Racism, segregation, and inequality contribute to health outcomes and drive health disparities across the life course, including for newborn infants and their families. In this review, we address their effects on the health and well-being of newborn infants and their families with a focus on preterm birth. We discuss three causal pathways: increased risk; lower-quality care; and socioeconomic disadvantages that persist into infancy, childhood, and beyond. For each pathway, we propose specific interventions and research priorities that may remedy the adverse effects of racism, segregation, and inequality. Infants and their families will not realize the full benefit of advances in perinatal and neonatal care until we, collectively, accept our responsibility for addressing the range of determinants that shape long-term outcomes.

Pediatric Research (2020) 87:227-234; https://doi.org/10.1038/s41390-019-0513-6

INTRODUCTION

In The Color of Law, Richard Rothstein documents how, from Reconstruction to present day, local, state, and federal policies, regulations, and laws have been used to segregate Americans by race resulting in dramatic, persistent inequalities in social, economic, and educational opportunities.1 Black and other minority Americans live in poorer neighborhoods,2,3 attend lower-quality schools,4,5 and receive health care at lower-quality hospitals.6,7-10

Racism, segregation, and inequality contribute to disparities in health outcomes across the life course.11 In this review, we address their effects on the health and well-being of newborn infants and their families with a focus on preterm birth. We explore three causal pathways that adversely and differentially affect outcomes for newborn infants and their families: increased risk; lower-quality care; and socioeconomic disadvantages that persist into infancy, childhood, and beyond. We propose interventions that health professionals, multidisciplinary care teams, and health-care organizations can adopt to remedy the adverse effects of racism, segregation, and inequality (Fig. 1) and research priorities to inform more effective action.

An underlying theme is that we as health professionals have the responsibility to act. The old paradigm of “follow-up” must be replaced with a new paradigm of “follow through,” the understanding that our responsibility extends beyond the hospital walls. All infants and their families will not realize the full benefit of the dramatic technical advances in perinatal and neonatal care that will occur in the 21st century unless we accept our responsibility for addressing the full range of determinants of health that ultimately shape long-term outcomes.

INCREASED RISK

In 2018, 10.0% of infants in the United States (U.S.) were born before 37 completed weeks’ gestation,12 often requiring care in a neonatal intensive care unit (NICU).13 Overall, 9.1% of white non-Hispanic infants were born preterm compared to 14.1% of Black non-Hispanic infants and 9.7% of Hispanic infants.12 Structural racism describes mutually enforcing forms of discrimination14 such as neighborhood deprivation,15-16 economic inequalities,16,17 educational disparities,16,18-20 and differential access to health care21 with sequelae like nutritional deficiencies22 and unhealthy environmental exposures.23 These factors are the social determinants of health (SDH), defined as the "conditions in which people are born, grow, work, live, and age."24 Consistently, the SDH increase the risk for preterm birth and infant mortality.25 However, exposure to the SDH alone may be an insufficient explanation for increased risk for preterm birth.26 The early programming life course theory suggests that exposures during the perinatal period, such as maternal stress27,28 and depression,29 increase vulnerability to preterm birth. The cumulative pathway life course theory posits that additive stressors and exposures...
resulting in allostatic load and inflammation increase the risk for preterm birth.\textsuperscript{30–33} The weathering hypothesis, in which chronic stress leads to accelerated aging and earlier onset of adverse health conditions, may also be a risk factor in preterm birth.\textsuperscript{34} While the contribution of genetics alone likely plays a small role in differential risk for preterm birth,\textsuperscript{35} gene–environment interactions may explain differences.\textsuperscript{36–39} There is evidence of an association between DNA methylation and preterm birth among black women.\textsuperscript{40} Risk for preterm birth may be heritable among African-American women.\textsuperscript{41–43} Immigrant African women are less likely to give birth preterm than native-born black women.\textsuperscript{44,45} All of these findings illustrate the potential role of epigenetics in preterm birth. However, genetic and epigenetic studies must include people of diverse ancestry.\textsuperscript{46}

Numerous experts have suggested ways to reduce differences in preterm birth rates.\textsuperscript{47–50} Improving and adherence to 17-alpha hydroxyprogesterone caproate after a previous preterm birth,\textsuperscript{51} identifying and monitoring cervical insufficiency,\textsuperscript{28} cerclage placement in high-risk women,\textsuperscript{52} screening and treatment of specific genital infections,\textsuperscript{53,54} and increasing interpregnancy intervals\textsuperscript{55} will help. Research with standardized definitions of race, ethnicity, and discrimination that seeks to disentangle the contributions of structural racism and its sequelae may inform interventions but will be difficult to conduct. Instead, research into interventions to improve access to health care, financial, and social supports for families and children, to reduce stress, and to identify and treat infections, inflammation, mental health issues, and nutritional deficiencies may have greater short-term impacts. Regardless, while broader structural issues are evaluated and being addressed, there is a clear argument for ensuring that women and their families receive the highest quality care.
LOWER-QUALITY CARE

Substantial improvements in perinatal and neonatal care have led to decreasing rates of death before discharge and serious morbidities in NICUs across the U.S. Yet wide variation in mortality and serious morbidities persists.63,65 Black neonates die at more than twice the rate of non-Hispanic white neonates, and black and Hispanic infants remain at increased risk for severe neonatal morbidities.66–68 Further, the magnitude of these disparities is larger than previously reported.65 Disparities in severe neonatal morbidities are associated with later neurodevelopmental, behavioral, and physical impairments that disadvantage very preterm infants across the life course, perpetuating health and socioeconomic disparities.69,70

Quality of care is a crucial and potentially modifiable factor that contributes to disparities and is one of the causal pathways by which racism, segregation, and inequality adversely affect outcomes of very preterm infants. Differences in quality of care contribute to racial and ethnic disparities in two ways.63 First, white and minority preterm infants may receive care in different NICUs, and NICUs primarily serving minority infants may have structural characteristics associated with lower quality or have organizational models or clinical processes that lead to lower-quality care.63 Second, the quality of care received by preterm infants can differ by race and ethnicity in individual NICUs. These pathways may be related to organizational (e.g., culture) and clinical processes (e.g., communication) rather than structural characteristics and can confer disadvantages to minority infants. There is accumulating evidence that both mechanisms are at work.

Evidence over the past two decades has documented that black very low birth weight (VLBW) infants are cared for in a concentrated set of hospitals64 and that minority-serving hospitals have higher neonatal mortality rates for both black and white infants.65,66 More recent research has expanded beyond mortality to include severe neonatal morbidities with similar findings. Black and Hispanic very preterm infants are more likely to be born in hospitals with higher risk-adjusted neonatal morbidity and mortality rates than white VLBW infants, and disparate utilization patterns can explain 40% of the black–white disparity and nearly one third of the Hispanic–white disparity in very preterm neonatal morbidity and mortality rates.79 These findings have prompted investigators to evaluate the origins of outcome differences between hospitals and the potential role of differential quality of care. Investigators also found that minority infants were less likely to be born at hospitals that achieved Magnet status and infants at non-Magnet hospitals had significantly higher rates of morbidity and mortality.67 Similarly, VLBW infants born in hospitals that served higher proportions of minority patients had higher rates of infection, discharge without breast milk, nurse understaffing, and worse practice environments for nurses.68 The patient-to-nurse ratios and missed care in minority-serving hospitals were much higher than in other hospitals suggesting that improving staffing and workloads would improve quality of care at minority-serving hospitals.68 Recent investigations utilizing national data found significant segregation across NICUs and that black infants received care at lower-quality NICUs.68,69 These studies suggest that improving care provided at lower-quality hospitals would result in a significant narrowing of disparities.69

While previous studies primarily investigated between-hospital disparities, additional research has examined within-hospital disparities. Using a composite measure of quality (including both process and outcomes measures), investigators found that significant racial and ethnic variation in quality of care also existed within NICUs.70 Studies have highlighted within-hospital differences in the use of evidence-based practices. Hispanic mothers were found to be less likely than white mothers to receive antenatal steroids and to be feeding human milk at discharge within a given institution.71,72 Additional research suggests that adverse interactions with families may be a mechanism for disparate care, interactions that may be driven by clinician implicit bias and other failures in staff–family communication. These factors can affect the health of the neonate and have the potential to incite disparities that will last beyond discharge.73 While studies have examined the role of racism in preterm birth, few studies have examined the role of racism and bias in the NICU setting.74 A growing body of research in maternal health acknowledges the role that structural racism plays in generating disparities.74 More research on how behaviors and attitudes impact clinical outcomes in the NICU setting is needed.

There is an urgent need to go beyond describing disparities to understand their causes, and develop and implement interventions to eliminate their occurrence. An abundance of evidence demonstrates that the quality of care is one of the causal pathways by which segregation and inequality adversely affect outcomes of very preterm infants. Lessons from efforts in maternal health to reduce peripartum disparities can be applied to the NICU setting to help us further the shift from “follow-up” to “follow through.”74,75 Critical steps to such a shift include: enhancing communication with families by educating clinicians and staff about racial and ethnic disparities in neonatal outcomes, the importance of shared decision making, cultural competency, and assessing non-English language proficiency and providing interpreter services; education on implicit bias that can affect a clinician’s perceptions and decisions; creating disparities dashboards that stratify quality measures by race/ethnicity, insurance, language, and other attributes so that NICUs can assess how well they are taking care of infants from different backgrounds; applying quality improvement tools to narrow the disparities that are identified; engaging families and communities in quality and safety teams; and promoting a culture of equity by employing many of the tools we have used to emphasize a culture of safety. These strategies and steps must also consider, identify, and address socioeconomic disadvantages that will accompany an infant home as they are discharged from the NICU.

SOCIOECONOMIC DISADVANTAGES IN INFANCY AND CHILDHOOD

Infants of women of color begin life with greater risks of vulnerability through their well-documented increased risk for being delivered preterm and experiencing greater morbidity and mortality. In part, this disparity reflects higher parallel rates of socioeconomic disadvantage among women of color. Such disadvantage both increases the risk of preterm birth and of subsequent poorer infant health and developmental outcomes.76,77

Characterizing factors that underlie racial and ethnic disparities in the health and well-being among premature infants can be difficult. While the study of long-term outcomes of preterm infants has progressed over the past quarter century, much of this work has focused on simple associations between prematurity, birth weight, and NICU complications (exposures) and measures of neurodevelopment (outcomes), generally within the first few years. Because of the interest in attributing outcomes to “prematurity,” the research often employs restrictive study designs, rarely considers post-discharge factors influencing child health and development, and fails to consider variations in the quality of care on outcomes.78 More recent work has considered a broader range of outcomes including physical health, social and emotional well-being, and cognitive development. Studies have also examined the influence of neonatal care, illness severity, and NICU length of stay on outcomes. Nonetheless, the conceptualization of outcomes remains simplistic, failing to consider the joint effect of sociodemographic factors, chronic child illness, and maternal physical and mental health over time.

Of course, factors related to the NICU course can significantly influence physical health status in all preterm and low birth

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weight infants, but these infants remain vulnerable to the factors that worsen health in all children, like poverty and adverse childhood events. To the extent that infants of color are disproportionately exposed to these post-NICU stressors, they will continue to experience disparities in health outcomes of all types. Life course health development formulations suggest that the adverse effects of such stressors can be buffered by interventions that reduce the effect of stress and foster resiliency.80 To that effect, greater attention is now being paid to NICU-to-home transitions and to enhancing family caregiving skills via specific therapeutic and developmental interventions for the infant.

The American Academy of Pediatrics recommended guidelines for the discharge of high-risk newborns that focus on risk identification, the optimal timing of discharge, and planning for care after discharge.78 The guidelines emphasized discharge planning, highlighting objectives for parental education, routine health-care maintenance, management planning for unresolved medical problems, comprehensive home-care planning, identification and involvement of support services, and the determination and designation of follow-up care. These recommendations have led to greater emphasis placed on comprehensive discharge preparation programs to ensure a systematic and individualized approach for each family,86 including more consistent measurement, standardized discharge approaches, and the incorporation of quality improvement practices.86–89 As noted above, evidence suggests that some minority families experience lower-quality interactions with NICU staff, which might reduce effectiveness in discharge planning and family readiness.78 Deeper study of new and existing programs that support discharged preterm infants as they grow and age is also occurring.

In the years following NICU discharge, intervention programs that focus on family factors and the home environment are proving effective.83 Indeed, early intervention programs improve cognitive outcomes and, to a lesser extent, motor outcomes – even effective for very preterm infants.84,85 Cognitive improvements are seen to preschool age. Those programs that focus on both the parent–infant relationship and on infant development using standardized curricula have been shown to be the most effective; however, the longer-term benefits of these programs remain to be established. Despite the evidence of their effectiveness, research has documented poorer access to such services for minority infants.86

An important component of how longer-term outcomes are measured is to ensure adequate characterization of the functional impact of prematurity and its complications as well as the effects of interventions. Test scores and neurologic exams may fail to provide a complete picture of child well-being. Depending on a variety of circumstances, functioning may be influenced by comorbidities (e.g., asthma, obesity, sensory deficits), movement disorders, and behavioral (e.g., attention and autism disorders) factors. To the extent that children of color experience more of these morbidities,87 they are at greater risk for poor functional status.

The life course models mentioned above also suggest other targets for research and intervention. Most have incorporated advances in biological, behavioral, and social science disciplines to define child development as a dynamic process that begins before conception and continues through the life span.88 The incorporation of emerging investigations in gene–environment interactions and epigenetic mechanisms shows promise in maternal and child health by highlighting the importance of fetal development and early childhood on the life span. These models provide an opportunity to focus on the impact of a variety of risk and protective factors in early childhood for all children and foster a shift to the promotion of more effective prevention and intervention strategies to optimize healthy development. This, in turn, informs health measurement, health-care organization, and health systems financing.88 It also highlights ways in which health care can meaningfully connect and collaborate with the community.

COMMUNITY INTERVENTIONS

NICUs are increasingly turning toward community-based prevention strategies and partnerships to improve perinatal outcomes. Strategies often precede the birth of a child, support provision of high-value care during NICU stays, and optimize transitions as patients and families return to their medical and neighborhood homes after discharge.94,95,99 This pivot from "follow-up" to "follow through" is accelerated by patient-level social needs screening, the use of electronic health record (EHR) data in combination with complementary datasets to support population-level pattern recognition, and bolstered clinical–community partnerships driving both patient- and population-level action.93–96 Many such actions elevate the relevance of SDH, which, in many ways, are thought to be "fundamental causes" of disease, factors that place individuals and populations at "risk of [having] risks."97,98

The National Academy of Medicine and American Academy of Pediatrics, among others, now recommend routine SDH-related screening.99,100 They encourage routine assessment of factors like race/ethnicity, social isolation, parental educational attainment, financial strain, and residential address (to enable linkage to complementary information like neighborhood-level median household income).96,101–104 Evaluation of SDH-related risks and assets using one of the many available screening tools104 could influence clinical care from the moment of first patient/family contact, ensuring that supports are deployed to match identified needs. Assessments could start prenatally in obstetrical clinics and extend across the NICU stay to bring an awareness of context to the bedside. Indeed, families with a daily life that is characterized by racial discrimination, housing instability, food insecurity, and inflexible work schedules may have a far more difficult time adhering to complex medical regimens without additional, complementary supports.

Although screening is critical, a broader prevention mindset would benefit from strategies capable of discerning population-wide patterns in adverse health outcomes.105–107 EHR data can be leveraged to do just this, employed to inform data-driven action.98 For instance, EHR data can define numerators for key population health (or risk) measures. Pooling EHR data from networked birth hospitals, nurseries, NICUs, and health departments could enumerate preterm birth or infant mortality rates across geographies and in association with a range of potentially underlying sociodemographic factors. When mapped to certain neighborhoods or regions, geographic patterns could be identified. Where are “hot spots” of preterm birth and infant mortality? What are the characteristics of those areas with respect to racial segregation and socioeconomic disadvantages? By layering data elements atop one another, and by ensuring that findings are shared transparently with partners inside and outside health-care settings, new research questions and intervention strategies may be revealed.

Ultimately, the pivot from risk assessment and pattern recognition to action is what will ensure the shift from "follow-up" to "follow through." With this as the goal, we suggest that health-care systems, including academic health centers, consider their role in population health improvement and social justice.91,94 Such a re-envisioned mission, one that elevates the importance of community well-being, is likely to require the added, complementary expertise that comes from meaningful community integration.107,108 Indeed, although health-care systems and providers have expertise in managing medical complexity, they may not always have the expertise necessary to manage social complexity. This is where clinical–community partnerships become important, valuable adjacents. Henize et al. defined
how such clinical–community partnerships can be developed, enhanced, and sustained. They described the importance of jointly defining the problem, identifying program champions, agreeing upon mutually beneficial metrics, ensuring clear and consistent communication, and planning for sustainability.109

The Medical–Legal Partnership (MLP) model is an example of a successful clinical–community partnership, one where legal expertise is brought into health-care settings.110-112 Although MLPs look different from partnership to partnership and region to region, their focus on context is consistent. For example, consider a patient in the NICU requiring ongoing respiratory and nutritional support. The intensity of their medical needs would challenge any family; that challenge is sure to be magnified by co-existent social needs. Imagine if this child were discharged to a home filled with cockroaches, a building managed by a landlord who threatened eviction with any complaint. How might this social complexity affect a family’s ability to manage the child’s medical complexity? Although health-care providers may be able to advocate for this child’s well-being, a housing attorney would amplify their voices. Through an MLP, such legal experts can advocate for remediation of adverse housing exposures and guard against illegal evictions. They may also illuminate population-level patterns.113 Perhaps the family from the NICU may have similarly effected neighbors. There may be other children, or expectant mothers, in the unit next door or a building located across town with the same landlord. Partnerships like MLPs work best when they extend the reach of each partner, authentically engage key stakeholders, and consistently evaluate whether they are achieving the desired outcomes.114

Clinical–community partnerships can extend to broader community-based investment strategies considered by health-care systems. The Healthy Neighborhood, Healthy Families Initiative in Columbus, OH involves a hospital making direct housing interventions within a “blighted” neighborhood. A multi-disciplinary team identified a racially segregated neighborhood filled with vacant housing units and characterized by high rates of poverty that was home to many of the hospital’s patients and employees. The resulting initiative involved a series of interventions focused on enhancing housing stability and quality.115 Although a motivated, mission-driven anchor institution galvanized the program, it coincided with the hospital assuming financial risk for community children who were insured through Medicaid. With this reality, the team suggested that investments in communities and in clinical–community partnerships, are likely to be most successful when they have both mission- and margin-oriented goals.115

Despite meaningful progress in screening, pattern recognition, and clinical–community partnership and action, questions clearly remain.115 First, the ways in which we screen warrant further assessment. When is it best to screen? What are the most relevant, important questions? For patients who remain hospitalized for months, such as those in the NICU, how frequently should screens be revisited? Second, the EHR provides us with new opportunities to consider how to document SDH-related assessments, integrate complementary data streams, and recognize population-level patterns. With these new capabilities, are there shared metrics on which obstetricians, neonatologists, and primary care pediatricians can collectively track and hold one another to account? How should metrics be shared across the clinical–community continuum (e.g., with parent advisory boards, health departments, legal aid societies, school districts, city councils)? How can we ensure that such metrics add value and are not driven by entrenched biases? Third, as we move toward action, how can we maintain ongoing stakeholder engagement, a view that stretches across the life course, and strategies that pair rigor with the reality of the real world? How can we translate this engagement across the clinical–community continuum into lasting, value-added partnerships? And finally, what is the role of the health-care system in moving toward community well-being or toward a dismantling of generations-old structural issues at the heart of unjust health outcomes? Many patients encountered in health-care settings, particularly NICUs, are medically complex. Yet their social complexity frequently determines how they ultimately experience health across the life course. As we shift our focus from “follow-up” to “follow through” for patients and populations, we must, in parallel, shift toward a more balanced approach to medical and social needs. This may require health-care systems and providers to embrace social needs screening, population-level pattern recognition, and investment in social supports and community well-being. It may also push us to identify our limitations, gaps that could be filled by willing, capable, and complementary community partners.

DISCUSSION

We propose three causal pathways through which racial inequality has amplified the burden of preterm birth on the health and well-being of minority communities in the U.S.: increased risk; lower-quality care; and socioeconomic disadvantages that persist into infancy, childhood, and beyond.

To improve these pathways, we must increase health equity by removing obstacles, such as discrimination, poverty, and lack of access to quality education, housing, and health care.119 As health-care providers, we have the obligation to “follow through,” accepting that our responsibility extends beyond the hospital walls to encompass the long-term health and well-being of the infants and families we serve. We cannot do this alone. These injustices are rooted in unconstitutional governmental actions at the local, state, and federal levels.1 Developing appropriate remedies must start with a national conversation that honestly confronts this history.1,120,121 Physicians, nurses, other health-care providers, and researchers play an important role in this conversation.

Eliminating racial disparities in preterm birth and in outcomes among premature infants and their families is an ambitious agenda that will not be accomplished quickly. Although many of the interventions identified in this review can be implemented in the near term, the sheer scope of the work is daunting, especially for hospital-based specialists and researchers who may be unfamiliar in the activities like working with community organizations or advocating for structural change. There is no tool kit of specific activities and instruments, although a list of “potentially better” practices is available in an appendix to this review. Implementation of various aspects of this agenda must be tailored to the local context of community characteristics and resources. It is not expected that health-care providers know and be responsible for all the potential services. So where to begin?

First, consider developing at least three teams. One team must address the internal culture of the health care institution and the extent to which implicit bias and other aspects of care impinge on the quality of services for all patients. Many institutions already have such teams in place; if not, models are available.122 A second team involves parents who can help identify the needs and potential solutions for differences in care. Principles of diversity and inclusion should be applied to this team to assure that concerns of those whose voices have long been ignored are now heard.5,123 The third is a community resources team with a bidirectional purpose: health-care personnel can educate community partners about the health problems and needs of premature infants and their families, and community partners can engage with health-care personnel to develop a comprehensive list of resources and services in the community and to foster communication across the hospital community. Such a group might include early intervention workers, primary care physicians, child physical and occupational therapists, social services, education services, public health professionals, community leaders, and representatives from community-based agencies.
ACKNOWLEDGEMENTS

E.M.E., J.D.H. and E.A.H. receive support from the Eunice Kennedy Shriver National Institute of Child Health & Human Development of the National Institutes of Health, E.M.E. and J.D.H. by Award Number R01HD083688, J.D.H. by R01HD084667, and E.A.H. by Award Number R01MD078565. E.A.H. also receives support from the National Institute for Minority Health and Health Disparities by Award Number R01MD007652. A.F.B. receives support from the National Institute of Allergy and Infectious Diseases by Award Number 1K23AI112916.

AUTHOR CONTRIBUTIONS

The authors are listed alphabetically to indicate that they all made equal contributions to this work.

ADDITIONAL INFORMATION

The online version of this article contains supplementary material, which is available to authorized users.

Competing interests: The authors declare no competing interests.

Publisher’s note: Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

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