

соммент Subthreshold laser therapy guidelines for retinal diseases

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Subthreshold laser therapy has been utilised in clinical practice for more than three decades, and numerous randomised and real-life studies has proven its efficacy and safety in various retinal diseases [1–10]. However, despite decades of usage, we still do not have a standard protocol for subthreshold laser applications and settings. The Subthreshold Ophthalmic Laser Society (SOLS) is comprised of global experts (10) in subthreshold lasers and has a goal to establish subthreshold laser guidelines and establish level 1 evidence on subthreshold laser in ophthalmology.

Here we report consensus guidelines by SOLS for the subthreshold laser settings and applications in diabetic macular oedema (DMO) and central serous chorioretinopathy (CSCR). These guidelines were based on 43 questions based on different aspects of subthreshold laser applications in these diseases. Responses were collected from each expert in a masked fashion in first round and two virtual meetings were performed to discuss these questions to reach final consensus.

For DMO, the experts support the role of subthreshold laser in both centre-involving and non-centre involving macular oedema in conjunction with or without anti-VEGF therapy. Subthreshold laser settings for DMO include 5% duty cycle, 200 ms pulse duration, and 150-200 µm spot size with no spacing between laser spots using integrated pattern system. SOLS experts support titration for subthreshold laser application with 50% of threshold power (achieved with subthreshold laser). SOLS experts suggest treating oedematous area with subthreshold laser, however, does not mandate focal treatment of microaneurysm. SOLS experts consider transfoveal treatment safe with settings described earlier, however, they emphasis the careful application and reconfirmation of subthreshold laser safe settings before application. Considering no visible or structural changes following subthreshold laser treatment, experts do not recommend any specific structural imaging studies to evaluate laser spots, however, autofluorescence or OCT could be carefully evaluated during follow up visits for any laser scars. Experts recommend follow-up evaluation at 6-8 weeks after subthreshold laser application, however, repeat subthreshold laser is suggested after 2-3 months of initial application, in case of poor response. Experts recommend adjunctive therapies such as anti-VEGF or steroid therapy for diabetic macular oedema as per the physician's discretion.

For CSCR, SOLS experts recommend subthreshold laser for both acute as well chronic types. Unlike conventional observation for 3–4 months for acute CSCR, considering the safety profile of subthreshold laser, SOLS experts recommend subthreshold laser application in 1 month if there is no self
 Table 1.
 Subthreshold laser consensus guideline settings for diabetic macular oedema and central serous chorioretinopathy by the Subthreshold Ophthalmic Laser Society (SOLS).

| Subthreshold laser settings | Diabetic macular oedema | Central serous chorioretinopathy |
|--------------------------------|----------------------------|-------------------------------------|
| Duty cycle | 5% | 5% |
| Pulse duration | 200 ms | 200 ms |
| Spot size | 150–200 µm | 100–200 µm |
| Spacing between spots | No | No |
| Titration | Yes | Yes |
| Titration power | 50% of threshold power | 50% of threshold power |

resolution. For chronic CSCR, SOLS expert support subthreshold laser as first line as well as combination therapy with other treatment options. Recommended settings for subthreshold laser application are 5% duty cycle, 200 ms pulse duration and 100-200 µm spot size. Settings are same for both acute and chronic CSCR with no spacing between the spots using integrated pattern system. SOLS experts support these settings safe for the transfoveal laser application, with reconfirmation of subthreshold settings. SOLS experts suggest titration with 50% of threshold power (threshold estimation using subthreshold laser mode) (Table 1). Regarding area covered, for acute CSCR, focal leak and adjacent area should be treated with subthreshold laser. In case of chronic CSCR, SOLS experts suggest areas of focal as well as diffuse hyperfluorescence on fluorescein angiography. Considering no visible or structural changes following subthreshold laser, experts do not recommend any specific structural imaging studies to evaluate laser spots, however, autofluorescence or OCT could be carefully evaluated during follow up visits for any laser scars. Evaluation at 6-8 weeks is recommended and in case of poor response or persistence of the subretinal fluid, a repeat treatment with same settings is recommended, in addition to adjuvant treatment options as per physician discretion.

These consensus guidelines do not suggest management of these disease but suggest laser application guidelines using different subthreshold laser deliver systems. These guidelines would help to establish standard subthreshold laser applications in clinical practice.

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REFERENCES

- Guymer RH, Wu Z, Hodgson LAB, Caruso E, Brassington KH, Tindill N, et al. Subthreshold nanosecond laser intervention in age-related macular degeneration: the LEAD randomized controlled clinical trial. Ophthalmology. 2019;126:829–38.
- Lavinsky D, Cardillo JA, Melo LA Jr, Dare A, Farah ME, Belfort R Jr. Randomized clinical trial evaluating mETDRS versus normal or high-density micropulse photocoagulation for diabetic macular edema. Investig Ophthalmol Vis Sci. 2011;52:4314–23.
- Pei-Pei W, Shi-Zhou H, Zhen T, Lin L, Ying L, Jiexiong O, et al. Randomised clinical trial evaluating best-corrected visual acuity and central macular thickness after 532-nm subthreshold laser grid photocoagulation treatment in diabetic macular oedema. Eye. 2015;29:313–21.
- Passos RM, Malerbi FK, Rocha M, Maia M, Farah ME. Real-life outcomes of subthreshold laser therapy for diabetic macular edema. Int J Retin Vitreous. 2021;7:4.
- Wu Z, Luu CD, Hodgson LAB, Caruso E, Brassington KH, Tindill N, et al. Secondary and exploratory outcomes of the subthreshold nanosecond laser intervention randomized trial in age-related macular degeneration: a LEAD study report. Ophthalmol Retina. 2019;3:1026–34.
- Donati MC, Murro V, Mucciolo DP, Giorgio D, Cinotti G, Virgili G, et al. Subthreshold yellow micropulse laser for treatment of diabetic macular edema: Comparison between fixed and variable treatment regimen. Eur J Ophthalmol. 2021;31:1254–60.
- Zhou L, Chong V, Lai K, Huang C, Xu F, Gong Y, et al. A pilot prospective study of 577-nm yellow subthreshold micropulse laser treatment with two different power settings for acute central serous chorioretinopathy. Lasers Med Sci. 2019;34:1345–51.

- Maruko I, Koizumi H, Hasegawa T, Arakawa H, Iida T. Subthreshold 577 nm micropulse laser treatment for central serous chorioretinopathy. PLoS ONE. 2017;12:e0184112.
- Chhablani J, Alshareef R, Kim DT, Narayanan R, Goud A, Mathai A. Comparison of different settings for yellow subthreshold laser treatment in diabetic macular edema. BMC Ophthalmol. 2018;18:168.
- Al-Barki A, Al-Hijji L, High R, Schatz P, Do D, Nguyen QD, et al. Comparison of short-pulse subthreshold (532 nm) and infrared micropulse (810 nm) macular laser for diabetic macular edema. Sci Rep. 2021;11:14.

COMPETING INTERESTS

JC has received consulting a speaker fees from Salutaris, Novartis, Allergan, and Biogen.

ADDITIONAL INFORMATION

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