Abstract

The aim of our study is to evaluate the value of Argon Plasma Coagulation in the treatment of gastrointestinal vascular abnormalities.

Patients and methods  This is a descriptive and analytical retrospective study, from January 2009 to September 2020. 198 patients who have benefited from treatment with Argon Plasma Coagulation for vascular anomalies of the digestive tract divided into 2 groups:

- Group A: Patients with radial rectitis lesions (n = 107).
- Group B: Patients with lesions of digestive angiodysplasia (n = 91).

Results  The mean age of our patients was 64.95 ± 9.88 years [43 - 83] in group A, while in group B the mean age was 65.19 ± 14.29 years [40 - 91] with a clear male predominance in 72.5%. The majority of patients in group A were followed for prostate cancer in 33.8%, and 26.3% of patients in group B had chronic renal failure, followed by stomach cancer in 15.8%, and esophageal cancer in 10.5%. Clinical symptomatology was dominated by rectories in 40.2% in group A versus 46.8% in group B. Rectal involvement was dominated in group A in 98.1%, whereas in group B the lesions were mainly located in the stomach in 60.5%. The endoscopic evolution was favorable in all our patients with a clear improvement of rectal lesions and digestive angiodysplasia lesions. The total complication rate in our series was nil.

Conclusion  Plasma Argon coagulation is a very effective method in the endoscopic treatment of digestive haemorrhages with good tolerability and a low complication rate.

Keywords

- argon plasma coagulation
- radiation rectitis
- intestinal angiodysplasia

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Introduction

Digestive angiodysplasias are vascular malformations whose pathogenesis is poorly known. They are a rare cause of upper and lower occult or externalized digestive hemorrhages. Most of them are of acquired origin, most often affecting the right colon and the cecum, but can sit on the entire digestive tract.

On the other hand, radiation rectitis is a frequent complication, of variable severity, occurring a few months after the radiation of the pelvic-perineal organs has stopped, but which can manifest itself several years later.

Argon plasma coagulation (APC) is the technique of choice in the treatment of digestive vascular abnormalities, improving the quality of life of patients and reducing the need for repeated transfusions.1,2

The objective of our study is to evaluate the value of Argon Plasma Coagulation in the treatment of gastrointestinal vascular abnormalities.

Materials and Methods

Method

Type of study:

This is a descriptive and analytical retrospective study over an 11-year period, from January 2009 to September 2020.

Instruments

The equipment required to perform the APC includes:

- **The Argon bottle**: Argon can be easily ionized at high electrical frequency and is then transformed into a stable plasma of a pale yellow color.

- **The electric scalpel with APC program**.

- **The pedal**: connected to the unit and allowing the control of the electric arc.

- **The APC probe**: this is a teflon catheter in which a tungsten electrical guide is inserted. This electrode is connected at its distal end with a ceramic inserted in the catheter lumen. Some catheters are perforated laterally at their distal end their diameters are 1.5 mm, 2.3 mm, and 3.2 mm. The standard length is 220 cm, up to 300 cm for use in enteroscopy.

- **The endoscope**: The endoscopes used for the treatment of gastroduodenal and colonic lesions were video endoscopes. The lesions of the small bowels that were not accessible of oesogastroduodenal fibroscope or colonoscopy were treated by high or low enteroscopy.

- **Flow rate (power //flux)**:
  - Colon, Duodenum and Small Intestine: 40W// 0.6L/min.
  - Esophagus and Stomach: 50W// 0.8L/min.

Patients:

198 patients who have benefited from treatment with Argon Plasma Coagulation (APC) for vascular anomalies of the digestive tract, divided into 2 groups:

- **Group A**: Patients with radial rectitis lesions (n = 107).
- **Group B**: Patients with lesions of digestive angiodysplasia (n = 91).

Positive diagnosis was established by endoscopic examinations (Fibroscopy, enteroscopy, colonoscopy) or by endoscopic videocapsule.

All patients included in the study had a pre-anesthetic consultation. They were given a complete clinical examination and a biological assessment, mainly the hemostasis assessment.

All patients who were going to benefit from a colonoscopy or low enteroscopy were given a preparation per 4 liters of polyethylene glycol.

A clinical and biological control in 15 days to one month was systematic in order to stratify the patients requiring a second APC session.

Inclusion Criteria

For Group A:

- Age > 18 years.
- Pelvic radiotherapy
- Presence of rectorragies.
- To have a diagnosis of radiation rectitis established on the basis of the anamnesis data and the endoscopic aspect of the rectal mucosa.

For Group B:

- Age > 18 years.
- Any patient presenting an externalized (upper/lower) or occult digestive hemorrhage, with the presence of angiodysplasia lesions during the endoscopic assessment of the digestive tract (esogastroduodenal fibroscope, colonoscopy, enteroscopy, videocapsule).

Data Collection

The data collection was done on the medical records of the patients, who are hospitalized either 24 hours before the procedure, or on the same day (day hospital), on the other hand the technical part relating to the endoscopic procedure was filled in by the operator explaining the different steps of the endoscopic procedure.

The data that have been exploited are:

- **Demographic data**: - Age
- Sex

- **Throughout the procedure**:
  - **Diagnosis**: What is the diagnosis?
  - **Location**: Where is the diagnosis located?
  - **Type of treatment**: How was the treatment performed?
  - **Degree of coagulation**: What is the degree of coagulation?

- **In the follow-up**:
  - **Number of sessions**: How many sessions of treatment were needed?
  - **Duration of treatment**: How long did the treatment last?
  - **Complications**: Were there any complications during or after the treatment?

- **Long-term follow-up**:
  - **Recurrent bleeding**: Did the bleeding recur?
  - **Quality of life**: How did the quality of life change post-treatment?
  - **Rehospitalization**: Was rehospitalization necessary?
- Surgical history (Gastrectomy, operated gastric ulcer, lung cancer, prostate cancer, esophageal cancer, cervical cancer, rectal cancer, anal cancer, colon cancer, system disease, kidney failure, cirrhosis).
- Indication: Anemic syndrome, rectorragy, melena, hematemesis, epigastralgia.

*Endoscopic data:* - Number of lesions.
- Site of injury (Oesophagus, fundus, antrum, bulb, duodenum, jejunum, ileum, gastric stump, oeso-jejunal or gastro-jejunal anastomosis, afferent or efferent loop, cecum, colon, rectum).
- Presence or not of active bleeding
- Use of complementary endoscopic hemostasis (Clip) or not.

*Number of APC sessions*
*Prescription of an adjuvant medical treatment* (Corticotherapy or 5- aminosalicylated enemas) or not.

**Statistical Analysis**
We performed a two-step statistical analysis:

1. A descriptive analysis of our population including the different demographic, clinical and endoscopic variables:
   a. **Qualitative:** in numbers and percentage
      - Sex
      - Clinical signs
      - Site of the vascular malformation.
      - Presence or not of active bleeding
      - Use of complementary endoscopic hemostasis maneuvers
      - Adjuvant medical treatment

b. **Quantitative:**
   - Age (Mean ± standard deviation)
   - Number of lesions (Median [Interquartile range])
   - Number of APC sessions (Median [Inter-quartile range])

2. An analytical study was then carried out:
   - The comparison of the quantitative variables with normal distribution was done by the Student’s test.
   - The comparison of quantitative variables with an unsymmetrical distribution was done by the Man Whitney test.
   - The comparison of qualitative variables between two groups was done by the Chi-square or Fisher test.

   A value of $p < 0.05$ was considered statistically significant.

   The statistical analysis was performed using SPSS version 24.0 software.

**Table 1** Patient Demographics:

<table>
<thead>
<tr>
<th>Characteristics</th>
<th>Group A (n = 107)</th>
<th>Group B (n = 91)</th>
<th>$p$</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age</td>
<td>64.95 ± 9.88</td>
<td>65.19 ± 14.29</td>
<td>0.89</td>
</tr>
<tr>
<td>Sex: Men</td>
<td>48(45.3%)</td>
<td>66(73.3%)</td>
<td>0.0001</td>
</tr>
<tr>
<td>Woman</td>
<td>58(54.7%)</td>
<td>24(26.7%)</td>
<td></td>
</tr>
</tbody>
</table>

Our study has been approved by our hospital’s ethics committee.

**Results**

The mean age of our patients was $64.95 ± 9.88$ years [43 - 83] in group A, with a sex ratio M/F of 0.82, while in group B the mean age was $65.19 ± 14.29$ years [40 - 91] with a clear male predominance in 72.5%. (–**Table 1**)

The majority of group A patients were followed for prostate cancer in 33.8% (n = 28), followed by cervical cancer in 33.8% (n = 22), while 13.8% (n = 9) and 4.6% (n = 3) had cancer of the rectum and bladder, respectively; and one patient had squamous cell carcinoma of the anus.

On the other hand, 26.3% (n = 5) of patients in group B had chronic renal failure, followed by cancer of the stomach in 15.8% (n = 3), and cancer of the esophagus in 10.5% (n = 2) (–**Table 2**).

The clinical symptomatology was dominated in group A by rectorragies in 40.2% (n = 43), an anemic syndrome in 3.7% (n = 4) and only one patient had melena of low abundance.

On the other hand, in group B, 46.8% (n = 49) of the patients had rectorragies, 35.8% (n = 33) had iron deficiency anemia, 32.6% (n = 30) had melenas, two patients had hematemesis and one patient had epigastralgia. (–**Table 3**)

The median number of APC sessions in both groups was 2.1-2.

The median number of lesions in each patient was 2.1-3.

The rectal injury was dominated in group A in 98.1% (n = 105), whereas in group B the lesions were mainly located in the stomach in 58.3% (n = 55), followed by the coecum in 30.9% (n = 25), the jejenum in 19.1% (n = 13) and the duodenum in 17.7% (n = 16).

The presence of active bleeding was without significant difference between the two groups: 10.7% in group A versus 11% in group B.

The use of adjuvant corticosteroid therapy or 5-ASA enema was only in group A in 4.7% (n = 5). Use of hemostatic clip was only in group B in 3.3% (n = 3) (–**Table 4**).

The endoscopic evolution was favorable in all our patients with a clear improvement of rectal lesions and digestive angiodysplasia lesions. The total complication rate in our series was nil.

**Discussion**

Vascular abnormalities of the gastrointestinal tract are a rare cause of digestive hemorrhage and pose a diagnostic and
Table 2 Patient medical and surgical history

<table>
<thead>
<tr>
<th>Background</th>
<th>Group A (n = 107)</th>
<th>Group B (n = 91)</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>Gastrectomy</td>
<td>3(15,8%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Gastric ulcer operated</td>
<td>1(5,3%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Lung cancer</td>
<td>1(5,3%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Prostate cancer</td>
<td>28(43,1%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Cancer of the esophagus</td>
<td>2(2,2%)</td>
<td></td>
<td>&lt;0,0001</td>
</tr>
<tr>
<td>Cervical cancer</td>
<td>22(33,8%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Rectal cancer</td>
<td>9(13,8)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Anal cancer</td>
<td>1(1,5%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Colon cancer</td>
<td>1(1,1%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>System disease</td>
<td>2(2,2%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Kidney failure</td>
<td>5(5,5%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Cirrhosis</td>
<td>1(1,1%)</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Table 3 Reason for patient hospitalization:

<table>
<thead>
<tr>
<th>Clinical signs</th>
<th>Group A (n = 107)</th>
<th>Group B (n = 91)</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>Anemic Syndrome</td>
<td>4(3,7%)</td>
<td>33(35,8%)</td>
<td></td>
</tr>
<tr>
<td>Melena</td>
<td>1(2,2%)</td>
<td>30(32,6%)</td>
<td></td>
</tr>
<tr>
<td>Rectragies</td>
<td>43(40,2%)</td>
<td>49(46,8%)</td>
<td>0,25</td>
</tr>
<tr>
<td>Hematemesis</td>
<td>2(2,2%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Epigastralgies</td>
<td>1(1,1%)</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Table 4 Endoscopic characteristics of patients

<table>
<thead>
<tr>
<th>Characteristics</th>
<th>Group A (n = 107)</th>
<th>Group B (n = 91)</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>Number of lesions</td>
<td>2 [1–3]</td>
<td>2 [1–4]</td>
<td>0,019</td>
</tr>
<tr>
<td>Esophagus</td>
<td>4(4,4%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Fundus</td>
<td>27(29,7%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Antre</td>
<td>22(24,2%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Bulb</td>
<td>6(6, 6%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Duodenum</td>
<td>10(11,1%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Jejunum</td>
<td>13(19,1%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Gastric stump</td>
<td>4(4,4%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Related/efferent handle</td>
<td>5(5,6%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Oeso-jejunal anastomosis</td>
<td>1(1,3%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Coecum</td>
<td>25(30,9)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Straight colon</td>
<td>7(8,6%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Transverse colon</td>
<td>4(4,9%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Left colon</td>
<td>5(4,7%)</td>
<td>5(5,6%)</td>
<td></td>
</tr>
<tr>
<td>Rectum</td>
<td>105(98,1%)</td>
<td>6(6,7%)</td>
<td></td>
</tr>
<tr>
<td>Active bleeding</td>
<td>11(10,7)</td>
<td>9(11%)</td>
<td>0,9</td>
</tr>
<tr>
<td>Complementary endoscopic haemostasis maneuvers</td>
<td>3(3,3%)</td>
<td></td>
<td>0,09</td>
</tr>
</tbody>
</table>
They were first described in 1839, and in 1974 the term intestinal angiodysplasia (IA) was defined as a single or multiple superficial acquired vascular lesion developed in the mucosa and/or submucosa of the gastrointestinal tract wall, without being associated with a cutaneous or visceral angiomatous lesion.

Their prevalence in the general population is 0.83%, they are equally present in both sexes and are more frequent after the sixties.

Various equivalent terms can be found in the literature: “arteriovenous malformation”, “telangiectasia” or “vascular ectasia”. On the other hand, intestinal angiodysplasia are to be differentiated from benign vascular tumors called hemangiomas or malignant tumors called angiosarcomas.

The typical appearance of an IA is that of a bright red, rounded, irregularly contoured, slightly raised, bright red “spot”, usually small, less than 10 mm in size (Fig. 1). A halo around the lesion is sometimes described, corresponding to a paler peri-lesional mucosa.

An IA corresponds histologically to a dilatation of the mucosal capillaries in communication with ectasic and tortuous submucosal veins (Fig. 2).

They account for 4% of the aetiologies of upper digestive haemorrhages. Gastric and/or duodenal localization is preferred, but they are more frequent in the small intestine and the colon, particularly in the ascending portion and in the cecum.

In our study the lesions were mainly located in the stomach in 58.3% followed by the coecum in 30.9%, the jejunum in 19.1% and the duodenum in 17.7%.

The development of endoscopy techniques and the quality of the video endoscopes used, particularly in terms of image resolution, make endoscopy the first avenue to follow for the diagnosis of IA; when located in the small intestine, diagnosis is made using endoscopic videocapsule or high or low enteroscopy.

It is not uncommon to miss some IA due to localization behind a fold or in the small intestine which is highly mobile, so there should be no hesitation to repeat endoscopic examinations when clinical suspicion is strong.

Approximately 90% of IAs stop bleeding spontaneously, but the risk of recurrence is high. This risk is between 25% and 65% depending on the studies. Surgical treatment is infrequent, up to 12% in some studies. Mortality from hemorrhagic IA is approximately 2%.

In another retrospective study, haemorrhagic angiodysplasia was reported in 50% of the 261 patients investigated by double-balloon enteroscopy for unexplained digestive bleeding who were treated with argon plasma hemostatic therapy. The rate of hemorrhagic recurrence at 3 years was 46%, with a high number of vascular lesions and the presence of rhythmic and/or valvular heart disease being predictive factors for recurrence. In the colon, the rate of hemorrhagic recurrence after coagulation with argon plasma treatment ranged from 7% to 15% with a median follow-up of 6 to 20 months.

In our study active bleeding was present in 11% of our patients; thus the use of hemostatic clip was present in only 3.3%.

The endoscopic evolution was favorable in all our patients with a clear improvement of rectal lesions and digestive angiodysplasia lesions. The total complication rate in our series was nil.
On the other hand, the risk of occurrence of radiation rectitis increases above all with the dose of irradiation administered, the threshold dose is 45 grays. The risk is also influenced by the volume irradiated, the mode of radiotherapy (external irradiation or curietherapy), time fractionation and dose spread.\textsuperscript{19–21}

The main symptom of chronic radiation rectitis is rectorragies, but it can also be revealed by mucusy diarrhoea secondary to the chronic irritative phenomenon of the rectal wall, and can also be accompanied by anal incontinence or an anaemic syndrome.\textsuperscript{22}

The diagnosis of radiation rectitis is primarily endoscopic. The appearance of the rectal mucosa is variable. It may present in two ways: either by a mucosa covered with telangiectasia taking on the appearance of more or less regular, dilated and fragile, disseminated neo-vessels (\textsuperscript{\textendash}Fig. 3); or it may be diffusely congestive, friable and hemorrhagic at the slightest contact or spontaneously (\textsuperscript{\textendash}Fig. 4). In extreme cases one may find scattered ulcerations that may be superficial or deepening (\textsuperscript{\textendash}Fig. 5).

Sometimes recto-vaginal or recto-vesical fistulas may occur secondary to the mucosal remodeling of the rays and its healing.

The treatment of choice for both digestive angiodysplasias and radiation rectitis is Argon Plasma Coagulation (APC).\textsuperscript{24} except for asymptomatic non-haemorrhagic AIs, of incidental discovery, where therapeutic abstention is well recommended.\textsuperscript{25} The main goal is to control bleeding and prevent recurrence.

The principle is based on the transformation of the energy produced by a high electric current into thermal energy, resulting in coagulation, debridement and/or desiccation of the tissues.\textsuperscript{26} as the simultaneous release of an electric current and a gas argon, which in its ionized form will destroy, in a targeted and non-contact manner (3 to 5 mm of distance), the superficial mucosa (0.5 to 3 mm deep) by means of an electric arc (\textsuperscript{\textendash}Figs. 6 and 7).

The power of the generator should be adjusted according to the location of the angiodysplasia.\textsuperscript{27} The recommended settings are (expert opinion): in the stomach: 0.8 L/min,
50W; in the esophagus, duodenum, small intestine, and right colon/cecum: 0.6 L/min, 40W; in the left colon: 0.8 to 1 L/min, 50W.

May and al showed in a recent retrospective series a significant reduction in transfusion requirements in 50 patients treated with argon plasma coagulation. However, recurrence of bleeding occurred in 42% of patients after a mean follow-up of 4.5 years. Gerson and al showed a similar rate of hemorrhagic recurrence (45%) 30 months after treatment of angiodysplasias. Other studies involving at least 300 patients have all reported argon plasma coagulation efficacy rates in the treatment of radiation rectitis of more than 80% on bleeding after an average of 1 to 3 sessions (extremes of 1 to 8), with complete or partial cessation of bleeding. Several sessions are often required, with a correlation between the number of sessions and the extent of the lesions to be treated.

However, a minimum delay of two to four weeks between sessions is preferable in order to allow sufficient healing time.

The complication rate of argon plasma coagulation treatment ranges from 1.7% to 7% and includes fever with bacteremia, urinary disorders, ulceration, bleeding from pressure ulcer, rectal stenosis, microrectitis, recto-vaginal fistula, perforation, and intracolic explosion.

Perforation occurs preferentially in the cecum where the wall is thin. Some authors have proposed injecting saline, with or without adrenaline, into the submucosa to elevate the angiodysplasia before argon coagulation.

Finally, exceptional cases of explosions have been reported with the use of argon in ill-prepared patients. It is therefore recommended that patients be prepared orally in order to obtain a perfectly clean colonic lumen.

Conclusion

Endoscopic treatment of digestive vascular abnormalities by argon plasma coagulation gives good results in terms of efficacy, tolerance and safety, which proves its first-line indication in the management of digestive angiodysplasia and chronic haemorrhagic rectitis.

Conflict of Interest

No conflicts of interest

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