top underlying complications among cases of SMO; similarly, hypertensive disorders and obstetric hemorrhage were the most common underlying causes of PLTC. This is comparable to the findings from studies in other parts of the country (21), (22) (23) and other countries (25-29) including sub-Saharan countries (30) (31).

Emergency obstetric care use by pregnant women is influenced by a complex interaction of factors leading to delay in decision-making, accessing services and receipt of proper care once a health facility is reached(30) (31). Receiving appropriate care after reaching at the health facility (delay three) was the most common (58.6%) followed by delay in seeking care (delay one) (51.7%) then reaching at the appropriate health facility (delay two) (44.8%) were identified among SMO and more than half of (58.6%) SMO cases had encountered at least one of the delays which were similar to study done elsewhere in the country (21) (32) (33); however delay two and one is seen less frequently than the findings from other countries (25) (30) (31) (34-37). This can be justified by overloaded of cases, limited hospitals capacity, a difference in socio-demographic characteristics of the study population and the proximity of health facilities.

In this study, failure to receiving appropriate care after reaching at the health facility (delay three) was found to have the strongest association with SMO: with a two-fold increase in the risk. This supports the WHO hypothesis relating a high case fatality in the hospital as an indicator for the presence of delay in receiving an adequate and proper treatment (38) and it indicates the poor performance of obstetrics services (39) (40). Seeking care from a facility that is incapacitated health facility and system, poor leadership (mismanagements of hospital resources, poor co-ordinations and lack of understanding of obstetrics emergencies) contribute to significant delay after reaching the health facility. These factors were reported as significant contributors to delay in several studies (21) (32) (34) (35). Studies from Tigray, 88% of all maternal deaths were attributed to health system failure (33). In our study, 59 % of SMO cases and both maternal deaths (MD) had health system related factors as a possible reason for delay three.

The limitation of this study include that it was conducted in two referral hospitals which often receive complicated cases and referred mother with complications; and the results might not be representative of other institutions and the community. In addition, cases might be missed because if they transferred to health centers and other hospitals after delivery for different reasons.

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However, we believe that this study gives the extent of delays and SMO &its indicators experienced by women who were referred for an emergency delivery. In fact, if the delay is so severe in these wellestablished referral centers one may expect it to be worse in some not well staffed and equipped centers. We decided to focus on women with severe maternal outcomes (SMO) rather than less severe forms of obstetric complications because their situations are the closest to maternal deaths.

Conclusion

The burden of severe maternal outcomes (SMO) is high due to preventable and/or treatable direct obstetric. The majority of the women in this study had serious delays in both making decision to seek care for birthing and in actually receiving care once at a hospital. We recommend strengthening health referral systems and addressing specific health systems bottlenecks during labor and birth in order to ensure no mother will be endanger. We also recommend a qualitative method of study (including focus group discussion, in-depth interview) and observing the tertiary hospitals set-up and readiness to manage high-risk pregnancies.

Abbreviations

AAHB-IRB: Addis Ababa Health Bureau-Institutional Review Board
BEmOC: Basic Emergency Obstetrics Care
CEmOC: Comprehensive Emergency Obstetrics Care
GMH: Gandhi Memorial Hospital
HELLP: Hemolysis, Elevated liver enzyme, Low Platelets
ICU: Intensive Care Unit
LW: Labor Ward
MD: Maternal Death Ratio
MNM: Maternal Near-Miss
MWH: Maternity Waiting Home
OPD: Out-Patient Department
PIH: Pregnancy Induced Hypertension
PLTC: Potential Life Threatening Conditions
PPH: Post Partum Hemorrhage
SMO: Severe Maternal Outcomes

WHO: World Health Organization

ZMH: Zewditu Memorial Hospital

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Author details

¹Endalkachew Mekonnen Assefa (MD, MPH), Addis Ababa University-College of Health Sciences Department of Obstetrics and Gynecology; e-mail-<u>endmekon@gmail.com</u>; phone no. +251-913605674: corresponding author

²Yemane Berhane (MD, PhD), Addis Continental Institute of Public Health; <u>yemaneberhane@gmail.com</u>

Contributors: EMA did design of the work, drafting the work, analysis, interpretation of data for the work and write-up the manuscript. YB advised the paper's scope, the draft of manuscript and did revising it critically for important intellectual content. Both authors approved the final version.

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Data availability statement: data are available from the corresponding author upon reasonable request.

References

1. Campbell O, Graham WJ. Measuring maternal mortality and morbidity: levels and trends. London: Lond Sch Hyg Trop Med. 1990;

2. Li XF, Fortney JA, Kotelchuck M, Glover LH. The postpartum period: the key to maternal mortality. Int J Gynaecol Obstet.1996;54:1–10.

3. Khan KS, Wojdyla D, Say L, Gulmezoglu AM, Van Look PF. WHO analysis of causes of maternal death: a systematic review.2006;Lancet 367:1066–1074.

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4. Fortney J. Emergency obstetric care: the keystone in the arch of safe motherhood. .; IJOG. 2001;(74):95–97.

5. World Health Organization (WHO). Maternal Mortality. Fact Sheet. Media Centre. 2016;

6. World Health Organization (WHO). True magnitude of stillbirths and maternal and neonatal deaths underreported. Media Cent. 2016;

7. Thaddeus S, Maine D. Too far to walk: maternal mortality in context. Soc Sci Med. 1994;38:1091–110.

8. The Effectiveness of Emergency Obstetric Referral Interventions in Developing Country Settings: A Systematic Review [Internet]. [cited 2018 Jul 28]. Available from: http://journals.plos.org/plosmedicine/article?id=10.1371/journal.pmed.1001264

9. Knight HE, Self A, Kennedy SH. Why Are Women Dying When They Reach Hospital on Time? A Systematic Review of the 'Third Delay.' PLoS ONE [Internet]. 2013 May 21 [cited 2018 Sep 7];8(5). Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3660500/

10. JSI Research and Training Institute. The Last 10 Kilometers (L10K). Emergency referral for pregnant women and newborns: A Rapid Community and Health System Assessment. BETA Development Consulting Firm, Addis Ababa. 2012 Jun;

11. U. Onwudiegwu, O. C.Ezechi. Emergency obstetric admissions: late referrals, misdiagnoses and consequences. J J Obstet Gynaecol. 2001;Volume 21(Issue 6).

12. Bossyns P, Abache R, Abdoulaye MS, Miyé H, Depoorter A-M, Van Lerberghe W. Monitoring the referral system through benchmarking in rural Niger: an evaluation of the functional relation between health centres and the district hospital. BMC Health Serv Res. 2006 Apr 12;6(1):51.

13.Giovine A, Ostrowski C. Technical report on Improving Transportation and Referral for Maternal Health: Knowledge Gaps and Recommendations. Wilson Cent. 2010;

14. Ronsmans C, Graham WJ, Lancet. Maternal Survival Series steering group: Maternal mortality: who, when, where, and why. Lancet. 2006;(368(9542)):1189–2000.

15. Qureshi RN, Sikandar R, Hoodbhoy Z, Bano R, Jabeen N. Referral pattern of emergencies in obstetrics: Implications for defining scope of services and policy. JPMA J Pak Med Assoc. 2016;66(12):1606–10.

16. Von Elm E, Altman DG, Egger M, Pocock SJ, Gotzsche PC, Vandenbroucke JP. The Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) Statement: guidelines for reporting observational studies.

17. McNamee P, Ternent L, Hussein J. Barriers in accessing maternal healthcare: evidence from low-and middle-income countries. Expert Rev Pharmacoecon Outcomes Res.2009;9(1):41–8.

18. Gabrysch SC, Oona MR. Still to far to walk: Literature review of the determinants of derlivery service use. BMC Pregnancy Childbirth. 2009;9(34).http://www.biomedcentral.com/1471-2393/9/34.

 Federal Ministry of Health, Addis Ababa Regional Health Bureau. Emergency Obstetric and Newborn Care Services: Standard Operation Manual for Activities of Catchment Teams in Addis Ababa.
 2012.

20. Dile M, Abate T, Seyum T. Proportion of maternal near misses and associated factors in referral hospitals of Amhara regional state, Northwest Ethiopia: institution based cross sectional study. GynecolObstet (Sunnyvale). 2015; 5:308. https://doi.org/10.4172/2161-0932.1000308.

21. Wondimagegnehu Sisay Woldeyes, Dejene Asefa and Geremew Muleta. Incidence and determinants of severe maternal outcome in Jimma University teaching hospital, south-West Ethiopia: a prospective cross-sectional study. Woldeyes Al BMC Pregnancy Childbirth 2018. 18:255(18:255).

22. Gebrehiwot Y, Tewolde BT. Improving maternity care in Ethiopia through facility based review of maternal deaths and near misses. Int J GynecolObstet. 2014; 127:S29–34.

23. Liyew EF, Yalew AW, Afework MF, Esse. Incidence and causes of maternal near-miss in selected hospitals of Addis Ababa, Ethiopia. ONE. 2017;12(6):e0179013. https://doi.org/10.1371/journal.pone.0179013.

24. Jabir M, et al. Maternal near miss and quality of maternal health care in Baghdad, Iraq. BMC Pregnancy Childbirth. 2013;13:11.

25. Leonardo Antonio Chavane, Patricia Bailey, Osvaldo Loquiha, Martinho Dgedge, Marc Aerts, and Marleen Temmerman. Maternal death and delays in accessing emergency obstetric care in Mozambique.BMC Pregnancy and Childbirth. (2018) 18:71 https://doi.org/10.1186/s12884-018-1699-z.

26. Syeda B.Mazhar, AfshanBatool, AngelaEmanuel, Arif T.Khan, ShireenBhutta.Severe maternal outcomes and their predictors among Pakistani women in the WHO Multicountry Survey on Maternal and Newborn Health. International Journal of Gynecology & Obstetrics Volume 129, Issue 1, April 2015, Pages 30-33.

27. Özge Tunçalp, Michelle J. Hindin, Kwame Adu-Bonsaffoh , Richard M., Adanu. Assessment of maternal near-miss and quality of care in a hospital based study in Accra, Ghana. Int J Gynecol Obstet 123 (2013) 58–63.

28. Say L, Souza JP, Pattinson RC. WHO working group on Maternal Mortality and Morbidity classifications. Maternal near miss–towards a standard tool for monitoring quality of maternal health care. 2009. 23:287–96.

29. Nelissen EJ, et al. Maternal near miss and mortality in a rural referral hospital in northern Tanzania: a cross-sectional study. BMC Pregnancy Childbirth. 2013;13:141.

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30. Oladapo et al. When getting there is not enough: a nationwide cross-sectional study of 998 maternal deaths and 1451 near-misses in public tertiary hospitals in a low-income country. BJOG.2015.www.bjog.org.DOI: 10.1111/1471-0528.13450.

31. Dan K Kaye, Othman Kakaire, and Michael O Osinde. Systematic review of the magnitude and case fatality ratio for severe maternal morbidity in sub-Saharan Africa between 1995 and 2010. BMC Pregnancy Childbirth. 2011;11(65).doi: 10.1186/1471-2393-11-65.

32. Ayele Geleto, Catherine Chojenta, Abdulbasit Musaand Deborah Loxton. Barriers to access and utilization of emergency obstetric care at health facilities in sub-Saharan Africa: a systematic review of literature. (2018) 7:183.https://doi.org/10.1186/s13643-018-0842-2.

33. Hailu S, Enqueselassie F, Berhane Y. Health facility-based maternal death audit in Tigray, Ethiopia. Ethiop J Health Dev. 2009; 23:115–9.

34. Rodolfo Carvalho Pacagnella, Jose Guilherme Cecatti, Maria Jose Osis & João, Paulo Souza. The role of delays in severe maternal morbidity and mortality: expanding the conceptual framework.Reproductive Health Matters 2012;20(39):155–163. www.rhm-elsevier.com. Doi:0.1016/S0968-8080(12)39601-8.

35. Rodolfo C Pacagnella, José G Cecatti, Mary A Parpinelli, Maria H Sousa, Samira M Haddad, Maria L Costa, João P Souza, Robert C Pattinson and the Brazilian Network for the Surveillance of Severe Maternal Morbidity, study group. Delays in receiving obstetric care and poor maternal outcomes: results from a national multicentre cross-sectional study.BMC Pregnancy and Childbirth 2014, 14:159.

36. Bouchra Assarag, Bruno Dujardin, Alexandre Delamou, Fatima-Zahra Meski,, Vincent De Brouwere. Determinants of Maternal Near-Miss in Morocco: Too Late, Too Far, Too Sloppy?PLOS ONE | January 22, 2015.DOI:10.1371/journal.pone.0116675.

37. David M. Goodman, Emmanuel K. Srofenyoh, Adeyemi J. Olufolabi, Sung Min Kim and Medge D. wen. The third delay: understanding waiting time for obstetric referrals at a large regional hospital in Ghana:BMC Pregnancy and Childbirth (2017) 17:216; DOI 10.1186/s12884-017-1407-4.

38. World Health Organization. Evaluating the quality of care for severe pregnancy complications: the WHO near-miss approach for maternal health. Geneva: WHO; 2011.

39. Saira Yunus, Shahnaz Kauser, Shazia Ali. Three "Delays" as a framework for critical analysis of maternal near miss and maternal mortality. Journal of South Asian Federation of Obstetrics and Gynecology, May-August 2013;5(2):57-59.

40. Filippi V, Ronsmans C, Gohou V, et al. Maternity wards or emergency obstetric rooms? Incidence of near-miss events in African hospitals. Acta Obstetricia et Gynecologica Scandinavica 2005; 84(1):11–16.

Figure 1. Types of severe maternal outcomes among referred pregnant women, Addis Ababa, Ethiopia, 2019

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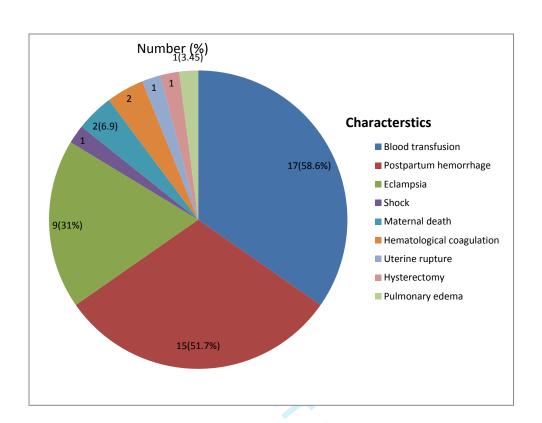


Figure 1. Types of severe maternal outcomes among referred pregnant women, Addis Ababa, Ethiopia, 2019.

Reporting checklist for cross sectional study.

Based on the STROBE cross sectional guidelines.

Instructions to authors

Complete this checklist by entering the page numbers from your manuscript where readers will find each of the items listed below. Your article may not currently address all the items on the checklist. Please modify your text to include the missing information. If you are certain that an item does not apply, please write "n/a" and provide a short explanation. Upload your completed checklist as an extra file when you submit to a journal. In your methods section, say that you used the STROBE cross sectional reporting guidelines, and cite them as: von Elm E, Altman DG, Egger M, Pocock SJ, Gotzsche PC, Vandenbroucke JP. The Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) Statement: guidelines for reporting observational studies. Page Reporting Item Number Title and abstract Title #i Indicate the study's design with a commonly used term in the title or the abstract Abstract #i Provide in the abstract an informative and balanced summary of

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6 7 8 9 10 11 12 13 14	Background /	<u>#1</u>	Explain the scientific background and rationale for the
	rationale		investigation being reported
	Objectives	<u>#2</u>	State specific objectives, including any prespecified hypotheses
15 16 17	Methods		
18 19 20	Study design	<u>#2</u>	Present key elements of study design early in the paper
21 22	Setting <u>#2</u>		Describe the setting, locations, and relevant dates, including
23 24 25			periods of recruitment, exposure, follow-up, and data collection
26 27 28	Eligibility criteria	<u>#3</u>	Give the eligibility criteria, and the sources and methods of
28 29 30 31			selection of participants.
32 33		<u>#3</u>	Clearly define all outcomes, exposures, predictors, potential
34 35			confounders, and effect modifiers. Give diagnostic criteria, if
36 37 38			applicable
39 40 41	Data sources /	<u>#3</u>	For each variable of interest give sources of data and details of
42 43	measurement		methods of assessment (measurement). Describe comparability
44 45			of assessment methods if there is more than one group. Give
46 47 48			information separately for for exposed and unexposed groups if
48 49 50			applicable.
51 52 53 54 55	Bias	<u>#n/a</u>	Describe any efforts to address potential sources of bias
56 57 58 59 60	Study size	<u>#3</u> For pe	Explain how the study size was arrived at eer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml

1 2	Quantitative	#4 Explain how quantitative variables were handled in			
3 4	variables		analyses. If applicable, describe which groupings were chosen,		
5 6 7			and why		
8 9 10	Statistical	<u>#4</u>	Describe all statistical methods, including those used to control		
11 12 13	methods		for confounding		
14 15	Statistical	<u>#n/a</u>	Describe any methods used to examine subgroups and		
16 17 18	methods		interactions		
19 20 21	Statistical	<u>#3</u>	Explain how missing data were addressed		
22 23 24	methods				
25 26	Statistical	<u>#4</u>	If applicable, describe analytical methods taking account of		
27 28 29	methods		sampling strategy		
51		<u>#n/a</u>	Describe any sensitivity analyses		
32 33 34 35	methods				
36 37 38	Results				
39 40	Participants	<u>#4</u>	Report numbers of individuals at each stage of study—eg		
41 42			numbers potentially eligible, examined for eligibility, confirmed		
43 44			eligible, included in the study, completing follow-up, and		
45 46 47			analysed. Give information separately for for exposed and		
47 48 49 50			unexposed groups if applicable.		
51 52 53			Note it was cross-sectional study		
54 55 56	Participants	<u>#n/a</u>	Give reasons for non-participation at each stage		
57 58			Note: all participants participated		
59 60		For pe	eer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml		

1 2 3	Participants	<u>#n/a</u>	Consider use of a flow diagram		
4 5 7 8 9 10 11	Descriptive data	<u>#4</u>	Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential		
12 13 14 15 16 17 18 19 20 21	Descriptive data	<u>#n/a</u>	confounders. Give information separately for exposed and unexposed groups if applicable. Indicate number of participants with missing data for each variable of interest		
22 23 24 25 26 27 28 29 30	Outcome data	<u>#6</u>	Report numbers of outcome events or summary measures. Give information separately for exposed and unexposed groups if applicable.		
 31 32 33 34 35 36 37 38 39 40 41 42 43 44 45 46 47 48 49 50 51 52 53 54 55 	Main results	#n/a	Give unadjusted estimates and, if applicable, confounder-		
	<u>man rooano</u>		adjusted estimates and their precision (eg, 95% confidence interval). Make clear which confounders were adjusted for and why they were included		
	Main results	<u>#n/a</u>	Report category boundaries when continuous variables were categorized		
	Main results	<u>#n/a</u>	If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period		
56 57 58 59 60	Other analyses	<u>#9</u> For p	Report other analyses done—e.g., analyses of subgroups and eer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml		

1			interactions, and sensitivity analyses			
2 3 4 5	Discussion					
6 7 8	Key results	<u>#10</u>	Summarise key results with reference to study objectives			
9 10 11	Limitations #1		Discuss limitations of the study, taking into account sources of			
12 13			potential bias or imprecision. Discuss both direction and			
14 15 16			magnitude of any potential bias.			
17 18	Interpretation	<u>#12</u>	Give a cautious overall interpretation considering objectives,			
19 20			limitations, multiplicity of analyses, results from similar studies,			
21 22 23 24			and other relevant evidence.			
24 25 26	Generalisability <u>#12</u>		Discuss the generalisability (external validity) of the study			
27 28			results			
29 30						
31 32	Other Information					
33 34	Funding	<u>#13</u>	Give the source of funding and the role of the funders for the			
35 36 37			present study and, if applicable, for the original study on which			
38 39 40			the present article is based			
41 42	 None The STROBE checklist is distributed under the terms of the Creative Commons Attribution License CC-BX. This checklist can be completed online using https://www.goodreports.org/.a.tool 					
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Delays in Emergency Obstetrics referrals in Addis Ababa hospitals: a facility-based, cross-sectional study

Endalkachew Mekonnen Assefa¹, Yemane Berhane²

¹ Addis Ababa University-College of Health Sciences (AAU-CHS); <u>endmekon@gmail.com</u>: corresponding author ² Addis Continental Institute of Public Health (ACIPH); <u>yemaneberhane@gmail.com</u>

Abstract

Objectives: To assess where the delays occur in the referral chain at most and maternal health outcomes based on the three delay model in Addis Ababa.

Design: The study was a facility-based cross-sectional study

Setting: Two public and tertiary hospitals in Addis Ababa

Participants: All pregnant women who were referred for only labor and delivery services after 28 weeks of gestation between December 2018 and February 2019 in Zewditu and Gandhi Memorial hospitals.

Primary and secondary outcome measures: The primary outcome was the type of delays from the three-delay model which met operationally defined time. Maternal health outcomes based on the three-delay model as a secondary outcome.

Results: A total of 403 pregnant women referred for delivery to the study hospitals were included in the study. Three-fourth (301, 74.7%) of referred pregnant women had the third delay (delayed receiving appropriate care); (211, 52.4%) had a first delay (delay in making a decision to seek care). Overall 366(90.8%) pregnant women had experienced at least one of the three delays and 71(17.6%) had all the three delays. Twenty-nine (7.2%) referred women had severe maternal outcomes (SMO). The most leading causes/diagnosis of SMO were blood transfusion 17 (58.6%) followed by postpartum hemorrhage 15 (52%) then eclampsia 9 (31%). In addition, women who experienced severe maternal outcomes were 2.9 times more likely to have at least one of the three delays.

Conclusion & recommendation: This study highlights the persistence of delays at all levels and especially delay three and its contribution to severe maternal outcomes. Strengthening health referral systems and addressing specific health system bottlenecks during labor and birth in order to ensure no mother will be endangered. We also recommend a qualitative method of study (focus group discussion and in-depth interview) and observing the tertiary hospitals set-up and readiness to manage obstetrics emergencies.

Key words: Addis Ababa, Emergency obstetrics referral, severe maternal outcomes, Three-delay model

Article summary

Strengths and limitations

This study gives the extent of delays and severe maternal outcomes experienced by women who were referred for labor and delivery.

This study focused on women with severe maternal outcomes (SMO) rather than less severe forms of obstetric complications because their situations are the closest to maternal deaths. Women after delivery might be missed if they transferred to health centers and to other hospitals after delivery for different reasons.

The results might not be representative of other institutions and the communities because it was conducted in two referral hospitals that often receive and treat complicated cases.

Introduction

The majority of maternal deaths are clustered around labour, delivery and the first 24 hours postpartum period (1),(2). It is estimated that just five conditions (postpartum hemorrhage; puerperal sepsis; preeclampsia and eclampsia; obstructed or prolonged labour, and complications of unsafe abortion), account for at least 60% of all maternal mortality (3). Life-threatening situations may develop rapidly and without warning, often in previously uncomplicated pregnancies(4).

Globally, it is recognized that significant inroads in maternal mortality cannot be made without dramatically increasing access to emergency obstetrical care (EmOC). The World Health Organization estimates that at least 88–98% of maternal deaths can be averted with timely access to existing, emergency obstetric interventions (5). This produces a triple return on investment, saving women and newborns and preventing stillbirths(6).

A number of factors can influence a woman's ability to access effective interventions to treat complications in the event of an obstetric emergency. Thaddeus and Maine (1994) group these into three broad categories using a classic, pathways-based framework known as the 'three delays model'(7).The 'three delays model' attempts to explain delays in women accessing emergency obstetric care as the result of 1) decision-making, 2) accessing services and 3) receipt of appropriate care once a health facility is reached.

Referral is often associated with the second delay of the three delays model– associated with reaching the appropriate level of care. But in fact, a referral system can reduce all three delays. If a population knows that a system is reliable and affordable, families may make the decision to seek care more quickly (the first delay(8).

The major obstacles that affect the referral system reported by both the health workers and women were: 1) financial barriers (for transportation and service payments at health facilities), 2) lack of means of transportation, 3) distance and 4) lack of awareness of services and the importance of services (9).

Factors associated with health-seeking behavior are multidimensional. Socio-cultural and economic problems, lack of awareness, the quality of health services, and infrastructure such as transport services all affect whether and where a woman will seek care, how long it will take to reach care, and whether she receives the appropriate care in a timely fashion(10).

Studies showed referrals in pregnancy and childbirth can be (1) institutional or self-referral, depending on the involvement of first line services; (2) antenatal, delivery or postnatal referral; and (3) elective or emergency referral. Pregnant women may referred due to demographic risks, obstetric historical risks, prenatal complications, and delivery and immediate postnatal complications(11). On the other hand, Studies show that high-risk prediction may not necessarily mean that the woman will have a complication and many women identified as being at risk go on to have normal deliveries(12).

Defining a framework and process for obstetric referrals may lead to reduction of maternal mortality and morbidity. Referral should be broadly defined to include not only transport, but it should be timely referral to minimize or prevent the delay for transportation (called second delay), and ensure prehospital care while transporting a patient to the referral facility(13), (14).

It is widely accepted that substantial reductions in maternal mortality and maternal near- miss are impossible to achieve without early decision-making to seek care, an effective referral system for complicated cases and receiving timely and appropriate care (9), (15) . Near miss cases represent most of the characteristics of maternal deaths, but occur more often (16). The near-miss approach assesses the gap between the actual use and optimal use of high-priority effective interventions in the prevention and management of severe maternal complications related to pregnancy and childbirth (17).

The objective of this study was to determine the types of delay and maternal health outcomes based on the three delay model among women who referred for labor and delivery. Results from this study may be an input for the hospitals, health bureau, policy-makers, and other stakeholders to act on bottlenecks of emergency obstetrics services by identifying the most common types of delay.

Methods

We used the STROBE cross-sectional reporting guidelines (18).

Study design

A facility-based cross-sectional study was conducted between December 10/2018-February 28/2019 in two government hospitals Zewditu Memorial Hospital (ZMH) and Gandhi Memorial Hospitals (GMH).

Study setting

This study conducted in tertiary hospitals located in the capital city of Ethiopia, Addis Ababa. Both hospitals are under Addis Ababa Health Bureau and affiliated with Addis Ababa University-College of Health Sciences. Gandhi Memorial Hospital is a referral maternity hospital and Zewditu Memorial Hospital is also a comprehensive referral hospital. Both hospitals are catchment hospitals for 40 (forty) health centers and other health facilities. Both hospitals are giving Comprehensive emergency obstetric care (CEmOC) and attending more than 17,000 deliveries per year.

The referral system for an obstetric emergency in Addis Ababa is organized to include Basic Emergency obstetric care (BEmOC) and CEmOC facilities. The referral system is developed to work both ways. Referral between health facilities is facilitated by the liaison office or Maternal Health Task Force. An ambulance system is organized to transport women accompanied by midwives. The midwife provides care during transportation and hand-over the mother to the receiving hospital care provider with a referral paper. In Addis Ababa, all maternity services including labour/delivery and the ambulance services are provided free of charge in all government health facilities. All components of BEmOC are expected to be given in the health centres.

Eligibility criteria

All pregnant women who were referred for only labor and delivery services after 28 weeks of gestation or baby outcome \geq 1 kg and delivered in the selected hospitals, and also who gave consent were included.

Data Collection Tools

After they gave birth, women were identified and interviewed from emergency OPD, labor ward, and inpatient wards every day before they discharged from the hospital by data collectors using pretested and structured questionnaires.

Data Collection Procedures

The referral papers reviewed and date, time and diagnosis of referral was recorded for each mother. The triage paper and patient chart are also reviewed including mode of transportation, date and time of arrival, sources of referrals, obstetrics performances, the time taken to admit/get the service after arrival, diagnosis at receiving hospital, gestational age, place and mode of delivery, newborn outcomes, and severe maternal complications types and managements. Women interviewed on socio-demographic characteristics, time interval to seek medical advice and reason for the delay to seek the care (if there was).

The completed questionnaires were reviewed by the principal investigator and supervisors. Incomplete questionnaires were filled if the women were not discharged; otherwise incomplete questionnaires were discarded.

Main outcomes and measures of the study

The three delays time frame was operationally defined through a consultative process involving six obstetrician & gynecologists (three from each hospital) who had working experience of 7-20 years in the selected hospitals. Accordingly, the first delay defined time elapsed between the recognition of complication/s, and the decision to transport and reach to a health facility, was considered if it took more than 60 minutes. The second delay was defined if the mother did not reach to referral hospitals within 60 minutes of referral. The third delay was if the mother didn't receive care or admitted within 30 minutes. Severe maternal outcomes (SMO) were any maternal complications including blood transfusion (any type & ≥ 2units), Post-partum hemorrhage, shock, eclampsia, uterine rupture, pulmonary edema, laparotomy, laboratory evidence of organ damage and/or maternal death during the process of delivery and/or before discharge from the hospital. Potentially life-threatening maternal conditions (PLTMC) was considered when the mother had at least one of the following; hemorrhagic complications, hypertensive disorders, and complications, end-organ injury, blood product transfusion, ICU admission, uterine rupture, hysterectomy/laparotomy.

Sample size: Single proportion formula used by assuming 50% of the referred women have a delay, degree of precision of a 5% (d), confidence interval of 95% (Z=1.96), assuming 5% non-response rate and the final the sample size was 403.

Statistical analysis

Data were entered in epi-info version 7.2.2.6 and transported to SPSS Version-21 statistics software for cleaning and analysis. Descriptive statistics were used to present women by their socio-demographic characteristics, referral diagnosis, diagnosis at receiving hospitals, obstetrics characteristics, mode of delivery, newborn outcomes, the three delays, and severe maternal outcomes. Severe maternal outcomes were analyzed for the three delays. The relationship between the three delays and SMO were examined using the multivariate logistic regression. The goodness of the model was tested by Hosmer-Lemeshow test for goodness-of-fit. Selected variables were included in the model to account for maternal characteristics differences other than delays in seeking and receiving care.

Ethical issues

Ethical clearance was obtained from both Addis Continental Institute of Public Health (ACIPH) and Addis Ababa Health Bureau (AAHB) Institution Review Board (IRB). Support letters were written to both study hospitals from AAHB-IRB to gain access.

Patient and public involvement:

Women asked for some questions after obtained informed consent on a voluntary basis. All the responses were given by the participants and the results obtained kept anonymous and confidential. There was no public involvement in the design, conduct, and interpretation of the study. Patients were not asked to advise on interpretation or writing up of results. We did not include patient involvement in the design of this study. We have presented a summary of the findings at medical and public health schools and among health providers in Addis Ababa and plan to continue presenting the results at professional society's conferences. Results were shared with the administration of both selected hospitals and Addis Ababa health Bureau to facilitate improved obstetrics services. There are no plans to disseminate the results of this research to study participants.

Results

Table 1 shows the descriptive information for socio-demographic and obstetric characteristics of referred pregnant women. The mean age of 403 pregnant women referred for labor delivery services was 26.47 ± 4.5 years and ranged from 18 to 43. The majorities were married (380,94.3%), and have completed at least secondary school (54.3%). Majority women were primigravida (56.1%), the mean gravidity was 1.77 ± 1.1 and ranged between 1 to 7. Most pregnant women (58.8%) were at term pregnancy (37 weeks- 41w6d) (Table1).

Table 1.Socio-demographic and obstetrics characteristics of respondents, Addis Ababa, Ethiopia, 2019 (n=403)

Characteristics n=403

Number (%)

Study Hospital	
Gandhi Memorial Hospital (GMH)	173 (42.9)
Zewditu Memorial Hospital (ZMH)	230 (57.1)
Age (years) median: 26 years (range 18-43)	
< 20	32 (7.9)
20-25	158 (39.2)
26-30	151 (37.5)
31-35	46 (11.4)
≥36	16 (4.0)
Marital status	
Married	380 (94.3)
Others(unmarried, divorced)	23 (5.7)
Educational level	
No formal education	49 (12.2)
Primary school	135 (33.5)
Secondary school	117 (29.0)
Preparatory	35 (8.7)
Vocational and above	67 (16.6)
Gravidity n=403	L
1	226 (56.1)
2-4	165 (40.9)
≥5	12 (3.0)
Parity n=146	
1	90 (61.6)
≥2	56 (38.4)

Abortion n=60	
1	48 (80)
≥2	12 (20)
Gestational age	
28-33w6d	9(2.2)
34-36w6d	22 (5.5)
37-41w6d	237 (58.8)
≥ 42	42 (10.4)
Unknown	93 (23.1)

The majority of the pregnant women were referred from health centers (387, 96%) and transported by the ambulance (72%) (Table 2).

The majority of the women delivered through vaginal route (254, 63.3%) followed by Cesarean-section (148, 36.7%). The most common indication for C-section was non-reassuring fetal heart rate pattern (27%) followed by cephalo-pelvic disproportion/malpresentation/malposition (24.3%) then meconium staining in the latent first stage of labor (22.3%). The majority of the babies were born alive (389, 96.5%). There were eight (2%) intrapartum fetal losses (Table 2).

Table 2. Source of referral, transportation, mode & place of delivery and perinatal outcome, among referred pregnant women for delivery, Addis Ababa, Ethiopia, 2019.

Characteristics	21	Number (%)
Source of referral N=403	1	
Health Center		387 (96.0)
Others		16 (4.0)
Transportation		
Ambulance		290 (72)
Others(Taxi, personal car)		113 (28)

Receiving hospital contacted before the women referred	
Yes	157 (39.0)
Νο	246 (61.0)
Mode of delivery	
Vaginal delivery	229 (56.8)
Assisted breech delivery	4 (1.0)
C-section	148 (36.7
Instrumental delivery	21(5.2)
Vacuum	16 (4.0)
Forceps	5 (1.2)
Laparatomy	1 (0.2)
C-section indication N=148	
Non-reassuring fetal heart rate (NRFHR)	40 (27.0)
Meconium in LFSOL	33 (22.3)
Cephalopelvic disproportion (CPD)	21 (14.2)
Malpresentation/malposition	15 (10.1)
Previous C-scar with labor/labor abnormality	10 (6.8)
Cord prolapse/presentation	7 (4.7)
Non reassuring biophysical profile (NRBPP)	7 (4.7) 5 (3.4)
АРН	4 (2.7)
Others	13(8.8)
Place of delivery	N=403
Labor ward/Operation room	386 (95.8)
Emergency OPD	17 (4.2)

Outcome	N=403	
Alive		389 (96.5)
Newborn referred to NICU	Yes	137 (34)
	No	252 (62.5)
Stillbirth		14 (3.5)
Fetal heart beat positive	on arrival	8(2.0)
Fetal heart beat negative	on arrival	6(1.5)

Among 403 referred women for childbirth, 71(17.6%) women experienced all the three delays. Almost three-fourth referred women (74.7%) were experienced the third delay followed by first delay (52.4%). The majority (366, 90.8%) women had at least one of the delays (Table 3).

Table 3. The frequency of the three delays among referred women, Addis Ababa, Ethiopia 2019.

Characteristics	Number (%)	
N=403	Yes	No
First delay	211 (52.4)	192 (47.6)
Second delay	163 (40.4)	240(59.6)
Third delay	301(74.7)	102(25.3)
At least one delay	366 (90.8)	37 (9.2)
All three delay	71 (17.6)	332 (82.4)

** multiple answer possible

Twenty-nine (7.2%) women had severe maternal outcomes. The most common severe maternal outcomes were blood transfusion (58.6%) followed by postpartum hemorrhage (51.7%) then eclampsia (31%). Nearly three-fourth women with SMO (78.5%) had more than one complication (Figure 1).

The most common delays in women with SMO were the third delay (58.6%) followed by first delay (51.7%). All most three-fourth women experienced at least one of the delays (79.3%) and one-quarter (24.1%) women had all the delays (Table 4).

Statistically, a significant association was observed between SMO and third delay, at least one of the delays. Referred women with the third delay had 2.2 times (95% CI of 1.025-4.840) more likely to be a

risk for severe maternal outcomes. Women who experienced at least one of the delays had 2.9 times (95% CI 1.093-7.620) more likely to be a risk for severe maternal outcomes (Table 4).

After adjusted for age, gravidity, parity, educational level, and marital status none of the delays were significant. This may be due to a small sample of women with SMO. Also, the three delays were not significant for age, gravidity, parity, educational level, gestational age, and marital status.

Characteristics		Severe maternal outcomes				
Types of delay		No	Yes	P-value	COR 95 % CI	
		Number (%)	Number (%)			
First delay	No	178 (47.6%)	14 (48.3)			
	Yes	196 (52.4)	15 (51.7)	0.944	1.028(0.483-2.189)	
Second delay	No	224 (59.9)	16 (55.2)			
	Yes	150 (40.1)	13 (44.8)	0.618	1.213(0.567-2.596)	
Third delay	No	90(24.1)	12 (41.4)			
	Yes	284 (75.9)	15 (58.6)	0.043	2.227(1.025-4.840)	
All delays	No	310 (82.9)	22 (75.9)			
	Yes	64 (17.1)	7 (24.1)	0.342	1.541(0.632-3.761)	
At least one	No	31 (8.3)	6 (20.7)			
delay	Yes	343 (91.7)	23(79.3)	0.032	2.889(1.093-7.620)	

Table 4. SMO and types of delays Addis Ababa, Ethiopia 2019.

**AOR not significant after adjusted for age, marital status, educational level, gestational age, gravidity, parity.

Eighty (19.9%) of referred women had at least one potential life- threatening conditions (PLTC). The most common complications were hypertensive disorders 56 (70%) followed by blood transfusion 17 (21.3%) then postpartum hemorrhage (Table 5).

Table 5. Potentially life threatening conditions among referred women Addis Ababa, Ethiopia, 2019

Characteristics* N=80	Number (%)

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Ante partum hemorrhage (AP,PP)	13 (16.3)
Post partum hemorrhage (PPH)	15 (18.8)
Ruptured uterus	1 (1.25)
Coagulopathy	2 (2.5)
Hypertensive disorders	
Severe hypertension/Preeclampsia	50 (62.5)
Eclampsia	9 (11.3)
HELLP syndrome	2 (2.5)
Others	
Pulmonary edema	1 (1.25)
Shock	1(1.25)
Thrombocytopenia	2 (2.5)
Management indication of severity	
Transfusion of blood derivatives	17 (21.3)
Major surgical intervention(hysterectomy)	1(1.25)
*Multiple response possible	0,

Discussion

All three types of delay were common in the study hospitals; the most severe being the delay within the receiving hospital. The third delay was significantly associated with severe maternal outcomes.

Studies have found that distance to facilities is a clear barrier to women accessing health facilities (19) (20) but in Addis Ababa, proximity to services does not appear to be a problem, as the median distance to a facility that provides surgical services is 5 kilometers, well below the national average of 45 kilometers (21). Two-fifth of women had the second delay. When we compare with other studies it may be low however this proportion of the second delay is not expected because referring facilities/catchment health centers are near to the receiving hospitals and expected to refer by ambulance (22).

This study showed the rate of occurrence of SMO indicators were higher than the findings of an earlier study done in other parts of the country (22) (23) (24) (25) and other countries (26-28). This high proportion SMO might be due to the fact that our study selected facilities are referral hospitals which are serving complicated cases and cases referred from other health facilities which were beyond their capacity/needed further interventions. This study showed that direct obstetric causes were the most common leading factors of SMO and the most common diagnosis were postpartum hemorrhage (52%) followed by eclampsia (31%) and the most common intervention was blood transfusion (58.6%). Obstetric hemorrhage and hypertensive disorders (eclampsia, hematological coagulation, and pulmonary edema) were found to be the top underlying complications among cases of SMO; similarly, hypertensive disorders and obstetric hemorrhage were the most common underlying causes of PLTC. This is comparable to the findings from studies in other parts of the country (23), (24) (25) and other countries (27-31) including sub-Saharan countries (32) (33).

Emergency obstetric care use by pregnant women is influenced by a complex interaction of factors leading to delay in decision-making, accessing services and receipt of proper care once a health facility is reached(32) (33). Receiving appropriate care after reaching the health facility (delay three) was the most common (58.6%) followed by a delay in seeking care (delay one) (51.7%) then reaching at the appropriate health facility (delay two) (44.8%) were identified among SMO and more than half of (58.6%) SMO cases had encountered at least one of the delays which were similar to study done elsewhere in the country (23) (34) (35); however, delay two and one is seen less frequently than the findings from other countries (27) (32) (33) (36-39). This can be justified by overloaded of cases, limited hospitals capacity, the difference in socio-demographic characteristics of the study population and the proximity of health facilities.

In this study, failure to receiving appropriate care after reaching the health facility (delay three) was found to have the strongest association with SMO: with a two-fold increase in the risk. This supports the WHO hypothesis relating a high case fatality in the hospital as an indicator for the presence of delay in receiving adequate and proper treatment (19) and it indicates the poor performance of obstetrics services (40) (41). Seeking care from a facility that is incapacitated health facility and system, poor leadership (mismanagements of hospital resources, poor co-ordinations and lack of understanding of obstetrics emergencies) contribute to significant delay after reaching the health facility. These factors were reported as significant contributors to delay in several studies (23) (34) (36) (37). Studies from Tigray, 88% of all maternal deaths were attributed to health system failure (35). In our study, 59 % of SMO cases and both maternal deaths (MD) had a health system related factors as a possible reason for delay three.

The limitation of this study include that it was conducted in two referral hospitals which often receive complicated cases and referred women with complications; and the results might not be representative of other institutions and the community. In addition, cases might be missed because if they transferred to health centers and other hospitals after delivery for different reasons.

However, we believe that this study gives the extent of delays and SMO &its indicators experienced by women who were referred for an emergency delivery. In fact, if the delay is so severe in these well-

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established referral centers one may expect it to be worse in some not well-staffed and equipped centers. We decided to focus on women with severe maternal outcomes (SMO) rather than less severe forms of obstetric complications because their situations are the closest to maternal deaths.

Conclusion

The burden of severe maternal outcomes (SMO) is high due to preventable and/or treatable direct obstetric. The majority of the women in this study had serious delays in both making decisions to seek care for birthing and in actually receiving care once at a hospital. We recommend strengthening health referral systems and addressing specific health systems bottlenecks during labor and delivery in order to ensure no woman will be endangered. We also recommend a qualitative method of study (including focus group discussion, in-depth interview) and observing the tertiary hospitals setup and readiness to manage high-risk pregnancies.

Abbreviations

AAHB-IRB: Addis Ababa Health Bureau-Institutional Review Board

BEmOC: Basic Emergency Obstetrics Care

CEmOC: Comprehensive Emergency Obstetrics Care

GMH: Gandhi Memorial Hospital

HELLP: Hemolysis, Elevated liver enzyme, Low Platelets

ICU: Intensive Care Unit

LW: Labor Ward

MD: Maternal Death Ratio

MNM: Maternal Near-Miss

MWH: Maternity Waiting Home

OPD: Out-Patient Department

PIH: Pregnancy Induced Hypertension

PLTC: Potential Life Threatening Conditions

PPH: Post Partum Hemorrhage

SMO: Severe Maternal Outcomes

WHO: World Health Organization

ZMH: Zewditu Memorial Hospital

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Author Details

¹Endalkachew Mekonnen Assefa (MD, MPH), Addis Ababa University-College of Health Sciences Department of Obstetrics and Gynecology; e-mail-<u>endmekon@gmail.com</u>; phone no. +251-913605674: corresponding author

²Yemane Berhane (MD, Ph.D.), Addis Continental Institute of Public Health; <u>yemaneberhane@gmail.com</u>

Contributors: EMA did design of the work, drafting the work, analysis, interpretation of data for the work and write-up the manuscript. YB advised the paper's scope, the draft of the manuscript and did revise it critically for important intellectual content. Both authors approved the final version.

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References

1. Campbell O, Graham WJ. Measuring maternal mortality and morbidity: levels and trends. London: Lond Sch Hyg Trop Med. 1990;

2. Li XF, Fortney JA, Kotelchuck M, Glover LH. The postpartum period: the key to maternal mortality. Int J Gynaecol Obstet.1996;54:1–10.

3. Khan KS, Wojdyla D, Say L, Gulmezoglu AM, Van Look PF. WHO analysis of causes of maternal death: a systematic review.2006; Lancet 367:1066–1074.

4. Fortney J. Emergency obstetric care: the keystone in the arch of safe motherhood. .; IJOG. 2001;(74):95–97.

5. World Health Organization (WHO). Maternal Mortality. Fact Sheet. Media Centre. 2016;

6. World Health Organization (WHO). True magnitude of stillbirths and maternal and neonatal deaths underreported. Media Cent. 2016;

7. Thaddeus S, Maine D. Too far too walk: maternal mortality in context. Soc Sci Med. 1994;38:1091– 110.

8. The Effectiveness of Emergency Obstetric Referral Interventions in Developing Country Settings: A Systematic Review [Internet]. [cited 2018 Jul 28]. Available from: http://journals.plos.org/plosmedicine/article?id=10.1371/journal.pmed.1001264

9. Knight HE, Self A, Kennedy SH. Why Are Women Dying When They Reach Hospital on Time? A Systematic Review of the 'Third Delay.' PLoS ONE [Internet]. 2013 May 21 [cited 2018 Sep 7];8(5). Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3660500/

10. JSI Research and Training Institute. The Last 10 Kilometers (L10K). Emergency referral for pregnant women and newborns: A Rapid Community and Health System Assessment. BETA Development Consulting Firm, Addis Ababa. 2012 Jun;

11. U. Onwudiegwu, O. C.Ezechi. Emergency obstetric admissions: late referrals, misdiagnoses and consequences. J J Obstet Gynaecol. 2001; Volume 21 (Issue 6).

12. Bossyns P, Abache R, Abdoulaye MS, Miyé H, Depoorter A-M, Van Lerberghe W. Monitoring the referral system through benchmarking in rural Niger: an evaluation of the functional relation between health centres and the district hospital. BMC Health Serv Res. 2006 Apr 12;6(1):51.

13.Giovine A, Ostrowski C. Technical report on Improving Transportation and Referral for Maternal Health: Knowledge Gaps and Recommendations. Wilson Cent. 2010;

14. Ronsmans C, Graham WJ, Lancet. Maternal Survival Series steering group: Maternal mortality: who, when, where, and why. Lancet. 2006;(368(9542)):1189–2000.

15. Qureshi RN, Sikandar R, Hoodbhoy Z, Bano R, Jabeen N. Referral pattern of emergencies in obstetrics: Implications for defining scope of services and policy. JPMA J Pak Med Assoc. 2016;66 (12):1606–10.

16. Ellen Nelissen, Estomih Mduma, Jacqueline Broerse, Hege Ersdal, Bjørg Evjen-Olsen, Jos van Roosmalen, Jelle Stekelenburg. Applicability of the WHO Maternal Near Miss Criteria in a Low-Resource Setting. PLOS ONE | www.plosone.org. April 2013 | Volume 8 |Issue 4 | e6124.

17. World Health Organization. Evaluating the quality of care for severe pregnancy complications: the WHO near-miss approach for maternal health. Geneva: WHO; 2011.

18. Von Elm E, Altman DG, Egger M, Pocock SJ, Gotzsche PC, Vandenbroucke JP. The Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) Statement: guidelines for reporting observational studies.

19. McNamee P, Ternent L, Hussein J. Barriers in accessing maternal healthcare: evidence from low-and middle-income countries. Expert Rev Pharmacoecon Outcomes Res.2009;9(1):41–8.

20. Gabrysch SC, Oona MR. Still to far to walk: Literature review of the determinants of delivery service use. BMC Pregnancy Childbirth. 2009;9(34).http://www.biomedcentral.com/1471-2393/9/34.

 Federal Ministry of Health, Addis Ababa Regional Health Bureau. Emergency Obstetric and Newborn Care Services: Standard Operation Manual for Activities of Catchment Teams in Addis Ababa.
 2012.

22. Dile M, Abate T, Seyum T. Proportion of maternal near misses and associated factors in referral hospitals of Amhara regional state, Northwest Ethiopia: institution-based cross-sectional study. GynecolObstet (Sunnyvale). 2015; 5:308. https://doi.org/10.4172/2161-0932.1000308.

23. Wondimagegnehu Sisay Woldeyes, Dejene Asefa and Geremew Muleta. Incidence and determinants of severe maternal outcome in Jimma University teaching hospital, south-West Ethiopia: a prospective cross-sectional study. Woldeyes Al BMC Pregnancy Childbirth 2018. 18:255(18:255).

24. Gebrehiwot Y, Tewolde BT. Improving maternity care in Ethiopia through facility based review of maternal deaths and near misses. Int J GynecolObstet. 2014; 127:S29–34.

25. Liyew EF, Yalew AW, Afework MF, Esse. Incidence and causes of maternal near-miss in selected hospitals of Addis Ababa, Ethiopia. ONE. 2017;12(6):e0179013. https://doi.org/10.1371/journal.pone.0179013.

26. Jabir M, et al. Maternal near miss and quality of maternal health care in Baghdad, Iraq. BMC Pregnancy Childbirth. 2013;13:11.

27. Leonardo Antonio Chavane, Patricia Bailey, Osvaldo Loquiha, Martinho Dgedge, Marc Aerts, and Marleen Temmerman. Maternal death and delays in accessing emergency obstetric care in Mozambique.BMC Pregnancy and Childbirth. (2018) 18:71 https://doi.org/10.1186/s12884-018-1699-z.

BMJ Open

28.Syeda B.Mazhar, AfshanBatool, AngelaEmanuel, Arif T.Khan, ShireenBhutta.Severe maternal outcomes and their predictors among Pakistani women in the WHO Multicountry Survey on Maternal and Newborn Health. International Journal of Gynecology & Obstetrics Volume 129, Issue 1, April 2015, Pages 30-33.

29. Özge Tunçalp, Michelle J. Hindin, Kwame Adu-Bonsaffoh , Richard M., Adanu. Assessment of maternal near-miss and quality of care in a hospital- based study in Accra, Ghana. Int J Gynecol Obstet 123 (2013) 58–63.

30. Say L, Souza JP, Pattinson RC. WHO working group on Maternal Mortality and Morbidity classifications. Maternal near miss-towards a standard tool for monitoring quality of maternal health care. 2009. 23:287–96.

31. Nelissen EJ, et al. Maternal near miss and mortality in a rural referral hospital in northern Tanzania: a cross-sectional study. BMC Pregnancy Childbirth. 2013;13:141.

32. Oladapo et al. When getting there is not enough: a nationwide cross-sectional study of 998 maternal deaths and 1451 near-misses in public tertiary hospitals in a low-income country. BJOG.2015.www.bjog.org.DOI: 10.1111/1471-0528.13450.

33. Dan K Kaye, Othman Kakaire, and Michael O Osinde. Systematic review of the magnitude and case fatality ratio for severe maternal morbidity in sub-Saharan Africa between 1995 and 2010. BMC Pregnancy Childbirth. 2011;11(65).doi: 10.1186/1471-2393-11-65.

34. Ayele Geleto, Catherine Chojenta, Abdulbasit Musaand Deborah Loxton. Barriers to access and utilization of emergency obstetric care at health facilities in sub-Saharan Africa: a systematic review of literature. (2018) 7:183.https://doi.org/10.1186/s13643-018-0842-2.

35. Hailu S, Enqueselassie F, Berhane Y. Health facility-based maternal death audit in Tigray, Ethiopia. Ethiop J Health Dev. 2009; 23:115–9.

36. Rodolfo Carvalho Pacagnella, Jose Guilherme Cecatti, Maria Jose Osis & João, Paulo Souza. The role of delays in severe maternal morbidity and mortality: expanding the conceptual framework.Reproductive Health Matters 2012;20(39):155–163. www.rhm-elsevier.com. Doi:0.1016/S0968-8080(12)39601-8.

37. Rodolfo C Pacagnella, José G Cecatti, Mary A Parpinelli, Maria H Sousa, Samira M Haddad, Maria L Costa, João P Souza, Robert C Pattinson and the Brazilian Network for the Surveillance of Severe Maternal Morbidity, study group. Delays in receiving obstetric care and poor maternal outcomes: results from a national multicentre cross-sectional study.BMC Pregnancy and Childbirth 2014, 14:159.

38. Bouchra Assarag, Bruno Dujardin, Alexandre Delamou, Fatima-Zahra Meski,, Vincent De Brouwere. Determinants of Maternal Near-Miss in Morocco: Too Late, Too Far, Too Sloppy?PLOS ONE | January 22, 2015.DOI:10.1371/journal.pone.0116675.

39. David M. Goodman, Emmanuel K. Srofenyoh, Adeyemi J. Olufolabi, Sung Min Kim and Medge D. wen. The third delay: understanding waiting time for obstetric referrals at a large regional hospital in Ghana:BMC Pregnancy and Childbirth (2017) 17:216; DOI 10.1186/s12884-017-1407-4.

40. Saira Yunus, Shahnaz Kauser, Shazia Ali. Three "Delays" as a framework for critical analysis of maternal near miss and maternal mortality. Journal of South Asian Federation of Obstetrics and Gynecology, May-August 2013;5(2):57-59.

41. Filippi V, Ronsmans C, Gohou V, et al. Maternity wards or emergency obstetric rooms? Incidence of near-miss events in African hospitals. Acta Obstetricia et Gynecologica Scandinavica 2005; 84(1):11–16.

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Figure 1. Types of severe maternal outcomes among referred pregnant women, Addis Ababa, Ethiopia,

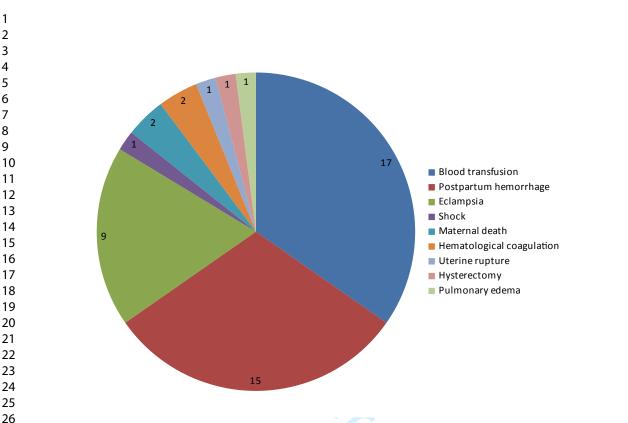


Figure 1. Types of severe maternal outcomes among referred pregnant women, Addis Ababa, Ethiopia, 2019.

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			Page
		Reporting Item	Number
Title and abstract			
Title	<u>#i</u>	Indicate the study's design with a commonly used term in the title or the abstract	
Abstract	<u>#i</u>	Provide in the abstract an informative and balanced summary of what was done and what was found	
Introduction			
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rationale			
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Methods			
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		follow-up, and data collection	
Eligibility criteria	<u>#3</u>	Give the eligibility criteria, and the sources and methods of selection of participants.	
	<u>#3</u>	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers.	
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1			Give diagnostic criteria, if applicable
2 3	Data sources /	<u>#3</u>	For each variable of interest give sources of data and details of methods of assessment
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5 6 7			Give information separately for for exposed and unexposed groups if applicable.
, 8 9 10 11	Bias	<u>#3</u>	Describe any efforts to address potential sources of bias
12 13 14	Study size	<u>#4</u>	Explain how the study size was arrived at
15	Quantitative	#4	Explain how quantitative variables were handled in the analyses. If applicable, describe which
16 17	variables		groupings were chosen, and why
18 19 20	Statistical methods	<u>#4</u>	Describe all statistical methods, including those used to control for confounding
21 22	Statistical methods	<u>#4</u>	Describe any methods used to examine subgroups and interactions
23 24 25	Statistical methods	<u>#3</u>	Explain how missing data were addressed
26 27	Statistical methods	<u>#3</u>	If applicable, describe analytical methods taking account of sampling strategy
28 29	Statistical methods	<u>#3</u>	Describe any sensitivity analyses
30 31 32	Results		
33	Participants	<u>#3</u>	Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined
34 35			for eligibility, confirmed eligible, included in the study, completing follow-up, and analysed.
36 37			Give information separately for for exposed and unexposed groups if applicable.
38 39 40			Note it was cross-sectional study Give reasons for non-participation at each stage
41 42	Participants	<u>#3</u>	Give reasons for non-participation at each stage
43 44 45			Note: all participants participated
45 46 47 48 49	Participants	<u>#n/a</u>	Consider use of a flow diagram
50 51 52 53 54 55	Descriptive data	<u>#4</u>	Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders. Give information separately for exposed and unexposed groups if applicable.
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1	Outcome data	<u>#6</u>	Report numbers of outcome events or summary measures. Give information separately for					
2 3 4 5			exposed and unexposed groups if applicable.					
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12 13	Main results	<u>#10</u>	Report category boundaries when continuous variables were categorized					
14 15 16 17	Main results	<u>#n/a</u>	If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period					
18 19 20 21	Other analyses	<u>#10</u>	Report other analyses done—e.g., analyses of subgroups and interactions, and sensitivity analyses					
22 23 24	Discussion							
24 25 26	Key results	<u>#11</u>	Summarise key results with reference to study objectives					
27 28 29 30	Limitations	<u>#12</u>	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias.					
30 31 32 33 34	Interpretation	<u>#12</u>	Give a cautious overall interpretation considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence.					
35 36	Generalisability	<u>#12</u>	Discuss the generalisability (external validity) of the study results					
37 38	Other							
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41 42 43 44	Funding	<u>#13</u>	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based					
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Delays in Emergency Obstetrics referrals in Addis Ababa hospitals, Ethiopia: a facility-based, cross-sectional study

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Keywords:	Human resource management < HEALTH SERVICES ADMINISTRATION & MANAGEMENT, Health policy < HEALTH SERVICES ADMINISTRATION & MANAGEMENT, Quality in health care < HEALTH SERVICES ADMINISTRATION & MANAGEMENT		





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Delays in Emergency Obstetrics referrals in Addis Ababa hospitals, Ethiopia: a facility-based, crosssectional study

Endalkachew Mekonnen Assefa¹, Yemane Berhane²

¹ Addis Ababa University-College of Health Sciences (AAU-CHS); <u>endmekon@gmail.com</u>: corresponding author ² Addis Continental Institute of Public Health (ACIPH); <u>yemaneberhane@gmail.com</u>

Abstract

i

Objectives: To assess where the delays occur in the referral chain at most and maternal health outcomes based on the three delay model in Addis Ababa, Ethiopia.

Design: The study was a facility-based cross-sectional study

Setting: Two public and tertiary hospitals in Addis Ababa

Participants: All pregnant women who were referred for only labor and delivery services after 28 weeks of gestation between December 2018 and February 2019 in Zewditu and Gandhi Memorial hospitals.

Primary and secondary outcome measures: The primary outcome was the type of delays from the three-delay model which met operationally defined time. Maternal health outcomes based on the three-delay model as a secondary outcome.

Results: A total of 403 pregnant women referred for delivery to the study hospitals were included in the study. Three-fourth (301, 74.7%) of referred pregnant women had the third delay (delayed receiving appropriate care); (211, 52.4%) had a first delay (delay in making a decision to seek care). Overall 366(90.8%) pregnant women had experienced at least one of the three delays and 71(17.6%) had all the three delays. Twenty-nine (7.2%) referred women had severe maternal outcomes (SMO). The most leading causes/diagnosis of SMO were blood transfusion 17 (58.6%) followed by postpartum hemorrhage 15 (52%) then eclampsia 9 (31%). In addition, women who experienced severe maternal outcomes were 2.9 times more likely to have at least one of the three delays.

Conclusion & recommendation: This study highlights the persistence of delays at all levels and especially delay three and its contribution to severe maternal outcomes. Strengthening health referral systems and addressing specific health system bottlenecks during labor and birth in order to ensure no mother will be endangered. We also recommend a qualitative method of study (focus group discussion and in-depth interview) and observing the tertiary hospitals set-up and readiness to manage obstetrics emergencies.

Key words: Addis Ababa, Emergency obstetrics referral, Severe maternal outcomes, Three-delay model

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-	Article summary
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9	Strengths and limitations
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12	This study gives the extent of delays and severe maternal outcomes experienced by
13	women who were referred for labor and delivery.
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16	This study focused on women with severe maternal outcomes (SMO) rather than less
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Introduction

According to the World Health Organization's (WHO) report on maternal mortality trends, about 295, 000 women died during and following pregnancy and childbirth in 2017 (1). Similarly, the 2019 WHO maternal mortality fact sheet reported that approximately 810 women die every day from pregnancy related complications. The vast majority of these deaths (94%) occurred in low-resource settings, and most could have been prevented (2). Sub-Saharan Africa alone accounted for roughly two-thirds (196, 000) of maternal deaths and Ethiopia is among thus countries (1, 2).

Globally, it is recognized that significant inroads maternal mortality cannot be made without dramatically increasing access to emergency obstetrical care (EmOC). The World Health Organization estimates that at least 88–98% of maternal deaths can be averted with timely access to existing, emergency obstetric interventions (3). This produces a triple return on investment, saving women and newborns and preventing stillbirths(4).

A number of factors can influence a woman's ability to access effective interventions to treat complications in the event of an obstetric emergency. Thaddeus and Maine (1994) group these into three broad categories using a classic, pathways-based framework known as the 'three delays model' (5). The 'three delays model' attempts to explain delays in women accessing emergency obstetric care as the result of 1) decision-making, 2) accessing services and 3) receipt of appropriate care once a health facility is reached.

Referral is often associated with the second delay of the three delays model– associated with reaching the appropriate level of care. But in fact, a referral system can reduce all three delays. If a population knows that a system is reliable and affordable, families may make the decision to seek care more quickly (the first delay(6).

The major obstacles that affect the referral system reported by both the health workers and women were: 1) financial barriers (for transportation and service payments at health facilities), 2) lack of means of transportation, 3) distance and 4) lack of awareness of services and the importance of services (7).

Factors associated with health-seeking behavior are multidimensional. Socio-cultural and economic problems, lack of awareness, the quality of health services, and infrastructure such as transport services all affect whether and where a woman will seek care, how long it will take to reach care, and whether she receives the appropriate care in a timely fashion(8).

Studies showed referrals in pregnancy and childbirth can be: (1) institutional or self-referral, depending on the involvement of first-line services; (2) antenatal, delivery or postnatal referral; and (3) elective or emergency referral. Pregnant women may be referred due to demographic risks, obstetric historical risks, prenatal complications, and delivery and immediate postnatal complications(9). On the other hand, Studies show that high-risk prediction may not necessarily mean that the woman will have a complication and many women identified as being at risk go on to have normal deliveries(10).

Defining a framework and process for obstetric referrals may lead to a reduction of maternal mortality and morbidity. Referral should be broadly defined to include not only transport, but it should be a timely referral to minimize or prevent the delay for transportation (called the second delay) and ensure prehospital care while transporting a patient to the referral facility(11), (12).

It is widely accepted that substantial reductions in maternal mortality and maternal near- miss are impossible to achieve without early decision-making to seek care, an effective referral system for complicated cases and receiving timely and appropriate care (7), (13). Near- miss cases represent most of the characteristics of maternal deaths but occur more often (14). The near-miss approach assesses the gap between the actual use and optimal use of high-priority effective interventions in the prevention and management of severe maternal complications related to pregnancy and childbirth (15).

The objective of this study was to determine the types of delay and maternal health outcomes based on the three delay model among women who referred for labor and delivery. Results from this study may be an input for the hospitals, health bureau, policy-makers, and other stakeholders to act on bottlenecks of emergency obstetrics services by identifying the most common types of delay.

Methods

We used the STROBE cross-sectional reporting guidelines (16).

Study design

A facility-based cross-sectional study was conducted between December 10/2018-February 28/2019 in two government hospitals Zewditu Memorial Hospital (ZMH) and Gandhi Memorial Hospitals (GMH).

Study setting

This study conducted in tertiary hospitals located in the capital city of Ethiopia, Addis Ababa. Both hospitals are under Addis Ababa Health Bureau and affiliated with Addis Ababa University-College of Health Sciences. Gandhi Memorial Hospital is a referral maternity hospital and Zewditu Memorial Hospital is also a comprehensive referral hospital. Both hospitals are catchment hospitals for 40 (forty) health centers and other health facilities. Both hospitals are giving Comprehensive emergency obstetric care (CEmOC) and attending more than 17,000 deliveries per year.

The referral system for an obstetric emergency in Addis Ababa is organized to include Basic Emergency obstetric care (BEmOC) and CEmOC facilities. The referral system is developed to work both ways. Referral between health facilities is facilitated by the liaison office or Maternal Health Task Force. An ambulance system is organized to transport women accompanied by midwives. The midwife provides care during transportation and hand-over the mother to the receiving hospital care provider with a referral paper. In Addis Ababa, all maternity services including labour/delivery and the ambulance services are provided free of charge in all government health facilities. All hospitals (including primary, secondary and tertiary), Maternity and Child hospitals are expected to give CEmOC: On the other hand, all components of BEmOC are expected to be given in the health centres, medium clinics, and specialty clinics.

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Eligibility criteria

All pregnant women who were referred for only labor and delivery services after 28 weeks of gestation or baby outcome \geq 1 kg and delivered in the selected hospitals, and also who gave consent were included.

Data Collection Tools

After they gave birth, women were identified and interviewed from emergency OPD, labor ward, and inpatient wards every day before they discharged from the hospital by data collectors using pretested and structured questionnaires.

Data Collection Procedures

The referral papers reviewed and date, time and diagnosis of referral was recorded for each mother. The triage paper and patient chart are also reviewed including mode of transportation, date and time of arrival, sources of referrals, obstetrics performances, the time taken to admit/get the service after arrival, diagnosis at receiving hospital, gestational age, place and mode of delivery, newborn outcomes, and severe maternal complications types and managements. Women interviewed on socio-demographic characteristics, time interval to seek medical advice and reason for the delay to seek the care (if there was).

The completed questionnaires were reviewed by the principal investigator and supervisors. Incomplete questionnaires were filled if the women were not discharged; otherwise incomplete questionnaires were discarded.

Main outcomes and measures of the study

The three delays time frame was operationally defined through a consultative process involving six obstetricians & gynecologists (three from each hospital) who had working experience of 7-20 years in the selected hospitals. Accordingly, the first delay defined time elapsed between the recognition of complication/s, and the decision to transport and reach a health facility, was considered if it took more than 60 minutes. The second delay was defined if the mother did not reach to referral hospitals within 60 minutes of referral. The third delay was if the mother didn't receive care or admitted within 30 minutes. Severe maternal outcomes (SMO) were any maternal complications including blood transfusion (any type & ≥ 2units), Post-partum hemorrhage, shock, eclampsia, uterine rupture, pulmonary edema, laparotomy, laboratory evidence of organ damage and/or maternal death during the process of delivery and/or before discharge from the hospital. Potentially life-threatening maternal conditions (PLTMC) was considered when the mother had at least one of the following; hemorrhagic complications, hypertensive disorders, and complications, end-organ injury, blood product transfusion, ICU admission, uterine rupture, hysterectomy/laparotomy.

Sample size: Single proportion formula used by assuming 50% of the referred women have a delay, degree of precision of a 5% (d), confidence interval of 95% (Z=1.96), assuming 5% non-response rate and the final the sample size was 403.

Statistical analysis

Data were entered in epi-info version 7.2.2.6 and transported to SPSS Version-21 statistics software for cleaning and analysis. Descriptive statistics were used to present women by their socio-demographic characteristics, referral diagnosis, diagnosis at receiving hospitals, obstetrics characteristics, mode of delivery, newborn outcomes, the three delays, and severe maternal outcomes. Severe maternal outcomes were analyzed for the three delays. The relationship between the three delays and SMO were examined using the multivariate logistic regression. The goodness of the model was tested by the Hosmer-Lemeshow test for goodness-of-fit. Selected variables were included in the model to account for maternal characteristics differences other than delays in seeking and receiving care.

Ethical issues

Ethical clearance was obtained from both Addis Continental Institute of Public Health (ACIPH) and Addis Ababa Health Bureau (AAHB) Institution Review Board (IRB). Support letters were written to both study hospitals from AAHB-IRB to gain access. Women asked for some questions after obtained informed consent on a voluntary basis. All the responses were given by the participants and the results obtained kept anonymous and confidential.

Patient and public involvement:

There was no public involvement in the design, conduct, and interpretation of the study. Patients were not asked to advise on interpretation or writing up of results. We did not include patient involvement in the design of this study. We have presented a summary of the findings at medical and public health schools and among health providers in Addis Ababa and plan to continue presenting the results at professional society's conferences. Results were shared with the administration of both selected hospitals and Addis Ababa health Bureau to facilitate improved obstetrics services. There are no plans to disseminate the results of this research to study participants.

Results

Table 1 shows the descriptive information for socio-demographic and obstetric characteristics of referred pregnant women. The mean age of 403 pregnant women referred for labor-delivery services was 26.47 ± 4.5 years and ranged from 18 to 43. The majority were married (380,94.3%), and have completed at least secondary school (54.3%).The majority of women were primigravida (56.1%), the mean gravidity was 1.77 ± 1.1 and ranged between 1 to 7. Most pregnant women (58.8%) were at term pregnancy (37 weeks- 41w6d) (Table1).

Characteristics n=403	Number (%)
Study Hospital	
Gandhi Memorial Hospital (GMH)	173 (42.9)
Zewditu Memorial Hospital (ZMH)	230 (57.1)
Age (years) median: 26 years (range 18-43)	
< 20	32 (7.9)
20-25	158 (39.2)
26-30	151 (37.5)
31-35	46 (11.4)
≥36	16 (4.0)
Marital status	
Married	380 (94.3)
Others(unmarried, divorced)	23 (5.7)
Educational level	
No formal education	49 (12.2)
Primary school	135 (33.5)
Secondary school	117 (29.0)
Preparatory	35 (8.7)
Vocational and above	67 (16.6)
Gravidity n=403	
1	226 (56.1)
2-4	165 (40.9)
≥5	12 (3.0)

Table 1. Socio-demographic and obstetrics characteristics of respondents. Addis Ababa. Ethiopia. 2019

Parity n=146	
1	90 (61.6)
≥2	56 (38.4)
Abortion n=60	
1	48 (80)
≥2	12 (20)
Gestational age	
28-33w6d	9(2.2)
34-36w6d	22 (5.5)
37-41w6d	237 (58.8)
≥ 42	42 (10.4)
Unknown	93 (23.1)

The majority of the pregnant women were referred from health centers (387, 96%) and transported by the ambulance (72%) (Table 2).

The majority of the women delivered through vaginal route (254, 63.3%) followed by Cesarean-section (148, 36.7%). The most common indication for C-section was non-reassuring fetal heart rate pattern (27%) followed by cephalo-pelvic disproportion/malpresentation/malposition (24.3%) then meconium staining in the latent first stage of labor (22.3%). The majority of the babies were born alive (389, 96.5%). There were eight (2%) intrapartum fetal losses (Table 2).

Table 2. Source of referral, transportation, mode & place of delivery and perinatal outcome, among referred pregnant women for delivery, Addis Ababa, Ethiopia, 2019.

387 (96.0)
16 (4.0)

Ambulance	290 (72)
Others(Taxi, personal car)	113 (28)
Receiving hospital contacted before the women referred	
Yes	157 (39.0)
Νο	246 (61.0)
Mode of delivery	
Vaginal delivery	229 (56.8)
Assisted breech delivery	4 (1.0)
C-section	148 (36.7
Instrumental delivery	21(5.2)
Vacuum	16 (4.0)
Forceps	5 (1.2)
Laparatomy	1 (0.2)
C-section indication N=148	
Non-reassuring fetal heart rate (NRFHR)	40 (27.0)
Meconium in LFSOL	33 (22.3)
Cephalopelvic disproportion (CPD)	21 (14.2)
Malpresentation/malposition	15 (10.1)
Previous C-scar with labor/labor abnormality	10 (6.8)
Cord prolapse/presentation	7 (4.7)
Non reassuring biophysical profile (NRBPP)	5 (3.4)
АРН	4 (2.7)
Others	13(8.8)

Place of delivery		N=403	
Labor ward/Operation roo	m	386 (95.8)	
Emergency OPD		17 (4.2)	
Outcome	N=403		
Alive		389 (96.5)	
Newborn referred to NICU Yes		137 (34)	
	No	252 (62.5)	
Stillbirth		14 (3.5)	
Fetal heart beat po	sitive on arrival	8(2.0)	
Fetal heart beat neg	gative on arrival	6(1.5)	

Among 403 referred women for childbirth, 71(17.6%) women experienced all the three delays. Almost three-fourth referred women (74.7%) were experienced the third delay followed by the first delay (52.4%). The majority (366, 90.8%) women had at least one of the delays (Table 3).

Table 3. The frequency of the three delays among referred women, Addis Ababa, Ethiopia 2019.

Characteristics	Number (%)	
N=403	Yes	No
First delay	211 (52.4)	192 (47.6)
Second delay	163 (40.4)	240(59.6)
Third delay	301(74.7)	102(25.3)
At least one delay	366 (90.8)	37 (9.2)
All three delay	71 (17.6)	332 (82.4)

** multiple answer possible

Twenty-nine (7.2%) women had severe maternal outcomes. The most common severe maternal outcomes were blood transfusion (58.6%) followed by postpartum hemorrhage (51.7%) then eclampsia (31%). Nearly three-fourth women with SMO (78.5%) had more than one complication (Figure 1).

The most common delays in women with SMO were the third delay (58.6%) followed by the first delay (51.7%). All most three-fourth women experienced at least one of the delays (79.3%) and one-quarter (24.1%) women had all the delays (Table 4).

Statistically, a significant association was observed between SMO and third delay, at least one of the delays. Referred women with the third delay had 2.2 times (95% CI of 1.025-4.840) more likely to be a risk for severe maternal outcomes. Women who experienced at least one of the delays had 2.9 times (95% CI 1.093-7.620) more likely to be a risk for severe maternal outcomes (Table 4).

After adjusted for age, gravidity, parity, educational level, gestational age, and marital status none of the delays were significant. This may be due to a small sample of women with SMO.

Characteristics		Severe maternal outcomes			
Types of delay		No	Yes	P-value	COR 95 % CI
		Number (%)	Number (%)		
First delay	No	178 (47.6%)	14 (48.3)		
	Yes	196 (52.4)	15 (51.7)	0.944	1.028(0.483-2.189)
Second delay	No	224 (59.9)	16 (55.2)		
	Yes	150 (40.1)	13 (44.8)	0.618	1.213(0.567-2.596)
Third delay	No	90(24.1)	12 (41.4)		
	Yes	284 (75.9)	15 (58.6)	0.043	2.227(1.025-4.840)
All delays	No	310 (82.9)	22 (75.9)		
	Yes	64 (17.1)	7 (24.1)	0.342	1.541(0.632-3.761)
At least one	No	31 (8.3)	6 (20.7)		
delay	Yes	343 (91.7)	23(79.3)	0.032	2.889(1.093-7.620)

Table 4. SMO and types of delays Addis Ababa, Ethiopia 2019.

**AOR not significant after adjusted for age, marital status, educational level, gestational age, gravidity, parity.

Eighty (19.9%) of referred women had at least one potential life- threatening conditions (PLTC). The most common complications were hypertensive disorders 56 (70%) followed by blood transfusion 17 (21.3%) then postpartum hemorrhage (Table 5).

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Characteristics* N=80	Number (%)	
Hemorrhagic complications		
Ante partum hemorrhage (AP,PP)	13 (16.3)	
Post partum hemorrhage (PPH)	15 (18.8)	
Ruptured uterus	1 (1.25)	
Coagulopathy	2 (2.5)	
Hypertensive disorders		
Severe hypertension/Preeclampsia	50 (62.5)	
Eclampsia	9 (11.3)	
HELLP syndrome	2 (2.5)	
Dthers		
Pulmonary edema	1 (1.25)	
Shock	1(1.25)	
Thrombocytopenia	2 (2.5)	
Management indication of severity	2	
Transfusion of blood derivatives	17 (21.3)	
Major surgical intervention(hysterectomy)	1(1.25)	
*Multiple response possible		

Discussion

All three types of delays were common in the study hospitals; the most severe being the delay within the receiving hospital. The third delay was significantly associated with severe maternal outcomes.

Studies have found that distance to facilities is a clear barrier to women accessing health facilities (17) (18) but in Addis Ababa, proximity to services does not appear to be a problem, as the median distance to a facility that provides surgical services is 5 kilometers, well below the national average of 45 kilometers (19). Two-fifth of women had the second delay. When we compare with other studies it may be low however this proportion of the second delay is not expected because referring facilities/catchment health centers are near to the receiving hospitals and expected to refer by ambulance (20).

This study showed the rate of occurrence of SMO indicators were higher than the findings of an earlier study done in other parts of the country (20-23) and other countries (24-26). This high proportion SMO might be due to the fact that our study selected facilities are referral hospitals that are serving complicated cases and cases referred from other health facilities that were beyond their capacity/needed further interventions. This study showed that direct obstetric causes were the most common leading factors of SMO and the most common diagnosis were postpartum hemorrhage (52%) followed by eclampsia (31%) and the most common intervention was blood transfusion (58.6%). Obstetric hemorrhage and hypertensive disorders (eclampsia, hematological coagulation, and pulmonary edema) were found to be the top underlying complications among cases of SMO; similarly, hypertensive disorders and obstetric hemorrhage were the most common underlying causes of PLTC. This is comparable to the findings from studies in other parts of the country (23), (24) (25) and other countries (25-29) including sub-Saharan countries (30) (31).

Emergency obstetric care use by pregnant women is influenced by a complex interaction of factors leading to delay in decision-making, accessing services and receipt of proper care once a health facility is reached(30) (31). Receiving appropriate care after reaching the health facility (delay three) was the most common (58.6%) followed by a delay in seeking care (delay one) (51.7%) then reaching at the appropriate health facility (delay two) (44.8%) were identified among SMO and more than half of (58.6%) SMO cases had encountered at least one of the delays which were similar to study done elsewhere in the country (21) (32) (33); however, delay two and one is seen less frequently than the findings from other countries (25) (30) (31) (34-37). This can be justified by the overloaded cases, limited hospital capacity, the difference in socio-demographic characteristics of the study population and the proximity of health facilities.

In this study, failure to receiving appropriate care after reaching the health facility (delay three) was found to have the strongest association with SMO: with a two-fold increase in the risk. This supports the WHO hypothesis relating a high case fatality in the hospital as an indicator for the presence of delay in receiving adequate and proper treatment (17) and it indicates the poor performance of obstetrics services (38) (39). Seeking care from a facility that is incapacitated health facility and system, poor leadership (mismanagements of hospital resources, poor co-ordinations and lack of understanding of

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obstetrics emergencies) contribute to significant delay after reaching the health facility. These factors were reported as significant contributors to delay in several studies (21) (32) (34) (35). Studies from Tigray, 88% of all maternal deaths were attributed to health system failure (33). In our study, 59 % of SMO cases and both maternal deaths (MD) had health system-related factors as a possible reason for delay three.

The limitation of this study include that it was conducted in two referral hospitals which often receive complicated cases and referred women with complications; the results might not be representative of other institutions and the community. In addition, cases might be missed because if they transferred to health centers and other hospitals after delivery for different reasons.

However, we believe that this study gives the extent of delays and SMO &its indicators experienced by women who were referred for an emergency delivery. In fact, if the delay is so severe in these wellestablished referral centers one may expect it to be worse in some not well-staffed and equipped centers. We decided to focus on women with severe maternal outcomes (SMO) rather than less severe forms of obstetric complications because their situations are the closest to maternal deaths.

Conclusion

The burden of severe maternal outcomes (SMO) is high due to preventable and/or treatable direct obstetric causes. The majority of the women in this study had serious delays in both making decisions to seek care for birthing and in actually receiving care once at a hospital. We recommend strengthening health referral systems and addressing specific health systems bottlenecks during labor and delivery in order to ensure no woman will be endangered. We also recommend a qualitative method of study (including focus group discussion, in-depth interview) and observing the tertiary hospitals set-up and readiness to manage high-risk pregnancies.

Abbreviations AAHB-IRB: Addis Ababa Health Bureau-Institutional Review Board BEmOC: Basic Emergency Obstetrics Care

CEmOC: Comprehensive Emergency Obstetrics Care

GMH: Gandhi Memorial Hospital

HELLP: Hemolysis, Elevated liver enzyme, Low Platelets

ICU: Intensive Care Unit

LW: Labor Ward

MD: Maternal Death Ratio

MNM: Maternal Near-Miss

MWH: Maternity Waiting Home

OPD: Out-Patient Department

PIH: Pregnancy Induced Hypertension

PLTC: Potential Life Threatening Conditions

PPH: Post Partum Hemorrhage

SMO: Severe Maternal Outcomes

WHO: World Health Organization

ZMH: Zewditu Memorial Hospital

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Author Details

¹Endalkachew Mekonnen Assefa (MD, MPH), Addis Ababa University-College of Health Sciences Department of Obstetrics and Gynecology; e-mail-<u>endmekon@gmail.com</u>; phone no. +251-913605674: corresponding author

²Yemane Berhane (MD, Ph.D.), Addis Continental Institute of Public Health; <u>yemaneberhane@gmail.com</u>

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References

1. WHO, et al. Trends in maternal mortality: 2000 to 2017. Geneva: WHO; 2019. [https://www.who.int/reproductivehealth/publications/maternal-mortality-2000-2017/en/]

2. WHO. Maternal mortality; fact sheets: WHO. 19 September 2019. [https://www.who.int/newsroom/fact-sheets/detail/maternal-mortality]

3. World Health Organization (WHO). Maternal Mortality. Fact Sheet. Media Centre. 2016;

4. World Health Organization (WHO). True magnitude of stillbirths and maternal and neonatal deaths underreported. Media Cent. 2016;

5. Thaddeus S, Maine D. Too far too walk: maternal mortality in context. Soc Sci Med. 1994;38:1091– 110.

6. The Effectiveness of Emergency Obstetric Referral Interventions in Developing Country Settings: A Systematic Review [Internet]. [cited 2018 Jul 28]. Available from: http://journals.plos.org/plosmedicine/article?id=10.1371/journal.pmed.1001264

7. Knight HE, Self A, Kennedy SH. Why Are Women Dying When They Reach Hospital on Time? A Systematic Review of the 'Third Delay.' PLoS ONE [Internet]. 2013 May 21 [cited 2018 Sep 7];8(5). Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3660500/

8. JSI Research and Training Institute. The Last 10 Kilometers (L10K). Emergency referral for pregnant women and newborns: A Rapid Community and Health System Assessment. BETA Development Consulting Firm, Addis Ababa. 2012 Jun;

9. U. Onwudiegwu, O. C.Ezechi. Emergency obstetric admissions: late referrals, misdiagnoses and consequences. J J Obstet Gynaecol. 2001; Volume 21 (Issue 6).

10. Bossyns P, Abache R, Abdoulaye MS, Miyé H, Depoorter A-M, Van Lerberghe W. Monitoring the referral system through benchmarking in rural Niger: an evaluation of the functional relation between health centres and the district hospital. BMC Health Serv Res. 2006 Apr 12;6(1):51.

11. Giovine A, Ostrowski C. Technical report on Improving Transportation and Referral for Maternal Health: Knowledge Gaps and Recommendations. Wilson Cent. 2010;

12. Ronsmans C, Graham WJ, Lancet. Maternal Survival Series steering group: Maternal mortality: who, when, where, and why. Lancet. 2006;(368(9542)):1189–2000.

13. Qureshi RN, Sikandar R, Hoodbhoy Z, Bano R, Jabeen N. Referral pattern of emergencies in obstetrics: Implications for defining scope of services and policy. JPMA J Pak Med Assoc. 2016;66 (12):1606–10.

14. Ellen Nelissen, Estomih Mduma, Jacqueline Broerse, Hege Ersdal, Bjørg Evjen-Olsen, Jos van Roosmalen, Jelle Stekelenburg. Applicability of the WHO Maternal Near Miss Criteria in a Low-Resource Setting. PLOS ONE | www.plosone.org. April 2013 | Volume 8 | Issue 4 | e6124.

15. World Health Organization. Evaluating the quality of care for severe pregnancy complications: the WHO near-miss approach for maternal health. Geneva: WHO; 2011.

16. Von Elm E, Altman DG, Egger M, Pocock SJ, Gotzsche PC, Vandenbroucke JP. The Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) Statement: guidelines for reporting observational studies.

17. McNamee P, Ternent L, Hussein J. Barriers in accessing maternal healthcare: evidence from low-and middle-income countries. Expert Rev Pharmacoecon Outcomes Res.2009;9(1):41–8.

18. Gabrysch SC, Oona MR. Still to far to walk: Literature review of the determinants of delivery service use. BMC Pregnancy Childbirth. 2009;9(34).http://www.biomedcentral.com/1471-2393/9/34.

 Federal Ministry of Health, Addis Ababa Regional Health Bureau. Emergency Obstetric and Newborn Care Services: Standard Operation Manual for Activities of Catchment Teams in Addis Ababa.
 2012.

20. Dile M, Abate T, Seyum T. Proportion of maternal near misses and associated factors in referral hospitals of Amhara regional state, Northwest Ethiopia: institution-based cross-sectional study. GynecolObstet (Sunnyvale). 2015; 5:308. https://doi.org/10.4172/2161-0932.1000308.

21. Wondimagegnehu Sisay Woldeyes, Dejene Asefa and Geremew Muleta. Incidence and determinants of severe maternal outcome in Jimma University teaching hospital, south-West Ethiopia: a prospective cross-sectional study. Woldeyes Al BMC Pregnancy Childbirth 2018. 18:255(18:255).

22. Gebrehiwot Y, Tewolde BT. Improving maternity care in Ethiopia through facility based review of maternal deaths and near misses. Int J GynecolObstet. 2014; 127:S29–34.

23. Liyew EF, Yalew AW, Afework MF, Esse. Incidence and causes of maternal near-miss in selected hospitals of Addis Ababa, Ethiopia. ONE. 2017;12(6):e0179013. https://doi.org/10.1371/journal.pone.0179013.

24. Jabir M, et al. Maternal near miss and quality of maternal health care in Baghdad, Iraq. BMC Pregnancy Childbirth. 2013;13:11.

25. Leonardo Antonio Chavane, Patricia Bailey, Osvaldo Loquiha, Martinho Dgedge, Marc Aerts, and Marleen Temmerman. Maternal death and delays in accessing emergency obstetric care in Mozambique.BMC Pregnancy and Childbirth. (2018) 18:71 https://doi.org/10.1186/s12884-018-1699-z.

BMJ Open

26.Syeda B.Mazhar, AfshanBatool, AngelaEmanuel, Arif T.Khan, ShireenBhutta.Severe maternal outcomes and their predictors among Pakistani women in the WHO Multicountry Survey on Maternal and Newborn Health. International Journal of Gynecology & Obstetrics Volume 129, Issue 1, April 2015, Pages 30-33.

27. Özge Tunçalp, Michelle J. Hindin, Kwame Adu-Bonsaffoh , Richard M., Adanu. Assessment of maternal near-miss and quality of care in a hospital- based study in Accra, Ghana. Int J Gynecol Obstet 123 (2013) 58–63.

28. Say L, Souza JP, Pattinson RC. WHO working group on Maternal Mortality and Morbidity classifications. Maternal near miss-towards a standard tool for monitoring quality of maternal health care. 2009. 23:287–96.

29. Nelissen EJ, et al. Maternal near miss and mortality in a rural referral hospital in northern Tanzania: a cross-sectional study. BMC Pregnancy Childbirth. 2013;13:141.

30. Oladapo et al. When getting there is not enough: a nationwide cross-sectional study of 998 maternal deaths and 1451 near-misses in public tertiary hospitals in a low-income country. BJOG.2015.www.bjog.org.DOI: 10.1111/1471-0528.13450.

31. Dan K Kaye, Othman Kakaire, and Michael O Osinde. Systematic review of the magnitude and case fatality ratio for severe maternal morbidity in sub-Saharan Africa between 1995 and 2010. BMC Pregnancy Childbirth. 2011;11(65).doi: 10.1186/1471-2393-11-65.

32. Ayele Geleto, Catherine Chojenta, Abdulbasit Musaand Deborah Loxton. Barriers to access and utilization of emergency obstetric care at health facilities in sub-Saharan Africa: a systematic review of literature. (2018) 7:183.https://doi.org/10.1186/s13643-018-0842-2.

33. Hailu S, Enqueselassie F, Berhane Y. Health facility-based maternal death audit in Tigray, Ethiopia. Ethiop J Health Dev. 2009; 23:115–9.

34. Rodolfo Carvalho Pacagnella, Jose Guilherme Cecatti, Maria Jose Osis & João, Paulo Souza. The role of delays in severe maternal morbidity and mortality: expanding the conceptual framework.Reproductive Health Matters 2012;20(39):155–163. www.rhm-elsevier.com. Doi:0.1016/S0968-8080(12)39601-8.

35. Rodolfo C Pacagnella, José G Cecatti, Mary A Parpinelli, Maria H Sousa, Samira M Haddad, Maria L Costa, João P Souza, Robert C Pattinson and the Brazilian Network for the Surveillance of Severe Maternal Morbidity, study group. Delays in receiving obstetric care and poor maternal outcomes: results from a national multicentre cross-sectional study.BMC Pregnancy and Childbirth 2014, 14:159.

36. Bouchra Assarag, Bruno Dujardin, Alexandre Delamou, Fatima-Zahra Meski,, Vincent De Brouwere. Determinants of Maternal Near-Miss in Morocco: Too Late, Too Far, Too Sloppy?PLOS ONE | January 22, 2015.DOI:10.1371/journal.pone.0116675.

37. David M. Goodman, Emmanuel K. Srofenyoh, Adeyemi J. Olufolabi, Sung Min Kim and Medge D. wen. The third delay: understanding waiting time for obstetric referrals at a large regional hospital in Ghana:BMC Pregnancy and Childbirth (2017) 17:216; DOI 10.1186/s12884-017-1407-4.

38. Saira Yunus, Shahnaz Kauser, Shazia Ali. Three "Delays" as a framework for critical analysis of maternal near miss and maternal mortality. Journal of South Asian Federation of Obstetrics and Gynecology, May-August 2013;5(2):57-59.

39. Filippi V, Ronsmans C, Gohou V, et al. Maternity wards or emergency obstetric rooms? Incidence of near-miss events in African hospitals. Acta Obstetricia et Gynecologica Scandinavica 2005; 84(1):11–16.

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Figure 1. Types of severe maternal outcomes among referred pregnant women, Addis Ababa, Ethiopia,

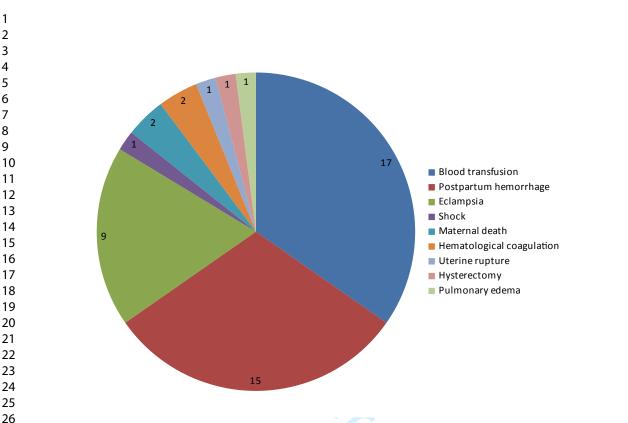


Figure 1. Types of severe maternal outcomes among referred pregnant women, Addis Ababa, Ethiopia, 2019.

Reporting checklist for cross sectional study.

Based on the STROBE cross sectional guidelines.

Instructions to authors

Complete this checklist by entering the page numbers from your manuscript where readers will find each of the items listed below.

Your article may not currently address all the items on the checklist. Please modify your text to include the missing information. If you are certain that an item does not apply, please write "n/a" and provide a short explanation.

Upload your completed checklist as an extra file when you submit to a journal.

In your methods section, say that you used the STROBE cross sectional reporting guidelines, and cite them as:

von Elm E, Altman DG, Egger M, Pocock SJ, Gotzsche PC, Vandenbroucke JP. The Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) Statement: guidelines for reporting observational studies.

		Reporting Item	Number	
Title and abstract				
Title	<u>#i</u>	Indicate the study's design with a commonly used term in the title or the abstract		
Abstract	<u>#i</u>	Provide in the abstract an informative and balanced summary of what was done and what was found		
Introduction				
Background /	<u>#1</u>	Explain the scientific background and rationale for the investigation being reported		
rationale				
Objectives	<u>#2</u>	State specific objectives, including any prespecified hypotheses		
Methods				
Study design	<u>#2</u>	Present key elements of study design early in the paper		
Setting	<u>#2</u>	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure,		
		follow-up, and data collection		
Eligibility criteria	<u>#3</u>	Give the eligibility criteria, and the sources and methods of selection of participants.		
	<u>#3</u>	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers.		
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1			Give diagnostic criteria, if applicable
2 3	Data sources /	<u>#3</u>	For each variable of interest give sources of data and details of methods of assessment
4	measurement		(measurement). Describe comparability of assessment methods if there is more than one group.
5 6 7			Give information separately for for exposed and unexposed groups if applicable.
, 8 9 10 11	Bias	<u>#n/a</u>	Describe any efforts to address potential sources of bias
12 13 14	Study size	<u>#3</u>	Explain how the study size was arrived at
15	Quantitative	<u>#3</u>	Explain how quantitative variables were handled in the analyses. If applicable, describe which
16 17 18	variables		groupings were chosen, and why
19 20	Statistical methods	<u>#4</u>	Describe all statistical methods, including those used to control for confounding
21 22 23	Statistical methods	<u>#4</u>	Describe any methods used to examine subgroups and interactions
24 25	Statistical methods	<u>#3</u>	Explain how missing data were addressed
26 27	Statistical methods	<u>#3</u>	If applicable, describe analytical methods taking account of sampling strategy
28 29 30	Statistical methods	<u>#n/a</u>	Describe any sensitivity analyses
31 32	Results		
33 34	Participants	<u>#4</u>	Report numbers of individuals at each stage of study-eg numbers potentially eligible, examined
35			for eligibility, confirmed eligible, included in the study, completing follow-up, and analysed.
36 37 38 39 40			Give information separately for for exposed and unexposed groups if applicable. Give reasons for non-participation at each stage
41 42 43 44 45	Participants	<u>#4</u>	Give reasons for non-participation at each stage
46 47	Participants	<u>#n/a</u>	Consider use of a flow diagram
48 49			
48 49 50 51 52 53 54	Descriptive data	<u>#4</u>	Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders. Give information separately for exposed and unexposed groups if applicable.
48 49 50 51 52 53	Descriptive data	<u>#4</u> <u>#5</u>	exposures and potential confounders. Give information separately for exposed and unexposed

be

1	Outcome data	<u>#5</u>	Report numbers of outcome events or summary measures. Give information separately for
2 3 4 5			exposed and unexposed groups if applicable.
6 7 8 9 10 11	Main results	<u>#9</u>	Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 95% confidence interval). Make clear which confounders were adjusted for and why they were included
12 13	Main results	<u>#10</u>	Report category boundaries when continuous variables were categorized
14 15 16 17	Main results	<u>#n/a</u>	If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period
18 19 20 21	Other analyses	<u>#10</u>	Report other analyses done—e.g., analyses of subgroups and interactions, and sensitivity analyses
22 23 24	Discussion		
25 26	Key results	<u>#11</u>	Summarise key results with reference to study objectives
27 28 29 30	Limitations	<u>#12</u>	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias.
31 32 33 34	Interpretation	<u>#12</u>	Give a cautious overall interpretation considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence.
35 36	Generalisability	<u>#12</u>	Discuss the generalisability (external validity) of the study results
37 38	Other		
39 40	Information		
41 42 43 44	Funding	<u>#13</u>	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based
45	None The STROBI	E checkli	st is distributed under the terms of the Creative Commons Attribution License CC-BY. This checklist can
 46 47 48 49 50 51 52 53 54 55 56 57 58 			s://www.goodreports.org/, a tool made by the <u>EQUATOR Network</u> in collaboration with <u>Penelope.ai</u>
59 60			For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml