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# Factors Associated with Survival to Discharge of Newborns in a Middle-Income Country

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#### Abstract

Clinical and mortality audit is an essential part of quality improvement in health care; information obtained in this process is used to develop targeted interventions to improve outcome. This study aimed to determine predictors of short-term survival in neonates. An existing neonatal database was reviewed. A total of 5018 neonates > 400 g admitted to a tertiary hospital (Johannesburg South Africa) between 1 January 2013 and 31 December 2015 were analysed. Mean birth weight was 2148 g (standard deviation [SD]: 972) and mean gestational age was 34.2 weeks (SD: 4.8). Overall survival was 85.6% (4294/5018). The most common causes of death were prematurity (46.2%), hypoxia (19.5%) and infection (17.2). The strongest predictors of survival were birth weight (OR 1.0; 95% confidence intervals (CI): 1.0–1.01) and gestational age (OR = 1.1, 95% CI: 1.05–1.17). Other predictors of survival included metabolic acidosis (OR = 0.14, 95% CI: 0.09–0.20), hyperglycemia (OR = 0.31, 95% CI: 0.23–0.41), mechanical ventilation (OR = 0.35, 95% CI: 0.28–0.46), major birth defect (OR = 0.12, 95% CI: 0.08–0.18), resuscitation at birth (OR = 0.39, 95% CI: 0.31–0.49) and Caesarean section (OR = 1.8, 95% CI: 1.44–2.25). In conclusion, resources need to be focused on improved care of VLBW infants.

Keywords: neonatal mortality, clinical audit, very low birth weight, premature infants

#### 1. Introduction

The fourth Millennium Development Goal (MDG) was a two-third reduction in the mortality of children under the age of 5 years, which sub-Sahara African countries (including South Africa) failed to achieve this [1]. In 2015, 1 million children died within the first day of life, a further million in the first week of life and yet another 2.8 million in the first 28 days of life –



© 2016 The Author(s). Licensee InTech. This chapter is distributed under the terms of the Creative Commons Attribution License (http://creativecommons.org/licenses/by/3.0), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited. 4.8 million of the almost 6 million children under the age of five years who died in 2015, died within the neonatal period [1]. Concentrating resources on newborns is therefore essential to further reduce childhood mortality.

The causes of neonatal mortality vary considerably among different units and different countries. The United Nations MDG 2015 report [1] states that "better data are needed for the post-2015 development agenda" and "real-time data are needed" to guide policy makers. Most data have a time lag of between 2 and 3 years before the policies are implemented. The MDGs formed the foundation of the so-called Sustainable Development Goals (SDG) [2]. The SDGs are less specific than the MDGs, but include health targets, one of which is to reduce both neonatal mortality and mortality of children under the age of 5 years.

Regular audits of neonatal mortality are required to identify the causes of death so that proper interventions can be implemented to reduce neonatal deaths. It is essential to have local data to address local health issues; transposing mortality data from another country will not necessarily solve local problems. This is particularly true when using data from a high-income country to address problems experienced in low- to middle-income countries (LMICS). A recent review of the mortality rates in neonatal intensive care units showed that the rate varied considerably between different countries [3]; the mortality rate was generally high, but greater in developing than developed countries. Issues such as the lack of antenatal care and inadequate health facilities are the causes of neonatal mortality rate -35% of admitted neonates died. The important causes of neonatal death included lack of antenatal care, birth weight below 1500 g, hypothermia at birth, and delivery outside a teaching hospital.

Previous studies done in very low birth weight (VLBW) neonates – birth weight below 1500 g – at Charlotte Maxeke Johannesburg Academic Hospital (CMJAH) have shown that birth weight was the most significant predictor of survival [5, 6]. Resuscitation at birth, the use of nasal continuous positive airways pressure (NCPAP) and the mode of delivery were also important factors affecting survival. Survival of extremely low birth weight (ELBW) neonates was particularly low at CMJAH [7]. The provision of NCPAP to this category of neonates more than doubled their survival to discharge [6].

#### 2. Determinants of neonatal survival at a tertiary hospital in Johannesburg, South Africa

Although VLBW mortality at CMJAH has been studied, the overall neonatal survival has not been audited. The aim of this study is to review neonatal survival at CMJAH and to determine important modifiable factors to inform protocols and budgeting for neonatal care. The objectives of this study were to:

• Describe the patient population with regard to demographic information, clinical characteristics, and outcome at discharge.

- Determine the survival rate for different birth weight categories.
- Establish factors associated with neonatal survival.

#### 2.1. Subjects and methods

The study was conducted in the neonatal unit of a tertiary academic hospital (CMJAH) in Johannesburg, South Africa. All neonates admitted within 48 h of birth, between 1 January 2013 and 31 December 2015, were included in the study. Neonates with a birth weight below 400 g and those with important missing data, particularly birth weight, gestational age, and outcome at discharge were excluded.

#### 2.2. Study design

This was a secondary analysis of an existing neonatal database. Data were collected upon discharge for each neonate admitted to the CMJAH neonatal unit and entered on to a database. The database was managed using Research Electronic Data Capture (REDCAP) [8] hosted by the University of the Witwatersrand. The information collected included demographic details, maternal information, delivery room data, clinical information, and outcome at discharge. Data from VLBW neonates was contributed to the Vermont Oxford Network (VON) (www.vtoxford.org), a multinational neonatal collaboration. A paper computer summary form was completed for each patient, using the patient file. Data were checked against the patient file and then entered on to the database. The information on the database was then checked against the paper form. Any discrepancies noted were verified against the patient files. Definitions and codes for congenital defects or surgical procedures were obtained from the VON.

Neonates were classified by weight using standard definitions — term large for gestational age (TLGA) neonates weighed above 4000 g at birth, term appropriate for gestational age (TAGA) infants weighed between 2500 and 3999 g at birth, low birth weight (LBW) neonates had a birth weight less than 2500 g, very low birth weight (VLBW) included those weighing less than 1500 g at birth and extremely low birth weight (ELBW) less than 1000 g at birth. Term was considered to be a gestation age between 37 and 42 weeks, preterm below 37 weeks, and post-term to be above 42 weeks.

The unit participated in a national perinatal mortality audit – the perinatal problem identification programme (PPIP)(www.ppip.co.za). The broad causes of neonatal death were categorized using standard PPIP definitions.

#### 2.2.1. Neonatal unit

The neonatal unit was situated in large tertiary academic hospital in a metropolitan setting. Neonatal facilities included a transitional nursery in labour ward, a shared paediatric/neonatal intensive care unit (PNICU) with 15 ventilator beds, a neonatal high care unit with 40 beds, low-care facility with 25 beds, and nine kangaroo mother-care (KMC) beds. Nasal continuous positive airways pressure (NCPAP) and therapeutic hypothermia for perinatal asphyxia were

provided in high care. The neonatal unit was staffed by neonatologists, registrars, and house staff. There were various paediatric sub-specialities in the hospital including nephrology, neurology, cardiology, endocrinology, and infectious diseases. There was a large paediatric surgery service and paediatric surgical neonates were admitted to the neonatal unit and jointly managed with the neonatal staff.

Neonates who were observed in the transitional unit and then discharged to their mothers were not included in the study. Neonates who died in the delivery room and transitional nursery were considered to be admissions and were included in the study. Owing to resource constraints, there were insufficient ventilator beds for the number of neonates requiring ventilation. The PNICU functioned essentially as a ventilator unit; high-care observation was not possible due to limited facilities. The neonatal unit had a policy of rationing care based on birth weight—babies weighing below 750 g at birth would not be offered surfactant or NCPAP, but only given supplemental oxygen, intravenous fluids, and antibiotics; babies weighing between 750 and 900 g would be given surfactant and NCPAP, but would not be provided with mechanical ventilation if required. All neonates with respiratory distress syndrome were initially managed with NCPAP and early rescue surfactant; those who failed would be transferred to the PNICU for mechanical ventilation. The use of NCPAP at CMJAH has recently been reviewed [9].

#### 2.2.2. Statistical analysis

Data were exported to IBM SPSS version 22 for the purpose of analysis. The standard statistical methods were used to describe the data — continuous variables were described using measures of central tendency and dispersion, mean and standard deviation (SD), or median and interquartile range (IQR) as appropriate. Categorical variables were described using frequency and percentages.

The primary endpoint was whether a neonate survived to hospital discharge. Univariate analysis was done considering different maternal, demographic, and clinical variables as independent factors of survival. Differences in outcome for continuous variables were compared using unpaired *t*-tests or Mann Whitney *U*-test as appropriate. Associations of outcome with categorical variables were investigated using Chi-squared test. A factor with a *p*-value of 0.05 was considered statistically significant. Variables with a *p*-value <0.1 on the univariate analysis were entered into a multiple logistic regression model considering whether a child survived to discharge as the outcome variable. Factors associated with neonatal mortality were determined separately for VLBW and bigger babies.

The possible sources of bias were identified and excluded from the analysis. Conditions which were only present in neonates who were survivors and approaching discharge were identified and excluded from the analysis of deaths. These conditions included supplementary oxygen at 28 days, home oxygen and steroids for chronic lung disease. Maternal and delivery room conditions were compared between those neonates who died in the delivery room and those who died in the neonatal ward.

#### 2.3. Ethics

Data were de-identified and the key to patient details was kept separately and only known to the principal investigator. Ethical clearance for the study was obtained from the Human Research Ethics Committee of the University of the Witwatersrand. Permission to conduct the study was obtained from the Chief Executive Officer of CMJAH. One of the authors was the gatekeeper of the neonatal database; additional permission to access the database was not required.

#### 3. Results

The database was accessed on 20 February 2016, and there were 5695 neonatal records on the database; 5386 records were for neonates born within the study period. There were 26 records with missing outcome data, four babies who had a birth weight below 400 g and 338 neonates who were admitted to the unit after 48 h. Thus, 5018 records were included in the review. The mean birth weight was 2148 g (SD 972) and the mean gestational age was 34.2 weeks (SD 4.8). The mean duration of stay was 13.75 days (SD 18.0).

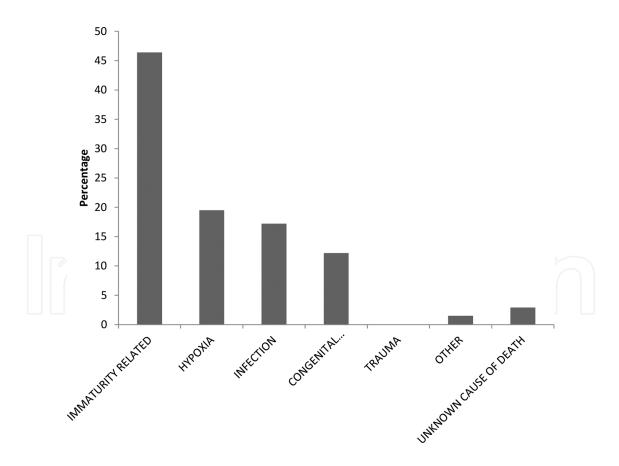


Figure 1. Causes of neonatal deaths in Johannesburg, South Africa, between 01 January 2013 and 31 December 2015.

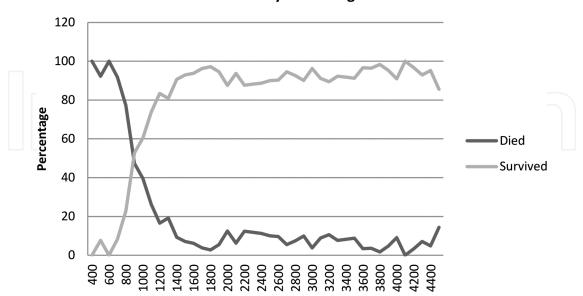
There were 724 deaths, giving an overall mortality rate of 14.4%, alternatively expressed as a percentage surviving to discharge of 85.6. Seventy-three percent (530/724) of neonates died in the early neonatal period, within seven days of birth. There were 147 (20.3%) deaths in the delivery room and seventy neonates (9.6%) died within the first 12 hours of admission to the neonatal ward. The various causes of neonatal death according to the PPIP classification are shown in **Figure 1**.

#### 3.1. Birth weight

The mortality rate was strongly associated with birth weight. There were 3134 LBW neonates, with a mortality rate of 18.6% (586/3134). The majority of deaths in LBW neonates occurred in VLBW neonates (30.1% (479/1590)). Significantly more VLBW neonates died than babies >1500 g (30.1% vs. 7.1%; p < 0.001). The number of neonates and those who died in each birth weight category is shown in **Table 1**.

| Birth weight (g) | Number | Died | % Mortality |  |
|------------------|--------|------|-------------|--|
| <1000 (ELBW)     | 524    | 315  | 60.1        |  |
| 1000–1499        | 1066   | 164  | 15.4        |  |
| 1500–2499        | 1544   | 107  | 6.4         |  |
| 2500–3999 (TAGA) | 1730   | 130  | 7.5         |  |
| >4000 (TLGA)     | 154    | 8    | 5.2         |  |

Table 1. Distribution of deaths by birth weight category for neonates at CMJAH between 2013 and 2015.



#### Survival by birth weight

Figure 2. Percentage surviving by birth weight for neonates at CMJAH between 2013 and 2015.

The highly significant association between decreasing birth weight and increasing mortality is shown in **Figure 2** which depicts how the proportion surviving increases as birth weight increases. The percentage survival for a birth weight of 900 g is 52.8.

#### 3.1.1. Demographic and clinical characteristics in VLBW neonates compared to bigger babies

Further results are reported for VLBW neonates compared to bigger babies. Demographic, maternal, and clinical characteristics are shown in **Table 2**. Certain conditions only occur in bigger babies and were thus not reported for VLBW neonates, namely meconium aspiration syndrome (MAS), persistent pulmonary hypertension of the neonate (PPHN), hypoxic ischemic encephalopathy (HIE), and cerebral cooling.

| Factor                        | Cases | %    | <1500 | g    | >1500 | g    | P-value |
|-------------------------------|-------|------|-------|------|-------|------|---------|
|                               |       |      | n     | %    | n     | %    |         |
| Birth defect                  | 264   | 5.4  | 29    | 1.9  | 226   | 6.8  | < 0.001 |
| Delivery room death           | 147   | 2.9  | 54    | 3.5  | 33    | 1    | < 0.001 |
| Birth place                   |       |      |       |      |       |      | < 0.001 |
| -Other unit                   | 718   | 14.4 | 162   | 10.5 | 552   | 14.5 |         |
| -Born outside health facility | 194   | 3.9  | 85    | 5.5  | 104   | 3.8  | < 0.001 |
| -Inborn                       | 4059  | 81.6 | 1302  | 84.1 | 2707  | 80.5 |         |
| Antenatal care                | 4017  | 85.6 | 1150  | 77.4 | 2829  | 89   | < 0.001 |
| Antenatal steroids            | 889   | 27.8 | 657   | 74.5 | 225   | 25.5 | < 0.001 |
| Antenatal magnesium sulfate   | 102   | 2.3  | 65    | 4.7  | 37    | 1.2  | < 0.001 |
| Chorioamnionitis              | 127   | 2.8  | 47    | 3.4  | 79    | 2.5  | 0.118   |
| Maternal hypertension         | 651   | 14.3 | 377   | 27   | 269   | 8.7  | < 0.001 |
| Maternal HIV                  | 1410  | 29.4 | 456   | 31   | 946   | 28.9 | 0.147   |
| Maternal syphilis             | 79    | 1.6  | 29    | 2.1  | 48    | 1.5  | 0.164   |
| Maternal diabetes             | 134   | 2.9  | 7     | 0.5  | 127   | 4    | <0.001  |
| Maternal TB                   | 44    | 1    | 11    | 0.8  | 33    | 1.1  | 0.416   |
| Teenage mother                | 114   | 2.3  | 39    | 2.7  | 75    | 2.4  | 0.529   |
| Vaginal delivery              | 2191  | 44.7 | 661   | 43.3 | 1489  | 44.9 | 0.288   |
| Male gender                   | 2752  | 55   | 720   | 46.5 | 1995  | 58.7 | < 0.001 |
| Multiple gestation            | 586   | 11.8 | 272   | 17.8 | 301   | 8.9  | < 0.001 |
| Delivery room resuscitation   | 1532  | 31   | 643   | 43.7 | 850   | 25   | < 0.001 |
| Early onset sepsis            | 173   | 3.4  | 62    | 4.2  | 111   | 3.3  | 0.133   |
| Oxygen on day 28              | 404   | 8.1  | 347   | 25.6 | 57    | 1.7  | < 0.001 |
|                               |       |      |       |      |       |      |         |

| Factor                        | Cases | %    | <1500 | 00 g >1500 g |     | P-value |         |
|-------------------------------|-------|------|-------|--------------|-----|---------|---------|
|                               |       |      | n     | %            | n   | %       | _       |
| IVH 3/4                       | 61    | 1.2  |       |              | N/A | N/A     |         |
| PVL                           | 11    | 0.2  | 8     | 0.7          | 3   | 0.1     | < 0.001 |
| Died within 12 h of admission | 70    | 1.4  | 39    | 2.6          | 31  | 0.9     | < 0.001 |
| Pneumothorax                  | 36    | 0.7  | 10    | 0.7          | 26  | 0.8     | 0.444   |
| Pulmonary hemorrhage          | 32    | 0.7  | 27    | 1.8          | 5   | 0.1     | < 0.001 |
| HIE 2/3                       | 174   | 3.6  | N/A   | N/A          |     |         |         |
| Cerebral cooling              | 103   | 38.1 | N/A   | N/A          |     |         |         |
| Meconium aspiration syndrome  | 264   | 7.8  | N/A   | N/A          |     |         |         |
| PPHN                          | 54    | 1.6  | N/A   | N/A          |     |         |         |
| HMD                           | 2004  | 41.1 | 1347  | 89.6         | 657 | 19.5    | < 0.001 |
| NCPAP                         | 1565  | 32.5 | 1015  | 70.1         | 550 | 32.5    | < 0.001 |
| IPPV                          | 692   | 14.4 | 299   | 21           | 393 | 11.7    | < 0.001 |
| NCPAP without IPPV            | 1228  | 24.5 | 795   | 78.9         | 433 | 79.4    | 0.816   |
| Surfactant therapy            | 1580  | 33.1 | 1038  | 69.8         | 542 | 16.5    | < 0.001 |
| Steroids for CLD              | 216   | 5.7  | 199   | 13.8         | 17  | 0.7     | < 0.001 |
| PDA                           | 245   | 5    | 152   | 10.2         | 93  | 2.8     | < 0.001 |
| NEC                           | 156   | 3.2  | 107   | 7.2          | 49  | 1.5     | < 0.001 |
| Other surgery                 | 136   | 2.9  | 29    | 2            | 107 | 3.3     | 0.014   |
| Packed cell transfusion       | 674   | 13.4 | 527   | 35.9         | 147 | 4.4     | < 0.001 |
| Exchange transfusion          | 24    | 1.5  | 9     | 1            | 15  | 2.2     | 0.039   |
| Hypoglycemia                  | 525   | 10.8 | 185   | 12.3         | 340 | 10      | 0.02    |
| Hyperglycemia                 | 375   | 7.7  | 287   | 19.1         | 88  | 2.6     | <0.001  |
| Hypernatraemia                | 169   | 3.5  | 148   | 9.8          | 21  | 0.6     | <0.001  |
| Metabolic acidosis            | 185   | 3.8  | 92    | 6.1          | 93  | 2.8     | <0.001  |
| Late onset sepsis             | 608   | 12.6 | 421   | 28.3         | 187 | 5.6     | < 0.001 |

**Table 2.** Demographic, maternal and clinical characteristics by birth weight for neonates at CMJAH between 2013 and 2015.

#### 3.2. Risk factors for neonatal death

Neonates who survived were born at a significantly more mature gestational age than those who died (34.8 weeks [SD 4.4] vs. 30.5 weeks [SD 5.5]; p < 0.001). Similarly, the birth weight of neonates who survived was significantly greater than those who died (2260 g [SD 932] vs. 1495

g [SD 940]; p < 0.001). Survivors stayed in hospital for a longer period of time than those neonates who died (14.8 days [SD 18.5] vs. 7.1 days [SD 13.3]; p < 0.001). Body temperature on admission was significantly higher in neonates who survived compared to those who died (36.3°C [SD 8.0] vs. 35.6°C [SD 1.6]).

Conditions significantly associated with death in all the neonates, including those who died in the delivery room, are shown in **Table 3**. Only data for babies who died are reported. The percentages refer to the number of babies who died with and without the various conditions. For example, 38.5% (102) of babies who had a major birth defect died and 12.8% (596) of babies without a major birth defect died. Percentages are reported per the total number of complete cases for each condition—missing data were excluded. All other conditions were not significantly associated with death in the whole group of neonates.

| Factor                   | Condition present |       | Condition a | Condition absent |         |  |
|--------------------------|-------------------|-------|-------------|------------------|---------|--|
|                          | # Died            | %     | # Died      | %                |         |  |
| Birth defect             | 102               | 38.5  | 596         | 12.8             | < 0.001 |  |
| Antenatal care           | 503               | 12.5  | 162         | 23.0             | < 0.001 |  |
| Maternal HIV             | 209               | 14.8  | 433         | 12.8             | 0.061   |  |
| Maternal diabetes        | 9                 | 6.7   | 607         | 13.5             | 0.023   |  |
| Vaginal delivery         | 404               | 18.4  | 303         | 11.2             | < 0.001 |  |
| Birth place              |                   |       |             |                  |         |  |
| Another unit             | 100               | 13.9  |             |                  |         |  |
| Outside health facility  | 53                | 27.3  |             |                  | < 0.001 |  |
| Inborn                   | 569               | 34.0  |             |                  |         |  |
| Multiple gestation       | 101               | 17.2  | 612         | 14.0             | 0.035   |  |
| Initial resuscitation    | 393               | 25.7  | 312         | 9.2              | < 0.001 |  |
| MAS                      | 34                | `12.9 | 156         | 5.0              | < 0.001 |  |
| Pneumothorax             | 12                | 33.3  | 559         | 11.6             | < 0.001 |  |
| Pulmonary haemorrhage    | 24                | 75.0  | 558         | 11.6             | < 0.001 |  |
| PPHN                     | 25                | 46.3  | 165         | 5.0              | < 0.001 |  |
| Hyaline membrane disease | 399               | 19.9  | 183         | 6.4              | < 0.001 |  |
| NCPAP                    | 304               | 19.4  | 266         | 8.2              | < 0.001 |  |
| IPPV                     | 219               | 31.6  | 348         | 8.5              | < 0.001 |  |
| NCPAP without IPPV       | 205               | 16.7  | 98          | 30.2             | < 0.001 |  |
| Surfactant therapy       | 310               | 19.6  | 256         | 8.0              | < 0.001 |  |

| Factor             | Condition p | oresent | Condition a | Condition absent |         |
|--------------------|-------------|---------|-------------|------------------|---------|
|                    | # Died      | %       | # Died      | %                |         |
| PVL                | 4           | 36.4    | 424         | 9.5              | 0.003   |
| IVH grade 3/4      | 29          | 47.5    | 32          | 52.5             | < 0.001 |
| HIE grade 2/3      | 48          | 27.9    | 124         | 72.1             | < 0.001 |
| NEC                | 65          | 41.7    | 508         | 10.9             | <0.001  |
| Surgery (not NEC)  | 38          | 27.7    | 524         | 11.4             | < 0.001 |
| Blood transfusion  | 148         | 22.0    | 422         | 10.1             | < 0.001 |
| Hypoglycemia       | 78          | 14.9    | 504         | 11.6             | 0.029   |
| Hyperglycemia      | 165         | 44.0    | 417         | 9.3              | < 0.001 |
| Hypernatraemia     | 64          | 37.9    | 518         | 11.0             | < 0.001 |
| Metabolic acidosis | 101         | 54.6    | 481         | 10.3             | < 0.001 |
| Late onset sepsis  | 135         | 22.0    | 434         | 10.3             | < 0.001 |

**Table 3.** Factors associated with death in all neonates who died (*n* = 724), including delivery room deaths.

The results of binary logistic regression, considering whether neonate survived to discharge as the outcome variable, are shown in **Table 4**. The chances of survival decreased with metabolic acidosis, hyperglycemia, mechanical ventilation, major birth defect and the need for resuscitation at birth, while increasing birth weight and gestational age and delivery by Caesarean section were associated with an increased chance of survival.

| Condition              | Odds ratio |       | OR    |
|------------------------|------------|-------|-------|
|                        |            | Lower | Upper |
| Metabolic acidosis     | 0.135      | 0.09  | 0.204 |
| Hyperglycemia          | 0.307      | 0.23  | 0.409 |
| Mechanical ventilation | 0.357      | 0.278 | 0.46  |
| Birth weight           | 1.001      | 1     | 1.001 |
| Major birth defect     | 0.118      | 0.079 | 0.175 |
| Gestational age        | 1.109      | 1.054 | 1.167 |
| Caesarean section      | 1.803      | 1.444 | 2.251 |
| Resuscitated at birth  | 0.395      | 0.315 | 0.495 |
| Constant               | 0.21       |       |       |

**Table 4.** Results of binary logistic regression model for factors associated with survival in all neonates at CMJAH between 2013 and 2015.

#### 3.2.1. Binary logistic regression: VLBW neonates

The results of binary logistic regression considering survival to discharge as the outcome variable were performed for VLBW neonates (see **Table 5**). The percentage survival increased with increasing birth weight, delivery by Caesarean section and the use of NCPAP without the need for mechanical ventilation. Maternal HIV, hyperglycemia, resuscitation at birth, pulmonary hemorrhage, NEC, and metabolic acidosis were associated with a reduced chance of survival.

| Factor                    | Odds ratio | 95% CI |       |
|---------------------------|------------|--------|-------|
|                           |            | Lower  | Upper |
| Birth weight (g)          | 1.005      | 1.004  | 1.006 |
| Maternal HIV              | 0.582      | 0.394  | 0.861 |
| Caesarean section         | 1.81       | 1.242  | 2.638 |
| Resuscitated at birth     | 0.589      | 0.405  | 0.858 |
| Pulmonary haemorrhage     | 0.176      | 0.063  | 0.493 |
| Necrotising enterocolitis | 0.252      | 0.139  | 0.459 |
| Hyperglycaemia            | 0.489      | 0.325  | 0.737 |
| Metabolic acidosis        | 0.098      | 0.051  | 0.191 |
| NCPAP without ventilation | 2.032      | 1.314  | 3.142 |
| Constant                  | 0.022      |        |       |

**Table 5.** Binary logistic regression for factors associated with survival to discharge in VLBW neonates at CMJAH between 2013 and 2015.

#### 3.2.2. Binary logistic regression: bigger neonates

The results of binary logistic regression considering survival to discharge as the outcome are shown in **Table 6**. Birth weight was not significantly different between survivors and non-survivors in this weight category. Decreasing gestational age, the need for resuscitation at birth, mechanical ventilation, metabolic acidosis, and hyperglycemia were all associated with a reduced chance of survival.

| Factor                  | Odds ratio | 95% CI for | OR    |
|-------------------------|------------|------------|-------|
|                         |            | Lower      | Upper |
| Gestational age (weeks) | 0.937      | 0.881      | 0.996 |
| Resuscitated at birth   | 0.375      | 0.249      | 0.564 |
| Mechanical ventilation  | 0.12       | 0.078      | 0.184 |
| Metabolic acidosis      | 0.244      | 0.122      | 0.486 |
| Hyperglycaemia          | 0.16       | 0.08       | 0.321 |
| Constant                | 668.746    |            |       |

**Table 6.** Binary logistic regression for factors associated with survival to discharge in bigger neonates at CMJAH between 2013 and 2015.

#### 3.2.3. Delivery room deaths

Neonates who died in the delivery room were less likely to have received antenatal steroids and be delivered to mothers with hypertension or HIV, compared to neonates who died in the neonatal wards. Delivery room deaths were associated with vaginal delivery and were more likely in neonates who had been resuscitated at birth (see **Table 7**). Neonates who died in the delivery room had a lower body temperature on admission than those who died in the neonatal wards (34.6°C [SD 2.8] compared to 35.8°C [SD 1.2]; p < 0.001). All other variables including birth weight and gestational age were not different between neonates who died in the delivery room compared to the neonatal wards.

| Condition present     | Delivery room death | Percentage | Neonatal ward death | Percentage | <i>P</i> -value |
|-----------------------|---------------------|------------|---------------------|------------|-----------------|
| Antenatal steroids    | 15                  | 9.4        | 144                 | 32.3       | 0.001           |
| Maternal hypertension | 12                  | 12.2       | 86                  | 17.3       | 0.032           |
| Maternal HIV          | 25                  | 21.6       | 184                 | 35.2       | 0.004           |
| Caesarean section     | 51                  | 35.4       | 252                 | 44.8       | 0.042           |
| Resuscitated at birth | 95                  | 64.6       | 297                 | 53.2       | 0.013           |

**Table 7.** Maternal and delivery room factors compared between babies who died in the delivery room and those who died in the neonatal wards.

#### 4. Discussion

The ongoing audit of neonatal mortality and neonatal care to determine risk factors for poor outcome is essential so that correct interventions can be implemented. The MDG 2015 report states that better readily available data is urgently needed to guide health policies [1]. There is a slogan in the report that says "together we can measure what we treasure". The so-called "Plan Do Study Act [PDS] cycle is a tool for quality improvement projects [10]. Ongoing clinical audit is fundamental to quality improvement projects, both for planning the intervention and then measuring the benefit of the intervention [11, 12]. It is also essential to have appropriate local data available; different NICUs and neonatal populations have different problems and need tailored solutions. For example, maternal HIV is an important issue in the current study, but would not apply in a European setting.

The best example of clinical audit and quality improvement in neonatal care is the Vermont Oxford Network [VON] (www.vtoxford.org). The VON is a multinational multicenter collaboration of neonatal units established in 1989 with the aim of improving quality and effectiveness of neonatal care by research, education and quality improvement projects [13]. There are currently more than 1000 neonatal units from around the world that participate in the VON. Collaborative multi-disciplinary quality improvement projects [NIC/Q] are conducted annually [14].

The present study was an audit of neonatal survival and risk factors for poor outcome in Johannesburg, South Africa. The overall neonatal survival rate in the present study was 85.6%.

Birth weight greatly influenced survival with 69.1% of VLBW surviving compared to 92.1% of neonates above 1500 g birth weight. The VLBW survival in our unit was significantly less than that reported in the VON [www.vtoxford.org] for the same period (69.1% vs. 85.6%). Neonatal mortality rates among different neonatal units are highly variable, but the rates reported in the present study are within the reported range for developing nations [3]. The current neonatal survival rates are better than those reported from NICUs in The Gambia [4] and Ethiopia [15], but worse than those reported from a NICU in Thailand [16]. It must be noted that different mortality rates will be reported depending on which neonates are included in the audit—the present study included neonates from 400 g birth weight and those who died within the delivery room—omission of these would improve the results.

The most important causes of neonatal death in the present study were complications of prematurity, perinatal asphyxia, infection, and birth defects. These findings are similar to other studies evaluating risk factors for neonatal mortality [1, 17], although the contribution of prematurity to neonatal death is considerably higher than that reported in United Nations Millennium Development Goal Report 2015 [1] (42.3% vs. 35%). Birth weight is closely linked to gestational age in LBW neonates; the higher mortality with decreasing birth weight in the present study corresponds to increasingly premature neonates. It is interesting to note that in bigger babies, gestational age, rather than birth weight, was associated with survival. Almost 15% of deaths in the present study were due to congenital abnormalities; this reflects the fact that the unit was a referral centre for pediatric surgery; so many neonates with major congenital abnormalities were referred in for surgery.

The present results are also similar to a report from a private healthcare group in South Africa, who found that birth weight, Apgar score, and mode of delivery were all associated with neonatal mortality [18]. This is interesting, as the majority of patients in the private health care group were of White and Indian ethnicity, whereas those in the current report were almost exclusively Black African.

Most of the neonatal deaths in the current study occurred in VLBW neonates; therefore resources need to be focused on this group of neonates in order to reduce childhood mortality. Decreasing birth weight, maternal HIV, the need for resuscitation at birth, pulmonary hemorrhage, NEC, hyperglycemia, and metabolic acidosis were all associated with a decreased chance of survival in VLBW neonates, while delivery by Caesarean section and the use of NCPAP without the need for mechanical ventilation significantly increased survival. These findings are similar to reports from the same unit [5, 6]. Interventions need to be devised to address these specific risk factors, such as ensuring prevention of mother to child transmission of HIV, providing proper prompt neonatal resuscitation, maintaining normoglycemia, and promoting breastfeeding. All preterm neonates, irrespective of birth weight, should be provided with NCPAP. The use of surfactant and mechanical ventilation may not be available in all NICUs in LMICS due to resource limitations. If necessary, surfactant and mechanical ventilation can be rationed using prognostic criteria. The association of better survival with Caesarean section is a more difficult one - it is possible that neonates delivered by Caesarean section are the "better babies." These mothers may have attended antenatal care, been admitted earlier in labor, and received antenatal steroids. It is therefore possible that Caesarean section

is a confounding variable. It is certainly not feasible to suggest that all preterm neonates in LMICS be delivered by Caesarean section. Other factors such as antenatal care, antenatal steroid use, and neonatal infection were not significantly predictive of survival in the present study. This does not mean, however, that regular antenatal care attendance, the use of antenatal steroids, and infection control should be omitted from interventions to improve VLBW survival.

The factors associated with poor survival in bigger neonates included decreasing gestational age, the need for resuscitation at birth, mechanical ventilation, metabolic acidosis, and hyperglycemia. This emphasizes the need for all birth attendants to be skilled in neonatal resuscitation. It is possible that mechanical ventilation will not be available in many NICUs in LIMICs, but bigger preterm infants can be successfully managed with surfactant therapy and NCPAP [9].

A recent report from Burundi showed that the neonatal survival rates were significantly improved in a low resourced district hospital, without specialist care [19]. This was achieved by integrating neonatal and obstetric services, with an emphasis on prompt referral and transfer of mothers in preterm labor, the ongoing on-site training of staff with clear protocols for case management, provision of essential equipment, and providing complementary kangaroo mother care and NICU facilities.

In conclusion, ongoing clinical audit is integral to the process of quality improvement, to develop appropriate health care policies and to monitor the impact of these policies. Focus on neonatal care and especially that of VLBW neonates is essential if we are to achieve the SDG goal of reducing neonatal mortality to 12 per 1000 births.

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#### References

 United Nations. The Millenium Development Goals Report 2015. United Nations, 2015 July 2015. Report No.

- [2] Kumar S, Kumar N, Vivekadhish S. Millennium Development Goals [MDGs] to Sustainable Development Goals [SDGs]: addressing unfinished agenda and strengthening sustainable development and partnership. Indian J Community Med. 2016;41[1]: 1–4.
- [3] Chow S, Chow R, Popovic M, Lam M, Popovic M, Merrick J, et al. A selected review of the mortality rates of neonatal intensive care units. Front Public Health. 2015;3:225.
- [4] Okomo UA, Dibbasey T, Kassama K, Lawn JE, Zaman SM, Kampmann B, et al. Neonatal admissions, quality of care and outcome: 4 years of inpatient audit data from The Gambia's teaching hospital. Paediatr Int Child Health. 2015;35[3]:252–264.
- [5] Ballot DE, Chirwa TF, Cooper PA. Determinants of survival in very low birth weight neonates in a public sector hospital in Johannesburg. BMC Pediatr. 2010;10:30.
- [6] Ballot DE, Chirwa T, Ramdin T, Chirwa L, Mare I, Davies VA, et al. Comparison of morbidity and mortality of very low birth weight infants in a Central Hospital in Johannesburg between 2006/2007 and 2013. BMC Pediatr. 2015;15:20.
- [7] Kalimba EM BD. Survival of extremely low-birth-weight infants. S Afr J Child Health. 2013;7[1]:13–16.
- [8] Harris PA, Taylor R, Thielke R, Payne J, Gonzalez N, Conde JG. Research electronic data capture [REDCap]—a metadata-driven methodology and workflow process for providing translational research informatics support. J Biomed Inform. 2009;42[2]:377– 381.
- [9] Jardine C, Ballot DE. The use of nasal CPAP at Charlotte Maxeke Johannesburg Academic Hospital. S Afr J Child Health. 2015;9[1]:4.
- [10] Speroff T, James BC, Nelson EC, Headrick LA, Brommels M. Guidelines for appraisal and publication of PDSA quality improvement. Qual Manag Health Care. 2004;13[1]: 33–39.
- [11] Dalal PG, Porath J, Parekh U, Dhar P, Wang M, Hulse M, et al. A quality improvement project to reduce hypothermia in infants undergoing MRI scanning. Pediatr Radiol. 2016.
- [12] Read B, Lee DS, Fraser D. Evaluation of a practice guideline for the management of respiratory distress syndrome in preterm infants: a quality improvement initiative. Paediatr Child Health. 2016;21[1]:e4-9.
- [13] Horbar JD. The Vermont Oxford Network: evidence-based quality improvement for neonatology. Pediatrics. 1999;103[1 Suppl E]:350–359.
- [14] Horbar JD, Carpenter JH, Buzas J, Soll RF, Suresh G, Bracken MB, et al. Collaborative quality improvement to promote evidence based surfactant for preterm infants: a cluster randomised trial. BMJ. 2004;329[7473]:1004.

- [15] Kokeb M, Desta T. Institution Based prospective cross-sectional study on patterns of neonatal morbidity at Gondar University Hospital Neonatal Unit, North-West Ethiopia. Ethiop J Health Sci. 2016;26[1]:73–79.
- [16] Sritipsukho S, Suarod T, Sritipsukho P. Survival and outcome of very low birth weight infants born in a university hospital with level II NICU. J Med Assoc Thai. 2007;90[7]: 1323–1329.
- [17] Dhaded SM, Somannavar MS, Vernekar SS, Goudar SS, Mwenche M, Derman R, et al. Neonatal mortality and coverage of essential newborn interventions 2010–2013: a prospective, population-based study from low-middle income countries. Reprod Health. 2015;12 Suppl 2:S6.
- [18] Pepler PT, Uys DW, Nel DG. Predicting mortality and length-of-stay for neonatal admissions to private hospital neonatal intensive care units: a Southern African retrospective study. Afr Health Sci. 2012;12[2]:166–173.
- [19] Ndelema B, Van den Bergh R, Manzi M, van den Boogaard W, Kosgei RJ, Zuniga I, et al. Low-tech, high impact: care for premature neonates in a district hospital in Burundi. A way forward to decrease neonatal mortality. BMC Res Notes. 2016;9[1]:28.

