Analysis of radiofrequency lesions in egg whites in vitro produced by application of the Tew electrode for different temperatures and times

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BACKGROUND: Understanding the size and shape of radiofrequency lesions is important to reduce side effects when applied to patients.

OBJECTIVES: To investigate the radiofrequency lesions produced by the application of the Tew electrode for different temperatures and times.

METHODS: The white from a fresh hen's egg was placed in a rectangular glass container and warmed to 37°C. After immersion of the Tew electrode in the egg white, radiofrequency lesions were produced at 65°C, 70°C, 75°C, 80°C, 85°C and 90°C. For each temperature, photographs were taken at 10 s, 20 s, 30 s, 40 s, 50 s, 60 s, 70 s, 80 s, 90 s, 100 s, 110 s and 120 s. The size of the lesion was measured at each temperature and time. A mixed model was used to analyze the data.

RESULTS: The size of the lesion increased with increasing temperature and time. There were statistically significant differences in the size of the internal radius between the 65°C and 70°C groups and the 70°C and 75°C groups, as well as in the 70°C and 75°C groups in the size of the external radius and the 60°C to 80°C groups in the size of the distal radius. The maximum lesion size was produced at 90°C and 120 s, and was 1.06±0.16 mm in internal radius, 0.37±0.15 mm in external radius, 0.39±0.04 mm in distal radius.

CONCLUSION: The Tew electrode produces lesions following the contour of the tip, and the internal radius is larger than the external and distal radius. The best combination of temperature and time for lesioning using the Tew electrode is 80° C, for 60 s to 90 s.

Key Words : Egg white; Electrode; Lesion size; Radiofrequency

 $R_{\rm a}$ adiofrequency (RF) thermocoagulation has been used clinically as a modality to treat chronic pain since the 1950s (1); presently, it is one of the most frequently used neurolytic techniques for the relief of pain.

High-frequency electrical current is applied adjacent to the structure of the nerve that is to be ablated, leading to ionic oscillation and frictional dissipation of the ions and electrolytes, which produces heat. Thus, the main advantage of RF is more reliable and reproducible lesion production at a desired specific neurological location compared with electrocautery (2). The lesion size and shape produced by RF thermocoagulation is determined by various parameters, such as temperature, electrode configuration and the characteristics of the tissue (3). Information regarding the lesion size generated using different parameters would increase the efficacy of the technique and reduce the incidence of side effects.

L'analyse des lésions de radiofréquence dans des blancs d'œuf *in vitro*, produites par l'application de l'électrode Tew à différentes températures et pendant des durées plus ou moins longues

HISTORIQUE : Il est important de comprendre la dimension et la forme des lésions de radiofréquence pour en limiter les effets secondaires chez les patients.

OBJECTIFS : Examiner les lésions de radiofréquence produites par l'application de l'électrode Tew à différentes températures et pendant des durées plus ou moins longues.

MÉTHODOLOGIE : Les chercheurs ont placé le blanc d'un œuf de poule frais dans un contenant rectangulaire en verre et l'ont chauffé à 37 °C. Après y avoir immergé l'électrode Tew, ils ont produit des lésions de radiofréquence à 65 °C, 70 °C, 75 °C, 80 °C, 85 °C et 90 °C. À chaque température, ils ont pris des photos au bout de 10 s, 20 s, 30 s, 40 s, 50 s, 60 s, 70 s, 80 s, 90 s, 100 s, 110 s et 120 s. Ils ont mesuré la dimension de la lésion à chaque température et au bout de chaque durée. Ils ont utilisé un modèle mixte pour analyser les données.

RÉSULTATS : La dimension de la lésion augmentait proportionnellement à la température et à la durée. Il y avait des différences statistiquement significatives dans la dimension du rayon interne entre les groupes de 65 °C et de 70 °C et entre ceux de 70 °C et 75 °C, de même que dans la dimension du rayon externe dans les groupes de 70 °C et 75 °C et dans la dimension du rayon distal dans les groupes de 60 °C à 80 °C. Les lésions atteignaient leur dimension maximale à 90 °C, au bout de 120 s, d'un rayon interne de 1,06±0,16 mm, d'un rayon externe de 0,37±0,15 mm et d'un rayon distal de 0,39±0,04 mm.

CONCLUSION : Les électrodes Tew produisent des lésions qui suivent le contour de leur pointe, et le rayon intérieur est plus grand que les rayons externe et distal. La meilleure combinaison de température et de durée pour effectuer une lésion avec l'électrode Tew est de 80 °C pendant 60 s à 90 s.

Many studies involving RF lesions have been reported, including those regarding lesion shapes generated using various electrodes (4-7). Egg whites have been used in many studies to determine the size and shape of RF lesions. Because egg whites do not precisely model neural tissue, their use in in vitro studies is limited. However, egg whites are composed primarily of pure protein, a composition similar to that of human nerve tissue, and its transparency facilitates observation of changes in the lesions (8).

Since 1974, when Sweet and Wepsic (9) first described a technique for generating RF lesions in the Gasserian ganglion for the treatment of trigeminal neuralgia, RF thermocoagulation of the Gasserian ganglion has been used for treatment of idiopathic trigeminal neuralgia, atypical trigeminal neuralgia, atypical facial pain, intractable ocular pain, cluster headache and cancer pain (10). Due to the unique anatomical position of the Gasserian ganglion, the Tew electrode was developed as a new

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Figure 1) Determination of the radius of a lesion generated using the Tew electrode (Tew-CTC, Radionics, USA) in egg white. DR Distal radius; ER External radius; IR Internal radius

flexible curved electrode, virtually eliminating complication (11). The flexible curved electrode permits more precise positioning of the electrode tip than a straight electrode, resulting in generation of discrete lesions and decreasing the incidence of sensory and motor side effects (12). Despite the above-mentioned advantage of the Tew electrode, its lesion size and shape have not been investigated extensively.

Therefore, using egg whites, we measured and analyzed the size and shape of lesions produced by the application of the Tew electrode for different temperatures and times. The resulting data will facilitate the use of RF in chronic pain management.

METHODS

The white from a fresh hen's egg was placed in a rectangular glass container and warmed in a water bath to 37°C. The Tew electrode set was fixed to a sponge on the upper surface of the container, and a thin metal plate, the ground electrode, was inserted into the egg white and suspended in a distant corner of the container. The Tew cannula and electrode (Tew-CTC; Radionics, USA) were inserted into the egg white. The Tew electrode is a flexible curved electrode 0.5 mm in diameter with an active tip 5 mm in length. RF was applied through the electrode using an RF lesion generator system (RFG-3C; Radionics, USA). RF lesions were produced by the Tew electrode at 65°C, 70°C, 75°C, 80°C, 85°C and 90°C. For each temperature, photographs were taken at 10 s, 20 s, 30 s, 40 s, 50 s, 60 s, 70 s, 80 s, 90 s, 100 s, 110 s and 120 s. Lesion production was repeated five times for a total of 360 lesions ($12 \times 6 \times 5 = 360$). The target temperature was reached by elevating the temperature at a rate of 6°C/s to 9°C/s. Measuring the lesion size began when the target temperature was reached. Coagulation was defined when the transparent egg white became hazy and whitish. The transparency of the egg white and glass container enabled the changes in size of the RF lesions over time to be photographed. The magnification ratio was calculated by comparing the diameter of the real electrode with the diameter of the electrode measured on the photograph. The internal radius (IR) and external radius (ER) of the lesions were then measured, and the distal radius (DR) from the tip of the lesion from the active tip of the electrode was determined. The IR of the lesion was the perpendicular distance from the internal side of the midpoint of the electrode to the furthest point on the perimeter of the lesion. The ER of the lesion was the perpendicular distance from the external side of the lesion. The DR was the distance from the tip of the electrode to the furthest point on the lesion in line with the long axis of the electrode (Figure 1). A mixed model in SPSS version 21.0 (IBM Corporation, USA) was used to compare lesion size according to temperature and time of application. P<0.05 was deemed to indicate statistical significance.

RESULTS

The Tew electrode produced lesions following the contour of the tip (Figure 2). The size of the lesions increased with increasing



Figure 2) Photograph of a lesion generated in egg white using the Tew electrode (Tew-CTC, Radionics, USA) at 75° C for 30 s

temperature and time, and the IR was larger than the ER and DR in egg white (Figure 3). The maximum lesion size was generated at 90°C and 120 s, and was 1.06 ± 0.16 mm in IR, 0.37 ± 0.15 mm in ER and 0.39 ± 0.04 mm in DR from the tip. The IR of the lesion increased with temperature, and there were statistically significant differences between the 65°C and 70°C groups and the 70°C and 75°C groups (Figure 4A). A similar pattern was observed for the ER, and there was a statistically significant difference between the 70°C and 75°C groups (Figure 4B). As the temperature of electrodes were increased, DR of the tip gradually increased from 60°C to 80°C. However, there were no significant differences of DR above 80°C (Figure 4C). The IR, ER and DR of lesions increased significantly with application time, from 10 s to 120 s, at the same temperature. However, the lesion sizes increased with application time at the same temperature without statistical significance among the time groups.

DISCUSSION

RF thermocoagulation has been used for the treatment of conditions including chronic pain, malignant tumour and cardiac arrhythmia. RF thermocoagulation enables reliable formation of discrete thermal lesions in a desired specific neurological location (10). As a mechanism of RF, some authors have demonstrated that the compound action potentials of A-delta and C fibres in nerves are blocked at lower temperatures compared with those of the larger A-alpha and A-beta fibres (13-15). However, this concept is controversial and was contradicted by multiple histological studies (16,17). The current rationale is that RF lesions lead to nonselective heat destruction of all fibres in the centre of the lesion; at the periphery of the lesion, there is selective destruction of more vulnerable, small fibres with preservation of the larger fibres (18,19).

Although RF for the treatment of chronic pain has proven to be an effective, minimally invasive treatment, it has inherent risks and complications. Because the heating radius of a small needle is limited, the needle must be placed exactly on the nerve to guarantee its coagulation (8). The lesion size varies depending on the shape, thickness and length of the noninsulated electrode tip, and with the time and temperature of application (3). Therefore, information regarding the lesion size, generated using different parameters, would increase the efficacy of the technique and reduce the incidence of side effects.



Figure 3) The size of lesions generated in egg white by radiofrequency using the Tew electrode (Tew-CTC, Radionics, USA) at different temperatures and times. Changes in size (mean) of the (A) internal radius, (B) external radius and (C) distal radius. Figures were created using Microsoft Excel 2007 (Mircosoft Corporation, USA)



Figure 4) Analysis of the size of lesions generated in egg white using the Tew electrode (Tew-CTC, Radionics, USA) at different temperatures. A The internal radius increased with temperature and statistically significant differences were noted between the 65°C and 70°C groups and between the 70°C and 75°C groups. B The external radius increased with temperature and there was a statistically significant difference between the 70°C and 75°C groups. C As the tempeature of electrodes were increased, distal radius of the tip gradually increased from 60°C to 80°C. However, there were no significant differences of distal radius above 80°C. *P<0.05

The goal of RF treatment of trigeminal neuralgia is to achieve total pain relief while preserving trigeminal sensation (20). The most prevalent complications of RF treatment are sensory loss in the treated branch or paralysis of the masseter muscle. In the long term, anesthesia dolorosa, corneal hypoesthesia and keratitis, and temporary paralysis of the third and fourth cranial nerve can occur (21). Several improvements have been developed to minimize unwanted side effects. The innovation of RF thermocoagulation by the Tew, van Loveren, and Keller group included both the introduction of the Tew curved-tip electrode (11), and the modification of the technique for cannulation of the foramen ovale using image-guided fluoroscopy (22). The Tew electrode was developed as a new flexible curved-tip electrode that has virtually eliminated motor root dysfunction. The tip of the curved electrode extends 5 mm in an axial direction from the distal end of the cannula, and the electrode has a rounded distal end to avoid any cutting action and is 0.5 mm in diameter (9). Manipulation of the electrode about its 360° axis enables an infinite variety of position adjustments of the electrode tip. The incidence of masseter weakness and paresthesia has decreased compared with that in patients treated with a straight electrode (23). However, despite the above-mentioned advantages, the size and shape of lesions generated using the Tew curved-tip electrode have not been reported. Therefore, our data may facilitate the design of clinical trials that produce an adequate lesion during RF treatment using the Tew electrode.

The size of the lesions may vary in vivo, depending on factors such as vascularity and tissue heat conductivity (24). Computed tomography and magnetic resonance imaging are the only modalities for estimating the size and shape of the lesions in the human brain. However, these techniques are not sufficient to evaluate correlations between lesion shape and electrode configuration; additionally, the results of early postoperative imaging of lesions are difficult to interpret because of tissue swelling and edema (25,26). The in vitro study by Eriksson et al (27), revealed that the real lesion size in tissue may be greater than that in albumin because electrical resistance is greater in tissue than in albumin. Nevertheless, egg whites seemed a suitable medium in which to examine lesion production because RF treatment denatures proteins, of which an egg white is predominantly comprised. In a previous report (8), monopolar lesions produced in both egg whites and skeletal muscle were similar in geometry and size to those produced in brain tissue. Moreover, the translucency of the egg white allows visualization and measurement of the lesions as they are created (6). There are many combined studies involving in vitro lesioning and its clinical application (28,29), and lesion size in egg whites was useful as a reference for clinical application.

Using a straight or curved electrode, the lesion extends radially around the electrode tip in the shape of an oblate spheroid (30). The lesion is an oval shape parallel to the needle that usually does not exceed the tip of the needle; thus, ideally, the needle should be parallel to the targeted nerve (31). Theoretically, the lesion size is dependent on the heat generated, the heat loss and the electrode size. The generated heat is inversely proportional to the fourth power of the distance from the electrode, proportional to the second power of the current, and linearly proportional to the duration of lesioning (32). However, in our study where a Tew electrode was used, the lesion size was not an exact function of temperature or time. Our results reveal that the Tew curved electrode produced lesions following the contour of the tip, and the lesions exceed the tip of the needle. Lesion size differed depending on the IR, ER and DR from the tip. The IR was larger than the ER and DR. The latter finding is noteworthy because of the possibility for manipulation of the Tew electrode about its 360° axis (33).

Previous reports have demonstrated the importance of temperature monitoring for predicting lesion size (34). Our data also show that temperature was an important determinant of lesion size. Lesion size increased significantly with increment of temperature. Lesion IR increased significantly between the 65°C and 70°C groups and 70°C and 75°C groups, and lesion ER between the 70°C and 75°C groups. DR of the tip gradually increased from 60°C to 80°C, but there were no significant differences of DR above 80°C. These results suggest that temperatures >80°C would not result in a significant change in the ER, IR and DR of the lesion. Consistency of lesion size is achieved by letting the lesion come to thermal equilibrium. The nominal radius of the lesion approaches a maximum value as equilibrium between the RF heat into and the heat conduction away from the tissue around the tip is established. In our experiment, the peak effect appears as a result of the thermal equilibrium process, and there was no statistical significance in changes of the lesion size above 80°C. Higher temperatures may result in gas formation, tissue adherence to the probe, tearing and hematoma formation with changes in electrode position (35). Popped lesions may be caused by tissue boiling and gas formation (36), resulting in damage to adjacent neural structures. Popped lesions are associated with the rate of increase in tip temperature (2). When heat spreads too rapidly, scattered areas of explosive water vapourization are produced, resulting in an irregularly shaped lesion larger than the desired volume. Bogduk et al (8) increased the temperature of the tip at a rate of 1°C to 2°C per second in their study; however, Eriksson et al (27) and the present study both increased the temperature of the tip at a rate of 6°C/s to 9°C/s. Because there were no popped lesions in the present study, the rate of increase in temperature did not lead to gas formation or tissue boiling.

Increased duration of application has a lesser effect on lesion size than increasing temperature (2). In the present study, lesion size increased significantly as application duration was raised from 10 s to 120 s, but no significant differences were found among the time groups. Because the time of application at a constant temperature is less critical in determining the lesion's size, the use of longer durations is not recommended. However, a minimum time has to be exceeded before the lesion approaches its maximum possible size. As a general rule, satisfactory lesion durations are 30 s for small lesions (eg, percutaneous cordotomy) and 60 s to 90 s for large lesions (eg, Gasserian ganglionectomy, thalamotomy) (2).

Larger lesions can reduce the likelihood of missing a target nerve and increase the number and extent of target nerves captured by each lesion (37). This may help to avoid incomplete neurolysis, to improve the degree and duration of pain relief, and to reduce procedure time and x-ray exposure. However, larger lesions may also increase the likelihood and severity of damage to nontarget nerves, skin burns in thin patients and postprocedural paresthesia (38). The ideal technique places the smallest lesion possible in the nervous system, exactly at the target structure, with full lesion controllability and without causing complications and discomfort to the patient (10). Therefore, in our experiment, the best combination of temperature and time was 80°C for 60 s to 90 s.

In our study, the lesion size attained using the Tew electrode in an egg white increased with increasing temperature and time, but time had a lesser effect on lesion size than temperature. The Tew electrode produced lesions following the contour of the tip, and the IR was larger than the ER and DR. The best combination of temperature and time for lesioning was 80°C for 60 s to 90 s. Although our study was an in vitro experiment using egg whites, we believe that our results will be valuable data for RF treatment in a clinical setting.

CONCLUSION

The Tew electrode produces lesions following the contour of the tip, and the IR is larger than the ER and DR in egg whites. The lesion size increases with increasing temperature and time. The best combination of temperature and time for lesioning using a Tew electrode is 80°C for 60 s to 90 s. We believe that our data will facilitate expansion of the applications of Tew curved electrodes and provide information useful for clinical trials.

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