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In vitro compressive properties of skeletal muscles and inverse finite element analysis: comparison of human versus animals

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Abstract: Virtual finite element human body models have been widely used in 1 biomedical engineering, traffic safety injury analysis, etc. Soft tissue modeling like 2 3 skeletal muscle accounts for a large portion of a human body model establishment, and its modeling method is not enough explored. The present study aims to investigate the 4 compressive properties of skeletal muscles due to different species, loading rates and 5 fiber orientations, in order to obtain available parameters of specific material laws as 6 references for building or improving the human body model concerning both modeling 7 accuracy and computational cost. A series of compressive experiments of skeletal 8 muscles were implemented for human gastrocnemius muscle, bovine and porcine hind 9 leg muscle. To avoid long-time preservation effects, all experimental tests were carried 10

out in 24 hours after that the samples were harvested. Considering computational cost 11 and generally used in the previous human body models, one-order hyperelastic Ogden 12 model and three-term simplified viscoelastic quasi-linear viscoelastic (QLV) were 13 selected for numerical analysis. Inverse finite element analysis was employed to obtain 14 corresponding material parameters. With good fitting records, the simulation results 15 presented available material parameters for human body model establishment, and also 16 indicated significant differences of muscle compressive properties due to species, 17 18 loading rates and fiber orientations. When considering one-order Ogden law, it is worthy of noting that the inversed material parameters of the porcine muscles are 19 similar to those of the human gastrocnemius regardless of fiber orientations. In 20 conclusion, the obtained material parameters in the present study can be references for 21 global human body and body segment modeling. 22

Keywords: Skeletal muscle, Human gastrocnemius, Inverse analysis, Finite element
 simulation, Parameter identification

25

26 **1. Introduction**

27 Finite element (FE) Human body models (HBMs) are widely used for biomechanical analysis in biomedical engineering, transport safety, sport science, and other human 28 involved activities (Florio, 2018; Halloran, et al., 2010; Hedenstierna, et al., 2008; 29 Lenhart, et al., 2015). Skeletal muscle accounts for about 40% of the body mass, 30 therefore an understanding of its mechanical properties is important for virtual human 31 32 body modeling and related biomechanical application (Chomentowski et al., 2011, Takaza, et al., 2013). Muscle tissues can be roughly characterized to be highly non-33 linear, viscoelastic, anisotropic and strain rate dependency like most biological tissues, 34 35 while their properties can also vary significantly due to diversity of species. Thus, the experimental test and numerical modeling of muscle tissues are challenging works. 36

37

38 Skeletal muscle is composed of about 70-80% water, 3% fat and 10% collagen (Vignos and Lefkowitz, 1959). As it is very soft, most of the previous studies often tested it in 39 compression setups. Partly due to experimental condition limitation, many previous 40 studies primarily investigated mechanical behaviors of skeletal muscles in compression 41 with different animal species, and obtained corresponding parameters with inverse 42 analysis. For example, several previous studies (McElhaney, 1966; Van Loocke et al., 43 44 2006, 2008; Sligtenhorst and Cronin, 2006; Song et al., 2007; Chawla et al., 2009) investigated high strain rate effects of compression loading on skeletal muscles till 45 46 3700/s, while other studies presented experimental results with relatively lower strain rates till 0.001/s (Sacks, 2000; Sligtenhorst et al., 2006; Song et al., 2007; Xu et al., 47 2011). With regard to fiber orientations, continuous researches were implemented with 48 different animal muscles primarily considering the fiber orientations of 0° , 45° and 90° 49 50 (Bosboom et al., 2001 a, b; Van Loocke et al., 2006; Böl et al., 2012; Mohammadkhah et al., 2016). Although the uniform macro feature of experimental curves can be noted 51 52 for different animal muscles, the mechanical properties in detail during compression 53 change a lot due to the species as well as experimental conditions. Thus, the limitations of these animal muscle experiments for FE human body modeling have always existed.

55

56 To our knowledge, previous researchers on human skeletal muscle properties under compression loading were not enough and often limited to cadaver specimens with 57 long-time preservation. Vannah et al. (1996) conducted quasi-static compression 58 experiments of human muscles to provide design data for lower limb prostheses and 59 other soft tissue supporting devices. Dhaliwal et al. (2002) conducted low energy 60 impact tests on human cadaver muscles and obtained corresponding dynamic 61 compressive mechanical properties. These previous studies primarily used the samples 62 63 harvested from cadavers without indicating details of preservation conditions, like 64 preservation time. However, the previous investigations have indicated that long term formalin preservation can significantly change mechanical properties of biological 65 tissues, even for bone tissues (Kikugawa et al., 2004; Ohman et al., 2008; Zhang et al., 66 2019). Hence, we believe that implementing the experiments with short time 67 preservation samples should be more available for soft tissues like skeletal muscle. 68

69

Hence, the objective of the present study was to investigate the compressive properties 70 of human skeletal muscles with short term preservation less than 24 hours and compare 71 them with those of other animals, in order to provide available modeling parameters for 72 FE human modeling or related biomechanical application. In addition, many virtual FE 73 74 human body or body segment models with skeletal muscle modeling were developed 75 in the previous studies for various considerations, where the Ogden or QLV laws were usually used for these complicated model establishment (Mo et al., 2019). Here, we 76 believe that in a FE human body modeling both accuracy and calculating efficiency 77 should be considered for engineering application. Thus, two generally used material 78 types for the global human body or segment models were selected to obtain specific 79 80 material parameters by inverse finite element analysis in order to use them in existing or future FE human body models. 81

82 **2. Materials and methods**

2.1 Sample preparation and experimental procedure

84 All experiments were approved by the ethics committee of the Second Xiangya 85 Hospital of Central South University (NO. 2012-S231) and the informed consent was obtained from the patients participating in the study. Cubic muscle samples with a 86 characteristic edge length of 10 mm were obtained from human gastrocnemius muscle 87 as well as bovine and porcine hind legs. Summary of sample sizes is listed in Table 1. 88 89 The bovine samples were harvested from the hindquarter of a large Yorkashire, and the porcine samples were harvested from the hindquarter of a Chinese yellow cattle. The 90 human gastrocnemius muscle was obtained from two amputee donators (age 69 ± 3 years 91 old) without any leg muscle disorders or diseases. The tissues were fixed using formalin 92 93 to prevent tissue dissolution for no more than 24 hours before the experiments.

94

Before the experiment, the skin, fat and fascia were firstly removed from the muscle 95 96 tissues to obtain standard cubic testing specimens. All operations were carried out by a surgeon through a scalpel and an assistive block with ruler to control the size. 97 Subsequent compressive experiments were performed using different loading rates and 98 along different fiber orientations $(0^\circ, 45^\circ \text{ and } 90^\circ)$ as shown in Fig. 1. The MTS E45 99 100 electronic universal testing machine was used to compress the specimens up to 50% strain at rates of 0.05%/s, 0.5%/s and 5%/s. The measurement range of the selected 101 force sensor is 30kN with a resolution of 0.001N. As pre-compression may lead to 102 unpredictable permanent tissue deformation, each specimen was tested once and no 103 104 precondition was applied to the specimen. At least three successful specimen tests were achieved at each testing condition. A total of successful 99 compression experiments 105 106 were conducted at different loading rates and fiber directions.

107 Fig.1. Schematic of compression experiments with different loading rates and fiber

108

directions

Specimen	Fiber direction	Average height \pm std (mm)	Average width \pm std (mm)	Average thickness ±std (mm)	Sample size (N)
Bovine	0 °	10.5±0.6	10.7±0.5	10.5±0.6	12
	45°	10.8±1.2	11.1±1.2	10.6±0.6	12
	90°	10.9±0.9	10.8±0.7	10.8±0.7	12
Porcine	0 °	11.3±0.9	10.8±0.8	10.9±0.9	12
	45°	11.1±1.4	10.5±1.1	10.5±0.5	12
	90°	11.7±0.8	11.3±0.9	10.8±0.7	12
Human	0°	10.0±1.0	10.7±0.5	10.3±0.5	9
	45°	9.9±0.9	10.4±0.8	10.1±0.7	9
	90°	12.0±2.1	10.6±1.9	10.3±0.9	9

Table 1. Sample size and average specimen dimensions at different fiber orientations

110 **2.2 Mathematical modeling**

Following the previous studies (Comley et al. 2012, Takaza et al. 2013, Mo et al, 2018, 2019a, b), one-order Ogden hyperelastic material law and three-terms quasi-linear viscoelastic (QLV) material law were usually selected for modeling passive behavior of muscle tissues in human body models. The Ogden hyperelastic material law is expressed as follows,

116
$$W = \sum_{i=1}^{n} \frac{\mu_i}{\alpha_i} \left(\lambda_1^{\alpha_i} + \lambda_2^{\alpha_i} + \lambda_3^{\alpha_i} - 3 \right) + K(J - 1 - \ln J)$$
(1)

117 Where *W* is the strain energy of the model; μ is the initial shear modulus, α is the 118 deviatoric exponent, λ_i is the deviatoric principal stretches in three directions: x, y and 119 z; *J* represents the bulk modulus. The skeletal muscle was considered incompressible, 120 thus, *J* equaled to 1. Therefore, the simple one-order Ogden law is expressed as follows,

121
$$W = \frac{2\mu}{\alpha^2} (\lambda_1^{\alpha} + \lambda_2^{\alpha} + \lambda_3^{\alpha} - 3)$$
(2)

122 The viscoelastic QLV law is expressed as equation (3) and (4),

123
$$\sigma(t) = \int_0^t G(t-\tau) \frac{\partial \sigma^e(\lambda)}{\partial \lambda} \cdot \frac{\partial \lambda}{\partial \tau} d\tau$$
(3)

124 Where $\partial \sigma^e(\lambda)/\partial \lambda$ is the instantaneous elastic response, $\partial \lambda/\partial \tau$ is the stretch history; 125 $G(t - \tau)$ is the relaxation function.

Based on the QLV law, we used a simplified model focusing on instantaneous elastic
response in compression. And three terms of model coefficients were adopted. The
equation is expressed as follows,

129
$$\sigma_{\varepsilon}(\varepsilon) = C_1 \varepsilon^1 + C_2 \varepsilon^2 + C_3 \varepsilon^3 \tag{4}$$

130 Where C_1 , C_2 and C_3 are the coefficients of instantaneous elastic responses; ε^1 , ε^2 131 and ε^3 are the principal strains in three directions.

132 **2.3 Inverse analysis**

Inverse analysis has been applied for defining material parameters of soft tissues (Ahn, et al., 2010; Chawla, et al., 2009; Böl, et al., 2012; Nava, et al., 2008; Samur, et al., 2005, 2007). The purpose of the inverse analysis was to minimize the error between the experimental results and simulation responses by using an optimization algorithm to identify optimal material parameters (Böl, et al., 2012). The agreement between experimental results and simulation results were evaluated by the objective function as follows,

140
$$f(x) = \sqrt{\frac{\sum_{i}^{n} \left(F_{i}^{exp} - F_{i}^{FEM}\right)^{2}}{n}}$$
(5)

141 Where F_i^{exp} is the experimental force value, F_i^{FEM} is the responsive force of finite 142 element simulation, and *n* is the number of data points. The process of inverse analysis 143 for material parameter identification is described in the following section.

144 **2.4 Identification of material parameters**

To identify material parameters of the tested skeletal muscles, a method combing 145 inverse solve and finite element optimization of material parameters were employed, as 146 shown in Fig. 2. The errors between the simulation and experimental results are 147 calculated by equation (5). A custom python program using a least-squares algorithm 148 149 was firstly used to fit experimental results to acquire initial parameters of the corresponding laws. Then, their values were optimized and finally determined by 150 inversed finite element analysis. Adaptive response surface method (ARSM) was 151 adopted in the FE simulations to minimize the error and obtain optimal parameters until 152 153 calculation convergence referring to the previous study (Wang et al., 2007), which is a direct optimization method based on response surface. The basic idea of ARSM method 154 is to construct a response surface with fewer sample points first and determine the 155 optimization direction. During the optimization process, new design points are obtained 156 157 along the gradient direction of the response surface, and introduced into the design space so that the response surface model can be gradually updated to improve 158 optimization efficiency and fitting accuracy. By optimizing and updating the response 159 160 surface, the optimal solution can be continuously approached to the real optimal solution. 161

162

Fig.2. Schematic illustration of material parameter identification

The FE simulations were implemented with Ls-Dyna codes. The specimen was meshed 163 with hexahedral solid elements of the size 1 mm by Hypermesh software. Element 164 quality was checked to avoid distorted elements and to satisfy the minimum time step 165 criteria. The time step was set as 1.0e-5. Surface to surface contacts were defined 166 167 between the platens and the specimen, and a self-contact of the specimen was defined 168 to avoid solid collapse during large deformation. The lower platen was constrained with no freedom in all directions, while the upper platen can freely translate along the 169 compression direction. The friction coefficient between the platen and specimen was 170 171 set as 0.3 according to the previous study (Wu, et al., 2004).

172 **3. Results**

3.1 Experimental results

174 Characteristic compressive stress-strain responses are classified based on tissue fiber 175 orientations, compression strain rates and species (Fig. 3-Fig. 5). The bovine and porcine muscle properties are illustrated in Fig. 3 and Fig. 4, respectively. Compressive 176 responses of the human gastrocnemius muscles are shown in Fig. 5. As general soft 177 biological tissues, the experimental data of the muscle tissues scatter on a large scale in 178 179 the present study. When comparing compressive responses of different species as shown in Fig. 6, the substantial difference can be noted due to species, fiber orientations 180 and strain rates. The compressive properties of the porcine muscles seem to be closer 181 to the human gastrocnemius muscles than those of the bovine muscles. 182

183

In most testing conditions except for 90° bovine muscle, potential strain effects are 184 noted in the testing range from 0.05% to 5% for all species and fiber orientations, while 185 186 the Cauchy stress value inclines significantly with the increasing strain rate at 50% strain as shown in Fig. 6 (p<0.005). With the increase of the strain rate, the scattering 187 range of the compressive curves in most of testing conditions tends to enlarge. The 188 stiffest tissue response was obtained when the muscle fibers were oriented in 189 190 perpendicular to the loading direction (90° fiber orientation), followed by a fiber orientation of 45° . Fiber orientation of 0° in parallel with the loading direction yielded 191 the softest response. 192

193

(a) 0° (b) 45° (c) 90°

194	Fig.3. Compressive stress-stretch responses of the bovine muscles at various rates and fiber
195	orientations
196	(a) 0° (b) 45° (c) 90°

197 Fig.4. Compressive stress-stretch responses of the porcine muscles at various rates and fiber

198	orientations
199	(a) 0° (b) 45° (c) 90°
200	Fig.5. Compressive stress-stretch responses of the human gastrocnemius muscles at various
201	rates and fiber orientations
202	Fig.6. Comparison of compressive responses of different species under a strain level of 50%

3.2 Identification of material parameters

We used the averaged experimental curves of different fiber orientations and different species as the objective curves for the optimization of material parameters. Through the above-mentioned method, the typical fitting results of numerical simulations to experimental results considering the human skeletal muscles of three fiber orientations are shown in Fig. 7. All simulation results are in good agreement with the experimental responses.

210

Inversed material parameters of different muscle tissues using one-order Ogden law 211 212 were illustrated in Table 2. The corresponding material parameters of three-terms simplified QLV law were listed in Table 3. For both material laws, the obtained material 213 parameters for different species and fiber orientations present some differences. 214 Although significant differences (p < 0.05) of material parameters existed between the 215 porcine and human muscles considering different fiber orientations, the inversed 216 217 material parameters of the averaged curves of all three fiber orientations show similar values for both human (μ =3.43 kPa, α =8.74) and porcine (μ =3.63 kPa, α =8.74) muscles 218 219 concerning one-order Ogden law.

220

(a) 0° (b) 45° (c) 90°

Fig.7. Typical fitting results of the finite element simulations to the experimental results of the human skeletal muscles.

- 223
- 224

Species	Fiber orientation	Material parameters	R ² (FE)	RMSE
	0°	μ=0.26 kPa α=14.00	0.994	1.21
D '	45°	μ=0.65kPa α=13.83	0.998	1.67
Bovine	90°	μ=2.00 kPa α=10.11	0.997	8.91
	Average	μ=1.18 kPa α=11.97	0.998	2.12
	0°	μ=0.54 kPa α=14.00	0.983	11.07
Donoina	45°	μ=3.80 kPa α=9.00	0.969	115.50
Porcine	90°	μ=3.96 kPa α=8.92	0.994	23.08
	Average	μ=3.63 kPa α=8.74	0.983	40.59
	0°	μ=2.24 kPa α=8.97	0.989	15.72
Ilumon	45°	μ=3.77 kPa α=9.00	0.982	63.88
Human	90°	μ=5.07 kPa α=9.14	0.994	43.79
	Average	μ=3.43 kPa α=8.74	0.934	173.33

Table 2 Optimized muscle material parameters of one-order Ogden law

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227

Table 3 Optimized muscle material parameters of three-terms QLV law

Species	Fiber orientation	Material parameters	R^2 (FE)	RMSE
	0°	C1=88.61 C2=-772.05 C3=1716.00	0.937	18.74
Bovine	45°	C1=169.70 C2=-1500.86 C3=3607.17	0.957	45.73
	90°	C1= 132.00 C2= -1140.00 C3=3320.00	0.960	79.09
	Average	C1= 122.18 C2= -1065.46 C3=280800	0.978	26.77
Porcine	0°	C1=90.00 C2= -1088.22 C3=2849.00	0.977	22.26
	45°	C1=99.00 C2= -994.16 C3=3366.99	0.992	22.51
	90°	C1=189.00 C2= -1477.87 C3=4488.00	0.991	36.25
	Average	C1= 132.00 C2=-990.00 C3=2981.00	0.982	40.32
Human	0°	C1= 115.36 C2= -965.06 C3=2758.01	0.967	42.45

45°	C1= 165.00 C2= -1160.00 C3=3310.00	0.964	76.57
90°	C1=436.05 C2= -3124.03 C3=7616.21	0.941	367.21
average	C1=253.00 C2=-1810.00 C3=4590.00	0.945	152.51

228 Notes: RMSE represents Root Mean Squared Error.

229 4. Discussion

230 Although some previous studies have investigated skeletal muscle properties in different loading types using different animal specimens (Mohammadkhah et al., 2016, 231 Nie et al., 2011, Simms et al., 2012, Takaza et al., 2013a), the experimental results of 232 human skeletal muscles especially the data with short preservation time in formalin or 233 234 in status were rare. In the present study, we aim to provide material properties for human body modeling, especially in the application of comfort and impact analysis during 235 seating and car crash. In these two loading conditions, human body muscles generally 236 experience a compression loading. In addition, when we compared the published 237 238 muscle properties of different animals (Song et al., 2007, Dhaliwal et al., 200, Mohammadkhah et al., 2016, Van Sligtenhorst et al., 2006, Takaza et al., 2014), 239 substantial differences of mechanical behaviors in compression can be noted. Hence, 240 the first purpose of this study was to investigate compressive behaviors of human 241 242 skeletal muscle without long formalin storing time, then obtain their material parameters of generally used and computationally efficient material laws for the 243 improvement or establishment of human body models. By the way, the dominant fiber 244 directions can be different considering anatomic structures of different muscles. Muscle 245 246 compressive properties from three representative fiber directions were measured and analyzed. Compressive properties of muscle tissues in other fiber directions should be 247 located between them. In the establishment of a related human body model, we 248 recommended either establishing or employing an anisotropic constitutive material law 249 250 in the model, or optimizing the corresponding muscle parameters of the available 251 isotropic laws considering fiber distribution and boundary loading conditions.

252

The Cauchy stress-strain relationship presented in this study provided a reference for 253 investigating mechanical behaviors of human skeletal muscles. The results showed that 254 the Cauchy stress during compression increased with the increasing strain rate, which 255 was similar to the previous researches (Chawla, et al., 2009; Sligtenhorst, et al., 2006; 256 Song, et al., 2007; Van Loocke, et al., 2006; Van Loocke, et al., 2008). The Cauchy 257 stress of 50% compression strain at the compression loading rate of 5%/s was about 3 258 times larger than that at the compression rate of 0.05%/s, which was qualitatively 259 260 similar to the results at other strain rates (Van Loocke, et al., 2006; Van Loocke, et al., 2008). The cross-fiber orientation mechanical response was found to be stiffer than 261 other fiber orientations, which was similar to the findings of the previous researches 262 with different loading strain rates (Song, et al., 2007; Böl, et al., 2012). 263

264

As the inversed material parameters shown in Table 2, the optimized shear modulus μ 265 of 5.07 kPa for human gastrocnemius was close to in vivo experimental results using 266 supersonic shear wave elastography (SWE) of Maïsetti et al. (2012) study or the results 267 of our verifying measurements. This can indicate the reliability of our results, and that 268 the obtained tissue properties are close to in vivo muscle tissue properties. In addition, 269 the average inversed material parameters of one-order Ogden law regardless of fiber 270 271 orientations for the porcine and human muscles are similar. But those of the bovine muscle are largely different with them. To some extent, it could indicate that using 272 porcine muscle for experimental tests could provide more useful information for human 273 muscle tissue modeling if the human tissue experiment is limited. But it can be also 274 275 influenced by native species' difference. To extend experimental possibility, we believe that it is important to find proper alternative animal tissues with similar mechanical 276 properties to human tissues for experimental researches. 277

278

The present study selected one-order Ogden law for skeletal muscle modeling, as it isa simple and reasonable solution for soft tissue modeling. Especially concerning its

281 application in a global biomechanical model, the high computational efficiency was extremely appreciated in our previous foot model and lower limb modeling (Mo et al., 282 2018, 2019). We compared the computational cost of one-order Ogden law and three-283 terms simplified QLV law in the present compression modeling using Ls-Dyna solver. 284 With 8-core Xeon[®] CPU, the calculation time for a simple compression simulation is 285 23 min and 27 mins, respectively. The one-order Ogden law has some advantages in 286 saving computational cost as well as controlling the stability of a global human body 287 288 model for loading conditions without large concerns on viscoelasticity. Considering a global human body or large body segment modeling, we believe the balance of 289 modeling accuracy and computational cost should be achieved properly with regard to 290 291 research objectives.

292

The present study combined experimental tests and inverse analysis method to 293 investigate the effects of loading rate, fiber orientation and species differences on 294 295 compressive behaviors of skeletal muscles. A large number of experiments and numerical simulations were implemented, and available material parameters were 296 297 obtained for human body modeling reference. There are still several limitations. First, the limited sample size for each testing condition may partly limit part of the robustness 298 299 of our results. As limited pure muscle tissues can be obtained after removing other fiber tissues in one donator, we had to harvest samples from two amputees not as we can 300 obtain all muscle samples from one animal. But if we enlarge our sample size, the 301 effects of individuals' difference would be also extended. Second, we use an ideal and 302 303 standard geometry for inverse finite element analysis to obtain muscle material parameters. As indicated by the previous study (Böl et al, 2012), to further precisely 304 define the muscle parameters, an inverse finite element method with detailed geometry 305 306 definition could be more powerful. Additionally, we also believe that using short-time preservation tissues for experiments or adopting in-vivo measuring methods like the 307 SWE method for obtaining or evaluating material parameters can be further 308 309 investigated for virtual FE human model establishment.

310 **5.** Conclusion

Inspired by the objective of human body modeling, the present study investigated the 311 compressive properties of skeletal muscles with different species, loading rates and 312 fiber orientations. Two commonly used material laws of Ogden and QLV constitutive 313 models were selected and simplified for numerical analysis in the consideration of 314 computational efficiency and modeling accuracy. Optimized material parameters were 315 316 well obtained through inverse analysis based on custom-made codes and optimal finite element simulations, which can be available references for human body modeling. The 317 experimental results show substantial differences between species, loading rates and 318 fiber orientations. Referring to the material parameters of one-order Ogden law, similar 319 values between the porcine muscles and the human gastrocnemius were found 320 regardless of fiber orientations. The inversed shear modulus was further verified by in 321 vivo SWE measurement, thus it also indicated that the present parameters can reflect 322 323 the properties of in vivo human gastrocnemius muscle. Coupling experimental and simulation investigations on different human muscles could be extended in the future 324 to further evaluate the present method and findings. 325

326 **Conflict of interest statement**

327 The authors assert that there are no conflicts of interest of any type.

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Reference

- Ahn, B., and J. Kim. 2010. Measurement and characterization of soft tissue behavior with surface deformation and force response under large deformations [J]. Med. Image. Anal. 14(2):138-148, 2010.
- Böl, M., R. Kruse, A. E. Ehret, K. 2012. Leichsenring, and T. Siebert. Compressive properties of passive skeletal muscle-The impact of precise sample geometry on parameter identification in inverse finite element analysis [J]. J. Biomech. 45(15):2673-2679.
- Bosboom, E. M. H., M. K. C. Hesselink, C. W. J. Oomens, C. V. C. Bouten, M. R. Drost, and F. P. T. Baaijens. 2001. Passive transverse mechanical properties of skeletal muscle under in vivo compression [J]. J. Biomech. 34(10):1365-1368.
- Bosboom, E. M. H., J. A. M. Thomassen, C. W. J. Oomens, C. V. C. Bouten, and F. P. T. Baaijens. 2001. A numerical-experimental approach to determine the transverse mechanical properties of skeletal muscle [C], J. Middleton, M.L. Jones, N.G. Shrive(Eds.), Computer Methods In Biomechanics And Biomedical Engineering, 3, Gordon and Breach Sciences Publishers, Ne York, pp. 187-192.
- Chawla, A., S. Mukherjee, and B. Karthikeyan. 2009. Characterization of human passive muscles for impact loads using genetic algorithm and inverse finite element methods [J]. Biomech. Model. Mechan. 8(1):67-76, 2009.
- Chomentowski, P., P. M. Coen, Z. Radikova, B. H. Goodpaster, and F. G. S. Toledo. 2011. Skeletal Muscle Mitochondria in Insulin Resistance: Differences in Intermyofibrillar Versus Subsarcolemmal Subpopulations and Relationship to Metabolic Flexibility [J]. J. Clin. Endocrinol. Metab. 96(2):494-503.
- Comley, K., and N. Fleck. 2012. The compressive response of porcine adipose tissue from low to high strain rate [J]. Int. J. Impact. Eng. 46:1-10.
- 8. Dhaliwal, T. S., P. Beillas, C. C. Chou, P. Prasad, K. H. Yang, and A. I. King. 2002.

Structural response of lower leg muscles in compression: a low impact energy study employing volunteers, cadavers and the hybrid III [J]. Stapp. Car . Crash. J. 46:229-243.

- Florio, C. S. 2018. Effectiveness of various isometric exercises at improving bone strength in cortical regions prone to distal tibial stress fractures [J]. Int. J. Numer Method. Biomed. Eng. 34(6): e2976.
- Halloran, J. P., M. Ackermann, A. Erdemir, and A. J. van den Bogert. 2010. Concurrent musculoskeletal dynamics and finite element analysis predicts altered gait patterns to reduce foot tissue loading [J]. J. Biomech. 43(14):2810-2815.
- Hedenstierna, S., P. Halldin, and K. Brolin. 2008. Evaluation of a combination of continuum and truss finite elements in a model of passive and active muscle tissue [J]. Comput. Methods. Biomech. Biomed. Eng. 11(6):627-639.
- Kikugawa, H., and T. Asaka. 2004. Effect of long-term formalin preservation on bending properties and fracture toughness of bovine compact bone [J]. Mater. Trans. 45(10):3060-3064.
- Lenhart, R. L., J. Kaiser, C. R. Smith, and D. G. Thelen. 2015. Prediction and Validation of Load-Dependent Behavior of the Tibiofemoral and Patellofemoral Joints During Movement [J]. Ann. Biomed. Eng. 43(11):2675-2685.
- Maisetti, O., F. Hug, K. Bouillard, and A. Nordez. 2012. Characterization of passive elastic properties of the human medial gastrocnemius muscle belly using supersonic shear imaging [J]. J. Biomech. 45(6):978-984.
- McElhaney, J. H. 1966. Dynamic response of bone and muscle tissue [J]. J. Appl. Physiol. 21(4):1231-1236.
- Mo, F., F. Li, M. Behr, Z. Xiao, G. Zhang, and X. Du. 2018. A Lower Limb-Pelvis Finite Element Model with 3D Active Muscles [J]. Ann. Biomed. Eng. 46(1):86-96.
- 17. Mo, F., J. Li, M. Dan, T. Liu, and M. Behr. 2019. Implementation of controlling

strategy in a biomechanical lower limb model with active muscles for coupling multibody dynamics and finite element analysis [J]. J. Biomech. 91:51-60.

- Mo, F., J. Li, Z. Yang, S. Zhou, and M. Behr. 2019. In Vivo Measurement of Plantar Tissue Characteristics and Its Indication for Foot Modeling [J]. Ann. Biomed. Eng. 47: 2356-2371.
- 19. Mohammadkhah, M., P. Murphy, and C. K. Simms. 2016. The in vitro passive elastic response of chicken pectoralis muscle to applied tensile and compressive deformation [J]. J. Mech. Behav. Biomed. Mater. 62:468-480.
- 20. Nava, A., E. Mazza, M. Furrer, P. Villiger, and W. H. Reinhart. 2018. In vivo mechanical characterization of human liver [J]. Med. Image. Anal. 12(2):203-216.
- Nie, X., J.-I. Cheng, W. W. Chen, and T. Weerasooriya. 2011. Dynamic Tensile Response of Porcine Muscle [J]. J. Appl. Mech. Mar 2011, 78(2): 021009-14.
- 22. Ohman, C., E. Dall'Ara, M. Baleani, S. V. S. Jan, and M. Viceconti. 2008. The effects of embalming using a 4% formalin solution on the compressive mechanical properties of human cortical bone [J]. Clin. Biomech. 23(10):1294-1298.
- Sacks, M. S. 2000. Biaxial mechanical evaluation of planar biological materials [J].
 J. Elast. 61(1-3):199-246.
- 24. Samur, E., M. Sedef, C. Basdogan, L. Avtan, and O. Duzgun. 2005. A robotic indenter for minimally invasive characterization of soft tissues [C], in CARS 2005: Computer Assisted Radiology and Surgery, edited by H. U. Lemke, K. Inamura, K. Doi, M. W. Vannier, and A. G. Farman, 713-718.
- 25. Samur, E., M. Sedef, C. Basdogan, L. Avtan, and O. Duzgun. 2007. A robotic indenter for minimally invasive measurement and characterization of soft tissue response [J]. Med. Image. Anal. 11(4):361-373.
- 26. Simms, K.C., Van Loocke, M., Lyons, C.G., 2012. Skeletal muscle in compression:modelling approaches for the passive muscle bulk [J]. Int. J.

Multiscale. Com. 10 (2), 143 - 154.

- Song, B., W. Chen, Y. Ge, and T. Weerasooriya. 2007. Dynamic and quasi-static compressive response of porcine muscle [J]. J. Biomech. 40(13):2999-3005.
- 28. Takaza, M., Cooney, G.M., Mcmanus, G., Stafford, P., Simms, C.K.,2014. Assessing the microstructural response to applied deformation in porcine passive skeletal muscle.[J]. J. Mech. Behav. Biomed. Mater. 40: 115-126.
- 29. Takaza, M., Moerman, K.M., Gindre, J., Lyons, G., Simms, C.K.,2013a. The anisotropic mechanical behaviour of passive skeletal muscle tissue subjected to large tensile strain.[J]. J. Mech. Behav. Biomed. Mater. 17: 209-220.
- Takaza, M., K. M. Moerman, and C. K. Simms. 2013. Passive skeletal muscle response to impact loading: Experimental testing and inverse modelling [J]. J. Mech. Behav. Biomed. Mater. 27:214-225.
- Van Loocke, M., C. G. Lyons, and C. K. Simms. 2006. A validated model of passive muscle in compression [J]. J. Biomech. 39(16):2999-3009.
- 32. Van Loocke, M., C. G. Lyons, and C. K. Simms. 2008. Viscoelastic properties of passive skeletal muscle in compression: Stress-relaxation behaviour and constitutive modelling [J]. J. Biomech. 41(7):1555-1566.
- 33. Van Sligtenhorst, C., D. S. Cronin, and G. W. Brodland. 2006. High strain rate compressive properties of bovine muscle tissue determined using a split Hopkinson bar apparatus [J]. J. Biomech. 39(10):1852-1858.
- 34. Vannah, W. M., and D. S. Childress. 1996. Indentor tests and finite element modeling of bulk muscular tissue in vivo [J]. J. Rehabil. Res. Dev. 33(3):239-252.
- 35. Vignos, P. J., Jr., and M. Lefkowitz. 1959. A biochemical study of certain skeletal muscle constituents in human progressive muscular dystrophy [J]. J. Clin. Invest. 38(6):873-881.
- 36. Wang, H., G. Li and Z. Zhong. 2008. Optimization of sheet metal forming processes

by adaptive response surface based on intelligent sampling method [J]. J. Mater. Process. Tech. 197(1-3), 77-88.

- 37. Wu, J. Z., R. G. Dong, and A. W. Schopper. 2004. Analysis of effects of friction on the deformation behavior of soft tissues in unconfined compression tests [J]. J. Biomech. 37(1):147-155.
- 38. Zhang, G., S. Wang, S. Xu, F. Guan, Z. Bai, and H. Mao. 2019. The Effect of Formalin Preservation Time and Temperature on the Material Properties of Bovine Femoral Cortical Bone Tissue [J]. Ann. Biomed. Eng. 47(4):937-952.

Figure legends

Figure 1. Schematic of compression experiments with different loading rates and fiber directions.

Figure 2. Schematic illustration of material parameter identification.

Figure 3. Compressive stress-stretch responses of the bovine muscles at various rates and fiber orientations. (a) 0° (b) 45° (c) 90°

Figure 4. Compressive stress-stretch responses of the porcine muscles at various rates and fiber orientations. (a) 0° (b) 45° (c) 90°

Figure 5. Compressive stress-stretch responses of the human gastrocnemius muscles at various rates and fiber orientations. (a) 0° (b) 45° (c) 90°

Figure 6. Comparison of compressive responses of different species under a strain level of 50%.

Figure 7. Typical fitting results of the finite element simulations to the experimental results of the human skeletal muscles. (a) 0° (b) 45° (c) 90°

































Conflict of Interest Statement

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Title: In vitro compressive properties of skeletal muscles and inverse finite element analysis: comparison of human versus animals

The authors assert that there are no conflicts of interest of any type. This assertion is also included in the manuscript. Fuhao MO: Conceptualization, Methodology, Writing- Reviewing and Editing, Supervision;
Zhefeng ZHENG: Data curation, Formal analysis, Writing- Original draft preparation;
Haotian ZHANG, Software, Validation;
Guibing Li: Investigation, Visulization;
ZuRong YANG: Validation;
Deyi Sun: Resources, Writing- Reviewing and Editing, Supervision.