

**Review Article** 

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# Discogenic Low Back Pain: A Topical Review



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#### Abstract

**Background:** Discogenic low back pain (LBP) is a significant medical condition that carries a heavy socioeconomic burden. The pathophysiology, diagnosis and treatment modalities are reviewed in this article.

Methods: A review of the English literature addressing discogenic LBP in PubMed until 2015 is undertaken.

**Results:** The various methods of confirming the diagnosis of discogenic pain is elucidated in this article. With regards to the effectiveness of percutaneous treatments, intradiscal methylene blue (MB) and biacuplasty have a weakly positive recommendation. The evidence for intradiscal electrothermal therapy (IDET) is equivocal. Nucleoplasty, intradiscal pulsed radiofrequency (pRF) and rami communicans radiofrequency (RF) have too few studies to allow them to be assessed. Intradiscal RF, steroids, DiscTRODE and L2 block/RF have negative recommendations.

**Conclusion:** More evidence is needed for to allow for a better assessment of the effectiveness of treatment for discogenic pain. Future studies need to adhere to the IASP definition of discogenic pain.

Keywords: Low Back Pain; Intradiscal; Intradiscal Electrothermal Annuloplasty; Radiofrequency; Discogram; Back Pain

### Introduction

Chronic Low back pain (LBP) remains a significant medical problem and is a major cause of disability worldwide. In a study published in 2014, the global age-standardized point prevalence of LBP was 9.4%, with it being higher in men at 10.1% compared with women at 8.7% [1]. The prevalence increases with age, peaking at 80 years old with a prevalence of up to 40% in males and 35% in females. In a retrospective chart review by Depalma et al. [2], the intervertebral disc contributed to 41.8%. The aim of this topical review is to give an overview on the pathophysiology, diagnosis and the evidence for the various treatment modalities for discogenic pain. A search in Pubmed for English articles up to 2015 was used to provide the literature contained in this review.

### **Pathophysiology**

From the rationale that a structure must have a sensory nerve supply for it to be capable of causing pain, the innervation of the intervertebral disc has been extensively studied and documented. Bogduk described the innervation of the lumbar intervertebral disc in 1980 [3]. The anterior and lateral portions of the annulus fibrosis (AF) are supplied by branches of the grey rami communicantes of the sympathetic trunk. The posterior aspect of the AF is supplied by the sinuvertebral nerve, which is

a combination of a branch from the ventral ramus and a branch of the grey ramus communicantes of the corresponding segment.

In a normal disc, only the outer one third of the AF is innervated. However, in patients with discogenic pain, it is found that there is innervation all the way into the inner third of the AF and even into the nucleus pulposus (NP) [4-6]. The tears in the AF, which are commonly in the posterior part, caused the formation of a vascularized granulation tissue starting from the outer part of the AF into the disc. Within this granulation tissue, nerve fibers are found and these abnormal nerves are the likely cause for the nociception of the intervertebral disc [7]. In addition to the abnormal sensory innervation described above, inflammation within the intervertebral disc has also been implicated in the pathogenesis of discogenic pain [8,9].

### **Definition of Discogenic Low Back Pain**

According to the International Association for the Study of Pain (IASP), the definition of lumbar discogenic pain is lumbar spinal pain, with or without referred pain to the lower limb girdle or lower limb, stemming from a lumbar intervertebral disc [10].

The diagnosis hinges on the fact that the pain must be shown to be conclusively from the intervertebral disc. This can be done in 3 ways.

- i. Selective anesthetization of the putatively symptomatic intervertebral disc completely relieves the patient of the accustomed pain.
- ii. Selective anesthetization of the putatively symptomatic intervertebral disc substantially relieves the patient of the accustomed pain for a period consonant with the expected duration of the local anaesthetic used.
- iii. Provocation discography of the putatively symptomatic intervertebral disc reproduces the patient's accustomed pain but provided that provocation of at least 2 adjacent intervertebral discs clearly does not reproduce the patient's pain.

Of note, none of the above criteria is based on MRI findings. Physiological aging of the intervertebral disc will appear abnormal on MRI even though that disc is not symptomatic [11]. As such, an MRI by itself cannot be used to diagnose the symptomatic disc and the above interventional diagnostic procedures are indicated. Though the IASP states that selective anaesthesia of the disc is a valid method of diagnosing discogenic pain, only a few studies attempt to use selective blockade of the disc [12-17] as part of their inclusion criteria.

### **Provocative Discography**

The most commonly used method of diagnosing discogenic LBP hinges on a positive provocative discography. The procedure is briefly outlined and the criteria for a diagnosis of discogenic LBP are highlighted. The Spine Intervention Society (SIS), formerly known as the International Spine Intervention Society (ISIS), recommends the following criteria to diagnose discogenic pain during provocative discography [18].

- i. Provocation of the suspect disc induces concordant pain.
- ii. The pain is at least 7/10 on a numeric rating scale.
- iii. The pain is provoked by less than 50 psi over opening pressure.
- iv. Provocation of at least 1 adjacent disc does not induce concordant pain.

The rate of infusion should not exceed 0.05 ml/s as this allows for uniform pressurization within the disc. If too high an infusion rate is used, false positives may occur because of the pressure peaks. These pressure peaks may cause vertebral end plate compression and distention of the adjacent facet joint [19]. The patient should be blinded as to which disc is being stimulated and the disc which is expected to be most painful should be stimulated last.

During provocative discography, the following events should be noted.

- i. Opening pressure (OP) which is defined as the pressure in which contrast is first seen entering the disc.
- ii. Provocation pressure which is defined as the pressure greater than the OP in which the patient first complains of pain
- iii. The peak pressure or final pressure at the end of the procedure which is greater than the OP.

The procedure is terminated if the following is reached.

- a. Concordant pain is reproduced to a level of 7 out of 10 on a numeric rating scale
- b. The volume injected reaches 3 ml (or up to 4 ml in very degenerated disc in which the pressure remains less than 15psi)
- c. The pressure rises to 50 psi above OP in discs with a grade 3 annular tear
- d. If contrast leaks through the outer annulus fibrosis or through the endplates in which it might not be possible to pressurize the disc sufficiently to test its sensitivity.

### Selective Blockade of the Intervertebral Disc

The first study to use selective blockade of the disc was performed in 1996 [17]. 1.5-2 milliliters of a mixture of two milliliters of 2% lignocaine and 1 ml of contrast media were injected into the disc and pain relief was assessed 30 minutes later. Pain relief of >50% was judged to be a positive test for discogenic LBP. The same group of authors then used the same criteria in a subsequent study in 2001 [14]. The term "Discoblock" was first used by Ohtori [13]. 0.75ml of 0.5% bupivacaine was injected into the intervertebral disc and if pain was reduced thereafter, the diagnosis of discogenic pain was made. This method was then compared to that of conventional provocative discography in patients who underwent surgery based on the diagnosis of discogenic pain obtained from the 2 different diagnostic methods. They found that patients whose diagnosis was obtained from Discoblock were more satisfied with surgery.

2 studies from the same authors on intradiscal pulsed radiofrequency (pRF) used Discoblock in conjunction with provocative discography as part of their inclusion criteria [15,16]. They required a 70% reduction in pain for it to be positive. A modification of this technique is used by Alamin et al. [12] to confirm the diagnosis obtained from provocative discography. In patients who had a positive provocative discography, they inserted a catheter into the nucleus pulposus of the symptomatic disc. They then randomized the patient into either an infiltration of 0.6ml of 4% lignocaine or normal saline via the catheter. The results of the 2nd part of the test were then compared with the

initial provocative discography findings and the results of the functional disc block and provocative discography were different in 46% of the patients. This is the only study that has used this protocol and in the absence of a gold standard diagnostic test, it is difficult to comment on its utility as compared to provocative discography.

#### **Interventional Procedures**

#### Intradiscal Steroids

Steroids were introduced into the disc to decrease the inflammatory reaction in patients with discogenic pain. As early as the 1960s, clinicians were already introducing steroids into the disc [20]. The last 3 studies published were randomized controlled trials (RCT) and these will be reviewed in this article. In 1992, Simmons et al conducted a randomized double blind active control study (Jadad scoring = 3, no mention of method of randomization or blinding) [21]. Either methylprednisolone 80mg (14 patients) or Bupivacaine 7.5mg (11 patients) was injected intradiscally after a positive single-level provocative discogram. A follow-up of only 10-14 days showed that 21% of patients who had methylprednisolone injected versus 9% of bupivacaine reported improvement. There was no statistical superiority in the steroid group.

The RCT by Khot et al. in 2004 (Jadad score = 4, method of blinding inadequate) compared methylprednisolone 40mg against 1ml of normal saline with 60 patients in each group. Only the patient was blinded and follow-up was for 1 year. There was a high loss to FU rate of 14 patients in the steroid group and 8 patients in the saline group. The results of this trial showed no difference between the 2 groups in terms of disability or pain scores. There was a median change of 0 in pain scores in both groups.

Peng et al. [22] published the latest RCT in 2011 in which they looked at patients with positive provocative discography and end plate Modic changes (Jadad score = 4, method of randomization not stated). They divided the group into 2 based on Modic change Type I or II then subsequently randomized them into 3 treatment groups. The first group received intradiscal saline, the 2nd group intradiscal betamethasone, and the 3rd group intradiscal betamethasone and songmeilie (Chinese herbal medicinal ingredient that reduces inflammation). At 6-month follow-up, in both types of end plate Modic changes, the intradiscal betamethasone and intradiscal betamethasone with songmeilie groups had significant drops in VAS and ODI scores. In contrast to the previous RCTs, intradiscal steroids showed some short-term benefit in patients with discogenic pain and end plate inflammatory changes. This has been suggested in another study [23].

Based on the above RCTs, there is a limited role of intradiscal steroids for patients with discogenic LBP. Older observational studies were not included in this review. Based on the results of the RCTs, only in patients with inflammatory end plate changes would intradiscal steroids have a possibility of short-term benefit and this still needs further clarification with more trials.

## Intradiscal Electrothermal Annuloplasty (IDEA)/ Intradiscal Electrothermal Therapy (IDET)

The IDET procedure utilizes a navigable intradiscal catheter (Spine CATH, Oratec Interventions Inc., Menlo Park, CA, USA) with a 6-cm active tip that is introduced through a 17G introducer. The 17G introducer is inserted into the disc in a similar fashion to performing a discography (Table 1). The flexible catheter is then inserted and it directed to make a loop such that the active tip lies against the posterior AF. In which layer the catheter lies exactly is a matter of debate as some studies state it should lie in the NP-AF junction whereas others say that it should lie within the AF itself [24-26]. It is then postulated that the RF current which is applied across the catheter causes heating of the active tip which ablates the aberrant nociceptive nerve endings in the posterior AF.

In the first observational study, Saal et al. [27] reported that 80% of the patients had at least a 2 point reduction on VAS with a mean reduction of 3.74 at 7 months follow-up. There were also improvements in analgesic use, SF-36 scores and sitting tolerance. Multiple prospective studies were then published and selected studies are presented in Table 1. Outcomes at 2 years were also published [25,28] and these were in favour of IDET with 57-72% having a decrease of at least a 2-point decrease in VAS at 2 years. However, several studies showed disappointing results with IDET [29-31] and so there was a need for randomized studies to be performed to better ascertain the utility of IDET.

The first RCT was performed by Pauza et al. [32]. In this well-conducted study, patients either had IDET treatment or a sham procedure. It must be noted that the inclusion criteria for this study required patients to only have a loss of <20% of disc height. The results of the study showed that both the placebo and IDET group improved at 6 months but more marked improvement was seen in the IDET group (56% vs. 38%). IDET was also more effective in patients who had poorer function preprocedure. Even in this select group of patients with relatively preserved disc height and reasonable function, almost 50% did not benefit from IDET.

The results from a  $2^{nd}$  RCT were even less impressive. Freedman et al. [33] conducted a placebo controlled trial in which placebo consisted of inserting the introducer needle and the catheter into treatment position but not heating the catheter. They included patients with up to 50% loss in disc height and patients on worker's compensation. Compared to the Pauza study, the patients had poorer ODI and SF-36 scores. In this study, both the IDET and placebo groups had no improvement at 6 months compared to baseline. It should be noted that in both the RCTs on IDET, the targeted sample size for 80% power were not reached.

Table 1: Intradiscal Electrothermal Therapy (IDET).

Author and publication year of study	Type of study, number of patients, length of follow-up	Inclusion Criteria	Exclusion Criteria	Intervention Performed	Results	Comments
Saal et al. [27]	Prospective. 25 patients. Mean FU of 7 months.	•LBP > 6 months • Failure of conservative management • Failed at least 1 fluoroscopically guided epidural steroid injection • Normal neurological exam • MRI did not show a neural compressive lesion • Positive provocative discography	Inflammatory arthritis     Non-spinal conditions that can mimic lumbar pain     Medical disorder that would preclude FU and participation     Prior surgery at the symptomatic level	Temperature increased to 90°C over duration of 13 minutes and maintained at 90°C for 4 minutes.	80% had reduction in VAS score of at least 2 points 72% had increase in sitting tolerance and reduction of analgesic use.	3 patients treated with oral prednisolone for pain flare within 2 weeks of IDET, with 1 patient also needing an epidural steroid injection. The authors are the developers of the Spine Cath catheter
Derby et al. [86]	Prospective, 32 patients, 12 months FU	LBP > 6 months     LBP > 60% of overall symptoms     Failure of conservative treatment     Normal neurological examination     Positive provocative discography     Declined or not suitable for surgery	Infections     Bleeding diatheses     Unstable medical conditions     Radiculopathy	Up to 2 levels  Final study protocol: 80-90°C, 13.5-16.5 minutes	Outcome assessed on improvements in VAS, Rolland-Morris Disability Questionnaire, and NASS LBP Outcome assessment instrument patient satisfaction index and modified general activity questionnaire.  For a favourable outcome, 3 of the 4 most have improved. Non-favourable would require a decrease in 3 out of the 4 outcomes. The rest assessed as no change.  62.5% had a favourable outcome, 12.5% non favourable and 25% no change.	All patients experienced a flare of their pain after procedure with a mean duration of 5 days.
Karasek et al. [26]	Prospective case-control study. 36 treatment, 17 controls. Controls were patients who were diagnosed with IDD on provocative discography but denied IDET by their insurance companies. 12 months FU	Back pain > 3     months     IDD was     diagnosed by     provocative discography and CT     scan.	• No evidence of disc prolapse, tumour, infection or neurological disease	1-2 levels  90°C for 17 minutes  If patient unable to tolerate the pain of heating, temperature dropped to 85°C.	Mean VAS score of IDET group dropped from 8 to 3 60% had greater than 50% reduction in VAS and returned to work. 23% obtained complete pain relief. No improvement of control group in VAS at 3 months.	Control group only had FU until 3 months Majority of patients had single level treated (30/36) and preserved disc heights.

Saal et al. [87]	Prospective. 62 patients. 16 months mean FU.	•LBP > 6 months •Failure of conservative management •Failed at least 1 fluoroscopically guided epidural steroid injection •Normal neurological exam •MRI did not show a neural compressive lesion •Positive provocative discography	•Inflammatory arthritis •Non-spinal conditions that can mimic lumbar pain •Medical disorder that would preclude FU and participation •Prior surgery at the symptomatic level	30 patients treated at 1 level, 24 patients at 2 levels, and 8 at 3 levels.  Temperature increased to 90°C over duration of 13 minutes and maintained at 90°C for 4 minutes.	Mean drop of 3 on VAS scores  71-74% had improvement in either the SF-36 physical function, SF-36 bodily pain or the VAS scores.  19% had no improvement in any score.	No complications.  Disc space narrowing of at least 30% associated with worse outcome.
Singh [88]  Gertzen et al.  [89]	Prospective. 23 patients. 6 months FU.  Prospective, 27 patients. 1 year FU.	LBP > 6 months	Inflammatory arthritis     Non-spinal conditions that can mimic lumbar pain     Medical disorder that would preclude FU and participation     Evidence of instability on imaging studies     Active infection     Malignancy     Metabolic disorder that would preclude appropriate FU and participation	Up to 2 levels treated.  16.5 minutes with final temperature of 80 to 90°C.  16 patients had treatment at 1 level,  11 patients treated at multiple levels.  Temperature increased to 90°C over 13 minutes and maintained for 4 minutes.	2 lost to FU.  67% had greater than 50% pain relief.  Statistically significant increase in standing and walking time. The improvement in sitting did not attain statistical significance.  Pain scores not collected as part of study.  47% had significant improvement in SF-36 scores.  75% had improvement in ODI scores.	No complications. No complications reported.  Failed IDET occurred in 1 patient as the catheter could not be inserted likely because of scar tissue from previous discectomy.
Saal et al. [28]	Longer follow- up study of the prospective trial done above [87]. Mean FU of 28 months.	• As stated above	• As stated above	As stated above	4 of the initial 62 lost to FU at >24 months. 58 patients assessed.  72% had at least a 2-point improvement in VAS, 50% had at least 4 points.  78% had at least a 7-point improvement in SF bodily pain subscale.  Improvements in mean VAS and SF-36 bodily pain subscale sustained at 24 months.	No complications

Spruit et al. [31]	Prospective. 20 patients. 6 months FU.	Degenerative disc disease     Affected level L1-S1     Predominant low back pain     Intolerance for sitting     Neurological examination normal     Conservative treatment applied for at least 6 months and failed.     Positive provocative discography	Spondylolysis or spondylolisthesis     Infection     Active malignancy     Pregnancy     Previous lumbar surgery	16 patients treated at 1 level, 4 patients at 2 levels  Temperature slowly increased to 90°C then maintained for 4 minutes	1 patient lost to FU.  Drop of 14mm on VAS (p=0.046) but much variation noted between patients.  ODI did not improve significantly.  SF36 vitality and bodily pain subscales improved significantly but not others.	No complications reported.
Bogduk et al. [25]	Longer term outcomes of the study [26] mentioned above.	• As above	• As above	• As above	At 24 months, the control group was not significantly better as compared to baseline.  At 24 months, 57% of patients had at least 50% pain relief and 23% had complete pain relief.  Results at 6 months FU is stable as 12 and 24 months FU in the majority of patients.	Interestingly, though the initial study stated that the control group was lost to FU after 3 months, this study managed to get results from them at both 12 months and 24 months.
Freed man et al. [30]	Prospective. 36 patients who are active military personnel. Mean FU of 29.7 months.	Failed 6 months of non-operative treatment of chronic LBP     Positive provocative discography     MRI showing no nerve compression, tumour/infection or osteoligamentous trauma     Absence of prominent radicular symptoms and signs	Non-active duty status Previous lumbar fusion history Severe spinal stenosis Spondylolisthesis, inflammatory arthritis or neoplastic disease Predominant radicular pain or positive neurological findings Medical or metabolic condition that would preclude appropriate FU and participation.	IDET protocol not mentioned.  17 patients had 1 level treated. 14 patients had 2 levels treated.	5 patients lost at >6 months FU.  47% of 36 patients had >50% reduction at 6 months FU but only 16% at final FU.  52% had a 2 point or greater decrease in NRS.  29% stated they had improved at final FU and only 16% said they were satisfied with IDET.	Additional 5 patients were treated that was not in statistical analysis. 3 had another IDET procedure done at another setting at an adjacent disc after good success from first IDET. 2 had a repeat IDET done at the same level due to poor result.  5 complications (16%): Foot drop, increased disc herniation, decreased sphincter tone and faecal incontinence and increased or new nondermatomal leg pain x 2.

Cohen et al. [37]	Retrospective. 79 patients. 6 month FU.	• LBP > 6 months • Failure of conservative therapy • Positive provocative discography • Absence of radicular symptoms	• Age >60yo • Severe spinal stenosis • <50% disc height • Significant spondylolisthesis, inflammatory arthritis • Unstable medical conditions	1 disc treated in 50 patients, 2 discs in 29 patients.  Catheter head to 90°C over 16.5 minutes.	48% had >50% reduction at 6 months FU.  Drop in NRS from 5.9 to 2.1 in successful patients. Unsuccessful patients decreased from 6.2 to 5.1.	In 11 cases in which the catheter could not adequately cover the posterior annulus, the procedure was performed from the other side.  10% complication rate with obesity being a risk factor.
Lee et al. [90]	Prospective. 62 patients. Average FU of 34 months.	Moderate to severe LBP > 6 months     Sitting>standing pain     Normal neuro examination     Failure of conservative care     MRI/CT showing no nerve compression     Positive provocative discogram and IDD on CT	• Disc space narrowing >50% • Disc extrusion or sequestrated segment • Severe spinal stenosis • Spondylolisthesis > grade 1 • Segmental instability	32 patients had a single level treated. 19 patients had 2 levels treated.  Temperature increased gradually to 90°C over 12.5 minutes and maintained for another 4 minutes.	11 patients lost by final follow-up.  Back pain visual numerical scale (VNS) improvement of 3.2 (p<0.001).  53% demonstrated clinically significant VNS and RM improvements.	2 out of 51% underwent fusion which was also unsuccessful. Nucleoplasty was done in 2 patients and RF of the medial branches was done in 3 patients. 2 patients underwent repeat IDET.
Davis et al. [29]	Retrospective. 60 patients. Mean FU of 20.4 months.	IDD with an annular fissure     Contained disc herniation     Positive provocative discography     Disc height >50%     Symptoms >6 months     Failed conservative treatment	Stenosis     Disc herniation or sequestrated disc     Neural compression on MRI     Previous lumbar surgery     Psychological issues     Segmental instability	Not stated.	44 patients responded (73%)  14% underwent a post- IDET lumbar surgery within 1 year with another 4 in the 2nd year.  97% still had pain after IDET at 1 year though 39% had less pain. No change in .disability pre and post -IDET.  37% were satisfied with procedure.	1 report of discitis and 1 report of anterolisthesis.

Pauza et al. [32]	RCT of IDET vs. sham placebo. 6 months FU. 37 IDET, 27 placebo	• 18-65yo • LBP > leg pain for > 6 months • Failure to improve after 6 weeks of conservative management • LBP exacerbated by sitting/standing and relieved by lying down • < 20 on Beck Depression scale • No surgical intervention within last 3 months • < 20% disc height loss • Positive provocative discography and posterior annular tear on CT discogram.	Previous lumbar surgery Abnormal neurological exam Radicular pain Structural deformities Vertebral canal stenosis or scoliosis Herniations >4mm or sequestrated herniations Concomitant cervical or thoracic pain >2 on VAS Uncontrolled medical conditions Injury litigation Disability remuneration	15 patients were treated at 2 levels, 22 at 1 level.  Catheter heated to 90°C over 13 minutes and maintained for another 4 minutes.	56% of IDET and 38% of placebo had >2 points improvement in VAS. Only 40% of patients with IDET had >50% reduction in pain vs. 33% in placebo.  Both groups had significant improvements in pain scores but the IDET group had statistically greater improvements.  Mean change in VAS is 2.4.  IDET group had better outcomes in ODI.  NNT for IDET for >75% relief = 5.	Jadad scoring = 5  Per Protocol analysis.  Moderate loss to FU rate with the results of 86% in IDET and 89% in placebo analyzed.  Study initially wanted to have 40 patients in IDET and 27 patients in placebo but this was not obtained. Power dropped from 80 to 60%.
Freeman et al. [33]	RCT. IDET vs. Sham Placebo. 57 patients in total. 6 months FU. 38 IDET and 19 to placebo.	• >18yo • Single or 2 level disease • Degenerative lumbar disc disease >3 months • Failure of conservative management of 6 weeks • Marked functional limitation • Sitting intolerance > standing • Predominantly LBP rather than leg pain • MRI shows disc disease • No evidence of nerve compression	Large herniation or sequestrated disc     >50 disc height loss     Severely disrupted disc     Spinal stenosis     >2 symptomatic levels     Previous lumbar surgery     Spondylolisthesis at symptomatic level          Psychological conditions          Medical conditions that would affect follow-up          Current injury litigation	Temperature increased to 90°C over 12.5 minutes and maintained for 4 minutes.	Successful outcome is defined as no neurological deficit, improvement of at least 7 points in Low Back Pain Outcome score and improvement of 1 standard deviation from mean in SF-bodily pain and physical functioning subscales at 6 months.  No patient in either group achieved a successful outcome.  No difference between the IDET and placebo group in primary and secondary outcomes.	Jadad Scoring: 4 (method of randomization not stated)  Target sample size was 50 IDET and 25 Placebo for 80% power.  Compared to Pauza's study, the patients were more disabled at baseline.  Pain scores not an outcome measure.  50% of patients in both groups on Worker's Compensation
Maurer et al. [91]	Prospective. 56 patients. Mean FU of 20.5 months.	LBP and impaired function > 6 months     <50% height loss	• Severe disc degeneration • Extruded or sequestrated herniation • Previous lumbar surgery • Chronic lower limb radiculopathy • Spinal canal stenosis >30% • Spondylolisthesis	30 patients treated at 1 level, 26 at 2 levels  Standardized heating procedure used until 60-65°C detected adjacent to the coil.	16% receiving worker's compensation 75% treatment success. VAS improved from 6.1 to 2.4 at final FU. Sitting, standing and walking tolerance all had statistically significant improvements.	No complications noted.

Nunley et al. [92]	Prospective. 53 patients with worker's compensation. 12 months FU.	LBP > 6 months         Failure of         conservative         management         Exclusion of         radiculopathy,         pseudo-         radiculopathy, facet         disease         Positive         provocative         discography	Prior spine surgery Abnormal neurological exam Structural deformities Spinal stenosis Loss of >60% disc height	38% had 1 level treated, 58% 2 levels and 4 % had 3 levels treated.  Temperature raised to 90-95°C and maintained for 5 minutes.	Significant reduction in both VAS (62.6%) and Oswestry scores (69.3%) Significant increase in economic productivity post-IDET.	1 complication stated but not elaborated.
Assietti et al. [93]	Prospective. 50 patients. 24 months FU.	LBP > 6 months	Extruded or sequestrated disc     Previous lumbar surgery at affected level     IDET performed within last 6 months     Nerve root impingement     Spinal stenosis     Spondylolisthesis     Cervical degenerated discs     Major psychological impairment	Temperature increased to 90°C over 12.5 minutes and maintained for 4 minutes.	78% success rate at 24 months.  Pain score improved from 7.6 (baseline) to 3.0 (12 months) and 2.4 (24 months.  ODI improved from 59.0 (baseline) to 27.0 (12 months) and 20.1 (24 months)  9 patients still had severe back pain at 6 months FU and underwent surgery (not factored into results pain score and ODI)	No complications.  Patients who have concordant pain at low pressures on discography or who had HIZ on MRI had better outcomes.

Complications stated in the literature includes cauda equina syndrome [30,34,35], increased disc herniation [30,36,37], vertebral osteonecrosis [38,39], a broken catheter that migrated intradurally causing radiculopathy [40], nerve root injury [30,37] anterolisthesis and discitism [29]. Though severe complications can occur, the overall rate of complications is low. Based on multiple observational studies that show positive results, an RCT that showed fair results in selected patients and a RCT that showed a negative result, there is still clinical equipoise over this modality.

### **DiscTRODE**

The discTRODE (Radionics, Burlington, MA, USA) is also based on a similar working principle as the IDET. First, a 17G curved and electrically insulated introducer needle is inserted into outer annulus from the contra-lateral side to the annular tear. Electrical impedance is checked and the needle is further advanced in the annulus until an impedance of  $300\text{-}400\Omega$  is obtained, which usually corresponds to that of a mid-annulus depth. The catheter is then inserted and it is navigated through the posterior annulus and upon contacting the lateral wall of the annulus, it travels anteriorly. As such, the electrode covers the posterolateral portion of the AF. The catheter is then heated to a maximum temperature of  $65^{\circ}\text{C}$  for 10 minutes.

The first published study on DiscTRODE was in 2005 by Finch et al. [41]. 31 patients underwent the procedure and another 15 who were refused funding became the control group. At 12-month follow-up, VAS scores had dropped 37% in the treated

group with a 3% increase in the control group. 9 out of 31 patients had >50% drop in VAS scores. The ODI decreased significantly over the same period in treated patients. Interestingly, though VAS and ODI scores dropped in the treatment group, medication use did not decrease accordingly.

A comparative non-randomized study between IDET and DiscTRODE was published in 2005 by Kapural et al. [42]. 21 patients underwent the DiscTRODE procedure, in which the heating protocol consists of 55°C for 4 minutes, 60°C for 5 minutes then 65°C for 5 minutes. Another 28 patients underwent IDET and the patients in both groups were matched against each other. Patients in the DiscTRODE group had a decrease in VAS from 6.6 to 4.4 at 1-year follow-up but patients in the IDET group had a larger drop of 7.4 to 1.4. Though DiscTRODE had a significant drop, it was much smaller compared to IDET. 81% of patients in the IDET group had a 60% improvement in VAS as compared to 29% in the DiscTRODE group. As such, based on the results of this study, DiscTRODE is inferior to IDET for the treatment of discogenic LBP.

A well conducted (Jadad score = 5) randomized, double blinded, placebo controlled trial was then conducted by Kvarstein et al. [43]. The placebo was a sham procedure in which the introducer needle and electrode were both inserted into the appropriate position but the RF current was not delivered. In the treatment group, incremental heating started at  $50^{\circ}$ C and this was increased by  $5^{\circ}$ C every 2 minutes. At  $65^{\circ}$ C, the heating was maintained for 4 minutes. The authors intended to have 25 patients in each group but planned an interim analysis

when 10 patients in each group had reached 6-month follow-up. Unfortunately, the interim analysis did not show any significant benefit to the patients who underwent DiscTRODE and so the study was discontinued. As such, this study did not attain the predetermined sample size and analysis of this study was based only on the 20 patients recruited. The VAS scores between the sham and treated group were not different. The treatment group had only a slight reduction of 1 compared to 0 in the sham group. There were more patients in the sham group who reported a 2 or greater decrease in VAS. In terms of complications, there is a case report of Complex Regional Pain Syndrome after the DiscTRODE procedure [44] but nothing else can be found in the literature.

As such, based on a well-conducted underpowered RCT which showed no significant difference between DiscTRODE and sham procedure, a comparative study that showed inferiority

evidence is not supportive of  $\ensuremath{\mathsf{DiscTRODE}}$  for the treatment of discogenic pain.

of DiscTRODE to IDET and a positive prospective study, the

# Intradiscal Radiofrequency (RF)/Percutaneous intradiscal radiofrequency thermocoagulation (PIRFT)

Intradiscal RF first appeared in the literature in 1996. The hypothesis for its effectiveness in treating discogenic pain was that the heat produced would destroy the nerves in the AF. The intervertebral disc has poor internal circulation and the vertebral endplates act as insulators to trap the heat generated by the RF current within the disc [45]. The procedure consists of inserting a cannula with a 10mm active tip into the centre of the NP. An RF electrode is then inserted and a lesion is made. The studies are presented in Table 2.

Table 2: Intradiscal Radiofrequency.

Author and publication year of study	Type of study, number of patients, length of FU	Inclusion Criteria	Exclusion Criteria	Intervention performed	Results	Comments
Van Kleef et al. [17]	Prospective, 39 patients.  Mean FU of 16 months	Age 25 60yo     LBPwith or without sciatica for >12 months     Failed conservative management     Positive anaesthetic discography	Herniated disc with nerve root compression     Spinal stenosis     Extensive multi-level spondylosis     Previous spinal fusion     Psychological problems     Positive lumbar facet blocks	1 level 20G C15 cannula with 10mm active tip 70°C for 90 seconds	54% reported adequate reduction of pain at 8 weeks, 41% at long-term FU  Better results seen in patients without previous disc surgery  Repeat MRI after the procedure in 10 patients did not reveal disc degeneration attributable to procedure	No complications or adverse events
Barendse et al. [14]	RCT, control group had needle inserted into the disc but no RF current delivered. 28 patients. 1 year FU.	Age 30-65yo     LBP with or without sciatica for >12 months     Failed conservative management     Positive anaesthetic discography	Spinal stenosis     Extensive multi-level spondylosis     Previous spinal fusion     Psychological problems     Positive lumbar facet blocks     Pregnancy     Coagulation disturbances      VAS score less than 5     Diabetes mellitus     >1 pain syndrome	Single level  20G C15 cannula with 10mm active tip  70°C for 90 seconds	2-point reduction in VAS and >50% reduction on global perceived effect considered a success  Only 1 out of 13 patients in the treatment group and 2 out of 15 patients in the control group had success at 8 weeks. At 1 year, only 1 patient in the treatment group still had success.  No complications	Jadad scoring: 5  Sample size calculation not mentioned so unable to determine if study was adequately powered.  This study showed no difference between control or intradiscal RF group.

Ercelen et al. [46]	RCT, 2 different durations of intradiscal RF compared. Group A: 20 patients Group B: 19 patients 6 months FU	Chronic low back pain for >2 years     Failed conservative therapy     Positive manual provocative discography	Spinal stenosis     Spinal instability     Spondylolisthesis     Diabetes mellitus     Tumor infiltration     Coagulation disorders     Clinical radiculopathy     Other neurologic abnormalities     Systemic inflammatory disease	20G RFK C15 cannula with 10mm active tip Group A: 80°C for 120 seconds Group B: 80°C for 360 seconds	No difference between Groups A or B.  Patients in both groups had improvement in VAS and ODI scores compared to pretreatment till 1 month but then returned to baseline by 6 months 1 patient had discitis.	Jadad scoring: 3 (no mention of blinding) Sample size calculation not performed. 1 patient lost to FU.
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The first single cohort prospective observational trial was published by van Kleef et al. [17]. After positive anesthetic discography, the patients had RF of the intervertebral disc to achieve a temperature of 70°C for 90 seconds. This study showed positive results with 70% of patients having pain reduction at 8 weeks and 55% on longer follow up (mean of 16 months) if they have not had previous surgery on the affected disc. The results for patients who have had surgery on the affected intervertebral disc were worse with only 37% of patients having pain relief at 8 weeks and 27% on longer follow-up. A good quality RCT was then conducted based on the positive results from the previous prospective study [14]. Barendse et al. [14] randomized 28 patients with positive anaesthetic discography into a treatment group which used the same settings for RF as above, and a control group which had a needle inserted into the intervertebral disc but no RF applied. In contrast to the previous study, only 2 patients in the control group and 1 patient in the treatment group had a positive response at 8 weeks. By 1 year, only 1 patient in the treatment group still had a successful outcome. This RCT showed no difference between the treatment group and the control group.

Another group conducted a RCT of moderate quality in 2003. Ercelen et al. [46] randomized 39 patients into 2 groups, the first of which had RF delivered to obtain 80°C for 120 seconds and the second group had it maintained for 360 seconds. At 6-month follow-up there was no difference between the 2 groups. Both the groups had significant pain reduction up to 1-month post procedure but they returned to pre-treatment levels by 6

months. With regards to complications, there was 1 case of discitis reported in the trial by Ercelen et al. [46].

In summary, there are 2 RCTs which showed that intradiscal RF has no long-term utility in the treatment of discogenic pain. Both the RCTs did not show sample size calculation and so it is uncertain if the lack of efficacy of intra-discal RF is due to a lack of power in the trials or that it is truly ineffective. It is unlikely that any more trials would be conducted for this modality as results from both the RCTs were disappointing. In addition, IDET and biacuplasty delivers a targeted thermocoagulation of the posterior AF as compared to intradiscal RF which should theoretically produce better results. Based on current evidence, there is no role for intradiscal RF.

### Intradiscal Pulsed Radio Frequency (pRF)

pRF has been gaining popularity in recent years and several studies has used it intradiscally for discogenic pain (Table 3). The highest temperature attained with pRF is limited to 42°C and so thermal destruction of the aberrant nociceptive nerves in the disc does not occur with pRF. The most likely pathway for the therapeutic effect is that the electric field generated induces changes in the nerves targeted and so generate a long term analgesic effect [47]. The technique of inserting the cannula is similar to that for provocative discography. An RF electrode is then inserted and the current delivered. Teixeira et al. [48] was the first to publish an article on the use of pRF intradiscally. There were significant decreases in NRS in the 8 patients in his study and at least a 4-point drop in NRS at 3 months.

Table 3: Intradiscal Pulsed Radiofrequency (pRF).

Author and publication year of study	Type of study, number of patients, length of follow-up	Inclusion Criteria	Exclusion Criteria	Intervention performed	Results	Comments
Teixeira et al. [48]	Prospective, 8 patients, median FU of 9 months	LBP > 6 months     Normal neurological exam     Negative medial branch block     Single level discogenic pain diagnosed by manually controlled low pressure discography	Not described	20G SMK C15 cannula 15mm active tip 2Hz 20ms 60V for 20 minutes	Significant fall (p<0.0001) in NRS in the 1st week and this is also seen by the 3rd month (p<0.0001)  All patients had a fall of at least 4 points in NRS by 3 months.  Follow-up data of 5 patients at a mean of 12.8 months showed 4 were pain free, 1 has an NRS of 2.	First study to use pRF intradiscally.
Fukui et al. [16]	Prospective, 16 patients in the IDET group (Apr 2003 to March 2009), 15 patients in the intradiscal pRF group (April 2009 to March 2011), 6 months FU	•LBP >6 months     • Failed conservative therapy including epidural steroid     • Normal neurological exam     • Negative SLR     • MRI did not show a nerve compressive lesion     • Concordant pain at low pressurization during discography     • Positive discoblock with 1ml of 2% lignocaine	Disc sequestration or extrusion     Canal stenosis     Psychological issues     Previous lumbar surgery     History of opioid abuse      Chronic lower limb radiculopathy     Systemic infection or localized infection over needle entry points	Diskit II needle. 20G, 20mm active tip 5Hz 5ms 60V 15 min max temp 40°C	IDET: NRS decreased from 7.5 (pre) to 4.6 (1 month) to 1.7 (6 months) (p<0.01)  RMDQ scores: 10.4 (pre) to 8.9 (1 month) to 2.8 (6 month) (p<0.01)  pRF: NRS decreased from 7.2 (pre) to 3.4 (1 month) to 2.5 (6 month) (p<0.01). RMDQ scores: 10.8 (pre) to 3.5 (1 month) to 2.3 (6 month) (p<0.01)  No complications	87.5% of IDET patients complained of flare-up pain after procedure that lasted 1-8 weeks. Discoblock was used as part of diagnostic criteria.
Jung YJ et al. [49]	Prospective, 26 patients, 12 months FU	LBP >6 months with VAS>4 and ODI>30%     Pain provoked on prolonged sitting     Normal neurological examination     Concordant pain by low pressure automated discography	50% decrease in height of disc  • Extruded or sequestrated disc  • Previous back surgery  • Chronic lower limb radiculopathy  • Spinal canal stenosis  • Instability  • Psychiatric disease	SMK C15 cannula, 20 G 15mm active tip 2Hz, 20ms 60V for 20 minutes	VAS: 6.4 (pre) to 4.1 (1 month) to 4.4 (12 months) (p<0.01)  ODI: 47.3(pre) to 33.7 (1 month) to 36.7(12 months) (p<0.05)  35% had >50% reduction, 7% had improvement of 2 in the VAS but <50%, 58% no improvement  Sitting tolerance: 27.8 min(pre) to 70.8 (1 month) to 71.5 (12 months)  No complications	pRF done within 1 week of discography Though drop in VAS and ODI were statistically significant but the absolute improvements are small. 12 patients had 1 disc treated, 12 at 2 levels and 2 at 3 levels.

Rohof et al. [50]	Prospective, 76 patients, FU 12 months	LBP >6 months     refractory to     conservative     management     >18 years old     Manual positive     provocative discography	Conditions like tumour, infection, fracture and nerve compression	Diskit needle. 20G with 20 mm active tip 2Hz 10ms 60V for 15 minutes	3 month FU: 28.9% no effect, 30% had improvement of at least 2 NRS but <50% reduction, 38% had >50% improvement, 2.6% had surgery  12 month FU: 26.3% no effect, 14.4% improvement of 2 on NRS but <50% reduction, 56.5% had >50% reduction, 2.6% surgery No complications	pRF done immediately after discography. After 3 months, if results are unsatisfactory, an additional pain procedure may have been done.
Fukui et al. [15]	Prospective, 23 patients, FU 12 months	LBP >6 months      Failed conservative therapy including epidural steroid      Normal neurological exam      Negative SLR      MRI did not show a nerve compressive lesion      Concordant pain at low pressurization during discography + positive discoblock with 1ml of 2% lignocaine	Disc sequestration or extrusion     Canal stenosis     Psychological issues     Previous lumbar surgery     History of opioid abuse     Chronic lower limb radiculopathy     Systemic infection or localized infection over needle entry points	Diskit II needle. 20G, 20mm active tip 5Hz 5ms 60V 15 min, max temp 40°C	NRS dropped from 7.47 down to 3.47 at 1 month FU and sustained at 3.13 at 12 month FU (p<0.01) Rolland-Morris Disability Questionnaire (RMDQ) scores improved from 11.4 to 5 at 1 month and 2.90 at 12 months (p<0.01) 4 out of 23 had no improvements. 4 out of 23 had a good outcome.  No complications reported	One of the few studies that used discoblock in addition to provocative discography for inclusion criteria  3 patients had 2 discs treated Only 4 out of 23 had a pain reduction >50% at 12 months

Jung YJ et al. [49] then used intradiscal pRF in 26 patients [49]. More than half the patients had multiple levels treated. There were significant but small drops in VAS and ODI at 12-month follow-up. Only 35% of the patients had a reduction of >50% in VAS at 12 months though. A relatively large prospective study of 76 patients was done by Rohof [50]. At the 3-month follow-up after intradiscal pRF, patients whose pain reduction was <50% may have additional diagnostic and therapeutic interventions like pRF of the dorsal root ganglion or a RF ablation of the medial branches of the spinal nerves. At the 12-month follow-up, 56.5% had a reduction of >50% in NRS and this includes the group that just had intradiscal pRF alone and those in whom multiple procedures had been done. For the patients who only had intradiscal pRF, only 30% had a successful outcome at 1 year.

A comparative study of IDET against intradiscal pRF was then done by Fukui et al. [16]. The first group of patients underwent IDET from 2003 to 2009, then the 2nd group of patients underwent intradiscal pRF from 2009 to 2011. The inclusion and exclusion criteria for the 2 groups were the same. This study showed that pRF was comparable to IDET with significant decreases in NRS and in the Roland-Morris Disability Questionnaires. The same group of authors then performed intradiscal pRF for a prospective study of 23 patients [15]. 82.6% had at least a drop of 2 in the NRS, with 65.2% having >50% reduction at 12-month follow-up.

In summary, there is limited evidence for the efficacy of intradiscal pRF. Only prospective observational studies have been done and the results differ substantially between studies. More studies are needed and at the current time intradiscal pRF cannot be routinely recommended.

### Rami Communicans Radiofrequency Ablation

There has only been 1 study published for this technique and the aim of the technique is to disrupt the nociceptive input from the AF by creating a thermal lesion in the rami communicans. In 2004, Oh et al. [51] published a RCT (Jadad score = 2) comparing radiofrequency ablation of the rami communicans above and below to the pathological disc versus a lignocaine 2% infiltration. No blinding or method of randomization was mentioned in the article. The study group consisted of patients who had a poor outcome following IDET at a single disc after a positive provocative discography. A diagnostic block of the rami communicans is first performed. If pain decreases >50%, the RF procedure is repeated one day later. A curved needle with a 10mm active tip is directed to the inferior third of the vertebral body. The needle should then lie in the postero-lateral 1/3 of the vertebral body where the rami communicans runs. The control group receives 2 millilitres of 1% lignocaine but the treatment group gets RF thermocoagulation at 65°C for 60 seconds. 26 patients were in the treatment group and 23 patients in the control group. All patients completed 4 months of follow-up.

The patients in the treatment group had significantly better improvements in the VAS and SF-36 scores. Mean VAS dropped from 7.1 to 3.8 in the treatment group.

Though the results from this RCT were positive, it was not well conducted and the inclusion criteria of patients who failed IDET are not generalizable to all patients with discogenic pain.

### L2 Selective Blockade and Radiofrequency Ablation

Another possible route of nociceptive transmission of a painful intervertebral disc back to the spinal cord is via the lumbar sympathetic chain. It is postulated that nociceptive fibers from the AF of the lower lumbar intervertebral discs run through the rami communicans to the lumbar sympathetic chain. It then ascends the sympathetic chain, travels via the L2 rami communicans which then joins the ventral root of the L2 spinal nerve before reaching the spinal cord. Nakamura et al. [52] were the first group to investigate this pathway in 1996. 33 patients with LBP of at least 1 month and diagnosed with discogenic pain based only on physical examination, plain radiographs and MRI had a unilateral L2 nerve block with 1.5ml of 1.5% lignocaine. After 15 minutes, VAS scores (max of 20 points) and pain provocation were assessed. 26 patients had complete relief of ipsilateral pain and the remaining 7 had some relief. Mean VAS scores dropped from 10 to 1.7. The average duration of pain relief was 20.7 days.

Simopoulos et al. [53] then reported on a case series of 5 patients who underwent RF of the L2 rami communicans with good results. Patients had to have both a positive provocative discography and greater than 50% pain relief from diagnostic L2 rami communicans block to be considered for a RF treatment. This consisted of reaching a temperature of  $80^{\circ}\text{C}$  for one minute. There was improvement in the VAS scores, sitting tolerance and medication use. However, the pain relief only lasted for about 4 months following which the procedure had to be repeated to maintain the pain relief.

A negative prospective cohort study was then published by Richardson et al. [54]. Following positive manual provocative discography, 12 patients underwent bilateral L1 and L2 dorsal root ganglion (DRG) blocks using methylprednisolone 80mg,

clonidine 75 mcg and 4 ml of 0.5% bupivacaine in total. After 1 month, there were no differences in the pain scores and pain interference scores regarding daily activities pre- and post-procedure. The study was thus terminated. No complications were reported in any of the above studies. Conflicting results are seen in the studies on L2 nerve blockade and the numbers of patients are small in each study. It is uncertain if there is any utility in treating discogenic LBP with a L2 nerve root intervention. It is likely that the sympathetic chain is not the main route for the transmission of discogenic pain and that the traditional route via the cauda equina contains most of the nociceptive pathways. As such, there is no role for it in routine practice currently.

### **Nucleoplasty**

The Spine Wand (Arthrocare Inc., Sunnyvale, CA, USA) utilizes coblation technology to decrease intradiscal pressure by removing a small amount of volume from the NP [55]. The coblation is meant to result in a decrease of about 1 millilitre of NP volume and thus causing a subsequent large fall in intradiscal pressure. Though it was initially used for radicular pain secondary to a herniated disc, it was also utilized in patients with discogenic LBP.

Only observational studies are found for the use of nucleoplasty in discogenic LBP (Table 4). Sharps et al. [55] first published this in 2002 where they had positive results up to 1 year in duration. Singh et al then published 3 prospective studies where all showed positive results with about 50% of patients obtaining greater than 50% pain relief at 1 year [56-58]. Reddy et al. [59] published a retrospective analysis of 49 patients with a mix of axial and radicular pain. The criteria for inclusion for axial back pain seemed to suggest that a positive provocative discography is not a requirement. As such, this study was not included in the Table. Similarly, the study by Masala et al. [60] had similar problems in their study and was not included in the table. Both these studies though had positive results. Another study by Kumar et al. [61] had positive results but concordant pain during discography was not an inclusion criteria though it was performed prior to nucleoplasty. They found that concordant pain had no predictive effect on the success of the procedure.

Table 4: Nucleoplasty.

Author and publication year of study	Type of study, number of patients. Length of FU.	Inclusion Criteria	Exclusion Criteria	Intervention Performed	Results	Comments
Sharps et al. [55]	Prospective, 49 patients with 1 year FU	LBP with/without radicular pain     Failure of 6 weeks of conservative treatment + epidural steroid injections     Positive provocative discography only in patients without radicular pain	Sequestrated herniation  Herniation > 1/3 spinal canal  Spinal stenosis  Presence of progressive neurological deficits  Tumour, infection, spinal fracture	17G introducer needle, 6 channels created with Spine Wand	VAS scores dropped from 7.9 (pre-) to 3.6 (1 month), 3.1 (3 months), 3.2 (6 months) and 4.3 (12 months) 79% success rate	1 patient lost to FU and only 13 patients have 12 month FU data.  No complications  Provocative discography not done in patients with LBP and radicular pain.

Singh et al. [58]	Prospective, 67 patients with 12 months FU	Contained disc herniation with axial ±leg pain > 3months     No neurological deficit     Failure of conservative management and injection therapies     Positive provocative discography	• Litigation • Heavy opioid usage • Uncontrolled psychological disorders • Sequestrated disc • Herniation >1/3 spinal canal • Infection • Spinal instability • Spinal stenosis	As above	56% of patients reported >50% pain relief at 12 months 59% of patients had >2 point reduction in VAS at 12 months.  Improvement in sitting, standing and walking tolerances.	No data available for 20 patients at 12 month FU as not contactable, in addition to 6 patients lost to FU.  No complications
Singh et al. [57]	Prospective, 80 patients with 1 year FU.	As above	As above	As above	75% had decrease in numeric pain scores at 12 months with a statistically significant reduction of 2.43.  54% of patients had >50% pain relief.	11 patients' data not included at 12 months. No complications
Singh et al. [56]	Prospective, 47 patients with 1 year FU.	LBP with failure of conservative therapy >3 months      Positive provocative discography     No neurological deficit      Average pain score of at least 5	As above	As above	53% had >50% pain relief at 12 months.  Improvements in the ability to sit, stand and walk at 12 months.	Data for 37 patients at 1 year FU. No complications
Yakovlev et al. [94]	Retrospective, 22 patients with 1 year FU	Axial/radicular pain >6 months     Failed conservative management     <50% disc height loss     No neurological deficit     Positive provocative discography     Contained disc protrusion on MRI	Infection     Spinal tumour/facture     > 2 symptomatic levels     Previous surgery at affected level     Psychological disorders	As above	68.2% of patients had >50% pain relief at 12 months.  Mean decrease of 3.98 points in VAS scores at 12 months  72.7% of patients had >50% daily opioid usage reduction at 12 months  Significant improvement in functional status at 12 months	

Zhu et al. [95]	Retrospective, 42 patients with 2 year FU	Back pain ± leg pain Failure of conservative therapy for 6 months  > 50% disc height Positive provocative discography Contained disc protrusion on MRI	<ul> <li>Spinal stenosis</li> <li>Disc protrusion &gt; 1/3 spinal canal</li> <li>Previous surgery</li> <li>Severe neurological deficits</li> <li>Spinal tumour</li> <li>Infections</li> </ul>	As above.  In addition, patients with radicular symptoms also received a nerve root injection of betamethasone and lidocaine	VAS scores for back pain improved from 7.7 (pre-) to 4.2 (2 year FU) (p<0.05)  VAS scores also significantly improved for leg pain though not for numbness at 2 year FU.  ODI dropped form 68.2 (pre-) to 39.4 (2 year)	Method of discography did not mention negative control discs Patients with predominantly radicular pain included.
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Lastly, He et al. [62] published an interesting study in which they modified the technique of nucleoplasty and named it coblation annuloplasty. Instead of creating channels in the NP, they purposely left the introducer needle at the edge of the outer annulus, and created 6 channels in the annulus instead. The selection criteria were good as only patients with discogenic LBP confirmed by provocative discography was included. In 17 patients, the mean VAS scores dropped from 6.5 to 3.2 at 6-month follow-up. In addition, 58.8% of patients had greater than 50% pain relief.

The complications of nucleoplasty have been presented in another article [63] and it includes increased back pain, increased radicular pain, new onset transient lower limb neurological symptoms and epidural fibrosis. No complications were reported in the articles for discogenic pain. In summary, based only on observational studies in which some did not diagnose discogenic pain per IASP criteria before the procedure, nucleoplasty showed a consistently positive result. Based on the current literature, it is not possible to recommend this technique for routine clinical practice.

### **Intradiscal Biacuplasty**

Intradiscal biacuplasty is essentially a cooled radiofrequency ablation of the posterior annulus. Two 17G introducers are inserted into the posterior annulus on both sides of the disc using fluoroscopic guidance. The electrodes, which are 18G in size with a 6mm active tip, are then inserted through the introducers and these electrodes are cooled internally using circulating water. A bipolar current is then created between the 2 electrodes, thus creating a strip lesion between and around the electrodes, ablating the posterior and posterolateral annulus. The aim of biacuplasty is to destroy the abnormal nociceptors that are responsible for discogenic pain in the posterior aspect of the disc. A porcine study [64] and a cadaveric study [65] have shown that there is minimal thermal damage to the anterior disc or posterior longitudinal ligament.

2 prospective cohort studies and 1 RCT with a follow-up report at 1 year is found. Biacuplasty was first described in a

case report by Kapural et al. [66]. The main author then enrolled 15 patients into a pilot study with a 6-month follow-up [67] with positive results. Another group then conducted a prospective study in 2011, again with 15 patients and 6 months follow-up, and improvements in the VAS and ODI were seen [68].

The only RCT on biacuplasty was done by Kapural et al. [69]. In this double blinded sham-controlled study consisting of 32 patients in each group with a follow-up of 6 months, positive results were seen in the treatment group in terms of statistically significant improvements in SF-36, NRS and ODI scores with no complications reported. However, 9 patients were lost to follow-up and the results stated were from a per protocol analysis. The same authors recently published a followup study to the RCT mentioned above [70]. The patients who underwent the treatment had follow-up to 1 year and the sham patients who elected to cross over to the treatment arm had follow-up to 6 months. Sustained improvements were seen at 12 months in the NRS and SF-36 scores but the improvement in ODI score became insignificant. For the cross-over patients, significant improvement in the SF-36 score was obtained but the improvements in NRS, ODI and opioid usage did not attain statistical importance. Similar to the criticism of the RCT, there was a high drop-out rate and a per-protocol analysis was used.

In summary, there is 1 RCT of moderate quality and 2 prospective studies that show positive results for intradiscal biacuplasty. No complications have been reported that is directly related to this procedure. There is certainly a need for more studies to be done on this promising procedure. There is some evidence that this modality may be useful for the treatment of discogenic pain.

### Methylene Blue (MB)

The latest addition to the treatment of discogenic LBP is intradiscal MB. The postulated mechanism of action is that MB is neurolytic and it denervates the small nociceptive fibers that grow into a degenerated disc's annulus fibrosis [71,72]. MB is also a direct inhibitor of nitric oxide which is involved in the inflammatory process in the intervertebral disc [73,74].

5 studies have been published on it with the first being reported in 2005. In this prospective study [72] with a follow up of 18 months, Peng et al had very good results with intradiscal MB in 24 patients. The authors then went on to perform a well conducted multi-centre placebo controlled RCT [71] (Jadad score = 5) in which similarly impressive results were replicated. Patients were followed for 24 months and large decreases in NRS and ODI were seen. These improvements were seen by 6 months and sustained up to 24 months.

Unfortunately, these results were not seen when other authors utilized MB in their own centers. Gupta et al. [75] had a small retrospective case series of 8 patients in which only 1 patient has 100% relief at 6 months. 3 patients had relief for shorter durations of between 2 weeks to 5 months. The other 4 had no pain relief at all. Kim et al. [76] conducted a prospective study in 20 patients with a follow-up of 12 months. At 3 months' post-procedure, 55% had an improvement but this reduced to 25% at 12 months. The difference in this study as compared to the previously mentioned ones is that there was a 1 week lapse between PD and MB injection, and that a significant number of their patients (75%) had treatment at 2 or more levels. There is a possibility that the patients in this study had more severe disease as compared to the other studies and as such their patients could not attain the same results as in Peng's study. The most recent study published was by Kallewaard et al. [77] as a pilot study to assess the efficacy of MB, and if it was effective to carry on and conduct a larger placebo controlled RCT. In their study, they defined success as a 30% decrease in pain at 6 months which 6 out of 15 patients (40%) managed to achieve.

In addition to questions over its overall efficacy, it is a relatively recent development of intradiscal therapy and so it is unknown if there is any utility or harm in repeating intradiscal MB injection in situations in which it provided short term pain relief.

In summary, there is a well-conducted RCT that showed very good results for intradiscal MB therapy but conflicting results were then shown in the other trials. There is some evidence that it may be efficacious but more studies will be needed to confirm this.

#### Other Therapies

A recent article describes a new navigable catheter, L'DISQ (U&I Co. Ltd., Uijeongbu, Korea) that uses coblation technology to ablate the torn annulus. L'DISQ allows coblation to be performed directly to the annular tear as the catheter is navigable so it should theoretically be more effective. In this pilot study of 20 patients, which was conducted by the inventor of the device, 11 had greater than 50% pain reduction at 48-week follow-up [78]. More studies are needed to properly assess this modality.

Several articles have already been published utilizing percutaneous lumbar disc decompression (PLDD) for lumbar radicular pain but the literature of its effectiveness on discogenic

LBP is scanty. A prospective study was published in 2011 [79] which described its usage in discogenic pain. 11 patients with LBP secondary to a single level disease that had a selective anaesthetic blockade of 1 ml of 1% lidocaine were included. At a follow-up of 2 years, there were significant improvements in ODI and VAS scores. 1 patient developed a discitis that needed surgery. The other study on the use of PLDD for discogenic pain was a retrospective report in which patients were included either based on clinical findings or a discography [80]. 32 patients responded to a telephone interview in which 28 had a good or fair response. More studies will be needed before this modality can be considered a viable treatment for discogenic LBP.

There are 2 articles that have been published investigating the efficacy of using etanercept, a TNF- $\alpha$  inhibitor, intradiscally. The first study was a double blind, placebo controlled, dose response RCT (Jadad score = 5) in which intradiscal saline was compared against a range of etanercept doses with the maximum being 1.5mg [81]. At 1-month follow-up there was no difference between any of the etanercept groups and placebo. The 2nd article, also an RCT (Jadad score = 4), compared 2 groups. The control group had 10mg bupivacaine infiltrated intradiscally and the other group had 10 mg etanercept introduced in addition to that [82]. The study showed lower NRS scores from one to 4-week follow-up in the etanercept group but this difference disappeared at 8-week follow-up. Intradiscal etanercept seems to provide short term pain relief at best.

#### Discussion

As presented in our topical review, most of the studies on percutaneous treatments to discogenic pain are observational in nature. Only 7 out of more than 40 articles presented are RCTs. Though all are of reasonable to good quality (Jadad score all  $\geq$ 3), 5 out of 7 either did not calculate a sample size or did not meet it at the end of the study [14,32,43,46,83]. As such, the negative results obtained in the study could have been due to a lack of power. The difficulty of obtaining a proper blinded control group may also be a reason for the lack of RCTs as subjecting patients to a sham interventional procedure is neither palatable to patients and ethical boards.

The assessment of the various modalities is dependent largely on the results of prospective observational studies with all their inherent biases. As such, it is difficult for any modality to have a strong positive recommendation and none were found in our review. Even among the modalities with a weakly positive recommendation i.e. biacuplasty and MB, more well conducted studies are certainly needed. IDET has the most number of publications though most of it is observational in nature. As mentioned above, 1 RCT was a negative study and the other RCT showed a slight advantage of IDET over placebo in well selected patients.

MB and biacuplasty are still in its infancy as compared to IDET but the initial results show promise. More evidence will be

needed to ascertain their role in the treatment of discogenic pain. Though the mechanism of action of nucleoplasty for discogenic LBP is still in doubt and with some of the trials not fulfilling IASP criteria, all the observational studies show a positive result. More literature specifically addressing nucleoplasty for the management of discogenic LBP are needed to evaluate if it is a viable treatment option.

For the other modalities, it has been shown in this article that intradiscal steroids, DiscTRODE, intradiscal RF and L2 blockade/RF are not effective and there are better alternatives in the treatment of discogenic LBP. Intradiscal pRF has conflicting results and the methodology differs substantially between the studies, making it difficult to evaluate and should only be used in the context of a study. Rami communicantes RF only has 1 study in a very select group of patients and so no conclusions can be made on this modality.

It should be remembered that in this group of patients that have failed conservative management of discogenic LBP, the options for treatment are limited. Fusion surgery for this condition is not without controversy [84,85] and its complications significant. However, allowing the patients to continue on long-term analgesics with its attendant side effects is not optimal, and many of these patients will still have significant disability [86]. Percutaneous treatments may allow a subset of these patients to have a significant improvement and the risk of complications as compared to surgery is much lower [87-90]. It is therefore critical to appraise the evidence for such modalities and adhere to strict patient selection. The limitations of this article are that only literature in PubMed and published in English were considered [91,92]. A broad overview, though not exhaustive, of all available percutaneous treatments for discogenic back pain has been discussed in sufficient detail to identify a need to have better designed studies and modalities for the indication of discogenic back pain [93-95].

### Conclusion

Biacuplasty and intradiscal MB have a weakly positive recommendation for the treatment of discogenic pain. Nucleoplasty may potentially be a viable treatment though more evidence and basic research is needed. The evidence for IDET is equivocal. There are negative recommendations for intradiscal steroids, intradiscal RF, DiscTRODE or L2 blocks/RF. Future trials on discogenic pain should ensure that the IASP definition of discogenic pain is adhered to and the methodology of confirming the diagnosis of discogenic pain clearly elucidated.

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