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Consistency Among Musculoskeletal Models: Caveat Utilitor

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Abstract—Musculoskeletal simulation software and model repositories have broadened the user base able to perform musculoskeletal analysis and have facilitated in the sharing of models. As the recognition of musculoskeletal modeling continues to grow as an engineering discipline, the consistency in results derived from different models and software is becoming more critical. The purpose of this study was to compare eight models from three software packages and evaluate differences in quadriceps moment arms, predicted muscle forces, and predicted tibiofemoral contact forces for an idealized knee-extension task spanning -125 to $+10^\circ$ of knee extension. Substantial variation among models was observed for the majority of aspects evaluated. Differences among models were influenced by knee angle, with better agreement of moment arms and tibiofemoral joint contact force occurring at low to moderate knee flexion angles. The results suggest a lack of consistency among models and that output differences are not simply an artifact of naturally occurring inter-individual differences. Although generic musculoskeletal models can easily be scaled to consistent limb lengths and use the same muscle recruitment algorithm, the results suggest those are not sufficient conditions to produce consistent muscle or joint contact forces, even for simplified models with no potential of co-contraction.

Keywords—Musculoskeletal models, Muscle moment arm, Joint contact force, Muscle recruitment, Musculoskeletal simulation, Knee flexion.

INTRODUCTION

Software packages specifically designed to facilitate the development and analysis of musculoskeletal models

(e.g., AnyBody,¹³ BoB,⁵⁷ LifeModeler (<http://www.lifemodeler.com>), Opensim,¹⁵ SIMM¹⁶) have led to the expansion of musculoskeletal simulations. Additionally, model repositories (e.g., AnyBody Repository (<http://forge.anyscript.org/gf/>), PhysiomeSpace (www.physioimespace.com/), Simtk.org) have made possible the sharing and distribution of musculoskeletal models, which have allowed different researchers and users to more easily expand or incorporate previous work not developed locally. One early example of such a repository that contained model parameters of the lower limb (<http://isbweb.org/data/delp/index.html>) demonstrates the potential and impact that musculoskeletal data, made available to the research community, can have with the primary manuscript associated with the dataset¹⁷ currently having 533 citations (Scopus, accessed 5/9/2013). The widespread use of this data set over the past two decades can in part be explained by the considerable time and effort required to develop mathematical representations of anatomical structures.

Musculoskeletal models have been used to investigate a wide range of research topics including physiological loading,^{33,45,57,58,64} wheelchair propulsion,²⁰ reaching,⁶⁰ ergonomic evaluation,^{1,49,63} and design optimization.^{31,50} Musculoskeletal simulation software, which can be used to estimate quantities difficult to measure non-invasively (e.g., muscle force, joint contact force), has not only been developed to quantify absolute internal body forces, but also with the intent of examining the effect of an environmental or postural change on model performance (e.g., stability, muscle function).^{13,55} Analysis of such cause-effect relationships has great potential for incorporating internal body measures into device and component design.^{30,32} The same relationships have also been

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80 proposed as a method for validating certain compo-
 81 nents of musculoskeletal simulations.⁴² Generic human
 82 figure models widely used in the related field of ergo-
 83 nomics can be scaled to population-based anthropo-
 84 metric measurements to evaluate accommodation and
 85 other engineering-based design goals.²² In a similar
 86 capacity, the use of scaled generic musculoskeletal
 87 models has the potential to be used as an engineering
 88 tool in which individualized patient assessment is not
 89 required. Additionally, compared to image-based
 90 models defined using individual-specific scan data,
 91 analyses with generic models are not burdened by
 92 expensive scan costs and lengthy image processing
 93 times.^{9,62}

94 Verification and validation of newly developed
 95 and currently available musculoskeletal models are
 96 non-trivial tasks and remain topics of ongoing
 97 research.^{14,27,42} Recent studies have investigated the
 98 comparative accuracy of scaled generic musculoskele-
 99 tal models to that of subject-specific geometry, and the
 100 effect of those differences on computed muscle
 101 moment arm,^{6,52–54} functional roles of muscles during
 102 gait,¹² and joint contact force.^{27,45} Validation among
 103 models is also necessary, with the expectation of users
 104 that the same analyses performed with different models
 105 or software will produce consistent results.⁶¹ It is not
 106 known whether this expectation is currently being met
 107 and/or to what capacity users of different models must
 108 scale or adapt those models to yield consistent results.

109 Mathematical models of the knee joint and its sur-
 110 rounding muscles have been used to better understand
 111 a wide array of topics including cruciate ligament
 112 function,² the interaction of muscle activation and
 113 knee injury during frontal car crashes,¹¹ and knee joint
 114 reaction loading during walking.²⁷ One application
 115 relevant to our laboratory is the use of generic mus-
 116 culoskeletal models for evaluating exercise therapies
 117 and interventions for individuals with spinal cord
 118 injury (SCI). Joint reaction force at the knee has pre-
 119 viously been used to compare different exercises and
 120 quantify internal loading during exercise participation,
 121 including those with a functional electrical stimulation
 122 component.^{5,21,28,35,44} In the context of skeletal health,
 123 an issue particularly relevant to individuals with SCI,³⁶
 124 both trend and absolute estimates of knee force can aid
 125 in the design or adaptation of an exercise. To our
 126 knowledge, there exist no directly measured data (e.g.,
 127 instrumented endoprostheses) that can be used to
 128 compare to the knee joint reaction force output of
 129 musculoskeletal models simulating exercise therapies
 130 or interventions for individuals with SCI. Therefore,
 131 indirect validation of the overall musculoskeletal
 132 model appears to remain the optimal method for
 133 gaining confidence in the simulation results. The model
 134 may in fact provide the best available estimate to the

135 internal loading within the actual system. However, in
 136 the context of this application, it remains unclear if the
 137 selection of the generic model substantially influences
 138 the accuracy and/or interpretation of the results.

139 The overall goal of this study was to compare the
 140 results of several commonly available generic muscu-
 141 loskeletal models, scaled to consistent anthropometry,
 142 in determining moment arms, muscle force contribu-
 143 tions, and predicted knee joint contact force during an
 144 idealized knee-extension task for postures spanning an
 145 extended and substantially flexed knee. To simplify the
 146 comparisons, simplified musculoskeletal models that
 147 only included the quadriceps muscles were used. Our
 148 first study aim was to quantify the differences in the
 149 lengths of the quadriceps moment arms between
 150 models, particularly at postures of high knee flexion.
 151 Our second study aim was to explore absolute and
 152 trend differences in simulated muscle recruitment and
 153 joint contact force between models. Our final aim was
 154 to identify future research questions and topics that
 155 will aid in the consistency of results produced by dif-
 156 ferent musculoskeletal modeling models and software
 157 packages.

158 MATERIALS AND METHODS

159 Quadriceps muscle moment arms and tibiofemoral
 160 joint contact for a simulated knee extension task were
 161 computed for several musculoskeletal models spanning
 162 three unique musculoskeletal simulation software
 163 environments. Models were anthropometrically scaled
 164 to have consistent limb length dimensions. Muscle
 165 moment arms were computed for eight unique mus-
 166 culoskeletal models. Tibiofemoral joint contact loads
 167 were computed for a subset of five models that had the
 168 capability for computing tibiofemoral joint loading
 169 during a simplified isotonic knee extension task.
 170 Results are presented over knee angles ranging from
 171 -125° to $+10^\circ$ knee extension. Knee angles of -20° ,
 172 corresponding to peak knee flexion during mid-stance
 173 of normal gait,⁴⁶ and -100° , corresponding to peak or
 174 sub-peak knee flexion during activities that include
 175 stair ascent, stair descent, cycling, leg press, sit to
 176 stand, power lifting, squatting and FES row-
 177 ing,^{23–26,34,43,67} are also used to compare intra and
 178 inter-model differences for minimal and deep knee
 179 flexion postures.

180 *Musculoskeletal Models*

181 Eight musculoskeletal models that included lower
 182 extremity musculature (Table 1) were evaluated (see
 183 Appendix—Table 7 for model accessibility). The se-
 184 lected models were implemented in the AnyBody (<http://>

185 www.anybodytech.com), OpenSim (<http://opensim.stanford.edu/>), or Biomechanics of Bodies (BoB) modeling software packages. Prior to testing, each model was scaled to the joint-to-joint dimensions listed in Table 2. Off-axis bone dimensions were scaled isometrically. Each model was simplified to only include representations for four quadriceps muscle groups (vastus lateralis, vastus intermedius, vastus medialis, and rectus femoris). All muscles were modeled using a Hill-type representation.⁷⁰ The model-defined values of maximum muscle strength at optimal fiber length (Table 3) were not changed. Additional differences between muscle model representations and parameters (e.g., optimal fiber length, pennation angle, *etc.*) are not presented.

199 Muscle path representation, a component that contributes to the effective muscle moment arm, varied among models. The AnyBody and Biomechanics of Bodies musculoskeletal models represented muscle paths as line segments defined by insertion, origin, and intermediate *via* points. *Via* points are frictionless constraints at one or more locations along the path of the muscle. The Delp 1990, Gait 2392, and Steele 2012 models used *via* points that depended on posture. The London Lower Limb and Lower Limb 2010 models defined the path of each quadriceps muscle based on insertion and origin points and idealized surface geometry used to represent underlying physiological structures around which a muscle wraps.⁶ The AnyBody—LegTD and London Lower Limb models, based on the same cadaver dataset,³⁷ represent each quadriceps muscle using multiple muscle fascicles while the remaining models represent each quadriceps muscle with a single muscle unit. For example, in both models with multiple muscle fascicles, the vastus intermedius is represented as 6 separate fascicles attached at two insertion points on the proximal aspect of the patella, and multiple muscle origins along the femur. The reported muscle strengths are the sum of all the muscle fascicles representing that single muscle (Table 3).

224 The kinematic knee joint definition, another component that contributes to the effective muscle moment arm, was not consistent among all models. The AnyBody—Leg, AnyBody—LegTD, and London Lower Limb models define the tibiofemoral joint kinematics as an idealized hinge (revolute) joint. The Delp 1990, Gait 2392, and Steele 2012 models define the tibiofemoral kinematics as a single coordinate with coupled rotation and translation.⁶⁹ The Lower Limb 2010 model defines the tibiofemoral kinematics based on experimental data presented in Walker *et al.*⁶⁵ The BoB model defines the tibiofemoral kinematics as two rolling cylinders with radii approximated from Leszko *et al.*⁴¹ The AnyBody—LegTD and London Lower Limb models define the patellar kinematics as a circular path defined in the local femur reference frame and is prescribed by the tibiofemoral knee angle. For those models, the patellar position maintains a constant patellar tendon length throughout the knee range of motion. The AnyBody—Leg model does not have a patellar body but includes a quadriceps muscle *via* point in the approximate location of the patella with the quadriceps muscles attached to the proximal tibia. The Gait 2392 model does not include a patella. The Delp 1990 model includes a patella body with its position defined by 4 coordinates (3 translational, 1 rotation), each functionally prescribed by the tibiofemoral knee angle, with respect to the local tibial

TABLE 2. Lower extremity scaled model dimensions.

| Scaled dimension ^a | Value | Definition |
|-------------------------------|-------|---|
| Pelvis width (m) | 0.166 | Left to right hip joint center |
| Thigh length (m) | 0.434 | Hip to knee joint center |
| Shank length (m) | 0.428 | Knee to ankle joint center |
| Body mass (kg) | 74 | Whole body mass |
| Body height (m) | 1.75 | Not used in scaling, for reference only |

^aDimensions based on scaled 'AnyBody—Leg' model to 50th percentile male by stature.

TABLE 1. Musculoskeletal models used to compute model-predicted moment arms.

| Model name | Software package | References |
|-----------------------------------|-------------------|---|
| AnyBody—Leg | AnyBody (v 4.1.0) | Damsgaard <i>et al.</i> ¹³ |
| AnyBody—LegTD | AnyBody (v 4.1.0) | Andersen <i>et al.</i> ⁴ |
| Biomechanics of Bodies (BoB v3.0) | Matlab (v 7.12) | Shippen and May ⁵⁷ |
| Delp 1990 ^a | Opensim (v 2.4.0) | Delp <i>et al.</i> ¹⁷ |
| Steele 2012 | Opensim (v 2.4.0) | Steele <i>et al.</i> ⁵⁸ |
| Gait 2392 | Opensim (v 2.4.0) | http://simtk-confluence.stanford.edu:8080/x/54Mz |
| London Lower Limb | Opensim (v 2.4.0) | Modenese <i>et al.</i> ⁴⁵ |
| Lower Limb 2010 | Opensim (v 2.4.0) | Arnold <i>et al.</i> ⁷ |

^aAs implemented in the Opensim model 'BothLegs.osim'.

TABLE 3. Maximum quadriceps muscle strengths for the different models.

| Model name | Maximum isometric strength at optimal fiber length (N) | | | |
|--------------------------------|--|--------------------|-----------------|----------------|
| | Vastus lateralis | Vastus intermedius | Vastus medialis | Rectus femoris |
| AnyBody—Leg ^a | 1852 | 1224 | 1283 | 773 |
| AnyBody—LegTD ^b | 1882 | 1029 | 1617 | 780 |
| Biomechanics of Bodies | 1870 | 1235 | 1295 | 780 |
| Delp 1990 | 1871 | 1365 | 1294 | 779 |
| Steele 2012 | 1871 | 1365 | 1294 | 1169 |
| Gait 2392 | 1871 | 1365 | 1294 | 1169 |
| London Lower Limb ^b | 2579 | 1410 | 2216 | 1069 |
| Lower Limb 2010 | 2255 | 1024 | 1444 | 849 |

^aMuscle strengths were scaled based on thigh mass using standard software pipeline.

^bMuscle strengths are the sum of individual fascicles used to represent each muscle.

252 reference frame. The Steele 2012 and Lower Limb 2010
253 models include a patella body with its kinematics
254 defined by 3 coordinates (2 translational, 1 rotational),
255 each functionally prescribed by the tibiofemoral knee
256 angle, with respect to the local femur reference frame.
257 The BoB model includes a patella with its kinematics
258 defined from Azmy *et al.*⁸ with the patella translations
259 and rotations defined as a function of knee flexion angle
260 encoded using a cubic interpolating look-up table.

261 *Muscle Moment Arms—Quadriceps*

262 Model-predicted moment arm data were obtained
263 using the same method for all models using a direct
264 load measurement method, previously summarized by
265 An *et al.*³ Sub-models of each musculoskeletal model
266 were constructed with only the single muscle (or group
267 of muscle fascicles representing a single muscle) to be
268 evaluated. An external unit torque was applied about
269 the rotational axis of the knee. Knee flexion was varied
270 between -125° and $+10^\circ$ (knee extension) over a time
271 of 1000 s to approximate a quasi-static analysis at each
272 analyzed posture. Hip flexion, abduction, and internal
273 rotation were defined to be 90° , 0° , and 0° , respectively.
274 Muscle and tendon force for each model was computed
275 using a static optimization procedure incorporated
276 into each software package that the models were con-
277 structed in. Although an optimization procedure was
278 used for the moment-arm analysis, the results are
279 deterministic since only one muscle was included in
280 each model and the muscle and connected skeletal
281 linkage was modeled as a deterministic system (as
282 opposed to a stochastic representation). The muscle
283 moment arm at each knee angle was computed as the
284 applied torque divided by the computed tendon force.
285 The force of gravity was reduced to zero for each
286 model. The computed moment-arms for the models
287 implemented in OpenSim were essentially equivalent to
288 the moment-arms given by the software's muscle
289 moment arm calculation function.⁵⁶

Tibiofemoral Joint Contact Force

290
291 Model-predicted tibiofemoral joint contact forces
292 were obtained for a simulated task of knee extension.
293 The method of load application and evaluated knee
294 postures was similar to the muscle moment arm deri-
295 vation previously described. A constant external knee
296 flexion torque of 90 N-m was used in each simulation.
297 Each musculoskeletal model included representations
298 of all four components of the quadriceps. Individual
299 muscle strengths, paths, and muscle model parameters
300 were not changed from their default values following
301 anthropometric scaling. Muscle forces were computed
302 using a static optimization procedure that minimized
303 the sum of squared muscle activations. Tibiofemoral
304 joint contact forces were computed within each mus-
305 culoskeletal software program and reported in the
306 local tibial reference frame defined by each model. The
307 overall magnitude of the joint contact force is reported
308 here to facilitate comparisons between models.

RESULTS

Moment Arms

311 The difference between the moment arms for the
312 individual quadriceps muscles within a single model
313 was relatively small. The maximum intra-model
314 moment arm difference was 1.33 cm and occurred for
315 the BoB model with a knee extension angle of 10° . At
316 each knee angle, the quadriceps moment arms were
317 equal for the AnyBody-Leg model with the exception
318 of the rectus femoris, which was not able to produce a
319 knee extension torque between -22° and $+10^\circ$ of knee
320 extension. The mean intra-model quadriceps muscle
321 moment arm difference across models (excluding the
322 AnyBody-Leg model) over the range of motion tested
323 (-125° to $+10^\circ$ knee extension) was 0.44 cm. For knee
324 flexion angles greater than 20° , the maximum moment
325 arm difference for all models was 0.68 cm, which

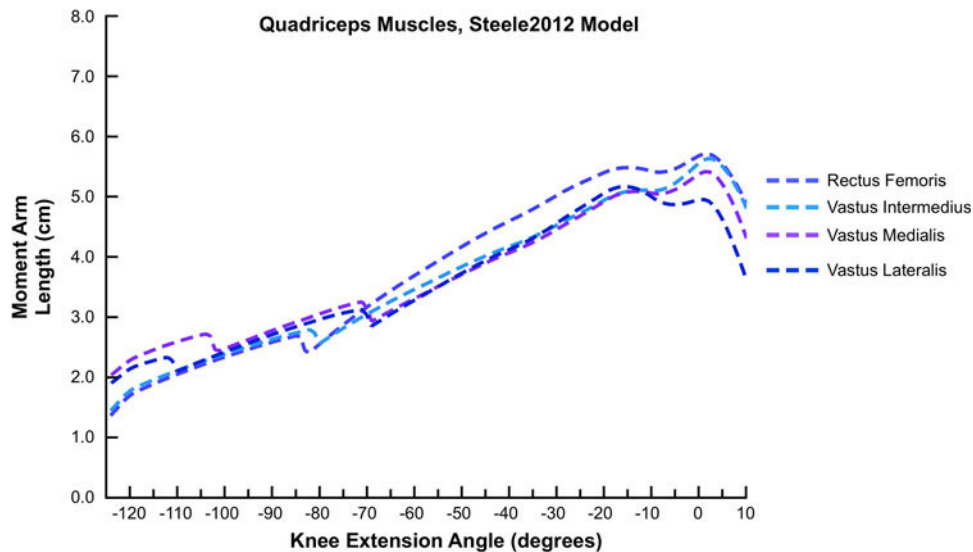


FIGURE 1. Quadriceps muscle moment arms for the Steele 2012 model.

TABLE 4. Maximum difference in quadriceps muscle moment arm for each musculoskeletal model at 20° and 100° of knee flexion.

| Model | Moment arm difference at 20° (cm) | Moment arm difference at 100° (cm) |
|-------------------|-----------------------------------|------------------------------------|
| AnyBody—Leg | 0.0 ^a | 0.0 |
| AnyBody—LegTD | 0.15 | 0.35 |
| BoB | 0.55 | 0.30 |
| Delp 1990 | 0.46 | 0.19 |
| Steele 2012 | 0.48 | 0.14 |
| Gait 2392 | 0.57 | 0.08 |
| London Lower Limb | 0.65 | 0.12 |
| Lower Limb 2010 | 0.40 | 0.27 |

^aExcluding rectus femoris, which could not produce a knee extension moment in this posture.

TABLE 5. Maximum muscle moment arm change observed in the quadriceps muscle group for knee extension angles spanning -125 to +10° for each model.

| Model | Muscle(s) | Minimum (cm) | Maximum (cm) | Range (cm) |
|-------------------|------------------|--------------|--------------|------------|
| AnyBody—Leg | All vasti | 1.50 | 5.03 | 3.53 |
| AnyBody—LegTD | Vastus medialis | 1.65 | 6.17 | 4.53 |
| BoB | Vastus medialis | 3.07 | 3.84 | 0.78 |
| Delp 1990 | Rectus femoris | 2.02 | 5.11 | 3.09 |
| Steele 2012 | Rectus femoris | 1.27 | 5.70 | 4.43 |
| Gait 2392 | Rectus femoris | 4.73 | 7.53 | 2.80 |
| London Lower Limb | Vastus medialis | 2.67 | 6.87 | 4.20 |
| Lower Limb 2010 | Vastus lateralis | 1.33 | 4.93 | 3.60 |

326 occurred in the Steele 2012 model at maximum knee
327 flexion (-125°) between the vastus medialis and rectus
328 femoris (Fig. 1). Table 4 summarizes the intra-model
329 moment arm differences at 20 and 100° knee flexion.

330 In general, the quadriceps moment arms decreased
331 as the knee extended beyond -20° . The exception to
332 this trend occurred in the BoB model, which exhibited
333 consistent moment arms throughout the evaluated
334 range of motion. Within each model, the maximum
335 length change of a single quadriceps muscle moment
336 arm over the evaluated knee range of motion (Table 5)
337 spanned from 0.78 cm (BoB) to 4.53 cm (Any-
338 Body—LegTD). Table 6 summarizes the computed
339 quadriceps moment arms for each model at 20 and
340 100° knee flexion.

341 The eight scaled musculoskeletal models have both
342 different absolute lengths of the quadriceps moment
343 arms and different trends over the evaluated knee

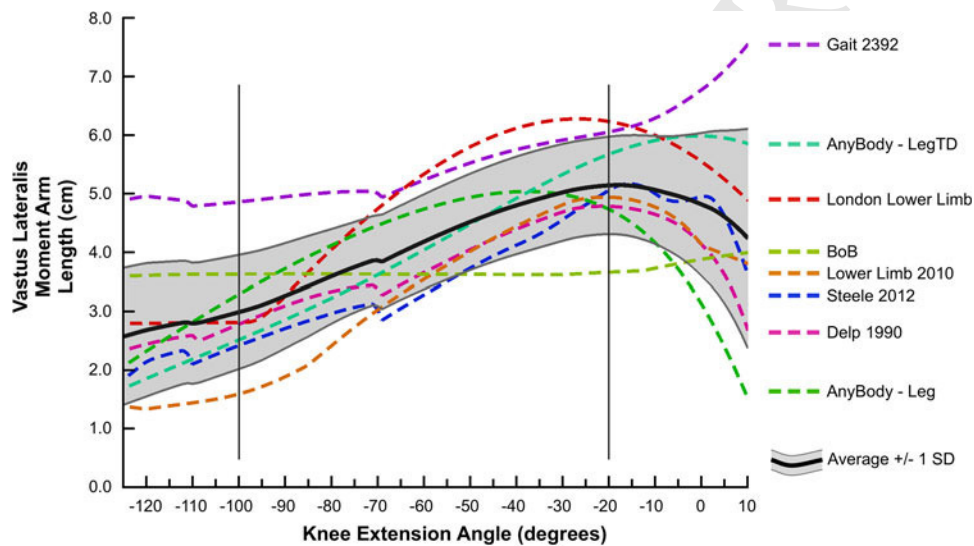
range of motion. The moment arms of the vastus 344
lateralis for the different models are presented in 345
Fig. 2. Similar results were observed for the vastus 346
medialis, vastus intermedius, and the rectus femoris 347
(not shown). No single model resulted in either the 348
highest or lowest moment arm limits over the range of 349
knee angles evaluated. 350

351 The greatest inter-model agreement, identified by the
352 coefficient of variation (COV), was observed between
353 knee flexion angles of -10 and -60° , angles nearly
354 spanning those observed in normal gait⁴⁶ (Fig. 3). For
355 knee flexion angles approaching either end of the range
356 of motion limits, the coefficient of variation exceeded
357 2.5 times the minimum value observed at 23° knee
358 flexion. Excluding the BoB and Gait 2392 models,
359 which have different qualitative trends for the moment
360 arm versus knee extension angle as the other models
361 and previously reported data,¹⁰ the minimum coeffi-
362 cient of variation value decreases from 0.16 to 0.11, the
363 maximum coefficient of variation for deep knee flexion

TABLE 6. Quadriceps muscle moment arms for each musculoskeletal model at 20° and 100° of knee flexion.

| Model | Moment arms at 20° knee flexion (cm) | | | | Moment arms at 100° knee flexion (cm) | | | |
|-------------------|--------------------------------------|------|------|------|---------------------------------------|------|------|------|
| | VL ^a | VM | VI | RF | VL | VM | VI | RF |
| AnyBody—Leg | 4.72 | 4.72 | 4.72 | 0.0 | 3.28 | 3.28 | 3.28 | 3.28 |
| AnyBody—LegTD | 5.66 | 5.61 | 5.72 | 5.77 | 2.50 | 2.35 | 2.64 | 2.70 |
| BoB | 3.65 | 3.68 | 3.91 | 4.20 | 3.62 | 3.81 | 3.91 | 3.92 |
| Delp 1990 | 4.77 | 4.64 | 4.64 | 5.10 | 2.77 | 2.87 | 2.77 | 2.69 |
| Steele 2012 | 5.05 | 4.91 | 4.93 | 5.39 | 2.41 | 2.46 | 2.38 | 2.33 |
| Gait 2392 | 6.04 | 5.66 | 5.63 | 6.20 | 4.85 | 4.93 | 4.93 | 4.85 |
| London Lower Limb | 6.22 | 6.87 | 6.57 | 6.55 | 2.80 | 2.68 | 2.80 | 2.75 |
| Lower Limb 2010 | 4.93 | 4.80 | 4.82 | 5.20 | 1.58 | 1.57 | 1.76 | 1.84 |

^aVL—Vastus Lateralis, VM—Vastus Medialis, VI—Vastus Intermedius, RF—Rectus Femoris.

**FIGURE 2. Muscle moment arms for the vastus lateralis.**

364 decreases from 0.46 to 0.25, and the inter-model
365 agreement remains relatively unchanged for straight
366 and hyper-extended knee postures. The variation
367 among moment arms between similar anthropometri-
368 cally scaled (isometrically) musculoskeletal models is
369 comparable to the variation previously reported
370 between subjects from previous studies (Fig. 3).

371 *Muscle Force*

372 Quadriceps muscle recruitment was compared for
373 seven of eight musculoskeletal models. Muscle forces
374 for the vastus lateralis, vastus medialis, vastus inter-
375 medius, and rectus femoris were computed for each
376 model during the same simulated knee-extension task.
377 For each knee angle, the distribution of quadriceps
378 muscle forces to produce a 90 N-m knee extension
379 torque was computed. Results are presented for all the
380 models for knee flexion angles of 20 and 100° (Fig. 4).

381 All models had an increase in the combined quadriceps
382 muscle force between 20 and 100° knee flexion, with an
383 average increase of 1351 N. At 100° knee flexion, the
384 contribution of the vastus lateralis to the combined
385 quadriceps force was reasonably consistent between 30
386 and 49 percent. In contrast, at 20° knee flexion, the
387 contribution of the vastus lateralis ranged from 14 to
388 82 percent. Of the combined 14 evaluated models and
389 postures, the vastus lateralis contributed the largest
390 percentage of all muscles in 11 of the analyses. Within
391 each model, the contribution of force associated
392 with the vastus medialis and vastus intermedius was
393 fairly consistent. The difference in force contribution
394 between those muscles within each model was always
395 less than 6%, with the exception of the AnyBody—Leg
396 (20° knee flexion) and the AnyBody—LegTD models,
397 which had differences between the force contribution
398 from the vastus intermedius and vastus medialis
399 upwards of 20%.

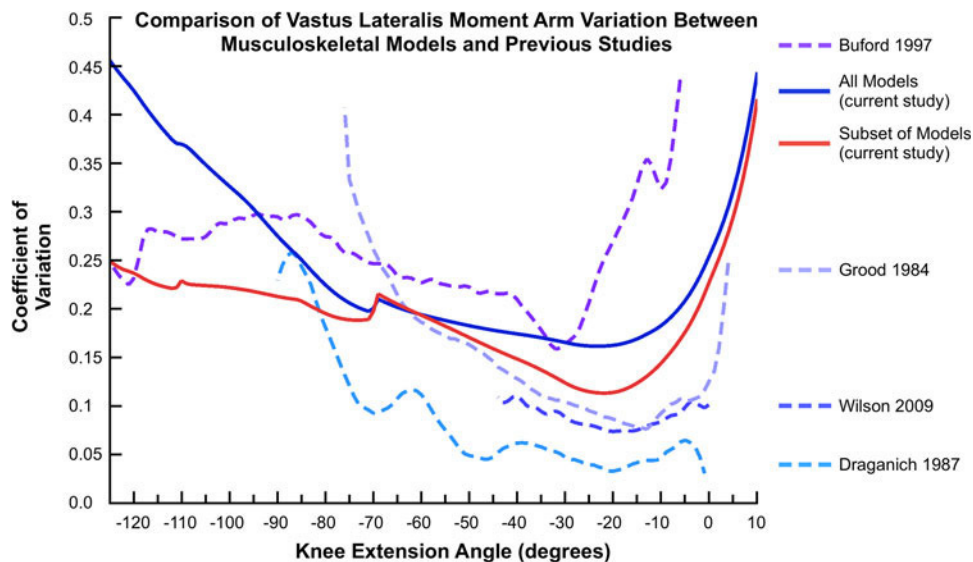


FIGURE 3. Coefficient of variation for the vastus lateralis moment arm for the musculoskeletal models and previous studies. The 'subset of models' group excludes the Gait 2392 and BoB musculoskeletal models.

400

Tibiofemoral Joint Contact Force

401 The magnitude of the resultant force vector of the
 402 tibiofemoral joint contact force was calculated for a subset
 403 of the models for the 90 N-m knee extension torque task.
 404 Results from the five musculoskeletal models that could
 405 be directly used to compute the knee joint contact force
 406 are presented in Fig. 5. The within model range of knee
 407 joint contact force spanned 219 N and 4204 N for the
 408 BoB and AnyBody—Leg models, respectively, over the
 409 range of knee angles evaluated. Both the Steele 2012 and
 410 the AnyBody—Leg models exhibited a substantial
 411 increase in tibiofemoral contact force as knee extension
 412 angle decreased past -50° . In contrast, the remaining
 413 three models had only slight changes in joint reaction
 414 force above and below -50° knee extension. At -100°
 415 knee extension, the knee joint reaction force ranged from
 416 1839 to 3754 N between models, a difference of 2.6 body
 417 weights. In contrast, the knee joint reaction force ranged
 418 from 1525 to 2269 N at -20° knee extension, a difference
 419 of approximately one body weight.

420

DISCUSSION

421 The study compared knee extensor moment arms,
 422 muscle force predictions, and knee joint contact force
 423 predictions for several similarly scaled musculoskeletal
 424 models available to the biomechanics community.
 425 Substantial variation among models was observed for
 426 all aspects evaluated. The one exception was the rela-
 427 tively consistent (among models) within-model
 428 moment arm range spanned by the quadriceps muscle
 429 group of each model (e.g., Fig. 1). Differences between
 430 models were influenced by knee angle, with better

inter-model agreement occurring at knee flexion angles
 in the range from 10 to 60° .

The within-model moment arm range was relatively
 small and always less than 1.33 cm for each of the
 models evaluated. A slightly smaller value was observed
 in data presented by Klein Horsman,³⁸ which showed a
 maximum range between individual quadriceps muscle
 moment arms of less than 1 cm for knee extension
 angles spanning -135 to 0° for a single cadaver speci-
 men. Similarly, the maximum range of the averaged
 moment arms (15 cadaver specimens) for the different
 quadriceps muscles presented by Buford *et al.*¹⁰ was also
 less than 1 cm for a similar range of knee angles. In both
 studies the maximum range occurred at small angles of
 knee extension (i.e., near full leg extension), similar to
 the models evaluated in this study. The average of the
 maximum moment arm differences sampled at each knee
 posture for the Buford *et al.*¹⁰ and Klein
 Horsman³⁸ cadaver studies were 0.47 and 0.44 cm,
 respectively, similar to the average of 0.44 cm of the
 eight evaluated musculoskeletal models. The results
 suggest the musculoskeletal models appear to be
 reasonably consistent, with each other and previous
 cadaveric studies, in representing the moment arm intra-
 specimen variability of the quadriceps muscle group.

In attempting to apply the formal concepts of veri-
 fication and validation to musculoskeletal modeling,
 Lund *et al.*⁴² states that, "...verification is a pre-
 requisite for validation. Verification provides the evi-
 dence that the computer code correctly solves the
 underlying mathematical model. Absence of verifica-
 tion creates the risk of mixing modeling errors and
 errors caused by implementation." Verification is an
 important topic; however the study conducted here

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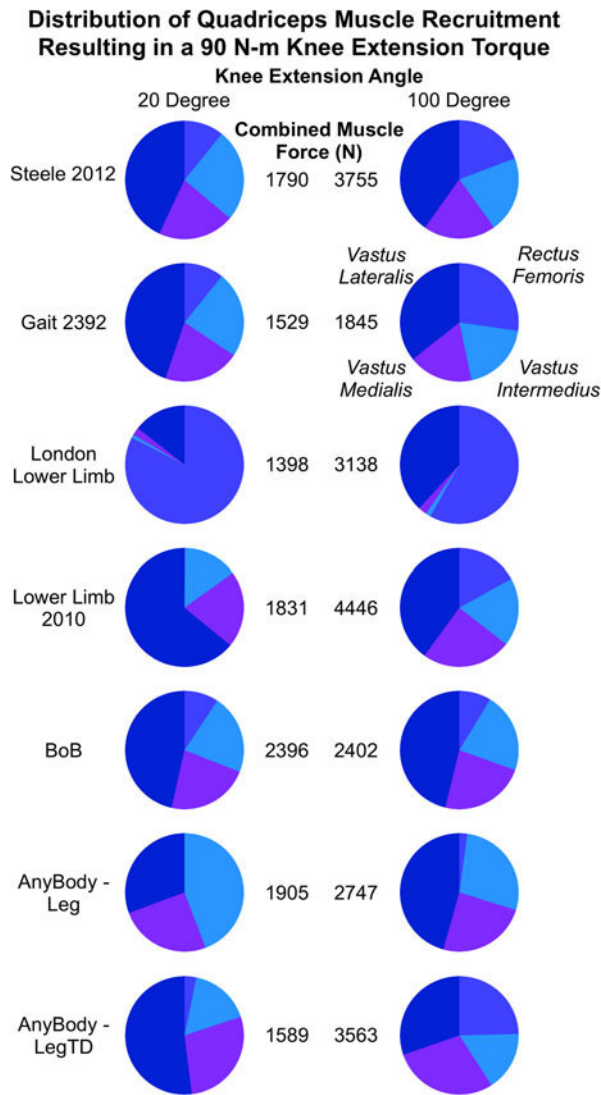


FIGURE 4. Contribution of quadriceps muscle force to produce a 90 N-m knee extension torque at 20 and 100° knee flexion.

465 focused on validation and not verification. For example, the results from the static optimization analyses performed in the current study were not explicitly checked and it was assumed they were consistent with the equations of motion for the defined system. Lund *et al.*⁴² further defined the examination of the “correctness of variable interaction” as trend validation, a concept that has been previously used to evaluate musculoskeletal model performance and understand changes in knee loads for different walking styles.^{14,47} Comparing the models tested in this study in the context of variable interaction, the majority of models (7 of 8) did exhibit smaller muscle moment arms at large angles of knee flexion compared to moderate or low knee flexion angles, a result consistent with previous studies.^{10,29,68} Two of the eight models had maximum moment arm values for the vastus lateralis

482 at +10° knee extension (hyper-extension), the maximum knee extension angle evaluated. The remaining models exhibited maximum vastus lateralis moment arms at slightly flexed knee postures, an observation more consistent with previous studies.^{29,68} The general consistency of these variables between musculoskeletal models is encouraging and suggests similar interpretations from a trend type analysis may be achieved when using the majority of the available models.

491 The results suggest a lack of absolute consistency in the tested musculoskeletal models and that model differences are not simply an artifact of naturally occurring inter-individual differences. Although the data used to develop the individual musculoskeletal models in this study were not from a consistent or nominal population, the expectation of the generically scaled models evaluated in this study is that they each represent the mean anatomy of a male individual with 50th percentile stature. It is currently difficult to evaluate whether a single musculoskeletal model accurately represents such mean anatomy, potentially explaining the differences between models observed here, as there is limited data available and differences due to inter-individual variation are unknown. The available data quantifying the variability for the vastus lateralis moment arm is not consistent. Using data from previous studies, moment arm standard deviations (averaged across the available knee angles) for subjects include values of: 3.02 cm,¹⁰ 2.30 cm,¹⁹ 0.43 cm,⁶⁸ and 0.38 cm.²⁹ In comparing a musculoskeletal model to literature values, Klein Horsman³⁸ assumed absolute differences smaller than 2 cm could be attributed to inter-individual differences. In contrast to the previous literature, the average standard deviation (over all knee angles) of the vastus lateralis moment arm for the models evaluated in this study was 0.95 cm, with the maximum inter-model difference ranging from 2.0 to 6.0 cm.

519 Isometric scaling was applied to scale the off-axis skeletal dimensions using the same scaling factors applied to define the limb lengths in an effort to generate consistent musculoskeletal models. However, width and breadth anthropometric dimensions are not as well correlated with stature as limb length dimensions⁵¹ suggesting that advanced scaling methods may be necessary to improve model consistency. As the quadriceps muscle moment arms have been shown to be well correlated with femoral condyle width,³⁹ consistent scaling between models along that dimension may reduce model differences. The AnyBody Modeling System has body-scaling functions that incorporate body mass and percent fat, which are used to influence the mediolateral and anteroposterior skeletal dimensions. As those same functions were not available in the other modeling programs, they were not investigated in this study. Previous studies have investigated the use of patient-

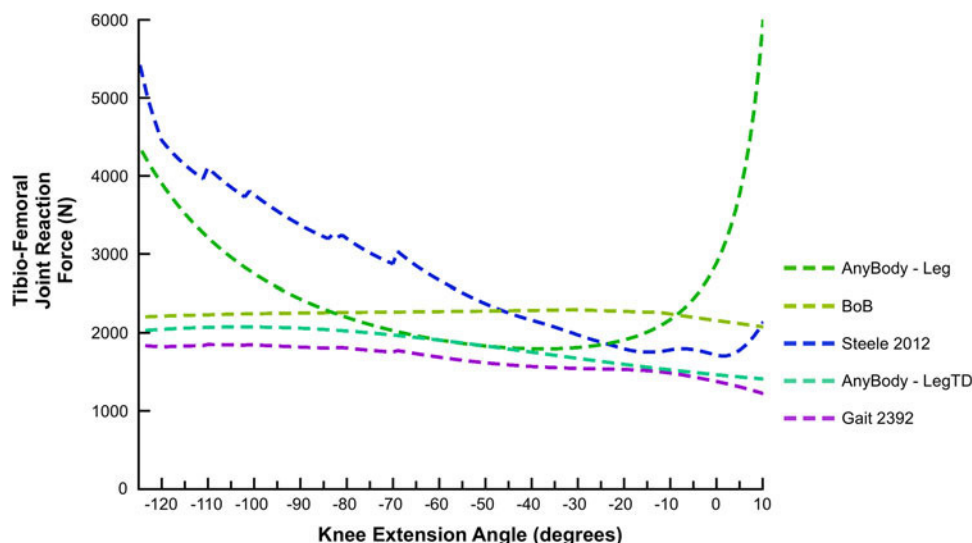


FIGURE 5. Tibiofemoral joint reaction force from quadriceps muscles resisting a 90 N-m flexion torque.

537 specific anatomy (derived from MR or CT imaging) to
 538 dimension and develop musculoskeletal models.^{6,12,53,59}
 539 However, custom scaling and definition of the muscle
 540 path based on imaging was not performed here, as the
 541 intent of this study was to compare differences in
 542 available generic models that could be used without the
 543 need for data from a specific subject.

544 The vastus lateralis was the largest contributor to
 545 the overall quadriceps muscle force in the majority of
 546 models and postures evaluated. This result was
 547 expected considering that the vastus lateralis had the
 548 largest maximum isometric strength at optimal fiber
 549 length in all the models, a result consistent with pre-
 550 vious studies which have shown the vastus lateralis to
 551 have the largest physiologic cross-sectional area of the
 552 four quadriceps muscles.⁶⁶ One notable exception was
 553 observed for the muscle recruitment results associated
 554 with the London Lower Limb model, which recruited
 555 the majority of the quadriceps force from the rectus
 556 femoris despite the vastus lateralis having a larger
 557 effective moment arm, maximum isometric strength,
 558 and maximum torque producing capability at 20 and
 559 100° knee flexion. Upon further investigation, the rel-
 560 atively large rectus femoris force can be primarily
 561 explained by the large force contribution from the
 562 passive element of those muscle fascicles. At low
 563 muscle activation, comparatively high forces can be
 564 transmitted to the muscle tendon. The large passive
 565 force contribution suggests the London Lower Limb
 566 model may not have appropriately scaled muscle fiber
 567 lengths for the muscles investigated here. Upon further
 568 inspection, the normalized fiber length for one of the
 569 rectus femoris muscle fascicles ranged from 3.3 to 4.5
 570 for the corresponding knee flexion angles of 0 and
 571 125°, respectively, suggesting a potential modeling

error with the defined optimal fiber length. Similar
 normalized fiber length values were also observed for
 the un-scaled model.

In a study analyzing the sensitivity of individual
 muscle parameters on computed muscle force from a
 static optimization procedure, Raikova and Prilutsky⁴⁸
 concluded that the non-zero optimal force of each
 muscle was non-linearly related to the moments at all the
 joints, the muscle moment arms, and the physiological
 cross sectional areas of all the muscles, which were used
 to normalize the predicted forces to compute muscle
 activity in the static optimization objective function.
 The differences between the models analyzed here sup-
 port those conclusions and further identify that the
 parameters of the muscle model (although not directly
 analyzed here), particularly those that define the force-
 length curve and the contribution between the passive
 and active elements, also substantially influence the
 subsequently recruited muscle force. Further research,
 methods, and protocols for reliably producing consis-
 tent muscle forces between musculoskeletal models
 under the same boundary conditions are necessary.

The London Lower Limb and the Lower Limb 2010
 models, which did have muscle forces computed to resist
 the simulated 90 N-m flexion torque, could not be used
 to calculate accurate tibiofemoral reaction forces using
 the Joint Reaction analysis tool in OpenSim.⁵⁸ Both
 models utilized a kinematic constraint to define the
 position of the patella as a function of knee angle. This
 constraint acts in place of the patellar tendon force such
 that the force exerted by the quadriceps muscles acting
 through the patella and patella tendon are not trans-
 mitted to the proximal tibia. For both models, the tibi-
 ofemoral reaction force computed using the Joint
 Reaction analysis tool was zero, as the weight of the

607 lower limb was neglected (gravity was set to zero) and a
 608 pure torque was applied about the knee joint axis of
 609 rotation. It should be highlighted that if a force vector
 610 were applied distally to the knee (e.g., a ground reaction
 611 force) for either of these models, the Joint Reaction
 612 analysis tool would report a tibiofemoral joint reaction
 613 force that would be consistent with the inter-segmental
 614 forces of the linkage. If the user were unaware of the
 615 modeling implications of the kinematic constraint and
 616 the assumptions associated with the Joint Reaction
 617 analysis tool, reported reaction forces may be misin-
 618 terpreted as true joint reaction forces.

619 The results suggest that although musculoskeletal
 620 models can fairly easily be scaled to have the same limb
 621 lengths and use the same muscle recruitment algo-
 622 rithm, those are not sufficient conditions to produce
 623 consistent muscle or joint contact forces (globally or by
 624 trends), even for simplified models with idealized
 625 boundary conditions and with no potential of
 626 co-contraction. However, between -10 and -50° knee
 627 extension, joint contact forces from all models were
 628 fairly consistent and ranged between 2.0 and 3.3 body
 629 weights (BW). Two models exhibited increased joint
 630 reaction forces as knee flexion angle increased, a result
 631 consistent with the observation that “tibial forces
 632 peaked at increasing knee flexion angle” from three
 633 subjects with instrumented endoprosthesis during a
 634 knee extension task.¹⁸ The remaining three musculo-
 635 skeletal models did not exhibit that same trend, but did
 636 have joint contact forces that converged between 2.5
 637 and 3.0 BW at 125° knee flexion. Trepczynski *et al.*⁵⁹
 638 identified considerable subject-specific variation in
 639 peak tibiofemoral joint loads during a variety of
 640 activities (e.g., walking and stair climbing), particularly
 641 those involving large knee flexion like squatting, a
 642 result consistent with the increased variation observed
 643 here between models as knee flexion increased.

647 scaling, model parameters, and underlying model con-
 648 structs must be matched to produce consistent results
 649 between musculoskeletal models? Is this possible? At
 650 what level should a musculoskeletal modeling user be
 651 expected to adapt a generic model to achieve “average”
 652 population results? These questions can be complicated
 653 to address considering the difficulties in identifying
 654 appropriate methods for model validation (e.g., what is
 655 the expected average behavior the models should be
 656 matching?). Additionally, differences that do exist
 657 between models can be difficult to interpret, as differ-
 658 ences resulting from natural inter-individual variation
 659 remain unknown. Although this study focused on the
 660 differences between generic musculoskeletal models and
 661 did not investigate models scaled to match patient-
 662 specific data, the answer to many of the questions above
 663 may rely on additional patient-specific data being made
 664 available to the musculoskeletal simulation commu-
 665 nity.⁶² For example, average and inter-subject variation
 666 may have to be defined based on analysis from
 667 patient-specific models (e.g., Scheys *et al.*⁵²) with the
 668 accuracy of those models being further evaluated
 669 using additional data available from instrumented en-
 670 doprotheses.^{27,40} For musculoskeletal simulation to be
 671 widely adopted and incorporated as an engineering
 672 discipline, verification and validation methods that are
 673 common to other computer aided engineering modali-
 674 ties must be more widely incorporated.⁴² Consistent
 675 results between generic musculoskeletal models is one
 676 step toward accomplishing that goal such that a bio-
 677 mechanical analysis performed by one investigator at
 678 one location with one piece of software produces
 679 the same reliable and repeatable results as the same
 680 analysis performed by another individual, at another
 681 location, with another musculoskeletal simulation
 682 software package.

644 FUTURE RESEARCH

645 The results presented here raise several questions and
 646 potential topics for future research including: What

APPENDIX

See Table 7.

TABLE 7. Musculoskeletal software and model download locations.

| Model name | Model accessibility | Software | Software accessibility |
|------------------------|---|-------------------------|---|
| AnyBody—Leg | http://forge.anyscript.org/gf/project/ammr/ | AnyBody Modeling System | www.anybodytech.com |
| AnyBody—LegTD | http://forge.anyscript.org/gf/project/ammr/ | | |
| Biomechanics of Bodies | http://www.marlbroom.com/download | Matlab | www.matlab.com |
| Delp 1990 | https://simtk.org/home/low-ext-model | Opensim | https://simtk.org/home/opensim |
| Steele 2012 | https://simtk.org/home/mattdemersstuff | | |
| Gait 2392 | https://simtk.org/home/torso_legs | | |
| London Lower Limb | https://simtk.org/home/low_limb_london | | |
| Lower Limb 2010 | https://simtk.org/home/lowlimbmodel09 | | |

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689 the Paralyzed Veterans of America Endowment for
690 Spinal Cord Injury at Stanford University.

691

CONFLICT OF INTEREST

692 Matthew S. DeMers collaborated on the develop-
693 ment of the Steele 2012 model evaluated in this man-
694 uscript. James M. Shippen is the primary developer of
695 the Biomechanics of Bodies software used in this
696 manuscript.

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REFERENCES

698 ¹Ahmed, S., and K. Babski-Reeves. Assessment of upper
699 extremity postures in novice and expert during simulated
700 carpentry tasks. *Proc. Hum. Factors Ergonomics Soc.*
701 *Annu. Meet.* 56:1173–1177, 2012.

702 ²Alkjaer, T., M. R. Wieland, M. S. Andersen, E. B.
703 Simonsen, and J. Rasmussen. Computational modeling of
704 a forward lunge: towards a better understanding of the
705 function of the cruciate ligaments. *J. Anat.* 221:590–597,
706 2012.

707 ³An, K. N., K. Takahashi, T. P. Harrigan, and E. Y. Chao.
708 Determination of muscle orientations and moment arms. *J.*
709 *Biomech. Eng.* 106:280–282, 1984.

710 ⁴Andersen, M. S., M. de Zee, S. Dendorfer, B.
711 MacWilliams, and J. Rasmussen. Validation of a detailed
712 lower extremity model based on the klein horsman data set.
713 In: *The 12th International Symposium On Computer*
714 *Simulation In Biomechanics*, edited by Proceedings Of.
715 Cape Town: South Africa, 2009, pp. 27–28.

716 ⁵Andrews, B. J., J. Shippen, R. S. Gibbons, B. May, and G.
717 Wheeler. FES rowing biomechanics: fixed and floating
718 stretcher ergometers. In: *17th Annual International FES*
719 *Society Conference: Smart Machines—Neural Evolution*,
720 Banff, Alberta, Canada, 2012.

721 ⁶Arnold, A. S., S. Salinas, D. J. Asakawa, and S. L. Delp.
722 Accuracy of muscle moment arms estimated from MRI-
723 based musculoskeletal models of the lower extremity.
724 *Comput. Aided Surg.* 5:108–119, 2000.

725 ⁷Arnold, E. M., S. R. Ward, R. L. Lieber, and S. L. Delp. A
726 model of the lower limb for analysis of human movement.
727 *Ann. Biomed. Eng.* 38:269–279, 2010.

728 ⁸Azmy, C., S. Guérard, X. Bonnet, F. Gabrielli, and W.
729 Skalli. EOS orthopaedic imaging system to study patel-
730 lofemoral kinematics: assessment of uncertainty. *Orthop.*
731 *Traumatol. Surg. Res.* 96:28–36, 2010.

732 ⁹Blemker, S. S., D. S. Asakawa, G. E. Gold, and S. L. Delp.
733 Image-based musculoskeletal modeling: applications,
734 advances, and future opportunities. *J. Magn. Reson.*
735 *Imaging* 25:441–451, 2007.

736 ¹⁰Buford, W. L., Jr., F. M. Ivey, Jr., J. D. Malone, R. M.
737 Patterson, G. L. Peare, D. K. Nguyen, and A. A. Stewart.
738 Muscle balance at the knee-moment arms for the normal
739 knee and the ACL-minus knee. *IEEE Trans. Rehabil. Eng.*
740 5:367–379, 1997.

741 ¹¹Chang, C.-Y., J. D. Rupp, M. P. Reed, R. E. Hughes, and
742 L. W. Schneider. Predicting the effects of muscle activation
743 on knee, thigh, and hip injuries in frontal crashes using a
744 finite-element model with muscle forces from subject testing
745 and musculoskeletal modeling. *Stapp Car Crash J.* 53:291–
746 328, 2009.

747 ¹²Correa, T. A., R. Baker, H. K. Graham, and M. G. Pandy.
748 Accuracy of generic musculoskeletal models in predicting
749 the functional roles of muscles in human gait. *J. Biomech.*
750 44:2096–2105, 2011.

751 ¹³Damsgaard, M., J. Rasmussen, S. T. Christensen, E.
752 Surma, and M. de Zee. Analysis of musculoskeletal systems
753 in the AnyBody Modeling System. *Simul. Model. Pract.*
754 *Theory* 14:1100–1111, 2006.

755 ¹⁴de Zee, M., M. Lund, C. Schwartz, C. Olesen, and J.
756 Rasmussen. Validation of musculoskeletal models: the
757 importance of trend validations. Leuven, Belgium: IUTAM
758 Symposium on Human Movement Analysis and Simula-
759 tion, 2010.

760 ¹⁵Delp, S. L., F. C. Anderson, A. S. Arnold, P. Loan, A.
761 Habib, C. T. John, E. Guendelman, and D. G. Thelen.
762 OpenSim: open-source software to create and analyze
763 dynamic simulations of movement. *IEEE Trans. Biomed.*
764 *Eng.* 54:1940–1950, 2007.

765 ¹⁶Delp, S. L., and J. P. Loan. A graphics-based software
766 system to develop and analyze models of musculoskeletal
767 structures. *Comput. Biol. Med.* 25:21–34, 1995.

768 ¹⁷Delp, S. L., J. P. Loan, M. G. Hoy, F. E. Zajac, E. L.
769 Topp, and J. M. Rosen. An interactive graphics-based
770 model of the lower extremity to study orthopaedic surgical
771 procedures. *IEEE Trans. Biomed. Eng.* 37:757–767, 1990.

772 ¹⁸D’Lima, D. D., N. Steklov, S. Patil, and C. W. Colwell, Jr.
773 The Mark Coventry Award: in vivo knee forces during
774 recreation and exercise after knee arthroplasty. *Clin. Ort-*
775 *hop. Relat. Res.* 466:2605–2611, 2008.

776 ¹⁹Draganich, L. F., T. P. Andriacchi, and G. B. Andersson.
777 Interaction between intrinsic knee mechanics and the knee
778 extensor mechanism. *J. Orthop. Res.* 5:539–547, 1987.

779 ²⁰Dubowsky, S. R., J. Rasmussen, S. A. Sisto, and N. A.
780 Langrana. Validation of a musculoskeletal model of
781 wheelchair propulsion and its application to minimizing
782 shoulder joint forces. *J. Biomech.* 41:2981–2988, 2008.

783 ²¹Dudley-Javoroski, S., A. E. Littmann, S.-H. Chang, C. L.
784 McHenry, and R. K. Shields. Enhancing muscle force and
785 femur compressive loads via feedback-controlled stimula-
786 tion of paralyzed quadriceps in humans. *Arch. Phys. Med.*
787 *Rehabil.* 92:242–249, 2011.

788 ²²Duffy, V. G. *Handbook Of Digital Human Modeling:*
789 *Research For Applied Ergonomics And Human Factors*
790 *Engineering* 1st ed. Boca Raton: CRC Press, Inc., 2008.

791 ²³Escamilla, R. F., G. S. Fleisig, T. M. Lowry, S. W.
792 Barrentine, and J. R. Andrews. A three-dimensional bio-
793 mechanical analysis of the squat during varying stance
794 widths. *Med. Sci. Sports Exerc.* 33:984–998, 2001.

795 ²⁴Escamilla, R. F., G. S. Fleisig, N. Zheng, J. E. Lander,
796 S. W. Barrentine, J. R. Andrews, B. W. Bergemann, and
797 C. T. Moorman, III. Effects of technique variations on
798 knee biomechanics during the squat and leg press. *Med.*
799 *Sci. Sports Exerc.* 33:1552–1566, 2001.

800 ²⁵Farrell, K. C., K. D. Reisinger, and M. D. Tillman. Force
801 and repetition in cycling: possible implications for iliotibial
802 band friction syndrome. *Knee* 10:103–109, 2003.

803 ²⁶Fotoohabadi, M. R., E. A. Tully, and M. P. Galea. Kine-
804 matics of rising from a chair: image-based analysis of the

- 805 sagittal hip-spine movement pattern in elderly people who
806 are healthy. *Phys. Ther.* 90:561–571, 2010.
- 807 ²⁷Fregly, B. J., T. F. Besier, D. G. Lloyd, S. L. Delp, S. A.
808 Banks, M. G. Pandy, and D. D. D’Lima. Grand challenge
809 competition to predict in vivo knee loads. *J. Orthop. Res.*
810 30:503–513, 2012.
- 811 ²⁸Frey Law, L. A., and R. K. Shields. Femoral loads during
812 passive, active, and active-resistive stance after spinal cord
813 injury: a mathematical model. *Clin. Biomech. (Bristol, Avon)*
814 19:313–321, 2004.
- 815 ²⁹Good, E. S., W. J. Suntay, F. R. Noyes, and D. L. Butler.
816 Biomechanics of the knee-extension exercise. Effect of
817 cutting the anterior cruciate ligament. *J. Bone Joint Surg. Am.*
818 66:725–734, 1984.
- 819 ³⁰Grujicic, M., G. Arakere, X. Xie, M. LaBerge, A. Grujicic,
820 D. W. Wagner, and A. Vallejo. Design-optimization and
821 material selection for a femoral-fracture fixation-plate
822 implant. *Mater. Des.* 31:3463–3473, 2010.
- 823 ³¹Grujicic, M., B. Pandurangan, X. Xie, A. K. Gramopad-
824 hye, D. W. Wagner, and M. Ozen. Musculoskeletal com-
825 putational analysis of the influence of car-seat design/
826 adjustments on long-distance driving fatigue. *Int. J. Ind. Ergon.*
827 40:345–355, 2010.
- 828 ³²Grujicic, M., X. Xie, G. Arakere, A. Grujicic, D. W.
829 Wagner, and A. Vallejo. Design-optimization and material
830 selection for a proximal radius fracture-fixation implant. *J. Mater. Eng. Perform.* 19:1090–1103, 2010.
- 831 ³³Hamner, S. R., A. Seth, and S. L. Delp. Muscle contribu-
832 tions to propulsion and support during running. *J. Biomech.*
833 43:2709–2716, 2010.
- 834 ³⁴Hettinga, D. M., and B. J. Andrews. The feasibility of
835 functional electrical stimulation indoor rowing for high-
836 energy training and sport. *Neuromodulation* 10:291–297,
837 2007.
- 838 ³⁵Iwami, T., K. Miyawaki, K. Hiramoto, M. Takeshima, T.
839 Matsunaga, Y. Shimada, and G. Obinata. Biomechanical
840 analysis and muscle tension estimation of the lower
841 extremities using EMG data. International Symposium on
842 Micro-NanoMechatronics and Human Science (MHS),
843 2010, 2010, pp 175–180.
- 844 ³⁶Kiratli, B. J. Immobilization osteopenia. In: Osteoporosis,
845 2nd edition. San Diego: Academic Press, 2001, pp. 207–
846 227.
- 847 ³⁷Klein Horsman, M. D., H. F. J. M. Koopman, F. C. T. van
848 der Helm, L. P. Prosé, and H. E. J. Veeger. Morphological
849 muscle and joint parameters for musculoskeletal modelling
850 of the lower extremity. *Clin. Biomech. (Bristol, Avon)*
851 22:239–247, 2007.
- 852 ³⁸Klein Horsman, M. D. The Twente Lower Extremity
853 Model: Consistent Dynamic Simulation of the Human
854 Locomotor Apparatus [dissertation]. Department of Engi-
855 neering Technology. Enschede, The Netherlands: Univer-
856 sity of Twente, 2007.
- 857 ³⁹Krevolin, J. L., M. G. Pandy, and J. C. Pearce. Moment
858 arm of the patellar tendon in the human knee. *J. Biomech.*
859 37:785–788, 2004.
- 860 ⁴⁰Kutzner, I., B. Heinlein, F. Graichen, A. Bender, A.
861 Rohlmann, A. Halder, A. Beier, and G. Bergmann.
862 Loading of the knee joint during activities of daily living
863 measured in vivo in five subjects. *J. Biomech.* 43:2164–2173,
864 2010.
- 865 ⁴¹Leszko, F., K. R. Hovinga, A. L. Lerner, R. D. Komistek,
866 and M. R. Mahfouz. In vivo normal knee kinematics: is
867 ethnicity or gender an influencing factor? *Clin. Orthop. Relat. Res.*
868 469:95–106, 2011.
- ⁴²Lund, M. E., M. de Zee, M. S. Andersen, and J. 870
871 Rasmussen. On validation of multibody musculoskeletal
872 models. *Proc. Inst. Mech. Eng. H.* 226:82–94, 2012.
- ⁴³McFadyen, B. J., and D. A. Winter. An integrated bio-
873 mechanical analysis of normal stair ascent and descent. *J. Biomech.*
874 21:733–744, 1988.
- ⁴⁴McHenry, C. L., and R. K. Shields. A biomechanical
875 analysis of exercise in standing, supine, and seated posi-
876 tions: Implications for individuals with spinal cord injury.
877 *J. Spinal Cord Med.* 35:140–147, 2012.
- ⁴⁵Modenese, L., A. T. M. Phillips, and A. M. J. Bull. An
878 open source lower limb model: hip joint validation. *J. Biomech.*
879 44:2185–2193, 2011.
- ⁴⁶Perry, J. *Gait Analysis : Normal And Pathological Func- tion.*
880 Thorofare, N.J.: SLACK inc., 1992.
- ⁴⁷Pontonnier, C., M. de Zee, A. Samani, G. Dumont, and P.
881 Madeleine. Trend Validation of a Musculoskeletal Model
882 with a Workstation Design Parameter. ISB Technical
883 Group on Computer Simulation Symposium 2011, Leuven,
884 Belgium, 2011.
- ⁴⁸Raikova, R. T., and B. I. Prilutsky. Sensitivity of predicted
885 muscle forces to parameters of the optimization-based
886 human leg model revealed by analytical and numerical
887 analyses. *J. Biomech.* 34:1243–1255, 2001.
- ⁴⁹Rasmussen, J., M. Boockock, and G. Paul. Advanced
888 musculoskeletal simulation as an ergonomic design meth-
889 od. *Work: J. Prev. Assess. Rehabil.* 41:6107–6111, 2012.
- ⁵⁰Rasmussen, J., and M. de Zee. Design optimization of
890 airline seats. In: Sae Transactions: Journal of Passenger
891 Cars—Electronic and Electrical Systems, 2008.
- ⁵¹Roebuck, J. A. *Anthropometric Methods : Designing To Fit The Human Body.* Santa Monica, CA, USA: Human
892 Factors and Ergonomics Society, 1995.
- ⁵²Scheys, L., K. Desloovere, P. Suetens, and I. Jonkers. Level
893 of subject-specific detail in musculoskeletal models affects
894 hip moment arm length calculation during gait in pediatric
895 subjects with increased femoral anteversion. *J. Biomech.*
896 44:1346–1353, 2011.
- ⁵³Scheys, L., A. Spaepen, P. Suetens, and I. Jonkers. Cal-
897 culated moment-arm and muscle-tendon lengths during
898 gait differ substantially using MR based versus rescaled
899 generic lower-limb musculoskeletal models. *Gait Posture*
900 28:640–648, 2008.
- ⁵⁴Scheys, L., A. Van Campenhout, A. Spaepen, P. Suetens,
901 and I. Jonkers. Personalized MR-based musculoskeletal
902 models compared to rescaled generic models in the presence
903 of increased femoral anteversion: effect on hip moment arm
904 lengths. *Gait Posture* 28:358–365, 2008.
- ⁵⁵Seth, A., M. Sherman, J. A. Reinbolt, and S. L. Delp.
905 OpenSim: a musculoskeletal modeling and simulation
906 framework for in silico investigations and exchange. *Pro- cedia IUTAM* 2:212–232, 2011.
- ⁵⁶Sherman, M., A. Seth, and S. Delp. How to compute muscle
907 moment arm using generalized coordinates, Rev. 0.2.
908 <http://simtk-confluence.stanford.edu:8080/x/yoQz>. 2010.
- ⁵⁷Shippen, J. M., and B. May. Calculation of muscle loading
909 and joint contact forces during the rock step in Irish dance.
910 *J. Dance Med. Sci.* 14:11–18, 2010.
- ⁵⁸Steele, K. M., M. S. Demers, M. H. Schwartz, and S. L.
911 Delp. Compressive tibiofemoral force during crouch gait.
912 *Gait Posture* 35:556–560, 2012.
- ⁵⁹Trepczynski, A., I. Kutzner, E. Kornaropoulos, W. R.
913 Taylor, G. N. Duda, G. Bergmann, and M. O. Heller.
914 Patellofemoral joint contact forces during activities with
915 high knee flexion. *J. Orthop. Res.* 30:408–415, 2012.

- 935 ⁶⁰Vandenberghe, A., L. Bosmans, J. De Schutter, S. Swinnen, and I. Jonkers. Quantifying individual muscle contribution to three-dimensional reaching tasks. *Gait Posture* 35:579–584, 2012. 956
- 936 937 938 939 ⁶¹Viceconti, M. A tentative taxonomy for predictive models in relation to their falsifiability. *Philos. Trans. A Math. Phys. Eng. Sci.* 369:4149–4161, 2011. 957
- 940 941 942 ⁶²Viceconti, M., D. Testi, F. Taddei, S. Martelli, G. J. Clapworthy, and S. V. S. Jan. Biomechanics Modeling of the Musculoskeletal Apparatus: Status and Key Issues. *Proc. IEEE* 94:725–739, 2006. 958
- 943 944 945 ⁶³Wagner, D. W., J. Rasmussen, and M. P. Reed. Assessing the importance of motion dynamics for ergonomic analysis of manual materials handling tasks using the AnyBody Modeling System. In: *Sae Transactions: Journal Of Passenger Cars—Mechanical Systems*, 2007, pp. 2092–2101. 959
- 946 947 948 949 950 ⁶⁴Wagner, D. W., K. Divringi, C. Ozcan, M. Grujicic, B. Pandurangan, and A. Grujicic. Combined musculoskeletal dynamics/structural finite element analysis of femur physiological loads during walking. *Multidiscip. Model. Mater. Struct.* 6:417–437, 2010. 960
- 951 952 953 954 955 ⁶⁵Walker, P. S., J. S. Rovick, and D. D. Robertson. The effects of knee brace hinge design and placement on joint mechanics. *J. Biomech.* 21:965–974, 1988. 956
- 957 958 959 ⁶⁶Ward, S. R., C. M. Eng, L. H. Smallwood, and R. L. Lieber. Are current measurements of lower extremity muscle architecture accurate? *Clin. Orthop. Relat. Res.* 467:1074–1082, 2009. 960
- 961 962 963 ⁶⁷Wilk, K. E., R. F. Escamilla, G. S. Fleisig, S. W. Barrentine, J. R. Andrews, and M. L. Boyd. A comparison of tibiofemoral joint forces and electromyographic activity during open and closed kinetic chain exercises. *Am. J. Sports Med.* 24:518–527, 1996. 961
- 964 965 966 967 968 ⁶⁸Wilson, N. A., and F. T. Sheehan. Dynamic in vivo 3-dimensional moment arms of the individual quadriceps components. *J. Biomech.* 42:1891–1897, 2009. 962
- 969 970 971 ⁶⁹Yamaguchi, G. T., and F. E. Zajac. A planar model of the knee joint to characterize the knee extensor mechanism. *J. Biomech.* 22:1–10, 1989. 963
- 972 973 974 975 976 977 ⁷⁰Zajac, F. E. Muscle and tendon: properties, models, scaling, and application to biomechanics and motor control. *Crit. Rev. Biomed. Eng.* 17:359–411, 1989. 964

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