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CLINICAL EFFECTIVENESS OF GLUCOSAMINE AND CHONDROITIN SULPHATE IN TREATMENT OF OSTEOARTHRITIS

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ABSTRACT: Osteoarthritis is a form of arthritis, and is the most common form of arthritis. Persons suffering from osteoarthritis have symptoms of pain, stiffness, decreased range of motion of affected joints. Although NSAIDS are the most commonly prescribed agents for this disorder but can cause of serious adverse effects. Two compounds Glucosamine and chondroitin which are extracted from animal products have been used in various forms for OA. To assess the clinical effectiveness of glucosamine and chondroitinsulphate in treatment of osteoarthritis symptoms like joint pain, joint space narrowing, reduced walking time, swelling etc. We searched articles separately for glucosamine and chondroitin sulphate using internet. Fifteen articles met the inclusion criteria. Data from articles was extracted using a standardized data extraction tables i.e. table 1 and table 2. Glucosamine and Chondroitin sulphate are effective in the treatment of Osteoarthritis because these canreducepain, prevent further joint space narrowing and solve other related problems of this disease. The two agents can be used in osteoarthritis treatment as their safety is already assured as compared to other symptomatic treatment for OA. But these agents can take more time to treat disease as compared to conventional medicine like NSAIDS.

INTRODUCTION: Osteoarthritis is a form of arthritis, and is the most common form of arthritis. ^{1, 2}. In 2010-2011, over 4.6 million Canadians (16.7% of those 15 years and older) reported suffering from arthritis ³⁹. Osteoarthritis is classified on the basis of its cause. Two classes of osteoarthritis are primary or idiopathic osteoarthritis and secondary osteoarthritis. Primary osteoarthritis occurs due to unknown causes but is strongly associated with age.

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Secondary OA develops as a result of joint injury, infection, hereditary, developmental, metabolic or neurologic disorders. Secondary osteoarthritis occurs less frequently ^{3, 4}. Friction between the bones is resulted by gradual wear and loss of cartilage in the joints, which Causes pain and swelling in affected joints. For a long time it was thought that in osteoarthritis only cartilage is affected. But, now it is known that the underlying bone synovium also undergoes changes ⁵⁻⁷.

In Osteoarthritis joint movement suffers additional restriction due to the reaction of prearticular bone with osteophyte formation. It predominates in weight-bearing joints, such as the knee and hip ⁸. There are many risk factors known of OA which include; age ⁹, Over weight and obesity ¹⁰, genetic determinants ^{11, 12}. Persons suffering from

osteoarthritis have symptoms of pain, stiffness, decreased range of motion of affected joints ¹³. It is the leading cause of pain and physical disability in older people ¹⁴. A biomechanical abnormality to the joint or limb may be present in osteoarthriris ⁴⁰. There are still questions concerning the causal factors of OA. The nature of the initiating event is often unknown, although many processes involved in the progression of OA are known. Due to disruption of the cartilage collagen matrix, the water content of the cartilage increases ⁴³.

Osteoarthritis-affected joints are commonly tender. Patients suffer from morning and/or prolonged fixed body position stiffness. Swelling and crepitus may also be evident. Generally, pain escalates with increasing activity throughout the day and many patients need frequent breaks to rest the involved joint ⁴¹. The use of NSAIDs has a palliative effect and can cause adverse effects in the long-term. Therefore, effective and safe treatments for the control and management of osteoarthrosis of the Temporomandibular Joint are the use Chondroitin sulphate and Glucosamine 44. For treatment of osteoarthritis only few effective remedies are available. 15 Primary concern of currently available medical therapies osteoarthritis is treatment of joint pain in patient.¹⁶

Analgesics well as traditional and as cyclooxygenase-2-selective non-steroidalantiinflammatory drugs (NSAIDs) are effective and are widely used ^{17, 18}. Although NSAIDS are the most commonly prescribed agents for this disorder but can cause of serious adverseeffects 19, 20. Two compounds Glucosamine and chondroitin which are extracted from animal products have been used in various forms for OA ²¹. These compounds are found modestly effective but because of their safety, these would have high utility in the treatment of OA ^{22, 23}. Chondroitin sulphate reduces both cartilage volume loss and bone marrow lesions in knee osteoarthritis patients starting as early as 6 months after initiation of therapy. ⁴²

MATERIALS AND METHODS:

Data Sources

To search the original articles of both glucosamine and chondroitin sulphate we searched the electronic data bases from 1980 to 2011 including: Science direct.com, American college of rheumatology (arthritis and rheumatism), Pub Med and American medical association. From there we selected the articles which met our inclusion criteria.

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Inclusion criteria

All the published trials on arthritis of various parts of body and in which preparations were given orally (in form of tablets or powder) were included. Comparisons in trials of glucosamine and chondroitin were mostly with placebo but trials for comparison of glucosamine and chondroitin with NSAIDS were also included. Duration of study should be at least one month because these agents may take time to produce effect.

Data extraction

Thirteen articles met the inclusion criteria. Data from that articles were extracted using a standardized data extraction table. In **Table 1** and **Table 2**, we notified the author/year of article, duration of study, dose of agent tested, outcome measured and conclusion of the article. And from the **Table 1** and **Table 2**, having all material for review, we drew conclusion of our review.

RESULTS:

Table 1 and table 2 summarizes prospective data based on the use of glucosamine and chondroitin sulfate in the treatment of osteoarthritis in which names of authors along with years have been given. Table 1 and table 2 also contains number of patients, their dosage, duration of intervention, type of intervention and conclusion based on these interventions.

DISCUSSIONS: Objective of this review is to assess the clinical effectiveness of glucosamine and chondroitin sulphate in reducing pain and preventing joint space narrowing and other problems that OA patients face, hence the overall effectiveness of these agents for osteoarthritis treatment and their role in progression of osteoarthritis disease. To collect articles we set inclusion criteria, according to which published articles of glucosamine and chondroitin sulphate were collected and reviewed by forming standard tables i.e. **Table 1** and **Table 2**.

In three articles of glucosamine sulphate ^{24, 25, 29} outcome measure is joint pain and from the conclusions of two articles in which study was

conducted by comparing glucosamine to placebo we can see that glucosamine is superior to placebo to reduce pain. In one article study was conducted by comparing glucosamine to ibuprofen both agents showed almost equal success (glucosamine: 48%, ibuprofen: 52%) but ibuprofen showed effect sooner than glucosamine sulphate.

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TABLE 1: SHOWS THE USE OF GLUCOSAMINE IN OSTEOARTHRITIS TREATMENT

		Glucosamine			
Author, year	No. of patients	Dosage	Duration	Type of intervention	Conclusion
Drovanti A et al. 1980	80	500mg t.i.d	30 days	Articular Joint pain, Joint tenderness Swelling, and Range of motion	Interventions were found to be significantly improved in the GS group than in the placebo group.
Pujalte JM et al. 1980 ²⁵	20	500mg t.i.d	6 to 8 weeks	Joint pain, Joint tenderness, and Swelling.	GS is superior to placebo in improving outcome measures.
Noack W et al.1994 ²⁶	252	500mg t.i.d	4 weeks	Lequesne's index	Decreased by 3.2 points in the GS group and only 2.2 points in the placebo group.
Reginster JY et al.2001	212	1500mg o.d	3 years	Joint space width	Joint space narrowing In GS group: 0 In placebo:31mm
Pavelka K et al.2002 ²⁸	202	1500mg t.i.d	3 years	Worsening osteophytes	20% in placebo group and 6% in GS group
Fassbender HM et al.1994 ²⁹	200	GS: 1500mg o.d Ibuprofen: 1200mg o.d	3 months	Joint pain	52% pain reduction observed in ibuprofen group while 48% in GS group but effect occurred sooner in ibuprofen group
Bruyere et al. 2004 ³⁰	319 postmenopausal Women	1500 mg o.d	3 years	Radiographs of the knee: joint space narrowing.	GlcN·S: no significant joint space loss Plac.: Progressive joint space narrowing.
Kawasaki 2008 ³¹	142	1500 mg o.d	18 months	Joint space width	0.0 mm in GS group and - 0.31 mm in control group

TABLE 2: SHOWS THE USE OF CHONDROITIN SULPHATE IN OSTEOARTHRITIS TREATMENT

Author, year	No. of patients	Dosage	duration	Type of intervention	Conclusion
Bourgeois P et al.1998 32	127	1200mg o.d	3 months	Joint pain	Improved mean spontaneous
Uebelhart D et al. 1998 33	46	800 mg o.d	1 year	Joint space width	joint pain was observed 0 mm in Cs group but
Gebeniait D et al. 1998	40	ooo iiig o.u	i yeai	Joint space width	increased by 0.4 mm in
					placebo group
Author, year	No. of patients	Dosage	Duration	Outcome measures	Conclusion
Bucsi L et al. 1998 34	80	800 mg o.d		Pain VAS,	Pain and lequense's index
				Lequense's index	decreased in CS group.
				Walking time	Walking time improved in CS
Verbruggen G et al.	119	400 mg t.i.d	3 years	New erosive OA of finger	group. CS protect from the
1998 ³⁵	119	400 Hig t.i.u	3 years	joints	development of erosive
1,7,0				Joines	changes in patients with
					finger joint OA.
Cem Gabay et al. 2011 36	162	800 mg o.d	6 months	Hand pain	Decrease in the hand pain in
					the CS group than in the
Kahan et al. 2009 37	622	900ma a d	2 *****	icint angas namawina	placebo group is observed.
Kanan et al. 2009	022	800mg o.d	2 yrs	joint space narrowing	28% CS pts. Versus 41% Plac. pts.
					showed joint space
					Narrowing.
Michel et al. 2005 38	300	800 mg o.d	2 years	Joint space narrowing	In CS group no significant
					joint space loss and
					significant joint space
					narrowing in placebo group.

In three included studies ^{27, 30, 31} improvement in joint space narrowing is observed and conclusion of those articles show that GS is very effective in preventing joint space narrowing.

Two studies ^{24, 25} show improvement in joint tenderness and swelling. One study ²⁶ concluded decrease in lequence's index by 3.2 points. One article ²⁸ has outcome measure of worsening

osteophytes which is very less in placebo group than in GS group. Among included articles of Chondroitin sulphate outcome measure is joint space width in three articles ^{33, 37, 38} which showed that chondroitin sulphate is prominently superior to placebo in preventing further joint space narrowing.

Three studies ^{32, 34, 36} included were conducted to know the effect of chondroitin sulphate in pain reduction and conclusion of that studies showed that chondroitin sulphate effectively decrease the joint pain and was found to be better than placebo. One study ³⁴ show decrease in lequence's index and improved walking time in the CS group. Another study ³⁵ concluded that CS may protect against the development of erosive changes in patients with finger joint OA.

CONCLUSIONS: According to conducted review it is concluded that Glucosamine and Chondroitin sulphate are effective in the treatment of Osteoarthritis because these are found to be better placebo in reducing pain and more prominently effective in preventing further joint space narrowing already present in patients of OA. Other problems which the patients of this disease have to face like swelling and walking time are also improved by these chondroprotective agents. The two agents are also found to be effective in reducing lequence's index. So the two agents can be used in osteoarthritis treatment as their safety is already assured as compared to other symptomatic treatment for OA (NSAIDS cause severe damage to gastro protective layer). But these agents can take more time to treat disease than the conventional medicine like NSAIDS.

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