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JOGNN

Current Resources for Evidence-Based Practice, September 2020

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ABSTRACT

An extensive review of new resources to support the provision of evidence-based care for women and infants. The current column includes a discussion of "spin" in scientific reporting and its effect on summaries and syntheses of the literature and commentaries on reviews about early versus late amniotomy as part of labor induction protocols and the economic burden associated with maternal morbidity.

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On the Perils of Skimming Articles

This column's literature searches yielded an interesting systematic review by Ghannad et al. (2019). Rather than reporting results of a clinical question per se, the authors analyzed how those results were reported. Specifically, they looked at the literature on biomarkers as screening, diagnostic, and prognostic tests for ovarian cancer. Drawing on a robust existing literature on "spin" in presentation of research findings for their analysis, Ghannad and colleagues (2019) evaluated the most recent 200 studies on biomarkers for ovarian cancer and looked specifically at titles, abstracts, and conclusions compared to the main findings as reported in the results sections.

Spin of research findings comes in two forms: misrepresentation, defined as "misreporting and/ or distorted presentation of the study results in the title, abstract, or the main text, in a way that could mislead the reader," and misinterpretation, defined as "an interpretation of the study results in the abstract or main-text conclusion that is not consistent and/or is an extrapolation of the actual study results" (Ghannad et al., 2019, p. 11). In other words, authors conducted their study and found X, but in the title, abstract, and/or article conclusion they reported X+Y.

I assume readers of a column entitled "Current Resources for Evidence-based Practice" are familiar with the concept of publication bias, wherein studies without exciting results (often defined, unfortunately, as results that are not statistically significant) are rejected for publication. This leads to errors when one is attempting to conduct systematic reviews or meta-analyses because the search of published literature only produces papers that were indeed published. Thus, the review can only include *some* of all studies actually conducted because the rest never made it into print. The resulting review is therefore biased because by definition it does not truly encompass what is known about a topic. Publication bias as described here applies to the entire literature on a particular topic, not to any one study specifically.

Spin is likely a direct consequence of the process that leads to publication bias. When faced with research findings not deemed interesting enough to be worthy of publication, an author has a few options. One is to give up completely on publishing the work, which leads, of course, directly to publication bias. This is absolutely not ideal; publication bias aside, who wants to see months, sometimes years of work collecting, analyzing, and reporting data lead to nothing? Unfortunately, this is the simplest option and chosen not infrequently (hence, publication bias).

A second possibility is to reformat the work in some way to try and make it worthy of publication. You could combine results from two studies, for instance, or rework the paper so it makes a methodologic rather than substantive point. This latter approach is one I used for a paper stemming from my dissertation work several years ago. With help from my advising committee, I found maternal physical activity during pregnancy had little bearing on risk of cesarean. The results were decidedly not statistically significant. Despite this being a worthwhile thing to know-as evidence-based practitioners we want to know what does not work as well as what does-seven journals rejected this so-called negative results paper. Eventually tiring of beating my head against that wall, I rewrote it to focus on our analytic choices instead of our findings, after which it was published immediately. Importantly, I was still able to present the findings that physical activity did not correlate with cesarean birth in our sample. These findings were just now couched within a larger discussion of how to analyze data on physical activity during pregnancy instead of within a discussion on whether physical activity could prevent cesareans (Bovbjerg et al., 2015). Lest I portray myself as some paragon of publishing virtue, I'll admit the other paper that stemmed from my dissertation also did not contain exciting results, and when faced with the huge amount of work to redo it, I availed myself of Option 1.

The final option for a troublesome paper is spin. Here, the authors rework their paper slightly so that the results seem more exciting than they really were, hoping these slights-of-hand are not called out by peer reviewers (assuming they were done consciously, which might not always be the case). This almost certainly happens behind the scenes as well, as authors choose perhaps only to present the more interesting results instead of all results. In the Ghannad et al. (2019) article on the biomarkers for ovarian cancer literature, 70% of studies included one or more forms of spin. Paper or abstract conclusions claimed a biomarker worked when the results were not statistically significant or when p-values and confidence intervals were not presented in the results section (and one must assume they were not favorable), numbers in the abstract did not match numbers in the text or tables, the abstract or title mischaracterized the extent of biomarker testing (e.g., claiming it predicts ovarian cancer deaths when the study endpoint was 6-month recurrence), and so on. The end result is that the literature is saturated with claims of biomarkers that can assist with diagnosis or prognosis for ovarian cancer, but few of these have panned out in practice.

It is tempting to place the blame for spin squarely on authors, and indeed they are not blameless. However, much like when we consider how to reduce cesareans, we must look at the system as a whole rather than blaming any one class of person or any one individual person. I think the issue is larger than individual authors. It must be, for 140 out of 200 examined studies to have been written in a misleading way (Ghannad et al., 2019). Ghannad et.al. did not report on funding sources for these studies, but I wonder how commercial interests affect spin.

The system of publish-or-perish in academia has led to an ever-increasing number of global publications, as university administrators and granting agencies demand ever-more research productivity (Kovanis et al., 2016). Not only does this system subtly encourage authors to take shortcuts so another paper can go out the door, but also the burden placed on peer reviewers (who are not paid) has increased to a level that may be unsustainable (Kovanis et al., 2016). Journal editors in turn are evaluated based on the impact factors of their journals-a measure of how many citations, on average, each published paper in that journal receives. The impact factor contributes to publication bias because it incentivizes editors to disproportionately accept only the exciting papers that may be cited. The entire system is flawed.

What does this mean for evidence-based practice if so much of the evidence is potentially questionable? Systematic reviews and metaanalyses help dilute the effect of any one perhaps suboptimal study, whether written with spin or otherwise problematic (bias, uncontrolled confounding, etc.). However, publication biasbeing unable to include null results in reviews because null results tend not to get publishedmight still lead one to the wrong conclusion. In well-done systematic reviews, reviewers formally assess publication bias. While I do not check all reviews included in these bibliographies, I do make sure the authors of the highlighted reviews have assessed publication bias to a satisfactory degree. Clinicians can and should allow more fluidity in their practice; the term "evidenceinformed" rather than "evidence-based" implies more room for clinical experience, patient and provider preferences, and unique characteristics (Nevo & Slonim-Nevo, 2011). If they are involved with their universities' governance structures, researchers can work to align promotion and tenure expectations with the realities of doing good science. More importantly, researchers can collectively admit when a research question does not seem answerable. Treatment of women with ovarian cancer would be infinitely easier if we had reliable diagnostic or prognostic biomarkers; however, an extremely large body of literature suggests such a thing is currently—and perhaps permanently—out of reach.

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From Cochrane Database of Systematic Reviews (CDSR) Issues 05–06 (2020)

Systematic Reviews in CDSR: Women's Health

• Pelvic floor muscle training for preventing and treating urinary and faecal incontinence in antenatal and postnatal women.

Systematic Reviews in CDSR: Fertility, Contraception, and ART

 Aspirin or heparin or both for improving pregnancy outcomes in women with persistent antiphospholipid antibodies and recurrent pregnancy loss.

Systematic Reviews in CDSR: Pregnancy and Birth

 Factors influencing the uptake and use of nicotine replacement therapy and e-cigarettes in pregnant women who smoke: A qualitative evidence synthesis.

- Interventions to prevent women from developing gestational diabetes mellitus: An overview of Cochrane Reviews.
- Probiotic treatment for women with gestational diabetes to improve maternal and infant health and well-being.
- Skin preparation for preventing infection following caesarean section.
- Strategies for optimising antenatal corticosteroid administration for women with anticipated preterm birth.

Systematic Reviews in CDSR: Infant Health and Breastfeeding

- Higher versus lower protein intake in formula-fed low birth weight infants.
- Interventions for congenital talipes equinovarus (clubfoot).
- Newborn screening for galactosaemia.

Systematic Reviews in CDSR: SARS-CoV-2

- Antibody tests for identification of current and past infection with SARS-CoV-2.
- Convalescent plasma or hyperimmune immunoglobulin for people with COVID-19: A rapid review.
- Personal protective equipment for preventing highly infectious diseases due to exposure to contaminated body fluids in healthcare staff.

Recent Guidelines

U.S. Preventive Services Task Force

U.S. Preventive Services Task Force, Owens, D. K., Davidson, K. W., Krist, A. H., Barry, M. J., Cabana, M., ... Wong, J. B. (2020). Screening for bacterial vaginosis in pregnant persons to prevent preterm delivery: US Preventive Services Task Force recommendation statement. *JAMA*, 323(13), 1286–1292. https://doi.org/1 0.1001/jama.2020.2684

Evidence-Based Reviews From Other Sources

Recent Evidence-Based Reviews: Women's Health

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Recent Evidence-Based Reviews: Fertility, Contraception, and ART

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Featured Review: De Vivo, V., Carbone, L., Saccone, G., Magoga, G., De Vivo, G., Locci, M., ... Berghella, V. (2020). Early amniotomy after cervical ripening for induction of labor: A systematic review and meta-analysis of randomized controlled trials. *American Journal of Obstetrics and Gynecology*, *222*(4), 320–329. https://doi. org/10.1016/j.ajog.2019.07.049

In this meta-analysis of four randomized, controlled trials including 1,273 women undergoing mechanical or pharmaceutical cervical ripening followed by synthetic oxytocin (Pitocin) for induction of labor, De Vivo et al. (2020) summarized the literature about early amniotomy vs. late amniotomy or spontaneous rupture of membranes. They found early amniotomy, defined as before active labor begins, does not increase the risk of cesarean (31.1% vs. 30.9%; pooled relative risk 1.05, 95% confidence interval [0.71, 1.56]), but it does shorten the induction to birth interval by just under 5 hours (weighted mean difference, –4.95 hours, 95% confidence interval [–0.812, –1.78]).

Comment: Shortening the interval between induction and birth is a laudable goal, and in this relatively small number of women, early amniotomy appears to do so without a concurrent

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increase in risk of cesarean. The meta-analysis does not indicate whether women whose labors ended in cesarean were excluded (or otherwise censored) from the labor duration analysis; this would be a potential source for bias. However, given the similarity in cesarean rates between the groups, it is less of a concern.

Much more concerning is the notion that when faced with a problem (cesareans) possibly arising from an intervention considered over-used (induction; Declercq et al., 2013; Seijmonsbergen-Schermers et al., 2020), we attempt to solve it by intervening further in the normal physiologic process (amniotomy). Used in standard induction protocols, synthetic oxytocin causes intense uterine contractions but does not cross the blood-brain barrier as the endogenous hormone does (Buckley & Moberg, 2019). Thus, women who undergo induction have substantially more painful labors without the concurrent pleasure center activation/pain reduction effects of endogenous oxytocin, which infiltrates the brain (Buckley & Moberg, 2019). Women whose labors are induced are thus more likely to request epidural anesthesia (90% of participants in the De Vivo et al. [2020] meta-analysis used epidurals), which in turn further slows labor, possibly because the numbed pelvic nerves are then unable to transmit cues to release additional oxytocin as the head of the fetus descends (Buckley & Moberg, 2019). Thus begins the cascade of interventions in obstetrics that has so infiltrated our sense of normal we now often see this sort of false dichotomy in the literature: one intervention compared to another without a physiologic comparison group as if birth is not possible without intervention.

What if we instead choose to support physiologic labor processes as much as possible, even under circumstances including genuinely unavoidable interventions such as induction of labor? I'll assume here we have already reduced unnecessary inductions to the extent possible. There exists a large body of literature on continuous labor support, usually via doulas, as a low-cost, highly effective way to improve birth outcomes, including lowering cesarean rates (Bohren et al., 2017; Kozhimannil et al., 2016). It seems very likely that providing doulas for all women who undergo induction would fare even better than early amniotomy in terms of labor duration and cesarean rate. As nurses, midwives, and public health professionals, we must push back against the idea that adding interventions to birth is the only solution or even the best solution.

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Featured Review: Moran, P. S., Wuytack, F., Turner, M., Normand, C., Brown, S., Begley, C., & Daly, D. (2020). Economic burden of maternal morbidity—A systematic review of cost-of-illness studies. *PLOS ONE*, *15*(1), e0227377. https:// doi.org/10.1371/journal.pone.0227377

In this systematic review, Moran and colleagues (2020) summarized the cost of illness literature on "the economic burden of common health problems women experience over the course of their pregnancy and postpartum, excluding acute complications of labour and birth, or severe acute adverse maternal outcomes" (Moran et al., 2020, p. 3). All cost results from included studies were converted to 2018 Euros.

The authors found 16 studies on the costs associated with gestational diabetes, 13 for obesity, four for poor maternal mental health, four for hypertensive disorders, two for severe nausea and vomiting, and two for epilepsy. Results for these conditions are summarized in Table 1. The authors identified no studies on costs associated with urinary incontinence, back or pelvic girdle pain, exhaustion, or sexual function.

Comment: It is unsurprising that costs for these conditions are noticeably higher in the United States, which reinforces that the American maternity care system is unsustainable. The mental health results were surprising; I would have

		Average Cost,	Average Cost,	Average Cost,	Average
Outcome	Range	US vs. non-US	Woman Only	Neonate Only	Cost Both
Gestational Diabetes	€263 to E13,680	€4,607 vs. €1,444	€2,048	_	€3,388
Obesity	€191 to €16,046	€6867 vs. €768	€1,612	_	€8,964
Mental Health	€452 to €794	_	_	_	—
Hypertension	€2,860 to €8,595	_	_	_	_
Nausea	€191 to €454	_	_	_	_
Epilepsy	—	_	€6,033	€1,694	—
Note. All costs are in 2018 Euros. — = no data.					

Table 1: Summary of Results Reported by Moran et al. (2020)

predicted much higher costs for these conditions. The low costs may be related to lack of access to mental health services, stigma associated with seeking mental health services, or higher rates of reimbursement for other services (e.g., more billing for obesity than depression in women with comorbidities). Most studies included in this review also included only payer costs not societal costs such as lost work days, which led to underestimates for all conditions but perhaps more so for mental health.

Finally, it is interesting that no studies appear to have been conducted on common pregnancy sequelae such as incontinence. Perhaps these are so common they are not considered "problems" per se, and therefore not worth studying? Any nursing graduate students in need of a thesis or dissertation topic, this would be a good one! Regardless, it is clear that existing, published numbers on costs of childbearing vastly underestimate the true costs, and here in the United States, we pay too much for care.

Recent Evidence-Based Reviews: Infant Health and Breastfeeding

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