

Appendix A - Risk Of Bias

| Study | Confounding | Selection | Intervention Measurment | Missing Data | Outcome Measurement | Reported Results | Overall |
|-------------------------|-------------|-----------|----------------------------|--------------|------------------------|---------------------|----------|
| Castellani et al. | moderate | Low | moderate | Low | Low | Low | Low |
| Akgun et al. | moderate | Low | moderate | Low | Low | Low | Low |
| Ji et al. | moderate | Low | moderate | Low | Low | Low | Low |
| Li H et al. | moderate | Low | moderate | Low | low | low | low |
| Massin et al. | moderate | Low | moderate | Low | Low | moderate | moderate |
| Jenny et al. (2016) | moderate | Low | moderate | Low | low | moderate | moderate |
| Cochran et al. | low | Low | low | Low | Low | Low | Low |
| Zahar A et al | moderate | Low | low | Low | Low | Low | Low |
| Haddad et al. | moderate | Low | moderate | Low | Low | Low | Low |
| Cury Rde P et al. | moderate | Moderate | Low | moderate | low | low | moderate |
| Tibrewal et al. | low | low | low | low | low | low | low |
| Klatte et al. | moderate | low | low | moderate | low | low | low |
| Shanmugasundaram et al. | low | low | low | low | low | low | low |
| Baker et al. | low | low | moderate | moderate | low | low | moderate |
| Jenny et al. (2013) | moderate | low | low | low | low | moderate | moderate |
| Singer et al. | moderate | low | low | low | low | low | low |
| Whiteside et al. | moderate | low | low | low | low | low | low |

Note: Moderate=the study is sound for a non-randomized study with regard to this domain but cannot be considered comparable to a well-performed randomized trial;
 Low=the study is comparable to a well-performed randomized trial with regard to this domain

Appendix B – GRADE Quality

| Author | Study design | Risk of Bias | Inconsistency | Indirectness | Imprecision | Publication bias | Large effect (RR%) | Plausible confounding | Quality |
|-------------------|-------------------------------|--------------|---------------|--------------|-------------|------------------|--------------------|-------------------------------|-----------------|
| Castellani et al. | Retrospective Cohort | Low | Not serious | Not serious | serious | Undetected | N/A | Would suggest spurious effect | ⊕⊕⊕⊕ High |
| Ji et al. | Retrospective Case Series | Low | Not serious | Not serious | serious | serious | N/A | | ⊕⊕⊕ Moderate |
| Li H et al. | Retrospective Cohort | Low | Not serious | serious | Not serious | Undetected | N/A | No | ⊕⊕⊕⊕ High |
| Massin et al. | Retrospective Cohort | Moderate | Serious | Not serious | Not serious | Undetected | N/A | No | ⊕⊕⊕ Moderate |
| Jenny et al. 2016 | Retrospective case-control | Moderate | serious | Not serious | Not serious | Not serious | N/A | No | ⊕⊕⊕ Moderate |
| Cochran et al. | Retrospective Cohort registry | Low | Not serious | Not serious | Not serious | serious | N/A | No | ⊕⊕⊕⊕ High |

| | | | | | | | | | |
|-------------------------|-------------------------------------|----------|-------------|-------------|-------------|-------------|-----|--------------------------------------|-----------------|
| Zahar A et al. | Retrospective Cohort | Low | Not serious | Not serious | serious | Not serious | N/A | No | ⊕⊕⊕⊕ High |
| Haddad et al. | Retrospective Cohort | Low | serious | Not serious | Not serious | Not serious | N/A | No | ⊕⊕⊕⊕ High |
| Cury Rde P et al. | Retrospective Cohort | Moderate | Not serious | Not serious | serious | serious | N/A | No | ⊕⊕⊕ Moderate |
| Tibrewal et al. | Prospective Cohort | low | serious | Not serious | Not serious | Not serious | N/A | No | ⊕⊕⊕⊕ High |
| Klatte et al. | Retrospective Cohort | Low | Not serious | Not serious | serious | serious | N/A | Would reduce the demonstrated effect | ⊕⊕⊕ Moderate |
| Shanmugasundaram et al. | Retrospective Cohort | Low | Not serious | Not serious | serious | serious | N/A | No | ⊕⊕⊕ Moderate |
| Baker et al. | Prospective Cohort | Moderate | Not serious | Not serious | Not serious | Not serious | N/A | No | ⊕⊕⊕⊕ High |
| Jenny et al. (2013) | Observational Cohort Prospective | Moderate | Not serious | Not serious | Not serious | Not serious | N/A | No | ⊕⊕⊕⊕ High |

| | | | | | | | | | |
|------------------|------------------------------|-----|-------------|-------------|-------------|-------------|-----|----|--------------|
| Singer et al. | Descriptive Retrospective | Low | Not serious | Not serious | Not serious | Not serious | N/A | No | ⊕⊕⊕⊕ High |
| Whiteside et al. | Retrospective Cohort | Low | Not serious | Not serious | Not serious | Not serious | N/A | No | ⊕⊕⊕⊕ High |

“Very low”; the true effect is probably markedly different from the estimated effect, “Low”; The true effect might be markedly different from the estimated effect, “Moderate”; the authors believe that the true effect is probably close to the estimated effect, “High”; the authors have a lot of confidence that the true effect is similar to the estimated effect