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A View of the Literature

The most influential articles in critical care medicine[☆]

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Citation classics;
Landmark articles

Abstract

Purpose: The study aimed to examine query strategies that would provide an exhaustive search method to retrieve the most referenced articles within specific categories of critical care.

Material and Methods: A comprehensive list of the most cited critical care medicine articles was generated by searching the Science Citation Index Expanded data set using general critical care terms keywords such as “critical care,” critical care journal titles, and keywords for subsubjects of critical care.

Results: The final database included 1187 articles published between 1905 and 2006. The most cited article was referenced 4909 times. The most productive search term was *intensive care*. However, this term only retrieved 25% of the top 100 articles. Furthermore, 662 of the top 1000 articles could not be found using any of the basic critical care search terms. Sepsis, acute lung injury, and mechanical ventilation were the most common areas of focus for the articles retrieved.

Conclusion: Retrieving frequently cited, influential articles in critical care requires using multiple search terms and manuscript sources. Periodic compilations of most cited articles may be useful for critical care practitioners and researchers to keep abreast of important information.

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1. Introduction

Modern methods to search for important medial literature have recently flourished with the dissemination and near ubiquity of Internet-based search engines (eg, PubMed, MEDLINE, OVID). Articles that have been cited most often, referred to as “citation classics,” are frequently important sources of both detailed information that has changed clinical/research practices as well as of useful connectors to other relevant literature that cites these articles. Finding landmarks or classic articles still requires an understanding of the literature and how articles are referenced. Different

databases may categorize disciplines and journals from the same clinical areas under different terms and yield different results when using the same search term.

These referencing differences as well as other conceptual arguments have led to some debate regarding attributing the number of times an article is cited to its quality [1-4]. However, there is general agreement that the number of times an article is cited does reflect the “impact of that article on the scientific market” [5].

Garfield [6] first published citation classics of the *Journal of the American Medical Association* in 1987. Similar studies (citations classic or impact factors) have been published with respect to various clinical specialties [7-15]. In 2004, Baltussen and Kindler [16] published citation classics of the critical care literature and enumerated the 71 top-cited articles in critical care journals and the 45 top-cited critical care articles in non-critical care journals. This article was somewhat limited by the number of citations referenced as

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well as the absence of specific critical care topic citations groups. The first and primary aim of this study was to expand on Baltussen's earlier work and provide an updated and more comprehensive catalogue of the most influential articles pertaining to critical care medicine and *within* specific categories of critical care, something not done in previous studies of this type. We also examined the methods used to retrieve articles to provide an exhaustive search method that future researchers may use in modern electronic databases to find frequently cited critical care articles.

2. Methods

2.1. Creation of master comprehensive list

To develop the most comprehensive master list of most often cited critical care medicine-related articles, we first focused on querying the ISI Web of Knowledge (2008 Thompson Corporation) using the Science Citation Index Expanded (SCI Expanded) data set. The SCI Expanded is a database that provides bibliographic information, abstracts, references, and citations counts for more than 6650 scholarly science and technology journals. The SCI Expanded search was limited to articles published from 1900 to December 31, 2007. To find any article among the broadest selection of

scientific journals related to critical care, we searched for any article that contained the keywords "critical-care," "intensive-care," "ICU," or "critically-ill OR critical-illness" and that had been cited more than 100 times. A total of 472 articles were retrieved with this technique after duplicates were removed (see Fig. 1A).

Because many critical care articles found in the SCI Expanded were not tagged by the general keywords noted above, we also searched *within* critical care journals. To do this, we used the Journal Citation Report (JCR) 2006 to identify journals specializing in critical care medicine from more than 7500 scholarly journals. We first reduced the target list to 16 English-language journals of 18 critical care journals identified by the JCR. We reviewed any article that had been cited more than 100 times and saved articles pertaining to critical care medicine (Fig. 1B). This search resulted in 470 hits; however, we removed 159 duplicate articles that were already found by keyword searches to bring the total database to 783 articles (Fig. 1).

2.2. Creation of disease/condition-specific lists

A representative list of most cited articles for each of the specific critical care topics was created from the existing database and by searching each topic individually using the terms in Table 1. Using these search terms, articles pertinent to

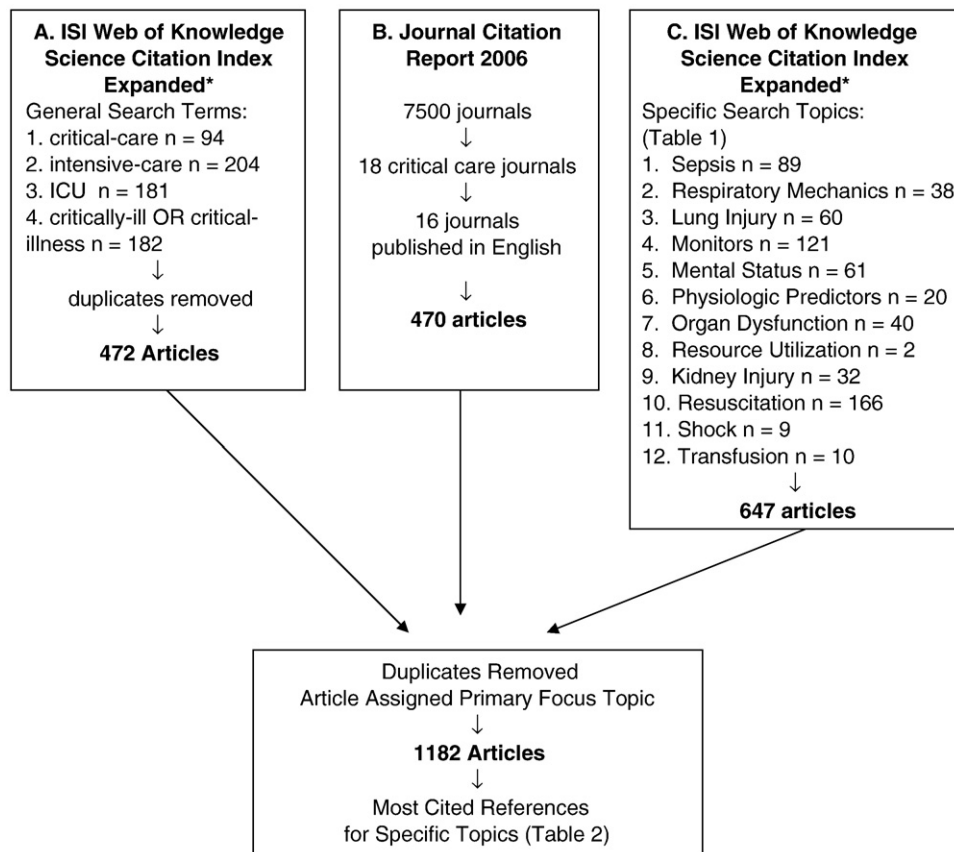


Fig. 1 Methods.

critical care and referenced more than 100 times were added to the database. These searches by specific disease or conditions added 404 articles to the master database as 194 articles were already in the database from either the initial search or from the journal search. From this master database of 1187 articles, we then generated a list of all articles cited more than 150 times (Table 2). We organized this table by topics particularly relevant to critical care. Within each category we listed both the most cited articles within that topic as well as that article's ranking within the top 200 most cited articles. Finally, we also listed the absolute number of citations for each article.

3. Results

The final database included 1187 articles (Fig. 1). The articles collated in this study came from almost 100 different

Table 1 Specific critical care subjects

Subject area	Search terms
Sepsis and infectious disease	"sepsis" "septic*" "nosocomial" "blood stream"
Mechanical ventilation and lung injury	"ventilation" "respiratory failure" "pulmonary failure" "weaning" "extubation" "ventilator" "PEEP" "positive end expiratory pressure" "noninvasive" "ARDS" "respiratory-distress" "lung injury"
Monitors	"hemodynamic-monitor" "CVC" "physiologic-monitor" "noninvasive-monitor" "catheter*" "central venous" "pulmonary-artery-catheter" "Swan-Ganz" "ultrasound" "Doppler" "pressure variation"
Mental status	"coma" "brain-damage" "brain-injury" "ICU AND delirium" "ICU AND sedation"
Physiologic predictors	"physiology score" "severity of illness" "severity of disease" "mortality-predict" "severity-score"
Organ dysfunction	"organ-failure" "organ-dysfunction" "MODS" "MSOF"
Resource utilization	"intensivist" "leap frog" "ICU-administration" "critical care management" "ICU management"
Kidney injury	"kidney-injury" "renal-failure" "dialysis" "renal-replacement-therapy" "CVVH*" "CRRT"
Resuscitation	"life-support" "cardiac-arrest" "cpr" "resuscitation" "cardiac massage" "cardiac-life-support" "fluid" "saline" "lactated" "albumin" "colloid" "crystalloid"
Shock (excluding infectious causes)	"shock" "hypoperfusion" "CHF" "heart-failure"
Transfusions	"transfusion" "fluid-therapy"

*Denotes any form of the word preceding the symbol, i.e., septic shock, septic.

journals with publication years ranging from 1905 (#149) to 2006 (average publication year = 1991). Surprisingly, of the top 1000 most cited critical care articles, 662 were *not* found using any of the basic critical care search terms—critical care, intensive care, ICU, or critically ill/critical illness. These articles were discovered only after searching within a specific critical care journal (n = 227) or by specific critical care topics (Table 1) (n = 373). In fact, only 4 of the top 1000 articles could be found using any 1 of the 4 basic critical care terms listed above. Of these search terms, the most productive was "intensive care" (25 of top 100 articles), then "critically ill" or "critical illness" (19/100), "ICU" (11/100), and finally "critical care" (5/100). Sepsis/systemic inflammatory response syndrome was the most common topic (193 articles), followed by acute lung injury and mechanical ventilation (177 articles). Overall, 41% of the most cited articles came from critical care journals as designated by the JCR. The most articles were from *Critical Care Medicine* (23%) followed by *American Journal of Respiratory and Critical Care* (7%), *Journal of Trauma* (5%), *Intensive Care Medicine* (4%), and *Shock* (1%). Among the non-critical care journals, the *New England Journal of Medicine* (11%) followed by *JAMA* (9%) provided the most articles.

4. Discussion

One method to quickly retrieve scholarly articles related to critical care medicine is by using readily available, Internet-based, bibliographic database search tools supported by most medical libraries. We have found, however, that commonly used basic search terms retrieve under only 50% of the most cited and therefore arguably most influential articles in critical care. Therefore, although searches may have become much faster and easier, they may not be sufficient to retrieve a comprehensive list of articles.

We found that there is no single term that exhaustively retrieves all critical care medicine articles. The best generic search term, "intensive care," only retrieved 25% of what one would consider the 100 of the most influential articles in critical care. Furthermore, the search term "critical care" was linked to only 5 of the top 100 articles, although it is a common keyword used by many search engines. Not to mention the fact that it is in the title of 8 of the 18 "critical care" journals.

One goal of this study was to determine the most efficient search terms for future literature searches. The capriciousness of how efficient a search term is perhaps no better illustrated than with our experience retrieving articles on ICU mortality prediction models. That not one of the basic critical care search terms in the Science Citation Index was tagged to these most influential articles highlights some of the difficulties in finding the most cited studies in critical care. It is clear that even for other types of articles, the best yield of

Table 2 Most cited articles in critical care medicine by subject

Overall Citation Rank
A. Sepsis/systemic inflammatory response syndrome (no. of times cited)
1. (#6) Tracey, KJ et al. Anti-cachectin TNF monoclonal-antibodies prevent septic shock during lethal bacteremia. <i>Nature</i> 1987. (1983)
2. (#7) Bone, RC et al. Definitions for sepsis and organ failure and guidelines for the use of innovative therapies in sepsis. <i>Chest</i> 1992. (1838)
3. (#8) Bernard, GR et al. Efficacy and safety of recombinant human activated protein C for severe sepsis. <i>N Engl J Med</i> 2001. (1827)
4. (#10) Bone, RC et al. American-College of Chest Physicians Society of Critical Care Medicine Consensus Conference—definitions for sepsis and organ failure and guidelines for the use of innovative therapies in sepsis. <i>Crit Care Med</i> 1992. (1724)
5. (#12) Van den Berghe, G et al. Intensive insulin therapy in critically ill patients. <i>N Engl J Med</i> 2001. (1608)
6. (#19) Ziegler, EJ et al. Treatment of gram-negative bacteremia and septic shock with HA-1a human monoclonal-antibody against endotoxin—a randomized, double-blind, placebo-controlled trial. <i>N Engl J Med</i> 1991. (1072)
7. (#22) Angus, DC et al. Epidemiology of severe sepsis in the United States: analysis of incidence, outcome, and associated costs of care. <i>Crit Care Med</i> 2001. (1016)
8. (#25) Bone, RC. The pathogenesis of sepsis. <i>Ann Intern Med</i> 1991. (982)
9. (#28) Waage, A et al. The complex pattern of cytokines in serum from patients with meningococcal septic shock—association between interleukin-6, interleukin-1, and fatal outcome. <i>J Exp Med</i> 1989. (892)
10. (#31) Wichterman, KA et al. Sepsis and septic shock—a review of laboratory models and a proposal. <i>J Surg Res</i> 1980. (846)
11. (#34) Bone, RC et al. A controlled clinical-trial of high-dose methylprednisolone in the treatment of severe sepsis and septic shock. <i>N Engl J Med</i> 1987. (791)
12. (#37) Parrillo, JE. Mechanisms of disease—pathogenetic mechanisms of septic shock. <i>N Engl J Med</i> 1993. (756)
13. (#41) Annane, D et al. Effect of treatment with low doses of hydrocortisone and fludrocortisone on mortality in patients with septic shock. <i>JAMA</i> 2002. (712)
14. (#43) Petros, A et al. Effect of nitric-oxide synthase inhibitors on hypotension in patients with septic shock. <i>Lancet</i> 1991. (705)
15. (#52) Cannon, JG et al. Circulating interleukin-1 and tumor necrosis factor in septic shock and experimental endotoxin fever. <i>J Infect Dis</i> 1990. (642)
16. (#55) Casey, LC et al. Plasma cytokine and endotoxin levels correlate with survival in patients with the sepsis syndrome. <i>Ann Intern Med</i> 1993. (624)
17. (#56) Vanzee, KJ et al. Tumor-necrosis-factor soluble receptors circulate during experimental and clinical inflammation and can protect against excessive tumor-necrosis-factor- α in vitro and in vivo. <i>Proc Nat Acad Sci USA</i> 1992. (600)

Table 2 (continued)

Overall Citation Rank
A. Sepsis/systemic inflammatory response syndrome (no. of times cited)
18. (#59) Hack, CE et al. Increased plasma-levels of interleukin-6 in sepsis. <i>Blood</i> 1989. (589)
19. (#62) Moore, FA et al. Early enteral feeding, compared with parenteral, reduces postoperative septic complications—the results of a metaanalysis. <i>Ann Surg</i> 1992. (578)
20. (#64) Lowenstein, CJ et al. Nitric-oxide—a physiological messenger. <i>Ann Intern Med</i> 1994. (574)
21. (#69) Fry, DE et al. Multiple system organ failure—role of uncontrolled infection. <i>Arch Surg</i> 1980. (549)
22. (#70) Parker, MM et al. Profound but reversible myocardial depression in patients with septic shock. <i>Ann Intern Med</i> 1984. (548)
23. (#72) Kudsk, KA et al. Enteral versus parenteral-feeding—effects on septic morbidity after blunt and penetrating abdominal-trauma. <i>Ann Surg</i> 1992. (543)
24. (#75) Fisher, CJ et al. Treatment of septic shock with the tumor necrosis factor receptor:Fc fusion protein. <i>N Engl J Med</i> 1996. (540)
25. (#77) Hotchkiss, RS et al. Medical progress: the pathophysiology and treatment of sepsis. <i>N Engl J Med</i> 2003. (538)
26. (#78) Greenman, RL et al. A controlled clinical-trial of E5 murine monoclonal IgM antibody to endotoxin in the treatment of gram-negative sepsis. <i>JAMA</i> 1991. (536)
27. (#81) Martin, GS et al. The epidemiology of sepsis in the United States from 1979 through 2000. <i>N Engl J Med</i> 2003. (529)
28. (#83) Glauser, MP et al. Septic shock—pathogenesis. <i>Lancet</i> 1991. (522)
29. (#89) Dellinger, RP et al. Surviving sepsis campaign guidelines for management of severe sepsis and septic shock. <i>Crit Care Med</i> 2004. (513)
30. (#94) Bone, RC et al. Sepsis syndrome—a valid clinical entity. <i>Crit Care Med</i> 1989. (506)
31. (#96) Fisher, CJ et al. Recombinant human interleukin-1 receptor antagonist in the treatment of patients with sepsis syndrome—results from a randomized, double-blind, placebo-controlled trial. <i>JAMA</i> 1994. (499)
32. (#97) Parrillo, JE et al. Septic shock in humans—advances in the understanding of pathogenesis, cardiovascular dysfunction, and therapy. <i>Ann Intern Med</i> 1990. (499)
33. (#105) Assicot, M et al. High serum procalcitonin concentrations in patients with sepsis and infection. <i>Lancet</i> 1993. (481)
34. (#109) Ochoa, JB et al. Nitrogen-oxide levels in patients after trauma and during sepsis. <i>Ann Surg</i> 1991. (464)
35. (#114) Moore, FA et al. TEN versus TPN following major abdominal-trauma—reduced septic morbidity. <i>J Trauma</i> 1997. (435)
36. (#120) Abraham, E et al. Efficacy and safety of monoclonal-antibody to human tumor necrosis-factor- α in patients with sepsis syndrome—a randomized, controlled, double-blind, multicenter clinical-trial. <i>JAMA</i> 1995. (425)

Table 2 (continued)

Overall Citation Rank

A. Sepsis/systemic inflammatory response syndrome
(no. of times cited)

37. (#121) Meakins, JL et al. Delayed-hypersensitivity—indicator of acquired failure of host defenses in sepsis and trauma. *Ann Surg* 1977. (425)
38. (#122) Pinsky, MR et al. Serum cytokine levels in human septic shock—relation to multiple system organ failure and mortality. *Chest* 1993. (422)
39. (#131) Danner, RL et al. Endotoxemia in human septic shock. *Chest* 1991. (409)
40. (#132) Eickhoff, TC et al. Neonatal sepsis + other infections due to group b beta-hemolytic streptococci. *N Engl J Med* 1964. (408)
41. (#134) Wheeler, AP et al. Treating patients with severe sepsis. *N Engl J Med* 1999. (405)
42. (#151) Sprung, CL et al. The effects of high-dose corticosteroids in patients with septic shock—a prospective, controlled-study. *N Engl J Med* 1984. (386)
43. (#153) Damas, P et al. Cytokine serum level during severe sepsis in human il-6 as a marker of severity. *Ann Surg* 1992. (382)
44. (#157) Askanazi, J et al. Influence of total parenteral-nutrition on fuel utilization in injury and sepsis. *Ann Surg* 1980. (377)
45. (#162) Docke, WD et al. Monocyte deactivation in septic patients: restoration by IFN-gamma treatment. *Nature Med* 1997. (372)
46. (#168) Petros, A et al. Effects of a nitric-oxide synthase inhibitor in humans with septic shock. *Cardiovasc Res* 1994. (366)
47. (#173) BrunBuisson, C et al. Incidence, risk-factors, and outcome of severe sepsis and septic shock in adults—a multicenter prospective-study in intensive-care units. *JAMA* 1995. (361)
48. (#174) Nava, E et al. Inhibition of nitric-oxide synthesis in septic shock—how much is beneficial. *Lancet* 1991. (361)
49. (#178) Szabo, C. The pathophysiological role of peroxynitrite in shock, inflammation, and ischemia-reperfusion injury. *Shock* 1996. (350)
50. (#180) Warren, BL et al. High-dose a randomized antithrombin III in severe sepsis—a randomized controlled trial. *JAMA* 2001. (348)
51. (#181) Bone, RC. Sir Isaac Newton, Sepsis, SIRS, and cars. *Crit Care Med* 1996. (348)
52. (#182) Schumer, W. Steroids in treatment of clinical septic shock. *Ann Surg* 1976. (348)
53. (#183) Damas, P et al. Tumor necrosis factor and interleukin-1 serum levels during severe sepsis in humans. *Crit Care Med* 1989. (346)
54. (#185) Cohen, J et al. The immunopathogenesis of sepsis. *Nature* 2002. (345)
55. (#187) Rush, BF et al. Endotoxemia and bacteremia during hemorrhagic-shock—the link between trauma and sepsis. *Ann Surg* 1988. (341)
56. (#190) Fourrier, F et al. Septic shock, multiple organ failure, and disseminated intravascular coagulation—compared

Table 2 (continued)

Overall Citation Rank

A. Sepsis/systemic inflammatory response syndrome
(no. of times cited)

- patterns of antithrombin-III, protein-C, and protein-S deficiencies. *Chest* 1992. (339)
57. (#192) Munoz, C et al. Dysregulation of in vitro cytokine production by monocytes during sepsis. *J Clin Invest* 1991. (336)

B. Outcome/benchmarking/prediction model articles
(no. of times cited)

1. (#1) Knaus, WA et al. APACHE-II—a severity of disease classification-system. *Crit Care Med* 1985. (4909)
2. (#4) Baker, SP et al. Injury severity score—method for describing patients with multiple injuries and evaluating emergency care. *J Trauma* 1974. (2637)
3. (#13) Legall, JR et al. A new simplified acute physiology score (SAPS-II) based on a European North-American multicenter study. *JAMA* 1993. (1402)
4. (#14) Knaus, WA et al. The APACHE-III prognostic system—risk prediction of hospital mortality for critically ill hospitalized adults. *Chest* 1991. (1357)
5. (#21) Knaus, WA et al. A controlled trial to improve care for seriously ill hospitalized-patients—the study to understand prognoses and preferences for outcomes and risks of treatments (support). *JAMA* 1995. (1056)
6. (#26) Fine, MJ et al. A prediction rule to identify low-risk patients with community-acquired pneumonia. *N Engl J Med* 1997. (965)
7. (#33) Deitch, EA. Multiple organ failure—pathophysiology and potential future therapy. *Ann Surg* 1992. (793)
8. (#36) Knaus, WA et al. APACHE-acute physiology and chronic health evaluation—a physiologically based classification-system. *Crit Care Med* 1981. (758)
9. (#38) Vincent, JL et al. The SOFA (sepsis-related organ failure assessment) score to describe organ dysfunction/failure. *Intens Care Med* 1996. (735)
10. (#40) Knaus, WA et al. An evaluation of outcome from intensive-care in major medical centers. *Ann Intern Med* 1986. (715)
11. (#47) Legall, JR et al. A simplified acute physiology score for ICU patients. *Crit Care Med* 1984. (667)
12. (#49) Boyd, CR et al. Evaluating trauma care—the TRISS method. *J Trauma* 2001. (658)
13. (#58) Marshall, JC et al. Multiple organ dysfunction score—a reliable descriptor of a complex clinical outcome. *Crit Care Med* 1995. (592)
14. (#63) Carrico, CJ et al. Multiple-organ-failure syndrome. *Arch Surg* 1986. (575)
15. (#80) Baker, SP et al. Injury severity score—update. *J Trauma* 1997. (532)
16. (#87) Champion, HR et al. A revision of the trauma score. *J Trauma* 1997. (516)
17. (#98) Pollack, MM et al. Pediatric risk of mortality (prism) score. *Crit Care Med* 1988. (498)

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Table 2 (continued)

 B. Outcome/benchmarking/prediction model articles
 (no. of times cited)

18. (#103) Champion, HR et al. Trauma score. *Crit Care Med* 1981. (486)
19. (#112) Keene, AR et al. Therapeutic intervention scoring system—update 1983. *Crit Care Med* 1983. (444)
20. (#115) Higgins, TL et al. Stratification of morbidity and mortality outcome by preoperative risk-factors in coronary-artery bypass patients—a clinical severity score. *JAMA* 1992. (435)
21. (#149) Lemeshow, S et al. Mortality probability-models (MPM-II) based on an international cohort of intensive-care unit patients. *JAMA* 1993. (388)
22. (#150) Champion, HR et al. The major trauma outcome study—establishing national norms for trauma care. *J Trauma* 1996. (386)
23. (#169) Takala, J et al. Increased mortality associated with growth hormone treatment in critically ill adults. *N Engl J Med* 1999. (365)
24. (#184) Vincent, JL et al. Use of the SOFA score to assess the incidence of organ dysfunction/failure in intensive care units: results of a multicenter, prospective study. *Crit Care Med* 1998. (346)
25. (#197) Faist, E et al. Multiple organ failure in polytrauma patients. *J Trauma* 1995. (332)
26. Moore, EE et al. Organ injury scaling—spleen, liver, and kidney. *J Trauma* 1995. (317)
27. Cockburn, F et al. The CRIB (clinical risk index for babies) score—a tool for assessing initial neonatal risk and comparing performance of neonatal intensive-care units. *Lancet* 1993. (273)
28. Greenspan, L et al. Abbreviated injury scale and injury severity score—a scoring chart. *J Trauma* 1993. (267)
29. Ware, JE et al. Comparison of methods for the scoring and statistical-analysis of SF-36 health profile and summary measures—summary of results from the medical outcomes study. *Med Care* 1995. (259)
30. Richardson, DK et al. Score for neonatal acute physiology—a physiological severity index for neonatal intensive-care. *Pediatrics* 1993. (252)
31. Moore, EE et al. Organ injury scaling—spleen and liver [1994 revision]. *J Trauma* 1993. (233)
32. Copes, WS et al. The injury severity score revisited. *J Trauma* 1988. (217)
33. Pollack, MM et al. PRISM III: an updated pediatric risk of mortality score. *Crit Care Med* 1996. (212)

 C. Mechanical and positive pressure ventilation/airway articles
 (no. of times cited)

1. (#9) Brower, RG et al. Ventilation with lower tidal volumes as compared with traditional tidal volumes for acute lung injury and the acute respiratory distress syndrome. *N Engl J Med* 2000. (1750)
 2. (#27) Amato, MBP et al. Effect of a protective-ventilation strategy on mortality in the acute respiratory distress syndrome. *N Engl J Med* 1998. (911)
 3. (#42) Gregory, GA et al. Treatment of idiopathic respiratory-distress syndrome with continuous positive airway pressure. *N Engl J Med* 1971. (708)
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Table 2 (continued)

 C. Mechanical and positive pressure ventilation/airway articles
 (no. of times cited)

4. (#44) Suter, PM et al. Optimum end-expiratory airway pressure in patients with acute pulmonary failure. *N Engl J Med* 1975. (698)
 5. (#53) Dreyfuss, D et al. Ventilator-induced lung injury—lessons from experimental studies. *Am J Resp Crit Care* 1992. (641)
 6. (#65) Brochard, L et al. Noninvasive ventilation for acute exacerbations of chronic obstructive pulmonary-disease. *N Engl J Med* 1995. (561)
 7. (#85) Stauffer, JL et al. Complications and consequences of endotracheal intubation and tracheotomy—a prospective-study of 150 critically ill adult patients. *Am J Med* 1981. (517)
 8. (#93) Ranieri, VM et al. Effect of mechanical ventilation on inflammatory mediators in patients with acute respiratory distress syndrome—a randomized controlled trial. *JAMA* 1999. (508)
 9. (#106) Nash, G et al. Pulmonary lesions associated with oxygen therapy and artificial ventilation. *N Engl J Med* 1967. (478)
 10. (#108) Bendixen, HH et al. Impaired oxygenation in surgical patients during general anesthesia with controlled ventilation—a concept of atelectasis. *N Engl J Med* 1963. (465)
 11. (#111) Webb, HH et al. Experimental pulmonary-edema due to intermittent positive pressure ventilation with high inflation pressures. protection by positive end-expiratory pressure. *Amer Rev Resp Dis* 1974. (445)
 12. (#113) Boyd, O et al. A randomized clinical-trial of the effect of deliberate perioperative increase of oxygen delivery on mortality in high-risk surgical patients. *JAMA* 1993. (439)
 13. (#129) Muscedere, JG et al. Tidal ventilation at low airway pressures can augment lung injury. *Am J Resp Crit Care* 1994. (412)
 14. (#135) Kumar, A et al. Continuous positive-pressure ventilation in acute respiratory failure—effects on hemodynamics and lung function. *N Engl J Med* 1970. (403)
 15. (#138) Hickling, KG et al. Low mortality associated with low-volume pressure limited ventilation with permissive hypercapnia in severe adult respiratory-distress syndrome. *Intens Care Med* 1990. (402)
 16. (#139) Arora, NS et al. Respiratory muscle strength and maximal voluntary ventilation in undernourished patients. *Amer Rev Resp Dis* 1982. (402)
 17. (#142) Kolobow, T et al. Severe impairment in lung-function induced by high peak airway pressure during mechanical ventilation—an experimental-study. *Amer Rev Resp Dis* 1987. (398)
 18. (#147) Haldane, JS et al. The regulation of the lung-ventilation. *J Physiol-London* 1905. (390)
 19. (#154) Ashbaugh, DG et al. Continuous positive-pressure breathing (CPPB) in adult respiratory distress syndrome. *J Thorac Cardio Surg* 1969. (382)
 20. (#196) Bott, J et al. Randomized controlled trial of nasal ventilation in acute ventilatory failure due to chronic obstructive airways disease. *Lancet* 1993. (333)
 21. Falke, KJ et al. Ventilation with end-expiratory pressure in acute lung-disease. *J Clin Invest* 1972. (325)
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Table 2 (continued)

C. Mechanical and positive pressure ventilation/airway articles (no. of times cited)

22. Kramer, N et al. Randomized, prospective trial of noninvasive positive pressure ventilation in acute respiratory-failure. *Am J Resp Crit Care* 1995. (323)
23. Stewart, TE et al. Evaluation of a ventilation strategy to prevent barotrauma in patients at high risk for acute respiratory distress syndrome. *N Engl J Med* 1998. (322)
24. Yang, KL et al. A prospective-study of indexes predicting the outcome of trials of weaning from mechanical ventilation. *N Engl J Med* 1991. (322)
25. Gattinoni, L et al. Low-frequency positive-pressure ventilation with extracorporeal CO₂ removal in severe acute respiratory-failure. *JAMA* 1986. (321)

D. Acute lung injury/adult respiratory distress syndrome articles (no. of times cited)

1. (#11) Bernard, GR et al. The American-European Consensus Conference on ARDS—definitions, mechanisms, relevant outcomes, and clinical-trial coordination. *Am J Resp Crit Care* 1994. (1627)
2. (#16) Ashbaugh, DG et al. Acute respiratory distress in adults. *Lancet* 1967. (1283)
3. (#17) Murray, JF et al. An expanded definition of the adult respiratory-distress syndrome. *Amer Rev Resp Dis* 1988. (1133)
4. (#18) Rossaint, R et al. Inhaled nitric-oxide for the adult respiratory-distress syndrome. *N Engl J Med* 1993. (1083)
5. (#32) Ware, LB et al. Medical progress—the acute respiratory distress syndrome. *N Engl J Med* 2000. (794)
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Table 2 (continued)**J. Resuscitation and cardiac arrest articles (no. of times cited)**

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L. Transfusion medicine/hematology/bleeding articles (no. of times cited)

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Table 2 (continued)

L. Transfusion medicine/hematology/bleeding articles (no. of times cited)
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20. Hebert, PC et al. Does transfusion practice affect mortality in critically ill patients? <i>Am J Resp Crit Care</i> 1997. (125)
21. Cosgriff, N et al. Predicting life-threatening coagulopathy in the massively transfused trauma patient: hypothermia and acidosis revisited. <i>J Trauma</i> 1997. (118)
22. Corwin, HL et al. RBC transfusion in the ICU—is there a reason. <i>Chest</i> 1995. (118)
23. Fitzgerald, RD et al. Transfusing red blood cells stored in citrate phosphate dextrose adenine-1 for 28 days fails to improve tissue oxygenation in rats. <i>Crit Care Med</i> 1997. (100)

Article's rankings listed by subgroup then by its position within the overall top 200 citations.

only 25% indicates the need for a comprehensive search strategy such as the one used in this study.

Although this is a problem for those looking for research articles, it can also be an issue for the way published research becomes cited by others. For example, articles without abstracts or very brief abstracts were often found only after using exact topic words found in the article's title or by starting a search from within a specific critical care journal itself. Our experience has been previously demonstrated where the choice of keywords and abstract construction significantly impacts the likelihood that an article will be found using modern electronic searches [17].

This study does highlight the importance of sepsis and infectious disease as well as respiratory physiology in critical

care with more than half the articles falling into 1 of these 2 broad categories. On the other hand, the most cited article was related to physiologic predictors of morbidity and mortality and the next 3 articles were related to mental status.

4.1. Limitations

This study, like previous ones focusing on citations, has several limitations. Perhaps the most relevant is the debate as to what constitutes the most influential articles within a professional discipline [1,4,18]. Citation data found in all bibliographic databases rely not only on correctly acquiring cited references but also on the assumption that a primary study cites other references that are most relevant to the published article. It is likely that this study captured many of the classic articles in critical care as each article in our list of the top 200 articles was cited at least 300 times. Considering that 46% of articles published in medicine are never even cited, this is a remarkable observation [19]. Certainly, these articles deserve attention and must have made an indelible impact. However, this is not always the case. Authors may preferentially cite their or their colleagues' previous works both because of familiarity or to increase the citation of that article. Recent scholarship on social networks, especially in the age of accelerated knowledge transmission, suggests that there may be an enhanced level of "connectedness" related to these networks [20,21]. It is not known whether, or how, this connectedness may be influencing the dissemination and therefore the pattern of references within the critical care community or among specific critical care topics. However, it is possible that the pattern of some citations may reflect the influence of an individual, as some have referred to as the "ceremonial citation," rather than the specific findings within an article [16,22,23]. In addition, authors are more likely to cite articles of their own language [8,24] or articles that come from a highly cited journal. Not surprisingly, Baltussen found that among the most cited 45 articles, most of them came from non-critical care-focused journals [16]. Nevertheless, using citations as a proxy for influence and importance has both face validity and is supported by empiric data demonstrating that citation analyses correlate with articles with the highest quality hierarchies of evidence and research design [25].

Most importantly, the use of citations is also confounded by the effect of time from the year of publication [26] with peak of citations differing for different journals and areas of research. Once citations do peak, they eventually become part of common knowledge and are no longer cited. Previous studies suggest that articles peak 7 to 10 years after publication [15,26]. It has been suggested that "classic" articles are relevant to only a few decades and many important articles are lost to the passage of time [15]. These findings may be supported by the fact that among the top 200 articles in this study, 158 were published during or after 2000. It is probably too early to tell how electronic databases

will change the epidemiology of citation classics because it is easier to retrieve and perhaps cite influential articles than before the age of the Internet.

In conclusion, although Web-based search engines can produce lists of references within seconds to minutes, the utility of these searches can be quite limited as a complete end effective literature search remains an art. Periodic reviews of the literature may prove helpful to trainees mastering the most influential literature of our field as well as more established professionals searching for starting points for new investigations.

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