

ORIGINAL RESEARCH ARTICLE

Prevalence, associated factors, and pharmacotherapy of hypertensive disorders among parturients in Bono Region of Ghana: An analytical cross-sectional study

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Abstract

The burden of hypertensive disorders in pregnancy (HDP) in remote areas of Ghana remains understudied. This analytical cross-sectional study investigated the prevalence, associated factors, and pharmacotherapy of HDPs in the Bono Region of Ghana. Data from labor registers of nine public hospitals, from January to December 2021, were analyzed. Guidelines for administering magnesium sulfate and antihypertensives were assessed. Associations were examined using chi-square and multivariable binomial regression with odds ratios at 95% confidence intervals. $P \leq 0.05$ was statistically significant. Of 16,206 deliveries, 711 parturients (4.4%) were complicated by HDPs. Non-severe pre-eclampsia (30.5%) and gestational hypertension (28.0%) were most frequent, while eclampsia (6.2%) and superimposed pre-eclampsia (1.7%) were less common. Maternal age 15–25 years (cOR = 2.43), unemployment (cOR = 2.14), primigravidity (cOR = 2.88), and primiparity (cOR = 2.39) were significantly associated with pre-eclampsia/eclampsia. After adjustment for confounding variables, primiparity remained borderline significant (aOR = 1.83; $p = 0.05$). Oral nifedipine and intravenous hydralazine were the primary antihypertensive therapies. Magnesium sulfate was universally administered using the Pritchard regimen, though product concentrations for intramuscular use varied slightly. Findings highlight the need to standardize magnesium sulfate formulations to optimize intramuscular dosing and enhance treatment consistency in lower-level facilities managing HDPs. (*Afr J Reprod Health* 2025; 29 [11]: 65-78).

Keywords: Bono region; hypertensive disorders in pregnancy; magnesium sulfate; preeclampsia/eclampsia; prevalence and pharmacotherapy

Résumé

La charge des troubles hypertensifs de la grossesse (THG) dans les zones reculées du Ghana demeure peu explorée. Cette étude transversale a évalué la prévalence, les facteurs associés et la pharmacothérapie des THG dans la région du Bono. Les registres d'accouchements de neuf hôpitaux publics, de janvier à décembre 2021, ont été analysés. Les directives concernant l'usage du sulfate de magnésium et des antihypertenseurs ont été examinées. Les associations ont été évaluées par chi-carré et régression binomiale multivariée, avec des rapports de cotes (IC 95 %). Une valeur de $p \leq 0,05$ était significative. Parmi 16 206 accouchements, 711 (4,4 %) présentaient des THG. La prééclampsie non sévère (30,5 %) et l'hypertension gestationnelle (28,0 %) étaient les plus fréquentes, tandis que l'éclampsie (6,2 %) et la prééclampsie surajoutée (1,7 %) restaient rares. L'âge de 15–25 ans (RC = 2,43), le chômage (RC = 2,14), la primigestion (RC = 2,88) et la primiparité (RC = 2,39) étaient associés à la prééclampsie/éclampsie. Après ajustement, la primiparité demeurait borderline significative (RCajusté = 1,83; $p = 0,05$). La nifédipine orale et l'hydralazine intraveineuse constituaient les principaux traitements, et le sulfate de magnésium était administré selon le schéma de Pritchard, avec de légères variations de concentration intramusculaire. Ces résultats soulignent la nécessité d'harmoniser les formulations afin d'optimiser le traitement et renforcer la cohérence des soins dans les structures périphériques. (*Afr J Reprod Health* 2024; 29 [11]: 65-78).

Mots-clés: La Région Bono; troubles hypertensifs de la grossesse; sulfate de magnésium; prééclampsie/éclampsie; prévalence et pharmacothérapie

Introduction

Hypertensive disorders in pregnancy (HDPs) continue to hinder the course of many pregnancies resulting in abrupt interventions, complications and mortalities with a potential of undermining target 3.1 of the Sustainable Development Goals (SDG).¹ The spectrum of HDPs, which varies from chronic, gestational, pre-eclampsia/eclampsia and superimposed pre-eclampsia, impact women's health worldwide and have had significant repercussions for low-income nations.^{2,3} These disorders have gained increased attention recently due to the rise in cases that have been documented and the unfavorable consequences they cause in certain nations.^{4,5} The incidence of HDPs has increased by 10.92% globally over the last three decades, having risen from 16.30 million in 1990 to 18.08 million in 2019.⁴

Given that almost a third of affected women are 'near miss', particularly in nations with limited resources, this rising trend is extremely concerning and unnecessarily causes panic among parturients and medical personnel.⁶ Hypertensive disorders of pregnancy together with hemorrhage, sepsis, and unsafe abortions are major contributors to the maternal death ratio in low-income countries surpassing the SDGs' estimates by more than six-fold.^{1,7,8} According to the World Health Organization (WHO) data, Ghana still has a high maternal mortality ratio (263/100,000), and the country's rising HDP burden adds to these deaths.^{9,10} Studies available on the subject almost a decade ago suggest that the incidence of HDPs in Ghana's Upper West and Greater Accra regions was less than 10%.¹¹ However, the prevalence was high when explored in 2019 and 2017 at the Komfo Anokye Teaching Hospital (KATH) in Kumasi, Ghana, and the Korle-Bu Teaching Hospital (KBTH) in Accra respectively.^{12,13} In Ghana, it is reported that over a third of women with HDPs experience 'near-miss' episodes; the ratio of these incidents to death is approximately 12:1.¹⁴ Despite the negative impact, it appears that the majority of research looking for HDPs in Ghana concentrate on

tertiary hospitals, with little information published from lower-tier facilities, particularly those located in the rural and smaller administrative regions. There is a need to close this information gap because it is argued that the unavailability of published statistics on HDPs in smaller regions significantly affects regional and global estimations which may mislead policies.¹⁵

Many maternal deaths, including those caused by HDPs in sub-Saharan Africa, are listed by the WHO as avoidable, implicating care plans offered to maternal hypertensives in the sub-region.⁸ The essential ideas that underpin the management of HDPs worldwide include early antenatal screening for case identification, antihypertensive therapy, the administration of anticonvulsants, and timely delivery of the baby. However, the way these concepts are applied may vary based on the resources, patient characteristics, and skill set that are available in a given geographical area. Inconsistencies were found in post-implementation studies involving the administration of magnesium sulfate (MgSO₄) to women who were eclamptic. Some nations lacked the necessary skills to safely administer the product, while others had difficulties with product accessibility. Furthermore, some institutions lacked treatment-guided protocols for MgSO₄ application, which hindered the safe use of the anticonvulsant in nations that have high rates of HDPs.^{16,17} Ghana's standard treatment guideline (STG) incorporates all of the fundamental HDP management principles; nevertheless, there aren't many post-implementation studies assessing these in healthcare institutions. In this study, we reviewed practice guidelines on the use of MgSO₄ and antihypertensives for HDPs as well as the prevalence and factors associated with these disorders among parturients in the Bono Region.

Methods

Study design and setting

An analytical cross-sectional study design was employed in which obstetric records, particularly

labor registers, and institutional treatment protocols for HDPs were monitored retrospectively for data in nine public hospitals of the region. The Sunyani Municipal, Sunyani SDA, Bono Regional, Berekum Holy Family, Dormaa Presbyterian, Drobo St. Mary's Catholic, Sampa Government, Tain District, and the Wenchi Methodist Hospitals were the purposely chosen research facilities. These hospitals were selected because they provided important obstetric services in eight of the region's twelve administrative districts and were the most established and resourced facilities. Health facilities in four districts were at the time of data collection not included due to their status as health centers which could not fully manage HDPs. The nine hospitals had maternal bed capacity of 384 and conducted about 1800 deliveries annually.

The Bono Region is located on Ghana's middle belt zone, sharing an international boundary to the west with the Republic of Cote d'Ivoire and to the north and south with Ghana's Savannah and Ashanti regions, respectively. With a population of roughly 1,208,649, the region occupies an area of 11,113 km².¹⁸ Its residents are primarily farmers and traders.

Study population and sample

All pregnant women who were registered and delivered at the public hospitals in Bono Region of Ghana between 3rd January 2021, and 31st December 2021, were considered. Overall, 16,206 deliveries were conducted during the period, of which 711 women presented with various HDPs. All 711 parturients diagnosed with HDPs during the study period were included in the study. Women with incomplete records during the review were excluded.

Diagnoses of HDPs according to the standard treatment guideline of Ghana

According to Ghana's STG, a pregnant woman has a hypertensive disorder when her blood pressure (BP) is $\geq 140/90$ mmHg on two or more occasions, spaced at least five minutes apart, using an

appropriate BP monitoring device.¹⁹ The diagnoses of gestational hypertension, chronic hypertension, pre-eclampsia with or without severe features, superposed pre-eclampsia and eclampsia may be made based on the timing of the onset of the hypertensive BP relative to the gestational age, time to conception, the patient's signs and symptoms, and any biochemical assessment suggesting an organ dysfunction. Gestational hypertension is new onset, occurring after mid-gestation without proteinuria or classical signs and symptoms; chronic hypertension is similarly symptomless, carried into pregnancy or identified in the first 20 weeks of gestation, unassociated with proteinuria, and persists after labor. Non-severe and severe forms exist in pre-eclampsia. Usually, it begins during mid-gestation with a fresh onset of hypertension (SBP ≥ 140 /DBP ≥ 90 mmHg) and mild proteinuria. It becomes severe if there are neurological signs and symptoms, massive proteinuria, elevated BP (SBP > 160 /DBP ≥ 110 mmHg), or biomarkers related to target organ injury. Women with chronic hypertension are diagnosed with superimposed pre-eclampsia if their features align with pre-eclampsia while eclampsia is diagnosed if a pre-eclamptic woman experiences seizures without other identifiable reasons.¹⁹ It is highlighted that screening for soluble fm-like tyrosine kinase-1 (sFlt-1) and placental growth factors (PlGF) is also accessible for diagnosing HDPs in advance settings but non-existent in the region at the time of data collection.

Pharmacotherapy of HDPs per the guidelines of Ghana

Under Ghanaian treatment policy, methyl dopa and sustained release/retarded nifedipine are prescribed for mild hypertensive cases in pregnancy (BP 140/90-159/109 mmHg). The recommended treatments for severe maternal hypertension are IV hydralazine, and labetalol. To abort seizures or prevent them, MgSO₄ by the Pritchard approach is recommended. This protocol administers the drug in two phases: an IV loading and IM maintenance

doses. During loading, 14g of the anticonvulsant is administered as 4g IV in a 20% MgSO₄ solution at first, with 2 doses of 5g IM in a 50% solution into each buttock (10g). Subsequently, a series of 5g IM doses of a 50% solution are injected into alternate buttocks every four hours to initiate the maintenance phase. This begins four hours after the loading dose has been initiated and continuously administered for 24 hours for preventive therapies or 24 hours after the last fit.¹⁹

Data collection

Data of participants was extracted from the labor registers of the hospitals using a designed template. Women's age, level of education, occupation, parity, gestational ages at labor, and the kind of HDP identified were among the information collected. Additionally, each hospital's institutional guideline for treating the spectrum of HDPs was reviewed, and with informed consent, the heads of the various labor wards filled a questionnaire on the use of anti-HDP drugs. The questionnaire was about antihypertensive and anticonvulsant medication availability and their usage for treating HDPs, particularly pre-eclampsia/eclampsia. It required the names of all medicines, their dosage forms, pharmaceutical strengths, and how they were administered. Importantly, requests were made for institutional MgSO₄ protocols and administration procedures including loading and maintenance doses, and their concentrations when given intravenously (IV) or intramuscularly (IM). Further, unit leaders answered questions on serum concentration monitoring of MgSO₄, adverse effects, and toxicity monitoring through both open- and closed-ended questions.

Data analysis

The data was analyzed using Stata 17.0 (Stata Corporation, Texas, USA). Categorical variables were compared using Chi-square (χ^2) or Fisher's exact tests, as appropriate. Binomial regression with a log-link function was used to estimate crude

and adjusted odds with 95% confidence intervals (CIs) for factors associated with hypertensive disorders in pregnancy (HDPs). Missing data was excluded, and results with p-values ≤ 0.05 were considered statistically significant.

Ethical approval

Permission to obtain data from the institutions in the Bono Region was granted by the Bono Regional Health Directorate, and ethical approval was sought and obtained from the Committee on Human research Publications and Ethics of KNUST (Ref: CHRPE/AP/119/20). Authors had no access to the identities of individual participants whose data were collected, and informed consent was obtained from labor ward in-charges/doctors where questionnaires required completion.

Results

Level of care of the health facilities involved in the study

Primary care level health services were provided by eight of the nine hospitals in rural and semi-urban communities of the region. The ninth establishment was a regional hospital that provided specialist services and doubled as the referral center for the Bono Region of Ghana.

Women's sociodemographic and obstetric data

Table 1 presents the sociodemographic data of parturients and their diagnoses. The age range was 15-46 years and the mean age was 31.0 (± 6.8). Those aged 26-34 years made up the largest proportion (43.9%). The majority were those who attained basic education (61.7%), working in informal occupations (68.0%), multigravida (51.9%), and multiparous women (53.0%). In all, a sum of 16,206 deliveries were conducted in the nine hospitals: 711 were diagnosed with HDPs.

Table 1: Sociodemographic data of women involved in the study

Variable	Frequency (n=711) n(%)
Age group (years)	
15-25	151 (21.2)
26-34	312 (43.9)
35-46	248 (34.9)
Educational level	
Basic	439 (61.7)
Secondary	124 (17.4)
Post-secondary/tertiary	148 (20.8)
Occupation	
Unemployed	89(12.5)
Informal employment	483(68.0)
Formal employment	139(19.5)
Gravida	
Primigravida	135 (19.0)
Multigravida	369 (51.9)
Grand multigravida	207 (29.1)
Parity	
Primiparous	248 (34.9)
Multiparous	377 (53.0)
Grand multiparous	86 (12.1)
Gestational age at delivery	
First trimester	8 (1.1)
Second trimester	18 (2.5)
Third trimester	685 (96.3)
Types of hypertensive disorders diagnosed	
Chronic hypertension	90 (12.7)
Gestational hypertension	199 (28.0)
Pre-eclampsia without severe features	217 (30.5)
Pre-eclampsia with severe features	149 (21.0)
Eclampsia	44 (6.2)
Chronic hypertension with superimposed pre-eclampsia	12 (1.7)

A prevalence of 4.4% was observed. The commonest HDP among the parturients was pre-eclampsia. This occurred without severe features in 30.5% of the subjects but in 21.0%, it was associated with severe features.

Gestational hypertension was found among 28.0%, whereas 6.2% developed eclampsia.

Maternal demographic data compared with diagnoses

Table 2 summarizes the distribution of HDPs across parturients’ demographic variables. Chronic hypertension was conspicuous among women aged 35-46 years (53.3%), basic education attainers (60.0%) and multiparous women (62.2%). Gestational hypertension was prevalent among women aged 26-34 years (45.2%), the least educated (65.8%) and multiparous as well (58.8%). Pre-eclamptic disorders (non-severe, severe, and superimposed) were seen mostly in middle-aged women (46.8%) between 26-34 years, those with low literacy (59.0%), multigravida (54.2%) and multiparous (52.1%). The majority of eclamptics (68.2%) were in the 15-25 age group, the least educated (70.5%), and primiparous (77.3%) who had delivered for the first time.

Factors associated with hypertensive disorders during pregnancy

The associations between parturients’ characteristics and hypertensive disorders are presented in Table 3. Age was significantly associated with hypertensive disorders during pregnancy (p<0.001). Younger women aged 15-25 years (25.8%), were observed more frequently with pre-eclampsia/eclampsia, whereas older women aged 35-46 years (41.5%), were more common in the chronic/gestational hypertension group. Maternal employment status also showed a significant association (p=0.002). A higher proportion of the unemployed (15.6%) were in the pre-eclampsia/eclampsia category, while those in employment (92.0%) dominated in the chronic/gestational hypertension group. Gravidity and parity were strongly associated with hypertensive disorders (p<0.001). Primigravida (23.7%) or primiparous women (41.5%), were more linked to pre-eclampsia/eclampsia.

Table 2: A comparison of hypertensive disorders with maternal sociodemographic characteristics

Variable	Chronic hypertension (n=90) n(%)	Gestational hypertension (n=199) n(%)	Pre-eclampsia (n=378) n(%)	Eclampsia (n=44) n(%)
Age				
15-25	5 (5.6)	37 (18.6)	79 (20.9)	30 (68.2)
26-34	37 (41.1)	90 (45.2)	177 (46.8)	8 (18.2)
35-46	48 (53.3)	72 (36.2)	122 (32.3)	6 (13.6)
Educational level				
Basic	54 (60.0)	131 (65.8)	223 (59.0)	31 (70.5)
Secondary	19 (21.1)	25 (12.6)	70 (18.5)	10 (22.7)
Post-secondary/tertiary	17 (18.9)	43 (21.6)	85 (22.5)	3 (6.8)
Occupation				
Unemployed	2 (2.2)	21 (10.6)	47 (12.4)	19 (43.2)
Employed	88 (97.8)	178 (89.4)	331 (87.6)	25 (56.8)
Gravida				
Primigravida	5 (5.6)	30 (15.1)	74 (19.6)	26 (59.1)
Multigravida	45 (50.0)	105 (52.8)	205 (54.2)	14 (31.8)
Grand multigravida	40 (44.4)	64 (32.2)	99 (26.2)	4 (9.1)
Parity				
Primiparous	16 (17.8)	57 (28.6)	141 (37.3)	34 (77.3)
Multiparous	56 (62.2)	117 (58.8)	197 (52.1)	7 (15.9)
Grand multiparous	18 (20.0)	25 (12.6)	40 (10.6)	3 (6.8)
Gestational age				
First trimester	7 (7.8)	1 (0.5)	0 (0.0)	0 (0.0)
Second trimester	7 (7.8)	2 (1.0)	7 (1.9)	2 (4.5)
Third trimester	76 (84.4)	196 (98.5)	371 (98.1)	42 (95.5)

Table 3: Parturients' factors and their association with hypertensive disorders (n=711)

Variable	Chronic/gestational hypertension	Pre-eclampsia/ eclampsia	p-value
Age			<0.001
15-25	42 (14.5)	109 (25.8)	
26-34	127 (43.9)	185 (43.8)	
35-46	120 (41.5)	128 (30.3)	
Educational level			0.411
Basic	185 (64.0)	254 (60.2)	
Secondary	44 (15.2)	80 (19.0)	
Post-secondary/tertiary	60 (20.8)	88 (20.9)	
Occupation			0.002
Unemployed	23 (8.0)	66 (15.6)	
Employed	266 (92.0)	356 (84.4)	
Gravida			<0.001
Primigravida	35 (12.1)	100 (23.7)	
Multigravida	150 (51.9)	219 (51.9)	
Grand multigravida	104 (36.0)	103 (24.4)	
Parity			<0.001
Primiparous	73 (25.3)	175 (41.5)	

Multiparous	173 (59.9)	204 (48.3)	
Grand multiparous	43 (14.9)	43 (10.2)	
Trimester of pregnancy			0.001*
First trimester	8 (2.8)	0 (0.0)	
Second trimester	9 (3.1)	9 (2.1)	
Third trimester	272 (94.1)	413 (97.9)	

Note: * (Fisher's exact)

Table 4: Regression analysis of the association between maternal sociodemographic data with pre-eclampsia/eclampsia

Variable	Pre-eclampsia /Eclampsia	cOR	95% CI	p-value	aOR	95% CI	p-value
Age				0<0.001			0.37
15-25	109 (72.2)	2.43	1.57-3.75		1.51	0.84-2.72	
26-34	185 (59.3)	1.36	0.97-1.91		1.14	0.78-1.67	
35-46	128 (51.6)	1			1		
Education				0.40			
Basic	254 (57.86)	1					
Secondary	80 (64.52)	1.32	0.88-2.00				
Post-sec/tertiary	88 (59.46)	1.07	0.73-1.56				
Occupation				0.001			
Unemployed	66 (74.16)	2.14	1.29-3.53				
Employed	356 (57.23)	1					
Gravida				0<0.001			
Primigravida	100 (74.07)	2.88	1.80-4.62				
Multigravida	219 (59.35)	1.47	1.04-2.07				
Grand multigravida	103 (49.76)	1					
Parity				0<0.001			0.05
Primiparous	175 (70.56)	2.39	1.44-3.96		1.83	0.97-3.46	
Multiparous	204 (54.11)	1.17	0.73-1.88		1.09	0.66-1.80	
Grand multiparous	43 (50.00)	1			1		
Trimester of pregnancy				0.38			
Second trimester	9 (50.00)	1					
Third trimester	413 (60.29)	0.65	0.25-1.68				

aOR, adjusted odds ratio; cOR, crude odds ratio; CI, confidence interval; p, probability

In contrast, grand multigravida women (36.0%) and multiparous women (59.9%) were more represented in the chronic/gestational hypertension group. However, the educational status of a parturient did not significantly impact the type of a hypertensive disorder (p = 0.411).

Parturients' odds of developing a hypertensive disorder

Table 4 examines the odds of a parturient developing pre-eclampsia/eclampsia based on sociodemographic and obstetric factors. Early

maternal age (15-25 years) was 2.43 times more related to pre-eclampsia/eclampsia (cOR = 2.43; 95% CI: 1.57-3.75 p = < 0.001) compared to older parturient age (35-46 years). Unemployed parturients were twice likely to have pre-eclampsia/eclampsia (cOR = 2.14; 95% CI=1.29-3.53; p = 0.001). Primigravidity elevated the odds to almost three times that of grand multi-gravida women (cOR =2.88; 95% CI: 1.80-4.62, p < 0.001). Primiparity also doubled the odds (cOR = 2.39, 95% CI: 1.44 -3.96; p = <0.001) and after adjusting for potential confounders, primiparity remained a

significant predictor though with a marginal significance level (aOR = 1.83; 95% CI: 0.97–3.46, p-value = 0.05). The educational level of a parturient did not significantly impact the odds of developing pre-eclampsia/eclampsia.

Availability and use of antihypertensives and magnesium sulfate for managing pregnancy-related hypertensive disorders at the hospitals

From Table 5, methyldopa and sustained-release nifedipine were the common oral antihypertensives available in all facilities for HDPs. Sustained-release nifedipine (30 mg) was the first-choice agent for HDPs, particularly pre-eclampsia, in seven hospitals. Intravenous hydralazine (20 mg/ml) was available and used in all hospitals for severe pre-eclampsia. Four of the nine hospitals also had IV labetalol (100 mg/20 ml) as an alternative. All nine hospitals stocked MgSO₄ injectables for managing pre-eclampsia/eclampsia. The 50% solution (10 ml ampoule) was available in eight facilities. Administration protocols, typically displayed as posters, were visible in all labor wards. The Pritchard regimen was standard across all facilities. Loading doses were 14g; injected as 4g IV followed by 10g IM. Maintenance doses were 5g IM, administered every four hours. The 50% solution was used for IM doses, while IV doses used a 20% solution. One hospital also had a protocol allowing the 20% solution for 5g IM doses. MgSO₄ prophylaxis was completed 24 hours after the loading dose and six IM maintenance doses, totaling 44g per patient. Renal function testing at therapy initiation was not consistently performed. None of the facilities could monitor serum magnesium levels. Therapy monitoring relied on clinical signs such as knee-jerk reflex, respiratory rate, and urine output. Seven hospitals had calcium gluconate injectables for managing MgSO₄ toxicity.

Discussion

This study highlights the prevalence and pharmacotherapy of hypertensive disorders in pregnancy (HDPs) among parturients in the Bono

Region of Ghana. The research was motivated by a lack of published data from this region, despite the national adoption of WHO-recommended treatment protocols. Few post-implementation evaluations exist, particularly regarding the administration of MgSO₄ for pre-eclampsia/eclampsia, making this study a timely contribution. The strengths of this study include its large delivery dataset drawn from nine peripheral hospitals which enhances the reliability of regional prevalence estimates beyond tertiary centers. Moreover, its analysis of drug protocols offers practical insights into real-world implementation of treatment guidelines.

Our results indicate that the prevalence of HDPs in the region was 4.4%, a rate less than those reported in Zambia (6.7%), Ethiopia (6.82%), sub-Saharan Africa (8%), and Nigeria (25.8%).^{20–23} However, it was higher than Burkina Faso (1.4%) and Ghana's Upper West Region (3.2%).^{11,24} The observed regional variations could be due to differences in study methodologies, environmental and socioeconomic factors. Unlike studies that collected data from antenatal clinics (ANCs) attendees, our study focused on data from women in labor, potentially influencing prevalence estimates. Hospital deliveries in most Ghanaian facilities usually rank lower than ANC attendance which may contribute to differences in findings.^{25,26} This may be due to factors such as transportation challenges, financial constraints, fear of cesarean section, and some negative perceptions of hospital staff attitudes which could contribute to home deliveries in rural settings, leading to an underestimation of HDP prevalence in hospitals.^{25,26} However, in Burkina Faso and the Upper West Region of Ghana, where the occurrences were lower than our data, it seems that the effects of attendance and social characteristics were more prominent.^{11,24} Pre-eclampsia was the most prevalent HDP in this study, consistent with findings of Dassah *et al.* in Kumasi, Ghana.¹³ Among affected women, 30.5% had non-severe pre-eclampsia while one-fifth had pre-eclampsia with severe features.

Table 5: Availability and utilization of antihypertensives and magnesium sulfate

Medicines	No of hospitals that use the antihypertensive (n=9)	Used as first choice agent (n=9)
Oral antihypertensives		
Methyldopa 250mg	9	2
Nifedipine retard 20mg	5	0
Extended release nifedipine 30mg	9	7
Oral hydralazine 25mg	2	0
Parenteral antihypertensives		
IV hydralazine (20mg/ml)	9	9
IV labetalol(100mg/20ml)	4	0
Magnesium sulfate		
Presence of MgSO₄ injectable at hospital		
Yes	9	
Type of stocks present at the time		
50% MgSO ₄ (10ml ampoule)	8	
20% MgSO ₄ (10ml ampoule)	1	
Treatment protocol pasted at labor ward		
Yes	9	
Type of treatment regimen		
Loading dose + maintenance doses	9	
Composition of loading dose		
4g IV + 10g IM	9	
Concentration of MgSO₄ by IV route		
20%	9	
50%	0	
Maintenance doses		
5g IM 4 hourly	9	
Concentration of MgSO₄ by IM route		
50%	8	
20%	1	
Injection site for IM doses		
Buttocks	9	
Completion of treatment regimen		
After 6 IM maintenance doses injected 4 hourly	9	
Maximum dose of MgSO₄ administered to a patient		
44g	9	
Renal function test results seen before starting MgSO₄		
Always	0	
Sometimes	9	
Patient monitoring		
Solely based on patient signs and symptoms	9	
Serum concentrations of Mg ²⁺	0	
Availability of calcium gluconate to manage toxicity		
Yes	7	
Expired	2	

These rates exceed the prevalence (4.1%) estimated in a systematic review of sub-Saharan Africa indicating that our findings may reflect regional differences in healthcare access and resources.²² Previous researchers showed that HDPs were predicted by maternal age.^{27,28} Our multivariable regression analysis revealed a significant association between age and HDP. Younger parturients (15-25 years) were more prone, particularly to pre-eclampsia/eclampsia. Decreasing maternal age increased the odds, with over two-thirds of eclamptic cases occurring in this age group (see Table 2). This trend is supported by previous studies in Ghana and South Africa.^{29,30} The pathophysiological basis for this association remains unclear but may involve mothers' aberrant immune responses to paternally inherited fetal antigens during first pregnancies as hypothesized in earlier literature which could apply to younger women getting pregnant for the first time.³¹ Additionally, reduced ANC attendance among teenage mothers due to social stigma may lead to poor pregnancy monitoring, anemia, and inadequate preventive care, further increasing their risk.²⁸ Contrarily, our advanced-aged parturients (≥ 35 years) were more aligned with chronic hypertension which was consistent with existing literature.^{20,32,33} This is conceivable, given that obesity, vascular calcification, stiffness, and loss of distensibility correlate with old age and greatly impact hypertension.³⁴

The results of this study relate to previous authors who reported that parity plays a role in pre-eclampsia/eclampsia.^{28,35} While our crude odds ratio indicated a significant association between primigravidity and HDPs ($p < 0.001$), the adjusted odds showed significance for primiparity ($p = 0.05$). Known in literature, multigravidity has been associated with a more regulated immune response and adaptation to repeated exposure to paternally inherited antigens as opposed to primigravid/primiparous women, who are first-time encounters, less adapted and therefore prone to inflammatory response leading to pre-eclampsia.³¹ Whereas parity was a significant predictor,

educational level did not show an association. Unlike some prior studies, our findings did not establish a significant relationship between HDPs and illiteracy after adjusting for other variables.^{27,28} The lack of variability in educational attainment among study participants may have limited the statistical power to detect such an association.

Maternal hypertensives were primarily treated with the 30mg extended release nifedipine, 250mg methyldopa, and IV hydralazine (20 mg/ml). These agents were consistently available across all the nine hospitals. Among the available oral agents, nifedipine was the most commonly prescribed first-line treatment in seven out of nine hospitals, mirroring prescribing patterns observed at Tamale Teaching Hospital in Ghana.³⁶ Its preference over methyldopa may be attributed to its faster onset of action (30-45 minutes vs. 1-1.5 hours for methyldopa).³⁷⁻³⁹ Oral labetalol was rarely stocked, so its use for treatment was limited, likely due to cost and availability. Nifedipine's use was in line with Ghana's treatment policy, but some evidence suggests that nifedipine may increase the risk of disease progression to pre-eclampsia when used in non-severe maternal hypertension, prompting WHO to advocate for alternative oral agent such as labetalol in non-severe hypertensive cases.⁴⁰

Some recently revised guidelines and review publications appear to place IV hydralazine as second line agent for severe pre-eclampsia due to its perceived unfavorable kinetics and side effects, but its utilization as first choice agent was evident in this study.⁴¹ The widespread usage was also observed in Nigeria.⁴² The high utilization of IV hydralazine, despite recent guidelines favoring IV labetalol may be driven by staff familiarity, affordability, and availability.⁴³

All study sites adhered to national and international guidelines recommending MgSO₄ for managing severe pre-eclampsia/eclampsia.^{19,44} All hospitals had access to the product, along with its administration protocols. The Pritchard regimen was uniformly followed, likely due to its convenience and independence from infusion

pumps, which are often scarce in resource-limited settings.¹⁹ Magnesium sulfate is administered to severe pre-eclamptic women not because they have Mg²⁺ deficiency, but rather to raise Mg²⁺ plasma concentrations to counteract Ca²⁺-induced muscle twitches, vasospasms, and glutamate-mediated neuronal excitement in eclamptic and severely pre-eclamptic women.⁴⁵⁻⁴⁷ Moreover, the goal is to hinder cholinergic transmission at the neuromuscular junction to stop or avoid convulsive episodes.^{46,47}

Under the Pritchard method, patients receive an initial loading dose of 14g MgSO₄. This is implemented as 4g slow IV infusion using a 20% solution, which provides immediate anticonvulsant activity lasting about 30 minutes. This is followed by a 10g intramuscular (IM) injection using a 50% solution, extending the anticonvulsant effect for 3-4 hours. To maintain therapeutic levels, a 5g IM dose is injected into alternating gluteal muscles every 4 hours for up to 24 hours.^{46,47} Eight hospitals stocked the 50% w/v MgSO₄ in 10 ml ampoules in compliance with WHO requirements.⁴⁸ However, one hospital only had 20% w/v MgSO₄, presenting a significant clinical challenge. A 5g IM dose of 20% MgSO₄ requires 25 ml, exceeding the regular injection volume of 10 ml for gluteal muscles from a 50% solution.^{19,49} Some hospital pharmacists might have underestimated the health implications of substituting a 50% w/v MgSO₄ solution with a 20% for IM doses. Such an excess volume may cause pain, drug leakage, inflammation, and abscess formation.^{50,51} This issue, as highlighted by Babu *et al.* may contribute to poor patient compliance with IM maintenance therapy.⁵²

At the sites, severe pre-eclamptic/eclamptics received six IM maintenance doses based on the MgSO₄ protocols evaluated. These amounted to cumulative doses of 44g MgSO₄ for each patient in 24 hours as opposed to 40g reported in some literature.⁵³ Despite high adherence to the MgSO₄ guidelines, none of the hospitals had the capacity to monitor serum Mg²⁺ levels, relying instead on urine output, respiratory

rate, and patellar reflexes, a practice that may be insufficient for preventing toxicity. Given the risk of accumulation in patients with impaired renal function, dose adjustments based on weight or alternative regimens like the Dhaka regimen or a 12-hour protocol proposed by Beyuo *et al.* may be safer options for facilities without laboratory capacity.⁵⁴⁻⁵⁶

The study has implications for policy and practice in Ghana. Standardizing MgSO₄ formulations nationwide would enhance dosing safety, particularly in lower-level facilities. Increasing access to labetalol could reduce overreliance on nifedipine and hydralazine in practice, while continuous professional training is needed to strengthen monitoring of HDP management in settings without serum magnesium testing. Furthermore, targeted maternal health programs for younger, unemployed, and primigravid women should be intensified to reduce their elevated risk of pre-eclampsia/eclampsia. Collectively, these measures would support Ghana's efforts toward achieving SDG 3.1 by reducing preventable maternal deaths linked to HDPs.

The study had some drawbacks. For example, part of the data was collected by observing drug protocols, drug shelves, and medicine trays. In addition, procedural data were gathered from physicians, labor/maternity unit heads, and pharmacists rather than by watching actual procedures being performed which might differ. Moreover, inaccurate records might have been captured in labor registers, which could impact the quality of the data collected. Again, the study did not include health centers, private clinics, or maternity homes since it was assumed that many HDP patients would be directed to major hospitals because such smaller facilities were unable to handle maternal complications or conduct cesarean surgeries when urgent deliveries were required. If some HDPs were managed without referral, this would have an effect on the statistics that this study projects.

Conclusion

The prevalence of HDP in the Bono Region of Ghana was 4.4%. Pre-eclampsia was the most common HDP. Young parturient age, unemployment, primigravida and primiparity were the predictors of pre-eclampsia/eclampsia. The extended-release oral nifedipine and IV hydralazine were the main therapies for HDPs, especially for severe pre-eclamptic patients. Magnesium sulfate protocols for pre-eclampsia/eclampsia conformed to standard regimen. The supply of 20% MgSO₄ solution for IM maintenance doses was inappropriate due to its larger injection volumes that may be harmful to patients. Standardizing the concentration of magnesium sulfate solutions for pre-eclampsia/eclampsia could optimize intramuscular dosing under the Pritchard regimen, improving treatment consistency.

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Competing interest

No conflict of interest is declared by authors

Contribution of authors

FF, ETD and KOB conceived and designed the study; FF, BO and KAD collected the data; FF, JBY and ETD analyzed the data; FF, BO, JBY and ETD wrote the original draft; KAD, ETD, and KOB reviewed and edited the draft; ETD and KOB provided mentorship and supervision. All authors approved the manuscript

References

1. WHO. Acceleration towards the sustainable development goal targets for maternal health and child mortality. Report by the Director-General, 2023. Available from: https://apps.who.int/gb/ebwha/pdf_files/EB154/B154_12-en.pdf
2. Bajpai D, Popa C, Verma P, Dumanski S and Shah, S. Evaluation and management of hypertensive disorders of pregnancy. *Kidney* 2023; 4(10): 1512–1525.
3. Say L, Chou D, Gemmill A, Tunçalp O, Moller AB, Daniels J, Gülmezoglu AM, Temmerman M and Alkema L. Global causes of maternal death: A WHO systematic analysis. *Lancet Glob Heal* 2014; 2(6): e323–e333.
4. Wang W, Xie X, Yuan T, Wang Y, Zhao F, Zhou Z and Zhang H. Epidemiological trends of maternal hypertensive disorders of pregnancy at the global, regional, and national levels: A population-based study. *BMC Pregnancy Childbirth* 2021; 21(1): 364.
5. Noubiap JJ, Bigna JJ, Nyaga UF, Jingi AM, Kaze AD, Nansseu JR and Fokom Domgue J. The burden of hypertensive disorders of pregnancy in Africa: A systematic review and meta-analysis. *J Clin Hypertens (Greenwich)* 2019; 21(4): 479–488.
6. Naderi T, Foroodnia S, Omidi S, Samadani F and Nakhaee N. Incidence and correlates of maternal near miss in southeast Iran. *Int J Reprod Med* 2015; 2015: 914713.
7. Lotufo FA, Parpinelli MA, Haddad SM, Surita FG and Cecatti, J. Applying the new concept of maternal near-miss in an intensive care unit. *Clin. (Sao Paulo)* 2012; 67(3): 225–230.
8. WHO. Maternal mortality. 2024 Apr 26. Available from: <https://www.who.int/news-room/fact-sheets/detail/maternal-mortality>.
9. Maternal mortality ratio (modeled estimate, per 100,000 live births) -United States. WHO, UNICEF, UNFPA, World Bank Group, and UNDESA/Population Division. *Trends in Maternal Mortality 2000 to 2020*. Geneva, 2023. Available from: <https://data.worldbank.org/indicator/SH.STA.MMR.T?locations=US>
10. Adu-Bonsaffoh K, Oppong SA, Binlinla G and Obed SA. Maternal deaths attributable to hypertensive disorders in a tertiary hospital in Ghana. *Int J Gynaecol Obs* 2013; 123(2): 110–113.
11. Antwi E, Klipstein-Grobusch K, Quansah Asare G, Koram KA, Grobbee D and Agyepong IA. Measuring regional and district variations in the incidence of pregnancy-induced hypertension in Ghana: challenges, opportunities and implications for maternal and newborn health policy and

- programmes. *Trop Med Int Heal* 2016; 21(1): 93100.
12. Adu-Bonsaffoh K, Ntummy MY, Obed SA and Seffah JD. Prevalence of hypertensive disorders in pregnancy at Korle-Bu Teaching Hospital in Ghana. *J Gynecol Neonatal Biol* 2017; 3(1): 8–13.
 13. Dassah ET, Kusi-Mensah E, Morhe ESK and Odoi AT. Maternal and perinatal outcomes among women with hypertensive disorders in pregnancy in Kumasi, Ghana. *PLoS One* 2019; 14(10): e0223478.
 14. Drechsel KCE, Adu-Bonsaffoh K, Loohuis KMO, Srofenyoh EK, Boateng D and Browne JL. Severe preeclampsia adverse outcome triage (SPOT) studies consortium. Maternal near-miss and mortality associated with hypertensive disorders of pregnancy remote from term: a multicenter observational study in Ghana. *AJOG Glob Rep* 2022; 2(2): 100045.
 15. Abalos E, Cuesta C, Grosso AL, Chou D and Say L. Global and regional estimates of preeclampsia and eclampsia: A systematic review. *Eur J Obs. Gynecol Reprod Biol* 2013; 170(1): 1–7.
 16. Long Q, Oladapo OT, Leathersich S, Vogel JP, Carroli G, Lumbiganon P, Qureshi Z and Gülmezoglu AM. WHO multicountry survey on maternal and newborn health research network. Clinical practice patterns on the use of magnesium sulfate for treatment of pre-eclampsia and eclampsia: a multi-country survey. *BJOG* 2017; 124(12): 1883–1890.
 17. Eddy KE, Vogel JP, Zahroh RI and Bohren MA. Factors affecting use of magnesium sulphate for pre-eclampsia or eclampsia: A qualitative evidence synthesis. *BJOG* 2022; 129: 379–391.
 18. GSS. Ghana statistical service. Ghana 2021 population and housing census general report volume 3A. Populations of regions and districts 2021. Available from: https://statsghana.gov.gh/gssmain/fileUpload/pressrelease/2021_PHC_General_Report_Vol_3A_Population_of_Regions_and_Districts_181121.pdf
 19. Republic of Ghana: Ministry of health, national drugs program, standard treatment guidelines. 7 ed. Ghana Yamens Press Ltd, 2017.
 20. Chowa PE, Lin C, Goma F and South-Paul J. Hypertension among women of child bearing age in Zambia. *Med J Zambia* 2011; 38(3): 2011
 21. Tesfa E, Nibret E, Gizaw ST, Zenebe Y, Mekonnen Z, Assefa S, Melese M, Fentahun N and Munshea A. Prevalence and determinants of hypertensive disorders of pregnancy in Ethiopia: A systematic review and meta-analysis. *PLoS One* 2020; 15(9): e0239048.
 22. Gemechu KS, Assefa N and Mengistie B. Prevalence of hypertensive disorders of pregnancy and pregnancy outcomes in Sub-Saharan Africa: A systematic review and meta-analysis. *Womens Health (Lond)* 2020; 16: 1745506520973105.
 23. Azubuike A and Danjuma I. Hypertension in pregnancy among rural women in Katsina State, Nigeria. *J. basic clin. reprod. sci* 2017; 6(1): 140–146.
 24. Garanet F, Samadoulougou S, Baguiya A, Bonnechère B, Millogo T, Degryse JM, Kirakoya-Samadoulougou F and Kouanda S. Low prevalence of high blood pressure in pregnant women in Burkina Faso: A cross-sectional study. *BMC Pregnancy Childbirth* 2022; 22(1): 955.
 25. Adongo PA, Atanga RA and Yakong VN. Demographic characteristics of women that use traditional birth attendants in Bongo District, Ghana. *Eur J Midwifery* 2020; 4(1).
 26. Adatara P, Strumpher J and Ricks E. Exploring the reasons why women prefer to give birth at home in rural northern Ghana: A qualitative study. *BMC Pregnancy Childbirth* 2020; 20(1): 500.
 27. Chemedda WC, Gurmessa TS, Gedefa AG and Woldasemayat LA. Factors associated with hypertensive disorders among pregnant mothers attending antenatal care services at public health facilities in Gambella Town, Southwest Ethiopia: A cross-sectional study. *Int J Afr Nurs Sc* 2022; 17: 100478.
 28. Meazaw MW, Chojenta C, Muluneh MD and Loxton D. Factors associated with hypertensive disorders of pregnancy in sub-Saharan Africa: A systematic and meta-analysis. *PLoS One* 2020; 15(8): e0237476.
 29. Beyuo TK, Lawrence ER, Kobemik EK and Oppong SA. Clinical presentation and predictors of eclampsia among women with hypertensive disorders of pregnancy in Ghana. *Pregnancy Hypertens* 2022; 30: 171–176.
 30. Moodley J and Ngene N. Maternal deaths due to eclampsia in teenagers: Lessons from assessment of maternal deaths in South Africa. *Afr J Prim Heal. Care Fam Med* 2020; 12(1): a2305.
 31. Dimitriadis E, Rolnik DL, Zhou W, Estrada-Gutierrez G, Koga K, Francisco RPV, Whitehead C, Hyett J, da Silva Costa F, Nicolaides K and Menkhorst E. Preeclampsia. *Nat Rev Dis Primers* 2023; 16; 9(1): 8.
 32. Walle TA and Azagew A. Hypertensive disorder of pregnancy prevalence and associated factors among pregnant women attending ante natal care at Gondar town health institutions, North West Ethiopia. *Pregnancy Hypertens* 2019; 16: 79–84.
 33. Lopian M, Kashani-Ligumsky L and Many A. A balancing act: Navigating hypertensive disorders of pregnancy at very advanced maternal age, from preconception to postpartum. *J Clin Med* 2023; 12(14): 4701.
 34. Harvey A, Montezano AC and Touyz R. Vascular biology of ageing-Implications in hypertension. *J Mol Cell Cardiol* 2015; 83: 112–121.
 35. Luo ZC, An N, Xu HR, Larante A, Audibert F and Fraser

- W. The effects and mechanisms of primiparity on the risk of pre-eclampsia: a systematic review. *Paediatr Perinat Epidemiol* 2007; Suppl 1: 36–45.
36. Bugri AA, Gumanga SK, Yamoah P, Frimpong EK and Nlooto M. Prevalence of hypertensive disorders, antihypertensive therapy and pregnancy outcomes among pregnant women: A retrospective review of cases at Tamale Teaching Hospital, Ghana. *Int J Env. Res Public* 2023; 20(12): 6153.
 37. Awaludin A, Rahayu C, Daud NAA and Zakiyah N. Antihypertensive medications for severe hypertension in pregnancy: A systematic review and meta-analysis. *Healthcare* 2022; 10(2): 325.
 38. Smith P, Anthony J and Johanson R. Nifedipine in pregnancy. *BJOG* 2000; 107(3): 299–307.
 39. Cohan JA and Checcio L. Nifedipine in the management of hypertensive emergencies: report of two cases and review of the literature. *Am J Emerg Med* 1985; 3(6): 524–530.
 40. WHO. Recommendations on drug treatment for non-severe hypertension in pregnancy. Geneva: World Health Organization, Licence: CC BY-NC-SA 3.0 IGO, 2020. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK561249/table/ch1.tab1/%0A>
 41. Fordjour F, Dassah ET, Mensah KB, Appiah-Kubi A, Amoateng P, Buabeng KO and Darko KA. Antihypertensives therapy and monitoring in pre-eclampsia: Role of the pharmacist. *J Popul Ther Clin Pharmac* 2023; 30(17):354–367.
 42. Oriji PC, Allagoa DO, Ubom AE, Kattay AK, Briggs DC, Chika MN, Iloeje UN, Akanatei ID and Atemie G. Hypertensive disorders in pregnancy at the Federal Medical Centre, Yenagoa, South-South Nigeria: A 5-year review. *Int J Res Med Sci* 2021; 9(10): 2923–2929.
 43. Hricik DE, Smith MC and Wright, J. *Hypertension Secrets*. (Hanley & Belfus, Philadelphia), 2002.
 44. WHO. Recommendations for prevention and treatment of preeclampsia and eclampsia implications and actions, 2014. Available from: https://iris.who.int/bitstream/handle/10665/119627/WHO_RHR_14.17_eng.pdf
 45. Euser AG and Cipolla M. Magnesium sulfate for the treatment of eclampsia: A brief review. *Stroke* 2009; 40(4): 1169–1175.
 46. APA. *Lexicom Drug Information Handbook*. (Wolters Kluwer: USA), 2016
 47. AHFS. American Hospital Formulary Service (AHFS) Drug Information. American Society of Health-System Pharmacists, USA, 2003
 48. USAID. Manual for procurement and supply of quality-assured MNCH commodities: Magnesium sulfate injection, 500 mg/ml in 2-ml and 10-ml ampule 2019. Available from [https://www.ghsupplychain.org/sites/default/files/2022-11/MNCH Commodities Procurement-Magnesium Sulfate.pdf](https://www.ghsupplychain.org/sites/default/files/2022-11/MNCH%20Commodities%20Procurement-Magnesium%20Sulfate.pdf)
 49. Magnesium sulfate injection solution: HF acquisition Co LLC DBA healthfirst. 2017. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=945419b8-6184-f4f3-e053-2a95a90a6236&type=display>.
 50. Roldán-Chicano MT, Rodríguez-Tello J, Cebrián-López R, Moore JR and Del Mar García-López M. Adverse effects of dorsogluteal intramuscular injection versus ventrogluteal intramuscular injection: A systematic review and meta-analysis. *Nurs Open* 2023; 10(9): 597.
 51. Allen LV and Ansel H. *Ansel's Pharmaceutical Dosage Forms and Drug Delivery System*. (Wolters Kluwer Health, Philadelphia), 2013.
 52. Babu JJ, Venkadalakshmi V, Dhandapani M and Chopra S. Pain, drug leakage and inflammation associated with intramuscular injections of magnesium sulfate in preeclamptic women: A descriptive study. *Nurs Midwifery Res. J* 2022; 18(1): 35–44.
 53. Vallerand AH, Sanoski CA and Quiring C. Magnesium Sulfate (Parenteral). In *Davis's Drug Guide*. 18th Ed. F.A. Davis Company, 2023
 54. PubChem. Bethesda (MD): National library of Medicine (US), national center for biotechnology information. PubChem compound summary for CID 24083, magnesium sulfate, 2004. Available from: <https://pubchem.ncbi.nlm.nih.gov/compound/Magnesium-Sulfate>.
 55. Beum R, Begum A, Johanson R, Ali MN and Akhter S. A low dose (“Dhaka”) magnesium sulphate regime for eclampsia. *Acta Obs. Gynecol Scand* 2001; 80(11): 998–1002.
 56. Beyuo K, Lawrence ER, Kobernik EK and Oppong SA. A novel 12-hour versus 24-hour magnesium sulfate regimen in the management of eclampsia and preeclampsia in Ghana (MOPEP Study): A randomized controlled trial. *Int J Gynaecol Obs* 2022; 159(2): 495–504.