Expressed breast milk on a neonatal unit: A hazard analysis and critical control points approach

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With the increasing use of human milk and growing evidence of the benefits of mother’s milk for preterm and ill newborns, guidelines to ensure its quality and safety are an important part of daily practice in neonatal intensive care units. Operating procedures based on hazard analysis and critical control points can standardize the handling of mother’s expressed milk, thereby improving nutrition and minimizing the risk of breast milk–induced infection in susceptible newborns. Because breast milk is not sterile, microorganisms can multiply when the milk is not handled properly. Additional exogenous contamination should be prevented. Strict hygiene and careful temperature and time control are important during the expression, collection, transport, storage, and feeding of maternal milk. In contrast to formula milk, no legal standards exist for the use of expressed maternal milk. The need for additional measures, such as bacteriological screening or heat treatment, remains unresolved.

Key Words: NICU; infection control; mother’s milk; quality control.

Serious Enterobacter sakazakii infections reported in formula-fed preterm and term infants have led to the promulgation of guidelines for the preparation, storage, and handling of powdered infant formula in hospitals following the basic risk control principles of hazard analysis and critical control points (HACCP) to reduce the risk of infection.1,2 Legal criteria have been developed for the use of powdered formula milk for infants under 6 months of age.3 The implementation of these guidelines is mandatory and regulated by national food agencies.

Nonetheless, breast milk remains the preferred nutrition for all infants, including preterm and ill newborns, with rare exceptions.4 The beneficial effects of breast milk are related to improvements in host defense, absorption of specific nutrients, enteral tolerance, and neurodevelopmental outcome. Breast-feeding also enhances maternal involvement and mother–infant interaction and bonding, which can be compromised in the neonatal intensive care unit (NICU) setting. Efforts are underway worldwide to encourage and support mothers’ milk feeding in the NICU. If direct breast-feeding is not possible, then expressed breast milk can be fed by nasogastric tube.

Mother’s milk is not sterile and can be a vehicle for commensal and pathogenic microorganisms derived from the mother or the NICU environment.5,6 Human milk has the potential to carry infectious agents, occasionally causing late-onset sepsis, especially in preterm and very low birth weight infants. Transmission through breast milk of such pathogens as Staphylococcus aureus, group B streptococci, Eschericia coli, Pseudomonas spp, Klebsiella spp, Serratia spp, Salmonella spp, and cytomegalovirus have been documented.7-17 Of greatest concern, and the most widely studied, is human immunodeficiency virus.18

Recommended good hygienic practices have been developed for the expression, collection, transport, storage, and optimal handling of human milk in a neonatal unit. These recommendations are intended to ensure that human milk is microbiologically safe and nutritionally and immunologically complete. In contrast to formula milk, no microbiological quality standards or specific state regulations are available for expressed breast milk used exclusively for a mother’s own infant in a hospital.

The aim of the present study was to apply the principles of HACCP to standardize the handling of expressed breast milk at every stage and ensure the
milk’s quality and safety. Health care professionals and other caregivers need to pay attention to the risk for environmental contamination and to the cold chain management of breast milk.

METHODS

To apply the HACCP principles to the use of expressed breast milk in the hospital, a multidisciplinary team was formed consisting of an infection control doctor and nurse, a neonatologist, nurses from the neonatal unit, and ancillary and lactation staff members. The NICU is part of the Leuven University Hospital, which has earned the Baby-Friendly Hospital Initiative designation. Breast-feeding is promoted for the >850 infants admitted annually to the NICU.

The HACCP process

HACCP is a systematic preventive approach to identify, evaluate, and control significant hazards to food safety. The HACCP approach is aimed at unsafe practices during the entire processing and storage process and as such differs from traditional final product testing, such as bacteriological sampling. All steps are designed to limit the contamination of expressed breast milk by microorganisms and to prevent multiplication of organisms.

The HACCP process involves 4 steps. In step 1, the scope of the HACCP is described, from the expression of milk to the feeding of infants in the NICU. Step 2 involves identifying potential hazards for each process step. In step 3, control measures to eliminate or reduce each of the identified hazards to an acceptable level are defined. For each critical control point (CCP) considered to have a significant effect on overall hazard of the procedure, a critical limit is established based on current evidence. Finally, in step 4 a monitoring system for each CCP is established, and corrective actions for deviations from critical limits are devised. Documentation includes a written HACCP plan, CCP monitoring, and adjustments for deviations.

Identification of CCPs

Most data are adapted from the existing standards for powdered formula and guidelines for the operation of donor human milk banks. A literature search of the PubMed database was conducted. Searches of published research and policy documents were carried out, and references from selected articles and from reviews were screened for additional relevant articles.

RESULTS

Figure 1 presents a flow diagram showing the human milk processing and storage steps and equipment. Table 1 summarizes the HACCP plan, including measurable indicators for use in monitoring the most important aspects of human milk expression, transport, delivery, storage, and administration. For all steps in the process, potential hazards, control measures, CCPs, and possible adjustments based on the level of concern are listed. At some NICUs, it has become routine to pasteurize mothers’ own milk for feeding to susceptible infants. The most common procedure used is Holder pasteurization, in which the milk is heated to 62.5°C for 30 minutes. An HACCP-based plan indicating the potential hazards and CCPs during the pasteurization process is available from the authors on request.

DISCUSSION

Interest in feeding human milk to preterm infants has increased during the past decade. The benefits of using a mother’s own milk for her preterm infant outweigh any potential disadvantages. However, the use of expressed mother’s milk mandates strict control of storage and handling in accordance with hygienic practices, given milk’s potential as a vector for pathogens, especially when fed to immunologically vulnerable infants at high ambient temperatures.

When applying the HACCP principles to the management of expressed breast milk in a NICU, all steps of the process must be carefully analyzed. Because human milk is not sterile, each step in the collection and storage process can affect the final product. The identification of potential hazards implies considering the infant’s susceptibility to infection and estimating the minimal infectious dose. CCPs should be defined based on clinical experience and limited evidence. If effective control options can be devised by implementation of
infection control policies, then corrective actions can be identified if deviations occur.

**Hazard identification**

Does bacterial contamination of expressed breast milk really matter, and what level of contamination are we trying to avoid? Microbiological testing of expressed milk identifies bacterial growth in >75% of samples, with 7%-36% of all samples containing pathogens. Some believe that the inherent anti-infective properties of the human milk are sufficiently strong to protect any infant under any conditions, including preterm infants in NICUs, and assume the best-case scenario when developing recommendations. However, the belief that expressed milk is safe even when containing pathogens is challenged by reports linking outbreaks and case reports of sepsis or necrotizing enterocolitis in NICUs to ingestion of contaminated human milk. This link is considered plausible, but clear evidence of causality is rarely based on molecular typing as proof of transmission.

**Hazard characteristics**

It is not clear what effect different organisms and different levels of contamination of expressed milk have on recipient infants, and whether these effects differ according to the age or immune status of the consumer. Very little is known about the minimal infectious dose and how this varies in susceptible populations. For *E. sakazakii*, even low-level contamination (<3 cfu/100 g) is considered a risk factor given the potential for multiplication during the preparation and holding time before consumption. The risk of sepsis or necrotizing enterocolitis after ingestion of milk contaminated with even small inocula of microorganisms or microbial toxins is considered the greatest in very low birth weight neonates (<1,500 g) and those born at gestational age <32 weeks, who have limited transplacental transfer of antibodies and an immature intestinal mucosal immunity.

What is the risk that a feed that nearly always contains at least some organisms will lead to infection in the preterm neonate? During their first 2 weeks of feeding, neonates weighing <2,000 g showed surprisingly few adverse events of bacteremia and feeding intolerance that could be directly related to ingestion of bacteria in raw milk, with $10^{4.8} \pm 1.1$ cfu of gram-negative bacilli ingested per feed. Other investigators found a dose-response relationship, with feeding intolerance correlated with recovery of $\geq 10^5$ cfu of gram-negative bacilli per mL of expressed milk and septic episodes correlated with recovery of $\geq 10^6$ cfu/mL. Because regular opening of the refrigerator in the NICU does not guarantee a stable temperature at 4°C, storage times exceeding 48 hours are not considered bacteriologically safe for use in the NICU. Human milk loses its bactericidal capacity when refrigerated for >72 hours.

Additional risk reduction strategies: To screen or to heat?

By implementing the principles of HACCP listed in Table 1 to the daily practice of using expressed breast milk, the level of contamination can be reduced to acceptable standards and the growth of microorganisms before consumption can be minimized. Some CCPs are not yet completely defined and thus are not controllable, including a limit for total bacterial colony count per mL of milk and standards for the presence of skin commensals or specific pathogens. Although evidence-based steps and preventive measures cannot be applied, to further neutralize any microbial contamination, additional steps, such as microbiological testing and heat treatment, can be considered. These practices remain controversial issues in NICUs worldwide.

Mother’s own milk cannot be subjected to the stringent microbiological standards applied to banked donor milk. Donated milk fed to a biologically unrelated...
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<th>Steps in the process</th>
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<tr>
<td>Milk expression and collection</td>
<td>• Hands may touch breast and milk during pumping. Breasts or nipples may be colonized or infected.</td>
<td>• Teach mothers to perform careful hygiene of hands before expressing or handling milk, as well as daily hygiene of breasts.</td>
<td>• No compliance with hygienic advice.</td>
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<td>• Supply written and verbal instructions on personal hygiene and expression of milk, storage and transport.</td>
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<td>• Pump may be contaminated with pathogens (exterior and interior; backflow of aerosol of milk).</td>
<td>• Use a correctly designed type of pump with separated internal circuits and a safety valve. Perform regular pump cleaning and maintenance.</td>
<td>• Visibly not clean.</td>
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<td>• Accessory kits may be contaminated.</td>
<td>• Thermal disinfection of shields and other parts in contact with milk after each use. Use clean disposable or sterile bottles or containers.</td>
<td>• Visibly not clean.</td>
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<td>• Milk not placed in cool storage after expression.</td>
<td>• Educate parents to refrigerate or freeze the milk within 1 hour.</td>
<td>(Partially) thawed milk at arrival (visual inspection)</td>
<td></td>
<td>Do not accept and discard thawed milk. Repeat instructions to parents regarding transport and storage.</td>
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<td>Milk transfer to the unit</td>
<td>• Growth of microorganisms if a break in cold chain occurs.</td>
<td>• Provide information regarding appropriate storage conditions for transporting milk in an icebox or isothermal bag.</td>
<td>• Visibly not clean.</td>
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<td>Repeat parental education.</td>
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<td>• Contamination of bottles.</td>
<td>• Educate parents about general hygiene and the use of a clean transfer box.</td>
<td>• Missing labels.</td>
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<td>Add missing data on arrival.</td>
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<td>• No or poor identification on arrival at the unit.</td>
<td>• Check name, date and time of expression on each bottle.</td>
<td>• Storage &gt;48 hours if fresh milk; &gt;24 hours if thawed milk; &gt;3 months if frozen milk.</td>
<td></td>
<td>Discard milk if critical limit is exceeded.</td>
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<tr>
<td>Storage in the unit</td>
<td>• Exceeding storage time and risk for contamination.</td>
<td>• Use fresh milk within 48 hours. Freeze milk that will not be used within 48 hours. Use thawed milk within 24 hours. Use frozen milk within 3 months.</td>
<td>• Check weekly.</td>
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<td>• First-in, first-out principle may not be followed.</td>
<td>• Place newly delivered milk at the back of the drawer in the freezer. Label containers clearly with waterproof ink.</td>
<td>• Each deviation from chronologic rank.</td>
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<td>• Temperature of refrigerator is too high.</td>
<td>• Keep doors closed. Monitor core temperature continuously with central alarm connected to the hospital building management system. Calibrate the logger system regularly. Control environmental temperature. Place refrigerator in cool room.</td>
<td>Core temperature &gt; 5°C.</td>
<td></td>
<td>Discard milk if critical limit is exceeded. Use second refrigerator in case of defect. Analyze and document any temperature excess.</td>
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<td><strong>Temperature of freezer is too high.</strong></td>
<td>Keep doors closed. Monitor temperature continuously with central alarm connected to the hospital building management system. Clean and defrost periodically and whenever visually contaminated.</td>
<td>&gt;1 cm of ice or visibly not clean. Temperature &gt; -15°C.</td>
<td>(Partially) thawed milk.</td>
<td>Discard milk if partially defrosted. Analyze and document any temperature excess.</td>
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<td><strong>Other products or dirt in freezer or refrigerator may contaminate the milk.</strong></td>
<td>Keep a freezer and refrigerator in a secure room exclusively dedicated to milk. Clean daily.</td>
<td>Visibly not clean.</td>
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<td><strong>Exogenous contamination when dividing milk in aliquots.</strong></td>
<td>Clean surface and clean hands. Wear mask and cap. Do not handle formula milk when preparing maternal milk. Prepare individual feedings. Aliquot milk into sterile capped syringe or container and store in refrigerator before use.</td>
<td>No compliance with general hygiene precautions. Caps and masks not available. Task is interrupted.</td>
<td>Simultaneous handling of mother’s milk and formula milk.</td>
<td>Educate personnel. Organize divided preparations for mother’s milk and formula milk.</td>
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<td><strong>Bacterial growth at room temperature or during warming.</strong></td>
<td>Thaw frozen milk in refrigerator. Limit warming up time to 30 min. Discard any milk from a partially consumed feeding, do not refrost or reheat.</td>
<td>Time between start of warming the milk and end of administration &gt;1 hour.</td>
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<td>Do not use milk if critical limit is exceeded. Educate health care providers. Document temperature logging of bottle warmers.</td>
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<td><strong>External contamination of bottles during warming.</strong></td>
<td>Use dry heating systems. Clean bottle warmer daily and when visually contaminated.</td>
<td>Visibly not clean.</td>
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<td><strong>Improper use of additives (eg, fortifiers).</strong></td>
<td>Use capsules for individual use made under aseptic conditions and kept in dust-free container. Use fortified milk ideally immediately or within 24 hours if refrigerated.</td>
<td>&gt; Expiration date of fortifier. &gt; 24 hours for fortified milk.</td>
<td></td>
<td>Check expiration date. Discard milk with additives after 24 hours.</td>
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<td><strong>Expressed milk may be given to the wrong infant.</strong></td>
<td>Milk must be clearly labeled. Before feeding milk label and infant's ID should be double-checked.</td>
<td>Milk given to the wrong infant.</td>
<td></td>
<td>Post-exposure protocol must be available in case of improper use.</td>
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<td><strong>Growth of microorganisms in expressed milk given as continuous tube feeding.</strong></td>
<td>Change syringes or bottles every 4 hours and tubing every 24 hours.</td>
<td>&gt; 4 hours hangtime</td>
<td></td>
<td>Discard milk if critical limit is exceeded.</td>
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infant is either pasteurized before consumption or discarded if it contains > 10^6 cfu/mL of total viable microorganisms or > 10^4 cfu/mL of Enterobacteriaceae or S. aureus. There are no universally accepted upper safe limits for bacterial concentrations in expressed breast milk given to the mother’s own infant, and interpreting the results of milk cultures is difficult in the absence of appropriate standards. Testing milk can be useful when it is a suspected source of neonatal sepsis or feeding intolerance. Screening also ensures that the instructions for collection and transportation have been given, understood, and followed.

Those who believe that the number or type of bacterial contaminants ingested with milk can be assumed to affect the incidence and severity of illness, especially in preterm infants, should not attempt to define safe limits, but rather should consider not feeding raw expressed milk to high-risk infants. However, pasteurizing a mother’s own milk for her infant will partly destroy the milk’s unique immunologic protection.28,42 The process of heat treatment also leads to delays before consumption and extra costs.

In conclusion, the use of human milk in the NICU is not formally regulated under specific legislation as is formula. Thus, NICUs must be committed to the highest standards of self-regulation that are practically and scientifically warranted. The HACCP approach is an essential part of operational planning during milk management. If those caring for preterm infants wish to increase the proportion who receive the mother’s own expressed breast milk, there needs to be agreement regarding the need for bacteriologic screening of milk and the appropriate response to positive cultures or to the need for pasteurization.

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