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AcetaMINophen for Back and Osteoarthritis Pain: Is the effect in the name?

Clinical Question: Is acetaminophen effective for the management of back pain or osteoarthritis?

Bottom Line: Acetaminophen is not efficacious for back pain, and provides no clinically meaningful impact on osteoarthritis. There are many other better interventions for these conditions like: short-term oral NSAIDs for back pain; topical NSAIDs or intra-articular steroids for osteoarthritis; and exercise for both.

Evidence:

- All scales out of 100.
 - Back pain:
 - One high-quality Randomized Controlled Trial (RCT)¹ of 1,652 primary care acute low back patients randomized to acetaminophen regularly 3,990 mg/day versus as needed ≤4,000 mg/day versus placebo x4 weeks.
 - No effect on any outcome (time to recovery, pain intensity, disability, function, global symptom change or quality of life) at any time.
 - Five systematic reviews²⁻⁶ (3-10 RCTs, ≤1,825 patients), mostly acute back pain, acetaminophen.
 - Versus placebo: No effect in acute back pain.²⁻⁴
 - Versus other treatments:
 - NSAIDs better (~7.5 points) for pain.⁵
 - Amitriptyline or heat wraps better (~13 points) for pain.⁶
 - Only one small study (29 patients) of chronic pain: Likely inferior to NSAIDs.⁴
- Osteoarthritis: Eight systematic reviews (4-15 RCTs, ≤5,986 patients).^{2,7-13}
 - Versus placebo:
 - Pain:^{2,7,8,11,12} standard mean difference is 0.13-0.18 (likely equals ~3-4 points in 100).
 - Does not meet minimal important difference of 0.37 (~9 points).¹⁴
 - When average change in pain is not meaningful, it is important to consider if any patients could get meaningful change. However, modeling suggests none will with acetaminophen.¹¹
 - Versus NSAIDs:

- Pain: NSAIDs improve pain ~6 points versus acetaminophen.¹⁰
 - Patient assessed global improvement in pain,⁸ NSAIDs better with Number Needed to Treat (NNT)=6.

Context:

- Harms: Acetaminophen increases the risk of elevated liver enzymes (>1.5x normal),² Number Needed to Harm (NNH)=21.
- Back pain guidelines recommend acetaminophen as 1st-line therapy despite inefficacy.¹⁵
- Osteoarthritis guidelines have recommended both for¹⁶ and against¹⁷ acetaminophen.
- What works best with lowest harms (NNT for pain unless other mentioned):
 - Acute back pain: Staying active help return to work (~3 days sooner),¹⁸ NSAIDs¹⁹ NNT~11, cyclobenzaprine²⁰ NNT~4 (but often adds little after naproxen).
 - Chronic back pain: Exercise¹⁸ NNT=4-8.
 - Osteoarthritis: Topical NSAIDs (hands and knees)²¹ NNT~5, intra-articular corticosteroid injections²² NNT~4, Exercise^{23,24} NNT~4-6.

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References:

- 1. Williams CM, Maher CG, Latimer J, et al. Lancet. 2014; 384:1586-96.
- 2. Machado GC, Maher CG, Ferreira PH, et al. BMJ. 2015; 350:h1225.
- 3. Saragiotto BT, Machado GC, Ferreira ML, *et al.* Cochrane Database Syst Rev. 2016;6: CD012230.
- Agency for Healthcare Research and Quality: <u>www.effectivehealthcare.ahrq.gov/search-for-guides-reviews-and-</u> <u>reports/?pageaction=displayproduct&productid=2192</u>. Last accessed August 26, 2016.
- 5. Chou R, Huffman LH. Ann Intern Med. 2007; 147:505-14.
- 6. Davies RA, Maher CG, Hancock MJ. Eur Spin J. 2008; 17:1423-30.
- 7. Bannuru RR, Schmid CH, Kent DM, et al. Ann Intern Med 2015;162:46-54.
- 8. Towheed TE, Maxwell L, Judd MG, *et al.* Cochrane Database Syst Rev. 2006; 1:CD004257.
- 9. Ennis ZN, Dideriksen D, Vaegter HB, *et al.* Basic Clin Pharmacol Toxicol. 2016; 118:184-9.
- 10. Lee C, Straus WL, Balshaw R, et al. Arthritis Rheum. 2004; 51:746-54.
- 11. da Cost BR, Reichenbach S, Keller, et al. Lancet. 2016; 387:2093-105.
- 12. Zhang W, Nuki G, Moskowitz RW. Osteoarthritis Cartilage. 2010; 18:476-99.
- 13. Bjordal JM, Klovning A Luunggren AE, et al. Eur J Pain. 2007; 11:125-38.
- 14. Wandel S, Juni P, Tendal B, et al. BMJ. 2010; 341:c4675.
- 15. Toward Optimized Practice: <u>www.topalbertadoctors.org/download/1885/LBPguideline.pdf?_20160802102552</u>. Last accessed August 26, 2016.
- 16. Hochberg MC, Altman RD, April KT, et al. Arthritis Care Res. 2012; 64:465-74.
- 17. Jevsevar DS, Brown GA, Jone DL, et al. J Bone Joint Surg Am. 2013; 95:1885-6.
- 18. Mildenberger A, Allan GM. Tools for Practice. Available at: <u>https://www.acfp.ca/wp-content/uploads/tools-for-practice/1473693475_tfp170exerciseandbackpainfv.pdf</u>. Last accessed: September 13, 2016.

- 19. Roelofs PD, Deyo RA, Koes BW, *et al.* Cochrane Database Syst Rev. 2008; 1:CD000396.
- 20. Braschi E, Garrison S, Allan GM. Can Fam Physician. 2015; 61:1074.
- 21. Allan GM, Turner R, Lindblad A. Tools for Practice #40: <u>www.acfp.ca/wp-</u> <u>content/uploads/tools-for-practice/1427230512_40updated-topicalnsaids.pdf</u>. Last accessed August 26, 2016.
- 22. Jamieson J, Allan GM. Tools for Practice #135: <u>www.acfp.ca/wp-</u> <u>content/uploads/tools-for-practice/1427727062_tfp135steroidsforkneeoa.pdf</u>. Last accessed August 26, 2016.
- 23. Fransen M, McConnell S, Harmer AR, *et al.* Cochrane Database Syst Rev. 2015; 1:CD004376.
- 24. Fransen M, McConnell S, Hernandez-Molina G, *et al.* Cochrane Database Syst Rev. 2014; 4:CD007912.

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