EFFICACY AND SAFETY OF INTRA-ARTICULAR BOTULINUM TOXIN A INJECTION FOR KNEE OSTEOARTHRITIS: A SYSTEMATIC REVIEW, META-ANALYSIS, AND META-REGRESSION OF CLINICAL TRIALS

http://dx.doi.org/10.2106/JBJS.OA.22.00121

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Supplementary Table 1. Literature search strategy

## 1A. PubMed and Europe PMC Search String:

"(knee osteoarthritis OR OA knee OR gonarthrosis OR genu osteoarthritis OR OA genu) AND (botulinum toxin OR botox OR BTX OR BoNT OR dysport) AND (clinical trials OR randomized trials OR RCT)"

## 1B. Example Scopus and ClinicalTrials.gov Search Strategy:

- 1. knee osteoarthritis
- 2. OA knee
- 3. gonarthrosis
- 4. genu osteoarthritis
- 5. OA genu
- 6. botulinum toxin
- 7. botox
- 8. BTX
- 9. BoNT
- 10. dysport
- 11. clinical trials
- 12. randomized trials
- 13. RCT
- 14. 1 or 2 or 3 or 4 or 5
- 15. 6 or 7 or 8 or 9 or 10
- 16. 11 or 12 or 13
- 17. 14 and 15 and 16

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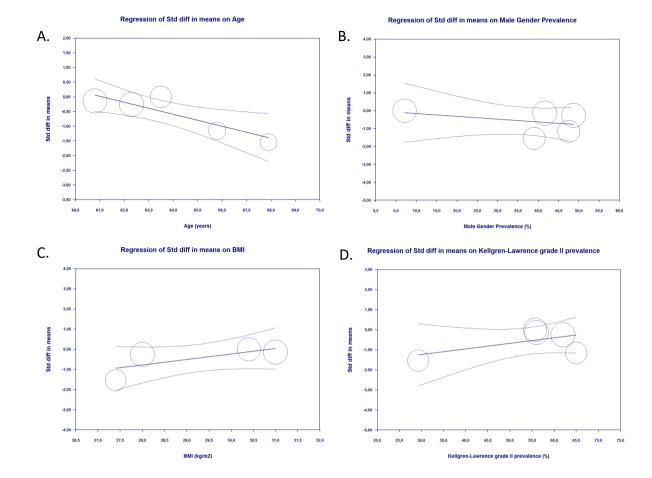
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## Supplementary Table 2. Results for the meta-regression models for each outcomes of

## interest

Change in Early VAS score (≤4 weeks)					
Covariate	Coefficient	95% CI (min)	95% CI (max)	S.E.	<i>p</i> -value
Age	-0.2046	-0.3332	-0.0760	0.0656	0.0018
Gender	-0.0157	-0.0554	0.0240	0.0203	0.4383
BMI	0.2721	-0.0876	0.6318	0.1835	0.1382
Kellgren-Lawrence grade	0.0276	-0.0172	0.0725	0.0229	0.2267
Change in Late VAS score (>4 weeks)					
Age	-0.1858	-0.3570	-0.0145	0.0874	0.0335
Gender	-0.0191	-0.0599	0.0217	0.0208	0.3584
BMI	0.1759	0.0321	0.3196	0.0734	0.0165
Kellgren-Lawrence grade	0.0028	-0.0538	0.0595	0.0289	0.9226
Change in Early WOMAC index (≤4 weeks)					
Age	-0.1973	-0.5117	0.1170	0.1604	0.2186
Gender	-0.0288	-0.0884	0.0308	0.0304	0.3442
Kellgren-Lawrence grade	-0.0116	-0.0951	0.0720	0.0426	0.7861
Change in Late WOMAC index (>4 weeks)					
Age	-0.3740	-1.1619	0.4139	0.4020	0.3522
Gender	0.0483	-0.1862	0.0895	0.0703	0.4919
BMI	0.1968	0.0534	0.3401	0.0731	0.0071
Kellgren-Lawrence grade	-0.0538	-0.2232	0.1155	0.0864	0.5332

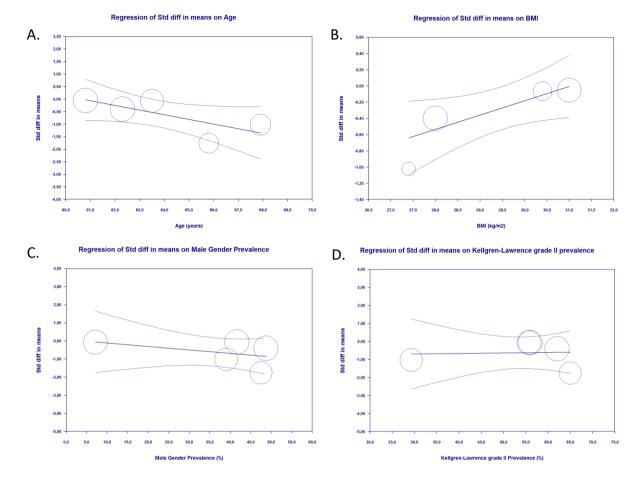
Efficacy and Safety of Intra-Articular Botulinum Toxin A Injection for Knee Osteoarthritis: A Systematic Review, Meta-Analysis, and Meta-Regression of Clinical Trials http://dx.doi.org/10.2106/JBJS.OA.22.00121 Page 3



Supplementary Figure 1. Bubble-plot for meta-regression. Meta-regression analysis showed that the association between botulinum toxin A intraarticular injection and changes in early VAS score was significantly affected by age (A), but not by gender (B), BMI (C), nor Kellgren-Lawrence grade (D)

 $\label{thm:condition} Efficacy \ and \ Safety \ of \ Intra-Articular \ Botulinum \ Toxin \ A \ Injection \ for \ Knee \ Osteoarthritis: A \ Systematic \ Review, \ Meta-Analysis, \ and \ Meta-Regression \ of \ Clinical \ Trials \ http://dx.doi.org/10.2106/JBJS.OA.22.00121$ 



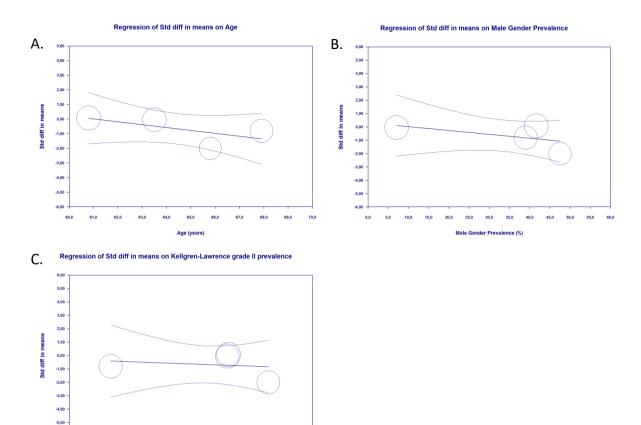


Supplementary Figure 2. Bubble-plot for meta-regression. Meta-regression analysis showed that the association between botulinum toxin A intraarticular injection and changes in late VAS score was significantly affected by age (A) and BMI (B), but not by gender (C) nor Kellgren-Lawrence grade (D)

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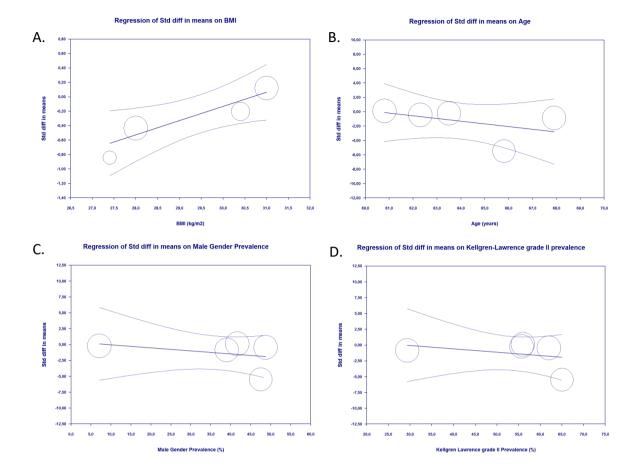
0 40,0 45,0 50,0 55,0

Kellgren-Lawrence grade II Prevalence (%)



Supplementary Figure 3. Bubble-plot for meta-regression. Meta-regression analysis showed that the association between botulinum toxin A intraarticular injection and changes in early WOMAC was not significantly affected by age (A), gender (B), nor Kellgren-Lawrence grade (C)

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Supplementary Figure 4. Bubble-plot for meta-regression. Meta-regression analysis showed that the association between botulinum toxin A intraarticular injection and changes in late WOMAC index was significantly affected by BMI (A), but not by age (B), gender (C), nor Kellgren-Lawrence grade (D)