

8. Naidoo RN. Occupational exposures and chronic obstructive pulmonary disease: incontrovertible evidence for causality? *Am J Respir Crit Care Med* 2012;185:1252–1254.
9. Darby AC, Waterhouse JC, Stevens V, Billings CG, Billings CG, Burton CM, Young C, Wight J, Blanc PD, Fishwick D. Chronic obstructive pulmonary disease among residents of an historically industrialised area. *Thorax* 2012;67:901–907.
10. Messing K, Punnett L, Bond M, Alexanderson K, Pyle J, Zahm S, Wegman D, Stock SR, de Grosbois S. Be the fairest of them all: challenges and recommendations for the treatment of gender in occupational health research. *Am J Ind Med* 2003;43:618–629.
11. Couper D, LaVange LM, Han M, Barr RG, Bleecker E, Hoffman EA, Kanner R, Kleerup E, Martinez FJ, Woodruff PG, *et al.*; SPIROMICS Research Group. Design of the Subpopulations and Intermediate Outcomes in COPD Study (SPIROMICS). *Thorax* 2014;69:491–494.
12. Regan EA, Hokanson JE, Murphy JR, Make B, Lynch DA, Beaty TH, Curran-Everett D, Silverman EK, Crapo JD. Genetic epidemiology of COPD (COPDGene) study design. *COPD* 2010;7:32–43.
13. Marchetti N, Garshick E, Kinney GL, McKenzie A, Stinson D, Lutz SM, Lynch DA, Criner GJ, Silverman EK, Crapo JD; COPDGene Investigators. Association between occupational exposure and lung function, respiratory symptoms, and high-resolution computed tomography imaging in COPDGene. *Am J Respir Crit Care Med* 2014;190:756–762.
14. Martinez CH, Kim V, Chen Y, Kazerooni EA, Murray S, Criner GJ, Curtis JL, Regan EA, Wan E, Hersh CP, *et al.*; COPDGene Investigators. The clinical impact of non-obstructive chronic bronchitis in current and former smokers. *Respir Med* 2014;108:491–499.
15. Han MK, Agustí A, Calverley PM, Celli BR, Criner G, Curtis JL, Fabbri LM, Goldin JG, Jones PW, Macnee W, *et al.* Chronic obstructive pulmonary disease phenotypes: the future of COPD. *Am J Respir Crit Care Med* 2010;182:598–604.
16. de Jong K, Boezen HM, Kromhout H, Vermeulen R, Vonk JM, Postma DS; LifeLines Cohort Study. Occupational exposure to vapors, gases, dusts, and fumes is associated with small airways obstruction. *Am J Respir Crit Care Med* 2014;189:487–490.
17. Dijkstra AE, de Jong K, Boezen HM, Kromhout H, Vermeulen R, Groen HJ, Postma DS, Vonk JM. Risk factors for chronic mucus hypersecretion in individuals with and without COPD: influence of smoking and job exposure on CMH. *Occup Environ Med* 2014;71:346–352.
18. Doney B, Hnizdo E, Graziani M, Kullman G, Burchfiel C, Baron S, Fujishiro K, Enright P, Hankinson JL, Stukovsky KH, *et al.* Occupational risk factors for COPD phenotypes in the Multi-Ethnic Study of Atherosclerosis (MESA) Lung Study. *COPD* 2014;11:368–380.
19. Caillaud D, Lemoigne F, Carré P, Escamilla R, Chanez P, Burgel PR, Court-Fortune I, Jebrak G, Pinet C, Perez T, *et al.*; initiative BPCO Scientific Committee. Association between occupational exposure and the clinical characteristics of COPD. *BMC Public Health* 2012;12:302.
20. Kuschner WG, Hegde S, Agrawal M. Occupational history quality in patients with newly documented, clinician-diagnosed chronic bronchitis. *Chest* 2009;135:378–383.
21. Lam KB, Yin P, Jiang CQ, Zhang WS, Adab P, Miller MR, Thomas GN, Ayres JG, Lam TH, Cheng KK. Past dust and GAS/FUME exposure and COPD in Chinese: the Guangzhou Biobank Cohort Study. *Respir Med* 2012;106:1421–1428.

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Scientific Evidence Supports Stronger Limits on Ozone

In 2007, 2010, and now again in 2015, the American Thoracic Society has recommended that the U.S. Environmental Protection Agency (EPA) adopt an 8-hour ozone national ambient air quality standard of 60 ppb to adequately protect public health (1, 2). Although the recommended standard endorsed by the American Thoracic Society has not changed during this time, the scientific evidence supporting this recommendation has significantly strengthened. The scientific evidence available 7 years ago justifying this recommendation has been supplemented by an even greater understanding of the health effects of ozone exposures, including infant respiratory problems, worse childhood asthma control, reduced lung function, and increased mortality in adults.

On November 25, 2014, the EPA proposed a standard in the range of 65–70 ppb, which is lower than the current standard of 75 ppb (the standard is defined as the annual fourth highest maximum daily 8-hour ozone average averaged over 3 years). Although we applaud the EPA for proposing a stricter standard, we believe the scientific evidence clearly calls for a standard of 60 ppb to protect human health. We are currently in the public comment period for the proposed ozone rule and urge the EPA to issue a more protective standard of 60 ppb. This is the second time the Obama Administration has reviewed the current ozone standard of 75 ppb. The previous administration established the current standard outside the range recommended by the Clean Air Science Advisory

Committee of 60–70 ppb (3). In 2010, the Clean Air Science Advisory Committee reaffirmed its initial recommendation as part of an early reassessment of the ozone standard, an effort that was ultimately abandoned in 2011 (4). Because a new science assessment was not conducted as part of that review, the current review of the ozone standard is the first to consider new scientific evidence since 2006.

Since 2006, much more evidence has accumulated that ozone exposures in the range of 60–75 ppb have adverse physiologic effects across the entire age spectrum, from infants to older adults. Although there is also some evidence of health effects of ozone exposure below 60 ppb, the strongest evidence supports the conclusion that serious adverse health effects occur across all ages at levels above 60 ppb.

Highlights of this new body of evidence include a study of emergency department visits among children aged 0 to 4 years in Atlanta, Georgia, which found that each 30-ppb increase in the 3-day average of ozone was associated with an 8% higher risk of pneumonia and a 4% higher risk for upper respiratory infection (5). Several studies have demonstrated dose–response relationships between ozone exposure and childhood asthma admissions at exposure levels in the 60–80 ppb range (6–9). Similar associations have been found for adult admissions for asthma (9–11) and chronic obstructive pulmonary disease (12, 13). A population-based cohort study of generally healthy adults found that FEV₁ was 56 ml lower after days

when ambient ozone ranged from 59 to 75 ppb compared with days with levels lower than 59 ppb (14). Controlled human exposure studies have reaffirmed lung function decrements in healthy adults after exposure to 60–70 ppb of ozone (15, 16). Perhaps of greatest concern, there is now stronger evidence of increased mortality in association with ozone (17–19), particularly among the elderly and those with chronic disease (20, 21).

In making this recommendation, we acknowledge that challenges exist in reducing ambient ozone concentrations. Unlike other pollutants regulated by the Clean Air Act, ozone is a secondary pollutant formed from precursor pollutants through photochemical reactions in the atmosphere. This presents challenges in designing successful abatement plans. For example, the natural presence of precursor chemicals and long-range transport of ozone from beyond U.S. jurisdictional boundaries can each contribute to background ozone levels (22). However, the adverse health effects of ozone do not discriminate on the basis of the source of precursor pollutants, nor do these complexities change the reality that serious adverse health effects occur at concentrations higher than 60 ppb. Although the science surrounding background ozone is still emerging, the evidence of adverse health risks of ozone is clear.

Another challenge for ozone management is that the secondary formation of ozone can result in higher concentrations downwind from the primary sources of precursor pollutants, cutting across jurisdictions. As a result, integrated planning across jurisdictional boundaries and compliance with the “good neighbor” provision of the Clean Air Act will be necessary to reduce regional ozone to levels that adequately protect public health (23).

Although controlling ambient ozone will be a challenge, it will also present opportunities for innovation and leadership. State Implementation Plans will vary in how counties across the United States plan to remediate unhealthy levels of ozone. Although all plans will take time to achieve the needed results, the best approaches will be identified and serve as a model that other regions can follow to protect public health.

The timing is excellent for revision of the ozone standard to 60 ppb, as the new standard can build on substantial recent progress. In March 2014, the EPA finalized new standards for motor vehicle emissions and cleaner fuels that are expected to reduce ozone levels significantly (24). Already-adopted revisions to standards for particulate matter and air toxics are also likely to further reduce ozone formation (25). Above all, however, we are entering an era of technological innovation, infrastructure reconstruction, and commitment to sustainability in which obsolete technologies are being replaced by more efficient and less-polluting innovations. This is exactly the right time to lay down the correct performance criteria and design specifications for the new technology before we commit to a new energy and transportation regime that could limit our choices in the future.

The U.S. EPA has taken significant actions in the past when justified by scientific evidence, most notably by reducing the permissible concentrations of airborne lead by 90% in 2008 and reducing the permissible annual concentrations of fine particle pollution by 20% in 2013 (26). We encourage

the EPA administrator, with the full support of the president, to acknowledge the large body of scientific evidence documenting the harms caused by ozone pollution and set a standard of 60 ppb to protect the health of the American public. ■

Author disclosures are available with the text of this article at www.atsjournals.org.

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References

1. Dey R, Van Winkle L, Ewart G, Balmes J, Pinkerton K; ATS Environmental Health Policy Committee. A second chance: setting a protective ozone standard. *Am J Respir Crit Care Med* 2010;181:297–299.
2. Pinkerton KE, Balmes JR, Fanucchi MV, Rom WN. Ozone, a malady for all ages. *Am J Respir Crit Care Med* 2007;176:107–108.
3. US Environmental Protection Agency. Clean Air Scientific Advisory Committee recommendations concerning the final rule for the national ambient air quality standards for ozone. Washington, DC: U.S. Environmental Protection Agency; 2008 [accessed 2015 Feb 6]. Available from: <http://nepis.epa.gov/Exe/ZyPDF.cgi/P1000JY2.PDF?Dockey=P1000JY2.PDF>
4. US Environmental Protection Agency. Review of EPA's proposed ozone national ambient air quality standard. Washington, DC: US Environmental Protection Agency; 2010 [accessed 2015 Feb 6]. Available from: [http://yosemite.epa.gov/sab/sabproduct.nsf/610BB57CFAC8A41C852576CF007076BD/\\$File/EPA-CASAC-10-007-unsigned.pdf](http://yosemite.epa.gov/sab/sabproduct.nsf/610BB57CFAC8A41C852576CF007076BD/$File/EPA-CASAC-10-007-unsigned.pdf)
5. Darrow LA, Klein M, Flanders WD, Mulholland JA, Tolbert PE, Strickland MJ. Air pollution and acute respiratory infections among children 0–4 years of age: an 18-year time-series study. *Am J Epidemiol* 2014;180:968–977.
6. Strickland MJ, Klein M, Flanders WD, Chang HH, Mulholland JA, Tolbert PE, Darrow LA. Modification of the effect of ambient air pollution on pediatric asthma emergency visits: susceptible subpopulations. *Epidemiology* 2014;25:843–850.
7. Strickland MJ, Darrow LA, Klein M, Flanders WD, Sarnat JA, Waller LA, Sarnat SE, Mulholland JA, Tolbert PE. Short-term associations between ambient air pollutants and pediatric asthma emergency department visits. *Am J Respir Crit Care Med* 2010;182:307–316.
8. Gleason JA, Bielory L, Fagliano JA. Associations between ozone, PM_{2.5}, and four pollen types on emergency department pediatric asthma events during the warm season in New Jersey: a case-crossover study. *Environ Res* 2014;132:421–429.
9. Silverman RA, Ito K. Age-related association of fine particles and ozone with severe acute asthma in New York City. *J Allergy Clin Immunol* 2010;125:367–373.
10. Glad JA, Brink LL, Talbott EO, Lee PC, Xu X, Saul M, Rager J. The relationship of ambient ozone and PM_{2.5} levels and asthma

- emergency department visits: possible influence of gender and ethnicity. *Arch Environ Occup Health* 2012;67:103–108.
11. Meng YY, Rull RP, Wilhelm M, Lombardi C, Balmes J, Ritz B. Outdoor air pollution and uncontrolled asthma in the San Joaquin Valley, California. *J Epidemiol Community Health* 2010;64:142–147.
 12. Ko FWS, Hui DSC. Air pollution and chronic obstructive pulmonary disease. *Respirology* 2012;17:395–401.
 13. Medina-Ramón M, Zanobetti A, Schwartz J. The effect of ozone and PM10 on hospital admissions for pneumonia and chronic obstructive pulmonary disease: a national multicity study. *Am J Epidemiol* 2006;163:579–588.
 14. Rice MB, Ljungman PL, Wilker EH, Gold DR, Schwartz JD, Koutrakis P, Washko GR, O'Connor GT, Mittleman MA. Short-term exposure to air pollution and lung function in the Framingham Heart Study. *Am J Respir Crit Care Med* 2013;188:1351–1357.
 15. Schelegle ES, Morales CA, Walby WF, Marion S, Allen RP. 6.6-hour inhalation of ozone concentrations from 60 to 87 parts per billion in healthy humans. *Am J Respir Crit Care Med* 2009;180:265–272.
 16. Kim CS, Alexis NE, Rappold AG, Kehrl H, Hazucha MJ, Lay JC, Schmitt MT, Case M, Devlin RB, Peden DB, et al. Lung function and inflammatory responses in healthy young adults exposed to 0.06 ppm ozone for 6.6 hours. *Am J Respir Crit Care Med* 2011;183:1215–1221.
 17. Peng RD, Samoli E, Pham L, Dominici F, Touloumi G, Ramsay T, Burnett RT, Krewski D, Le Tertre A, Cohen A, et al. Acute effects of ambient ozone on mortality in Europe and North America: results from the APHENA study. *Air Qual Atmos Health* 2013;6:445–453.
 18. Romieu I, Gouveia N, Cifuentes LA, de Leon AP, Junger W, Vera J, Strappa V, Hurtado-Díaz M, Miranda-Soberanis V, Rojas-Bracho L, et al.; HEI Health Review Committee. Multicity study of air pollution and mortality in Latin America (the ESCALA study). *Res Rep Health Eff Inst* 2012;5–86.
 19. Zanobetti A, Schwartz J. Mortality displacement in the association of ozone with mortality: an analysis of 48 cities in the United States. *Am J Respir Crit Care Med* 2008;177:184–189.
 20. Medina-Ramón M, Schwartz J. Who is more vulnerable to die from ozone air pollution? *Epidemiology* 2008;19:672–679.
 21. Zanobetti A, Schwartz J. Ozone and survival in four cohorts with potentially predisposing diseases. *Am J Respir Crit Care Med* 2011;184:836–841.
 22. US Environmental Protection Agency. Policy assessment for the review of the ozone national ambient air quality standards. Washington, DC: U.S. Environmental Protection Agency; 2014 [accessed 2015 Feb 6]. Available from: <http://www.epa.gov/ttn/naaqs/standards/ozone/data/20140829pa.pdf>
 23. Supreme Court of the United States. Environmental Protection Agency et al. v. Eme Homer City Generation, L.P., et al. 2013 [accessed 2015 Feb 6]. Available from: http://www.supremecourt.gov/opinions/13pdf/12-1182_553a.pdf
 24. U.S. Environmental Protection Agency. Control of air pollution from motor vehicles: tier 3 motor vehicle emission and fuel standards; final rule. *Federal Register* Vol. 79, No. 81. 2014 [accessed 2015 Feb 6]. Available from: <http://www.gpo.gov/fdsys/pkg/FR-2014-04-28/pdf/2014-06954.pdf>
 25. U.S. Environmental Protection Agency. Reconsideration of certain new source issues: national emission standards for hazardous air pollutants from coal- and oil-fired electric utility steam generating units and standards of performance for fossil-fuel-fired electric utility, industrial-commercial-institutional, and small industrial-commercial-institutional steam generating units. 2013 [accessed 2015 Feb 6]. Available from: <http://www.gpo.gov/fdsys/pkg/FR-2013-04-24/pdf/2013-07859.pdf>
 26. US Environmental Protection Agency. National ambient air quality standards for lead. *Federal Register* Vol. 73, No. 219. 2008 [accessed 2015 Feb 6]. Available from: <http://www.gpo.gov/fdsys/pkg/FR-2008-11-12/pdf/E8-25654.pdf>

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