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# Letter A modified relation between the intraocular and intracranial pressures



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# Hanjing Tian, Ruigi Du, Fan Song\*

State Key Laboratory of Nonlinear Mechanics (LNM), Institute of Mechanics, Chinese Academy of Sciences, Beisihuanxi Road No.15, Beijing, 100190, China

#### HIGHLIGHTS

- A novel relation between the intraocular and intracranial pressures.
- The present relation is in good agreement with the existing experiments.
- This relation overcomes the induced singularity of conventional models.

#### ARTICLE INFO

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## ABSTRACT

A modified relation between the intraocular and intracranial pressures is presented by employing the least square method to fit the existing experiments. Relative analysis here indicates that this modified relation not only is better than the previous relation by comparing with the existing experimental data but also overcomes the induced singularity in applying the existing mechanical models to compute the mechanical properties of the lamina cribrosa. The present study will be a beneficial help to understanding the relationship between the intraocular and intracranial pressures and even glaucomatous developing.

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Glaucoma is the most common cause of irreversible blindness, affecting more than 60 million people worldwide, and increasing to about 80 million by 2020 [1]. In the course of developing glaucoma, the difference between the intraocular pressure (IOP) and the intracranial pressure (ICP) plays a key role. It is this difference that exerts on the primary site of glaucoma-the lamina cribrosa (LC), then results in the irreversible deformation of LC, and finally induces the damage of the optic nerves passing through LC, thus triggers the visual field defect, which is medically considered to be the main pathological mechanism of glaucoma [2,3]. The existing studies already prove that the glaucomatous visual field defect is positively correlated with the difference between IOP and ICP and inversely correlated with the ICP [4]. Because directly observing and determining the deformation of LC and the damage of the optic nerves in LC in a living body is scarcely possible in the clinical diagnosis and treatment of glaucoma, building a mechanical model of LC now still remains the most efficient way to understand the states of LC in vivo [5]. Obviously, the relation between IOP and ICP is of the essence in correctly making the mechanical model of LC. However, the exact relation between IOP and ICP has not been understood very well yet so far.

In the existing studies of the relation between IOP and ICP, when IOP is greater than 20 mmHg, Quigley [3] experimentally and clinically proved that ICP is roughly equal to 10 mmHg, i.e.

$$ICP = 10 \text{ mmHg} \text{ for } IOP \ge 20 \text{ mmHg}.$$
 (1a)

When IOP is less than 20 mmHg, on the one hand, there is not a deterministic result as Eq. (1a) deriving directly from clinic and experiment up to now, but lots of investigations indicate that ICP is significantly and positively correlated with IOP [6–10]. On the other hand, in order to analyze the deformation of LC, all the existing mechanical models of LC greatly simplify the relation between IOP and ICP as a linear relation when IOP is less than 20 mmHg [5,11,12], i.e.

$$ICP = \frac{1}{2}IOP$$
 for  $IOP < 20$  mmHg. (1b)

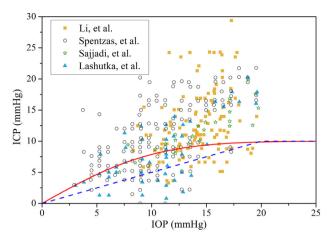
However, this simplification results directly in a serious problem: the derivatives of Eq. (1) at the point of IOP = 20 mmHg are inexistence, or the rate of change of ICP at the point is discontinuous according to Eq. (1), as shown in Fig. 1, which further induces that the singularity of the deformation appears at the

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E-mail address: songf@lnm.imech.ac.cn (F. Song).

\* Corresponding author.

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**Fig. 1.** The relations between IOP and ICP. The broken line stands for Eq. (1), the solid line expresses Eq. (3), and different figurate points indicate the different experimental data from Refs. [6–9]. The derivatives of the broken line do not exist at IOP = 20 mmHg.

point in the existing mechanical model of LC [11,12], as shown in Fig. 2. This can never happen in a living body. It is because that IOP is produced by the aqueous humor and ICP derives from the cerebrospinal fluid pressure. The changes of the two liquids that generate the two pressures have to be continuous in a live body. Therefore, exactly determining the relation between IOP and ICP becomes a key point in understanding the behaviors of LC. In addition, there is a big error between the values that the existing mechanical models [11,12] compute the deformations of LC according to Eq. (1) and the existing experiments measure the deformations of LC [13] when ICP is less than 20 mmHg, as shown in Fig. 2.

Here we modify the relation between IOP and ICP, and make the modified relation satisfy the three conditions: (i) at the point of IOP = 20 mmHg, its derivatives are continuous; (ii) when IOP  $\geq$  20 mmHg, its values are equal to the values of Eq. (1a); (iii) when IOP < 20 mmHg, its values are better than those of Eq. (1b) in fitting the existing experiments.

Based on the characteristics and topography of Eq. (1), we choose a modified error function to fit the relation between IOP and ICP, namely,

$$ICP = A \cdot erf(\alpha \cdot IOP), \tag{2}$$

where  $\operatorname{erf}(\xi) = (2/\sqrt{\pi}) \int_0^{\xi} e^{-\eta^2} d\eta$  is Gauss error function, *A* and  $\alpha$  are undetermined coefficients.

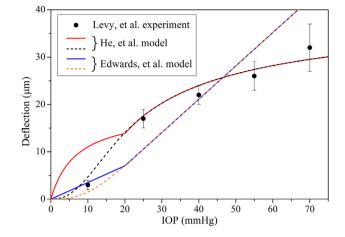
First of all, according to the properties of Gauss error function and Eq. (1a), the modulus of Eq. (2) is readily determined as A = 10.

Secondly, we use the existing experimental data between ICP and IOP to fit the other undetermined coefficient. By synthesizing the four group experimental data [6–9], we obtain  $\alpha = 0.088$  by virtue of employing the least square method, as shown in Fig. 1. Therefore, Eq. (2) is written by

$$ICP = 10 \cdot erf(0.088 \cdot IOP). \tag{3}$$

The relative error between Eqs. (1a) and (3) at the point of IOP = 20 mmHg is readily calculated to be 1.28%, which reaches a very good approximate in the theory of biomechanics [14]. And with IOP increasing, the error will rapidly decrease according to the property of Gauss error function.

Further, we separately compare the root mean squared errors (RMSEs) of Eqs. (1b) and (3) with the experimental data from Refs. [6–9]. All RMSE values of Eq. (3) are better than those of Eq. (1b), as shown in Table 1.



**Fig. 2.** The deflections of LC. The solid curves stand for the results of the different mechanical models from Refs. [11,12] by employing Eq. (1), and the broken curves display the results of the different mechanical models from Refs. [11,12] by employing Eq. (3). There are the derivative singular points of the solid curves at IOP = 20 mmHg.

 Table 1

 The RMSEs of Eqs. (1) and (3) with respect to the experiments [5–8].

	RMSE of Eq. (3) (mmHg)	RMSE of Eq. (1) (mmHg)
Lashutka, et al.	4.3244	4.7691
Sajjadi, et al.	3.8741	5.0320
Spentzas, et al.	5.9195	7.3052
Li, et al.	6.3158	7.5901

Finally, employing the existing mechanical models of LC and the modified relation between IOP and ICP, Eq. (3), we compute the deformations of LC, and compare these results with the existing experiments, as shown in Fig. 2. Our computed results display that at the point of IOP = 20 mmHg, the derivative of the deformation is continuous; when IOP  $\geq$  20 mmHg, the deformations are the same as the values obtained by using Eq. (1a); when IOP < 20 mmHg, the deformation that the modified relation between IOP and ICP, Eq. (3), is closer to the real states than Eq. (1a) in clinic experiments.

In conclusion, based on the existing experiments, we use the least square method to obtain a modified relation between IOP and ICP. The modified relation not only is better than the previous relation by virtue of comparing with the existing experimental data, but also thoroughly overcomes the induced singularity in applying the existing mechanical model of LC to compute the mechanical properties of LC. Obviously, the present result is a beneficial help to understanding the relationship between IOP and ICP and even glaucomatous developing.

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