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Health equity in the implementation of genomics and precision medicine: A public health imperative

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Abstract

Recent reviews have emphasized the need for a health equity agenda in genomics research. To ensure that genomic discoveries can lead to improved health outcomes for all segments of the population, a health equity agenda needs to go beyond research studies. Advances in genomics and precision medicine have led to an increasing number of evidence-based applications that can reduce morbidity and mortality for millions of people (tier 1). Studies have shown lower implementation rates for selected diseases with tier 1 applications (familial hypercholesterolemia, Lynch syndrome, hereditary breast and ovarian cancer) among racial and ethnic minority groups, rural communities, uninsured or underinsured people, and those with lower education and income. We make the case that a public health agenda is needed to address disparities in implementation of genomics and precision medicine. Public health actions can be centered on population-specific needs and outcomes assessment, policy and evidence development, and assurance of delivery of effective and ethical interventions. Crucial public health activities also include engaging communities, building coalitions, improving genetic health literacy, and building a diverse workforce. Without concerted public health action, further advances in genomics with potentially broad applications could lead to further widening of health disparities in the next decade.

Conflict of Interest

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Keywords

Genomics; Health equity; Precision medicine; Public health genomics

Introduction

Health equity, defined as everyone having the opportunity to be as healthy as possible,¹ has long been a primary goal of public health. Attention to disparities in health outcomes during the COVID-19 pandemic has intensified interest in public health action to address health equity.^{2–4} Health disparities affect length and quality of life; rates of disease, disability, and death; severity of disease; and access to health care and treatments. Health equity in genomic medicine can be viewed as "the global applicability of genomic knowledge, fair and even access to genomic services such as testing and counseling, and unbiased implementation of genomic medicine."⁴ Health equity in genomics is based on improved understanding of the interaction among biological, social, and environmental factors in disease occurrence.

Recent articles^{5,6} have included calls to action for a health equity agenda in genomics and precision medicine. Such calls to action have mostly focused on addressing underrepresentation of minority and ethnic populations in genomic research. For example, this cross-sectional study documents the underrepresentation of racial and ethnic minority groups in precision oncology studies and discusses the need to increase enrollment of participants from diverse racial and ethnic background.⁷ Another example is the disparities in research studies for sickle cell disease as compared with other less common genetic disorders, such as cystic fibrosis and hemophilia.⁸

However, to ensure that genomic discoveries can lead to improved health outcomes for all segments of the population, a health equity agenda needs to go beyond basic and clinical research. Advances in genomics and precision medicine have led to an increasing number of evidence-based applications in clinical practice and disease prevention (or tier 1, per the Centers for Disease Control and Prevention 3-tiered classification⁹). Tier 1 genomic applications can improve health outcomes and reduce morbidity and mortality for millions of people with various diseases across the lifespan. Examples¹⁰ of tier 1 genomic applications in clinical practice today include newborn screening, hereditary breast and ovarian cancer, Lynch syndrome, familial hypercholesterolemia (FH), hereditary hemochromatosis, and hypertrophic cardiomyopathy. Although we use tier 1 genomic applications as examples of evidence-based genomic medicine that require public health action, we recognize that issues of health equity in access and implementation apply to thousands of genetic disorders and to precision medicine as a whole.

Disparities in Implementation of Genomic Medicine

Implementation disparities for 3 genetic disorders with tier 1 guidelines—FH, Lynch syndrome, and hereditary breast and ovarian cancer—are summarized in Table 1. For all 3 conditions, the overall implementation of current guidelines is suboptimal in the population at large, but most especially among racial and ethnic minority groups, women, people living in rural communities, people who are uninsured or underinsured, and those with

lower education and income. For example, among breast cancer survivors, non-Hispanic Black women are less likely to have BRCA testing than non-Hispanic White women. Lack of testing can be driven in part by lack of discussions with providers about testing. Black women are less likely to have discussions with their health care providers about genetic testing than non-Hispanic White women. In addition, physicians primarily serving minorities are less likely to refer a patient to a genetic counselor or to a genetics center for genetic testing. Lack of testing can affect preventive care, both for individuals and families. Black women have lower rates of risk-reducing mastectomy and risk-reducing salpingo-oo-phorectomy than non-Hispanic White women, and cascade screening rates are lower among Black families with BRCA pathogenic variants. Even among those diagnosed, inequities in access to screening and treatments exist, which affect health outcomes. Among those with FH, those in racial and ethnic minority groups, those with lower incomes, and women are less likely to start treatment with PCSK9 inhibitors and more likely to be denied insurance coverage for PCSK9 inhibitors. Asian persons and Black persons are less likely than White persons to achieve optimal low-density lipoprotein cholesterol. Women are also less likely to achieve optimal low-density lipoprotein cholesterol reduction than men.

Access to health care is critically important for the implementation of evidence-based genomics and precision medicine applications. Access to testing and interventions involves a combination of psychosocial and structural factors, including availability, accessibility (eg, ability to get to testing services, have appropriate level of genetics health literacy, and language access), and acceptability (eg, people wanting the test, trusting the institution and provider(s), and perceiving that getting tested as beneficial and consistent with their health beliefs and goals). Barriers to health care differentially affect racial and ethnic minority groups, rural communities, people with disabilities, and people with lower incomes, among others. As a result, access to and utilization of health services and specialty care, including genetic services, are lower among these groups, which can lead to lower rates of diagnosis, suboptimal care, and worse health outcomes.

Public Health Action in Addressing Disparities in Implementation of Genomic Medicine

"Public health is what we do together as a society to ensure the conditions in which everyone can be healthy. Although many sectors play key roles, governmental public health is an essential component."⁵² There are 3 broad themes for government public health action at the intersection of genomics and health equity, based on the core functions of public health—assessment, policy development, and assurance.⁵³ Health equity is at the center of public health action. A special issue of the *American Journal of Public Health*, "COVID-19, Racism, and Public Health Infrastructure,"⁵⁴ sounds the alarm for dealing with the disparate effect of the pandemic on racial and ethnic minority groups and low-income populations. If genomics and precision medicine are to improve health for all, generational inequities embedded in society at large and the US health system have to be acknowledged and addressed. Any success resulting from a siloed equity approach to genomics could be overshadowed by other poor health outcomes unless the underlying determinants of health inequities, such as lack of access to health care, healthy food, air, and water; inadequate

housing; chronic stress; and exposures to environmental toxins are addressed, together with core drivers such as structural racism.

Long-standing health disparities and social inequities led to the articulation of Public Health 3.0, as described in a 2016 Health and Human Services report.⁵² The framework sought to revitalize the US approach to governmental public health with strong and diverse workforce, strategic partnerships, sustained funding, specific population-level data and metrics, and foundational infrastructure.

In the context of genomics and precision medicine, we view public health as an important part of the solution in dealing with equity challenges in genomics, just as with other fields of health care. It is important to note that although a focused population health approach to genomics might broaden access to genomic innovations, a public health approach including community engagement will be needed to address issues of trust, many of which are rooted in long-standing community experiences with structural racism.

Current public health practice is not optimally integrating genomics into the essential functions of assessment, policy development, and assurance. Here, and in Table 2, we summarize our vision and opportunities for specific public health actions that can be conducted by the Centers for Disease Control and Prevention and its many partners to help reduce disparities in the implementation of genomics and precision medicine. This framework lays the groundwork for the next steps, including identification of specific goals and measurable outcomes in the design and implementation of specific community-based interventions.

Public health assessment through data modernization and applied research

An important function of public health is to collect real-time data through surveillance and applied research to drive policies, guidelines, and programs. National, state, local, and community-specific data systems are needed to evaluate disparities in genomic application utilization, interventions, and outcomes to inform evidence-based guidelines and implementation of genomic medicine.

Public health is currently undergoing strategic innovation as part of a data modernization initiative⁵⁵ by assessing the opportunities for and limitations of new and nontraditional data sources and data science approaches to inform public health decision making. The need for tracking progress and outcomes in genomics and precision medicine is a high priority for action, and novel approaches could increase public health's ability to track health disparities, for example, by using geocoding to better measure area of residence to identify areas of high need and low resources. New and innovative approaches are needed to integrate tracking for genetic disorders into surveillance systems, surveys, claims, and administrative databases. For example, employer-sponsored health insurance claims data were used to track trends and differences in *BRCA1/2* genetic testing and receipt of preventive interventions in women aged 18 to 64 years between nonmetropolitan and metropolitan areas in the United States.⁴⁹ The analysis documented the existence of geographic differences in *BRCA* test utilization as a proxy for rural-urban differences in access to care.

Several groups are currently using next-generation tools, such as machine learning approaches, and nontraditional data sources for conducting public health surveillance of genetic disorders. A case in point is the use of the FIND-FH algorithm in health systems.³⁸ Such efforts need to be expanded for various diseases and cover broader population samples than traditional databases, such as those from employer-sponsored health plans. These approaches may facilitate identification of disparities in diagnosis, interventions, and outcomes and inform programs, practice, and policy.

Another example is the emerging integration of genomics into population-based cancer registries and surveillance systems.⁵⁶ Cancer surveillance traditionally is conducted based on tumor anatomic location and histology. However, molecular markers, such as gene expression profiles, can identify heterogeneous subgroups associated with different risk factors, treatment responses, recurrences, and survival patterns. In addition, conducting surveillance for hereditary cancer will allow stratification of reporting and tracking of population cancer incidence, outcomes, and disparities by underlying genetic cause.⁵⁶

In addition to public health surveillance, applied research based on principles of implementation science⁵⁷ is needed to evaluate multisectorial interventions (individual, health care providers, health systems, policy interventions) that can facilitate implementation of evidence-based genomic applications and drive uptake of recommended services. These studies would have a special emphasis on communities and subpopulations with the largest gaps in implementation. Until recently, implementation science had not been a strong focus in the genomic research portfolio.⁵⁸ Recent initiatives by the National Institutes of Health^{59,60} are attempting to leverage implementation science to close these gaps in genomic medicine. Accelerating implementation science can go a long way in identifying best practices for ensuring equity in the implementation of genomic medicine.

Furthermore, there is an important role for social, behavioral, and communication sciences in assessing best approaches for effective translation of genomic discoveries into population health benefits.⁶¹ Such approaches will require consideration and evaluation of multiple systemic and psychosocial factors occurring within and outside health care settings, as well the broader challenges of information diffusion, health literacy, and action in families and communities.⁵⁹

Evidence Synthesis, Guidelines, and Policy Development

Public health leadership is critically needed to establish an evidentiary foundation and guidance for implementing genomic applications to improve health outcomes in all segments of the population. It is imperative to assess the unique challenges of communities negatively affected by social determinants of health, including racism. Public health action is also needed to address policy barriers to effective, widespread implementation and monitoring of genomic applications.

Public health can build on previous successes of genomic evaluation initiatives, such as the Evaluation of Genomic Applications in Practice and Prevention initiative⁶² and the ACCE (Analytic validity, Clinical validity, Clinical utility, and Ethical legal and social

implications) framework⁶³ to develop processes for systematically evaluating evidence on the implementation of tier 1 genomic applications in different communities. Necessary actions include convening methods development and evidence review panels for topics of interest and developing and testing novel evidence synthesis strategies to reduce health disparities. The focus of reviews will be on implementation of science-based evidence regarding how to most effectively use genomics applications that have already been deployed in multiple communities and health systems. Considerations for implementation specific to certain communities and populations negatively affected by social determinants of health should be prioritized. Such evidence synthesis can drive the development of nationally credible, evidence-based implementation guidelines for genomic testing applications tailored to different communities and health care settings to facilitate broader understanding and adoption of evidence-based services.

A specific example of an implementation challenge is the low cascade screening uptake in relatives of affected patients, especially within communities of color. Cascade screening is an important component of guidelines for several tier 1 genomic applications, but its implementation falls short. A recent systematic review⁶⁴ assessed barriers to and facilitators of cascade screening. The review identified individual-, interpersonal-, and environmentallevel barriers that currently interfere with cascade screening and may be associated with health disparities. At the individual level, the review identified barriers related to demographics, knowledge, attitudes, beliefs, emotional responses of the individual, and attitudes toward relatives. Examples of identified barriers include low income, lack of access to health care, and lack of effective approaches that reach relatives. The review also identified factors associated with communication, support, and dynamics with family members and clinicians, such as communication and language barriers and lack of provider awareness and engagement. At the environmental level, the review identified barriers correlated with accessibility of genetic services, such as cost and insurance coverage. The review identified the need for implementation studies to further investigate these factors and inform future interventions for improving the implementation of cascade testing for genetic disorders in all populations. This systematic review can influence the development of guidelines and recommendations for best practices for implementation of cascade screening in different communities and promote equity in implementation.

To achieve the goal of equitable access to and uptake of genetic tests and services, policy barriers must be overcome. Many people who could benefit from these services are unor underinsured; expanded coverage and adequate reimbursement could help remedy this obstacle, especially among racial and ethnic minority populations, those with inadequate insurance coverage, and those who live in rural areas. Additional obstacles arise from data sharing and privacy policies that complicate the development of interoperable data systems, a critical source of real-world evidence to inform guidelines and enable cascade testing.

An example of an important policy consideration in cascade screening is how the Health Insurance Portability and Accountability Act (HIPAA),⁶⁵ considered by some providers to be a barrier to cascade screening, should be interpreted in the context of sharing of genomic information among relatives in different communities, especially those negatively affected by social determinants of health. A recent review⁶⁶ analyzed the HIPAA Privacy Rule and

developed multiple scenarios that could inform how HIPAA can be interpreted for patientand provider-mediated genetic risk notification. This analysis concluded that several forms of patient- or clinician-initiated contact of family members are permissible under HIPAA and consistent with ethical obligations of care to patients and their families. These include direct contacts of patients' adult relatives with patients' permission and direct contact of the relatives' providers. It is important to acknowledge, however, that effective and equitable cascade testing heavily depends on the dynamics of family relationships and that health care providers may not be able to ethically contact relatives without explicit permission of patients. It is crucial to assess how to translate this policy analysis into guidelines and tools among racial and ethnic minority populations to help advance cascade screening for many conditions. Further research will be needed to determine best practices for implementing these guidelines broadly and addressing the specific needs of different communities.

Public Health Assurance: Programs, Resources, and Workforce

Development

There is an essential role for public health in addressing health equity in genomics by integrating genomic tools in different community and health care settings. Public health programs in collaboration with communities and health systems could support effective implementation and improve population health outcomes by engaging communities equitably and addressing documented disparities in genomic medicine implementation.

Public health programs can establish exemplar projects and networks for population-level implementation of genomic testing applications that focus on communities that are differentially affected by social determinants of health. Engaging with communities is essential to build trust and tailor approaches to meet specific needs, raise public awareness of genomic applications, and increase uptake. These projects can be integrated into existing community and health care efforts, such as health screening programs, to identify people at increased risk of poor health outcomes associated with hereditary conditions.

Public health can also work with communities to increase access to genetic services, for example, through telehealth. For example, the Maternal and Child Health Bureau of the Health Resources and Services Administration has established 7 Regional Genetics Networks and the National Coordinating Center as part of ongoing efforts to improve the health of populations that are medically underserved.⁶⁵ A current emphasis of the network, especially during the COVID-19 pandemic, is to enhance the delivery of genetic services using telehealth resources (telegenetics), an emerging tool to facilitate virtual access to medical geneticists and genetic counselors, especially among rural communities with limited access to genetic services. A recent commentary reviewed the successes and challenges in the implementation of virtual genetics visits during the pandemic and discussed genetic testing considerations in addressing health disparities.⁶⁷

Another example is the ongoing partnership between the Million Hearts initiative and the National Association for Health Centers to improve the use of statin therapy for patients at high risk of heart attack and stroke through an enhanced care process to prescribe statins to as many patients at high risk as possible.⁶⁸ An added component of this initiative could

include enhanced methods for finding people at high genetic risk, such as those with FH, and cascade testing of their relatives.

Furthermore, tools and resources for action need to be specific and tailored to health care organizations and communities. Tier 1 genomic application decision support tools for clinical practice will be needed, as well as culturally and community-appropriate educational materials for patients and families. Because family health history is a common risk factor for many conditions, simple family health history tools,⁶⁹ including the popular Surgeon General's My Family Health Portrait,⁷⁰ could be adapted to different communities and health care organizations to overcome barriers in the identification of genetic disorders and facilitate cascade screening.

Finally, there is an emerging need to build the clinical and public health genomics workforce and increase its diversity. A survey of American Board of Medical Genetics and Genomics board-certified/eligible diplomates in 2019⁷¹ characterized the US clinical genetics workforce to inform workforce planning and public policy development. The survey showed that most genetics specialists work in academic medical centers in major metropolitan areas, leaving many people in rural areas with no or limited access to genetics specialists. In the absence of concerted efforts to increase the number of genetics specialists through enhanced training and enhanced career and salary incentives, the current and future workforce will not meet the increasing patient needs in genomic medicine.⁷² Furthermore, the lack of diversity in the genetics workforce hinders its ability to meet the needs of all populations—90% of genetic counselors responding to the survey identified as non-Hispanic White. In public health, there is a paucity of professionals trained in human genomics. In state public health programs, genetics capacity is concentrated almost exclusively in newborn screening programs and maternal and child health programs. A survey conducted by the Council for State and Territorial Epidemiologists⁷³ revealed that the most state epidemiology positions focused on infectious diseases, with only 0.1% focused on human genomics. Overall, state epidemiologists assessed their state epidemiology capacity in genomics as none to minimal. The COVID-19 pandemic has expanded state and academic capacity in pathogen genomics (see example of a recent initiative⁷⁴), but more effort will be needed before there will be a spillover effect to human genomics capacity.

Concluding Remarks

An ambitious public health agenda is needed now to ensure that the entire population reaps the benefits of genomics and precision medicine. We presented an overall vision and opportunities for selected public health actions (Table 2), but a more in-depth analysis to identify specific goals and measurable outcomes will be needed to design and implement specific interventions tailored to reach persons with genetic disorders in populations and communities experiencing health inequities. Achieving health equity in genomics and precision medicine will depend on strong collaborations with community leaders, patient organizations, professional organizations, academia, health care systems, health care payers, industry, and charitable foundations. A unique and valuable role for public health is to serve as a convener of partners to ensure community engagement that is inclusive and participatory and ultimately helps to accelerate the equitable implementation of genomics

and precision medicine. There is also a unique opportunity to engage social and behavioral scientists in assessing the effect of multiple psychosocial and communication factors on the optimal implementation of genomics and precision medicine in all populations. Finally, it is important to recognize that even though many of the populations with less access to genomics are the same communities with significant negative effects of social determinants of health, genomics will not address disparities that are primarily caused by social determinants.⁷⁵ A health equity agenda in genomics should be just one component of an overall health equity approach to health and health care. Any success resulting from a siloed equity approach to genomics would be overshadowed by other poor health outcomes unless the underlying determinants of health inequities, such as lack of access to health care, inadequate housing, and exposures to environmental toxins, are addressed together with core drivers, such as structural racism.

Experience from COVID-19 demonstrated the need to have data collection, systems, and analyses that disaggregate data and make that data accessible to stakeholders, including communities experiencing disproportionate gaps in implementation of genomic medicine. This will require integration of utilization of genetic screening and interventions into electronic health records as well as careful assessment and solutions to privacy and confidentiality concerns associated with data sharing. To ensure that equity approaches are embedded in all stages of implementation planning, engagement of communities is essential for wider implementation of genomic medicine.

We have a real opportunity to get out in front of the emerging field of genomics and precision medicine before the existing disparities widen further as new technologies lead to more evidence-based applications. Many health disparities are deeply entrenched, but as a new field, we have a window of opportunity to address the implementation challenges early on. Because the pandemic has differentially affected subgroups of the population and laid bare the effect of social determinants of health, achieving health equity in the implementation of genomics and precision medicine is more important than ever.

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References

- 1. Health equity. Centers for Disease Control and Prevention, https://www.cdc.gov/chronicdisease/ healthequity/index.htm. Accessed July 24, 2021.
- Ndumbe-Eyoh S, Muzumdar P, Betker C, Oickle D. 'Back to better': amplifying health equity, and determinants of health perspectives during the COVID-19 pandemic. Glob Health Promot. 2021; 28(2):7–16. 10.1177/17579759211000975. [PubMed: 33761795]
- 3. Tai DBG, Shah A, Doubeni CA, Sia IG, Wieland ML. The disproportionate impact of COVID-19 on racial and ethnic minorities in the United States. Clin Infect Dis. 2021;72(4):703–706. 10.1093/cid/ ciaa815. [PubMed: 32562416]
- Jooma S, Hahn MJ, Hindorff LA, Bonham VL. Defining and achieving health equity in genomic medicine. Ethn Dis. 2019;29(suppl 1): 173–178. 10.18865/ed.29.S1.173. [PubMed: 30906166]
- 5. Precision medicine needs an equity agenda. Nat Med. 2021;27(5):737. 10.1038/s41591-021-01373y. [PubMed: 33990803]

- Balogun OD, Olopade OI. Addressing health disparities in cancer with genomics. Nat Rev Genet. 2021;22(10):621–622. 10.1038/s41576-021-00390-4. [PubMed: 34244675]
- Aldrighetti CM, Niemierko A, Van Allen E, Willers H, Kamran SC. Racial and ethnic disparities among participants in precision oncology clinical studies. JAMA Netw Open. 2021;4(11):e2133205. 10.1001/jamanetworkopen.2021.33205. [PubMed: 34748007]
- Kanter J, Meier ER, Hankins JS. Improving outcomes for patients with sickle cell disease in the United States. Making the case for more resources, surveillance, and longitudinal data. JAMA Health Forum. 2021; 2(10):e213467. 10.1001/jamahealthforum.2021.3467.
- Dotson WD, Douglas MP, Kolor K, et al. Prioritizing genomic applications for action by level of evidence: a horizon-scanning method. Clin Pharmacol Ther. 2014;95(4):394–402. 10.1038/ clpt.2013.226. [PubMed: 24398597]
- Tier-classified guidelines database. Centers for Disease Control and Prevention. https:// phgkb.cdc.gov/PHGKB/tierStartPage.action. Accessed July 25, 2021.
- Win AK, Jenkins MA, Dowty JG, et al. Prevalence and penetrance of major genes and polygenes for colorectal cancer. Cancer Epidemiol Biomarkers Prev. 2017;26(3):404–112. 10.1158/1055-9965.EP1-16-0693. [PubMed: 27799157]
- de Ferranti SD, Rodday AM, Mendelson MM, Wong JB, Leslie LK, Sheldrick RC. Prevalence of familial hypercholesterolemia in the 1999 to 2012 United States National Health and Nutrition Examination Surveys (NHANES). Circulation. 2016;133(11):1067–1072. 10.1161/ CIRCULATIONAHA.115.018791. [PubMed: 26976914]
- Abul-Husn NS, Soper ER, Odgis JA, et al. Exome sequencing reveals a high prevalence of BRCA1 and BRCA2 founder variants in a diverse population-based biobank. Genome Med. 2019;12(1):2. 10.1186/s13073-019-0691-1. [PubMed: 31892343]
- Berera S, Koru-Sengul T, Miao F, et al. Colorectal tumors from different racial and ethnic minorities have similar rates of mismatch repair deficiency. Clin Gastroenterol Hepatol. 2016; 14(8): 1163–1171. 10.1016/j.cgh.2016.03.037. [PubMed: 27046481]
- Amrock SM, Duell PB, Knickelbine T, et al. Health disparities among adult patients with a phenotypic diagnosis of familial hypercholesterolemia in the CASCADE-FH[™] patient registry. Atherosclerosis. 2017;267:19–26. 10.1016/j.atherosclerosis.2017.10.006. [PubMed: 29080546]
- Belay B, Racine AD, Belamarich PF. Underrepresentation of non-White children in trials of statins in children with heterozygous familial hypercholesterolemia. Ethn Dis. 2009; 19(2): 166–171. [PubMed: 19537228]
- Ford ES, Li C, Pearson WS, Zhao G, Mokdad AH. Trends in hypercholesterolemia, treatment and control among United States adults. Int J Cardiol. 2010;140(2):226–235. 10.1016/ j.ijcard.2008.11.033. [PubMed: 19081646]
- Sturm AC, Truty R, Callis TE, et al. Limited-variant screening vs comprehensive genetic testing for familial hypercholesterolemia diagnosis. JAMA Cardiol. 2021;6(8):902–909. [PubMed: 34037665]
- Garg A, Fazio S, Duell PB, et al. Molecular characterization of familial hypercholesterolemia in a North American cohort. J Endocr Soc. 2019;4(1):bvz015. 10.1210/jendso/bvz015. [PubMed: 31993549]
- Derington CG, Colantonio LD, Herrick JS, et al. Factors associated with PCSK9 inhibitor initiation among US Veterans. J Am Heart Assoc. 2021; 10(8):e019254. 10.1161/JAHA.120.019254. [PubMed: 33821686]
- Myers KD, Farboodi N, Mwamburi M, et al. Effect of access to prescribed PCSK9 inhibitors on cardiovascular outcomes. Circ Cardiovasc Qual Outcomes. 2019;12(8):e005404. 10.1161/ CIRCOUTCOMES.118.005404. [PubMed: 31331194]
- Cragun D, Weidner A, Lewis C, et al. Racial disparities in BRCA testing and cancer risk management across a population-based sample of young breast cancer survivors. Cancer. 2017;123(13):2497–2505. 10.1002/cncr.30621. [PubMed: 28182268]
- Reid S, Cragun D, Tezak A, et al. Disparities in BRCA counseling across providers in a diverse population of young breast cancer survivors. Genet Med. 2020;22(6): 1088–1093. 10.1038/ s41436-020-0762-0. [PubMed: 32066870]

- Levy DE, Byfield SD, Comstock CB, et al. Underutilization of BRCA1/2 testing to guide breast cancer treatment: Black and Hispanic women particularly at risk. Genet Med. 2011; 13(4):349– 355. 10.1097/GIM.0b013e3182091ba4. [PubMed: 21358336]
- 25. Shields AE, Burke W, Levy DE. Differential use of available genetic tests among primary care physicians in the United States: results of a national survey. Genet Med. 2008; 10(6):404–414. 10.1097/GIM.0b013e3181770184. [PubMed: 18496223]
- Fehniger J, Lin F, Beattie MS, Joseph G, Kaplan C. Family communication of BRCA1/2 results and family uptake of BRCA1/2 testing in a diverse population of BRCA1/2 carriers. J Genet Couns. 2013;22(5):603–612. 10.1007/s10897-013-9592A [PubMed: 23666114]
- McCarthy AM, Bristol M, Domchek SM, et al. Health care segregation, physician recommendation, and racial disparities in BRCA1/2 testing among women with breast cancer. J Clin Oncol. 2016;34(22):2610–2618. 10.1200/JCO.2015.66.0019. [PubMed: 27161971]
- Babatunde OA, Eberth JM, Felder TM, et al. Racial disparities and diagnosis-to-treatment time among patients diagnosed with breast cancer in South Carolina. J Racial Ethn Health Disparities. 2022;9(1):124–134. [PubMed: 33428159]
- Conley CC, Ketcher D, Reblin M, et al. The big reveal: family disclosure patterns of BRCA genetic test results among young Black women with invasive breast cancer. J Genet Couns. 2020;29(3):410–422. 10.1002/jgc4.1196. [PubMed: 31912597]
- Rubinsak LA, Kleinman A, Quillin J, et al. Awareness and acceptability of population-based screening for pathogenic BRCA variants: do race and ethnicity matter? Gynecol Oncol. 2019;154(2):383–387. 10.1016/j.ygyno.2019.06.009. [PubMed: 31239069]
- 31. Camacho FT, Tan X, Alcala HE, Shah S, Anderson RT, Balkrishnan R. Impact of patient race and geographical factors on initiation and adherence to adjuvant endocrine therapy in medicare breast cancer survivors. Medicine (Baltimore). 2017;96(24):e7147. 10.1097/MD.000000000007147. [PubMed: 28614244]
- 32. Muller C, Lee SM, Barge W, et al. Low referral rate for genetic testing in racially and ethnically diverse patients despite universal colorectal cancer screening. Clin Gastroenterol Hepatol. 2018;16(12):1911–1918.e2. 10.1016/j.cgh.2018.08.038. [PubMed: 30130624]
- Dharwadkar P, Greenan G, Stoffel EM, et al. Racial and ethnic disparities in germline genetic testing of patients with young-onset colorectal cancer. Clin Gastroenterol Hepatol. 2022;20(2):353–361.e3. 10.1016/j.cgh.2020.12.025. [PubMed: 33359728]
- 34. Carethers JM. Screening for colorectal cancer in African Americans: determinants and rationale for an earlier age to commence screening. Dig Dis Sci. 2015;60(3):711–721. 10.1007/ s10620-014-3443-5. [PubMed: 25540085]
- Tsai MH, Xirasagar S, de Groen PC. Persisting racial disparities in colonoscopy screening of persons with a family history of colorectal cancer. J Racial Ethn Health Disparities. 2018;5(4):737–746. 10.1007/s40615-017-0418-1. [PubMed: 28812255]
- Salyer CV, Dontsi M, Armstrong MA, Lentz S, Hoodfar E, Powell B. Variation in physiciandirected immunohistochemistry screening among women with endometrial cancer. Int J Gynecol Cancer. 2020;30(9): 1356–1365. 10.1136/ijgc-2020-001449. [PubMed: 32641393]
- Balia S, Ekpo EP, Wilemon KA, Knowles JW, Rodriguez F. Women living with familial hypercholesterolemia: challenges and considerations surrounding their care. Curr Atheroscler Rep. 2020;22(10):60. 10.1007/s11883-020-00881-5. [PubMed: 32816232]
- 38. Myers KD, Knowles JW, Staszak D, et al. Precision screening for familial hypercholesterolaemia: a machine learning study applied to electronic health encounter data. Lancet Digit Health. 2019;1(8):e393–e402. http://doi.org/10-1016/S2589-7500(19)30150-5. [PubMed: 33323221]
- Gaddam S, Heller SL, Babb JS, Gao Y. Male breast cancer risk assessment and screening recommendations in high-risk men who undergo genetic counseling and multigene panel testing. Clin Breast Cancer. 2021; 21(1):e74–e79. 10.1016/j.clbc.2020.07.014. [PubMed: 32828665]
- Jeong GW, Shin W, Lee DO, et al. Uptake of family-specific mutation genetic testing among relatives of patients with ovarian cancer with BRCA1 or BRCA2 mutation. Cancer Res Treat. 2021;53(1):207–211. 10.4143/crt.2020.364. [PubMed: 32777875]

- Adar T, Rodgers LH, Shannon KM, et al. Universal screening of both endometrial and colon cancers increases the detection of Lynch syndrome. Cancer. 2018;124(15):3145–3153. 10.1002/ cncr.31534. [PubMed: 29750335]
- Seppälä TT, Dominguez-Valentin M, Crosbie EJ, et al. Uptake of hysterectomy and bilateral salpingo-oophorectomy in carriers of pathogenic mismatch repair variants: a Prospective Lynch Syndrome Database report. Eur J Cancer. 2021;148:124–133. 10.1016/j.ejca.2021.02.022. [PubMed: 33743481]
- Hagger MS, Hardcastle SJ, Hu M, et al. Health literacy in familial hypercholesterolemia: a crossnational study. Eur J Prev Cardiol. 2018;25(9):936–943. 10.1177/2047487318766954. [PubMed: 29592531]
- 44. Hope HF, Binkley GM, Fenton S, Kitas GD, Verstappen SMM, Symmons DPM. Systematic review of the predictors of statin adherence for the primary prevention of cardiovascular disease. PLoS One. 2019;14(1):e0201196. 10.1371/journal.pone.0201196. [PubMed: 30653535]
- Gamble CR, Huang Y, Wright JD, Hou JY. Precision medicine testing in ovarian cancer: the growing inequity between patients with commercial vs Medicaid insurance. Gynecol Oncol. 2021; 162(1): 18–23. 10.1016/j.ygyno.2021.04.025. [PubMed: 33958212]
- Adams J, White M, Barker G, Mathers J, Burn J. Are there socioeconomic inequalities in age of resection of colorectal cancer in people with HNPCC? Fam Cancer. 2003;2(3–4): 169–173. 10.1023/b:fame.0000004624.71900.15. [PubMed: 14707528]
- Groth NA, Stone NJ, Benziger CP. Cardiology clinic visit increases likelihood of evidencebased cholesterol prescribing in severe hypercholesterolemia. Clin Cardiol. 2021;44(2): 186–192. 10.1002/clc.23521. [PubMed: 33355940]
- 48. Wong ND, Bang M, Block RC, Peterson ALH, Karalis DG. Perceptions and barriers on the use of proprotein subtilisin/kexin type 9 inhibitors in heterozygous familial hypercholesterolemia (from a survey of primary care physicians and cardiologists). Am J Cardiol. 2021; 152:57–62. 10.1016/ j.amjcard.2021.04.034. [PubMed: 34147211]
- Kolor K, Chen Z, Grosse SD, et al. BRCA genetic testing and receipt of preventive interventions among women aged 18–64 years with employer-sponsored health insurance in nonmetropolitan and metropolitan areas—United States, 2009–2014. MMWR Surveill Summ. 2017;66(15):1–11. 10.15585/mmwr.ss6615al.
- 50. Noll A, Parekh JP, Zhou M, et al. Barriers to Lynch syndrome testing and preoperative result availability in early-onset colorectal cancer: A national physician survey study. Clin Transl Gastroenterol. 2018;9(9):185. 10.1038/s41424-018-0047-y. [PubMed: 30237431]
- Beamer LC, Grant ML, Espenschied CR, et al. Reflex immunohistochemistry and microsatellite instability testing of colorectal tumors for Lynch syndrome among US cancer programs and follow-up of abnormal results. J Clin Oncol. 2012;30(10):1058–1063. 10.1200/JCO.2011.38.4719. [PubMed: 22355048]
- De Salvo KB, Wang YC, Harris A, Auerbach J, Koo D, O'Carroll P. Public health 3.0: a call to action for public health to meet the challenges of the 21st century. Prev Chronic Dis. 2017;14:E78. 10.5888/pcd14.170017. [PubMed: 28880837]
- 53. Sellers K, Fisher JS, Kuehnert P, Castrucci BC. Meet the revised 10 essential public health services: developed by the field, centering equity. Health Affairs, 10.1377/ forefront.20210319.479091. Accessed July 25, 2021.
- Borrell LN, Erwin PC, Fiala S. COV1D-19, racism, and public health infrastructure. Am J Public Health. 2021; 111(S3):S172. 10.2105/AJPH.2021.306505. [PubMed: 34709863]
- 55. Data modernization initiative. Centers for Disease Control and Prevention. https://www.cdc.gov/ surveillance/data-modernization/index.html. Accessed July 25, 2021.
- 56. Khoury MJ, Penberthy L. Integrating genomics into population-based cancer surveillance in the era of precision medicine. Centers for Disease Control and Prevention. https://blogs.cdc.gov/ genomics/2017/09/19/integrating-genomics-2/. Accessed September 7, 2021.
- Roberts MC, Mensah GA, Khoury MJ. Leveraging implementation science to address health disparities in genomic medicine: examples from the field. Ethn Dis. 2019;29(suppl 1): 187–192. 10.18865/ed.29.S1.187. [PubMed: 30906168]

- Roberts MC, Kennedy AE, Chambers DA, Khoury MJ. The current state of implementation science in genomic medicine: opportunities for improvement. Genet Med. 2017;19(8):858–863. 10.1038/gim.2016.210. [PubMed: 28079898]
- 59. Prevention and early detection for hereditary cancer syndromes. National Cancer Institute, https://cancercontrol.cancer.gov/research-emphasis/contribution-to-the-cancer-moonshot/ prevention-and-early-detection. Accessed July 25, 2021.
- 60. Khoury MJ, Mensah GA, Implementation science to improve case finding, cascade screening, and treatment for familial hypercholesterolemia: a prototype for precision public health research. Centers for Disease Control and Prevention, https://blogs.cdc.gov/genomics/2020/05/05/implementation-science-3/. Accessed July 25, 2021.
- Koehly LM, Persky S, Shaw P, et al. Social and behavioral science at the forefront of genomics: discovery, translation, and health equity. Soc Sci Med. 2021;271:112450. 10.1016/ j.socscimed.2019.112450. [PubMed: 31558303]
- Evaluation of Genomic Applications in Practice and Prevention (EGAPP) Working Group. The EGAPP initiative: lessons learned. Genet Med. 2014;16(3):217–224. 10.1038/gim.2013.110. [PubMed: 23928914]
- 63. ACCE model process for evaluating genetic tests. Centers for Disease Control and Prevention, https://www.cdc.gov/genomics/gtesting/acce/index.htm. Accessed July 25, 2021.
- 64. Srinivasan S, Won NY, Dotson WD, Wright ST, Roberts MC. Barriers and facilitators for cascade testing in genetic conditions: a systematic review. Eur J Hum Genet. 2020;28(12):1631–1644. 10.1038/s41431-020-00725-5. [PubMed: 32948847]
- 65. National Coordinating Center for the Regional Genetics Networks. https://nccrcg.org/. Accessed July 25, 2021.
- 66. Hendrikson NB, Wagner JK, Hampel H, et al. What guidance does HIPAA offer to providers considering familial risk notification and cascade genetic testing? J Law Biosci. 2020;7(1):lsaa071. [PubMed: 34221429]
- Uhlmann WR, McKeon AJ, Wang C. Genetic counseling, virtual visits and equity in the era of COVID-19 and beyond. J Genet Couns. 2021; 30(4): 1038–1045. 10.1002/jgc4.1469. [PubMed: 34291525]
- 68. NACHC Million Hearts® Initiative. National Association of Community Health Centers, https://www.nachc.org/clinical-matters/nachc-million-hearts-initiative/. Accessed August 17, 2021.
- Wildin RS, Messersmith DJ, Houwink EJF. Modernizing family health history: achievable strategies to reduce implementation gaps. J Community Genet. 2021;12(3):493–196. 10.1007/ si2687-021-00531-6. [PubMed: 34028705]
- My Family Health Portrait. Centers for Disease Control and Prevention. http:// kahuna.clayton.edu/jqu/FHH/html/index.html. Accessed July 25, 2021.
- Jenkins BD, Fischer CG, Polito CA, et al. The 2019 US medical genetics workforce: a focus on clinical genetics. Genet Med. 2021; 23(8): 1458–1464. 10.1038/s41436-021-01162-5. [PubMed: 33941882]
- 72. Garrison T, Truong T, Green RF. How accessible are genetics providers and how can access be increased? Centers for Disease Control and Prevention. https://blogs.cdc.gov/genomics/ 2020/10/05/how-accessible/. Accessed July 28, 2021.
- 73. Arrazola J, Binkin N, Israel M, et al. Assessment of epidemiology capacity in state health departments United States, 2017. MMWR Morb Mortal Wkly Rep. 2018;67(33):935–939. Published correction appears in MMWR Morb Mortal Wkly Rep. 2019;68(16):377. 10.15585/mmwr.mm6733a5. [PubMed: 30138304]
- 74. SARS-CoV-2 Innovation: Broad agency announcement awards. Centers for Disease Control and Prevention, https://www.cdc.gov/amd/whats-new/cdc-announces-awards-SARS-CoV-2-sequencing-SPHERES-initiative.html. Accessed July 25, 2021.
- 75. West KM, Blacksher E, Burke W. Genomics, health disparities, and missed opportunities for the Nation's research agenda. JAMA. 2017;317(18):1831–1832. 10.1001/jama.2017.3096. [PubMed: 28346599]

	FH	HBOC	ΓS
Tier 1 recommendation	For those diagnosed with FH, perform cascade testing of first-degree relatives for FH by measuring LDL-C concentration, genetic testing, or both.	Assess risk for women with a personal or family history of breast, ovarian, tubal, or peritoneal cancer or who have an ancestry associated with <i>BRCA</i> variants and refer those identified as at risk for genetic counseling and, if indicated after counseling, testing.	Screen turnor tissue from colorectal (and possibly endometrial) cancer using MSI and/or IHC to check for LS. In some cases, genetic testing might be needed to confirm LS diagnosis. Genetic testing to identify pathogenic variants can allow cascade testing to identify family members with FH.
Estimated number of people with condition in the United States	1,300,000	660,000 to 990,000	1,200,000 ¹¹
Prevalence by race/ r ethnicity H H	Non-Hispanic White: 0.4% Non-Hispanic Black: 0.5% Hispanic: 0.2% ¹² ies	Non-Hispanic White: 1.5% Non-Hispanic Black: 0.5% Hispanic: 0.4% ¹³	Among those with colorectal tumors screened for LS: Non-Hispanic White: 10.4% Non-Hispanic Black: 9.6% Hispanic: 12.6% ¹⁴
Ethnicity/race	 Disparities documented by race and ethnicity in research,¹⁵ registries,¹⁵ and clinical trials,¹⁶ cholesterol screening,¹⁷ FH awareness,¹⁶ age at diagnosis,¹⁵ are at diagnosis,¹⁵ are at diagnosis,¹⁵ are diagnosis,¹⁵ are of high-intensity statins¹⁵ and PCSK9 inhibitors,²⁰ and denial of insurance coverage for PCSK9 inhibitors.²¹ 	Among breast cancer survivors, disparities documented by race and ethnicity in ^{22–29} a wareness of <i>BRCA</i> testing, ³⁰ discussion of genetic testing with health care provider, referrals to genetics specialists and centers, rates of <i>BRCA</i> testing, itime to surgery and chemotherapy, rates of risk-reducing mastectomy and salpingo- ophorectomy, and adherence to endocrine therapy. ³¹	Disparities documented by race and ethnicity in • referrals for genetic evaluation, ³² • genetic counseling and genetic testing, ³³ • sharing of personal or family history, information that affects screening age, ³⁴ • colonoscopy screening at ages 40 to 49 for first- degree relatives of those with colorectal cancer, ³⁵ and • IHC tumor screening of endometrial cancer. ³⁶
Sex	 Disparities documented for women in^{15,37,38} eresearch, age at diagnosis, likelihood of FH being diagnosed after an ASCVD event, enheving target LDL-C levels, enheving target LDL-C levels, use of statins, high-intensity statins, and PCSK9 inhibitors, maintenance of statin therapy, including during childbearing, side effects associated with statin use, and denial of insurance coverage for PCSK9 inhibitors.²¹ 	Men with HBOC are less likely than women to receive implementation of • screening recommendations ³⁹ and • cascade testing in families. ⁴⁰	Disparities documented for • rates of LS screening for endometrial cancer compared with colorectal cancer ⁴¹ and • uptake of risk-reducing surgery for women with LS in alignment with clinical guidelines. ⁴²
SES	Disparities documented for SES in • health literacy. ⁴³ • adherence to statin therapy. ⁴⁴ • initiation of PCSK9 inhibitors. ²⁰ and • denial of insurance coverage for PCSK9 inhibitors. ²¹	Disparities documented for Medicaid-insured patients in rates of $BRCA$ testing among patients with ovarian cancer. ⁴⁵	Disparities documented for SES in age at surgical resection for colorectal cancer. ⁴⁶

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Table 1

ASCVD, atherosclerotic cardiovascular disease; FH, familial hypercholesterolemia; HBOC, hereditary breast and ovarian cancer; IHC, immunohistochemistry; LDL, low-density lipoprotein; LDL-C, low-density lipoprotein cholesterol; LS, Lynch syndrome; MSI, microsatellite instability; SES, socioeconomic status.

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Table 2

Essential public health services, vision, and selected opportunities to help close the health equity gap in genomics and precision medicine

Continuum of Essential Public Health Services	Opportunities for Implementation of Emerging Genomic Applications Using a Health Equity Lens ^a
Population health assessment	Vision: Toward more precision in measuring disparities in diagnosis, prevention, treatment, and outcomes in emerging genomic applications in different communities
Public health surveillance	Integrate genomics and precision medicine into public health data modernization
Applied and implementation research	Develop and implement a robust community-specific implementation science agenda for genomics and precision medicine in underserved communities
Guidelines and policies	Vision: Toward effective engagement of different communities to drive needs assessment, evidence synthesis, policies, and guidelines
Evidence synthesis	Develop a process for continuous evidence synthesis and prioritization for implementation of tier 1 genomic applications in different communities
National and community-based guidelines	Develop evidence-driven models for coverage and reimbursement, data sharing and learning health systems for different communities
Assurance and capacity building	Vision: Toward achieving learning health systems for implementation of emerging genomic applications in different communities
Model community programs	Develop and implement exemplar pilot projects that facilitate building trust and access to evidence-based genomic applications in different communities
Tools and resources	Develop and validate patient- and community-specific tools and resources for implementation
Workforce development	Integrate genomics into clinical and public health workforce development and focus on diversity and inclusion in recruitment and retention of next-generation professionals

<u>a</u> ry wi ą 5 5 the main rocus of a puotic nearing genomics agained is on persons and communications negative, communities, uninsured or underinsured people, and those with lower education and income.