

Theory of ac electrokinetic behavior of spheroidal cell suspensions with an intrinsic dispersion

Lei Gao,^{1,2} J. P. Huang,¹ and K. W. Yu¹

¹*Department of Physics, The Chinese University of Hong Kong, Shatin, New Territories, Hong Kong, China*

²*Department of Physics, Suzhou University, Suzhou 215006, China*

(Received 26 August 2002; published 24 February 2003)

The dielectric dispersion, dielectrophoretic (DEP), and electrorotational (ER) spectra of spheroidal biological cell suspensions with an intrinsic dispersion in the constituent dielectric constants are investigated. By means of the spectral representation method, we express analytically the characteristic frequencies and dispersion strengths both for the effective dielectric constant and the Clausius-Mossotti factor (CMF). We identify four and six characteristic frequencies for the effective dielectric spectra and CMF, respectively, all of them being dependent on the depolarization factor (or the cell shape). The analytical results allow us to examine the effects of the cell shape, the dispersion strength, and the intrinsic frequency on the dielectric dispersion, DEP, and ER spectra. Furthermore, we include the local-field effects due to the mutual interactions between cells in a dense suspension, and study the dependence of co-field or antifield dispersion peaks on the volume fractions.

DOI: 10.1103/PhysRevE.67.021910

PACS number(s): 87.18.-h, 82.70.-y, 77.22.Gm, 77.84.Nh

I. INTRODUCTION

The polarization of biological cells has a wide range of practical applications such as manipulation, trapping, or separation of biological cells [1,2]. Dielectric spectroscopy [3], dielectrophoresis (DEP) [4], and electrorotation (ER) [5] offer a unique capability of monitoring the dielectric properties of dispersions of colloids and biological cells. Under the action of external fields, these particles exhibit rich fluid-dynamic behaviors as well as various dielectric responses. Hence, it is of importance to investigate their frequency-dependent responses to ac electric fields, which yields valuable information on the structural (Maxwell-Wagner) polarization effects. The polarization is characterized by a variety of characteristic frequency-dependent changes known as the dielectric dispersion, whose spectra are helpful to analyze the inhomogeneous systems, including biological cell suspensions and tissues [6]. Many factors exert influences on the effective dielectric behavior of the system such as orientation of dipoles, surface conductance, and the cell shape [7]. However, some factors can be dominant at certain ranges of frequencies (for example, the experimental data revealed that the low-frequency subdispersions were dependent on the cell shape [8]).

The DEP is used to describe the motion of the particles caused by the dielectric polarization effects in nonuniform electric field [9]. The DEP force drives the particles towards high intensity (positive DEP) or towards regions of minimal field intensity (negative DEP). Due to the interaction (i.e., DEP force) between the induced dipole and the external electric field, the particles can be levitated in the medium [10].

The ER behavior is caused by the existence of a phase difference between the field-induced dipole moment and the external rotating field, which results in a torque to cause the particle to rotate. In the dilute limit, the ER of individual cell can be predicted by ignoring the mutual interaction between the cells, and hence may be considered as an isolated particle in rotation [11].

In this paper, we will apply the spectral representation theory [12] to investigate the ac electrokinetic behavior including the dielectric dispersion, DEP, and ER spectra of

biological cell suspensions. The object of the present investigation is threefold. First, it is instructive to consider the effect of the intrinsic dielectric dispersion. Actually, such an intrinsic dielectric dispersion often occurs due to the surface conductivity [13,14] or the inhomogeneous structure such as the coated shells of biological cells or colloidal suspensions [15]. Second, many cells exist in the form of the nonspherical shape such as the fission yeast cell [8] and the red blood cell [16], and hence the effect of cell shape needs to be considered [17,18]. Third, the volume fractions of the biological cell suspensions are generally not in the dilute limit and can even exceed 0.1 [6,8,19], thus the local-field effect due to the average electrostatic interaction between cells must be taken into account [20]. In view of this, the effective dielectric dispersion spectra will be studied based on Maxwell-Garnett-type approximation [21], which involves an exact calculation of the field induced in the host medium by a single ellipsoidal cell and an approximate treatment of its distortion by the electrostatic interaction between different cell suspensions. Moreover, the Clausius-Mossotti factor (CMF), which determines the DEP and ER spectra, will be modified by replacing the dielectric constant of the host medium with the effective one.

Being beyond this work, we noticed an alternative paper [22], in which the authors considered the electrorotation and levitation of spherical cells or colloidal particles with or without the membrane-covered shells in the dilute limit. However, our investigation can be valid not only for nonspherical cell suspensions without membrane-covered shells, but also for nondilute limit suspensions. Furthermore, we adopt the spectral representation method [12], which offers the advantage of the separation of material parameters from the geometric information, to simplify the derivation of the analytical expressions for the characteristic frequencies and dispersion strengths of the effective dielectric permittivity and the CMF, respectively.

II. FORMALISM

We consider a composite system in which biological cells of dielectric constant $\bar{\epsilon}_1$ with the volume fraction p are dis-

persed in an isotropic host medium with dielectric constant $\tilde{\epsilon}_2 = \epsilon_2 + \sigma_2/(j2\pi f)$ with $j = \sqrt{-1}$, f being the frequency of the applied field. $\tilde{\epsilon}_1$ is assumed to exhibit an intrinsic dielectric dispersion, i.e.,

$$\tilde{\epsilon}_1 = \epsilon_1 + \frac{\Delta\epsilon_1}{1 + jf/f_1} + \frac{\sigma_1}{j2\pi f}, \quad (1)$$

where ϵ_1 (σ_1) is the limiting high-frequency (low-frequency) dielectric constant (conductivity), $\Delta\epsilon_1$ represents the dielectric dispersion strength with a characteristic frequency f_1 . Biological cells are further assumed to be spheroidal in shape. Such an assumption is supported by Bohren and Huffman [23] who suggested that dielectric dispersion spectra of particles of arbitrary shape can be approximated by these of spheroidal particles.

Then we will adopt the spectral representation to investigate the ac electrokinetic behavior such as the dielectric dispersion, dielectrophoretic, and electrorotational spectra of an inhomogeneous system, in which spheroidal biological cells are randomly distributed.

A. Dielectric dispersion spectrum

Generally, for cell suspensions of arbitrary shape, the spectral representation can only be solved numerically. However, for randomly oriented spheroidal cell suspensions, within the mean-field theory, the effective complex dielectric constant $\tilde{\epsilon}_e$ can be written as [21]

$$\tilde{\epsilon}_e = \tilde{\epsilon}_2 \frac{1 + p[b_z(1 - L_z) + 2b_{xy}(1 - L_{xy})]}{1 - p(b_z L_z + 2b_{xy} L_{xy})}, \quad (2)$$

where $b_k \equiv (\tilde{\epsilon}_1 - \tilde{\epsilon}_2)/(3[\tilde{\epsilon}_2 + L_k(\tilde{\epsilon}_1 - \tilde{\epsilon}_2)])$ ($k = z, xy$) is called the CMF, and L_z [$L_{xy} = (1 - L_z)/2$] are the depolarization factors along the z (x or y) axis of the spheroids. These depolarization factors depend on the aspect ratio $q \equiv c/a$ [21], where $a (=b)$ and c are the semiaxes of a spheroid along the Cartesian axes. For the prolate-spheroidal cell ($q > 1$), we have $0 < L_z < 1/3$; while for the oblate-spheroidal one ($q < 1$), we have $1/3 < L_z < 1$. Actually, once a q is given, L_z can be obtained uniquely and thus used to indicate the shape of spheroidal particles.

By invoking the spectral representation and introducing the dimensionless parameter $\tilde{s} \equiv \tilde{\epsilon}_2/(\tilde{\epsilon}_2 - \tilde{\epsilon}_1)$, we rewrite Eq. (2) as

$$\tilde{\epsilon}_e = \tilde{\epsilon}_2 \left(1 - \frac{W_1}{s - x_1} - \frac{W_2}{s - x_2} \right), \quad (3)$$

where the poles x_1 and x_2 are given as

$$x_{1,2} = \frac{1}{12} [3 - 2p + 3L_z \pm \sqrt{(3 - 2p + 3L_z)^2 - 72(1 - p)L_z(1 - L_z)}]. \quad (4)$$

Correspondingly, the residues W_1 and W_2 have

$$W_1 = \frac{p(1 + 3L_z - 6x_1)}{6(x_2 - x_1)}, \quad W_2 = \frac{p(1 + 3L_z - 6x_2)}{6(x_1 - x_2)}. \quad (5)$$

It is easy to check that W_1 and W_2 satisfy the sum rule $W_1 + W_2 = p$.

In order to obtain the analytic expressions for the dielectric dispersion strengths and the characteristic frequencies of the dielectric dispersion spectra, we then introduce two contrast parameters $s = \epsilon_2/(\epsilon_2 - \epsilon_1)$ and $t = \sigma_2/(\sigma_2 - \sigma_1)$, defined by Lei *et al.* [7], and derive an equality for biological cells possessing an intrinsic dispersion

$$\frac{1}{s - x} = \frac{1}{s - x} + \frac{A(x)}{1 + jf/f_{c1}(x)} + \frac{B(x)}{1 + jf/f_{c2}(x)}, \quad (6)$$

where

$$f_{c1,c2}(x) = \frac{f_1 f_2 + f_2 f_3 + f_3 f_1 \mp \sqrt{(f_1 f_2 + f_2 f_3 + f_3 f_1)^2 - 4f_1 f_2 f_3^2}}{2f_3}, \quad (7)$$

and

$$A(x) = \left[\frac{s - t}{(s - x)(t - x)} \left(\frac{f_1 - f_{c1}}{f_1 f_{c1}} \right) + \frac{s}{(s - x)x} \frac{1}{f_3} \right] \frac{f_1 f_2}{f_{c2}(x) - f_{c1}(x)}, \quad (8)$$

$$B(x) = \left[\frac{s - t}{(s - x)(t - x)} \left(\frac{f_2 - f_{c2}}{f_1 f_{c2}} \right) - \frac{s}{(s - x)x} \frac{1}{f_3} \right] \frac{f_1 f_2}{f_{c2}(x) - f_{c1}(x)}, \quad (9)$$

with

$$f_2(x) = \frac{1}{2\pi} \frac{s\sigma_2(t - x)}{\epsilon_2 t(s - x)}, \quad f_3(x) = \frac{1}{2\pi} \frac{\sigma_2(t - x)}{x\Delta\epsilon_1 t}. \quad (10)$$

After simple manipulations, we can express Eq. (3) as

$$\tilde{\epsilon}_e = \epsilon_H + \sum_{i=1}^4 \frac{\Delta\epsilon_{ei}}{1 + j(f/f_{ei})} + \frac{\sigma_L}{j2\pi f} \equiv \epsilon_e + \frac{\sigma_e}{j2\pi f}, \quad (11)$$

with

$$\epsilon_H = \epsilon_2 \left(1 - \frac{W_1}{s - x_1} - \frac{W_2}{s - x_2} \right),$$

$$\sigma_L = \sigma_2 \left(1 - \frac{W_1}{t - x_1} - \frac{W_2}{t - x_2} \right), \quad (12)$$

and

$$\Delta\epsilon_{e1} = W_1 A(x_1) \left(\frac{\sigma_2}{2\pi f_{c1}(x_1)} - \epsilon_2 \right), \quad f_{e1} = f_{c1}(x_1), \quad (13)$$

$$\Delta \epsilon_{e2} = W_1 B(x_1) \left(\frac{\sigma_2}{2\pi f_{c2}(x_1)} - \epsilon_2 \right), \quad f_{e2} = f_{c2}(x_1). \quad (14)$$

Similarly, $\Delta \epsilon_{ei}$ and f_{ei} for $i=3,4$ can be easily obtained by our replacing W_1 and x_1 with W_2 and x_2 from the above two equations.

Thus, within the spectral representation, for spheroidal suspensions with one intrinsic dispersion, we successfully obtain four characteristic frequencies f_{ei} ($i=1,4$) with the dispersion strengths $\Delta \epsilon_{ei}$ in terms of the geometric parameters (L_z and p) as well as physical parameters (s , t , $\Delta \epsilon_1$, σ_2 , and ϵ_2).

In Fig. 1, we numerically calculate four characteristic frequencies and dispersion strengths against L_z . As is evident from the figure, the characteristic frequencies and the corresponding dispersion strengths are strongly dependent on the shape of cell suspensions and exhibit nonmonotonical behavior with increasing L_z . For these four characteristic frequencies, two of them are located at frequencies higher than 10^6 Hz, while another two are at frequencies lower than 10^6 Hz. Such a separation results from the fact that we take into account one intrinsic dispersion in the dielectric cell suspensions. Furthermore, due to the nonspherical shape of cell suspensions, there are two characteristic frequencies in the high-frequency (or low-frequency) region. When the shape is far from spherical (say $L_z \rightarrow 0$ or 1), the difference between these two frequencies becomes large. Conversely, for spherical inclusions ($L_z = 1/3$), only two dispersion strengths are nonzero, leading to two main steps in the dielectric dispersion spectra, as will be shown in the following section.

B. Dielectrophoretic and electrorotational spectra

The spectral form of Eq. (6) also allows us to investigate the dielectrophoretic and electrorotational behavior.

It is known that the time average DEP force $F(f)$ and the ER torque $\Gamma(f)$ exerted on the spheroidal cell suspensions with z orientation parallel to the electric field are expressed as [24,25]

$$F(f) = 1.5v \tilde{\epsilon}_2 \operatorname{Re}[b_z(\tilde{\epsilon}_1, \tilde{\epsilon}_2)] |\nabla E_{(rms)}^2|, \\ \Gamma(f) = -3v \tilde{\epsilon}_2 \operatorname{Im}[b_z(\tilde{\epsilon}_1, \tilde{\epsilon}_2)] E_0^2, \quad (15)$$

where v is the volume of spheroidal cell, $E_{(rms)}$ is the root-mean-square magnitude of the imposed ac electric field and $\operatorname{Re}[\dots]$, $\operatorname{Im}[\dots]$ represent the real and imaginary parts of b_z , which can be written under the spectral representation as

$$b_z = \frac{2(p-1)\tilde{s}^2 + (1-p)(1+L_z)\tilde{s} - (1-p)L_z(1-L_z)}{6\tilde{s}^3 - (3+4p+9L_z-6pL_z)\tilde{s}^2 + (p+6L_z-3pL_z)\tilde{s} + 3(p-1)L_z^2(1-L_z)} \\ = \frac{V_1}{\tilde{s}-s_1} + \frac{V_2}{\tilde{s}-s_2} + \frac{V_3}{\tilde{s}-s_3}, \quad (18)$$

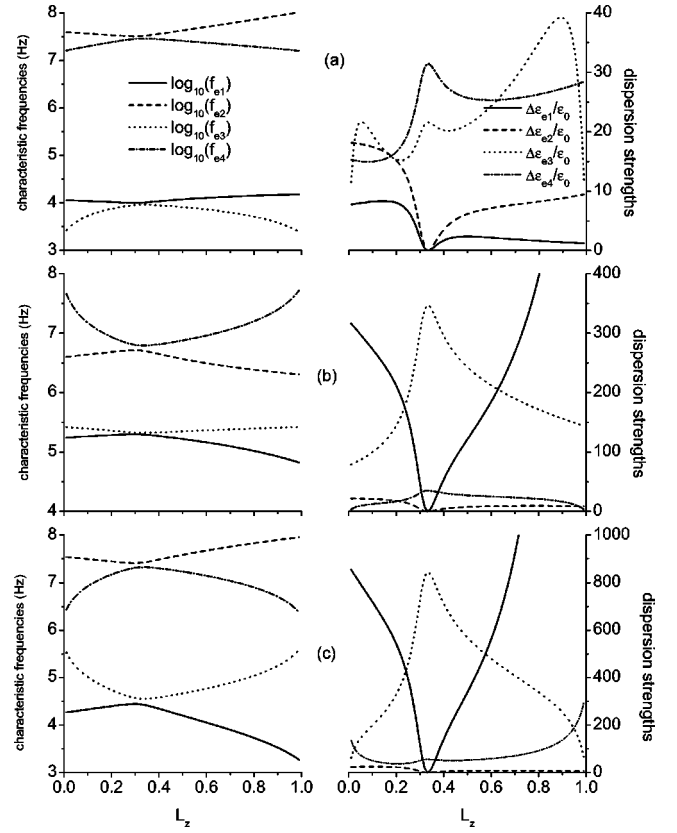


FIG. 1. The characteristic frequencies and the dispersion strengths against the depolarization factor L_z for $p=0.2$ and (a) $s = 1.625$, $t = -0.053$, $\epsilon_2 = 78\epsilon_0$, $\sigma_2 = 10^{-5}$ S/m, $\Delta \epsilon_1 = 200\epsilon_0$, and $f_1 = 1.59 \times 10^7$ Hz; (b) $s = -0.031$, $t = 1.25$, $\epsilon_2 = 6\epsilon_0$, $\sigma_2 = 0.02$ S/m, $\Delta \epsilon_1 = 1000\epsilon_0$, and $f_1 = 2.65 \times 10^5$ Hz; (c) $s = 2.78$, $t = 1.053$, $\epsilon_2 = 78\epsilon_0$, $\sigma_2 = 2 \times 10^{-3}$ S/m, $\Delta \epsilon_1 = 2500\epsilon_0$, and $f_1 = 1.9 \times 10^6$ Hz.

$$b_z(\tilde{\epsilon}_1, \tilde{\epsilon}_2) = \frac{1}{3} \frac{\tilde{\epsilon}_1 - \tilde{\epsilon}_2}{\tilde{\epsilon}_2 + L_z(\tilde{\epsilon}_1 - \tilde{\epsilon}_2)} = -\frac{1}{3} \frac{1}{\tilde{s} - L_z}. \quad (16)$$

Equation (16) is independent of the volume fraction p and thus is valid only in the dilute limit [22]. However, for non-dilute volume fractions, we must consider the local-field effect due to mutual interaction between the spheroidal cells and modify Eq. (16) as

$$b_z(\tilde{\epsilon}_1, \tilde{\epsilon}_2) = \frac{1}{3} \frac{\tilde{\epsilon}_1 - \tilde{\epsilon}_e}{\tilde{\epsilon}_e + L_z(\tilde{\epsilon}_1 - \tilde{\epsilon}_e)}. \quad (17)$$

Substituting Eq. (2) into Eq. (17), we have

where s_i ($i=1,2,3$) are three roots of polynomial equations

$$y^3 - \frac{3+4p+9L_z-6pL_z}{6}y^2 + \frac{p+6L_z-3pL_z}{6}y + \frac{(p-1)L_z^2(1-L_z)}{2} = 0, \quad (19)$$

while the residues V_1 , V_2 , and V_3 can be obtained from

$$V_1 + V_2 + V_3 = \frac{1}{3}(p-1),$$

$$V_1(s_2 + s_3) + V_2(s_3 + s_1) + V_3(s_1 + s_2) = \frac{1}{6}(p-1)(1+L_z),$$

$$V_1s_2s_3 + V_2s_1s_3 + V_3s_1s_2 = \frac{1}{6}(p-1)L_z(1-L_z). \quad (20)$$

Note that these residues again satisfy the sum rule $V_1 + V_2 + V_3 = -(1-p)/3$.

Introducing Eq. (6) into Eq. (18) leads to

$$b_z = \sum_{i=1}^3 \left[\frac{V_i}{s-s_i} + \frac{V_i A(s_i)}{1+jf/f_{c1}(s_i)} + \frac{V_i B(s_i)}{1+jf/f_{c2}(s_i)} \right]. \quad (21)$$

The normalized DEP force (f_N) and the normalized ER torque (Γ_N) can be defined as

$$\begin{aligned} f_N &= \text{Re}[b_z(\tilde{\epsilon}_1, \tilde{\epsilon}_2)] \\ &= \sum_{i=1}^3 \left[\frac{V_i}{s-s_i} + \frac{V_i A(s_i)}{1+[f/f_{c1}(s_i)]^2} + \frac{V_i B(s_i)}{1+[f/f_{c2}(s_i)]^2} \right], \end{aligned} \quad (22)$$

and

$$\begin{aligned} \Gamma_N &= -\text{Im}[b_z(\tilde{\epsilon}_1, \tilde{\epsilon}_2)] \\ &= \sum_{i=1}^3 \left[\frac{V_i A(s_i) f/f_{c1}(s_i)}{1+[f/f_{c1}(s_i)]^2} + \frac{V_i B(s_i) f/f_{c2}(s_i)}{1+[f/f_{c2}(s_i)]^2} \right]. \end{aligned} \quad (23)$$

Based upon the spectral representation, the expressions for the dispersion strengths and characteristic frequencies of the CMF are also explicitly obtained. Six CMF characteristic frequencies and dispersion strengths against L_z are shown in Fig. 2.

Again, L_z plays an important role in determining the CMF characteristic frequencies and the corresponding strengths. Within our model, six CMF characteristic frequencies are predicted; three of them are located below 10^6 Hz and another three are above 10^6 Hz, similar as that observed in Fig. 1. Due to this, the DEP (or ER) spectra are mainly characterized by two-step rapid changes (or two dominant peaks). For three typical cases, we also find that six CMF strengths are all positive in Fig. 2(a); all negative in Fig. 2(b), and three strengths are negative and three are positive in Fig. 2(c). Such behavior will result in co-field or antifield rotation of the particle for the ER spectra. We remark that as both the intrinsic dispersion and local-field effects are considered

simultaneously, we predict four characteristic frequencies, characterized by four nonzero dispersion strengths even for $L_z = 1/3$.

III. NUMERICAL RESULTS

We are now in a position to calculate the dielectric dispersion spectrum, DEP and ER spectra based on the model put forward in the preceding section.

In Figs. 3–5, we investigate the effect of depolarization factor L_z on the dielectric response [denoted by ϵ_e/ϵ_0 and σ_e defined in Eq. (11)], and the DEP and ER spectra using Eqs. (22) and (23).

The dielectric response mainly exhibits two subdispersions characterized by two-step rapid decrease in Figs. 3(a), 4(a), and 5(a). It is evident that the influence of L_z (the shape) on the effective dielectric constant (or the effective conductivity) is significant in the low-frequency region, where $f < 10^4$ Hz (or in the high-frequency region where $f > 10^8$ Hz). This is in agreement with previous experimental conclusions that the low-frequency subdispersion for ϵ_e depends on the cell shape, whereas the high-frequency is independent of it [8]. The dielectric permittivity for oblate spheroidal particles, say $L_z = 0.95$, in the low-frequency region can be four times of the one for spherical inclusions. Thus,

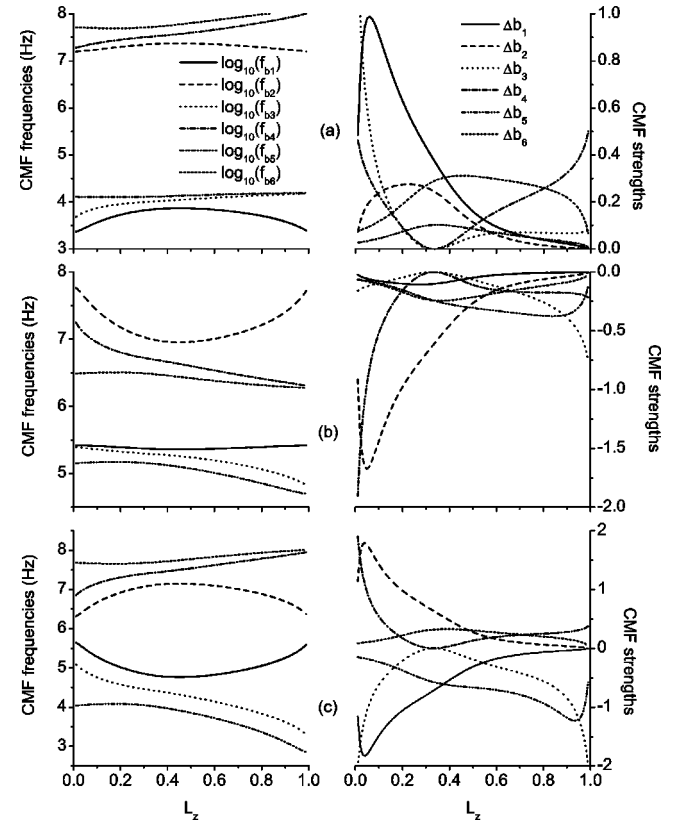


FIG. 2. The characteristic frequencies $f_{b1}=f_{c1}(s_1)$, $f_{b2}=f_{c2}(s_1)$, $f_{b3}=f_{c1}(s_2)$, $f_{b4}=f_{c2}(s_2)$, $f_{b5}=f_{c1}(s_3)$, $f_{b6}=f_{c2}(s_3)$, and the dispersion strengths $\Delta b_1=V_1A(s_1)$, $\Delta b_2=V_1B(s_1)$, $\Delta b_3=V_2A(s_2)$, $\Delta b_4=V_2B(s_2)$, $\Delta b_5=V_3A(s_3)$, and $\Delta b_6=V_3B(s_3)$ for the CMF (b_z) versus L_z . The parameters are the same as those in Fig. 1.

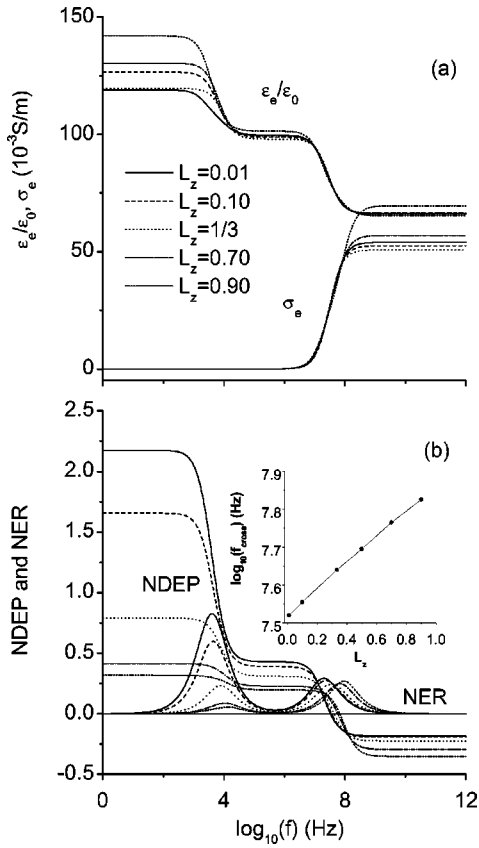


FIG. 3. Effective dielectric response (ϵ_e/ϵ_0 and σ_e) (a) and normalized dielectrophoresis (NDEP) and electrorotation (NER) (b) versus the frequency f for the parameters in Fig. 1(a). In the inset of (b), the crossover frequency f_{cross} is plotted against L_z .

by taking into account the shape of cell suspensions, it is possible to reduce some discrepancies between the theoretical and experimental results on the dielectric dispersion spectra [8,19].

For the DEP force, it exhibits strong sensitivity to the depolarization factor (the shape). When the DEP force is positive (the particles will be attracted towards the field-generating electrodes), with increasing L_z , such an attractive force becomes weak. However, when the DEP force is negative (the particles will be repelled from the electrodes), the larger L_z is, the stronger repulsion will be. Furthermore, we predict one or two shape-dependent crossover frequencies at which there is no net force on the cell particle. The crossover frequency is a monotonically increasing or decreasing function of L_z , dependent on whether the variation of the DEP force around it is negative or positive [see the insets in Figs. 3(b), 4(b), and 5(a)]. This is an interesting result. To the best of our knowledge, the dependence of crossover frequency on the spheroidal shape is reported here for the first time.

For the ER spectra, they exhibit two ER characteristic frequencies with both positive strengths in Fig. 3(b), both negative strengths in Fig. 4(b), and negative and positive strengths in Fig. 5(b) (This has been observed in experimental electrorotational spectra of the yeast cell [26,27]). Positive ER torque means the co-field rotation of the particle. In this case, with increasing L_z , the ER characteristic frequency

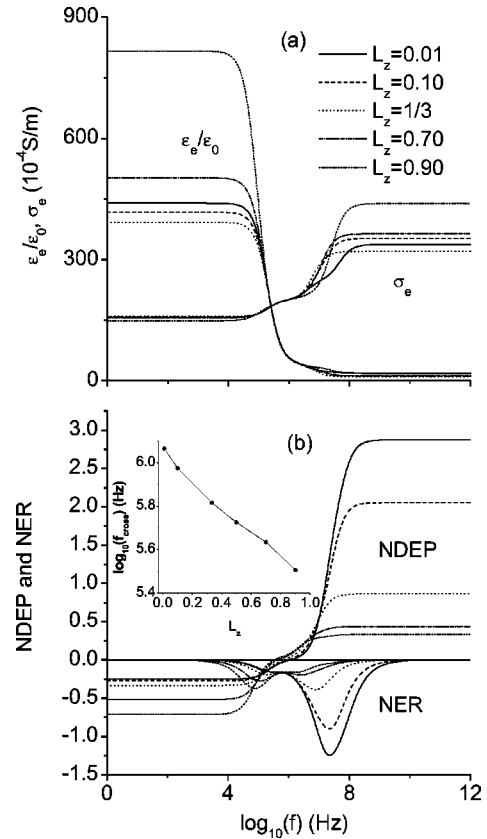


FIG. 4. Same as Fig. 3, but for the parameters in Fig. 1(b).

for the ER will be blue shifted accompanied with the decrease of the dispersion strengths. However, when the ER torque is negative (the particle exhibits the antifield rotation), both the ER characteristic frequencies and the strengths decrease concomitantly with L_z . Moreover, we find that the variation of the geometric parameter L_z does not change the polarity of the ER peak.

We emphasize to point out that, although we analytically predict four and six characteristic frequencies for the dielectric dispersion and ER spectra, these frequencies are mainly located in two regions or some of their dispersion strengths are quite small in comparison with others, resulting in only two effective characteristic frequencies at which the rate of change of the dielectric permittivity (or the DEP force), and the rotational torque peak attain their maxima. This can be well understood from Fig. 6.

As we have included the dielectric dispersion form in $\tilde{\epsilon}_1$, it would be interesting to investigate how the dispersion strength $\Delta\epsilon_1$ and the intrinsic frequency f_1 affect the dielectric dispersion and ER spectra.

In Fig. 7, we examine the effect of $\Delta\epsilon_1$ on the dielectric response (the left panel) and ER spectra (the right panel). Clearly, one common feature of all these dielectric spectra is that increasing $\Delta\epsilon_1$ yields increasing dielectric permittivity in the low-frequency region. As previous models do neglect the intrinsic dispersion effect, the theoretical results in the low-frequency region are less than experimental reports [8,19]. We think it would be helpful to give the closest fit with experimental data by suitable adjustment of both $\Delta\epsilon_1$

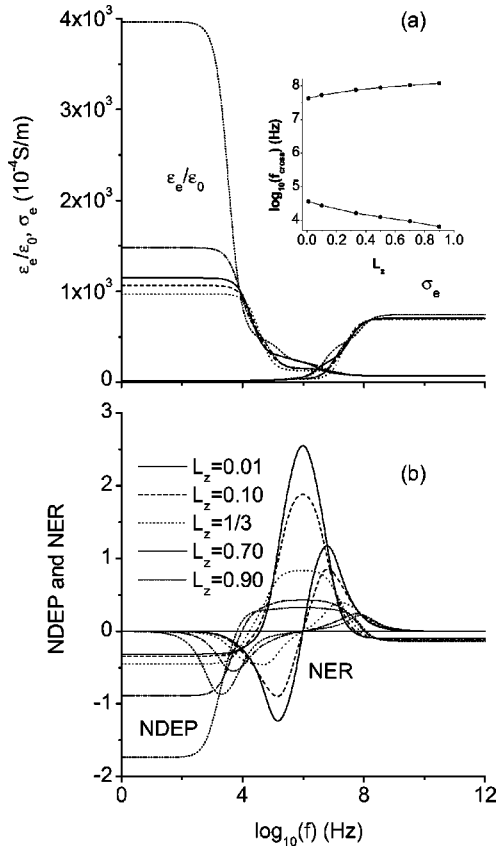


FIG. 5. Same as Fig. 3, but for the parameters in Fig. 1(c). Note that two shape-dependent crossover frequencies are predicted.

and L_z . For the ER spectra, when $\Delta\epsilon_1 = 0$, only one dominant peak is predicted as expected. For a finite $\Delta\epsilon_1$, two dominant peaks at low and high characteristic frequencies arise. With increasing $\Delta\epsilon_1$, the characteristic low frequency shifts towards a long wavelength; while the characteristic high frequency shifts towards a short one, leading to further separation between two characteristic frequencies for larger $\Delta\epsilon_1$. At the same time, the adjustment of $\Delta\epsilon_1$ can also change the peak value much or less and even dominate the co-field or antifield rotation through changing the polarity of the peak (see the upper right column).

In Fig. 8, we plot the dielectric dispersion and ER spectra versus the frequency f for different f_1 . It is evident that f_1 mainly play a role in the characteristic high frequency for both the effective dielectric dispersion and ER spectra. Generally speaking, the characteristic high frequency increases with increasing f_1 , in accord with the analytical formula from Eq. (7). Interestingly, the increase in f_1 can also lead to the change of the polarity of the ER peak in the high-frequency region (see the middle right panel).

In order to take into account the mutual interaction between the suspended particles, we adopt Eq. (17) instead of Eq. (16). When the mutual interaction is taken into account, the ER peak must be reduced, as expected in Fig. 9. With further increasing p , the mutual interaction becomes strong, leading to the serious depression of the ER peak and the

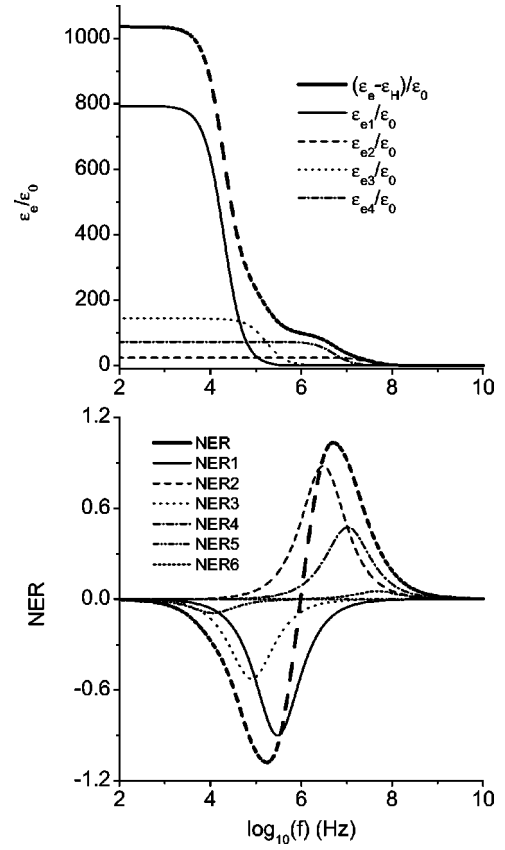


FIG. 6. Contributions of dielectric response $\epsilon_{ei}/\epsilon_0 \equiv \Delta\epsilon_{ei}/[1 + (f/f_{ei})^2]$ ($i=1,4$) to the dielectric spectra $(\epsilon_e - \epsilon_H)/\epsilon_0$ (a), and contributions of i th strengths $\Gamma_{Ni} \equiv \Delta b_i f f_{bi}/(f_{bi}^2 + f^2)$ to Γ_N (b) versus f for $L_z = 0.05$. Other parameters are the same as those in Fig. 1(c). Note that the effective dielectric spectra and the NER spectra mainly exhibit two-step rapid decrease and two dominant peaks, respectively, although four and six characteristic frequencies are analytically predicted.

slight blue shift (or redshift) in the characteristic low frequency (or high frequency).

IV. DISCUSSION AND CONCLUSION

In this work, we have investigated both analytically and numerically the ac electrokinetic behavior, i.e., the dielectric dispersion behavior, and the DEP the ER spectra of non-spherical cell suspensions with an intrinsic dispersion based on the spectral representation method. The dependence of the ac electrokinetic behavior on the depolarization factor is studied in detail. We find that an intrinsic dispersion in the dielectric constant of spheroidal cells can lead to four and six dispersions in the effective dielectric constant and the CMF, respectively. We also find that both the intrinsic dispersion strength $\Delta\epsilon_1$ and the characteristic frequency f_1 can change the polarity of the ER torque and thereby causing the co-field or antifield rotation. Furthermore, by taking into account the local-field effect from the mutual interaction, we examine the ER spectra for various volume fractions, and find that increasing volume fractions can result in the decrease of both the strengths and the difference between two characteristic

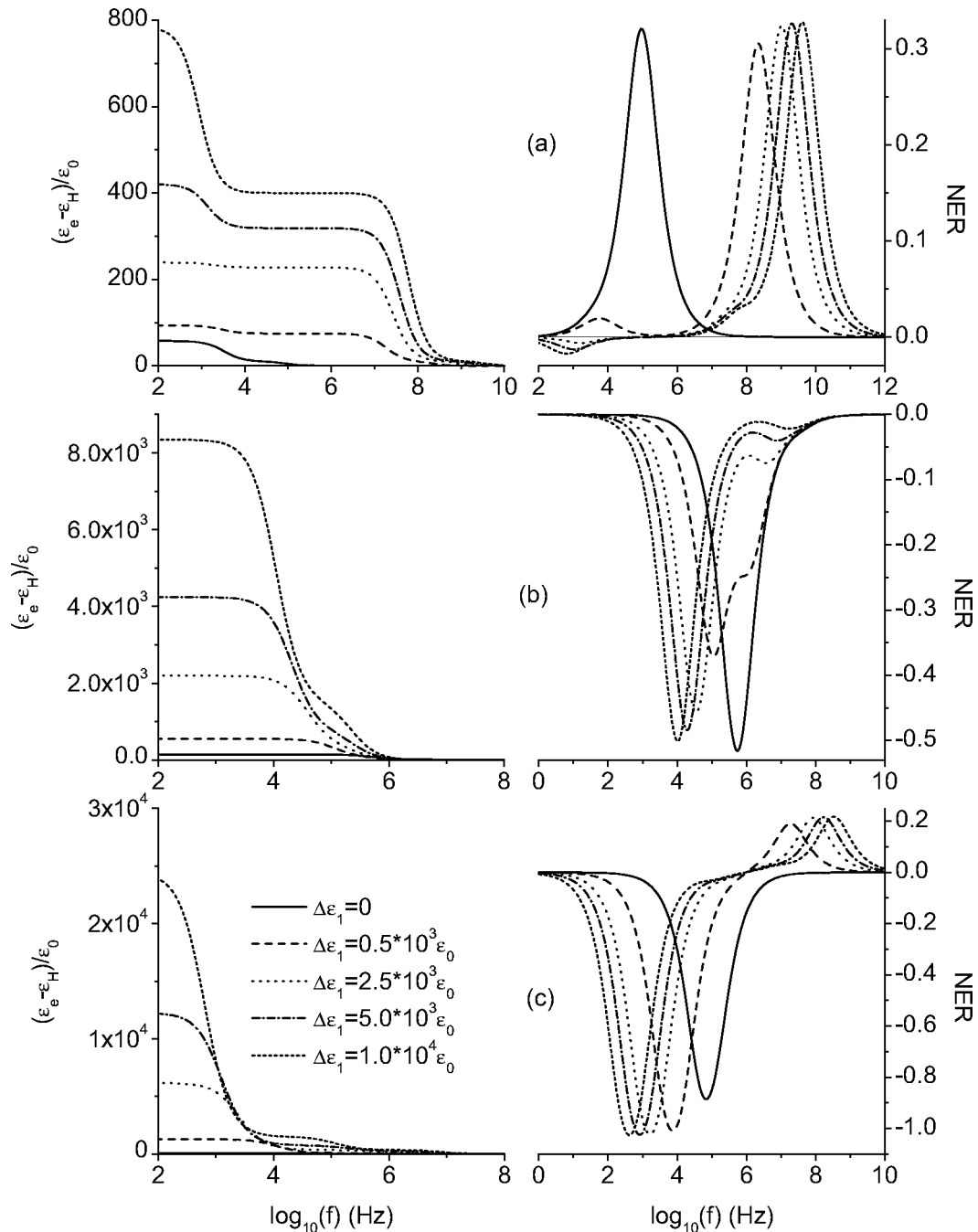


FIG. 7. $(\epsilon_e - \epsilon_H)/\epsilon_0$ and Γ_N versus f for $L_z = 0.95$ and various $\Delta\epsilon_1$. Other parameters are the same as those in Fig. 1.

low and high frequencies indeed. Thus, it is possible to obtain a good agreement between theoretical predictions and experimental data by the suitable adjustment of both the geometric parameters (for example, the particle shape) and the physical parameters (for example, the dispersion strength and the characteristic frequency of cell suspensions). On the contrary, such a fitting is very useful to obtain the relevant physical information of cell suspensions.

Here, a few comments are in order. We obtain four and six characteristic frequencies for the effective dielectric permittivity and CMF. Generally, none of the characteristic frequencies for the dielectric spectra is equal to those of CMF,

as the poles x_1 and x_2 [Eq. (4)] are quite different from the poles s_1 , s_2 , and s_3 of CMF. In the dilute limit and for spherical inclusions, both Eqs. (3) and (17) yield two characteristic frequencies; the smaller p is, the lesser are the differences between these frequencies [15]. Numerically, the dielectric dispersion, DEP, and ER spectra mainly exhibit two rapid changes or two dominant peaks. Here, we also show a four-step rapid decrease in dielectric dispersion spectra (see Fig. 10). However, due to small differences between the CMF frequencies or small strengths of some frequencies in comparison with others, it is difficult to show all six characteristic frequencies in the ER spectra clearly. A possible

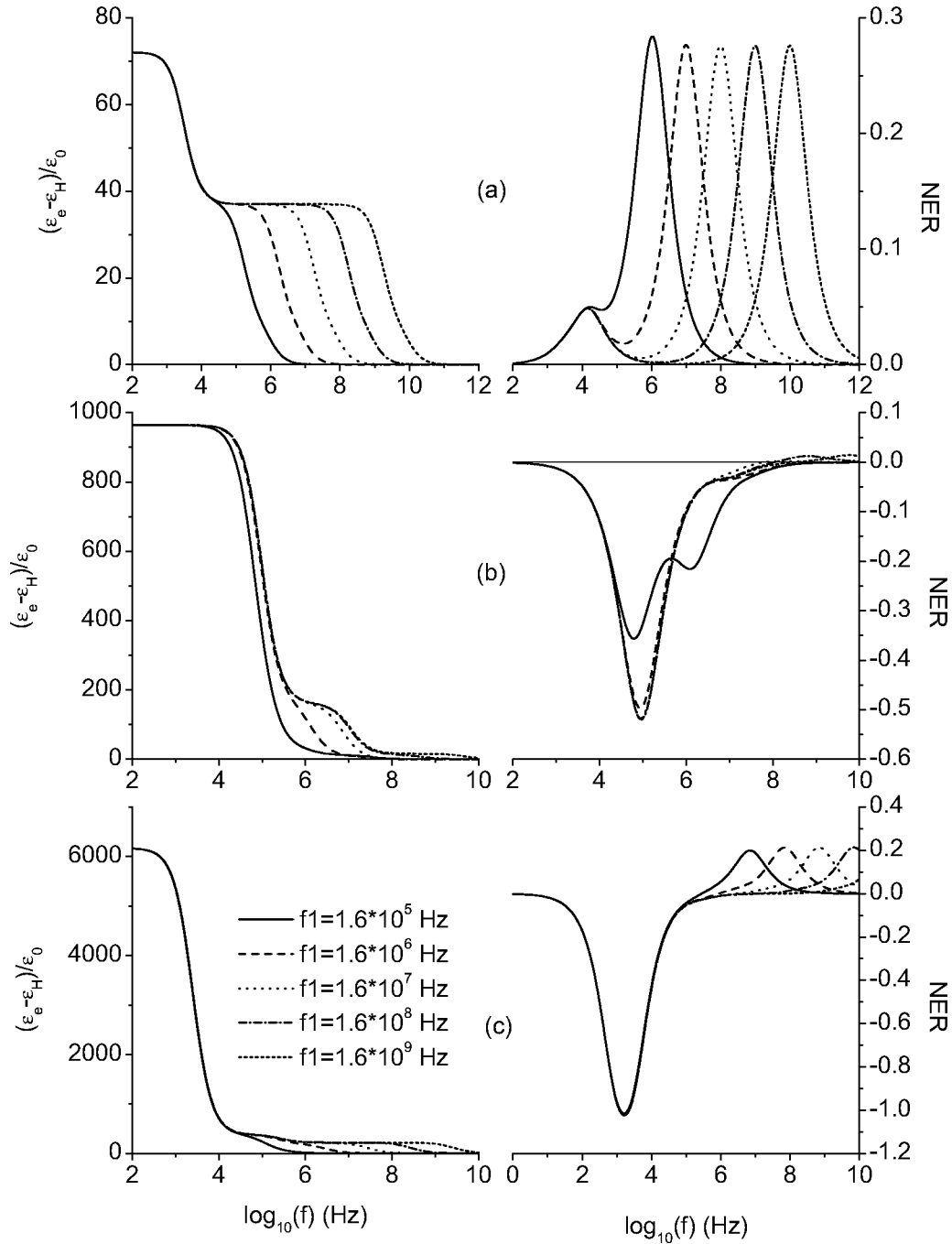


FIG. 8. $(\epsilon_e - \epsilon_H)/\epsilon_0$ and Γ_N versus f for $L_z=0.95$ and various f_1 . Other parameters are the same as those in Fig. 1.

way to achieve this is to consider cell suspensions with large volume fractions while being far from the spherical shape.

We demonstrate theoretically that the shape effect of the effective dielectric constant is significant in the low-frequency region, in accord with experimental conclusions [8]. In previous work, one of the authors have presented a first-principles study of the dielectric dispersion of fission yeast cell suspensions [7]. As the derivation is based on the assumption that cell suspensions are in the dilute limit and do not exhibit an intrinsic dispersion, the predicted two characteristic frequencies are independent of the volume fractions and some discrepancies between the theory and experi-

ment on fission yeast cells still exist. We believe that our present formula can be applied also and the discrepancies between theoretical and experimental results may further be reduced.

For shelled spheroidal particles dispersed in the host medium, we have found one peak in the electrorotation assay, under the assumption that the ratio of shell to the host dielectric constant is real [11]. The theory yields a good agreement with the experimental results only in the high-frequency region. In fact, for such kind of three-component system, the effective dielectric response ϵ_{cs} for the coated shells can be first obtained in a self-consistent way [21] and will exhibit at

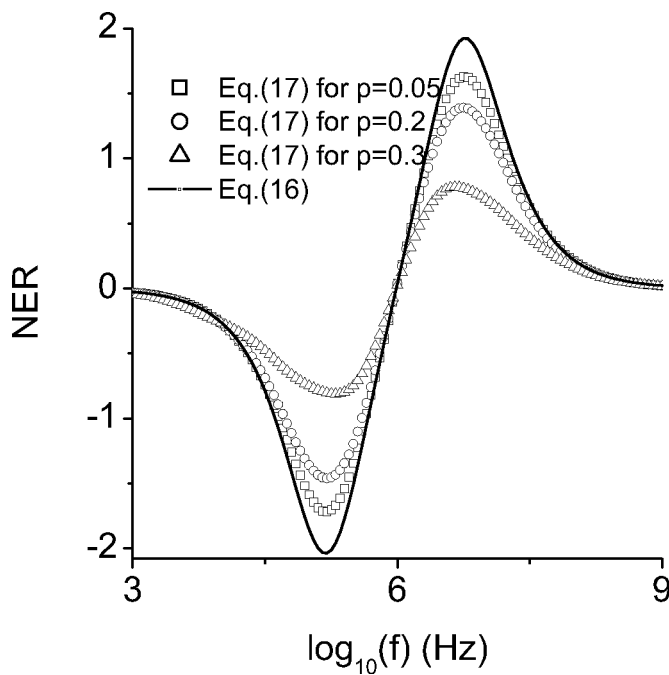


FIG. 9. Γ_N versus f with Eqs. (16) and (17) for $L_z=0.05$. Other parameters are the same as those in Fig. 5. Note that Eq. (16) is independent of p .

least one dispersion. The three-component composites can then be equivalent to the solid spheroidal particles with ϵ_{cs} including one dispersion term suspended in the host medium. Thus, our present model can be safely used, and the co-field rotational peak in the low-frequency region may also be predicted. Finally, we should remark that, for nonspherical cells, the intrinsic dispersion may depend on the orientation of a spheroid, which will yield dielectric anisotropy in cell suspensions. In this work, we take into account the geometric anisotropy as well as the dielectric isotropy. The work including both geometric and dielectric anisotropies is in progress and will be reported elsewhere.

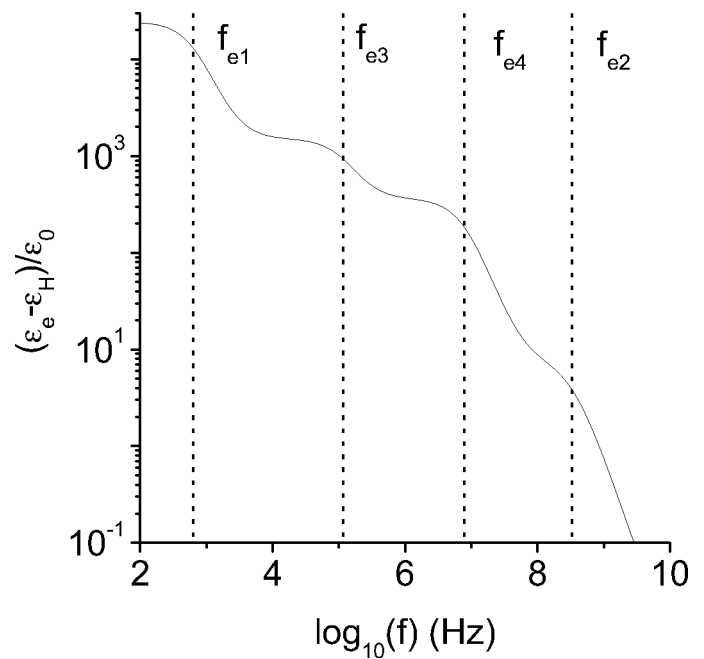


FIG. 10. $(\epsilon_e - \epsilon_H)/\epsilon_0$ and Γ_N versus f for $p=0.2$, $L_z=0.95$, and $\Delta\epsilon_1=1 \times 10^4\epsilon_0$. Other parameters are the same as those in Fig. 1(c). There are four characteristic frequencies, at which the dielectric spectra exhibit rapid decrease.

ACKNOWLEDGMENTS

This work was supported by the RGC Earmarked Grant under Project No. CUHK 4245/01P. L.G. acknowledges the financial support of the National Natural Science Foundation of China under Grant No. 10204017 and of the Science Foundation of Jiangsu Province under Grant No. BK2002038. K.W.Y. thanks Professor A.R. Day for his interest in our work and his suggestion to examine the effects of inhomogeneities on the dielectric response of biological cell suspensions.

-
- [1] For a review, see J. Gimsa, and D. Wachner, *Biophys. J.* **77**, 1316 (1999).
- [2] J. Gimsa, *Ann. N.Y. Acad. Sci.* **873**, 287 (1999).
- [3] K. Asami, T. Hanai, and N. Koizumi, *Jpn. J. Appl. Phys.* **19**, 359 (1980).
- [4] G. Fuhr, J. Gimsa, and R. Glaser, *Stud. Biophys.* **108**, 149 (1985).
- [5] J. Gimsa, P. Marszalek, U. Loewe, and T.Y. Tsong, *Biophys. J.* **60**, 749 (1991).
- [6] K. Asami, *J. Non-Cryst. Solids* **305**, 268 (2002).
- [7] J. Lei, Jones T.K. Wan, K.W. Yu, and H. Sun, *Phys. Rev. E* **64**, 012903 (2001).
- [8] K. Asami, *Biochim. Biophys. Acta* **1472**, 137 (1999).
- [9] H. A. Pohl, *Dielectrophoresis* (Cambridge University Press, Cambridge, 1978).
- [10] T.N. Tombs and T.B. Jones, *Rev. Sci. Instrum.* **62**, 1072 (1991).
- [11] J.P. Huang and K.W. Yu, *J. Phys.: Condens. Matter* **14**, 1213 (2002).
- [12] D.J. Bergman, *Phys. Rep.* **43**, 379 (1978).
- [13] G. Schwarz, *J. Phys. Chem.* **66**, 2636 (1962).
- [14] L. Gao, Jones T.K. Wan, K.W. Yu, and Z.Y. Li, *J. Appl. Phys.* **88**, 1893 (2000).
- [15] X.B. Wang, Y. Huang, R. Holzel, J.P.H. Burt, and R. Pethig, *J. Phys. D* **26**, 312 (1993).
- [16] R.D. Miller and T.B. Jones, *Biophys. J.* **64**, 1588 (1993).
- [17] R. Paul and M. Otwinowski, *J. Theor. Biol.* **148**, 495 (1991).
- [18] R. Paul and K.V.I.S. Kaler, *Phys. Rev. E* **48**, 1491 (1993).
- [19] F. Bordi, C. Cametti, and T. Gili, *J. Non-Cryst. Solids* **305**, 278 (2002).
- [20] J.P. Huang, K.W. Yu, and G.Q. Gu, *Phys. Rev. E* **65**, 021401 (2002).

- [21] L. Gao, Jones T.K. Wan, K.W. Yu, and Z.Y. Li, *J. Phys.: Condens. Matter* **12**, 6825 (2000).
- [22] K.R. Foster, F.A. Sauer, and H.P. Schwan, *Biophys. J.* **63**, 180 (1992).
- [23] C.F. Bohren and D.R. Huffman, *Absorption of Scattering of Light by Small Particles* (Wiley, New York, 1983).
- [24] J.M. Cruz and F.J. Garcia-Diego, *J. Phys. D* **31**, 1745 (1998).
- [25] X.B. Wang, R. Pethig, and T.B. Jones, *J. Phys. D* **25**, 905 (1992).
- [26] X.F. Zhou, G.H. Markx, R. Pethig, *Biochim. Biophys. Acta* **1281**, 60 (1996).
- [27] M. Kriegmaier, M. Zimmermann, K. Wolf, U. Zimmermann, and V.L. Sukhorukov, *Biochim. Biophys. Acta* **1568**, 135 (2001).