

REVIEW ARTICLE

Applications of vibrational energy in the treatment of sinonasal disease: A scoping review

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Abstract

Background: Acoustic energy and vibration therapy are emerging as helpful adjuncts among many disease states. There has been interest in how this technology can either serve as an alternative treatment or enhance delivery of medications to treat pathology within the nasal cavity and paranasal sinuses. Our objective was to perform a scoping review of the state of the science of vibration treatment used in sinonasal disease.

Methods: A search of Embase, PubMed, and CINAHL databases was performed in November 2021. Included studies evaluated acoustic energy as a means of treatment in sinonasal diseases. Data points collected included type of technology utilized, disease state treated, and outcomes.

Results: The initial search identified 2902 studies, of which 44 met inclusion criteria. A wide array of vibrational technology such as ultrasound, sonic aerosols, and phonophoresis, with varying frequency and amplitude were described. Twenty-six studies evaluated the use of acoustic energy to treat sinonasal disease itself, while 18 studies evaluated the use of acoustic energy to facilitate drug delivery to the sinonasal cavity. Outcome measures among studies were highly varied.

Conclusions: Vibration technology used in patients with sinonasal pathology has been shown to improve pain, sinonasal symptoms, and radiologic outcome measures in small studies. Given the heterogeneous study populations and outcomes, no conclusion could be reached regarding overall effectiveness of acoustic energy as a primary treatment. Further research is required to study specific treatment indications in larger patient populations to fully understand the potential clinical benefit and to determine optimal therapeutic characteristics of sound energy.

KEYWORDS

acoustic energy, chronic rhinosinusitis, rhinitis, ultrasound, vibration

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1 | INTRODUCTION

Acoustic energy is the disturbance of energy, which passes through a medium, in the form of a wave. Vibration specifically refers to a mechanical oscillation (ie, a repetitive movement) around a point of equilibrium. Sound and vibration are closely intertwined. For example, vibration can generate a wave of sound or, vice versa, a sound wave can cause an object to vibrate. Acoustic energy and vibrational technology have been used in the treatment of many different disease states. For example, vibration has been utilized in physical therapy for improvement in muscle disorders,¹ enhancement of wound healing,² and relief of arthritic pain.³ Ultrasound has been used to facilitate drug delivery via phonophoresis, a process by which ultrasound increases percutaneous absorption of medication. Vibration has also been used in patients with cystic fibrosis as part of chest physiotherapy to improve patency of their lower airway and reduce mucus plugging.⁴ With the successful application of vibrational therapy to lower airway diseases, there has been an emerging interest in evaluating applications of acoustic and vibrational technology for the treatment of upper respiratory inflammatory disorders.

Sinonasal inflammatory disorders encompass a wide array of diseases, most commonly chronic rhinosinusitis (CRS), allergic rhinitis (AR), and non-AR (NAR). These disorders are characterized by chronic symptoms including nasal congestion, nasal drainage, facial pain/pressure, and hyposmia. Patients with these disorders experience detriments in quality of life, decreased productivity, and exacerbation of comorbid diseases such as asthma.⁵ Disorders of tissue remodeling, mucociliary function, host immunity, cellular metabolism, and the inflammatory cascade have been studied as mechanisms that lead to the dysfunction seen in sinonasal inflammatory disorders.⁶ Additionally, neurogenic processes have also been implicated in pathophysiology leading to tissue edema and dysfunction.⁷

An emerging body of literature suggests that acoustic and vibrational technologies may offer therapeutic alternatives to traditional medical and surgical therapies for upper airway inflammatory conditions, or may enhance existing forms of medical therapy. We therefore performed a scoping review of the literature to evaluate the evidence for acoustic energy as a treatment of sinonasal inflammatory disorders.

2 | METHODOLOGY

The Preferred Reporting Items for Systematic reviews and Meta-Analysis extension for Scoping Reviews (PRISMA-ScR) was followed.⁸

2.1 | Search strategy

A literature search was performed on November 2, 2021, of Embase, PubMed, and CINAHL databases. Additional records were identified by examining the references of articles obtained for review. All databases were searched from inception to search date. The query used a combination of subject headings (eg, MeSH in PubMed) and keywords for sinonasal disease (pansinusitis, rhinosinusitis, sinus disease, sinus infection, sinus congestion, chronic rhinosinusitis, chronic sinusitis, rhinitis, sinonasal disease, facial pain, and facial neuralgia) and acoustic energy (ultrasonic, ultrasound, humming, vibrating, vibration, vibromassage, and kinetic oscillation). Full electronic search strategy is included in the supplemental material, Supplemental Table 1.

2.2 | Selection criteria

Inclusion criteria for review included articles that: (1) evaluated treatment of sinonasal disease (which broadly included facial pain) via the application of acoustic energy effectiveness (any outcome) for treatment purposes; (2) encompassed all levels of evidence; and (3) were published in all languages. Any non-English abstract was translated for screening and if inclusion criteria were met, the full article was then professionally translated for formal review. Exclusion criteria included: (1) nonsinonasal-related pathology or anatomic site, (2) use of acoustic energy for diagnostic rather than treatment purposes; and (3) reviews, commentaries, conference abstracts and proceedings, and nonhuman studies.

2.3 | Data extraction

Two authors (K.M.P. and P.R.) screened titles of identified abstracts to exclude articles that did not meet predetermined selection criteria or were duplicates. Articles that were agreed upon by both screeners to meet criteria were included for review; discordant judgments were adjudicated by a third author (P.H.H.). Following selection, two authors (K.M.P. and P.R.) independently extracted the following information when available: author, year of publication, study population, type of acoustic therapy intervention, and study outcome. Data were then synthesized, and results were divided into separate groups based on how sound energy was applied for treatment of sinonasal disease.

3 | RESULTS

3.1 | Search and study characteristics

The initial screening process returned 2902 citations. After removal of duplicates, titles and abstracts were then reviewed, which led to 131 articles for full review (Figure 1). Finally, 44 articles met our inclusion criteria and were included in the review. All studies evaluated the impact of acoustic energy on the treatment of sinonasal disease. Included studies were published between 1968 and 2021. Included articles were either originally published in English or professionally translated to English from Russian, Japanese, Polish, or German.

There were two main themes in which acoustic energy was used to treat sinonasal disease. The first was the application of acoustic energy to the sinonasal region to treat facial pain and sinonasal inflammation. The second theme was the use of acoustic energy to facilitate drug delivery to the nasal cavity and paranasal sinuses. The distribution of papers between the two themes, and levels of evidence are described in Tables 1 and 2 and Figure 2. For the sake of brevity, case reports are only detailed in the Tables.

3.2 | Acoustic energy applied to the sinonasal region to treat sinonasal disease

3.2.1 | Acoustic energy applied externally to address facial pain

Five studies used sound energy applied externally to address reduction in facial pain as the primary outcome of interest. The first study was a case report and is detailed in Table 1.⁹

Of the interventional studies, one study randomized 120 participants with myofascial pain dysfunction syndrome to medical therapy, shortwave diathermy, or ultrasound therapy (1.5 W/cm²) applied for 5 minutes daily for 2 weeks and found the most improvement in facial pain graded scales in the ultrasound group, relative to the shortwave diathermy and medical therapy groups.¹⁰ Another study followed 96 patients who were admitted to an inpatient service for orofacial acute pain secondary to dental pathology and were randomized to receive vibration stimulation of the face via a probe ($n = 76$) or placebo ($n = 20$). The authors found 71% of those who received vibration stimulation reported some pain relief (via reduction in visual analog scale [VAS]), whereas placebo stimulation was significantly less effective.¹¹ The next study evaluated facial pain reduction in a patient population with facial pain attributed to sinonasal causes, both acute and chronic. Mechanical vibratory stimulus (100 Hz) was applied to

the facial skeleton for 45 minutes, and 70% of participants reported reduction in pain.¹² The last study was a single-arm interventional study of 14 patients with CRS with facial pain who underwent multimodal frequency treatment administered via the AxioSonic device. This device operated at two simultaneous frequencies and was applied at 1 W/cm² for 5 minutes on the skin overlying the maxillary sinus and 0.5 W/cm² for 5 minutes overlying the skin on the frontal sinus. Mean 22-item Sino-Nasal Outcome Test (SNOT-22) improvement over 2 weeks was 14.11 ($p < 0.05$), exceeding the minimal clinically important difference (MCID). The authors of this study were associated with the company that developed the AxioSonic device.¹³

3.2.2 | Acoustic energy applied externally to address bacterial sinusitis

Five studies evaluated sound energy applied externally to address treatment of bacterial sinusitis. One study was a randomized clinical trial (RCT),¹⁴ one was an observational study,¹⁵ and the remaining three were interventional studies.^{16–18} The 2010 RCT was conducted in the primary care setting and randomized patients with acute bacterial sinusitis to either receive transcutaneous ultrasound therapy (1.0 W/cm² in continuous mode for 10 minutes each day for 4 days) or receive amoxicillin for 10 days. At day four of intervention, the group undergoing ultrasound therapy showed a 1.5-point greater reduction in pain relative to the antibiotic group. At day 21, there were no significant differences among symptoms between the two groups.¹⁴

A study from 1997 examined the ability of pulsed magnetic field (sinusoidal field shape, frequency of 35 Hz for 12 minutes at intensity of 100% (2.5 mT) once a day for 10 days) via a Magnetronic MF-IO apparatus to improve symptoms of a sinus infection in patients with both acute maxillary sinusitis and chronic maxillary sinusitis with exacerbation. Of the nine patients with acute maxillary sinusitis, all received antibiotics and pulsed magnetic field therapy, with six patients reporting improvement in nasal symptoms. For participants with chronic maxillary sinusitis with exacerbation, they were divided into three groups—those who underwent only pulsed magnetic field therapy, those who underwent surgery and pulsed magnetic field therapy, and those who underwent antibiotics and pulsed magnetic field therapy. In the group who had pulsed magnetic therapy alone, eight of 21 reported complete improvement, seven of 21 reported some improvement, and six of 21 reported no improvement in nasal symptoms. The study was limited by lack of randomization, lack of a control group, and lack of reported participant demographics.¹⁶ The next study evaluated the effect ultrasound had on the

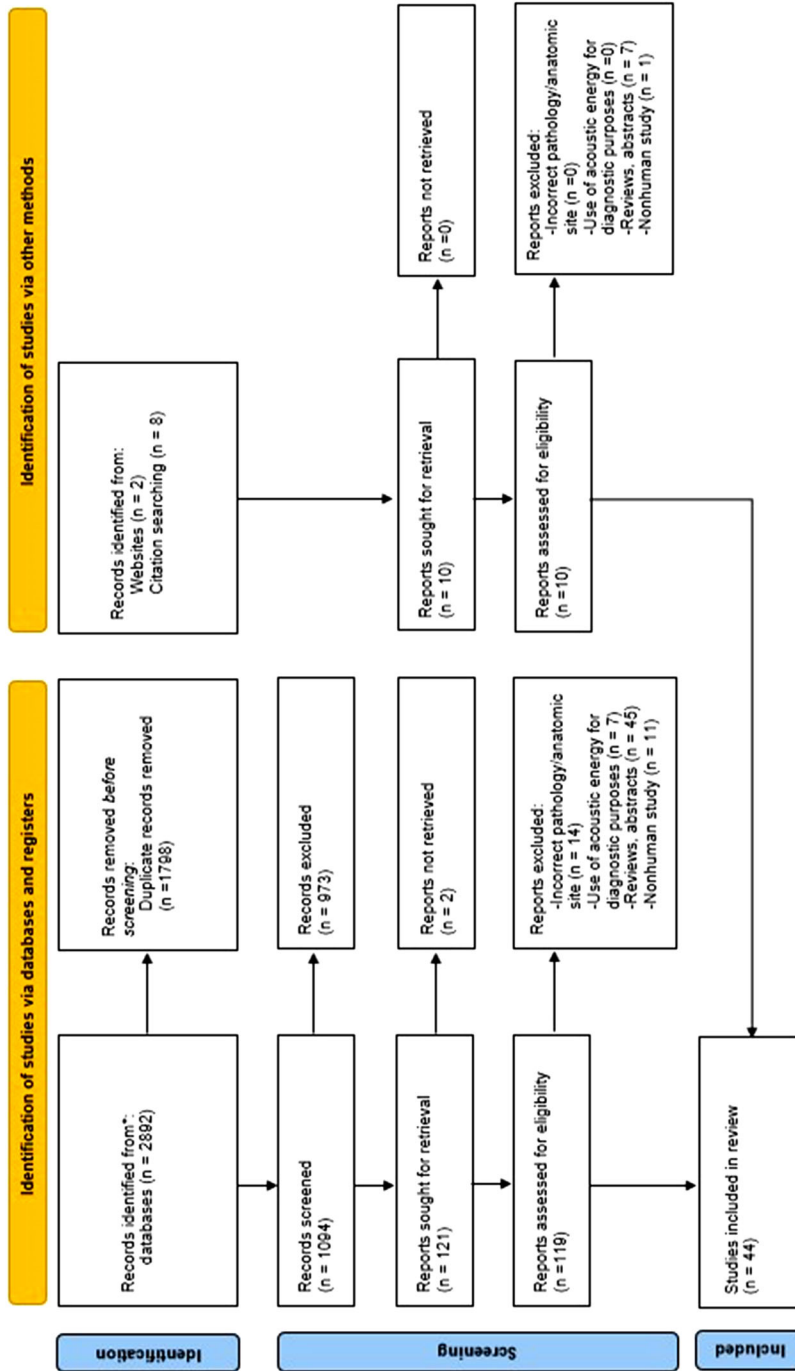


FIGURE 1 Flow chart of identification and screening as performed in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-Analyses guidelines.

TABLE 1 Acoustic energy used to treat pathology of the sinonasal cavity

Author	Year	Country	Study population	Type of acoustic intervention	Treatment group, n	Control group, n	Outcomes	LOE
Acoustic energy applied externally to address facial pain								
Lundeberg	1985	Sweden	Facial pain attributed to sinonasal cause	Vibratory stimulus to probe (100 Hz), external	20	0	70% reported reduction in pain via a pain graded scale	4
Hansson	1986	Sweden	Facial pain attributed to dental pathology	Vibratory stimulus via probe (10–200 Hz), external	76	20	71% in the vibration group reported significant reduction in pain via VAS	4
Talaat	1986	Egypt	Myofascial pain dysfunction syndrome	Ultrasound (1.5 W/cm ²), external	40	80	Better reduction in pain graded scale in the ultrasound group (2.40 ± 0.05–0.55 ± 0.75) vs (2.10 ± 0.74–1.90 ± 0.74, medical) and (electricity, 2.40 ± 0.67–0.78 ± 0.95)	3
Baduni	2017	India	Facial pain	Pulsed ultrasound (0.8W/cm ²), external	1	0	Three-point reduction in VAS for facial pain	5
Smith	2017	United States	CRS (facial pain)	Multimodal frequency device (low: 70–80 Hz, high: 1 MHz –0.5 to 1 W/cm ²)	14	0	Mean SNOT-22 improvement was 14.11 (p < 0.05)	4
Acoustic energy applied externally to address bacterial sinusitis								
Kantor	1997	Poland	Sinusitis (acute, chronic)	Pulsed magnetic field therapy (35 Hz [acute], 50 Hz [chronic, 2.5 mT), external	30	12	46% of the intervention group had complete resolution of sinusitis determined by resolution of nasal symptoms	4
Zelenkin	1998	Russia	Sinusitis (acute, chronic)	Ultrasound (64 Hz), external	128	0	86% of acute and 65% of chronic cases resolved determined by resolution of nasal symptoms and lack of purulence on last puncture	4
Zelenkin	2000	Russia	Acute maxillary sinusitis	Vibratory stimulus via probe (50–100 Hz), external	86	80	Vibration group with fewer punctures (67.8% had no purulence at second puncture) vs control (average 6 or 7 punctures) and LOS was 6 to 7 d in the vibration group vs control (8–10 d)	4

(Continues)

TABLE 1 (Continued)

Author	Year	Country	Study population	Type of acoustic intervention	Treatment group, n	Control group, n	Outcomes	LOE
Hosoien	2010	Norway	Acute bacterial sinusitis	Ultrasound (1.0 W/cm ²), external	24	24	1.5-point higher reduction in pain in the ultrasound group vs the control group	2
Feizabadi	2018	Iran	CRS and healthy controls	Ultrasound (0.5–1 W/cm ²), external	22	0	87% reduction in <i>Staphylococcus aureus</i> bacterial copy number after treatment	3
Acoustic energy applied intranasally to address bacterial sinusitis								
Zelenkin	1998	Russia	Acute frontal sinusitis	Vibration stimulus (50 Hz), intranasal	52	0	85% resolution of sinusitis by day 3 determined by patency of the frontal sinus tract via irrigation	4
Khudiev	2003	Russia	Chronic maxillary sinusitis	Ultrasound (3 W/cm ² , 26.5 Hz) external	61	10	72% of the ultrasound group had complete recovery, 40% had complete recovery in antibiotics alone	4
Acoustic energy applied externally to address CRS symptoms								
Ansari	2004	Iran	CRS	Ultrasound (1 W/cm ² , 1 MHz) external	1	0	Resolution of symptoms and CT findings of sinusitis	5
Eby	2006	United States	CRS	Humming (130 Hz, 18 hums per min)	1	0	Resolution of symptoms of sinusitis by day 4	5
Ansari	2007	Iran	CRS	Ultrasound (0.5–1 W/cm ²), external	57	0	81.3% improvement in sinus symptoms	4
Ansari	2007	Iran	CRS	Ultrasound (0.5–1 W/cm ²), external	10	10	86.56% (SD 20.76) improvement in sinus symptoms in the ultrasound group relative to 37.14% (SD 46.37) in control	3
Ansari	2010	Iran	CRS	Pulsed Ultrasound (1 MHz, 1 W/cm ²), external	1	0	Resolution of symptoms and CT findings of sinusitis	5
Young	2010	New Zealand	CRS	Pulsed Ultrasound (1 MHz, 0.5–1.0 W/cm ²), external	22	0	Median percent improvement in SNOT-20 was 34.1% after the sixth treatment session	4

(Continues)

TABLE 1 (Continued)

Author	Year	Country	Study population	Type of acoustic intervention	Treatment group, n	Control group, n	Outcomes	LOE
Rocha	2011	Brazil	CRS	Ultrasound (1 MHz, 1 W/cm ²), external	14	12	64% objective decrease in nasal obstruction in the intervention group vs. placebo	3
Ansari	2012	Iran	CRS	Pulsed vs. Cont. Ultrasound (1 MHz, 0.5–1 W/cm ²), external	15	15	No difference among the two groups in change in sinusitis symptom score	3
De Castro	2017	Philippines	CRSwNP	Pulsed ultrasound (1 MHz, 1.0 W/cm ²), external	21	21	At week 3, SNOT-22 score of controls was 16.57 (SD, 1.78) and of ultrasound was 10.45 (SD, 1.19) ($p = 1.07E-80$)	2
Ansari	2021	Iran	CRS (with olfactory dysfunction)	Pulsed ultrasound (1 MHz, 1 W/cm ²), external	15	0	Smell Identification Test improved from 13.1 (SD, 1.8) to 22.0 (SD, 1.3) SNOT-20 improved 49.3 (SD, 18.4) to 22.2 (SD, 16.4)	4
Khanwalkar	2021	United States	Nasal congestion	Vibrational headband and individualized sound file	50	0	90% with improved TNSS, facial pain VAS from 1.3 to 0.9, $p = 0.01$	4
Acoustic energy applied intranasally to address chronic sinonasal symptoms								
Juto	2014	Sweden	NAR	Intranasal kinetic oscillation (50 Hz)	35	36	Median RQSS stiffness from 2 to 1 in the treatment group, and no change in the placebo group	2
Cairns	2019	United States	Nasal congestion	Acoustic vibration (128 Hz at 80 decibels)	14	0	Statistically improved VAS in congestion ($Z = 3.1$, $p = 0.002$) and ease of breathing ($Z = 2.9$, $p = 0.003$)	4
Soler	2020	United States	Nasal congestion	Acoustic vibration (128 Hz at 80 decibels)	40	0	PNIF, nasal congestion VAS, TNSS, NOSE, and SNOT-22 all improved ($p < 0.001$)	4

Abbreviations: CRS, chronic rhinosinusitis; CT, computed tomography; LOE, level of evidence; LOS, length of stay; NAR, nonallergic rhinitis; NOSE, nasal obstruction symptom evaluation scale; PNIF, peak nasal inspiratory flow; RQSS, Rhinitis Questionnaire Symptom Score; SD, standard deviation; SNOT-20, 20-item Sino-Nasal Outcome Test; SNOT-22, 22-item Sino-Nasal Outcome Test; TNSS, Total Nasal Symptom Score; VAS, visual analog scale.

microbiome in the nasal cavity by culturing nasal swabs both before and after treatment with ultrasound (1 MHz, 1 W/cm² applied to the skin overlying the maxillary sinus and 0.5W/cm² applied to the skin overlying the frontal sinus). Using polymerase chain reaction as a detection method, *Staphylococcus aureus* was detected in 15 of the 22 patients. After ultrasound treatment, 87% of participants

who had *S aureus* detected were found to have a quantitative reduction in bacteria copy number.¹⁸

The remaining two studies were from the same group in Russia. The first was a study of 128 patients admitted to an inpatient service with acute maxillary ($n = 60$) and acute maxillary and ethmoid sinusitis ($n = 68$) treated first with sinus puncture and drainage, followed by

TABLE 2 Acoustic energy used to facilitate drug delivery to sinonasal cavity

Author	Year	Country	Study population	Type of acoustic intervention	Treatment group, n	Control group, n	Outcomes	LOE
Acoustic energy applied externally with phonophoresis								
Ansari	2013	Iran	CRS	EP (pulsed ultrasound, 1 MHz, 1 W/cm ²)	1	0	Resolution of symptoms and CT findings of sinusitis	5
Ansari	2015	Iran	CRS	EP (pulsed ultrasound, 1 MHz, 1 W/cm ²)	30	30	Percent improvement in symptom score was greater in the EP group (67.2 vs 49.3%, <i>p</i> = 0.03)	3
Acoustic energy applied to intranasal drug delivery devices to facilitate drug delivery								
Zippel	1968	Russia	Chronic maxillary sinusitis	Ultrasonic drug delivery	10	10	Mean (drug) in maxillary was 3.86×10^{-2} in the control group and 4.97×10^{-2} in the ultrasound group	4
Dañniak	1989	Russia	Chronic maxillary sinusitis	Vibrating device (26 KHz) with irrigation	65	35	Decreased need for repeat puncture, decreased duration of days until improvement, and greater resolution in bacterial culture in the vibration vs control groups	4
Saijo	2000	Japan	CRS	Ultrasonic nebulization vs jet nebulization	6	6	More drug delivered to maxillary with jet (average 16.66 ug/mL) vs ultrasound (average 6.5 ug/mL)	4
Gerber	2003	Russia	CRS wNP	Ultrasound probe intranasal	109	0	2.5% had relapsed disease (of note: 28% were lost to follow-up)	4
Maniscalco	2006	Italy	Healthy volunteers	Sonic enhancement of nebulized drug delivery	6	0	After delivery of L-NAME, 22% to 35% reduction in nasal NO after humming	4
Valentine	2008	Australia	Cadavers with maximal sinus surgery	PARI SINUS device (44-Hz vibration to aerosol)	7	7	Increase in intensity (2.06 vs 0.26), percentage of stain (49.96% vs 4.19%), and circumference stained (76.59% vs 12.7%) in the nasal rinse vs the PARI SINUS groups	3
Katsumi	2008	Japan	CRS	PARI SINUS device (44-Hz vibration to aerosol)	56	0	75% reported some level of symptom improvement relative to their baseline	4
Moller	2010	Germany	Healthy volunteers with normal anatomy	PARI SINUS device (44-Hz vibration to aerosol)	5	0	6.5% of drug in the sinuses with PARI, not seen with nasal spray 3× longer clearance with PARI vs spray	4

(Continues)

TABLE 2 (Continued)

Author	Year	Country	Study population	Type of acoustic intervention	Treatment group, n	Control group, n	Outcomes	LOE
Patel	2012	United States	Cadavers after sinus surgery	Pulsed ultrasound (25–27 KHz) drug delivery device	6	0	Drug in maxillary and sphenoid sinus 12 of 12, posterior ethmoid 8 of 12, and frontal 4 of 11	4
Goektas	2013	India	CRS olfactory dysfunction	AMSA device uses vibration (100 Hz) and pressure (10–50 mbar)	18	15	Improvement in olfactory function in both systemic steroid control group and topical steroid via drug delivery device groups	3
Ohki	2013	Japan	CRS	PARI SINUS device (44-Hz vibration to aerosol)	5	0	All patients deemed recovered at follow-up visit, and no specific outcome measures reported	5
Moller	2013	Germany	CRSsNP	PARI SINUS device (44-Hz vibration to aerosol)	11	0	Maxillary deposition was 4.0% ± 1.7% (before surgery) and 6.1% ± 2.2% (after)	4
Mainz	2014	Germany	CF-associated CRS	PARI SINUS device facilitated dornase alfa vs isotonic saline	23	23	SNOT-20 score improved after dornase alfa compared with isotonic saline ($p = 0.017$)	2
Raychler	2015	France	CRS (olfactory dysfunction)	Sonic enhancement of nebulized drug delivery	10	20	Objective olfactory tests (SST) improved (5.5, 5.8, and –1.1, $p = 0.01$) in the nebulization, oral, and nasal spray groups, respectively	3
Mainz	2016	Germany	CF-associated CRS	PARI SINUS device facilitated hypertonic vs isotonic saline	69	69	No statistical or clinical significance between the two groups in SNOT-20	2
Poletti	2017	Germany	CRS (olfactory dysfunction)	AMSA device uses vibration (100 Hz) and pressure (10–50 mbar)	13	16	Improvement in objective olfactory function in both groups at 2 weeks (2.2 with AMSA, 2.6 with nasal spray)	3

Abbreviations: CF, cystic fibrosis; CRS, chronic rhinosinusitis; CRSwNP, chronic rhinosinusitis with nasal polyps; CRSsNP, chronic rhinosinusitis without nasal polyps; CT, computed tomography; EP, erythromycin phonophoresis; L-NAME, nitric oxide synthase inhibitor; LOE, level of evidence; NO, nitric oxide; SNOT-20, 20-item Sino-Nasal Outcome Test; SST, Sniffin' Sticks Test.

low-frequency ultrasound (64 Hz for 2 minutes on consecutive days) applied to the external face. In the acute maxillary sinusitis group, 86% of patients had resolution of sinusitis at discharge. In the maxillary and ethmoid sinusitis patients, 65% had resolution of sinusitis at discharge. This study was limited by lack of a control group, eg, those receiving puncture and drainage only.¹⁵ The same group

then studied patients who were admitted to an inpatient service with acute sinusitis and underwent sinus puncture and drainage and then compared traditional treatment of sinusitis with antibiotics ($n = 86$) with intervention with vibrotherapy ($n = 80$), which was applied both extranasally and intranasally. The vibration therapy was performed with a Russian apparatus, Tonus-3, which requires a 220-V

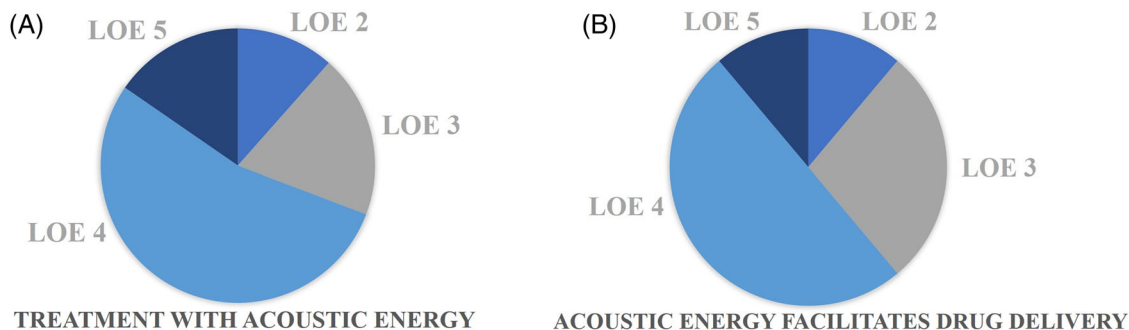


FIGURE 2 Level of evidence (LOE) of the 44 included articles divided by publications (A) that describe treatment with acoustic energy and (B) that describe acoustic energy which facilitates drug delivery to the sinonasal cavity.

circuit, and has an oscillation frequency of 50 Hz to 100 Hz with an amplitude of 0.2 mm to 0.5 mm. The patients who underwent vibrotherapy had a decreased incidence of purulence on second sinus puncture and had a decreased length of stay relative to the group who received antibiotics. The participants were not randomized, which may have introduced selection bias.¹⁷

3.2.3 | Acoustic energy applied intranasally to address bacterial sinusitis

Two studies in this category focused on the intranasal application of acoustic energy to treat bacterial sinusitis. The first study was an interventional study of 71 patients with chronic maxillary sinusitis. Sixty-one participants received intranasal low-frequency ultrasound (modified UZR-M ultrasound apparatus, which was set to 3 W/cm² with a working frequency of 26.5 Hz in amplitude for a duration of 3 minutes for 5 to 7 treatments) placed via maxillary sinus puncture; and 10 controls were given antibiotics only. Complete recovery (which was based on symptoms, repeat sinus puncture results, nasal endoscopy, bacterial culture data, and cytological evaluation of blood counts—although these data were not provided in the article) was noted in 72% of patients who received ultrasound alone and in 40% who received antibiotics alone. The study was limited by lack of reported clinical characteristics specific to each group, lack of randomization, and the absence of criteria constituting “complete recovery.”¹⁹

The second study was an observational investigation of 52 patients with acute frontal sinusitis who underwent puncture of their frontal sinus with removal of purulence, irrigation with an antibiotic solution, and then mechanical low-frequency (50 Hz) vibration via a device placed intranasally along the middle turbinate. Of the 52 patients, 85% had resolution of symptoms by day 3 and 93% of patients had patency of their frontal sinus tract at discharge. This study was limited by the lack

of a comparison group who underwent treatment with antibiotics and puncture without vibration.²⁰

3.2.4 | Acoustic energy applied externally to address CRS symptoms

Eleven publications addressed the application of acoustic energy to the external face to treat sinonasal symptoms in patients with CRS. One Iranian research group produced 6 of the 11 studies evaluating various forms of externally applied ultrasound therapy as it pertains to the treatment of chronic rhinosinusitis. This group published two separate case reports described in Table 1.^{21,22} The same group published a prospective interventional study that enrolled 57 patients with medically refractory CRS to receive low-intensity pulsed ultrasound (1 MHz) to the skin overlying the maxillary sinus (1W/cm² for 5 minutes) and to the skin overlying the frontal sinus (0.5 W/cm² for 4 minutes). The total improvement in sinonasal symptoms was 81.3%. Of the individual symptoms assessed, the greatest improvement was seen in the report of patients with nasal discharge, facial pain, and postnasal drip.²³ Next, the same group performed a placebo-controlled single-blinded RCT of 20 patients with medically refractory CRS to compare treatment with continuous ultrasound (1 MHz with similar settings to that described above regarding maxillary and frontal sinus settings) to placebo (mock ultrasound simulation). The authors reported an increased improvement in mean total sinus symptom score in the intervention group relative to placebo 1 month after intervention.²⁴ The group then conducted a double-blind RCT to compare the effects of continuous and pulsed ultrasound (same settings as above) in 30 patients with medically refractory CRS. No statistically significant difference was found in the main treatment outcome—change in sinusitis symptom score. Of note, this study was reported as a pilot study, as slow recruitment prevented the study from achieving adequate statistical power.²⁵ Finally, the last study evaluated 15 patients with CRS with olfactory dysfunction who under-

went treatment with therapeutic pulsed low-frequency ultrasound (same regimen as above) for 10 sessions, 3 days a week. Mean Smell Identification Test improved from 13.1 (standard deviation [SD], 1.8) at baseline to 22.0 (SD, 1.3) after 10 treatments and persisted at 1-month follow-up at 22.0 (SD, 1.3). Mean 20-item Sino-Nasal Outcome Test (SNOT-20) improved from 49.3 (SD, 18.4) to 22.2 (SD, 16.4) after 10 treatments and persisted at 1-month follow-up (21.0; SD, 15.9). No funding was disclosed.²⁶

Another group from New Zealand conducted a prospective interventional study of 22 patients with medically refractory CRS where participants underwent six sessions of pulsed ultrasound therapy (1 MHz, 0.5 W/cm² for 4 minutes overlying the frontal sinuses and 1 W/cm² for 5 minutes overlying the maxillary sinus). Median percent improvement in SNOT-20 was 34.1% after the sixth treatment session. Two patients reported worsening of symptoms, one of whom went on to develop acute bacterial sinusitis requiring antibiotics.²⁷ In Brazil, Rocha et al²⁸ conducted a cohort placebo-controlled study of 26 patients with medically refractory CRS where patients were randomized to ultrasound therapy (1 MHz, 1 W/cm² applied to the maxillary sinus) versus a sham procedure. The authors found improvement in reported nasal symptoms and an objective 64% decrease in nasal obstruction in the intervention group relative to the placebo group. From the Philippines, a single-blinded RCT of low-frequency ultrasound ($n = 21$) versus placebo ($n = 21$) was conducted in 42 patients with CRSwNP with medically refractory CRS who had undergone sinus surgery. The ultrasound regimen was started at 1 week postoperatively and low-frequency ultrasound (1 MHz) overlying the skin of the maxillary and frontal sinus was applied for 5 minutes for two sessions per week for a total of 3 weeks. The SNOT-22 and modified Lund-Mackay endoscopic scores were statistically improved relative to the sham group. No conflict of interest was disclosed.²⁹

Another study evaluated the efficacy of audible sound frequencies applied to the external face for the treatment of nasal congestion. A prospective, nonrandomized interventional study was performed in 50 patients with nasal congestion using the Soniflow vibrational headband device. With the aid of a mobile phone application, the Soniflow device generated an individualized sound file representing the resonant frequency of the sinuses, calculated from an individual's facial surface landmarks. Participants underwent two sequential 10-minute treatments. After two treatment cycles, 90% of patients showed improvement in Total Nasal Symptom Score, with statistically significant and clinically meaningful differences achieved. The nasal congestion subscore was significantly reduced and mean facial pain VAS also significantly decreased from baseline. Two of the authors did have equity stake in the device company.³⁰

The last study in this category was a case report of a patient who utilized humming as a CRS treatment strategy and is described in Table 1.³¹

3.2.5 | Acoustic energy applied intranasally to address chronic sinonasal symptoms

The following category describes three studies that evaluated acoustic energy applied intranasally to address chronic sinonasal symptoms. The first study was a single-blinded RCT of 71 patients with NAR that compared treatment with intranasal kinetic oscillation stimulation (KOS) with a sham procedure. Participants placed a device into their nasal cavity and inflated the device to 0.05 atm at home once daily for 2 weeks. Active treatment with KOS consisted of mechanical vibrations created using regular pressure oscillations at a frequency of 50 Hz. Improvement in Rhinitis Questionnaire Symptom Score (RQSS) was significant for the treatment group but not the placebo group. There was no significant difference in baseline peak nasal inspiratory flow (PNIF), but, after treatment, the placebo group had higher PNIF. Of note, one of the authors was a major shareholder in the company active in development of KOS products.³²

Two studies evaluated the SinuSonic device, which delivers a combination of acoustic vibration (128 Hz at 80 decibels) and oscillating expiratory pressure to the nasal cavity via a handheld nasal mask. The first was a pilot study that evaluated 14 participants with a history of nasal congestion who underwent application of the SinuSonic device for 2 to 5 minutes. Participants reported statistically improved VAS measures for congestion and ease of breathing after application.³³ The next study was an observational study of 40 participants with chronic nasal congestion who used the device for 3 minutes twice daily over 5 weeks. At 5-minute follow-up, improvement in PNIF and VAS symptoms (statistically and clinically significant) was noted. At the 2-week follow-up, PNIF showed a 31% increase. At the 5-week follow-up, all PROMs reached MCID improvement. No adverse effects were reported, although 10% of participants reported mild discomfort with the device.³⁴ Of note, both SinuSonic studies were funded by the device company.

3.3 | Acoustic energy used to facilitate drug delivery for treatment of sinonasal disease

3.3.1 | Acoustic energy applied externally with phonophoresis

Two studies investigated applications of phonophoresis in the treatment of CRS. Phonophoresis is a process in

which ultrasound is used to facilitate transcutaneous drug delivery. Ansari et al³⁵ published a case report described in Table 2. In 2015, the same group published a double-blind RCT comparing treatment of patients with medically refractory CRS who had pulsed ultrasound only versus pulsed ultrasound (1 MHz, 1 W/cm² for 5 minutes) with phonophoresis using 5% erythromycin ointment applied topically to the skin overlying the maxillary sinuses. Both groups had improvement in total symptom score relative to baseline, but the phonophoresis group had a statistically greater improvement relative to the ultrasound-only intervention group. There was no difference between the two groups in posttreatment computed tomography (CT) findings, although not all participants had a posttreatment CT.³⁶

3.3.2 | Acoustic energy applied to intranasal drug delivery devices to facilitate drug delivery

To understand whether sound energy can improve drug delivery to the paranasal sinuses, various groups have creatively conducted studies to evaluate this question. In one 2006 study, sonic enhancement of nebulized drug delivery (via a nebulizer connected to a rubber duck call that yielded sound with a pulsating airflow) was assessed by the efficacy of delivery of a nitric oxide (NO) synthase inhibitor (L-NG-Nitro arginine methyl ester, L-NAME) to the paranasal sinuses, where NO is synthesized. Nasal NO was induced in six healthy adults through the act of humming, which increases nasal NO levels via improved ventilation of the paranasal sinuses. When L-NAME was delivered transnasally via jet nebulizer without sonic enhancement, NO levels in the nasal cavity rose as expected after humming. Yet, when the same participants were then given L-NAME via the nebulizer with sonic enhancement, there was a 22% to 35% reduction in nasal NO after humming, likely as a result of L-NAME reaching the paranasal sinuses and blocking NO formation.³⁷ This study provided proof of concept for further drug delivery devices utilizing acoustic energy.

In 2012, a cadaver study used low-frequency ultrasound (25–27 KHz) through a handpiece inserted into the nasal vestibule to compare delivery of solution in cadaver heads before and after sinus surgery. Before sinus surgery, the drug was able to reach the central nasal cavity and ethmoids reliably. After surgery, the pulsed ultrasound device was able to deliver drug to the maxillary and sphenoid sinus reliably and the frontal sinus and skull base some of the time. Of note, this study had two authors with ownership interest in the device and another author on the advisory board of the device company.³⁸

A study then evaluated 20 patients with maxillary sinusitis, 10 of whom received ultrasonic aerosol drug delivery to their nasal cavity while the other 10 patients received aerosolized drug without ultrasound. All patients then underwent surgery, and their maxillary sinus tissue was sampled. The mean concentration of the drug was found to be 3.86×10^{-2} for those patients without ultrasound and 4.97×10^{-2} for those who had received the ultrasound-facilitated aerosolization before surgery.³⁹ On the contrary, in another study that evaluated six patients who had previously had sinus surgery, nebulized medication was administered first via an ultrasound nebulizer and then 2 days later administered via a jet nebulizer. Drug concentrations at the anterior head of the inferior turbinate, maxillary sinus, and posterior ethmoids were measured after each treatment. Both methods allowed good drug delivery to the aforementioned sites, with significantly more drug delivered to the anterior inferior turbinate and maxillary sinus in the jet nebulizer application relative to the nebulizer aided by ultrasound.⁴⁰

The next study described details of 109 patients with CRS with nasal polyps (NPs) who underwent a Caldwell-Luc procedure and intraoperatively had their maxillary sinus filled with a chlorhexidine solution to which ultrasonic energy was applied using the LORA-DON apparatus. Depending on the cause of sinusitis (infectious or inflammatory), the sinus was then infused with either an antibiotic- or dexamethasone-impregnated solution after the device was turned off. Of these 109 patients, long-term follow-up (6 months) was available for 78 patients. The main outcome followed was relapse of disease, which, while vaguely defined, was nevertheless reported in 2.5% of the study population at long-term follow-up.⁴¹ Another study then evaluated 100 patients with acute or chronic maxillary sinusitis and divided the participants into three treatment arms: (1) sinus puncture with application of a vibrating device (LORA apparatus, 26 KHz) using saline; (2) puncture, vibrating device with antibiotic irrigation; or (3) control, puncture and rinse with antibiotics only. The authors found decreased need for repeat puncture, shorter duration of days until improvement in symptoms, and greater resolution in bacterial culture in the vibration groups relative to control. There were no significant differences between the two vibration groups (saline and antibiotic solution). No funding sources were disclosed.⁴² Reychler et al⁴³ performed a prospective RCT of 30 patients with CRS with hyposmia who were randomized to treatment with oral steroids, budesonide nasal spray, or sonic nebulization of budesonide for 16 days. Objective olfactory tests improved with statistical significance in all three groups over baseline, but only the oral steroid and sonic nebulization groups achieved MCID. The study was funded in part

by the medical device company who made the sonic nebulizer.

The following two studies evaluated the AMSA nebulizer device, which provides vibration (100 Hz) and pressure impulses of 10 to 50 mbar. The first study evaluated 33 patients with CRS who had olfactory dysfunction. Eighteen participants were given pressure-pulsed inhalation of prednisolone for 20 minutes for six sessions, and 15 participants were given oral steroids. Both groups had improvement in VAS of olfactory function and objective measures of olfactory function at 2 months and 6 months. No funding source was disclosed.⁴⁴ The second study of the device by a different group, randomized patients with CRS to either receive conventional nasal spray (dexamethasone) or to pressure-pulsed nasal inhalation of dexamethasone for 12 days. Improvement in olfactory function was seen at 2 weeks in both groups, although MCID was not reached and decline in olfactory function was seen at 6 weeks relative to the 2-week mark. The AMSA device was provided by the company for the study.⁴⁵

The final studies evaluated the PARI SINUS drug delivery device, which works via vibration aerosol with 44-Hz vibration added to a spray aerosol. In 2008, a study of cadavers that had undergone maximal sinus surgery compared drug delivery with standard nasal douching to nebulization with the PARI SINUS device and found greater distribution and drug concentration to all sinuses with the standard nasal douching relative to the PARI SINUS device. This study was not funded by the drug device company.⁴⁶ Next, the device was studied in 56 patients with CRS who used the device to deliver topical steroids and antibiotics to the sinonasal cavity. Of all participants, 75% reported some improvement relative to baseline. This study was limited by lack of a comparison or control group. No funding source was listed for this study.⁴⁷ A case series of five patients using the device is detailed in Table 2.⁴⁸ Next, a group published two separate studies, the first a case series of five healthy patients with normal sinonasal anatomy who underwent drug delivery via a nasal spray and then via the PARI SINUS device on two separate occasions. Outcomes are shown in Table 2. Funding of the study was supported by the device company.⁴⁹ The same group went on to study the device in 11 patients with CRS without NPs before and after sinus surgery. Total nasal deposition was $56.7\% \pm 13.3\%$ and $46.7\% \pm 12.7\%$ before and after sinus surgery, respectively. Maxillary sinus deposition (as a percentage of nasal dose) was $4.0\% \pm 1.7\%$ and $6.1\% \pm 2.2\%$ before and after sinus surgery, respectively. No significant distribution was seen in the frontal sinus. This study was funded in part by the device company.⁵⁰

Finally, another group published two separate studies evaluating the PARI SINUS device in patients with cystic fibrosis (CF). The first study was a double-blind crossover

RCT. Twenty-three patients with CF were randomized to inhale either dornase alfa or isotonic saline (placebo) for 28 days with the PARI SINUS device. Participants then underwent a 28-day washout period and then crossed over to the alternative treatment. The authors found that primary nasal symptoms (via the nasal subdomain of SNOT-20) improved significantly with dornase alfa compared with no treatment, while a small improvement with isotonic saline did not reach significance. The SNOT-20 overall score improved significantly after dornase alfa compared with isotonic saline ($p = 0.017$). The device company did partially fund the study.⁵¹ The same group then went on to perform a multi-institutional RCT of 69 patients with CF who had CRS. Participants were randomized to receive sinonasal vibrating inhalation via the PARI SINUS device of either NaCl 6.0% or NaCl 0.9% for 28 days. Both therapeutic arms were well tolerated and showed trends towards significance in improvement of SNOT-20 scores (NaCl 6.0%: -3.1 ± 6.5 points, NaCl 0.9%: -5.1 ± 8.3 points), although did not reach statistical or clinical significance. This study was funded by the Association Luxembourgeoise de Lutte contre la Mucoviscidose and the German Cystic Fibrosis Association, although the device company provided the devices and medications.⁵² Notably, in both studies, the vibration-enhanced nebulizer was used in both treatment arms, so there was no ability to determine the specific contribution of the vibration component of therapy to the clinical outcome.

4 | DISCUSSION

Our scoping review found two main methods in which acoustic and vibrational technology was being utilized for chronic sinonasal disorders: (1) using the acoustic energy itself to treat the disease, and (2) using the technology to facilitate drug delivery.

In the first category where acoustic energy itself was used to treat sinonasal inflammation or facial pain, several studies reported success using ultrasound to treat facial pain, to aid in clearance of bacterial infections and to decrease CRS symptoms. There are several mechanisms of action that may explain the benefit of acoustic energy and vibration on sinonasal inflammation and infection as well as facial pain. First, it is known that sinonasal inflammation has a neurogenic component leading to autonomic nervous system dysfunction thought to be secondary to sympathetic hypofunction.⁵³ This autonomic dysfunction is hypothesized to cause tissue edema, increase mucus secretion within the sinonasal airway, and heighten sensation of pain.⁷ Vibration and acoustic energy have been shown to modify neuromodulators and reduce pain. A study in a rat model found that therapeutic ultrasound is

able to decrease upregulation of neuromodulators associated with pain including neurokinin 1 receptor, substance P, tumor necrosis factor α , and interleukin 6, with associated decreased pain behavior in rats.⁵⁴ Furthermore, there is evidence that vibration may help to eliminate biofilms—certainly a cause that may propagate or exacerbate sinonasal disease in some cases. In an *in vitro* study of patients with NPs, the polyps treated with low-frequency ultrasound showed decreased inflammatory cell count in the subepithelial layer and removal of bacterial biofilm from the surface of the epithelial layer relative to the control tissue not treated with ultrasound.⁵⁵ Moreover, ultrasound has also been shown to have an intrinsic antibacterial effect beyond disrupting biofilms.⁵⁶ Finally, vibration is also thought to mechanically reduce viscoelasticity of sputum and enhance mucociliary clearance and expectoration of mucus in the lower airway—as seen in CF.⁵⁷ It is reasonable to hypothesize that this might have a similar effect in the upper airway in patients without CF.

Limitations of the articles identified in this portion of the scoping review include small sample size, frequent lack of a control group, and heterogenous outcome measures, which made it difficult to compare interventions. Three separate devices, two that utilized acoustic vibration and the other that used KOS, had success in treating rhinitis symptoms. Yet, the sample size in the studies were small, and different outcomes measures were used, making comparisons difficult. In addition, some authors in each study had a financial interest in the success of the device.

In the second category—the use of acoustic energy and vibration to facilitate drug delivery—there were mixed results. In the L-NAME study, the authors found implied evidence that vibration-enhanced nebulizer devices improved delivery of pharmacologically active compounds to the sinuses compared with nebulizer alone—an important proof of concept for the potential success of this technology.³⁷ Yet, subsequent studies of vibration-enhanced drug delivery have shown varied outcomes. Only one study compared vibration-enhanced nebulization to standard nasal saline irrigations, in which saline irrigation achieved better drug delivery in patients who had previously undergone sinus surgery.⁴⁶ Other well-designed studies did show success in drug delivery to the sinuses and improved sinonasal symptoms, but these studies were often funded by the device company, therefore introducing a risk of bias. Phonophoresis with erythromycin applied transdermally overlying the maxillary sinus did show improvement in symptoms in the RCT,³⁶ yet the sample size of the study was small and no other group has successfully reproduced the results. Overall, articles reviewed for this component of the scoping review were limited by small sample size, different disease states studied, different outcomes measured, different

technology utilized, and conflicts of interest present in many of the device studies.

An additional consideration to be noted regarding this review is that two of the authors (K.M.P. and P.H.H.) have equity stake in a company focused on utilizing acoustic technology for sinonasal disease.

Despite the shortcomings of the body of evidence reviewed, this scoping review did identify 44 separate studies on this topic, of which provide mostly level 3 and 4 evidence.⁵⁸ The studies included in this review represented heterogenous device technologies, study designs, and clinical outcomes, and therefore no conclusion could be reached regarding the overall effectiveness of acoustic energy as primary treatment for sinonasal disease and facial pain, or for enhancement of drug delivery. However, there are well-designed studies that show promise in this field and speak to the potential for future growth of this emerging area of study towards the development of novel therapies. Studies directed at further elucidating underlying mechanisms of action and measuring rigorous clinical outcomes are needed to address current gaps in the existing body of evidence.

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CONFLICT OF INTEREST

Phillips: equity interest in Third Wave Therapeutics; Roodar: none; Hwang: equity interest in Lyra Therapeutics, Tivic Health, and Third Wave Therapeutics.

REFERENCES

1. Saquetto M, Carvalho V, Silva C, Conceição C, Gomes-Neto M. The effects of whole body vibration on mobility and balance in children with cerebral palsy: a systematic review with meta-analysis. *J Musculoskelet Neuronal Interact.* 2015;15:137–144.
2. Cullum N, Liu Z. Therapeutic ultrasound for venous leg ulcers. *Cochrane Database Syst Rev.* 2017;5:Cd001180.
3. Rutjes AW, Nuesch E, Sterchi R, Jüni P. Therapeutic ultrasound for osteoarthritis of the knee or hip. *Cochrane Database Syst Rev.* 2010:Cd003132.
4. Morrison L, Innes S. Oscillating devices for airway clearance in people with cystic fibrosis. *Cochrane Database Syst Rev.* 2017;5:CD006842.
5. Orlandi RR, Kingdom TT, Smith TL, et al. International consensus statement on allergy and rhinology: rhinosinusitis 2021. *Int Forum Allergy Rhinol.* 2021;11:213–739.
6. Kao SS, Bassiouni A, Ramezanzpour M, et al. Proteomic analysis of nasal mucus samples of healthy patients and patients with chronic rhinosinusitis. *J Allergy Clin Immunol.* 2021;147:168–178.
7. Baraniuk JN. Neurogenic mechanisms in rhinosinusitis. *Curr Allergy Asthma Rep.* 2001;1:252–261.

8. Tricco AC, Lillie E, Zarin W, et al. PRISMA extension for scoping reviews (PRISMA-ScR): checklist and explanation. *Ann Intern Med.* 2018;169:467–473.
9. Baduni A, Krishnamoorthy B. Treatment of hemifacial spasm in patient with hemifacial atrophy using combination therapy (ultrasound therapy and TENS): a case report. *Korean J Pain.* 2017;30:304–307.
10. Talaat AM, el-Dibany MM, el-Garf A. Physical therapy in the management of myofascial pain dysfunction syndrome. *Ann Otol Rhinol Laryngol.* 1986;95:225–228.
11. Hansson P, Ekblom A. Influence of stimulus frequency and probe size on vibration-induced alleviation of acute orofacial pain. *Appl Neurophysiol.* 1986;49:155–165.
12. Lundeberg T, Ekblom A, Hansson P. Relief of sinus pain by vibratory stimulation. *Ear Nose Throat J.* 1985;64:163–167.
13. Smith M, Berenger PG, Bonutti P, Ramakrishnan AP, Beyers J, Ramakrishnan V. Multimodal frequency treatment for facial pain caused by chronic rhinosinusitis: a pilot study. *Sinusitis.* 2017;2.
14. Høsoien E, Lund AB, Vasseljen O. Similar effect of therapeutic ultrasound and antibiotics for acute bacterial rhinosinusitis: a randomised trial. *J Physiother.* 2010;56:29–32.
15. Zelenkin EM, Prozorovskaia KN, Mirkin AS, Zavgorodniaia EG, Kandaurova AN. Low-frequency vibrations in the treatment of patients with acute maxillary sinusitis and ethmoiditis. Non-specific resistance and immunity. *Vestn Otorinolaringol.* 1998;41–43.
16. Kantor I, Szamborski R, Misztela A. Clinical application of magnetic pulsing fields in paranasal sinusitis treatment. *Otolaryngol Pol.* 1997;51:299–302. Suppl 25.
17. Zelenkin EM, Prozorovskaia KN, Petrovskaia AN, Zavgorodniaia EG, Kandaurova AN. Clinical and immunological aspects of vibration therapy for sinusitis. *Vestn Otorinolaringol.* 2000:66–69.
18. Feizabadi N, Sarrafzadeh J, Fathali M, et al. Quantitative analysis of staphylococcus aureus in patients with chronic rhinosinusitis under continuous ultrasound treatment. *Iran J Microbiol.* 2018;10:354–360.
19. Khudiev AM. Low-frequency ultrasonic therapy in combined treatment of chronic maxillary sinusitis. *Vestn Otorinolaringol.* 2003;44–46.
20. Zelenkin EM, Logunov AI, Tsyru'nikova LG. Course changes in acute inflammation of the frontal sinus based on the indices of lipid peroxidation and on the activity of enzyme during biovibration therapy. *Vestn Otorinolaringol.* 1998:32–34.
21. Nakhostin Ansari N, Naghdi S, Farhadi M. Therapeutic ultrasound as a treatment for chronic sinusitis. *Physiother Res Int.* 2004;9:144–146.
22. Ansari NN, Fathali M, Naghdi S, Hasson S. Effect of pulsed ultrasound on chronic rhinosinusitis: a case report. *Physiother Theory Pract.* 2010;26:558–563.
23. Ansari NN, Naghdi S, Farhadi M, Jalaie S. A preliminary study into the effect of low-intensity pulsed ultrasound on chronic maxillary and frontal sinusitis. *Physiother Theory Pract.* 2007;23:211–218.
24. nakhosin Ansari N, Naghdi S, Farhadi M. Physiotherapy for chronic rhinosinusitis: the use of continuous ultrasound. *Int J Ther Rehab.* 2007;14:306–310.
25. Ansari NN, Fathali M, Naghdi S, Hasson S, Jalaie S, Rastak MS. A randomized, double-blind clinical trial comparing the effects of continuous and pulsed ultrasound in patients with chronic rhinosinusitis. *Physiother Theory Pract.* 2012;28:85–94.
26. Nakhostin-Ansari A, Nazem A, Ansari NN, Fathali M, Naghdi S, Hasson S. Effects of pulsed ultrasound on olfactory dysfunction in patients with chronic rhinosinusitis: a pilot study. *Complement Ther Clin Pract.* 2021;44:101409.
27. Young D, Morton R, Bartley J. Therapeutic ultrasound as treatment for chronic rhinosinusitis: preliminary observations. *J Laryngol Otol.* 2010;124:495–499.
28. Rocha WA, Rodrigues KM, Pereira RR, Nogueira BV, Gonçalves WL. Acute effects of therapeutic 1-MHz ultrasound on nasal unblocking of subjects with chronic rhinosinusitis. *Braz J Otorhinolaryngol.* 2011;77:7–12.
29. De Castro RB, Cruz-Daylo MAB, Jardim MLA. Low Frequency ultrasound in chronic rhinosinusitis with nasal polyposis and recovery after endoscopic sinus surgery: a randomized controlled trial. *Phil J Otolaryngol Head Neck Surg.* 2017;32:6–13.
30. Khanwalkar A, Johnson J, Zhu W, Johnson E, Lin B, Hwang PH. Resonant vibration of the sinonasal cavities for the treatment of nasal congestion. *Int Forum Allergy Rhinol.* 2021;11:1308–1320.
31. Eby GA. Strong humming for one hour daily to terminate chronic rhinosinusitis in four days: A case report and hypothesis for action by stimulation of endogenous nasal nitric oxide production. *Med Hypotheses.* 2006;66:851–854.
32. Juto JE, Axelsson M. Kinetic oscillation stimulation as treatment of non-allergic rhinitis: an RCT study. *Acta Otolaryngol.* 2014;134:506–512.
33. Cairns A, Bogan R. The SinuSonic: reducing nasal congestion with acoustic vibration and oscillating expiratory pressure. *Med Devices (Auckl).* 2019;12:305–310.
34. Soler ZM, Nguyen SA, Salvador C, et al. A novel device combining acoustic vibration with oscillating expiratory pressure for the treatment of nasal congestion. *Int Forum Allergy Rhinol.* 2020;10:610–618.
35. Ansari NN, Fathali M, Naghdi S, Bartley J, Rastak MS. Treatment of chronic rhinosinusitis using erythromycin phonophoresis. *Physiother Theory Pract.* 2013;29:159–165.
36. Ansari NN, Naghdi S, Fathali M, Bartley J, Rastak MS. A randomized clinical trial comparing pulsed ultrasound and erythromycin phonophoresis in the treatment of patients with chronic rhinosinusitis. *Physiother Theory Pract.* 2015;31:166–172.
37. Maniscalco M, Sofia M, Weitzberg E, Lundberg JO. Sounding airflow enhances aerosol delivery into the paranasal sinuses. *Eur J Clin Invest.* 2006;36:509–513.
38. Patel ZM, Hwang PH, Chernomorsky A, et al. Low-frequency pulsed ultrasound in the nasal cavity and paranasal sinuses: a feasibility and distribution study. *Int Forum Allergy Rhinol.* 2012;2:303–308.
39. Zippel R, Conrad M, Mayer P. On the penetrability of the maxillary sinus ostium for common and ultrasonic vibration-aerosols in chronic maxillary sinusitis. *Z Laryngol Rhinol Otol.* 1968;47:610–620.
40. Saijo R, Majima Y, Hyo N, Takano H. Aerosol deposition in sinus cavities treated by functional endoscopic sinus surgery 2000;43:11–14.

41. Gerber V, Buskina AV, Dergachev VS. Application of ultrasonic cavitation in combined treatment of polypous rhinosinusitis. *Vestn Otorinolaringol.* 2003;25–27.
42. Dañniak LB, Sukhneva TP, Nikitina IM. Treatment of exudative maxillary sinusitis with low frequency ultrasound. *Vestn Otorinolaringol.* 1989;27–32.
43. Reychler G, Colbrant C, Huart C, et al. Effect of three-drug delivery modalities on olfactory function in chronic sinusitis. *Laryngoscope.* 2015;125:549–555.
44. Goektas O, Lau L, Olze H. Treatment of chronic rhinosinusitis with pressure-pulsed corticosteroid inhalation. *Indian J Otolaryngol Head Neck Surg.* 2013;65:402–405.
45. Poletti SC, Batashev I, Reden J, Hummel T. Olfaction in chronic rhinosinusitis: Comparing two different endonasal steroid application methods. *Eur Arch Otorhinolaryngol.* 2017;274:1431–1435.
46. Valentine R, Athanasiadis T, Thwin M, Singhal D, Weitzel EK, Wormald PJ. A prospective controlled trial of pulsed nasal nebulizer in maximally dissected cadavers. *Am J Rhinol.* 2008;22:390–394.
47. Katsumi N, Sugano S, Ishizuka Y. Clinical experience of using a pressurized and vibrating nebulizer-PARI sinus. *Oto-Rhino-Laryngology Tokyo.* 2008; 51:15–21.
48. Ohki M, Yoshikawa M, Yamaguchi S, et al. The effect of nebulization with a vibrating pulse in patients with resistive sinusitis. *Oto-Rhino-Laryngology Tokyo.* 2013; 56:7–11.
49. Möller W, Schuschnig U, Khadem Saba G, et al. Pulsating aerosols for drug delivery to the sinuses in healthy volunteers. *Otolaryngol Head Neck Surg.* 2010;142:382–388.
50. Möller W, Schuschnig U, Celik G, et al. Topical drug delivery in chronic rhinosinusitis patients before and after sinus surgery using pulsating aerosols. *PLoS One.* 2013;8:e74991.
51. Mainz JG, Schien C, Schiller I, et al. Sinonasal inhalation of dornase alfa administered by vibrating aerosol to cystic fibrosis patients: a double-blind placebo-controlled cross-over trial. *J Cyst Fibros.* 2014;13:461–470.
52. Mainz JG, Schumacher U, Schädlich K, et al. Sino nasal inhalation of isotonic versus hypertonic saline (6.0%) in CF patients with chronic rhinosinusitis – results of a multicenter, prospective, randomized, double-blind, controlled trial. *J Cyst Fibros.* 2016;15:e57–e66.
53. Ishman SL, Martin TJ, Hambrook DW, Smith TL, Jaradeh SS, Loehrl TA. Autonomic nervous system evaluation in allergic rhinitis. *Otolaryngol Head Neck Surg.* 2007;136:51–56.
54. Chen YW, Tzeng JI, Huang PC, Hung CH, Shao DZ, Wang JJ. Therapeutic ultrasound suppresses neuropathic pain and upregulation of substance P and neurokinin-1 receptor in rats after peripheral nerve injury. *Ultrasound Med Biol.* 2015;41:143–150.
55. Karosi T, Sziklai I, Csomor P. Low-frequency ultrasound for biofilm disruption in chronic rhinosinusitis with nasal polyposis: in vitro pilot study. *Laryngoscope.* 2013;123:17–23.
56. Runyan CM, Carmen JC, Beckstead BL, Nelson JL, Robison RA, Pitt WG. Low-frequency ultrasound increases outer membrane permeability of pseudomonas aeruginosa. *J Gen Appl Microbiol.* 2006;52:295–301.
57. Newbold ME, Tullis E, Corey M, Ross B, Brooks D. The flutter device versus the PEP mask in the treatment of adults with cystic fibrosis. *Physiother Can.* 2005;57:199–207.
58. Durieux N, Vandenput S, Pasleau F. OCEBM levels of evidence system. *Rev Méd Liège.* 2013;68:644–649.

SUPPORTING INFORMATION

Additional supporting information may be found in the online version of the article at the publisher's website.

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