

Table 1. Study overview

| Reference              | Population                                                                                                                                | Muscles Stimulated                                           | E-STIM Protocol                                                                                                                                                                                                              |
|------------------------|-------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Abitante <sup>24</sup> | 10 healthy adults (6M, 4F); age: 23.8 years                                                                                               | Quadriceps, hamstrings, tibialis anterior, and gastrocnemius | Frequency = 50Hz, width = 400 $\mu$ s. Performed 1s contractions with either 3s (75 reps), 4s (60 reps), or 5s (50 reps) rest periods. Amplitude was set to maximal tolerable comfort level. Each protocol lasted 5 minutes. |
| Ashley <sup>25</sup>   | 11 individuals with spinal cord injuries (10M, 1F); age: 32.1 years, 5 individuals without spinal cord injuries (3M, 2F); age: 21.5 years | Quadriceps                                                   | Square monophasic waves. Frequency = 50Hz, width = 250 $\mu$ s. 4 second contraction with 4 second rest. Amplitude set to reach full knee extension. 20 minute protocol.                                                     |
| Banerjee <sup>17</sup> | 10 healthy adults (8M, 2F); age: 37.6 years                                                                                               | Quadriceps, hamstrings, and gluteal muscles                  | Frequency = 4Hz. Amplitude went up to 40% of maximal stimulator output (120 mA).12 minute protocol.                                                                                                                          |
| Chen <sup>26</sup>     | 9 healthy male adults; age: 28 years                                                                                                      | Quadriceps and plantar flexors                               | Biphasic pulses. Frequency = 20 Hz, 1s on 1s off. Amplitude was set to maximal tolerable intensity without feeling discomfort. 2 hour protocol.                                                                              |
| Fornusek <sup>28</sup> | 9 males with spinal cord injuries; age: 48 years                                                                                          | Quadriceps, hamstrings, and gluteal muscles                  | Biphasic rectangular pulses. Frequency = 35 Hz, width = 300 $\mu$ s. Amplitude was increased from 40 to 140 mA over 10 min, and then held constant at 140 mA for the next 25 min. 15 minute protocol.                        |
| Grosset <sup>30</sup>  | 9 obese adults (3M, 6F); age = 43.8 years                                                                                                 | Quadriceps and hamstrings                                    | Biphasic symmetrical pulses. Frequency = 5 Hz, width = 760 – 857 $\mu$ s, Amplitude was increased to maximal tolerable intensity. 1 hour protocol.                                                                           |
| Galvan <sup>29</sup>   | 10 overweight or obese Hispanic adults (2M, 8F); age: 36.8 years                                                                          | Quadriceps                                                   | Biphasic pulses. Frequency = 50 Hz, width = 300 $\mu$ s, Amplitude set to maximal tolerable intensity. 30 minute protocol.                                                                                                   |
| Hamada <sup>32</sup>   | 6 healthy males; age: 24 years                                                                                                            | Quadriceps                                                   | Biphasic pulses. Frequency = 20 Hz, width = 200 $\mu$ s. Amplitude set to 80 volts for non-painful stimulation. 1 second on-off duty cycle. 20 min protocol.                                                                 |
| Hamada <sup>31</sup>   | 8 healthy males; age: 24.8 years                                                                                                          | Tibialis anterior, plantar flexors, quadriceps, hamstrings   | Biphasic pulses. Frequency = 20 Hz, width = 200 $\mu$ s. Amplitude set to 80 volts for non-painful stimulation. 1 second on-off duty cycle. 20 min protocol.                                                                 |
| Hirata <sup>33</sup>   | 10 healthy males; age: 27 years                                                                                                           | Quadriceps                                                   | Frequency = 20Hz, width = 200 $\mu$ s. 1 second on-off duty cycle. Stimulation voltage started at 20 volts and progressively increased until the maximum tolerable level.                                                    |
| Hsu <sup>13</sup>      | 40 healthy adults (18M, 22F); age: 28.2 years                                                                                             | Quadriceps, hamstrings, gluteal maximum, abdominal muscles   | Biphasic square pulses. Frequency = 20 Hz. Duty cycle on/off = 1:2. 10 min sensory level (subject felt NMES), 10 min motor threshold (visible contractions), and 10 min maximal intensity comfortably tolerated.             |
| Isakov <sup>14</sup>   | 3 individuals with spinal cord injuries; age: 36.3 years                                                                                  | Quadriceps and gluteus muscles                               | Frequency = 24Hz, width = 300 $\mu$ s, amplitude = 120 volts. 10-30 minute protocol.                                                                                                                                         |

|                                           |                                                                                                                               |                                                                                         |                                                                                                                                                                                                                                                                                                                                         |
|-------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Janssen <sup>19</sup>                     | 12 healthy males; age: 25.1 years                                                                                             | Quadriceps, hamstrings, plantar flexors, and tibialis anterior                          | Individuals completed each of the following: left leg at 35Hz, right leg at 35 Hz, both legs at 35 Hz, both legs at 3 Hz, both legs at 1Hz. All protocols involved rectangular pulses with a width of 300 $\mu$ s. Amplitude was set at the maximum tolerable. Duty cycle on/off 6:20. 5 minute protocols.                              |
| Lago <sup>35</sup>                        | 16 early phase septic shock patients (12 M, 4F); age: 56 years. 21 late phase septic shock patients (11M, 10F); age: 57 years | Plantar flexors                                                                         | Biphasic symmetrical pulses. Frequency = 50Hz, width = 250 $\mu$ s. Duty cycle on/off = 2:5. Amplitude was increased to generate a visible muscle contraction. 30 minute protocol.                                                                                                                                                      |
| Laskin <sup>36</sup>                      | 8 individuals (7M, 1F) with spinal cord injuries; age: 27.9 years                                                             | Quadriceps and hamstrings                                                               | 40 minute protocol.                                                                                                                                                                                                                                                                                                                     |
| Ma <sup>23</sup>                          | 9 males with spinal cord injuries; age: 50 years                                                                              | Quadriceps, hamstrings, gluteals, plantar flexors.                                      | Biphasic square pulses. Frequency = 75Hz, width = 500 $\mu$ s. Amplitude was set to induce visible muscle contractions at comfortable levels. Four different protocols. Two involved all 4 muscle groups, while two only involved the gluteals and hamstrings. Duty cycles on:off were either 1:4 or 1:8. Each protocol was 10 minutes. |
| Matsushima <sup>37</sup>                  | 13 healthy males; age: 28 years                                                                                               | Quadriceps, hamstrings, gluteus maximus, and plantar flexors.                           | Frequency = 4 Hz, width = 250 $\mu$ s. Amplitude was increased to achieve a $\text{VO}_2$ of 14 ml/kg/min. 20 minute protocols.                                                                                                                                                                                                         |
| Minogue <sup>39</sup>                     | 11 healthy males; age: 34.2 years                                                                                             | Quadriceps                                                                              | Biphasic pulses. Frequency = 4 Hz. Amplitude was set at the maximal tolerable. Phase duration of 600 $\mu$ s with an interphase interval of 100 $\mu$ s. 30 minute protocol.                                                                                                                                                            |
| Minogue <sup>38</sup>                     | 10 healthy males; age: 35 years                                                                                               | Quadriceps                                                                              | Biphasic pulses. Phase duration of 600 $\mu$ s with an interphase interval of 100 $\mu$ s. Several protocols were completed across different frequencies including 1, 2, 4, 5, 6, 8, 10, and 12 Hz. Amplitude was set at the maximal tolerable. 4 minute protocols.                                                                     |
| Miyamoto <sup>42</sup>                    | 13 healthy males; age: 21.7 years                                                                                             | Quadriceps, hamstrings, gluteus maximus, triceps, and plantar flexors.                  | Frequency = 4 Hz, width = 250 $\mu$ s. Amplitude was increased until reaching approximately 3 METs. 30 minute protocol.                                                                                                                                                                                                                 |
| Perez-De-Arrilucea-Le-Floch <sup>20</sup> | 10 healthy males; age: 21.6 years                                                                                             | Upper legs, upper arms, upper back, gluteal, abdomen, chest, lower back, and shoulders. | Width ranged from 200–400 $\mu$ s depending on muscle group. Several protocols were completed across different frequencies including 1, 2, 4, 6, 8, and 10 Hz. Duty cycle was 99% (continued impulse). Amplitude set to maximal tolerable intensity. 6 minute protocol.                                                                 |
| Sillen <sup>10</sup>                      | 13 individuals with COPD; age: 65 years                                                                                       | Quadriceps                                                                              | Biphasic symmetrical pulses. Frequency = 75 Hz, width = 410 $\mu$ s. Duty cycle on:off = 6:29. Amplitude increased to maximal tolerable. 21 minute protocol.                                                                                                                                                                            |
| Sillen <sup>21</sup>                      | 17 individuals with COPD; age: 67 years                                                                                       | Quadriceps                                                                              | Biphasic symmetrical pulses. Low frequency protocol used a frequency of 15 Hz, width = 390 $\mu$ s, and duty cycle on:off = 8:2. 29 minute protocol. High frequency protocol used a frequency of 75 Hz, width = 410 $\mu$ s, and duty cycle on:off = 6:29. 21 minute protocol. Both protocols used an amplitude of maximum tolerable.   |

|                        |                                                                  |                                                                                            |                                                                                                                                                                                                                                              |
|------------------------|------------------------------------------------------------------|--------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Takeda <sup>44</sup>   | 15 healthy males; age: 23 years                                  | Quadriceps, hamstrings, and gluteal muscles                                                | Biphasic square pulses. Frequency = 4 Hz, width = 100 $\mu$ s. Amplitude set to maximum tolerable. 3 minute protocol.                                                                                                                        |
| Theurel <sup>46</sup>  | 12 healthy males; age: 25 years                                  | Quadriceps                                                                                 | Biphasic symmetrical pulses. Frequency = 75 Hz, width = 400 $\mu$ s Duty cycle on:off = 6.25:20. Amplitude set to maximal tolerable. Protocol involved 40 contractions in 17.5 minutes.                                                      |
| Tseh <sup>47</sup>     | 20 healthy males; age: 35 years                                  | Quadriceps                                                                                 | Frequency ranged from 2-5 Hz. Duty cycle on:off = 3:3. Amplitude set to a comfortably tolerable level.                                                                                                                                       |
| Valli <sup>15</sup>    | 13 healthy individuals (7M, 6F); age: 50.6 years                 | Quadriceps                                                                                 | 7 second contraction with frequency between 60 to 90 Hz, 15 second rest with a 20 Hz frequency. Amplitude set to maximal tolerable. 30 minute protocol.                                                                                      |
| Woelfel <sup>22</sup>  | 10 individuals with spinal cord injuries (9M, 1F); age: 39 years | Quadriceps and hamstrings                                                                  | The protocol involved both frequencies of 1 and 3 Hz each completed with amplitudes of 50 and 100 mA. Protocols lasted 6 minutes.                                                                                                            |
| Zinner <sup>48</sup>   | 11 healthy individuals; age: 24 years                            | Upper arms, upper and lower back, chest, and stomach                                       | Frequency = 85 Hz, width = 400 $\mu$ s, duty cycle on:off = 10:2. Amplitude was increased to maximal tolerable. 3 minute protocol.                                                                                                           |
| Pollack <sup>43</sup>  | 11 individuals with spinal cord injuries (7M, 4F); age: 29 years | Quadriceps                                                                                 | Monophasic pulses. Frequency = 30 Hz, width = 400 $\mu$ s. Amplitude set to where individuals could lift a 3lb load through 75% range of motion.                                                                                             |
| Figoni <sup>27</sup>   | 14 individuals with spinal cord injuries; age: 32 years          | Quadriceps                                                                                 | Biphasic square pulses. Frequency = 35 Hz, width = 300 $\mu$ s, amplitude = 150 mA. Completed a 4 minute protocol performing knee extensions with each a 0, 5, 10, and 15 kg load.                                                           |
| Miyamoto <sup>41</sup> | 18 individuals with type 2 diabetes (10M, 8F); age: 64 years     | Quadriceps, hamstrings, and gluteal muscles                                                | Monophasic square pulses. Frequency = 4 Hz, width = 200 $\mu$ s. Amplitude set to reach 6.0 ml/kg/min. 30 minute protocol.                                                                                                                   |
| Kimura <sup>34</sup>   | 10 obese and 4 pre-obese males; age 45 years                     | Quadriceps, hamstrings, and gluteal muscles                                                | Monophasic square pulses. Frequency = 4 Hz, width = 200 $\mu$ s. Amplitude set to maximal level without discomfort.                                                                                                                          |
| Miyamoto <sup>40</sup> | 11 males with type 2 diabetes; age: 57 years                     | Quadriceps, hamstrings, and gluteal muscles                                                | Monophasic square pulses. Frequency = 4 Hz, width = 200 $\mu$ s. Amplitude set to maximal without discomfort. 30 minute protocol.                                                                                                            |
| Poole <sup>18</sup>    | 33 individuals with type 2 diabetes (16M, 17F); age: 55 years    | Quadriceps and abdominal muscles                                                           | Frequency = 50 Hz. Amplitude was set to a max of 40 mA which equated to about 10% MVC. Performed 8 seconds contractions.                                                                                                                     |
| Teschler <sup>45</sup> | 16 healthy individuals (13M, 3F); age: 33.5 years                | Upper legs, upper arms, gluteals, abdomen, chest, lower back, upper back, latissimus dorsi | Frequency = 35 Hz, width = 350 $\mu$ s, duty cycle on:off = 7:4. Performed 1-2 sets of 6-8 repetitions for 8 different movements. Amplitude increased to achieve a perceived exertion of 7-8 (very hard, to very hard+). 20 minute protocol. |

Table 2. Study quality assessment

| Reference                                  | Q1 | Q2 | Q3 | Q4 | Q5 | Q6 | Q7 | Q8 | Q9 | Q10 | Total |
|--------------------------------------------|----|----|----|----|----|----|----|----|----|-----|-------|
| Abitante <sup>24</sup>                     | 1  | 1  | 1  | 0  | 0  | 1  | 1  | 0  | 0  | 1   | 6     |
| Ashley <sup>25</sup>                       | 1  | 0  | 1  | 1  | 0  | 1  | 1  | 0  | 0  | 1   | 6     |
| Banerjee <sup>17</sup>                     | 1  | 1  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 1   | 7     |
| Chen <sup>26</sup>                         | 1  | 1  | 1  | 1  | 0  | 1  | 1  | 0  | 1  | 1   | 8     |
| Fornusek <sup>28</sup>                     | 1  | 0  | 1  | 1  | 0  | 1  | 1  | 0  | 1  | 1   | 7     |
| Grosset <sup>30</sup>                      | 1  | 1  | 1  | 1  | 0  | 1  | 1  | 0  | 1  | 1   | 8     |
| Galvan <sup>29</sup>                       | 1  | 1  | 1  | 1  | 0  | 1  | 1  | 0  | 1  | 1   | 8     |
| Hamada <sup>32</sup>                       | 1  | 0  | 1  | 0  | 0  | 1  | 0  | 0  | 1  | 1   | 5     |
| Hamada <sup>31</sup>                       | 1  | 1  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 1   | 7     |
| Hirata <sup>33</sup>                       | 1  | 0  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 1   | 6     |
| Hsu <sup>13</sup>                          | 1  | 0  | 1  | 0  | 1  | 1  | 1  | 0  | 0  | 1   | 6     |
| Isakov <sup>14</sup>                       | 1  | 1  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 0   | 6     |
| Janssen <sup>19</sup>                      | 1  | 0  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 1   | 6     |
| Lago <sup>35</sup>                         | 1  | 1  | 1  | 1  | 0  | 1  | 1  | 0  | 1  | 1   | 8     |
| Laskin <sup>36</sup>                       | 1  | 0  | 1  | 0  | 0  | 0  | 1  | 0  | 1  | 1   | 5     |
| Ma <sup>23</sup>                           | 1  | 1  | 1  | 1  | 0  | 0  | 1  | 0  | 1  | 1   | 7     |
| Matsushima <sup>37</sup>                   | 1  | 1  | 1  | 1  | 0  | 1  | 1  | 0  | 1  | 1   | 8     |
| Minogue <sup>39</sup>                      | 1  | 0  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 0   | 5     |
| Minogue <sup>38</sup>                      | 1  | 0  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 1   | 6     |
| Miyamoto <sup>42</sup>                     | 1  | 0  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 1   | 6     |
| Perez-De-Arrilucea-Le-Floc'h <sup>20</sup> | 1  | 1  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 1   | 7     |
| Sillen <sup>10</sup>                       | 1  | 0  | 1  | 0  | 0  | 1  | 1  | 0  | 0  | 1   | 5     |
| Sillen <sup>21</sup>                       | 1  | 1  | 1  | 0  | 1  | 1  | 1  | 0  | 1  | 1   | 8     |
| Takeda <sup>44</sup>                       | 1  | 1  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 1   | 7     |
| Theurel <sup>46</sup>                      | 1  | 0  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 1   | 6     |
| Tsch <sup>47</sup>                         | 1  | 1  | 1  | 1  | 1  | 1  | 1  | 0  | 1  | 1   | 9     |
| Valli <sup>15</sup>                        | 1  | 1  | 1  | 0  | 0  | 1  | 0  | 0  | 1  | 0   | 5     |
| Woelfel <sup>22</sup>                      | 1  | 1  | 1  | 1  | 0  | 1  | 1  | 0  | 1  | 1   | 8     |
| Zinner <sup>48</sup>                       | 1  | 0  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 1   | 6     |
| Pollack <sup>43</sup>                      | 1  | 1  | 1  | 1  | 0  | 1  | 1  | 0  | 1  | 1   | 8     |
| Figoni <sup>27</sup>                       | 1  | 0  | 1  | 1  | 0  | 1  | 1  | 0  | 1  | 1   | 7     |
| Miyamoto <sup>41</sup>                     | 1  | 0  | 1  | 0  | 1  | 1  | 1  | 0  | 1  | 1   | 7     |
| Kimura <sup>34</sup>                       | 1  | 0  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 1   | 6     |
| Miyamoto <sup>40</sup>                     | 1  | 0  | 1  | 0  | 0  | 1  | 1  | 0  | 1  | 1   | 6     |
| Poole <sup>18</sup>                        | 1  | 1  | 1  | 0  | 1  | 1  | 0  | 0  | 1  | 1   | 7     |
| Teschler <sup>45</sup>                     | 1  | 1  | 1  | 1  | 1  | 1  | 1  | 0  | 1  | 1   | 9     |

Q1: Was the study question or objective clearly stated?

Q2: Were eligibility/selection criteria for the study population prespecified and clearly described?

Q3: Were the participants in the study representative of those who would be eligible for the test/service/intervention in the general or clinical population of interest?

Q4: Were all eligible participants that met the prespecified entry criteria enrolled?

Q5: Was the sample size sufficiently large to provide confidence in the findings?

Q6: Was the test/service/intervention clearly described and delivered consistently across the study population?

Q7: Were the outcome measures prespecified, clearly defined, valid, reliable, and assessed consistently across all study participants?

Q8: Were the people assessing the outcomes blinded to the participants' exposures/interventions?

Q9: Was the loss to follow-up after baseline 20% or less? Were those lost to follow-up accounted for in the analysis?

Q10: Did the statistical methods examine changes in outcome measures from before to after the intervention? Were statistical tests done that provided p values for the pre-to-post changes?