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Journal: Journal of Neurosurgical Sciences

Paper code: J Neurosurg Sci-4753

Submission date: May 22, 2019

Article type: Original Article

Files:

1. Manuscript

Version: 1

Description: Manuscript

File format: application/vnd.openxmlformats-officedocument.wordprocessingml.document

Application of Indocyanine Green video angiography in vascular neurosurgery

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Keywords: vascular, aneurysm, indocyanine green, ICG-VA, AVM

1 **Abstract**

2
3 Indocyanine green video angiography (ICG-VA) is a non-invasive, easy to use and
4 very useful tool for various neurosurgical procedures.
5

6 The first application was in neurovascular surgery, because it was born as an
7 intravascular tracer for vessels visualization; this has been really useful in aneurysms,
8 AVMs and dural fistulas surgery where identification, obliteration or patency of
9 vessels is essential.
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11 Introduced in vascular neurosurgery since 2003, ICG-VA applications have broadened
12 over time, both in vascular and in other neurosurgical fields.
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14 As the first, in 2003 Raabe et al. [1] described the use of ICG-VA for intraoperative
15 assessment of cerebral vascular flow, enabling visualization of vessel patency and
16 aneurysm occlusion during aneurysm surgery.
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18 ICG-VA applications in vascular neurosurgery have significantly increased over time
19 including complex aneurysms, bypass, atero-venous malformations (AVM) artero-
20 venous fistulas (AVF), evaluation of cortical perfusion.[2-4].
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22 The procedure can be easily repeated after 5–10min. Adverse reactions are comparable
23 to those of other types of contrast media, with frequencies of 0.05% (hypotension,
24 arrhythmia, or, more rarely, anaphylactic shock) to 0.2% (nausea, pruritus, syncope, or
25 skin eruptions).
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27 The aim of the present study is to systematically analyses ICG-VA applications in
28 vascular neurosurgery, highlighting the reported advantages and disadvantages, and
29 discussing future perspectives.
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Overview of Indocyanine Green video angiography in vascular neurosurgery

Aneurysms

The first systematic report on ICG-VA application in cerebral aneurysm surgery was published in 2003 by Raabe et al.[1] Following this preliminary experience, several consistent patient series were published on ICG-VA use in aneurysm surgery by Roessler et al. in 2014[5]. ICG-VA is often compared to micro-doppler and digital subtraction angiography (DSA) to evaluate vascular anatomy, before and after clipping, and to assess correct position of the clip, presence of aneurysm residuals, patency of normal vessels. One interesting paper reports about a patient suffering from a giant aneurysm of the right MCA; indocyanine green was injected inside the aneurysm in order to identify a target middle cerebral artery branch (MCA) for bypass and allowing confident preservation of blood supply to distal areas to the sacrificed vessel.

The study published by Roessler et al. including 295 cases is nowadays the largest published series on this topic[5]. They reported a repositioning of aneurysm clips in 9% of the procedures because of parent vessel or adjacent perforating arteries occlusion not detected by micro-Doppler ultrasonography. Moreover, in 4.5% of the procedures residual perfusion was detected and one or more clips were applied. Nevertheless, postoperative angiography in 9.1% of successful ICG- VA guided clip applications demonstrated unexpected residual aneurysms.

A very interesting study was published by Hardesty et al. [6] where a comparison is between two “eras”: the intraoperative DSA one and the ICG-VA. They retrospectively evaluated whether the rates of perioperative stroke, unexpected postoperative aneurysm residual, or parent vessel stenosis differed in 100 patients from each era.

The issue of per-patient cost of intraoperative imaging was also estimated in a study published by Nishiyama et al. [7](patients undergoing ICG-VA and endoscopy in order to facilitate intra- operative real-time assessment of the patency of perforating arteries behind parent arteries or aneurysms). In a more recent paper by Bruneau et al. [8]endoscopic ICG-VA was used in anterior communicating artery aneurysm clipping

1 providing information regarding aneurysm occlusion and patency of parent and
2 branching vessels and small perforating arteries.
3

4
5 During aneurysms surgery, the use of ICG-VA is already consolidated, it's easy, rapid
6 to perform and non invasive[9]; it has good spatial resolution and allows a good
7 evaluation of the complete aneurysm exclusion, neck remnant, blood flow in the parent
8 vessels. A percentage of unexpected neck residuals and close vessels occlusion has
9 been reported (6% in the Dashti et al. study[10]).
10

11
12 Intraoperative ICG-VA is therefore considered a valuable and cost-effective
13 replacement to routine intraoperative diagnostic angiography. Nevertheless,
14 postoperative DSA still cannot be avoided because of a remaining little percentage of
15 inaccuracy of both intraoperative techniques [11]. Thus, care should be taken when
16 considering ICG-VA as the sole means for intraoperative evaluation of aneurysm clip
17 application. Thus, care should be taken when considering ICG-VA as the sole means
18 for intraoperative evaluation of aneurysm clip application.
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29 AVMs

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31 In AVMs surgery, ICG-VA is considered to give a moderate contribution. As reported
32 in the paper by Zaidi et al. [12] it can be useful for the intraoperative mapping of the
33 angio-architecture of superficial AVMs; it gives the possibility to visualize flow
34 variations directly on surgical field and to confirm the occlusion of nidus feeding
35 arteries. Unfortunately, it is considered useless in the residual detection [13], and the
36 difficult visualization in deep located AVMs [14] could limit its application
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42 Indeed in the most important series (56 patients) published by Zaidi et al.[12] do not
43 report any difference between patients undergoing or not ICG-VA in terms of residual
44 disease or clinical outcomes.
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50 Vascular bypasses and low flow malformations

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52 The principal aim of vascular bypass is the preservation of blood flow. Considering the
53 different type of bypass (i.e. STA-MCA OA-PICA, radial artery, saphenous vein, IC-
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1 IC, STA–PCA) we found that the application of ICG-VA is quite mandatory to
2 evaluate the patency of the vessels and thus the blood flow.[15]

3
4 Januszewski et al. [16] identified 3 different pattern on ICG-VA angiography to
5 evaluate the patency in the next 24–48 h. Prinz et al. [17]assessed the use of ICG-VA
6 to detect hemodynamic changes within the macro- circulation and microcirculation
7 after bypass. Schuette–Barrow in 2010 [18] together with other colleagues [2, 19-21]
8 published ICG-VA application as a tool in a minimally invasive approach for ligation
9 of dural AVFs [22]. Conversely, also in intramedullary cavernomas, ICG can provide
10 useful information about lesion margins and associated venous anomalies [23-26].

11
12 In this scenario, we can currently consider the use of ICG-VA of paramount importance
13 also in vascular bypasses and low flow vascular malformations.
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23 **Discussion**

24 Intraoperative ICG-VA, nowadays, is a standard visualization tool in vascular
25 neurosurgery.
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27 It was used for the first time as an intravascular tracer for vessels visualization [27];
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29 Thanks to its quickness and non invasiveness and providing real-time information it
30 has become an invaluable tool to intraoperative surgical decision-making;
31

32 in addition, recent experiences have shown ICG-VA potential use in a very different
33 set neurosurgical branches, including oncological surgery, endoscopy, pituitary,
34 cerebral hemodynamic studies
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36 During aneurysms surgery it is easy, rapid to perform and non invasive[27] and offers
37 a good spatial resolution[10, 28] and allows a good evaluation of the complete
38 aneurysm exclusion, neck remnant, blood flow in the parent arteries and perforating
39 arteries [27, 28].
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41 By the way, it has a limited view to the operating field [27, 29] and it can give a limited
42 ability to visualize the part of the base behind the aneurysm dome in deeply located
43 aneurysms in case of blood clot or intramural thrombi or calcifications [10, 30].
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1 Intraoperative ICG-VA is considered a valuable cost-effective replacement to routine
2 intraoperative diagnostic angiography but postoperative DSA wouldn't be avoided
3 because of a remaining little percentage of inaccuracy[6].
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6 ICG-VA had also in important role in bypasses surgery because of its definition, direct
7 evidence of graft patency and the identification of the parent and recipient arteries [30-
8 32]. On the other hand, it gives a limited and restricted visualization to the operating
9 field. Januszewski et al. established a classification to evaluate the type of flow through
10 the bypass graft: Type I flow (robust anterograde flow) strongly correlates with early
11 postoperative graft patency, Type II (anterograde flow but delayed compared to other
12 adjacent vascular structures) and Type III (anterograde flow but delayed with no
13 continuity to the bypass site) are both predicative of early graft failure and need to be
14 intraoperative revised in order to avoid postoperative complications.[16]
15

16 ICG-VA in the assessment of hemodynamic changes within the macrocirculation and
17 microcirculation after bypass surgery has been detected by Prinz et al. but currently,
18 there isn't a standard intraoperative method to obtain a visualization and quantitative
19 measurement of cortical microcirculatory perfusion. [17]
20

21 ICG-VA is also adopted in AVMs surgery: it can be suitable for the intraoperative
22 mapping of the angio-architecture of superficial AVMs giving the chance to visualize
23 flow variations directly on surgical field and to confirm the occlusion of nidus feeding
24 arteries. Beside that it is useless in the residual detection [33] and in deeply deep
25 located AVMs [34]. Basically, there are no difference between patients undergoing or
26 not ICG-VA in terms of residual disease or clinical outcomes as reported by Zaidi et
27 al[12].
28

29 Last, but not least, in dAVFs surgery, the main step is the identification of fistulous
30 site. Different studies report 100% correspondence to postoperative controls[19], the
31 identification of the fistulous site and confirmation of its obliteration during surgery
32 [19, 20, 22] and the possibility to identify both the early-filling fistula and the presence
33 of abnormal retrograde drainage thanks to the visualization of the timing and direction
34 of blood flow [20]. On the other hand, the use of ICG-VA, can get and increase of
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operating time and the limited visualization to the operating field requiring a larger exposition [2, 19].

Future Perspectives

ICG-VA allows evaluation of vessel patency but, unfortunately, does not allow quantitative, time-dependent and spatially precise analysis of intravascular blood flow. In fact, conventional ICG-VA gives information about vessel patency but not quantitative, time-dependent and spatially precise analysis of intravascular blood flow [28].

Faber et al. [35] and Kamp et al.[36] pioneered a color-coded maps of time to half-maximal peak that valuable for giving an overview of blood flow perturbations and distribution by extracting data from conventional ICG-VA[34].

Cortical ICG in the setting of acute SAH can also provides evidence of acute vasoconstriction after haemorrhage and measurement of CBF intraoperatively as stated by Shubert et al. [37].

Lastly, different mini-invasive approached and several high resolution tools are proposed for better intraoperative visualization[38-44]. Nishiyama et al. recently used ICG-VA for endoscopic-assisted aneurysm surgery, to better view perforating arteries, that are often not visible to the sole microscopic view[7].

Conclusions

We can conclude that ICG-VA has reached in recent years a wide utilization in various neurosurgical fields, mainly in vascular neurosurgery surgery. The technique allows future developments such as quantitative evaluation of cerebral blood flow or the combined use with the endoscope. It should be considered among the most promising easy and low cost tools towards the direction of a minimally invasive and safer neurosurgery.

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