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Cortical Bone Plate Properties Assessment using Inversion of Axially Transmitted Low Frequency Ultrasonic Guided Waves

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1 In the past few decades, early osteoporosis detection using ultrasonic bone qual-
2 ity evaluation has gained prominence. Specifically, various studies focused on axial
3 transmission using ultrasonic guided waves and have highlighted this technique's sen-
4 sitivity to intrinsic properties of long cortical bones. This work aims to demonstrate
5 the potential of low-frequency ultrasonic guided waves to infer the properties of the
6 bone inside which they are propagating. A proprietary ultrasonic transducer, tai-
7 lored to transmit ultrasonic guided waves under 500 kHz, was used for the data
8 collection. The gathered data underwent 2D Fast Fourier Transform processing to
9 extract experimental dispersion curves. The proposed inversion scheme compares
10 experimental dispersion curves with simulated dispersion curves calculated through
11 the semi-analytical iso-geometric analysis (SAIGA) method. The numerical model
12 integrates a bone phantom plate coupled with a soft tissue layer on its top surface,
13 mimicking the experimental bone phantom plates. Subsequently, the mechanical
14 properties of the bone phantom plates were estimated by reducing the misfit between
15 the experimental and simulated dispersion curves. This inversion leaned heavily on
16 the dispersive trajectories and amplitudes of ultrasonic guided wave modes. Re-
17 sults indicate a marginal discrepancy under 5% between the mechanical properties
18 ascertained using the SAIGA-based inversion and those measured using bulk wave
19 pulse-echo measurements.

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20 I. INTRODUCTION

21 Osteoporosis is a skeletal disorder characterized by compromised bone strength, leading
22 to an increased risk of fracture^{1,2}. The early diagnosis of osteoporosis is crucial as it allows
23 for timely intervention and prevention of fractures, and it has been extensively studied in the
24 past few decades. Bone density measurement is the gold standard to diagnose the disease.
25 However, the comprehensive evaluation of a person's bone health and fracture risk should
26 ideally consider other characteristics such as the bone stiffness, the cortical thickness and/or
27 the bone volume fraction³⁻⁵.

28 Osteoporosis is generally more prevalent in postmenopausal women, with nearly one in
29 two women affected (40%). It can also be found in 15% to 30% of the over 50 years old male
30 population for a combined total of approximately 200 million people affected worldwide⁶⁻⁹.
31 Comparatively, one in ten people will develop a cancer, across all ages, and the risk to de-
32 velop a cardiovascular disease is near 50%^{10,11}. Osteoporosis constitutes a major threat to
33 healthcare systems and leads to significant hospitalization costs worldwide. It is estimated
34 that there will be nearly 6.3 million hip fractures per year around the world by 2050, repre-
35 senting several billion dollars in healthcare costs⁴. Early detection of osteoporosis is crucial
36 to prevent fracture risk.

37 Recognized as the gold standard for diagnosing osteoporosis, dual-energy x-ray absorp-
38 tiometry (DEXA) uses two X-rays of different energy levels to estimate the bone density,
39 allowing to retrieve a 2D information on the bone mineral density (BMD)^{12,13}. To diagnose
40 osteoporosis in a patient, BMD is expressed in terms of standard deviation from a population

41 of healthy young adults called the T-score¹⁴. According to the World Health Organization,
42 a T-score of -2.5 or lower is considered as confirmed osteoporosis. The success of DEXA lies
43 in the fact that this technology and the interaction of X-rays with bones are well understood
44 and known¹⁵. Although this technology allows for obtaining BMD for different sites in the
45 human body, it remains difficult to retrieve the bone mechanical properties. The poor sen-
46 sitivity of BMD measurements makes the monitoring of the disease's progression difficult
47 to be carried out in the clinic. Moreover, DEXA is relatively expensive to implement and
48 requires adequate infrastructure.

49 Quantitative ultrasound (QUS) methods have been introduced as nonionizing alterna-
50 tives. The main motivation for the use of ultrasound lies in its ability to assess bone
51 mechanical properties, ultrasonic wave speed and attenuation being affected by the medium
52 of propagation¹⁶, which allows to go beyond a simple BMD (Bone Mineral Density)
53 estimation^{4,5,15,17,18}. QUS techniques have the advantage of being simple, safe, and cost-
54 effective. However, it should be noted that there is a lack of standardized quality control
55 and a great heterogeneity in the technologies used, making the results obtained by differ-
56 ent devices difficult to compare. The most widely QUS techniques used in the clinic are
57 transverse and axial transmission devices.

58 Ultrasonic guided waves are widely used in the field of nondestructive testing to character-
59 ize materials such as composite or bonded plates, for example¹⁹⁻²³. This method has been
60 adapted in recent years for cortical bones^{24,25}. Axial transmission uses ultrasonic guided
61 waves and is sensitive to intrinsic properties of cortical bone. This method is applied to
62 long bones such as the tibia, radius, or femur²⁶⁻²⁹. The clinical significance of ultrasound

63 velocities in reflecting various aspects of bone strength, such as cortical thickness, stiffness,
64 and porosity, has been partially confirmed. The initial studies on axial transmission were
65 primarily focused on measuring the time-of-flight (TOF) of the first arriving signal (FAS)³⁰.
66 The velocity of the fastest wave mode was then calculated and his typically associated with
67 the fundamental symmetrical mode of a plate³¹. Several clinical studies demonstrated that
68 the FAS velocity can be used to distinguish healthy from osteoporotic individuals^{29,32,33}.

69 However, assessing the mechanical and geometrical properties of bone cortical tissue us-
70 ing ultrasonic guided waves involves the complex issue of solving multiparametric inverse
71 problems^{3,18}. This resolution process relies on inversion algorithms that attempt to match
72 experimental data with simulated dispersion curves of ultrasonic guided wave modes, com-
73 puted for simplified models such as plates or cylinders^{34,35}. In previous studies, two ap-
74 proaches were proposed: operating at low frequencies (<500 kHz) and high frequencies (>1
75 MHz).

76 Low-frequency guided waves can penetrate more deeper inside cortical bone due to lower
77 attenuation and longer wavelengths. This ability to assess bone in its volume makes them
78 potentially useful to determine bone thickness, particularly in the early stages of osteoporosis
79 when the endosteal region is primarily affected³⁶. Moilanen et al.³⁷ introduced an inversion
80 scheme using a free isotropic plate model to estimate the cortical thickness of phantom
81 plates, which was later applied to tubular structures and ex-vivo bones^{35,38}. Similarly, Ta et
82 al.³⁹ used a hollow isotropic cylinder filled with a viscous liquid to determine the thickness of
83 bovine tibia specimens by manually matching experimental velocities with the phase veloci-
84 ties of simulated modes. Pereira et al.⁴⁰ studied the interaction of very low frequency (20-40

85 kHz) ultrasonic guided wave modes with an irregular, multi-layered, and heterogeneous bone
86 cross-section modeled with anisotropic and viscoelastic material properties. Their results
87 indicated that the FAS is not a good discriminator of intracortical bone properties, high-
88 lighting the need to target specific modes and not only the FAS to access bone properties.
89 Consequently, low-frequency methods face limitations, including the challenge of retrieving
90 multiple bone parameters beyond cortical thickness due to the limited information contained
91 in the few low-order modes typically utilized. Defining the appropriate operating frequency
92 range remains a key challenge in low-frequency assessments.

93 Higher frequencies are more likely to retrieve several parameters of the bone cortex
94 through inverse scheme. At these frequencies, the bone shape can be approximated by
95 a plate-like structure. Foiret et al.⁴¹ presented an inverse characterization method to es-
96 timate both the thickness and bulk wave velocities of *ex-vivo* cortical bone samples, using
97 a transverse isotropic free plate model with a user-parameter-dependent algorithm. Sub-
98 sequently, the same research group utilized inversion schemes based on genetic algorithms
99 to extract information from a multimode dataset, without the necessity of prior knowledge
100 regarding the mode orders³⁴. Several studies have highlighted the complexity introduced by
101 the presence of soft tissues when extracting dispersion curves, leading to a larger number
102 of modes and higher attenuation⁴²⁻⁴⁴. More recently, a bilayer plate model was introduced
103 to explore the accuracy of the inversion method when dealing with more complex models
104 and the presence of soft tissue³. Results indicated that while the free plate model (without
105 the soft tissue layer) allows for reliable retrieval of waveguide properties, the bilayer model
106 could become unmanageable for solving the inverse problem due to the additional modes

107 induced by the soft tissue layer. Li et al.⁴⁵ later introduced deep neural networks to re-
108 place traditional inverse algorithms for better estimation of cortical parameters. Despite the
109 potential to simplify the model geometry, higher frequencies may present significant chal-
110 lenges in managing the inverse problem due to the numerous modes generated, which can
111 accommodate almost any set of model parameters.

112 The originality of the present method lies in using low-frequency guided waves (from 50
113 kHz to 500 kHz) to retrieve the cortical thickness and properties of phantom plates topped
114 with soft tissue, employing a bilayer model representative of the experimental samples.
115 This frequency range reduces the number of propagating modes despite the presence of
116 soft tissue, thereby increasing the accuracy of the proposed inverse method. Additionally,
117 Pereira et al.⁴⁶ demonstrated that using the amplitude of ultrasonic guided wave modes can
118 significantly enhance the accuracy of the inversion, although their model did not account
119 for soft tissue. Thus, incorporating an excitability value into the inversion process of the
120 proposed method ensures that the inversion is based on both the dispersive properties of
121 ultrasonic guided waves and their amplitude, making the model sensitive to both cortical
122 thickness and cortical properties.

123 This paper aims to simulate a parameterized cortical bone phantom plate with a soft
124 tissue layer using the semi-analytical iso-geometric analysis (SAIGA)^{47,48} to perform the in-
125 verse characterization of the mechanical and the geometrical properties of soft tissue covered
126 bone phantom plates. Two cortical phantom plates coupled with a soft tissue mimicking
127 layer were used experimentally. A proprietary ultrasonic transducer specifically designed to
128 excite ultrasonic guided waves under 500 kHz was used for measurements. Acquired data

129 were processed using the 2D Fast Fourier Transform to extract the dispersion curves of the
130 propagating modes. The curves were then compared with the semi-analytical predictions.
131 For both plates, five parameters were estimated using the inversion scheme: (1-2) the longi-
132 tudinal and shear velocities in the cortical layer; (3-4) thicknesses of both layers; and (5) the
133 sound velocity in the soft tissue layer. To ensure that the inversion between simulated and
134 experimental dispersion curves was based on the dispersive properties of ultrasonic guided
135 waves and their amplitudes, excitability of the different guided wave modes is taken into
136 consideration in the cost function.

137 II. MATERIAL AND METHODS

138 A. Models and Simulations Models

139 A numerical model comprising a bone phantom plate coupled with a soft tissue layer,
140 mirroring experimental bone phantom plates, was used to match experimental dispersion
141 curves with ultrasonic guided wave modes calculated by SAIGA^{47,48}. Solving SAIGA equa-
142 tions enables the calculation of modes as well as their amplitude. Section II A 1 describes
143 the plate like model used as a forward model in this study. Section II A 2 then summarizes
144 background of the SAIGA method and detailed the formulation of the mode excitability
145 used.

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1. Plate-like Model

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The model geometry was defined based on the experimental bone phantom plates. Consider a homogeneous isotropic solid layer (Ω^S) of infinite extent and constant thickness h^S in the direction of propagation (e_1). The upper surface of the solid layer is loaded with soft tissue layer (Ω^f) modeled as fluid of thicknesses h^f . Fig. 1(a) shows a description of the model's geometry.

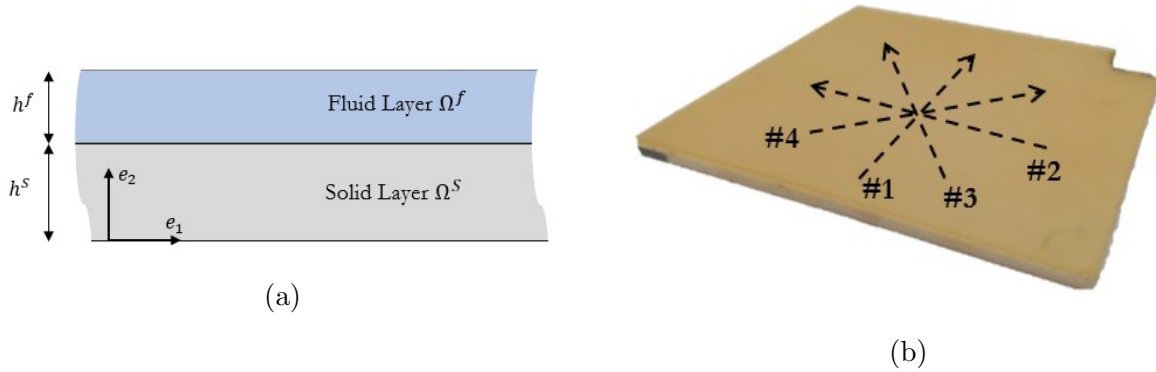


FIG. 1. (a) Geometry description of the cortical solid layer topped by a fluid finite space. (b) Experimental bone phantom plates with the four different directions of axial transmission measurements.

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The cortical layer of the waveguide was modeled as a homogeneous viscoelastic medium defined by its density (ρ), its Young's modulus (E) and Poisson's ratio (ν). Longitudinal and shear velocities of the solid layer are then derived from these parameters and used for the inversion process. This enable the respect of thermodynamic laws and avoid having results which have no physical meaning or values too far away from those found in the literature^{49–54}. The resulting model parameters, which reflect the waveguide's stiffness, consist of two bulk waves velocities C_P and C_S , incorporating the mass density within

159 the velocity parameters^{3,17,18,55}, and a stiffness ratio of elastic coefficients $C_{11}/C_{13} = 1 -$
 160 $2(C_S/C_P)^2$. For the longitudinal and shear waves, two attenuation coefficients α_P and α_S
 161 were added to the model and used for the calculation of the viscoelastic tensor. The ratio
 162 between α_S and α_P at a frequency of 1 MHz was fixed at 1.5^{56,57}, with $\alpha_P = 2.9$ dB/cm
 163 at this frequency in accordance with the manufacturer properties. Therefore, viscoelastic
 164 tensor coefficients can be calculated as^{58,59}:

$$\eta_{11} = \eta_{22} = \frac{\alpha_P}{8.686} \times 2C_P \times \frac{C_{11}}{(\omega_{ref})^2} \quad (1a)$$

$$\eta_{33} = \frac{\alpha_S}{8.686} \times 2C_S \times \frac{C_{33}}{(\omega_{ref})^2} \quad (1b)$$

$$\eta_{12} = \eta_{11} - 2 \times \eta_{33} \quad (1c)$$

165 where $\omega_{ref} = 2\pi \cdot f_{req}$ with $f_{req} = 1$ MHz the reference frequency used for the calculation of
 166 the viscoelastic tensor. The values of η_{11} , η_{33} and η_{12} are assumed to be constant within the
 167 frequency bandwidth of interest and determined by the value of the attenuation coefficients
 168 at frequency f_{req} . The effective stiffness tensor was then express as a complex tensor $C_{eff} =$
 169 $C + i\eta\omega_{ref}$. In the same way, the soft tissue layer was model as a fluid with its density (ρ^f),
 170 its sound velocity (C^f), and its attenuation coefficient $\alpha_f = 0.6$ dB/cm at 1 MHz, with its
 171 bulk modulus $K = \rho^f(C^f)^2$ and its viscoelastic coefficient $\eta^f = \frac{\alpha_f}{8.686} \times 2C^f \times \frac{K}{(\omega_{ref})^2}$. NURBS
 172 basis functions of order 8 were used for the SAIGA analysis. Three patches were used in
 173 each layer so that the number of degrees of freedom $N_{Dof} = 33$. Dispersion curves were
 174 computed within a frequency range from 50 to 500 kHz at a step of 5 kHz.

175 **2. Semi-Analytical Iso-geometric Analysis (SAIGA) Method**

176 The SAIGA method uses the same principle as the semi-analytical finite element (SAFE)
 177 method^{60–62}, but instead of using Lagrange polynomials for approximation, SAIGA uses
 178 B-splines and non-uniform rational B-Splines (NURBS) basis functions, providing higher
 179 continuity across the element boundaries, therefore significantly improving the precision,
 180 and reducing the computational cost^{63,64}. To determine the dispersion of guided waves, the
 181 method considers the waves propagating harmonically in the axial direction (e_1), which can
 182 be expressed for the displacement \mathbf{u} and the acoustic pressure \mathbf{p} as follows:

$$\mathbf{u}(x_1, x_2, t) = \mathbf{U}(x_2)e^{i(k_1x_1 - \omega t)} \quad (2a)$$

$$\mathbf{p}(x_1, x_2, t) = \mathbf{P}(x_2)e^{i(k_1x_1 - \omega t)} \quad (2b)$$

183 where ω is the angular frequency, t is the time, k_1 is the wave number in the propagation
 184 direction (e_1), vector $\mathbf{U}(x_2) = (U_1, U_2)^T$ and $\mathbf{P}(x_2) = P$ are respectively the amplitude of
 185 displacement in the solid domain Ω^S and pressure in the fluid domain Ω^f . Essentially, em-
 186 ploying SAIGA method leads to a quadratic eigenvalue problem to be solved for determining
 187 the relationship between the wavenumber (k_1) and angular frequency (ω). It could be solved
 188 with respect to k_1 as follows:

$$(-\omega^2\mathbf{M} + \mathbf{K}_0 + ik_1\mathbf{K}_1 + k_1^2\mathbf{K}_2)\mathbf{V} = \mathbf{0} \quad (3)$$

189 where $\mathbf{V} = (\mathbf{P}, \mathbf{U})^T$ contains the global eigenvectors of pressure (\mathbf{P}) and of displacement
 190 (\mathbf{U}), and $\mathbf{M}, \mathbf{K}_0, \mathbf{K}_1, \mathbf{K}_2$ are the global matrices of the system. More details can be found
 191 in the work of Seyfaddini and al.^{47,48}. By solving Eq. (3), it is possible to determine the

192 eigenvalues k_1 and the corresponding eigenvectors $\mathbf{V}(\omega, k_1)$ of guided modes for each value
 193 of angular frequency ω . The frequency dependent phase velocity (C_{ph}) and the attenuation
 194 (att) can be determined by:

$$C_{ph} = \omega / \Re(k_1) \text{ m/s} \quad att(\omega) = \Im(k_1) \text{ Np/m} \quad (4)$$

195 To ensure that the inversion between simulated and experimental dispersion curves was
 196 based on the dispersive properties of ultrasonic guided waves and their amplitudes, the
 197 excitability Ex of a given mode was calculated based on its mode shape as:

$$Ex(k_1, \omega) = \frac{\Re(k_1, \omega)}{\Im(k_1, \omega)} \times U_{norm}^2(k_1, \omega) \quad (5a)$$

$$U_{norm}(k_1, \omega) = \frac{abs(U_{2,surf})}{max(U_{1S}, U_{2S}, U_{1f}, U_{2f})} \quad (5b)$$

198 where U_{1S} , U_{2S} , U_{1f} , U_{2f} are respectively the in plane and out of plane displacement in the
 199 solid layer, and the in plane and out of plane displacement in the top fluid layer. $U_{2,surf}$ is
 200 the out of plane displacement at the top surface. An example of the excitability of guided
 201 wave modes in the bi-layer model as a function of frequency is displayed in Fig. 2(a). The
 202 color of each circle denotes the amplitude of the mode excitability associated with each
 203 mode. Fig. 2(b) represents the mode shapes associated with the selected mode on Fig. 2(a),
 204 with the different displacement used in the calculation of excitability.

205 B. Inversion scheme

206 It was assumed that matching experimental dispersion curves with the semi-analytical
 207 predictions generated via SAIGA may serve as a methodological approach for deducing the
 208 cortical bone properties (velocities C_P and C_S , thickness h^S) of the plate's solid layer. In

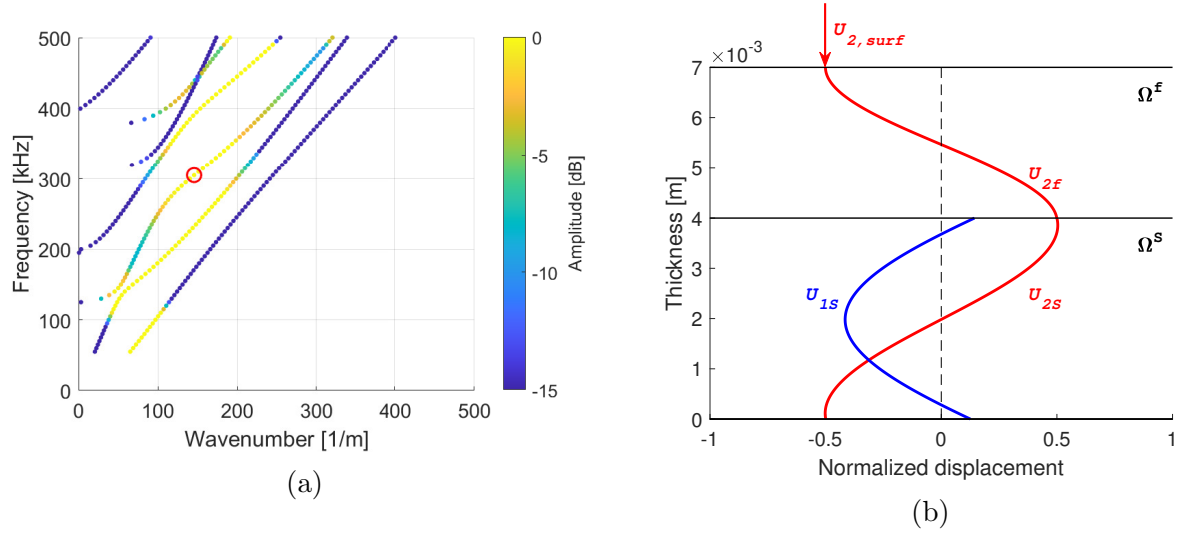


FIG. 2. (a) Guided modes dispersion curves with their frequency dependent excitability computed by SAIGA for a 2D cortical bone plate of 4mm ($\rho = 2310 \text{ kg/m}^3$, $C_P = 2900 \text{ m/s}$, $C_S = 1600 \text{ m/s}$) overlaid with 3mm soft tissue layer ($\rho^f = 1010 \text{ kg/m}^3$, $C^f = 1400 \text{ m/s}$). The color scale shows the excitability of each mode in dB normalized to the mode with the highest excitability at each frequency. (b) Mode shape of the selected mode (red circle on (a)).

209 pursuit of this objective, a cost function was formulated to ascertain the optimal fit between
 210 the experimental curves and a particular instantiation of the simulated dataset. This method
 211 offers a systematic way to estimates the cortical bone properties of the plates. The accuracy
 212 and robustness of this approach depends on several factors including the accuracy of the
 213 SAIGA method, the quality of the experimental dispersion curves and the choice of the cost
 214 function.

215 **1. Cost Function**

216 A cost function serves as a metric to evaluate the precision of a particular model in
 217 capturing the inherent associations within a dataset. A pronounced disparity between pre-
 218 dictions and actual outcomes leads to an elevated value of the cost function, while more
 219 accurate estimations correspond to reduced values. It represents the foundation for solving
 220 an inverse problem. The choice of the cost function can influence the performance of an
 221 inversion. It is important to use a function that represents the type of problem and the
 222 distribution of data. Using excitability enable to add information about the amplitude of
 223 the guided wave mode propagating in the model in addition with their dispersive trajectories
 224 in the frequency against wavenumber plot. Consequently, the cost function was designed to
 225 measure the difference of amplitude between experimental and simulated dispersion curves
 226 as shown in Fig. 3.

227 For a given set of parameters $\Lambda=(\rho E \nu h^S \rho^f C^f h^f)$, the amplitude difference at each
 228 frequency is given by:

$$d_n(\Lambda, f) = Ex(k_n(\Lambda), f) - A(k_n(\Lambda), f) \quad (6)$$

229 where $Ex(k_n(\Lambda), f)$ is the excitability of mode $k_n(\Lambda)$ of the simulated dispersion curves,
 230 $A(k_n(\Lambda), f)$ is the experimental amplitude observed at wavenumber k_n for the same fre-
 231 quency. Examples of distances d_n can be observed on Fig. 3(c).

232 To ensure that the inversion is based on modes with the highest amplitudes, a weighting
 233 factor χ_n is calculated as the excitability of a given mode divided by the maximum excitabil-
 234 ity of modes for each frequency⁴⁶. This ensures higher amplitude modes play a dominant

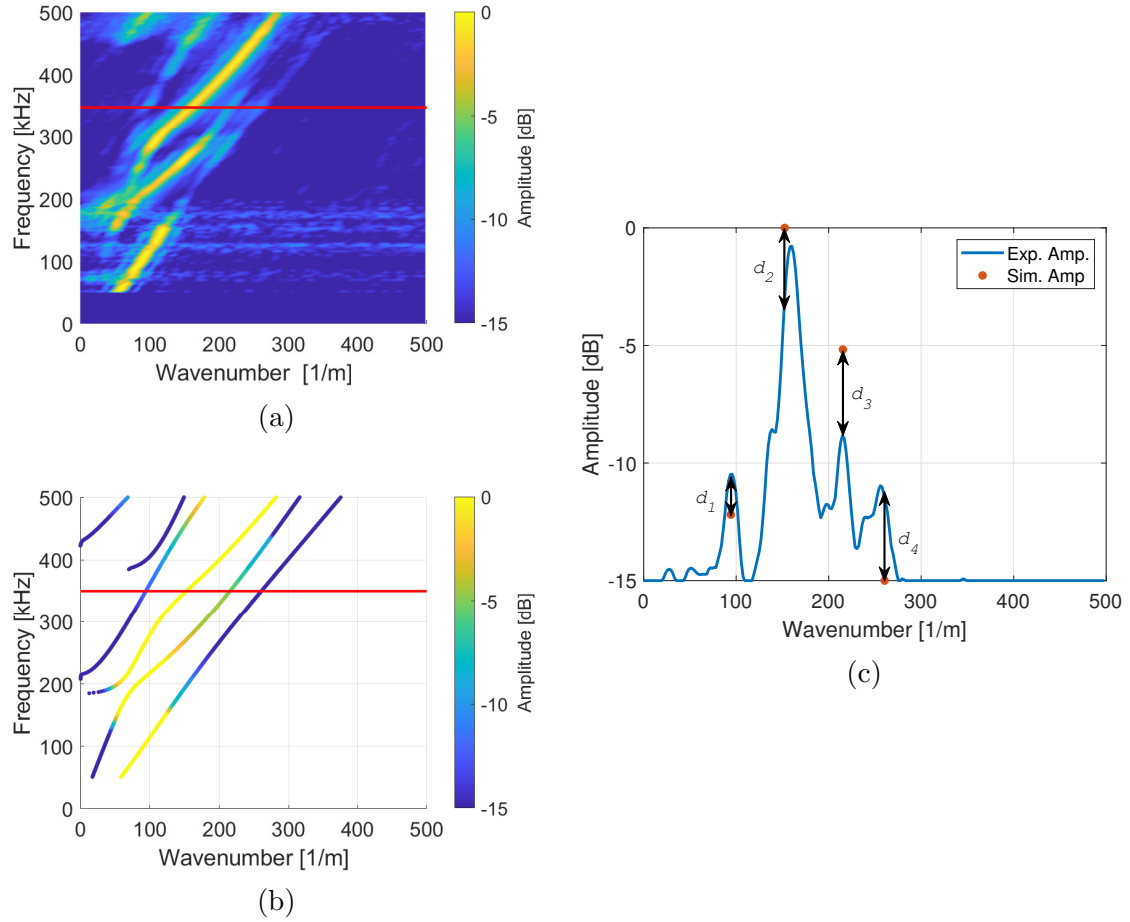


FIG. 3. (a) Surface plot representing experimental dispersion curves. (b) Simulated dispersion curves with mode excitability computed with SAIGA. (c) Projection of experimental amplitude at 350 kHz and simulated excitability at the same frequency, showing the distance measured for the calculation of the cost function.

235 role in the inversion process.

$$\chi_n(\Lambda, f) = \frac{Ex(k_n(\Lambda), f) - \xi}{-\xi} \quad (7)$$

236 where N is the total number of simulated modes for a given frequency. ξ is a user defined
 237 threshold setting the weighting factor equal to zero if the excitability of a given mode is
 238 inferior or equal to it. For example, by observing Fig. 3(c) which depicts distances between

239 simulated and experimental dispersion curves at 350 kHz, with a threshold $\xi = -15$ dB, the
 240 weighting factor $\chi_4 = 0$ as the excitability of the fourth mode is equal to ξ . For the other
 241 modes, we have $\chi_1 = 0.19$, $\chi_2 = 1.0$ as it's the higher amplitude mode, and $\chi_3 = 0.66$.

242 The cost function for a set of parameters $J(\Lambda)$ is then calculated as follow:

$$J(\Lambda) = \sum_f \sqrt{\sum_{n=1}^N (d_n(\Lambda, f) \times \chi_n(\Lambda, f))^2} \quad (8)$$

243 Such a cost function will therefore consider high amplitude modes and neglect those with
 244 an amplitude below the threshold. Doing so reduces the risk of using irrelevant information
 245 in the inversion and the solution can be considered as more accurate.

246 *2. Parameters sweeping*

247 To minimize the cost function and find the best fit between experimental and simulated
 248 data, the cost function was calculated for every model by sweeping each parameter along a
 249 multidimensional grid. Range of parameters were chosen to cover properties of both plates.
 250 This enables to test the effectiveness of our methodology in distinguishing between a bone
 251 in a healthy state and one afflicted by osteoporosis. Table I represents the set of parameters.
 252 Each range of parameters was defined large enough to ensure the measured properties from
 253 the plates are far enough from the range limits.

254 The grid steps were determined by balancing resolution fidelity and computational ef-
 255 ficiency. For the whole dataset, the total number of cases was 72 600. To ascertain the
 256 robustness of our method, each parameter within the model was subject to variation, en-
 257 compassing both values inside and outside the experimentally determined uncertainty range

TABLE I. Range of model parameters and their respective step used to perform the inversion.

	Parameters range	Step
ρ (kg/m ³)	[2050-2550]	50
E (GPa)	[11-21]	1
ν	[0.23-0.26]	0.01
h^S (mm)	[3.25-5.25]	0.5
ρ^f (kg/m ³)	1000	1
C^f (m/s)	[1275-1525]	50
h^f (mm)	[1.6-2.4]	0.2

258 of measured properties (see section II C 1 and III A). Based on the study of Mast⁶⁵, showing
 259 that density of soft tissue in the human body is comprised between 950 kg/m³ and 1100
 260 kg/m³, the choice was made to keep the soft tissue density ρ^f constant and equal to 1000
 261 kg/m³. As it will be shown in the results, the density of the soft tissue has virtually no
 262 effect on the dispersion curves. Emphasizing the significance of capturing all mode fea-
 263 tures, a frequency resolution increment of 5 kHz was selected, enabling 90 frequencies over
 264 the whole range of frequency [50 kHz – 500 kHz]. To counterbalance the computational
 265 demands inherent to such high-resolution, a comprehensive database encapsulating all pos-
 266 sible model parameter combinations was constructed over an extended, continuous modeling
 267 session lasting less than 48 hours, demonstrating the effectiveness of the SAIGA method.

268 The pre-existing database was utilized to calculate the cost function values within the 6-D
269 space, eliminating the need to rerun the model for every subsequent optimization scenario.

270 C. Experimental Measurements

271 1. *Cortical Bone Phantom Plates*

272 The measurements were performed with two cortical bone phantom plates layered with
273 soft tissue mimicking material provided by True Phantom Solutions (Windsor, ON, Canada).
274 One plate has material properties reflecting a healthy cortical bone and the other has prop-
275 erties similar to an osteoporotic bone. Both plates have the same material for the soft tissue
276 layer. They have the same dimension: 15x15 mm with approximately 4 mm of cortical layer
277 and 2 mm of soft tissue. Pulse-echo measurements were performed to acquire reference val-
278 ues for sound velocity in each layer which will be used in comparison with those obtained by
279 the inversion. To do so, longitudinal and shear single element piezoelectric probes were used
280 (Olympus V125-RM and Olympus V154-RM respectively). Small samples of each plate were
281 cut and used to measure the density of each layer. To do so, the sample was weighted and
282 its volume measure with a 3D laser scanner (Absolute Arm 85) from Hexagon (Stockholm,
283 Sweden). The measurement uncertainty of density is subsequently derived from the relative
284 uncertainties of the measuring instruments. The thicknesses of the samples were measured
285 20 times and pulse-echo measurements were then performed the same number of times to
286 measure velocities. Then, 20 pulse-echo measurements were realized at different locations
287 over the rest of the phantom plates to estimate the true thickness of each layer. Assumption

288 was made concerning the quasi-isotropic properties of the layers. However, it appeared that
289 the thicknesses of the layers were not homogeneous all over the plates, with 15% to 20%
290 variation for the cortical layers, and around 10% for the soft tissue layers. Therefore, mean
291 values will be used to compared with the results obtained by the inversion.

292 *2. Axial Transmission Measurements*

293 A proprietary ultrasonic probe was specifically built to perform ultrasonic excitation.
294 The probe had a central frequency of 400 kHz with a -6 dB bandwidth between 200 kHz
295 and 600 kHz approximately. Elements of the probe were specifically chosen to maximize the
296 emission of ultrasonic waves through soft tissue and cortical bone. The device was in contact
297 with the upper part of the soft tissue layer and coupled with an ultrasonic gel Ultragel®
298 II. A series of several Hann windowed tonebursts were used as input waveform, with central
299 frequencies varying between 50 kHz and 500 kHz with a step of 50 kHz. The number of
300 cycles evolved with the central frequencies to keep a bandwidth of 50 kHz for each toneburst.
301 Four consecutive acquisitions were realized on the plates (two in the length-width directions
302 and the two along the diagonals, see Fig. 1(b)) along the direction of propagation at 60
303 positions equally spaced by 1 mm using a laser Doppler vibrometer (Polytech OFV-505).
304 The probe was repositioned between each acquisition. Raw data were processed using the
305 2D Fast Fourier Transform⁶⁶, enabling the extraction of experimental dispersion curves for
306 each of the four acquisitions with the intensities of the propagating modes in a frequency-
307 wavenumber diagram.

308 **III. RESULTS**

309 **A. Phantom Plates Properties Measurement**

310 Due to uncertainties regarding the geometrical and material properties of the phantom
 311 plates, pulse-echo measurements were performed to determine the reference values of sound
 312 velocity and thickness of each layer. The plates were assumed homogeneous such that
 313 material properties and densities were measured on small samples of each layer. The same
 314 material was used for the soft tissue layer of both plates. As mentioned previously, it is
 315 worth noted that a discrepancy between 10% and 20% is observed for the different layer
 316 thicknesses over the whole surface of the plates. This can be explained by the method of
 317 fabrication of the plates. Table II summarizes the reference properties of each layer.

318 **B. Effect of the soft tissue density**

319 A study was conducted on the soft tissue parameters while maintaining other properties
 320 at a constant level. Subsequently, the maximal normalized cost function value was compared
 321 with the minimal value ($J(\Lambda) = 1$) corresponding to the best fit scenario. In accordance
 322 with Mast⁶⁵ and other relevant studies⁶⁷, the parameter range for the soft tissue density
 323 ρ^f was defined to be between 900 kg/m³ to 1200 kg/m³. The ranges for C^f and h^f are
 324 respectively [1250-1550] m/s and [1.0-2.5] mm. The study was performed on the healthy
 325 plate and each parameter can take 31 values.

326 Examining the outcomes yielded by altering solely the soft tissue density ρ^f , the normal-
 327 ized cost function pertaining to the least favorable fitting scenario registers at $J_{max}(\Lambda_{\rho^f}) =$

TABLE II. Bone Phantom Plates Properties.

	Healthy Cortical	Osteo. Cortical	Soft Tissue
	Layer	Layer	Layer
ρ (kg/m ³)	2416 ± 16	2139 ± 14	977 ± 12
C_P (m/s)	3000 ± 115	2655 ± 65	1390 ± 90
C_S (m/s)	1755 ± 50	1530 ± 35	-
h^S (mm)	4.3 ± 0.7	4.3 ± 0.8	-
	Soft Tissue	Soft Tissue	
	Healthy Plate	Osteo. Plate	
h^f (mm)	2.0 ± 0.2	2.1 ± 0.2	-

328 1.086. In contrast, the normalized cost function values for the least favorable fittings
329 when altering solely the soft tissue velocity C^f or the soft tissue thickness h^f stand at
330 $J_{max}(\Lambda_{C^f}) = 2.260$ and $J_{max}(\Lambda_{h^f}) = 2.948$, respectively. Furthermore, upon examining the
331 qualitative impact on the simulated dispersion curves (Fig. 4), ρ^f does not exert a substan-
332 tial influence on the trajectory of high-amplitude modes when contrasted with C^f and h^f ,
333 where variations in these parameters notably affect the modes of significance. Therefore,
334 fixing the soft tissue density can be considered as having virtually no effect on the inversion
335 results but reduces the parameter exploration in the fitting. The decision was taken to set
336 the soft tissue density ρ^f as a constant in our sweeping parameters.

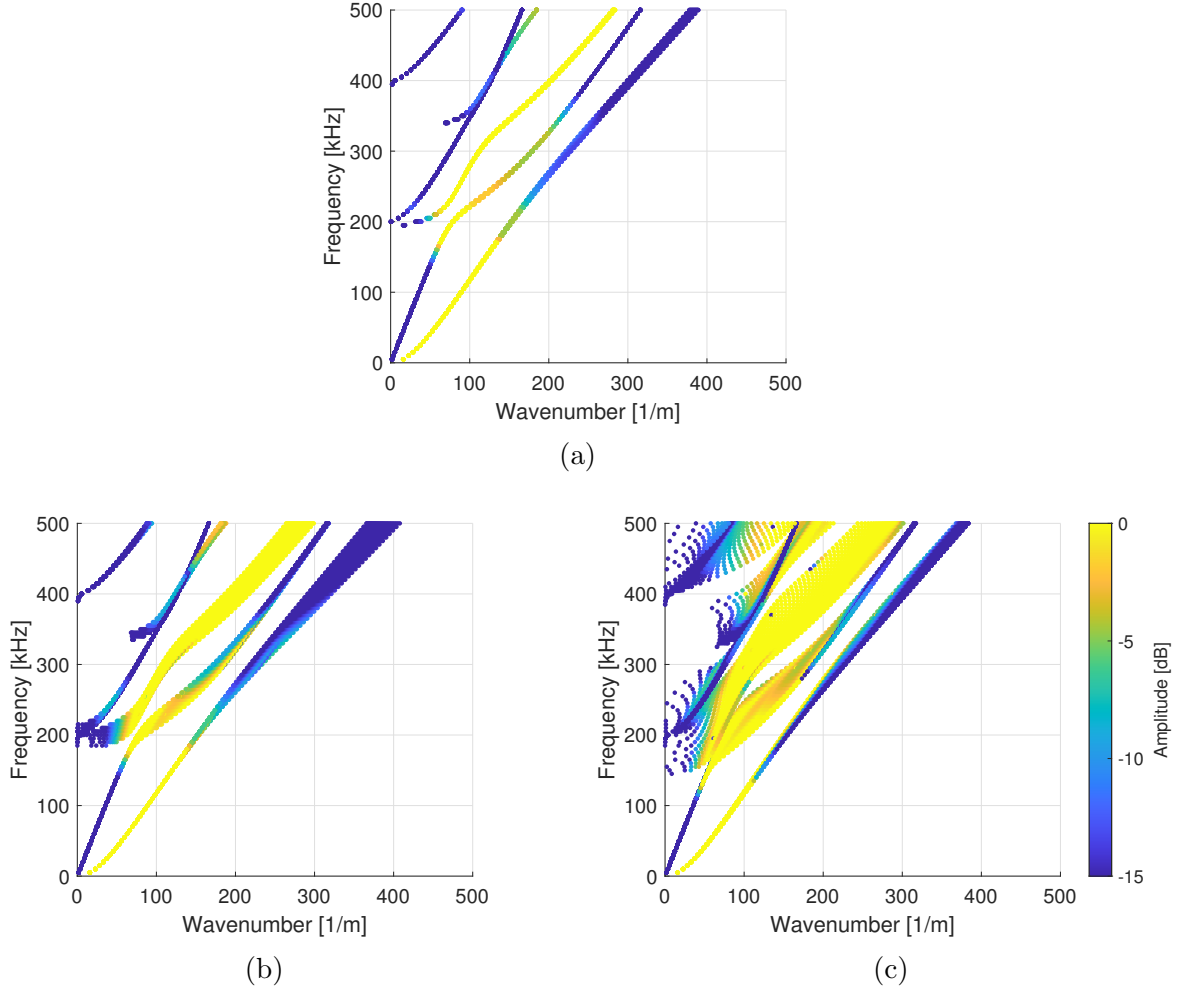


FIG. 4. Influence of property variations on the trajectory of ultrasonic modes: (a) soft tissue density ρ^f , (b) wave velocity in soft tissue C^f , and (c) thickness of the soft tissue h^f .

337 **C. Fitting of Simulated Data**

338 As previously mentioned, four acquisitions were conducted for each plate along various di-
 339 rections to measure experimental dispersion curves. The inversion procedure was executed
 340 for both plates to determine the optimal set of parameters from the database. Fig. 5(a-
 341 b) illustrates the most favorable outcomes achieved through the proposed method. Semi-
 342 analytical predictions are depicted as points overlaying the experimental surface plot. The

343 experimental plots depicted herein are the result of averaging the 2D-FFT plots obtained
344 from the four separate axial transmission measurements. This approach aims to consolidate
345 spatial frequency information across multiple measurements for enhanced clarity in visual
346 analysis. The alignment of experimental and simulated dispersion curves demonstrates a
347 highly satisfactory agreement, with all high excitability modes perfectly matched. A thresh-
348 old of -15dB was established to mitigate the impact of noise in experimental measurements,
349 facilitating clearer dispersion curves and facilitating a more straightforward fitting process.
350 Furthermore, the attenuation of modes in the simulated data closely aligns with the atten-
351 uation observed experimentally.

352 **D. Estimated Properties**

353 The mean of the inverse characteristics over the four axial transmission measurements
354 corresponding to the optimal fitting of each plate are detailed in Table III, alongside refer-
355 ence properties and associated relative errors. The findings indicate a strong agreement in
356 property values, revealing errors under 2.0% for the cortical layers. However, a more no-
357 table margin is observed for the soft tissue thickness, with errors up to 5%. The soft tissue
358 layer thickness is not uniform along the plates. Additionally, the pulse-echo measurements
359 introduce the potential for inaccuracies, as the layers may undergo compression during the
360 assessment.

361 Fig. 5(c-d) illustrates the inverted values in conjunction with the reference properties
362 and their respective uncertainties. The derived values are obtained through the averaging
363 of the reciprocals acquired from individual measurements across the four axial transmission

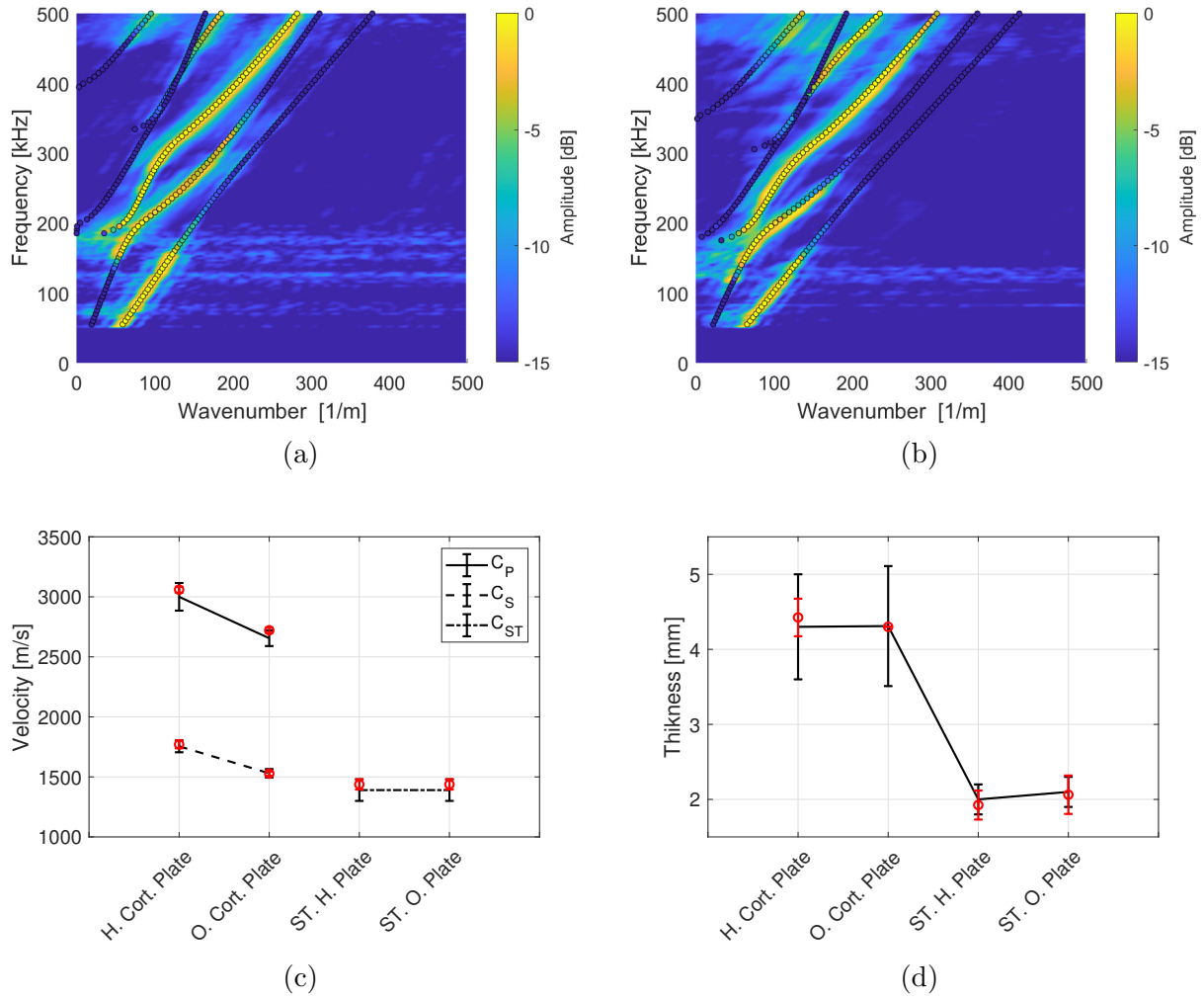


FIG. 5. Optimal fitting results between experimental and simulated dispersion curves for (a) healthy phantom plate, and (b) osteoporotic phantom plate. Average inverse values in conjunction with the reference properties and their respective uncertainties, (c) velocity properties, (d) thickness properties.

364 scenarios. Associated uncertainties are represented by their respective standard deviations.

365 Each inverse value falls within the uncertainty range of its corresponding property.

TABLE III. Average Inverse Properties of Phantom Plates with their Relative Errors.

	Healthy Plate			Osteo. Plate		
	Ref.	Inv.	Err.%	Ref.	Inv.	Err.%
C_P (m/s)	3000 ± 115	3052 ± 30	1.74	2655 ± 65	2653 ± 82	0.07
C_S (m/s)	1755 ± 50	1755 ± 29	0.02	1530 ± 35	1508 ± 45	1.39
h^S (mm)	4.30 ± 0.7	4.40 ± 0.25	1.74	4.30 ± 0.8	4.25 ± 0.0	1.16
	Soft Tissue Healthy Plate			Soft Tissue Osteo. Plate		
C^f (m/s)	1390 ± 90	1437 ± 25	3.42	1390 ± 90	1437 ± 48	3.42
h^f (mm)	2.00 ± 0.2	1.95 ± 0.3	2.50	2.10 ± 0.2	2.00 ± 0.16	4.76

366 **1. Estimated Bulk-Velocities**

367 Fig. 5(c) displays the mean and standard deviation obtained for longitudinal and shear
368 velocities in the cortical layer of both plates, as well as the wave velocities in their corre-
369 sponding soft tissue layer (red circles), alongside the reference average values obtained by
370 pulse-echo measurement (black lines). The predicted values fell within the uncertainty range
371 of the reference values for both plates, and their associated standard deviations were of a
372 comparable range to the reference ones. It is worth remembering that ultrasonic pulse-echo
373 measurements were conducted across the plates (dimension of 150 x 150 mm), and the four
374 axial transmission measurements were taken along a 60 mm length at four different locations
375 (two diagonals and two perpendicular axes). Therefore, the predicted values approximately

376 represent the average values across the plates, as well as the reference values. The error
377 was found to be less than 2% for cortical velocities and up to 3.4% for soft tissue velocity.
378 However, the average inverted values of the soft tissue velocity were found to be equal for
379 both plates, suggesting that the reference velocity of the soft tissue layer may have been
380 underestimated during pulse-echo measurement. Changes in the experimental environment
381 could have resulted in slight variations in the properties of the soft tissue sample, accounting
382 for this minor difference.

383 2. *Estimated Thicknesses*

384 Fig. 5(d) shows the comparison between the estimated thickness of each layer (red circles)
385 and their corresponding reference values (black lines). Similar to the velocities, a very
386 good agreement can be observed between the predicted and reference values. However, as
387 explained in section III A, the thickness values of each layer were not homogeneous over
388 the plates, which leads to uncertainties in the reference thicknesses measured by pulse-
389 echo measurement. Regarding the cortical phantom layer, these reference values can be
390 interpreted as the equivalent thickness across the entire plate area. Analogous to an *in-*
391 *vivo* radius, the predicted thickness represents an averaged thickness over the entire volume
392 corresponding to the investigated region of interest. Axial transmission measurements reflect
393 the mean thickness along the propagation direction. Four axial transmission measurements
394 were conducted in four different directions, each separated by 45 degrees, yielding a predicted
395 thickness that corresponds to the mean actual thickness of the plate. Inverse properties were
396 found to be close enough to the reference values to exhibit an error of less than 2% for the

397 cortical thicknesses and 5% for the soft tissue layers but remains in the standard deviation
398 of the reference values.

399 **IV. DISCUSSION**

400 The cost function value was computed across the multidimensional model grid, success-
401 fully identifying a global minimum for each acquisition and sample within the cost function
402 map. Sweeping through six dimensions is notably computationally intensive. Even when
403 using SAIGA, which enables a case calculation within a few seconds, it requires approxi-
404 mately 48 hours to run through the 72 600 scenarios. However, leveraging the existing
405 database enabled a gradual reduction in grid calculations to just a few minutes. Sweep-
406 ing the model parameters across a finite set (rather than employing an optimization search
407 routine) ensures that the global minimum is attained at each acquisition without needing ad-
408 ditional verification or concern regarding the optimization search’s performance and control
409 variables.

410 **A. Assessment of Cortical Layer Density**

411 Despite the promising match observed between the experimental data and the simulated
412 model, uncertainties persist regarding local minima. Upon investigation of cortical density,
413 the inverse values for the healthy cortical plate and the osteoporotic cortical plate are $2237 \pm$
414 165 kg/m^3 and $2337 \pm 221 \text{ kg/m}^3$, respectively, with errors of 7.4% and 9.3% compared to
415 reference values. Through analysis of the distribution of cost function values for experimental
416 axial transmission measurement #4 on both plates (see Fig. 6), a local minimum can be

417 identified for each density in our parameter set. These local minima remain very close to
 418 each other, with a maximal normalized cost function of $J_{max}(\Lambda_\rho) = 1.097$, and no visual
 419 differences can be seen between each solution. These uncertainties may arise concerning the
 420 inverted properties.

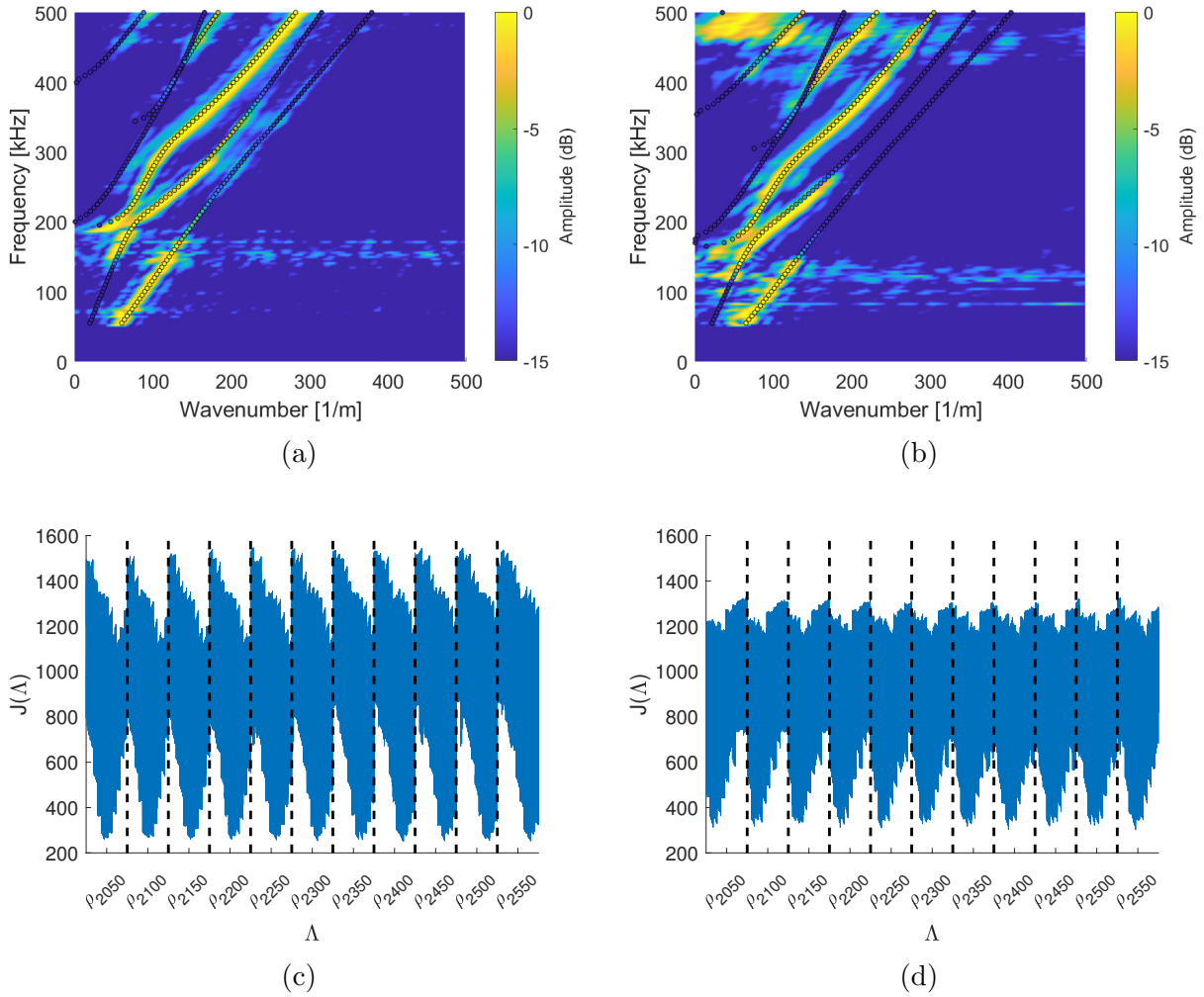


FIG. 6. Optimal fitting results between experimental axial transmission measurement #4 and simulated dispersion curves for (a) healthy phantom plate, and (b) osteoporotic phantom plate. Distribution of the cost function values with the corresponding density for experimental axial transmission measurement #4 (c) on the healthy plate and (d) on the osteoporotic plate.

421 However, upon inspection of the inverted cortical properties at each density (from 2050
 422 kg/m^3 to 2550 kg/m^3), the inverse values of longitudinal and shear velocities remain nearly
 423 unchanged, along with a significant portion of the other parameters, except for the Young's
 424 modulus (refer to Table IV). These phenomena can be elucidated by the reciprocal adjust-
 425 ment of the density and the Young's modulus, which serve to attain longitudinal and shear
 426 velocities consistent with the experimental dispersion curves. Moreover, looking at our range
 427 of parameters $\rho = [2050 : 50 : 2550] \text{ kg/m}^3$ and $E = [11 : 1 : 21] \text{ GPa}$, for a constant Poisson
 428 coefficient $\nu = 0.26$, target velocities can be obtained through a large number of pairs ($\rho - E$)
 429 as shown on Fig. 7.

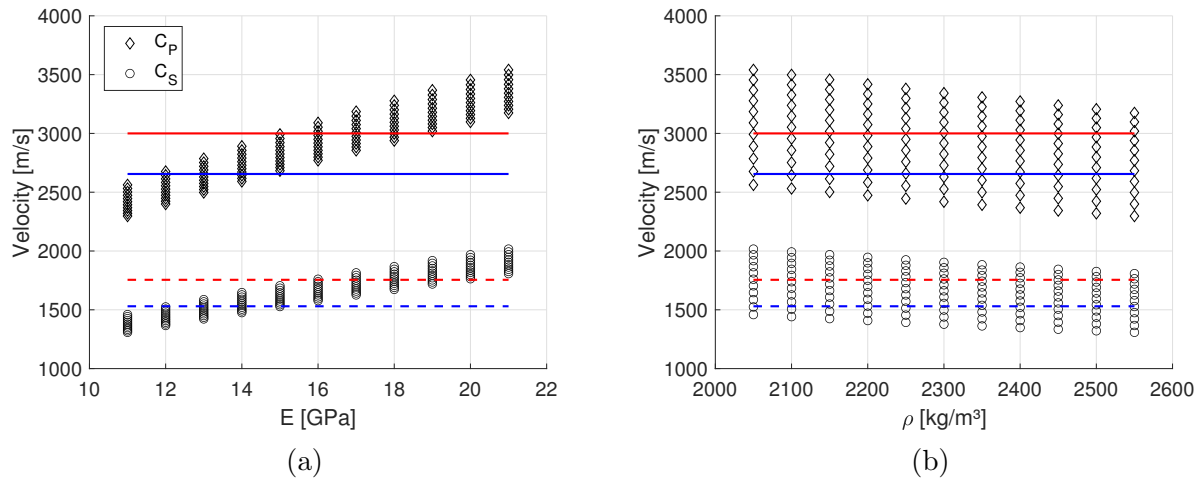


FIG. 7. Longitudinal and shear velocities possible values obtained through the set of parameters express as a function of (a) the Young's modulus E for different values of the density ρ , (b) the density ρ for different values of E . Red lines represent the reference values of C_P (full line) and C_S (dote line) for the healthy plate and blue lines for the osteoporotic plate.

430 As shown in Table IV, density and Young's modulus compensate each other to achieve
 431 the correct velocities. The higher the density, the higher the Young's modulus, for a given

TABLE IV. Mean inverted cortical properties for each frequency, relative to the optimal fit identified for axial transmission measurement #4.

	Healthy Plate Meas. #4			Osteo. Plate Meas. #4		
	Best fit	Mean over	Var.%	Best fit	Mean over	Var.%
	Meas. #4	densities	(σ/ref)	Meas. #4	densities	(σ/ref)
ρ (kg/m ³)	2400	2300 ± 166	6.90	2550	2300 ± 166	6.51
C_P (m/s)	3030	3023 ± 26	0.86	2683	2675 ± 29	1.08
C_S (m/s)	1725	1722 ± 15	0.87	1528	1523 ± 16	1.05
h^S (mm)	4.25	4.25 ± 0.0	0.00	4.25	4.25 ± 0.0	0.00
E (GPa)	18.0	17.2 ± 1.3	7.22	15	13.4 ± 1.0	6.67
ν	0.26	0.26 ± 0.0	0.00	0.26	0.26 ± 0.0	0.00

432 value of the Poisson coefficient. Consequently, the proposed method cannot directly assess
 433 the density of the cortical layer.

434 However, several studies have demonstrated the possibility of obtaining bone density
 435 through mathematical models^{53,68,69}. By adapting the work of Vu⁵³ and Pereira⁴⁶ for use
 436 with the phantom plate material used in this study, it is possible to estimate ρ from the
 437 bulk velocities.

438 For a 2D isotropic plate, the elasticity coefficients are given by $C_{11} = \rho C_P^2$, $C_{33} = \rho C_S^2$,
 439 $C_{12} = C_{11} - 2C_{33}$. Using experimental pulse-echo measurement values, a linear interpola-

440 tion can be made to express C_{11} , C_{12} and C_{33} as a function of the density. Consequently,
 441 longitudinal and shear bulk velocities can be expressed as functions of density (Fig. 8). This
 442 enables the determination of density for both plates corresponding to the inverse velocities
 443 depicted in Table III. The results yield $\rho = 2445 \pm 35 \text{ kg/m}^3$ and $\rho = 2130 \pm 14 \text{ kg/m}^3$ for
 444 the healthy plates and the osteoporotic plate, respectively, with relative errors of 1.2% and
 445 0.4%.

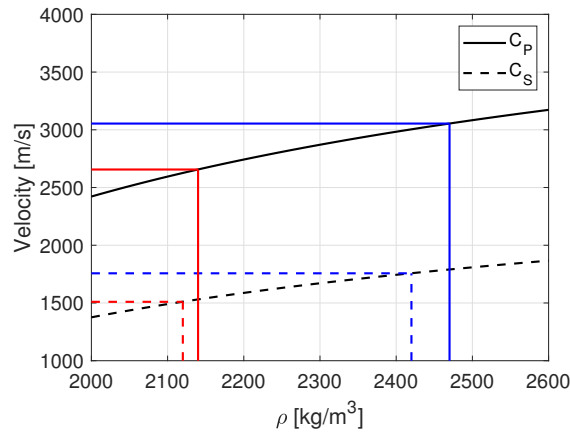


FIG. 8. Bulk velocities varying with the bulk density, derived from the values of the elastic properties for an isotropic medium adapting from the homogenization theory introduced in⁵³ and used in⁴⁶ to the phantom material. (Blue lines) Inverse values for longitudinal and shear velocities of Healthy phantom plate. (Red lines) Inverse values for longitudinal and shear velocities of Osteoporotic phantom plate.

446 **B. Importance of Excitability**

447 Utilizing excitability inherently prolongs computation time for each scenario, necessitat-
 448 ing the calculation of mode shape for every mode at each frequency across all parameters

449 sets. Nonetheless, excitability guarantees the consideration of all modes exhibiting high am-
 450 plitudes. Upon examining visually results from simulated datasets without the application
 451 of excitability, it becomes evident that inversion biases towards modes with high amplitudes
 452 across a broad frequency spectrum while overlooking modes of lower amplitudes. Conse-
 453 quently, certain modes are entirely disregarded, resulting in reduced thickness values and
 454 elevated velocities. As illustrated in Fig. 9, the second mode consistently disappears from the
 455 simulated curves. As depicted in Table V, the inverse values demonstrate an error exceed-
 456 ing 15% in comparison to the reference properties. Notably, several simulation parameters,
 457 particularly the thickness of cortical and soft tissue layers, approach the lower bounds of
 458 their respective ranges.

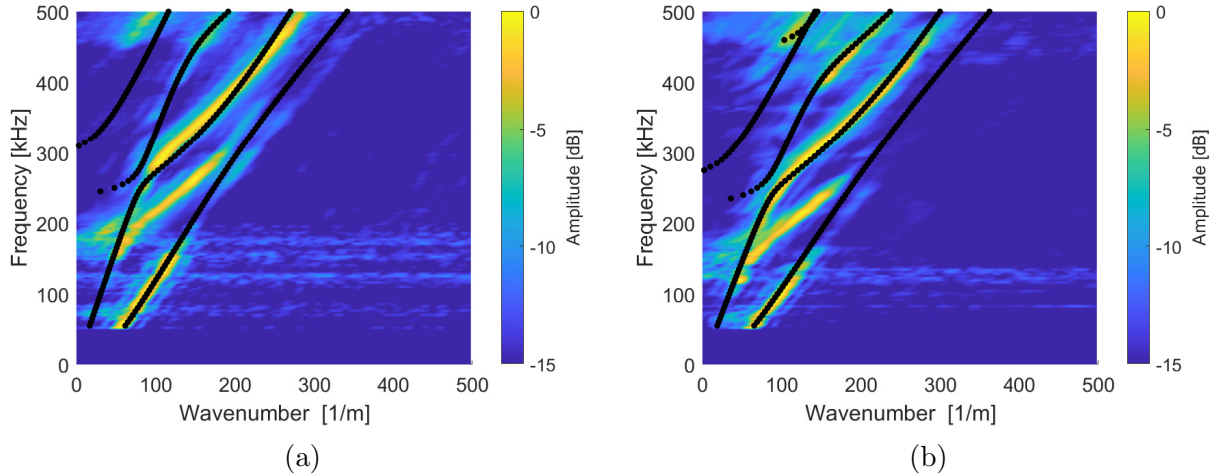


FIG. 9. Optimal fitting results between experimental and simulated dispersion curves without excitability for (a) healthy phantom plate, and (b) osteoporotic phantom plate.

TABLE V. Average Inverse Properties of Phantom Plates with their Relative Errors without using the excitability in the inversion scheme.

	Healthy Plate			Osteo. Plate		
	Ref.	Inv.	Err.%	Ref.	Inv.	Err.%
C_P (m/s)	3000 ± 115	3475 ± 0.0	15.83	2655 ± 65	3064 ± 52	15.39
C_S (m/s)	1755 ± 50	2032 ± 0.0	15.81	1530 ± 35	1785 ± 39	16.09
h^S (mm)	4.30 ± 0.7	3.25 ± 0.0	24.42	4.30 ± 0.8	3.25 ± 0.0	24.42
	Soft Tissue Healthy Plate			Soft Tissue Osteo. Plate		
C^f (m/s)	1390 ± 90	1487 ± 75	7.01	1390 ± 90	1500 ± 29	7.91
h^f (mm)	2.00 ± 0.2	1.7 ± 0.2	15.00	2.10 ± 0.2	1.6 ± 0.0	23.81

459 **V. CONCLUSION**

460 In this study, the inverse characterization of two cortical phantom plates covered with
461 a layer of soft tissue was conducted at low frequencies (between 50 and 500kHz). The
462 study focuses on the estimation of five parameters: longitudinal velocity, shear velocity, and
463 thickness of the cortical layer, as well as velocity and thickness of the soft tissue layer. The
464 proposed SAIGA model successfully aligned the amplitude in the experimental dispersion
465 curves by utilizing a combination of multiple high excitability mode segments.

466 The integration of modal excitability into the cost function played a pivotal role in the in-
467 version procedure, substantially reducing the number of relevant modes. This reduction was

468 achieved by assigning weights to the modes based on their excitabilities. The incorporation
469 of a limited number of modes prevents misfit of the data during the inversion process.

470 A high level of agreement was observed between the inversely determined properties of
471 cortical properties (bulk velocities and thickness) and their corresponding reference values
472 obtained through experimental measurements, with errors consistently below 2.0%. The
473 surrounding soft-tissue layer is critical during the inversion procedure due to the high num-
474 ber of modes involved. The model introduced in this study successfully extracted these
475 parameters even when a layer of soft tissue was present atop the cortical plate. Despite a
476 slightly larger error (inferior to 5%) in the determination of the soft tissue characteristics,
477 other properties remained accurately retrieved.

478 Additionally, the proposed low-frequency axial transmission configuration demonstrated
479 its efficacy in deducing equivalent bone properties within a multi-layer model. Remarkably,
480 this approach required fewer modes compared to conventional inversion techniques that
481 utilize higher frequencies. The current method serves as a promising proof of concept and
482 serves as a valuable reference for the development of a customized array probe, which could
483 be deployed for a comprehensive acquisition and reception in future extensive *in-vivo* and
484 clinical investigations. A portable device incorporating this specific probe could be utilized
485 in clinical settings for routine bone quality assessments, facilitating the early diagnosis of
486 osteoporosis at a reduced cost compared to DEXA or QCT-Scan technologies. However, a
487 significant challenge lies in developing a probe with a sufficiently wide bandwidth to capture
488 low-frequency guided waves propagating along the radius. Moreover, a dedicated design of
489 the setup will be required to ensure the repeatability of the location of the probe compared

490 to the radius. Furthermore, a clinical study will be necessary to compare the device's
491 results with reference values obtained through X-ray imaging. This comparison will help
492 establish a set of indicators representative of bone health, that can be used by practitioners
493 for diagnostic purposes.

494 The proposed method uses low-frequency ultrasonic guided waves to evaluate the bone
495 at the organ scale, while high frequency devices approximate bone by a plate-like structure.
496 The present study demonstrates the method's capability to accurately retrieve the properties
497 of a cortical phantom plate topped by a soft tissue layer, thus serving as a proof of concept.
498 Consequently, this model will evolve into a radius-shaped model, filled with bone marrow
499 and surrounded by soft tissue, following the work of Seyfaddini et al.⁴⁸. The computational
500 methodology using SAIGA for simulations will remain unchanged. To accurately represent
501 the bone cortex, the model will transition from an isotropic material to a transverse isotropic
502 medium^{18,46}. In such a model, it will be necessary to use a homogenization scheme to
503 determine the elastic stiffness coefficients as a function of the bone's porosity/density, as
504 described in the literature^{53,70,71}. Additionally, the inverse scheme could evolve to by using
505 AI techniques to enhance its robustness, which will assess bone properties based on the
506 dispersive trajectories and amplitudes of guided wave modes propagating in the waveguide.

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513 **AUTHOR DECLARATIONS**

514 **Conflict of Interest**

515 The authors have no conflicts to disclose.

516 **DATA AVAILABILITY**

517 The data that support the findings of this study are available from the corresponding
518 author upon reasonable request.

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