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# **CLINICAL STUDY**

The effect of flap thickness on the visual and refractive outcome of myopic laser *in situ* keratomileusis

### Abstract

Aims To study the effect of flap thickness on the visual and refractive outcome of myopic laser *in situ* keratomileusis (LASIK) surgery. *Methods* A total of 196 myopic eyes with spherical equivalent (SE) from -2.00 to -5.00 dioptres which underwent LASIK were studied retrospectively. Uncorrected visual acuity (UCVA), best-corrected visual acuity (BCVA) and manifest refraction were measured up to 6 months postoperatively. Logistic and linear regression analyses were used to examine the correlation of flap thickness to visual and refractive outcomes, postoperative complications and enhancement rate.

Results UCVA at 1 week and 1 month, but not at 1 day and at 6 months (P > 0.05), were negatively correlated to flap thickness (P < 0.05). BCVA was unrelated to flap thickness (P>0.05). At 1-month post-LASIK sphere and cylinder were not related to flap thickness (P > 0.05), but spherical equivalent was negatively correlated (P<0.05). Significant negative correlations with the sphere and SE were noticed at 6 months for the eyes not requiring enhancements (P<0.05). Flap thickness had no significant relationship to postoperative complications (P > 0.05) and no significant predictive value on the rate of enhancement procedures or the efficacy, safety and predictability indices (P > 0.05). Conclusions In myopic LASIK thinner flaps are associated with faster visual recovery and less myopic SE, but the BCVA and the final UCVA are independent of flap thickness. Postoperative complications are unrelated to flap thickness. Flap thickness does not affect the efficacy, safety and predictability, or the rate of enhancement procedures.

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*Keywords:* LASIK; flap; thickness; myopia; outcome

## Introduction

The advantages of laser *in situ* keratomileusis (LASIK),<sup>1</sup> with the fast and almost painless visual recovery have rendered this procedure very popular in the last decade.<sup>2</sup> In LASIK a corneal flap is created using a microkeratome and an excimer laser ablates the stromal bed. Flap dimensions vary considerably among the various microkeratomes.<sup>3–10</sup> Although flap diameter variability has not been reported to be a frequently significant problem in myopic laser ablations, flap thickness has been linked to a variety of postoperative complications, such as iatrogenic keratectasia,<sup>11,12</sup> epithelial ingrowth,<sup>13</sup> flap dislocation, striae,<sup>14</sup> and corneal opacities.<sup>15–17</sup>

This study sought to examine the relation of flap thickness to the visual and refractive results of myopic LASIK, in addition to assessing the safety, efficacy, predictability, complication, and enhancement rates in relation to flap thickness.

### Materials and methods

### Patient selection

In this retrospective analysis patients who underwent LASIK for myopia and compound myopic astigmatism in a single treatment centre between December 1999 and January 2001 were studied. Only patients that were 18 years of age or older, with at least 1 year stable refraction,

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spherical equivalent (SE) from -2.00 to -5.00 dioptres (D) at the spectacle plane were eligible for inclusion in this study. All patients were treated with the same excimer laser (Chiron Technolas 217C, Bausch and Lomb) aiming for emmetropia. Patients with a history of ocular disease, previous ocular surgery (including refractive surgery), intraoperative complications, connective tissue disorders or expecting and nursing mothers were also excluded. In order to eliminate the influence of treated optical zone variability on the postoperative results only eyes treated with a 6 mm optical zone were included in this study. After receiving informed consent all patients underwent a complete ophthalmic assessment that included manifest and cycloplegic subjective refraction, pupillometry, corneal topography, dilated fundoscopy, slit-lamp biomicroscopy, and corneal thickness measurement by contact ultrasound pachymetry (DGH 5100 Pachymeter, DGH Technology Inc, Exton, PA, USA). According to the manufacturer, this model pachymeter is accurate to within  $\pm 5 \,\mu$ m. Central corneal thickness measurements with the DGH have been reported to be accurate, reliable, and reproducible.<sup>18</sup>

# Surgical technique

Topical anaesthesia with oxybuprocaine was instilled in the eye prior to creation of a nasally hinged corneal flap using a manual microkeratome (LSK One, Moria, Antony, France). Microkeratome heads of 100 or 130  $\mu$ m were used according to the desired flap thickness. Intraoperative pachymetry of the central stromal bed (three measurements, considering the mean) was performed following flap elevation and the subtraction method (preoperative minus stromal bed thickness) was used to calculate the flap thickness. All operations were performed by the same experienced surgeon (JBS). After ablation the flap was repositioned on the stromal bed and aligned. Flap alignment and interface were checked immediately at the slitlamp. Interface debris were removed, when necessary, with the help of BSS using a Rycroft cannula mounted in a 5 ml syringe and flap misalignment was corrected similarly at the slitlamp in the theatre area. Postoperatively all patients received topical antibiotic and steroids for 1 week.

# Enhancement procedures

Eyes with stable residual refractive error underwent enhancement procedures, at least 1 month following the initial LASIK surgery. Arcuate keratotomies were performed in eyes with mixed astigmatism, whereas further laser ablation in the stromal bed (Re-LASIK) was performed for all the other residual refractive errors after the initial flap was lifted. All patients had at least 1-month follow-up following retreatment.

# Patient examinations

Postoperative data were collected retrospectively from follow-up visits of day 1, week 1, and months 1 and 6. For the study of the outcome, parameters such as uncorrected visual acuity (UCVA) (1 day, 1 week, 1 month, 6 months), best spectacle-corrected visual acuity (BCVA) (1 and 6 months), SE (1 and 6 months), presence and type of complications, rate of enhancement procedures required, were studied and their relation to flap thickness were examined. Mean visual acuity was calculated and expressed as the LogMAR equivalent of decimal equivalent as described by Holladay.<sup>19</sup>

# Statistical analysis

For statistical analysis SPSS for Windows version 10.0.7 was used. Measurements of continuous variables were described by their mean, standard deviation, and range. For continuous variables, mean values were compared by the paired or unpaired Student's *t*-test for normally distributed data according to the data under consideration. Differences for nominal variables were tested using the Fisher's exact test for independence. Correlations between flap thickness and continuous variables were determined by linear regression analysis. Logistic regression analysis was used to examine the effect of a 10  $\mu$ m increase in flap thickness on nominal variables. A *P*-value less than 0.05 was considered statistically significant.

# Results

# Demographic and preoperative data

In all, 196 eyes (108 patients) were included in this study. The demographic, preoperative, and operative data are shown in Table 1. In all, 47 eyes (24%) had flap thickness between 70 and 100  $\mu$ m, 93 (47%) between 101 and 130  $\mu$ m and 56 (29%) between 141 and 184  $\mu$ m.

# Visual and refractive results

The postoperative visual and refractive results and their relation to flap thickness are displayed in Tables 2 and 3 respectively.

Linear regression analysis revealed a statistically significant negative correlation between flap thickness and UCVA at the first week and the first month but not at 6 months (Table 3). Neither the UCVA of the eyes which underwent retreatment or those that did not show a significant correlation to flap thickness at 6 months (P > 0.05).

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The postoperative sphere was not correlated to flap thickness at 1 month but it was negatively correlated at 6 months. This was mainly due to the eyes that did not require enhancement procedures ( $\beta = -0.260$ , t = -2.875, P = 0.005) and not to the eyes which underwent enhancements (P = 0.695), although there was not a significant change (P > 0.05) at the sphere between 1 and 6 months in either group of eyes.

A statistically significant negative correlation was found between the SE and flap thickness at 1 but not at 6 months (Table 3). However, the SE was negatively correlated to the flap thickness at 6 months ( $\beta = -0.202$ , t = -2.203, P = 0.030) when eyes that underwent enhancement procedures were excluded. There was no significant change in the SE of the eyes that underwent enhancement procedures between 1 and 6 months (P = 0.316) as a result of these procedures, or in the eyes that did not required enhancements (P = 0.394).

## Postoperative complications

A total of 14 eyes developed postoperative complications. Logistic regression analysis did not reveal any statistically significant relationship between either the total number of complications or to individual postoperative complications and flap thickness (P > 0.05). These complications did not result in loss of lines of BCVA more than in uncomplicated eyes at 1 month (P > 0.99) and 6 months (P = 0.20).

### Loss or gain of lines of BCVA

In all, 3.57 and 2.19% of eyes lost two or more lines of BCVA at 1 and 6 months, respectively, whereas 89.8 and 90.51% maintained or gained lines of BCVA at the same times. The flap thickness had no statistically significant predictive value in the number of lines of BCVA lost or gained at 1 (P = 0.26) or 6 months (P = 0.229).

Table 1	Demographic,	preoperative,	and o	operative data
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Demographic and preoperative data			
196 (103 OD, 93 OS)			
108			
64/44			
$32.50 \pm 7.93$ (20–55) years			
$-2.89\pm0.86$ (-5.00 to -1.00) D			
$-0.79\pm0.53$ (-3.50 to 0) D			
$-3.29\pm0.82$ (-5.00 to -2.00) D			
$0.02 \pm 0.17$ (0.001–0.5)			
$0.96 \pm 0.86$ (0.3–1.5)			
$558.85 \pm 33.55$ (485–640) $\mu$ m			
118.09±22.52 (70–184)μm			
6 mm			
$67.76 \pm 15.03$ (32–103) $\mu m$			
$373 \pm 32.56 (302 - 456) \mu\mathrm{m}$			

UCVA: uncorrected visual acuity; BCVA: best-corrected visual acuity.

 Table 3
 Table demonstrates the relation of flap thickness to visual and refractive results

	Flap thickness relation to visual results				
	β	t	P-value		
UCVA at 1 day	-0.014	-0.191	0.848		
UCVA at 1 week	-0.149	-2.093	0.038		
UCVA at 1 month	-0.178	-2.522	0.012		
UCVA at 6 months	-0.005	-0.062	0.951		
BCVA at 1 month	-0.075	-1.050	0.295		
BCVA at 6 months	0.086	1.007	0.316		
Flap thickness relation to refractive results					
Sphere at 1 month	-0.104	-1.455	0.147		
Cylinder at 1 month	-0.088	-1.232	0.22		
SE at 1 month	-0.182	-2.581	0.011		
Sphere at 6 months	-0.208	-2.472	0.015		
Cylinder at 6 months	0.095	1.113	0.268		
SE at 6 months	-0.155	-1.827	0.070		

SE: spherical equivalent;  $\beta$ : standardised regression coefficient; t: t regression coefficient.

Table 2 Visual and refractive results

	Day 1 (n = 196)	Week 1 (n = 196)	<i>Month</i> 1 (n = 196)	Month 6 (n = 137)
UCVA BCVA Sphere Cylinder SE	0.87±0.77 (0.4–1.5)	0.93±0.77 (0.2–1.5)	$\begin{array}{c} 0.94 \pm \ 0.78 \ (0.3 - 1.5) \\ 1.01 \pm 0.84 \ (0.4 - 1.5) \\ -0.03 \pm 0.37 \ \mathrm{D} \ (-1.00 \ \mathrm{to} \ +1.50 \ \mathrm{D}) \\ -0.37 \pm 0.47 \ \mathrm{D} \ (-2.50 \ \mathrm{to} \ 0 \ \mathrm{D}) \\ -0.21 \pm 0.32 \ \mathrm{D} \ (-1.00 \ \mathrm{to} \ 0.88) \end{array}$	$\begin{array}{c} 0.97 \pm 0.82 \ (0.4 - 1.5) \\ 1.02 \pm 0.85 \ (0.4 - 1.5) \\ -0.09 \pm 0.30 \ \mathrm{D} \ (-1.00 \ \mathrm{to} \ + 0.75 \ \mathrm{D}) \\ -0.28 \pm 0.31 \ \mathrm{D} \ (-1.00 \ \mathrm{to} \ 0 \ \mathrm{D}) \\ -0.23 \pm 0.30 \ \mathrm{D} \ (-1.25 \ \mathrm{to} \ + 0.50) \end{array}$

UCVA: uncorrected visual acuity; BCVA: best-corrected visual acuity; SE: spherical equivalent.

# **Enhancement** procedures

Of the eyes, 4.08% had enhancement by laser and 6.63% by arcuate keratotomies for residual refractive errors. No eye had retreatment because of regression. Flap thickness did not have any significant effect on the rate of enhancements by laser (P = 0.58) or arcuate keratotomies (P = 0.84) individually, nor on the rate of enhancements procedures in total (P = 0.60). Eyes that underwent enhancement procedures had no significant difference in flap thickness, preoperative sphere, cylinder and SE (P > 0.05) from those they did not.

# Efficacy, safety, and predictability

There was no relation of flap thickness to efficacy, predictability, and safety at any time (P > 0.05) (Table 4).

### Discussion

The microkeratomes used in LASIK allow surgeons to create flaps with a frequent and well-documented difference between the predicted and the actual thickness.<sup>20–25</sup> According to manufacturers, flap thickness should ideally fall within 10–15% of the intended thickness. However, flap thickness shows often a wide variability,<sup>3–5,21,22</sup> thus intraoperative pachymetry is considered crucial for its measurement.

Flap thickness created by blade-based microkeratomes depends on multiple factors.<sup>3–6,15,22,26–34</sup> New devices using high-pressure water<sup>35</sup> and intrastromal lasers<sup>36–38</sup> have been recently developed to obtain more reproducible corneal flaps and appear to be more consistent than blade-based microkeratomes.<sup>39</sup>

**Table 4** Table demonstrates the relation of flap thickness to efficacy, safety, and predictability indices

	Flap thickness relation to refractive indices			
	Value (%)	OR	P-value	
Efficacy at 1 month	73.47	0.878	0.076	
Efficacy at 6 months	74.54	0.993	0.45	
Safety at 1 month	3.57	1.250	0.20	
Safety at 6 months	2.19	0.680	0.20	
Predictability (0.5 D) at 1 month	86.74	0.948	0.57	
Predictability (1 D) 1 month	100	NA	NA	
Predictability (0.5 D) at 6 months Predictability (1 D) at 6 months	87.59 99.27	0.825 1.878	0.119 0.292	

Efficacy: percentage of eyes with UCVA equal to or greater than preoperative BCVA; Safety: percentage of eyes with a loss of 2 or more Snellen lines of BCVA; Predictability  $_{(0.5\,D)}$ : percentage of eyes with spherical equivalent within  $\pm 0.5\,D$  from emmetropia; Predictability  $_{(1\,D)}$ : percentage of eyes with spherical equivalent within  $\pm 1\,D$  from emmetropia.

Although the lamellar flap does not contribute to the biomechanical stability of the cornea in LASIK, it remains a very important factor of this stability as flap thickness determines, along with the ablation depth and the preoperative corneal thickness, the thickness of the RSB. Reduced thickness of the RSB is implicated in the development of iatrogenic keratectasia in normal corneas following LASIK.<sup>11,40–43</sup> A minimum RSB thickness of  $250 \,\mu$ m is considered necessary for the prevention of keratectasia.<sup>11,44</sup>

Thicker corneal flaps may result in either excessively reduced stromal bed in cases of very deep lamellar cut<sup>12</sup> or deeper ablations weakening the biomechanical structure of the cornea,<sup>40,45</sup> whereas thinner flaps result in more superficial ablations maximising the RSB, thus reducing the risk of keratectasia.

Moreover, the creation of thinner flaps renders more eyes amenable to LASIK. In addition a thicker RSB is of great importance when laser enhancement is contemplated, eliminating the need for another microkeratome pass with the associated potential risks.<sup>46</sup>

It has been suggested by Dr Chayet that thinner flaps may have better UCVA on the first day as they develop less oedema hastening visual recovery (Cheryl Guttman, Ultra-thin flaps may guard against ectasia, Eurotimes, September 2002, pages 1 and 5). We did not notice a significant correlation between UCVA and flap thickness at day 1. However, a significant negative correlation was observed between flap thickness and UCVA at 1 week and 1 month. We believe that the presence of residual corneal flap oedema at the first postoperative day is responsible for the lack of a significant correlation. Nevertheless gradual resolution of flap oedema by the end of first week reveals the true relation. The reason for the better UCVA in eyes with thinner flaps after the first week is that they simply adopt the shape of the underlying stromal bed better. This may be an advantage in customised ablations over thicker flaps, which mask the fine changes of stromal curvature due to their rigidity. However, the advantage of the thinner flaps in UCVA is lost at 6 months as UCVA improved in eyes with thicker flaps although not significantly (P > 0.05), even in the eyes that did not have enhancement procedures. Therefore, one can conclude that eyes with thinner flaps achieve faster visual recovery following myopic LASIK than the thicker ones, but the final UCVA is independent of the flap thickness.

Unlike other investigators,<sup>4</sup> we found a statistically significant negative correlation between flap thickness and postoperative SE at 1 month, although we did not observe a significant correlation with the postoperative sphere and cylinder individually. Similarly, a significant negative correlation (P<0.05) was observed at 6 months with the SE and the sphere, but not

the cylinder, for those eyes that did not undergo enhancement procedures.

Thinner flaps have been associated with a higher incidence of complications.<sup>13,14,31</sup> Epithelial ingrowth has been seen more frequently in eyes with thinner flaps and was attributed to the difficulty in realigning the flap in the stromal bed.<sup>13</sup> Excessive manipulation under the flap to obtain proper alignment may result in implantation of loose epithelial cells in the interface and epithelial ingrowth. We believe that our technique of checking and realigning the flap, and, if necessary, cleaning the interface from debris and obvious epithelial cells, at the slitlamp are the main reasons for the low incidence of epithelial ingrowth in our cohort (0.51%).

It has also been suggested that thinner flaps may be displaced easier than thicker ones and may result in striae.<sup>14</sup> Excessive manipulations may lead to flap oedema, which in turn reduces the adherence of the flap and increases the risk of flap displacement and striae. In our study, we have observed a very low incidence of flap striae (0.51%) even though almost a quarter of the flaps were thinner than  $100 \,\mu$ m.

A number of confocal microscopy studies in myopic human eyes following LASIK have shown a negative correlation between flap thickness and subclinical haze,<sup>47</sup> and activated—cell area thickness.<sup>47,48</sup> Other confocal microscopy studies have confirmed the development of clinical haze in eyes with thin flaps but flap thickness is not the only aetiologic factor, as it does not happen exclusively in eyes with thin flaps.<sup>16</sup> Thinner flaps affect the integrity of the most anterior keratocyte layer that, through release of several cytokines from the corneal epithelium,<sup>49</sup> may control the keratocyte activation, myofibroblast transformation and haze formation after lamellar refractive surgery.<sup>48</sup>

Central corneal opacities in eyes with very thin flaps have been described,<sup>15–17</sup> attributed mainly to ablation of the Bowman's layer. We have not observed corneal opacities in any of our patients despite 24% of the eyes in our study had flaps between 70 and 100  $\mu$ m. We believe that the reason for this is that we did not violate the Bowman's layer. Although the term 'thin flap' has been used extensively in the literature, it appears that there is not a consensus of opinion regarding its definition. One should distinguish between pathologically thin flaps where there is dissection within the Bowman's layer, or even the epithelium (ie buttonholes), which may eventually lead to the creation of corneal scarring, and reduced thickness flaps with the lamellar cut below the Bowman's. Lamellar flaps thinner than  $60 \,\mu\text{m}$  are suspicious as average corneal epithelium are approximately  $50 \,\mu \text{m}^{50}$  and Bowman's layer  $10 \,\mu \text{m}$ . Appropriate measures to avoid pathologically thin flaps have been described.14

The enhancement rate in our cohort (10.71%) is within the range (5.5–36%) published in large series.<sup>51,52</sup> Logistic regression analysis revealed that increasing flap thickness had no significant effect on the rate of enhancement procedures.

Ambrosio and Wilson<sup>31</sup> have suggested that thin flaps may be associated with regression caused by epithelial hyperplasia due to the proximity of the epithelium to stromal wound healing response and epithelium modulating growth factor production. We did not observe a significant regression between 1 and 6 months in our group for the eyes which did not require enhancement procedures (P > 0.05).

The limitations of our study are the short follow-up and the relatively small number of patients. As complications after LASIK are rare, a large number of patients are required in order to determine whether there is a relationship between flap thickness and individual postoperative complications. In addition, intraoperative ultrasound pachymetry for the measurement of flap thickness by the subtraction method is subject to errors due to overhydration and dehydration of the cornea.<sup>53,54</sup>

In conclusion, in myopic LASIK thinner flaps are associated with faster visual recovery although the BCVA and the final UCVA are independent of flap thickness. In addition flap thickness is inversely related to SE, with eyes with thinner flaps achieving less myopic SE refraction. Postoperative complications are not related to flap thickness, provided that the lamellar cut and laser ablation respect the Bowman's layer. Moreover, enhancement rate, efficacy, safety, and predictability are not influenced by flap thickness.

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