| Author Year<br>Country<br>Research Design<br>Total Sample Size<br>AMSTAR Score                | Methods   | Outcome   |
|---|---|---|
| Arnold et al., (2017)<br>USA<br>Review of<br>published articles<br>up to February 2015<br>N=9 | Method: A comprehensive literature search<br>was conducted to identify randomized<br>controlled trials (RCT) evaluating the<br>efficacy and safety of antithrombotic<br>strategies. The strength of evidence was<br>evaluated using the Grading of<br>Recommendations Assessment,<br>Development and Evaluation (GRADE)<br>system.<br>Databases: MEDLINE; Cochrane<br>Collaboration Library.<br>Level of evidence: High quality study<br>designs such as RCTs and one prospective<br>controlled trial, were the only studies<br>included.<br>Questions/measures/hypothesis:<br>1. What is the effectiveness and safety of<br>anticoagulant thromboprophylaxis<br>compared to no prophylaxis, placebo, or<br>another anticoagulant strategy for<br>preventing deep vein thrombosis (DVT) and<br>pulmonary embolism (PE) after acute SCI?<br>2. What is the comparative effectiveness<br>and safety of mechanical prophylaxis<br>strategies alone or in combination with<br>other prophylactic strategies for preventing<br>DVT and PE after acute SCI?<br>3. What is the comparative effectiveness<br>and safety of prophylactic inferior vena cava<br>(IVC) filter insertion alone or in combination<br>with other prophylactic strategies for<br>preventing DVT and PE after acute SCI?<br>4. What is the optimal timing to initiate<br>and/or discontinue anticoagulant,<br>mechanical, and/or prophylactic IVC filter<br>following acute SCI?<br>What is the cost-effectiveness of the<br>treatment options mentioned above? | <ul> <li>Question one:</li> <li>Seven RCTs reported on the efficacy and/or safety of anticoagulant drug interventions.</li> <li>A single RCT reported the efficacy of LMWH versus no prophylaxis. Individuals treated with enoxaparin has a lower rate of DVT (5.4%) than those who received no LMWH prophylaxis (21.6%).</li> <li>Two RCTs assessed the risk of DVT in individuals receiving unfractionated heparin versus no treatment or placebo and found no significant difference between groups.</li> <li>A single RCT compared the efficacy and safety of two different LMWH drugs (enoxaparin or dalteparin). There was no significant difference in the rate of DVT or PE between groups.</li> <li>One RCT evaluated the efficacy and safety of fixed, low-dose versus adjusted-dose UFH. DVT and PE were observed in 9/29 (31%) and 2/29 (6.9%). The risk of DVT in the fixed, low-dose group was three times greater than the adjusted-dose group (RD=13.8, 95% CI=-3.6-31.2, RR=3.0, 95% CI=-3.6-31.2,</li></ul> |

| Author Year<br>Country<br>Research Design<br>Total Sample Size<br>AMSTAR Score | Methods  | Outcome   |
|--|--|---|
|  |  | <ol> <li>One RCT compared the<br/>efficacy and safety of<br/>mechanical prophylaxis<br/>versus mechanical<br/>prophylaxis plus<br/>antithrombotic drugs. No<br/>significant difference in<br/>safety or efficacy was<br/>observed between groups.</li> <li>Two RCTs compared<br/>outcomes between<br/>anticoagulant<br/>thromboprophylaxis and<br/>anticoagulant plus<br/>mechanical prophylaxis.<br/>Both studies reported<br/>significantly higher risk of<br/>DVT in the group that<br/>received anticoagulant<br/>prophylaxis only (50% and<br/>60.3% versus 6.7% and<br/>44.9%).</li> <li>Question three:</li> </ol> |
|  |  | 1. No RCTs were identified<br>that met inclusion criteria.<br>Question four:  |
|  |  | <ol> <li>One prospective<br/>controlled trial examined<br/>the timing of initiation of<br/>anticoagulant<br/>thromboprophylaxis in<br/>individuals with acute SCI.<br/>Combined anticoagulant<br/>and mechanical<br/>prophylaxis initiated<br/>within 72 hr of SCI resulted<br/>in significantly lower risk of<br/>DVT than treatment<br/>commenced 72 hr after<br/>injury.</li> <li>Question five:</li> </ol>  |
|  |  | 1. No RCTs were identified  |
| <u>Fehlings et al</u><br>(2017)<br>Canada<br>Clinical Practice<br>Guideline    | <b>Method:</b> A comprehensive literature search<br>was conducted to address key questions<br>relating to thromboprophylaxis in SCI. The<br>strength of evidence was evaluated using<br>the Grading of Recommendations | <ol> <li>that met inclusion criteria.</li> <li>Three RCTs compared the<br/>risk of DVT in individuals<br/>treated with LMWH or<br/>UFH to those receiving no<br/>prophylaxis or placebo.<br/>Individuals treated with</li> </ol>  |

| Author Year<br>Country<br>Research Design<br>Total Sample Size<br>AMSTAR Score | Methods   | Outcome   |
|--|---|---|
|  | <ul> <li>Assessment, Development and Evaluation<br/>(GRADE) system.</li> <li>Databases: Not reported.</li> <li>Level of evidence:</li> <li>Questions/measures/hypothesis: <ol> <li>Should anticoagulant<br/>thromboprophylaxis be employed to<br/>reduce the risk of thromboembolic<br/>events in the acute period after SCI?</li> <li>What anticoagulant<br/>thromboprophylaxis should be<br/>employed to reduce the risk of<br/>thromboembolic events in the acute<br/>period after traumatic SCI?</li> <li>Should enoxaparin versus dalteparin<br/>be used to reduce the risk of<br/>thromboembolic events in the acute<br/>period after traumatic SCI?</li> <li>Should fixed, low-dose, versus<br/>adjusted-dose unfractionated heparin<br/>(UFH) be used to reduce the risk of<br/>thromboembolic events in the acute<br/>period after traumatic SCI?</li> <li>Should low weight molecular heparin<br/>(LWMH) versus UFH be used to reduce<br/>the risk of thromboembolic events in<br/>the acute period after traumatic SCI?</li> <li>Should hormboprophylaxis be<br/>initiated within 72 hr (vs after 72 hr) of<br/>SCI?</li> </ol> </li></ul> | <ul> <li>enoxaparin have a lower<br/>rate of DVT (5.45%) than<br/>those who received no<br/>anticoagulant prophylaxis<br/>(21.6%) (p=0.09).</li> <li>Rates of DVT did not<br/>significantly differ<br/>between the UFH and the<br/>placebo/no prophylaxis<br/>group (1.8% and 3% in one<br/>trial and 50% and 74% in<br/>another).</li> <li>Anticoagulant<br/>thromboprophylaxis<br/>should be offered routinely<br/>to reduce the risk of<br/>thromboembolic events in<br/>the acute period after SCI.</li> <li>There is little to no<br/>difference in the rate of<br/>DVT, PE, bleeding and<br/>mortality between<br/>individuals treated with<br/>enoxaparin versus<br/>dalteparin.</li> <li>There is low quality<br/>evidence that the risk of<br/>DVT is three times higher<br/>in individuals who<br/>received fixed, low-dose<br/>UFH compared to<br/>adjusted-dose heparin<br/>(RD=13.8, 95% CI=-3.6-31.2;<br/>RR=3.0, 95% CI=0.66 to 13.7;<br/>p=0.25).</li> <li>The rate of bleeding is<br/>significantly higher in<br/>individuals treated with<br/>adjusted-dose heparin<br/>(24.1%) than in those<br/>receiving low-dose (0%)<br/>(RD=24.1, 95% CI=8.6-39.7;<br/>p=0.01).</li> <li>Anticoagulant<br/>thromboprophylaxis,<br/>consisting of either<br/>subcutaneous LMWH or<br/>fixed, low-dose UFH,<br/>should be offered to<br/>reduce the risk of<br/>thromboembolic events in<br/>the acute period after SCI.</li> </ul> |

| Author Year<br>Country<br>Research Design<br>Total Sample Size<br>AMSTAR Score | Methods | Outcome  |
|--|---------|--|
|  |         | <ol> <li>8. The authors caution<br/>against use of adjusted-<br/>dose UFH, due to the<br/>potential pf increased<br/>bleeding events.</li> <li>9. One prospective<br/>observational study<br/>evaluated the risks of DVT<br/>and PE in individuals who<br/>received prophylaxis<br/>initiated within or after 72<br/>hr of injury. Based on low<br/>quality evidence, the rate<br/>of DVT was significantly<br/>lower in individuals<br/>treated early (n=2)<br/>compared with late (n=46).<br/>There was insufficient<br/>evidence to compare the<br/>groups.</li> <li>10. Anticoagulant<br/>thromboprophylaxis<br/>should be commenced<br/>within the first 72 hr after<br/>injury, if possible, to<br/>minimize the risk of VTE<br/>complications during<br/>acute hospitalization.</li> <li>11. Individuals who received a<br/>combination of UFH and<br/>electronic calf stimulation<br/>had a lower risk of DVT<br/>than individuals treated<br/>with UFH alone (RD=43.3,<br/>95% CI=1.06-53.03, p=0.02).</li> <li>12. Individuals treated with<br/>LMWH alone have a lower<br/>risk of PE compared with<br/>individuals who receive<br/>UFH plus IPC (RD=13.2, 95%<br/>CI=0.9-25.4; RR=0.28, 95%<br/>CI=0.9-25.4; RR=0.28, 95%<br/>CI=0.08-0.98; p=0.06).</li> <li>13. A higher percentage of<br/>individuals experienced a<br/>DVT when treated with<br/>IPC alone (40%) compared<br/>with IPC plus aspirin and<br/>dipyridamole (25%);<br/>however, this difference<br/>was not statistically<br/>significant.</li> </ol> |

| Author Year<br>Country<br>Research Design<br>Total Sample Size<br>AMSTAR Score                       | Methods   | Outcome   |
|--|---|---|
| Christie et al., (2011)<br>Canada<br>Date included in<br>the review not<br>stated<br>N=5<br>AMSTAR=5 | Method: Comprehensive literature search of<br>English RCT, Cohort studies, case series, and<br>review articles of relating to prophylaxis low<br>molecular unfractionated heparin (LMWH)<br>for deep venous thrombosis (DVT) in<br>traumatic SCI in adult age group (+18yr).<br>Databases: PubMed.<br>Questions/measures/hypothesis: Examine<br>the ideal time for initiation of deep venous<br>thrombosis (DVT) prophylaxis with LMWH<br>after SCI or after surgery. | <ol> <li>DVT prophylaxis should be<br/>instituted within 72hr post<br/>injury.</li> <li>LMWH should be held on<br/>the morning of surgery and<br/>resumed within 24hr<br/>following surgery.</li> </ol> |