

**The confidence in MD for efficacy at prophylaxis by GRADE system.**

| Comparison                           | Study limitation | Imprecision                                              | Inconsistency | Indirectness                                                            | Publication bias                                                                                                                          | GRADE    |
|--------------------------------------|------------------|----------------------------------------------------------|---------------|-------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------|----------|
| DXA-DAM vs. CG                       | No downgrade     | Upgrade due to large effect, MD= 18.48<br>13.14 to 23.69 | No downgrade  | The overall effect was significantly based on the primary outcome.      | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Moderate |
| DXA-PN vs. CG                        | No downgrade     | Upgrade due to large effect, MD= 9.75<br>7.51 to 12.01   | No downgrade  | The overall effect was significantly based on the primary outcome.      | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Moderate |
| DXA-IV-High dose vs. CG              | No downgrade     | Upgrade due to large effect, MD= 7.39<br>4.39 to 10.38   | No downgrade  | The overall effect was significantly based on the primary outcome.      | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Moderate |
| DXM vs.CG                            | No downgrade     | No downgrade, MD=7.15<br>3.44 to 10.80                   | No downgrade  | The overall effect was significantly based on the primary outcome.      | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-IV-Low dose vs. CG               | No downgrade     | No downgrade, MD= 6.51<br>3.57 to 9.42                   | No downgrade  | The overall effect was significantly based on the primary outcome.      | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-DAM vs. DXA-IV-Low dose          | No downgrade     | No downgrade, MD= 11.97<br>6.35 to 17.52                 | No downgrade  | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-PN vs. DXA-IV-Low dose           | No downgrade     | No downgrade, MD= 3.25<br>0.30 to 6.26                   | No downgrade  | The treatment effect was significantly based on the primary outcome     | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-IV-High dose vs. DXA-IV-Low dose | No downgrade     | No downgrade, MD= 0.89<br>-2.43 to 4.22                  | No downgrade  | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXM vs. DXA-IV-Low dose              | No downgrade     | No downgrade, MD= 0.64<br>-3.65 to 4.92                  | No downgrade  | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-DAM vs. DXM                      | No downgrade     | No downgrade, MD= 11.33<br>5.60 to 17.02                 | No downgrade  | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-PN vs. DXM                       | No downgrade     | No downgrade, MD= 2.61<br>-1.19 to 6.48                  | No downgrade  | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-IV-High dose vs. DXM             | No downgrade     | No downgrade, MD= 0.24<br>-4.21 to 4.69                  | No downgrade  | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-DAM vs. DXA-IV-High dose         | No downgrade     | No downgrade, MD= 11.09<br>5.53 to 16.53                 | No downgrade  | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-PN vs. DXA-IV-High dose          | No downgrade     | No downgrade, MD= 2.36<br>-0.74 to 5.50                  | No downgrade  | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |

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|---------------------------------------------|--------------|--------------------------------------------|--------------|----------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------|----------|
| DXA-DAM vs.<br>DXA-PN                       | No downgrade | No downgrade, MD= 8.72<br>3.20 to 14.12    | No downgrade | The treatment effect<br>was not significantly<br>based on the primary<br>outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXA-PN vs. CG                               | No downgrade | No downgrade, MD= 13.17<br>-20.55 to -6.26 | No downgrade | The treatment effect<br>was significantly<br>based on the primary<br>outcome     | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXA-DAM vs.<br>CG                           | No downgrade | No downgrade, MD= 10.98<br>-18.59 to -3.49 | No downgrade | The treatment effect<br>was significantly<br>based on the primary<br>outcome     | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXA-IV-Low<br>dose vs. CG                   | No downgrade | No downgrade, MD= -8.72<br>-15.08 to -2.39 | No downgrade | The treatment effect<br>was not significantly<br>based on the primary<br>outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXA-IV-High<br>dose vs. CG                  | No downgrade | No downgrade, MD= -8.42<br>-14.67 to 2.35  | No downgrade | The treatment effect<br>was not significantly<br>based on the primary<br>outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXM vs. CG                                  | No downgrade | No downgrade, MD= -3.26<br>-11.44 to 5.21  | No downgrade | The treatment effect<br>was not significantly<br>based on the primary<br>outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXA-PN vs.<br>DXM                           | No downgrade | No downgrade, MD= -9.91<br>-18.14 to 1.99  | No downgrade | The treatment effect<br>was not significantly<br>based on the primary<br>outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXA-DAM vs.<br>DXM                          | No downgrade | No downgrade, MD= -7.72<br>-15.61 to -0.25 | No downgrade | The treatment effect<br>was significantly<br>based on the primary<br>outcome     | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Moderate |
| DXA-IV-Low<br>dose vs. DXM                  | No downgrade | No downgrade, MD= -5.46<br>-12.83 to 1.54  | No downgrade | The treatment effect<br>was not significantly<br>based on the primary<br>outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXA-IV-High<br>dose vs. DXM                 | No downgrade | No downgrade, MD= -5.16<br>-13.49 to 2.81  | No downgrade | The treatment effect<br>was not significantly<br>based on the primary<br>outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXA-PN vs.<br>DXA-IV-High<br>dose           | No downgrade | No downgrade, MD= -4.75<br>-11.34 to 1.58  | No downgrade | The treatment effect<br>was not significantly<br>based on the primary<br>outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXA-DAM vs.<br>DXA-IV-High<br>dose          | No downgrade | No downgrade, MD= -2.56<br>-9.83 to 4.62   | No downgrade | The treatment effect<br>was not significantly<br>based on the primary<br>outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXA-IV-Low<br>dose vs. DXA-IV-<br>High dose | No downgrade | No downgrade, MD= -0.30<br>-5.75 to 5.10   | No downgrade | The treatment effect<br>was not significantly<br>based on the primary<br>outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXA-PN vs.<br>DXA-DAM                       | No downgrade | No downgrade, MD= -2.19<br>-10.26 to 5.55  | No downgrade | The treatment effect<br>was not significantly<br>based on the primary<br>outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |
| DXA-PN vs. CG                               | No downgrade | No downgrade, MD= -1.19<br>-2.73 to 0.26   | No downgrade | The treatment effect<br>was not significantly<br>based on the primary<br>outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot<br>for the network is not suggestive of<br>any dominant publication bias | Low      |

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|--------------------------------------|--------------|-------------------------------------------------------|--------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------|----------|
| DXA-DAM vs. CG                       | No downgrade | No downgrade, MD= -2.10<br>-8.24 to 4.19              | Only one head-to-head study, and no heterogeneity.<br>One direct comparison and no <code>_x005f</code> | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-IV-High dose vs. CG              | No downgrade | No downgrade, MD= -0.99<br>-3.50 to 1.39              | No downgrade                                                                                           | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXM vs. CG                           | No downgrade | No downgrade, MD= -0.42<br>-2.15 to 0.87              | No downgrade                                                                                           | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-IV-Low dose vs. CG               | No downgrade | No downgrade, MD= -0.01<br>-2.79 to 2.54              | No downgrade                                                                                           | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-PN vs. DXA-IV-Low dose           | No downgrade | No downgrade, MD= -1.18<br>-3.44 to 1.24              | No downgrade                                                                                           | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-DAM vs. DXA-IV-Low dose          | No downgrade | No downgrade, MD= -2.09<br>-8.61 to 4.75              | No downgrade                                                                                           | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-IV-High dose vs. DXA-IV-Low dose | No downgrade | No downgrade, MD= -0.98<br>-3.57 to 1.93              | No downgrade                                                                                           | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXM vs. DXA-IV-Low dose              | No downgrade | No downgrade, MD= -0.42<br>-3.37 to 2.60              | No downgrade                                                                                           | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-PN vs. DXM                       | No downgrade | No downgrade, MD= -0.77<br>-2.71 to 1.34              | No downgrade                                                                                           | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-DAM vs. DXM                      | No downgrade | No downgrade, MD= -1.68<br>-7.99 to 4.77              | No downgrade                                                                                           | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-IV-High dose vs. DXM             | No downgrade | No downgrade, MD= -0.57<br>-3.34 to 2.19              | No downgrade                                                                                           | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-PN vs. DXA-IV-High dose          | No downgrade | No downgrade, MD= -0.20<br>-2.28 to 1.96              | No downgrade                                                                                           | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-DAM vs. DXA-IV-High dose         | No downgrade | No downgrade, MD= -1.11<br>-7.34 to 5.02              | No downgrade                                                                                           | The treatment effect was not significantly based on the primary outcome | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Low      |
| DXA-PN vs. DXA-DAM                   | No downgrade | Upgrade due to large effect, MD=0.91<br>-5.35 to 7.16 | No downgrade                                                                                           | The treatment effect was significantly based on the primary outcome     | Undetectable by routine method.<br>The comparison-adjusted funnel plot for the network is not suggestive of any dominant publication bias | Moderate |