

No. 23-477

IN THE
Supreme Court of the United States

UNITED STATES OF AMERICA,
Petitioner,

v.

JONATHAN SKRMETTI, ET AL.,
Respondents.

On Writ of Certiorari to the United States
Court of Appeals for the Sixth Circuit

**BRIEF OF *AMICI CURIAE* AMERICAN
ACADEMY OF PEDIATRICS AND ADDITIONAL
NATIONAL AND STATE MEDICAL AND
MENTAL HEALTH ORGANIZATIONS IN
SUPPORT OF PETITIONER AND
RESPONDENTS IN SUPPORT OF PETITIONER**

Cortlin H. Lannin	D. Jean Veta
COVINGTON & BURLING LLP	<i>Counsel of Record</i>
Salesforce Tower	William R. Isasi
415 Mission Street	
Suite 5400	
San Francisco, CA 94105	

TABLE OF CONTENTS

TABLE OF AUTHORITIES..... iii

INTEREST OF

B. The Guidelines for Treating Gender Dysphoria Were Developed Through a Robust and Transparent Process, Employing the Same Scientific Rigor That Underpins Other Medical Guidelines.	15
Scientific Evidence Indicates the Effectiveness of Treating Gender Dysphoria According to the Guidelines.	17
III. The Sixth Circuit’s Opinion and the State Rely on Factually Inaccurate Claims and Ignore Recommendations of the Medical Community. .	21
IV. The Healthcare Ban Irreparably Harms Many Adolescents with Gender Dysphoria by Denying Them the Treatment They Need.	26
CONCLUSION	28

TABLE OF AUTHORITIES

	Page(s)
 Statutes	
Tenn. Code Ann. § 68-33-102(5)(B).....	3
Tenn. Code Ann. § 68-33-103(a).....	3
 Other Authorities	
Alexis A. Topjian et al., <i>Pediatric Basic and Advanced Life Support: 2020 American Heart Association Guidelines for Cardiopulmonary Resuscitation and Emergency Cardiovascular Care</i> , 142 <i>Circulation</i> S469 (2020).....	22
Alyson Sulaski Wyckoff, Am. Acad. of Pediatrics, <i>AAP Reaffirms Gender-Affirming Care Policy, Authorizes Systematic Review of Evidence to Guide Update</i> , AAP News (Aug. 4, 2023)	4
Am. Acad. of Pediatrics, <i>Policy Statement - Off-Label Use of Drugs in Children</i> , 133 <i>Pediatrics</i> 563 (2014)	24
Am. Psychiatric Ass'n, <i>Diagnostic and Statistical Manual of Mental Disorders: DSM-5-TR</i> (2022)	7

Florence Comite et al., <i>Short-Term Treatment of Idiopathic Precocious Puberty with a Long-Acting Analogue of Luteinizing Hormone-Releasing Hormone: A Preliminary Report</i> , 305(26) <i>New Eng. J. Med.</i> 1546 (1981)	13
Gordon H. Guyatt et al., <i>GRADE: An Emerging Consensus on Rating Quality of Evidence and Strength of Recommendations</i> , 336 <i>BMJ</i> 924 (2008)	15
Gordon H. Guyatt et al., <i>GRADE Guidelines: 1. Introduction - GRADE Evidence Profiles and Summary of Findings Tables</i> , 64 <i>J. Clinical Epidemiology</i> 383 (2011).....	15
Gordon H. Guyatt et al., <i>GRADE: What Is “Quality of Evidence” and Why Is It Important to Clinicians?</i> , 336 <i>BMJ</i> 995 (2008)	22
Hilary Cass, <i>Independent Review of Gender Identity Services for Children and Young People: Final Report</i> , Cass Review (Apr. 2024).....	23
Jack L. Turban et al., <i>Access to Gender-Affirming Hormones During Adolescence and Mental Health Outcomes Among Transgender Adults</i> , <i>J. 17(1) PLOS One</i> e0261039 (2022)	18, 27

- Jack L. Turban et al., *Pubertal Suppression for Transgender Youth and Risk of Suicidal Ideation*, 145 *Pediatrics* e20191725 (2020)..... 17, 19, 20
- James L. Madara, *AMA to States: Stop Interfering in Healthcare of Transgender Children*, *Am. Med. Ass'n* (Apr. 26, 2021).....6
- Jason Rafferty, *Ensuring Comprehensive Care and Support for Transgender and Gender-Diverse Children and Adolescents*, 142(4) *Pediatrics* e20182162 (2018) 4, 6, 7, 13, 14
- Jody L. Herman et al., *How Many Adults and Youth Identify as Transgender in the United States?*, *Williams Inst.* (2022).....6
- Ken C. Pang et al., *Long-Term Puberty Suppression for a Nonbinary Teenager*, 145(2) *Pediatrics* e20191606 (2019)..... 13
- Laura E. Kuper et al., *Body Dissatisfaction and Mental Health Outcomes of Youth on Gender-Affirming Hormone Therapy*, 145 *Pediatrics* e20193006 (2020)..... 17, 18

Luke R. Allen et al., <i>Well-Being and Suicidality Among Transgender Youth After Gender-Affirming Hormones</i> , 7 <i>Clinical Prac. Pediatric Psych.</i> 302 (2019)	18, 19
M. Hassan Murad et al., <i>Hormonal Therapy and Sex Reassignment: A Systematic Review and Meta-Analysis of Quality of Life and Psychosocial Outcomes</i> , 72 <i>Clin. Endocrinol.</i> 214 (2010)	27
Michael S. Irwig, <i>Detransition Among Transgender and Gender-Diverse People-An Increasing and Increasingly Complex Phenomenon</i> , <i>J. Clin. Endocrinol. & Metab.</i> 1 (June 2022)	26
National Academy of Sciences, <i>Clinical Practice Guidelines We Trust</i> (2011)	16
Polly Carmichael et al., <i>Short-Term Outcomes of Pubertal Suppression in a Selected Cohort of 12 to 15 Year Old Young People with Persistent Gender Dysphoria in the UK</i> , 16 <i>PLOS One</i> e0243894 (2021)	17
Richard J. Lilford & Jennifer Jackson, <i>Equipoise and the Ethics of Randomization</i> , 88 <i>J. R. Soc. Med.</i> 552 (1995)	23

- Rittakerttu Kaltiala et al., *Adolescent Development and Psychosocial Functioning After Starting Cross-Sex Hormones for Gender Dysphoria*, 74 *Nordic J. Psychiatry* 213 (2020) 18
- Rosalia Costa et al., *Psychological Support, Puberty Suppression, and Psychosocial Functioning in Adolescents with Gender Dysphoria*, 12(11) *J. Sexual Med.* 2206 (2015) 17
- Sari L. Reisner et al., *Advancing Methods for U.S. Transgender Health Research*, 23(2) *Curr. Opin. Endocrinol Diabetes Obes.* 198 (2016) 23
- Simona Martin et al., *Criminalization of Gender-Affirming Care-Interfering with Essential Treatment for Transgender Children and Adolescents*, 385(7) *New Eng. J. Med.* 579 (2021) 5, 12, 13, 14, 15, 17
- Stephen M. Rosenthal, *Challenges in the Care of Transgender and Gender-Diverse Youth: An Endocrinologist's View*, 17(10) *Nature Rev. Endocrinol.* 581 (2021) 20, 25

Stewart L. Adelson, *Practice Parameter on Gay, Lesbian, or Bisexual Sexual Orientation, Gender Nonconformity, and Gender Discordance in Children and Adolescents*, 51 *J. Am. Acad. Child & Adolescent Psychiatry* 957 (2012)..... 25

World Health Org., *International Classification of Diseases, Eleventh Revision (ICD-11)* (2019/2021)..... 7

Wylie C. Hembree et al., *Endocrine Treatment of Gender-Dysphoric/ Gender-Incongruent Persons: An Endocrine Society Clinical Practice Guideline*, 102 *J. Clin. Endocrinol. & Metab.* 3869 (2017)..... 8, 9, 10, 12, 13, 14, 15

Zoe Aldridge et al., *Long Term Effect of Gender-Affirming Hormone Treatment on Depression and Anxiety Symptoms in Transgender People: A Prospective Cohort Study*, 9 *Andrology* 1808 (2021) 19

INTEREST OF *AMICI CURIAE*

Amici curiae are the American Academy of Pediatrics, the Academic Pediatric Association, the American Academy of Child & Adolescent Psychiatry, the Association of American Medical Colleges, the American Academy of Family Physicians, the American Academy of Nursing, the American Association of Physicians for Human Rights, Inc. d/b/a GLMA: Health Professionals Advancing LGBTQ+ Equality, the American College of Obstetricians and Gynecologists, the American College of Osteopathic Pediatricians, the American College of Physicians, the American Medical Association, the American Pediatric Society, the American Psychiatric Association, the Association of Medical School Pediatric Department Chairs, Inc., the Endocrine Society, the National Association of Pediatric Nurse Practitioners, the Pediatric Endocrine Society, the Tennessee Chapter of the American Academy of Pediatrics, the Societies for Pediatric Urology, the Society for Adolescent Health and Medicine, the Society for Pediatric Research, the Society of Pediatric Nurses, and the World Professional Association for Transgender Health (collectively, “*amici*”).¹

Amici are professional medical and mental health organizations seeking to ensure that all adolescents, including those with gender dysphoria, receive the optimal medical and mental health care they need and

¹ Pursuant to Rule 37.6, *amici* affirm that no counsel for a party authored this brief in whole or in part and that no person other than *amici*, their staff, or their counsel made any monetary contributions intended to fund the preparation or submission of this brief.

deserve. *Amici* represent thousands of healthcare providers who have specific expertise with the issues raised in this brief. The Court should consider *amicus*'s brief because it provides important expertise and addresses misstatements about the treatment for transgender adolescents.

SUMMARY OF ARGUMENT

On March 23, 2023, the Governor of Tennessee signed S.B. 1 into law (the “Healthcare Ban”). The Healthcare Ban prohibits healthcare providers from providing patients under the age of 18 with critical, medically necessary, evidence-based care for gender dysphoria.² Denying such evidence-based medical care to adolescents who meet the requisite medical criteria puts them at risk of significant harm. Below, *amici* provide the Court with an accurate description of the relevant treatment guidelines and summarize the scientific evidence supporting the gender-affirming medical care for adolescents that is prohibited by the Healthcare Ban.³

Gender dysphoria is a condition that is characterized by clinically significant distress or impairment in social, occupational, or other important areas of functioning due to a marked incongruence between the patient’s gender identity (i.e., the innate sense of oneself as being a particular gender) and sex

² Tenn. Code Ann. §§ 68-33-102(5)(B), 68-33-103(a) prohibits medical treatments that are administered for the purpose of

identity. Empirical evidence indicates that gender-affirming care, including the prescription of puberty blockers and hormone therapy to carefully evaluated patients who meet diagnostic criteria, can alleviate clinically significant distress and lead to significant improvements in the mental health and overall well-being of adolescents with gender dysphoria.

ErJ0-31Tf12-7.29.205 TD.00-02 T5.0012527[(impT

harms adolescents with gender dysphoria by denying crucial care to those who need it.

I. Understanding Gender Identity and Gender Dysphoria

Gender identity refers to a person's deep internal sense of belonging to a particular gender.⁸ Most people have a gender identity that aligns with their sex assigned at birth.⁹ However, transgender people have a gender identity that does not align with their sex assigned at birth.¹⁰ In the United States, approximately 1.6 million individuals identify as transgender.¹¹ Of these individuals, approximately 10% are teenagers aged 13 to 17.¹² Individuals often start to understand their gender identity during prepubertal childhood and adolescence.

Today, there is an increasing acceptance of being transgender as a normal variation of human identity.¹³ However, many transgender people suffer

⁸ AAP Policy Statement, *supra* note 4, at 2 tbl.1.

⁹ See Am. Psychological Ass'n, *Guidelines for Psychological Practice with Transgender and Gender Nonconforming People*, 70(9) *Am. Psychologist* 832, 862 (2015), <https://perma.cc/6VQK-S3F8>. Such individuals are referred to as "cisgender." See *id.* at 861.

¹⁰ See *id.* at 832.

¹¹ See Jody L. Herman et al., *How Many Adults and Youth Identify as Transgender in the United States?* at 2, Williams Inst. (2022), <https://perma.cc/UCB4-346Z>.

¹² See *id.* at 3.

¹³ James L. Madara, *AMA to States: Stop Interfering in Healthcare of Transgender Children*, *Am. Med. Ass'n* (Apr. 26, 2021), <https://perma.cc/BKS6-QFQ8>; see also *Am. Psychological*

from gender dysphoria, a serious medical condition in which the patient experiences significant distress that can lead to “impairment in peer and/or family relationships, school perfor

II. The Widely Accepted Guidelines for Treating Adolescents with Gender Dysphoria Provide for Gender-Affirming Medical Care When Indicated.

The widely accepted view of the professional medical community is that gender-affirming care is the appropriate treatment for gender dysphoria and that, for some adolescents, puberty blockers and hormone therapy are necessary.¹⁸ Gender-affirming care greatly reduces the negative physical and mental health consequences that result when gender dysphoria is untreated.¹⁹

A. The Gender Dysphoria Treatment Guidelines Include Thorough Mental Health Assessments and, for Some Adolescents, Gender-Affirming Medical Care.

The treatment protocols for gender dysphoria are laid out in established, evidence-based clinical guidelines: (i) the Endocrine Society Clinical Practice Guideline for Endocrine Treatment of Gender-Dysphoric/Gender-Incongruent Persons, and (ii) the WPATH Standards of Care for the Health of Transgender and Gender Diverse People (together, the “Guidelines”).²⁰ The Guidelines have been

developed by expert clinicians and researchers who have worked with patients with gender dysphoria for many years.

The Guidelines provide that all youth with gender dysphoria should be evaluated, diagnosed, and treated by a qualified health care professional (“HCP”).²¹ Further, the Guidelines provide that each patient who receives gender-affirming care should receive only medically necessary and appropriate care that is tailored to the patient’s individual needs and that is based on the best evidence possible along with clinical experience.²²

1. The Guidelines Do Not Recommend Gender-Affirming Medical Care for Prepubertal Children.

For prepubertal children with gender dysphoria, the Guidelines provide for mental health care and support for the child and their family, such as through psychotherapy and social transitioning.²³ The

Clinical Practice Guideline, 102 *J. Clin. Endocrinol. & Metab.* 3869 (2017) [hereinafter, “Endocrine Soc’y Guidelines”], <https://perma.cc/3L9J-428B>; Eli Coleman et al., *Standards of Care for the Health of Transgender and Gender Diverse People*, 23 *Int’l J. Transgender Health S1* (8th ed. 2022) [hereinafter “WPATH Guidelines”], <https://perma.cc/7SU3-RPK9>.

²¹ See *infra* Section II.A.2.

²² See WPATH Guidelines, *supra* note 20, at S16–S18; *Endocrine Soc’y Guidelines*, *supra* note 20, at 3872–73.

²³ See WPATH Guidelines, *supra* note 20, at S73–S76; *Endocrine Soc’y Guidelines*, *supra* note 20, at 3877–78. “Social transition” refers to a process by which a child is acknowledged by others and has the opportunity to live publicly, either in all situations

relevant to gender diverse children, adolescents, and families.²⁵

Prior to developing a treatment plan, the HCP should conduct a robust diagnostic assessment—specifically, a “comprehensive biopsychosocial assessment”—of the adolescent patient.²⁶ The HCP conducts this assessment to “understand the adolescent’s strengths, vulnerabilities, diagnostic profile, and unique needs,” so that the resulting treatment plan is appropriately individualized.²⁷ This assessment must be conducted collaboratively with the patient and their caregiver(s).²⁸

the diagnostic criteria of gender dysphoria or gender incongruence according to an established taxonomy;²⁹ (2) the adolescent has demonstrated a sustained and persistent pattern of gender nonconformity or gender dysphoria; (3) the adolescent has demonstrated the emotional and cognitive maturity required to provide informed consent for treatment; (4) any coexisting psychological, medical, or social problems that could interfere with

characteristics—which may result in significant distress for transgender youth—until adolescents are old enough and have had sufficient time to make more informed decisions about whether to pursue further treatments.³³ Puberty blockers also can make pursuing transition later in life easier, because they prevent irreversible bodily changes such as protrusion of the Adam’s apple or breast growth.³⁴

Later in adolescence—and if the criteria below are met—hormone therapy may be used to initiate puberty consistent with the patient’s gender identity.³⁹ Hormone therapy involves using gender-affirming hormones to allow adolescents to develop secondary sex characteristics consistent with their gender identity.⁴⁰ Hormone therapy is only prescribed when a qualified mental health professional has confirmed the persistence of the patient’s gender dysphoria, the patient’s mental capacity to consent to the treatment, and that any coexisting problems have been addressed.⁴¹ A pediatric endocrinologist or other clinician experienced in pubertal induction must also agree with the indication, and the patient and their parents must be informed of the potential effects and side effects and give their informed consent.⁴² Although some of the changes caused by hormone therapy become irreversible after those secondary sex characteristics are fully developed, others are partially reversible if the patient discontinues use of the hormones.⁴³

The Guidelines contemplate that the prescription of puberty blockers and/or hormone therapy be coupled with education on the safe use of such medications and close monitoring to mitigate any potential risks.⁴⁴ Decisions regarding the appropriate

³⁹ Martin et al., *supra* note 7, at 2.

⁴⁰ See AAP Policy Statement, *supra* note 4, at 6.

⁴¹ Endocrine Soc’86eiatrndoc’86eiate2o(o)-2.0006 3.0008c’8 e4c45T4c45T8to se25 0 TD0 (o)-t32[(so

treatment for each patient with gender dysphoria are made in consultation with the patient, their parents, and the medical and mental health care team. There is “no one-size-fits-all approach to this kind of care.”⁴⁵

B. The Guidelines for Treating Gender Dysphoria Were Developed Through a Robust and Transparent Process, Employing the Same Scientific Rigor That Underpins Other Medical Guidelines.

The Guidelines are the product of careful and robust deliberation following the same types of processes—and subject to the same types of rigorous requirements—as other guidelines promulgated by *amici* and other medical organizations.

s 8t to

independent groups of professionals.⁴⁸ Reviewers were subject to strict conflict of interest rules, and there was ample opportunity for feedback and debate through the years-long review process.⁴⁹ Further, the Endocrine Society continually reviews its own guidelines and recently determined that the 2017 transgender care guidelines continue to reflect the best, most up-to-date available evidence.⁵⁰

First published in 1979, the WPATH Standards of Care are currently in their 8th Edition. The current Standards of Care are the result of a robust drafting, comment, and review process that collectively took five years.⁵¹ The draft guidelines went through rigorous review and were publicly available for discussion and debate, receiving a total of 2,688 comments.⁵² There were 119 authors ultimately involved in the final draft, including feedback from experts in the field as well as from transgender individuals and their families.⁵³

C. Scientific Evidence Indicates the Effectiveness of Treating Gender Dysphoria According to the Guidelines.

Multiple studies indicate that adolescents with gender dysphoria who receive gender-affirming medical care experience improvements in their overall well-being.⁵⁴ A number of studies have been published that investigated the use of puberty blockers on adolescents with gender dysphoria⁵⁵

⁵⁴ See Martin et al., *supra* note 7, at 580.

⁵⁵ See, e.g., Christal Achille et al., *Longitudinal Impact of Gender-Affirming Endocrine Intervention on the Mental Health and Well-Being of Transgender Youths: Preliminary Results*, 8 Int'l J. Pediatric Endocrinol. 1, 1–5 (2020), <https://perma.cc/K5SR-EE3G>; Polly Carmichael et al., *Short-Term Outcomes of Pubertal Suppression in a Selected Cohort of 12 to 15 Year Old Young People with Persistent Gender Dysphoria in the UK*, 16 PLOS One e0243894 (2021), <https://doi.org/10.1371/journal.pone.0243894>; Rosalia Costa et al., *Psychological Support, Puberty Suppression, and Psychosocial Functioning in Adolescents with Gender Dysphoria*, 12(11) J. Sexual Med. 2206–14 (2015), <https://pubmed.ncbi.nlm.nih.gov/26556015/>; Annelou L.C. de

including statistically significant reductions in anxiety, depression, and suicidal ideation.⁵⁷

For example, a longitudinal study of nearly 50 transgender adolescents found that suicidality was decreased by a statistically significant degree after receiving gender-affirming hormone treatment.⁵⁸ A study published in January 2023, following 315 participants age 12 to 20 who received gender-affirming hormone treatment, found that the treatment was associated with decreased symptoms of depression and anxiety.⁵⁹ Additionally, a 2020 study analyzed survey data from 89 transgender adults who had access to puberty blockers while adolescents and from more than 3,400 transgender adults who did not.⁶⁰ The study found that those who received puberty blocking treatment had lower odds of lifetime suicidal ideation than those who wanted puberty blocking treatment but did not receive it, even after adjusting for demographic variables and level of family support.⁶¹ Approximately *nine in ten* transgender adults who wanted puberty blocking

⁵⁷ The data likewise indicate that adults who receive gender-affirming care experience positive mental health outcomes. See, e.g., Zoe Aldridge et al., *Long Term Effect of Gender-Affirming Hormone Treatment on Depression and Anxiety Symptoms in Transgender People: A Prospective Cohort Study*, 9 *Andrology* 1808–16 (2021), <https://perma.cc/543U-HL5P>.

⁵⁸ See Allen et al., *supra* note 56.

⁵⁹ See Chen et al., *supra* note 56.

⁶⁰ See Turban et al., *Pubertal Suppression for Transgender Youth and Risk of Suicidal Ideation*, *supra* note 55.

⁶¹ See *id.*

treatment but did not receive it reported lifetime suicidal ideation.⁶²

As another example, a prospective two-year follow-up study of adolescents with gender dysphoria published in 2011 found that treatment with puberty blockers was associated with decreased depression and improved overall functioning.⁶³ A six-year follow-up study of 55 individuals from the 2011 study found that subsequent treatment with hormone therapy followed by surgery in adulthood was associated with a statistically significant decrease in depression and anxiety.⁶⁴ “Remarkably, this study demonstrated that these transgender adolescents and young adults had a sense of well-being that was equivalent or superior to that seen in age-matched controls from the general population.”⁶⁵

As clinicians and scientific researchers, *amici* always welcome more research, including on this crucial topic. However, the available data indicate that the gender-affirming medical care prohibited by the Healthcare Ban is effective for the treatment of gender dysphoria.

⁶² *See id.*

⁶³ *See de Vries et al., Puberty Suppression in Adolescents with Gender Identity Disorder: A Prospective Follow-Up Study*

III. The Sixth Circuit’s Opinion and the State Rely on Factually Inaccurate Claims and Ignore Recommendations of the Medical Community.

Both the Sixth Circuit and the State make several inaccurate assertions regarding gender-affirming medical care that are inconsistent with the existing evidence and standard practices in medicine. Contrary to the State’s arguments, (1) the Guidelines’ reliance on evidence is consistent with international best practices and the development of standards of care for treating adolescents in other medical contexts, and (2) the Guidelines already carefully address the safety of gender-affirming medical care and provide recommendations for the mitigation of any potential risks, as is typical in every area of medicine.

First, the State of Tennessee posits that the use of puberty blockers and hormone therapy to treat gender dysphoria in adolescents is “experimental in nature and not supported by high-quality, long-term medical studies.”⁶⁶ To conclude that these treatments are not supported by evidence ignores a significant body of research.

When Respondents criticize the studies supporting gender-affirming medical care,⁶⁷ they refer

⁶⁶ Pet. App. 10a.

⁶⁷ While the Respondents broadly criticize the evidence supporting the Guidelines, the reports they rely upon do not evaluate, nor do they purport to evaluate, all of the evidence supporting the Guidelines. C.A. Dkt. 64, at 14 (citing to declarations discussing the 2020 NICE studies).

clinicians rely on the best evidence possible and clinical experience to provide treatment for their patients. The evidence supporting gender-affirming medical care is consistent with the type of evidence relied on in other clinical practices throughout the medical community.⁷²

In addition, the Sixth Circuit appears to conflate the use of FDA-approved medications for off-label uses with the use of *non*-FDA-approved medications.⁷³ According to the FDA, “once the FDA approves a drug,

acceptable.” Sari L. Reisner et al., *Advancing Methods for U.S. Transgender Health Research*, 23(2) *Curr. Opin. Endocrinol Diabetes Obes.* 198, 199 (2016). With preexisting guidelines that recommend gender-affirming care for those with gender dysphoria, randomized controlled trials would violate the principle of equipoise, which safeguards the rights of individual trial participants. Richard J. Lilford & Jennifer Jackson, *Equipoise and the Ethics of Randomization*, 88 *J. R. Soc. Med.* 552, 552 (1995). Moreover, the ability to perform RCTs is complicated where participants may be able to easily discern trial placement due to biological changes from treatment.

⁷² *Amici* also are aware of the systematic review conducted by Hilary Cass which was submitted to NHS England. See Hilary Cass, *Independent Review of Gender Identity Services for Children and Young People: Final Report*, Cass Review (Apr. 2024), <https://perma.cc/A8UR-Q2WD> (the “Cass Review”). However, the Cass Review, like other systematic reviews, is simply a summary of some of the existing research, as selected by the author. It does not purport to offer any new studies or findings regarding the efficacy and safety of prescribing gender-affirming medical care for adolescents that conflict with the recommendations in the Guidelines, which themselves are based on the available existing studies and research, as well as clinical experience. See Endocrine Soc’y, *Endocrine Society Statement in Support of Gender-Affirming Care* (May 8, 2024), <https://perma.cc/J4Y2-RUJ2>.

⁷³ Pet. App. 21a.

healthcare providers generally may prescribe the drug for an unapproved use when they judge that it is medically appropriate for their patient.”⁷⁴ Such off-label drug use is common, particularly in disciplines such as pediatrics, where patients are less likely to be included in clinical trials.⁷⁵ “[I]n no way does a lack of labeling signify that therapy is unsupported by clinical experience or data in children.”⁷⁶ The FDA does not regulate the practice of medicine, and a lack of labeling should not be confused with a finding of contraindication or unsafety.⁷⁷ Medications used in providing gender-affirming care have been approved by the FDA. To suggest that using those medications to treat gender dysphoria is *disapproved*, or *should be prohibited* based on a lack of approval, is incorrect and contradicts common medical practices.

Second, the State claims that “the risks likely outweigh the benefits” of prescribing puberty blockers and hormone therapy to transgender adolescents.⁷⁸ *Amici* disagree, and observe that the State’s view is inconsistent with the existing research. This research, as well as the clinical experience of members of the *amici* who specialize in treating gender

⁷⁴ FDA, *Understanding Unapproved Use of Approved Drugs “Off Label”* (Feb. 5, 2018), <https://perma.cc/YW48-NZJS>.

⁷⁵ See, e.g., Christopher M. Wittich et al., *Ten Common Questions (and Their Answers) About Off-Label Drug Use*, 87(10) *Mayo Clin. Proc.* 982, 983 (2012) (reporting on study finding “78.9% of children discharged from pediatric hospitals were

dysphoria show that the use of these medications to treat adolescents is both safe and effective in

varies from study to study

less depression, anxiety, and suicidal ideation.

CONCLUSION

For the foregoing reasons, this Court should vacate the decision below and remand for the application of heightened scrutiny, or reverse the judgment.