Peripherally inserted central catheters versus non-tunnelled ultrasound-guided central venous catheters in newborns: a retrospective observational study

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ABSTRACT

Objectives We aimed to compare the success rates and other catheter-related parameters between peripherally inserted central catheters (PICCs) and non-tunnelled ultrasound-guided central venous catheters (USG-CVCs) in newborns including femoral, jugular, brachiocephalic and subclavian lines.

Design This was a retrospective observational study.

Setting The study was performed in a level III neonatal intensive care unit (NICU) in Qatar, as a single-site study.

Participants This study included 1333 neonates who required CVC insertion in the NICU from January 2016 to December 2018. Of those, we had 1264 PICCs and 69 non-tunnelled USG-CVCs.

Outcome measures The success rate and other catheter-related complications in the two groups.

Results The overall success rate was 88.4% in the USG-CVCs (61/69) compared with 90% in the PICCs (1137/1264) group (p=0.68). However, the first prick success rate was 69.4% in USG-CVCs (43/69) compared with 63.6% in the PICCs (796/1264) group. Leaking and central line-associated blood stream infection (CLABSI) were significantly higher in the USG-CVC group compared with the PICC group (leaking 16.4% vs 2.3%, p=0.0001) (CLABSI 8.2% vs 3.1%, p=0.03). CLABSI rates in the PICC group were 1.75 per 1000 catheter days in 2016 and 3.3 in 2017 compared with 6.91 in 2016 (p=0.0001) and 14.32 in 2017 (p=0.0001) for the USG-CVCs. USG-CVCs had to be removed due to catheter-related complications in 52.5% of the cases compared with 29.9% in PICCs, p=0.001.

Conclusions The overall complication rate, CLABSI and leaking are significantly higher in non-tunnelled USG-CVCs compared with the PICCs. However, randomised controlled trials with larger sample sizes are desired. Proper central venous device selection and timing, early PICC insertion and early removal approach, dedicated vascular access team development, proper central venous line maintenance, central line simulation workshops and US-guided insertions are crucial elements for patient safety in NICU.

INTRODUCTION

Peripherally inserted central catheters (PICCs) were described for the first time by Shaw in 1973. Since then, they have been used extensively due to their features.1 PICC insertion by direct superficial peripheral vein puncture offers long-term venous access for both term and preterm neonates and is often indicated in neonatal intensive care unit (NICU) for parental nutrition, long-term intravenous medications, antibiotic therapy and vesicant drug administration.2,3

Non-tunnelled ultrasound-guided central venous catheters (USG-CVCs) are inserted in neonates in special circumstances, for example, central venous pressure monitoring, blood withdrawal, haemodialysis and for all other infusions and medications when PICC insertion fails.4 They are inserted in the internal jugular, brachiocephalic, subclavian and femoral veins under ultrasound guidance.1,5–7

There is a limited number of studies comparing PICCs with USG-CVCs in neonates that necessitated further research and comparative analysis. This study aimed to compare the success rates and other catheter-related parameters in PICCs and the non-tunnelled USG-CVCs in NICU between 2016 and 2018.

METHODS

This single-centre retrospective study was conducted in the NICU at the Women’s Wellness and Research Centre (WWRC), Hamad Medical Corporation (HMC), Doha.
Qatar. WWRC is the main specialist hub for women and newborns health services in Qatar with more than 18,000 deliveries per year. The NICU in WWRC is a level III mainly medical unit with 112 beds and more than 2000 admissions per year with limited congenital cardiac or surgery cases.

Patient and public involvement

Patients and/or the public were not involved in the design, conduct, reporting or dissemination plans of this research.

Participants

A total of 1333 cases were evaluated in this study. This includes 1264 babies who had PICC insertion and 69 who had non-tunnelled USG-CVC insertion. Related information for all cases between January 2016 and December 2018 was collected from the electronic medical system at the NICU.

A fully dedicated PICC insertion team was launched in January 2017. The PICC team has been expanded over time to include 15 neonatologist physicians, 1 neonatal nurse practitioner as well as seven NICU nurses. The team was trained in central line simulation workshops to insert PICC by the catheter-over-needle technique and the modified Seldinger technique. The central line simulation workshop is a full-day workshop that was founded by the neonatal simulation simulation team and is accredited by the Department of Healthcare Professions in the Ministry of Public Health with a total of 7 continuous professional development (CPD) hours both category I (1/7) and category III (6/7). The PICC team works in harmony and collaboration with 30 well-trained NICU nurses who are members of the neonatal specialist nursing (NSN) team. The NSN determines the patient’s eligibility, takes care of the central line maintenance using transparent semipermeable dressing, enters the data in the electronic database and gets the blood samples. There is no difference in the central line type of care, frequency or personnel in all types of catheters. In addition to their role in central line insertion and maintenance, the NSN team attends high-risk deliveries and play a pivotal role in neonatal transportation.

In our NICU, the indications for PICC insertions are the birth weight of <1500 g, the requirement of intravenous fluids for >5 days, the requirement of intravenous medications for >7 days, the requirement of hyperosmolar intravenous fluid therapy >700 mOsmol/L and the requirement of >3 peripheral intravenous catheters (PIVC) insertions in the last 24 hours. A successful catheter insertion means a catheter inserted into a proper central venous position that can be used with its tip located either in the superior or inferior vena cava. As per our institutional guideline, two pricks are allowed per operator with a maximum of three in difficult lines. After three unsuccessful pricks, the procedure should be terminated.

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| Variables                        | Total n=1333 | USG-CVC 69 (5.2%) | PICC 1264 (94.8%) | P value |
|---------------------------------|--------------|-------------------|-------------------|---------|
| **Year**                        |              |                   |                   |         |
| 2016                            | 376 (28.2)   | 42 (60.9)         | 334 (26.4)        | 0.001   |
| 2017                            | 507 (38)     | 27 (39.1)         | 480 (38)          |         |
| 2018                            | 450 (33.8)   | 0 (0)             | 450 (35.6)        |         |
| **Side of the body**            |              |                   |                   |         |
| Left                            | 498 (41.1)   | 14 (22.6)         | 484 (42.1)        | 0.002   |
| Right                           | 715 (58.9)   | 48 (77.4)         | 667 (57.9)        |         |
| **Site of insertion**           |              |                   |                   |         |
| Upper extremities               | 360 (29.5)   | 37 (53.6)         | 323 (28)          | 0.001   |
| Lower extremities               | 861 (70.5)   | 32 (46.4)         | 829 (72)          |         |
| **Number of pricks**            |              |                   |                   |         |
| First prick                     | 839 (63.9)   | 43 (69.4)         | 796 (63.6)        | 0.001   |
| Second prick                    | 305 (23.2)   | 8 (12.9)          | 297 (23.7)        |         |
| Third prick                     | 145 (11)     | 6 (9.7)           | 139 (11.1)        |         |
| Fourth prick                    | 22 (1.7)     | 4 (6.5)           | 18 (1.4)          |         |
| Fifth prick                     | 1 (0.1)      | 0 (0)             | 1 (0.1)           |         |
| Sixth prick                     | 2 (0.2)      | 1 (1.6)           | 1 (0.1)           |         |
| **Reason for insertion**        |              |                   |                   |         |
| Difficult intravenous insertion | 8 (0.6)      | 0 (0)             | 8 (0.6)           | 0.47    |
| Hypoglycaemia                   | 10 (0.8)     | 0 (0)             | 10 (0.8)          |         |
| Long-term intravenous fluid     | 1286 (96.6)  | 68 (100)          | 1218 (96.4)       |         |
| medication therapy              | 27 (2)       | 0 (0)             | 27 (2.1)          |         |
| **Catheter insertion success rate** |           |                   |                   |         |
| Successful                      | 1198 (89.9)  | 61 (88.4)         | 1137 (90)         | 0.68    |
| Not successful                  | 135 (10.1)   | 8 (11.6)          | 127 (10)          |         |
| **Reason for removal**          |              |                   |                   |         |
| CLABSI                          | 40 (3.4)     | 5 (8.2)           | 35 (3.1)          | 0.031   |
| Leaking                         | 36 (3)       | 10 (16.4)         | 26 (2.3)          | 0.001   |
| Accidental removal              | 8 (0.7)      | 1 (1.6)           | 7 (0.6)           | 0.40    |
| Broken catheter                 | 7 (0.6)      | 0 (0)             | 7 (0.6)           | 0.69    |
| Local redness and swelling      | 104 (8.7)    | 5 (8.2)           | 99 (8.8)          | 0.88    |
| Occlusion                       | 42 (3.5)     | 0 (0)             | 42 (3.7)          | 0.13    |
| Malposition                     | 13 (1.1)     | 0 (0)             | 13 (1.1)          | 0.50    |
| Elective                        | 833 (69.9)   | 29 (47.5)         | 804 (71.1)        | 0.001   |
| Death                           | 39 (3.3)     | 5 (8.2)           | 34 (3)            | 0.03    |
| Phlebitis                       | 70 (5.9)     | 6 (9.8)           | 64 (5.7)          | 0.18    |
| **Gestational age (weeks)**     |              |                   |                   |         |
| Mean±SD                         | 29.55±4.30   | 33.88±6.34        | 29.32±4.03        | 0.001   |
| Median (IQR)                    | 29 (27, 31)  | 37 (26, 40)       | 29 (27, 31)       |         |
| **Gestational age**             |              |                   |                   |         |
| 22–28 weeks                     | 602 (45.2)   | 22 (31.9)         | 580 (45.9)        | 0.001   |
| >28–32 weeks                    | 490 (36.8)   | 4 (5.8)           | 486 (38.4)        |         |
| >32–36 weeks                    | 100 (7.5)    | 6 (11.6)          | 92 (7.3)          |         |
| >36 weeks                       | 141 (10.6)   | 35 (50.7)         | 106 (8.4)         |         |
| **Birth weight (g)**            |              |                   |                   |         |
| Mean±SD                         | 1282±692.1   | 2161±1140.3       | 1234±624.9        | 0.001   |
| Median (IQR)                    | 1095 (850, 1400) | 2530 (970, 3122) | 1080 (840, 1370) |         |
| **Birth weight**                |              |                   |                   |         |
| BW ≤1 kg                        | 561 (42.1)   | 21 (30.4)         | 540 (42.7)        | 0.001   |
| BW >1–2 kg                      | 618 (46.5)   | 10 (14.5)         | 608 (48.1)        |         |
| BW >2–3 kg                      | 87 (6.5)     | 16 (23.2)         | 71 (5.6)          |         |
| BW >3 kg                        | 67 (5)       | 22 (31.9)         | 42 (3.6)          |         |

This is a retrospective study design and for some parameters, the data values were incomplete due to the unavailability of the information in the patients' record files. All percentage (%) was computed using non-missing data values.

CLABSI, central line-associated bloodstream infection; PICCs, peripherally inserted central catheters; USG-CVCs, ultrasound-guided central venous catheters.
by the paediatric surgeon or the paediatric anaesthetist on-call physician under US guidance. Currently, we use the handheld wireless probe-type ultrasound scanner machine to guide the catheter insertion and for the catheter tip location.

We followed the Centers for Disease Control and Prevention (CDC) definition for central line-associated bloodstream infection (CLABSI) and CLABSI rate. CLABSI is defined as a laboratory-confirmed bloodstream infection (LCBI) where an eligible bloodstream infection (BSI) organism is identified, and an eligible central line is present on the LCBI date of the event or the day before. The infection cannot be related to any other infection the patient might have and must not have been present or incubating when the patient was admitted to the facility. CLABSI rate is the total number of CLABSI divided by the total number of device days 1000.11 12

The differential time to positivity (DTP) is defined as a difference in time to positivity of ≥2 hours between peripheral blood culture and a CVC blood culture (peripheral DTP) or between two CVC blood cultures from different lumens of a multilumen catheter (CVC DTP).13 Due to its limitation reported in the literature, our unit does not prefer to use the DTP for the diagnosis of CLABSI.14

The authors designed an electronic system-based data collection sheet to collect all catheter-related parameters in the two groups.

**Statistical analysis**

Descriptive statistics were used to summarise and determine the sample characteristics and distribution of participants’ data. The normally distributed data and results were reported with mean and SD; the remaining results were reported with median and IQR. Categorical data were summarised using frequencies and proportions. Associations between two or more qualitative data variables were assessed using $\chi^2$ test or Fisher’s exact test as appropriate. Quantitative data between the two independent groups (USG-CVC and PICC) were analysed using unpaired $t$-test (for normally distributed data) or Mann-Whitney U test (for skewed or non-normally distributed data) as appropriate. Univariate and multivariate logistic regression analysis was applied to determine and assess the potential factors and predictors associated with the catheter insertion success rate adjusted for potential factors and predictors such as catheter types, gestational age, birth weight, the reason for catheter insertion, side of the body, site of insertion and number of pricks. For multivariate logistic regression models, predictor variables were included considering both statistical and clinical significance. The results of logistic regression analysis were presented as ORs with corresponding 95% CIs. Thereafter, we used the receiver operating characteristic curve (ROC) to evaluate the discriminative ability (predictive accuracy of the developed logistic regression model) of potentially significant variables associated with catheter insertion success rate. Box plots were
constructed depicting the distribution of gestational age and birth weight across two catheter types. All p values presented were two-tailed, and p values ≤ 0.05 were considered statistically significant. All statistical analyses were performed using statistical packages SPSS V.27.0 (IBM Corp) and Epi Info (CDC) software.

### RESULTS
Among the 3 years that this study covered, the usage of USG-CVC has progressively declined to zero in 2018, on which the catheter insertion success rate increased to 97%. Shown in table 1 are the distribution of patients and catheter-related variables associated with the types
of catheters. When USG-CVC was compared with PICC about gestational age, the former was significantly higher (33.88±6.34 vs 29.32±4.03, p=0.0001). Birth weight was also significantly higher among USG-CVC compared with PICC (2161.25±1140.26 vs 1234.57±624.90, respectively, p=0.0001). Figure 3 shows the distribution of gestational age (weeks) and birth weight (g) across two catheter types. The duration of catheter insertion was however not significant (USG-CVC 11.69±9.23, PICC 14.57±12.56, p=0.14). Further comparisons between USG-CVC and PICC on several parameters. PICC had a higher success rate (90% vs 88.4%), however, the difference did not reach statistical significance.

We performed univariate and multivariate logistic regression analysis testing for potential factors and predictors and their possible association with catheter insertion success rates are presented in tables 3 and 4. Univariate results indicated that year of catheter insertion, birth weight and the number of pricks had a significant effect on the likelihood of catheter insertion success rates. In patients who had 2 pricks (unadjusted OR 0.03; 95% CI 0.01 to 0.07, p=0.028) and ≥3 pricks (unadjusted OR 0.01; 95% CI 0.01 to 0.03, p=0.013) were significantly associated with a decreased likelihood of catheter insertion success rates compared with patients who had 1 prick. In addition, it was noted that catheter type PICC was associated with a higher rate of catheter insertion success rates, however, this difference was statistically insignificant (p=0.679).

The multivariate logistic regression analysis showed that duration of gestation (weeks) and the number of pricks were remained significantly (p<0.05) associated with the catheter insertion success rate after controlling and adjusting potential factors and predictors as shown in table 4. The higher catheter insertion success rates were associated with increasing gestational age (adjusted OR 1.23; 95% CI 1.03 to 1.44, p=0.015). Whereas, in patients who had 2 pricks (adjusted OR 0.07; 95% CI 0.0 to 0.57, p=0.014) and ≥3 pricks (adjusted OR 0.02; 95% CI 0.01 to 0.13, p=0.001) were significantly associated with a reduction in the likelihood of catheter insertion success rates when compared with patients who had 1 prick. Thereafter, we computed a prediction model to evaluate the discriminative ability of potentially significant predictors (observed in the developed multivariate logistic regression model) associated with catheter insertion success rates using ROC curve analysis. The value of area under the curve observed was found to be 0.841 (95% CI 0.81 to 0.87), which is indicating that this developed regression model demonstrated an excellent fit, figure 5.
DISCUSSION

Both PICCs and non-tunneled USG-CVCs have risks associated with their usage. Immediate risks include injury to local structures, accidental arterial puncture, phlebitis at the insertion site, air embolism, haematoma, arrhythmia and catheter damage and malposition. Late complications include infection, occlusion, thrombosis, infiltration, extravasation and catheter migration.16–18 Infection, thrombosis, embolisation, hydrocephalus, are complications reported in premature babies receiving central venous lines.3 The current study compared PICC to USG-CVC in a sample of cases from Qatar. The results also showed a progressive reduction in the usage of USG-CVC across the 3 years until reached 0% in 2018. This is due to the implementation of a PICC insertion team in early 2017 with a progressive build-up of the team skills.19 Since then, overall success and first prick rates have significantly increased. Reports of an overall success rate of 94% were indicated elsewhere.20 A systematic review highlighted the importance and necessity of a vascular access team in the NICU, as it reflects positively on the rate of BSI.21

Table 3  Univariate logistic regression analyses with potential significant factors and predictors associated with catheter insertion success rates

| Variables                        | Catheter insertion success rate, n (%) | Unadjusted OR | 95% CI for OR | P value |
|----------------------------------|--------------------------------------|---------------|---------------|---------|
| Catheter types                   |                                      |               |               |         |
| USG-CVC                          | 61 (88.4)                            | 1.0 (reference) | 0.55 to 2.51  | 0.679   |
| PICC                             | 1137 (90)                            | 1.17          |               |         |
| Year                             |                                      |               |               |         |
| 2016                             | 309 (81.7)                           | 2.0 (reference) | 1.18 to 2.52  | 0.004   |
| 2017                             | 450 (88.6)                           | 1.73          | 4.64 to 17.12 | 0.001   |
| 2018                             | 439 (97.6)                           | 8.91          |               |         |
| Gestational age (week)           | 29.56±4.20 vs 29.53±5.12              | 1.01          | 0.96 to 1.04  | 0.954   |
| Birth weight (g)                 | 1270.1±677.5 vs 1394.2±803.3          | 0.98          | 0.98 to 0.99  | 0.045   |
| Reason for catheter insertion    |                                      |               |               |         |
| Long-term intravenous fluid therapy| 1156 (89.9)                         | 2.0 (reference) | 0.48 to 1.51  | 0.453   |
| Others*                          | 42 (93.3)                            | 1.57          |               |         |
| Side of the body                 |                                      |               |               |         |
| Left                             | 491 (98.6)                           | 2.0 (reference) | 0.41 to 3.03  | 0.826   |
| Right                            | 706 (98.7)                           | 1.12          |               |         |
| Site of insertion                |                                      |               |               |         |
| Upper extremities                | 353 (98.1)                           | 2.0 (reference) | 0.43 to 2.57  | 0.920   |
| Lower extremities                | 845 (98.1)                           | 1.05          |               |         |
| Number of pricks                 |                                      |               |               |         |
| 1 prick                          | 833 (99.3)                           | 1.0 (reference) | 0.01 to 0.07  | 0.028   |
| 2 pricks                         | 244 (79.5)                           | 0.03          | 0.01 to 0.03  | 0.013   |
| ≥3 pricks                        | 121 (63.7)                           | 0.01          |               |         |

*Others includes: Difficult intravenous insertion, hypoglycaemia and long-term intravenous medication therapy. PICC, peripherally inserted central catheter; USG-CVC, ultrasound-guided central venous catheter.

Table 4  Multivariate logistic regression analyses with potential significant factors and predictors associated with catheter insertion success rates

| Variables                        | Catheter insertion success rate N (%) | Adjusted OR | 95% CI for OR | P value |
|----------------------------------|--------------------------------------|-------------|---------------|---------|
| Gestational age (week)           | 29.56±4.20 vs 29.53±5.12              | 1.23        | 1.03 to 1.44  | 0.015   |
| Number of pricks                 |                                      |             |               |         |
| 1 prick                          | 833 (99.3)                           | 1.0 (reference) | 0.01 to 0.57  | 0.014   |
| 2 pricks                         | 244 (79.5)                           | 0.07        | 0.01 to 0.13  | 0.001   |
| ≥3 pricks                        | 121 (63.7)                           | 0.02        |               |         |
with femoral central venous catheterisation, the overall thrombosis might also lead to catheter leaking. Approximation or undesirable catheter-tip location. CLABSI and from mechanical or postural factors, catheter malpositioning in USG-CVCs might be due to occlusions resulting in infections. The authors concluded by recommending the usage of PICCs routinely when dealing with neonatal surgical procedures. In another study which is close to our PICC data.25 The higher rate of catheter leaking in USG-CVCs might be due to occlusions resulting from mechanical or postural factors, catheter malpositioning, or undesirable catheter-tip location. CLABSI and thrombosis might also lead to catheter leaking.24 Approximately one-third of PICCs were associated with complications in another study which is close to our PICC data.25 Ragavan et al described the advantages of using PICCs inserted in the cubital veins as to have a reduced complication incidence rate, as well as maintenance rates in comparison to USG-CVCs inserted in the internal jugular vein. The authors concluded by recommending the usage of PICCs routinely when dealing with neonatal surgical patients.26 On the other hand, a recent study reported a 100% success rate of 30 preterm babies who underwent an USG brachiocephalic CVC insertion. No case of accidental arterial or pleural puncture was noted by the researchers.27 In another study involving neonates with femoral central venous catheterisation, the overall success rate was 100% of neonates (n=82/82), first attempt 63/74 (85%), second attempt 8/74 (11%) and third attempt 3/74 (4%). Another two studies reported no statistical difference in the complication rate or efficacy between those who had PICC and those who had USG-CVC.28

The limitation of this study is being retrospective with potential risks of bias and confounding factors especially when single-centre studies. The imbalance in numbers between the two groups suggest that the inferences may not be robust. Another limitation of the study is that the PICC team was properly trained to insert PICCs while the USG-CVC were placed by operators not belonging to the team (surgeons or anaesthetists). Potential bias by indication might be an issue as percutaneous CVCs were considered if some attempts for a PICC insertion failed. As reported by other researchers, USG-CVCs sometimes needed multiple pricks to get the catheter successfully inserted as reported in our study. This might be related to the level of experience, the number of exposures and lack of training as this task is not the main task daily performed by the operators (surgeons and anaesthetists). Besides, being inserted as rescue mode, not for selected patients is a stressor that might be a factor in increasing the number of pricks.

No USG-CVC was inserted in our unit for the last 2 years, however, it might be needed in the future in certain indications. Randomised controlled trials (RCTs) to study the feasibility of intracavitary ECG in catheter insertion and tip location in neonates are strongly recommended. Also, the use of US guidance during PIVC insertion and the frequency of its use in tip location monitoring of correctly positioned central lines to confirm the tip positions and diagnose catheter migration are both rich areas for future prospective studies.

CONCLUSION

The overall complication rate, CLABSI and leaking are significantly higher in non-tunnelled USG-CVCs compared with the PICCs. However, RCTs with larger sample sizes are desired. Proper central venous device selection and timing, early PICC insertion and early removal approach, dedicated vascular access team development, proper central venous line maintenance, central line simulation workshops and US-guided insertions are crucial elements for patient safety in NICU.

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Contributors MAAB is the principal author responsible for the overall content as guarantor. He accepts full responsibility for the finished work, the conduct of the study, had access to the data, and controlled the decision to publish. MAAB and DS conceptualised and designed the study, RV and MAAB collected, cleaned and anonymised the data. PC designed and performed the data analysis. MAAB, PC and SH drafted the initial manuscript. MAAB and PC designed the figures. EEE, MAAB,
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