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Why?

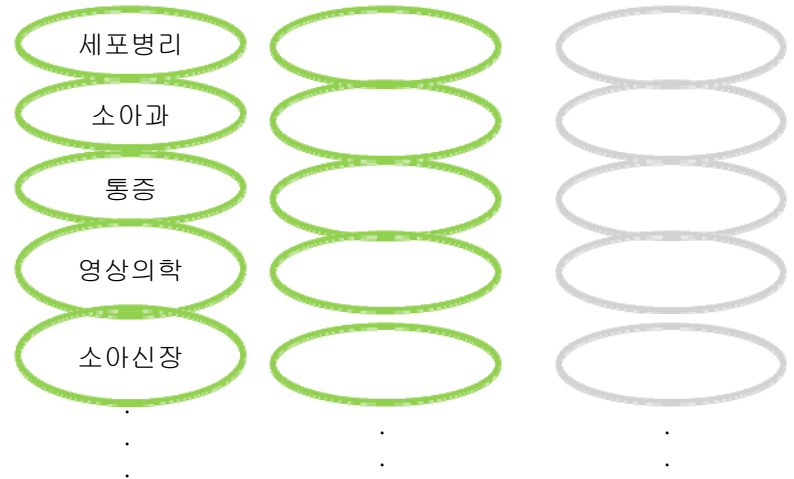
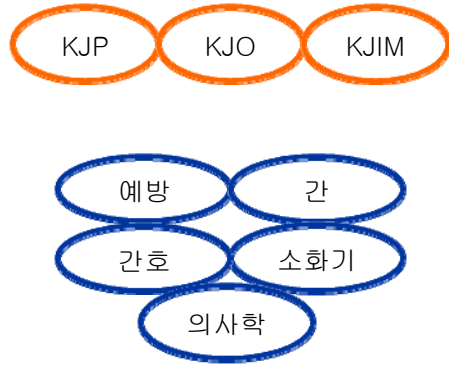
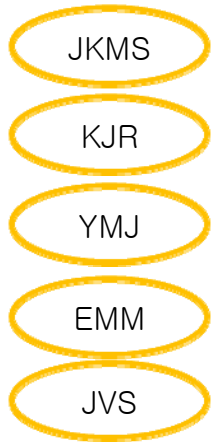
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KoMCI 사업단



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1: *Korean J Gastrointest Endosc.* 2007 May;34(5):244-250. Korean.

Full text article at the journal web site

The Usefulness of Endoscopic Ultrasonography for Discriminating Gastric Mucosal Cancer from Submucosal Invasion.

Lee SH, Jang YS, Jeon SH, Kim SY, Jang BK, Chung WJ, Cho KB, Park KS, Hwang JS.

Department of Internal medicine, Keimyung University School of Medicine, Daegu, Korea. seenae99@dsmc.or.kr

BACKGROUND/AIMS: Endoscopic ultrasonography (EUS) has been used to discriminate gastric mucosal cancer (T1m) from submucosal invasion (T1sm). Thus the aims of this study are 1) to determine the accuracy of EUS for diagnosing tumor depth, 2) to compare the accuracy of EUS with the endoscopic impressions of variously experienced endoscopists and 3) to compare the accuracy of performing EUS by one doctor according to the experience. METHODS: The EUS and pathologic reports of early gastric cancer patients were analyzed. The same endoscopic images were reviewed again by 3 endoscopists, who had one-, three- and five-years experience, respectively. The accuracies of EUS and conventional endoscopy were analyzed. RESULTS: 77 patients were included from November 2003 to October 2005. The kappa of the EUS for actual examiner and conventional endoscopy for reviewer 1, reviewer 2 and reviewer 3 were 0.421, 0.134, 0.359 and 0.307, respectively and accuracies were 68.8%, 45.5%, 67.5% and 62.3%, respectively. Of the 52 T1m patients, 23 (44.2%) were overstaged as T1sm with performing EUS. But of 25 T1sm patients, only 1 (4.0%) was understaged as T1m with performing EUS. The accuracy and kappa of the EUS for one doctor during the first-year experience were 60.6% and 0.316, respectively, and they were 75.0% and 0.508, respectively during the second-year experience. CONCLUSIONS: EUS is useful for complementing the conventional endoscopic discrimination of gastric mucosal cancer from submucosal invasion. Yet physician should keep in mind the relatively common overstaging.

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1: *J Korean Med Sci*. 2007 Jun;22(3):588-594. Eng

JKMS FULL-TEXT ARTICLE
Journal of Korean Medical Science

Successful Combined Treatment with I

Cho KY, Lee SJ, Burm JS, Park EA.

Department of Pediatrics, Ewha Womans University
Department of Plastic Surgery, Ewha Womans University

Extravasation injuries in the neonatal intensive care unit are not rare during parenteral hyperalimentation. There have been many different methods of management. We report five premature infants with wounds of hyperalimentation fluid extravasation managed by the antibacterial ointment (Terramycin ophthalmic ointment(TM)) and sesame oil and an antiinflammatory herbal mixture (HEBO(TM)). The mean gestational age of patients was 31(+2) weeks (range, 28 (+4) to 35(+6) weeks), and the mean weight at extravasation was 1,930 g (range, 1,140 to 2,680 g). Extravasation occurred within the mean of 32 days (range, 17 to 50 days). The method of dressing was application of a thick layer of this mixture covered by vaseline and wet gauze renewed at an interval of 8-12 hr after irrigating the wounds thoroughly with normal saline. The mean duration of dressing was 30 days (range, 20-50 days). The wounds had healed completely leaving a small size of contracture without functional abnormality. We conclude that this therapy may be considered for an alternative treatment and warrants clinical trials for the confirmation of the local management of extravasation injury.

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Journal of Korean Medical Science

1031-9924 Print
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2007 Vol. 22 No.1 No.2 No.3 No.4 No.5 No.6

Abstract Abstract + references Full-text PDF

J Korean Med Sci. 2007 Jun;22(3):588-594.

Successful Combined Treatment with Total Parenteral Nutrition Fluid Extravasation Injuries in Preterm Infants

Ky Young Cho, Soo Jung Lee, Jin Sik Burm, Eun Ae Park

Department of Pediatrics, Ewha Womans University College of Medicine, 911-1 Mok-dong, Yangcheon-gu, Seoul, Korea. pea8639@ewha.ac.kr, Department of Plastic Surgery, Ewha Womans University College of Medicine, Seoul, Korea.

Extravasation injuries in the neonatal intensive care unit are not rare during parenteral hyperalimentation. There have been many different methods of management. We report five premature infants with wounds of hyperalimentation fluid extravasation managed by the antibacterial ointment (Terramycin ophthalmic ointment(TM)) and sesame oil and an antiinflammatory herbal mixture (HEBO(TM)). The mean gestational age of patients was 31(+2) weeks (range, 28 (+4) to 35(+6) weeks), and the mean weight at extravasation was 1,930 g (range, 1,140 to 2,680 g). Extravasation occurred within the mean of 32 days (range, 17 to 50 days). The method of dressing was application of a thick layer of this mixture covered by vaseline and wet gauze renewed at an interval of 8-12 hr after irrigating the wounds thoroughly with normal saline. The mean duration of dressing was 30 days (range, 20-50 days). The wounds had healed completely leaving a small size of contracture without functional abnormality. We conclude that this therapy may be considered for an alternative treatment and warrants clinical trials for the confirmation of the local management of extravasation injury.

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주소(D) http://koreamed.org/SearchBasic.php?RID=94726&DT=1&QY=%22Yonsei+Med+J%22+%5B%5D%5D%5D++AND+1960+%5BDPY%5D 이동

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1: *Yonsei Med J.* 1960 Dec;1(1):9-16. English

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Hong PW.
Department of Surgery, Yonsei University College of Medicine, Seoul, Korea

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Yonsei Medical Journal

Yonsei Med J 1960 December, 1(1): 9-16

Title	Influence of Epinephrine, Cortisone Acetate and Adrenocorticotrophic Hormone on Gastric Secretion Full Text PDF
Authors	Pill Whoon Hong
Affiliation	Department of Surgery, Yonsei University College of Medicine, Seoul, Korea
Abstract	Not available
Keywords	Not available
Corresponding Author info	Dr. Pill Whoon Hong, Department of Surgery, Yonsei University College of Medicine, Seoul, Korea

Yonsei University College of Medicine, 134 Shinchon-dong, Seodaemun-gu, Seoul 120-752, Korea
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오후 6:01

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- 2007년 7월 현재:
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 - 45종 학술지 원문으로 LinkOut
 - 83종 학술지 Retro 사업 참여
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Enter information about the article you wish to find.

CITED AUTHOR: Enter the cited author's last name and initials (e.g., Jang YS for Jang Yeong Sil).

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CITED YEAR: Enter year Cited Journal was published as 2001 OR 2002.

SOURCE YEAR (Limit from 2000): Enter year Source Journal was published as 2001 OR 2002.

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- For Date, you may enter yyyy. For example, 2002.
- Author names are automatically truncated to account for varying initials, e.g., Jang Y will also match on Jang YA, Jang YB, Jang YC jr, etc. Enclose author names in double quotes to retrieve the exact match, e.g., "Jang Y".

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Cited Reference Search Results

567 references matched query:
Cited Author=Lee DH

The table lists all of the cited references that match your search request. Select all desired references (including variants) by clicking the checkboxes or . Then press .

Sort: Hits Display: 10
Items 1-10 of Page 1 of 57 Select page: [12345678910>>](#)

	Hits	Cited Author	Cited Journal	Volume	Page	Year
<input type="checkbox"/>	16	Lee DH	J Korean Pediatr Soc	34	445	1991
<input type="checkbox"/>	13	...Lee DH	J Korean Pediatr Soc	32	597	1989
<input type="checkbox"/>	11	...Lee DH	J Korean Ophthalmol Soc	39	658	1998
<input type="checkbox"/>	10	Lee DH	J Korean Pediatr Soc	39	1055	1996
<input type="checkbox"/>	9	...Lee DH	Korean J Anesthesiol	38	251	2000
<input type="checkbox"/>	8	Lee DH	J Korean Med Sci	17	457	2002
<input type="checkbox"/>	8	...Lee DH	Korean J Gastrointest Endosc	19	81	1999
<input type="checkbox"/>	7	...Lee DH	J Korean Ophthalmol Soc	39	2211	1998
<input type="checkbox"/>	7	...Lee DH	J Korean Pediatr Soc	32	597	1989
<input type="checkbox"/>	7	...Lee DH	Korean J Gastroenterol	36	457	2000

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Cited References Search Results - Full Record

Epidemiological Findings of Hepatitis B Infection Based Survey in Korea

Lee DH, Kim JH, Nam JJ, Kim H

J Korean Med Sci
17(4):457-462 Aug 2002. E

Total References:19 Cited Korean Referenc



The purpose of this study was to investigate the epidemiologic (HBV) infection in Korea based on the 1998 National Health and consisted of 9,771 aged 10 yr or over, who were selected from probability sampling design. The prevalence of hepatitis B surface age, sex, residency, household income, education, family history, out. The prevalence of HBsAg was 5.1% (95% confidence interval (CI): 4.5-5.7) in males and 4.1% (95% CI: 3.6-4.6) in females with a low prevalence in those under 20 yr old. Generally, HBsAg seropositivity by administrative area was similar with the exception of Jeju province. HBsAg seropositivity of Jeju island was approximately three times higher in both men and women, as compared with the national average. HBsAg seropositivity by socioeconomic status unexpectedly showed a very consistent positive association in both gender. Comparing HBsAg seropositivity by the frequency of eating out, in both gender, the more frequent they ate out, the higher it was. Our study suggested that there might be another transmission route of HBV, which is possibly related to diet.

Affiliation:

Department of Preventive Medicine, College of Medicine, Kosin University, Busan, Korea. hrshin@ncc.re.kr
Korea Institute of Health and Social Affairs, Seoul, Korea.

Division of Cancer Epidemiology & Control, National Cancer Center Research Institute, Koyang, Korea.

Cited References

Epidemiological Findings of Hepatitis B Infection Based on 1998 National Health and Nutrition Survey in Korea

Lee DH, Kim JH, Nam JJ, Kim HR, Shin HR.

J Korean Med Sci
17(4):457-462 Aug 2002. English.

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Items 1-7 of Page 1 of 1

Select page: 1

Cited Author	Cited Journal	Volume	Page	Year
	J Korean Med Assoc	26	425	1983
Kim JS	Korean J Epidemiol	8	115	1986
	Korean J Epidemiol	9	40	1987
Park BJ	Korean J Epidemiol	11	263	1989
	Korean J Epidemiol	11	98	1989
	Korean J Epidemiol	14	54	1992
Kim SJ	Korean J Intern Med	31	313	1986

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Items 1-7 of Page 1 of 1

Select page: 1

Citing Articles - Summary

Epidemiological Findings of Hepatitis B Infection Based on 1998 National Health and Nutrition Survey in Korea

Lee DH, Kim JH, Nam JJ, Kim HR, Shin HR.

J Korean Med Sci
17(4):457-462 Aug 2002. English.

Sort: Date

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Items 1-5 of 8 Page 1 of 2

These documents in the database cite the

- 1: Eun JR.
Comparison of Clinical Outcome Between Acute Exacerbation After Viral breakthrough and Chronic Hepatitis B Infection. Korean J Hepatol. 2006 Jun;12(2):184-188.
- 2: Shin BM, Yoo HM, Lee AS, Park SK.
Seroprevalence of Hepatitis B Virus among Korean Adults. J Korean Med Sci. 2006 Feb;21(1):58-61.
- 3: Kang HS, Song BC, Ji CX, Kim SY, Kim S.
Serologic Markers of Hepatitis B Virus in Korean Adults. Korean J Hepatol. 2004 Sep;10(3):191-194.
- 4: Lim YS, Suh DJ.
Current Antiviral Therapy for Chronic Hepatitis B. J Korean Med Sci. 2004 Aug;19(4):489-494.
- 5: Kim TH, Kim YS, Yeom JJ, Cho EY, Kim S.
Relevancy between Liver Injury, Serum ALT, and HBsAg Carriers. Korean J Gastroenterol. 2004 Aug;44(2):100-103.

Cited References Search Results - Full Record

Current Antiviral Therapy for Chronic Hepatitis B

Lim YS, Suh DJ.

J Korean Med Sci
19(4):489-494 Aug 2004. English.

Total References:43 Cited Korean References:3 Times Cited:1



During the past decade, major breakthroughs have been achieved in treatment of chronic hepatitis B. Currently, three therapeutic agents are approved for chronic hepatitis B: interferon-alpha, lamivudine and adefovir dipivoxil. In patients with HBeAg-positive chronic hepatitis B, all of these drugs achieve HBeAg loss (24-33%) and anti-HBe seroconversion (12-30%) rates that are superior to those observed in untreated controls. Interferon-alpha has several drawbacks, such as the parenteral administration and the development of frequent and potentially serious side effects. Lamivudine is a safe drug with rare and generally mild side effects. Lamivudine induces an initial virological remission in 70-90% of patients, but only 30-40% of patients remain in remission after the third year due to progressively increasing viral resistance. The main advantage of adefovir dipivoxil is the rare emergence of resistance, which has been identified in less than 2% of patients at 2 yr of treatment. Adefovir is also effective against lamivudine-resistant strains. This review will focus on the natural history and recently gained knowledge on the treatment of chronic hepatitis B.

Affiliation:

Department of Internal Medicine, University of Ulsan, College of Medicine, Asan Medical Center, Seoul, Korea. djsuh@amc.seoul.kr

probability sampling design. age, sex, residency, household. The prevalence of HBsAg (95% CI: 3.6-4.6) in female seropositivity by administrative of Jeju island was approximately national average. HBsAg seropositivity by socioeconomic status unexpectedly showed a very consistent positive association in both gender. Comparing HBsAg seropositivity by the frequency of eating out, in both gender, the more frequent they ate out, the higher it was. Our study suggested that there might be another transmission route of HBV, which is possibly related to diet.

Affiliation:

Department of Preventive Medicine, College of Medicine, Kosin University, Busan, Korea. hrshin@ncc.re.kr
Korea Institute of Health and Social Affairs, Seoul, Korea.
Division of Cancer Epidemiology & Control, National Cancer Center Research Institute, Koyang, Korea.

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8

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- ALLERGY
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IF: Impact Factor
ZIF: Z Impact Factor
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Citing Half-Life

KoMCI 2006

Bibliographic Data IF: Impact Factor
Citation Information ZIF: Z Impact Factor

Bibliographic Data

Full Journal Title	Journal of Korean Medical Science
Abbreviated Journal Title	J Korean Med Sci
ISSN	1011-8934
Issues / Year	6
Publisher	Korean Academy of Medical Science
Publisher Address	Korean Academy of Medical Science, 147-750, Seil-daero, Yongsu-gu, Seoul 140-721, South Korea
Subject Categories	MEDICINE, GENERAL AND INTERNAL; MEDICINE, RESEARCH

Citation Information

CITED JOURNAL DATA CITING JOURNAL DATA

No.	Abbreviated Journal Title	ISSN	Articles
1	J Korean Med Sci	1011-8934	216

KoMCI 2006

Cited Journal: Journal of Korean Medical Science (ISSN: 1011-8934)

Number of times articles published in 2006 (in journals below) cited articles published in J Korean Med Sci (in years below).

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Rank.	IF	Citing Journal	All	2006	2005	2004	2003	2002	2001	2000	1999	1998	1997	Rest
		All Journals	271	10	37	34	31	25	17	27	11	13	8	58
1	0.186	J Korean Med Sci	26	3	4	8	4	1	1	1	1	1	0	2
2	0.185	Korean J Med	15	0	3	2	1	0	0	2	0	1	1	5
3	0.088	Korean J Pathol	11	1	0	0	1	2	2	0	0	0	0	5
4	0.164	Korean J Pediatr*	10	0	3	0	2	0	0	0	0	1	0	4
5	0.213	Korean J Gastrointest Endosc	10	0	0	2	0	0	1	0	1	0	0	6
6	0.215	Korean J Dermatol	10	1	1	0	2	2	1	1	0	2	0	0
7	0.128	J Korean Neurol Assoc	9	1	1	0	0	1	0	4	0	0	0	2
8	0.241	Korean J Nephrol	9	0	1	0	0	1	3	1	0	0	1	2
9	0.146	Infect Chemother	8	0	1	1	2	0	0	0	1	1	1	1
10	0.045	J Korean Med Assoc	8	0	4	2	0	0	1	1	0	0	0	0

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Number of times articles published in journals below (in years below) where cited in J Korean Med Sci in 2006.

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Rank.	IF	Cited Journal	All	2006	2005	2004	2003	2002	2001	2000	1999	1998	1997	Rest
		All Journals	246	3	19	41	34	25	24	21	20	15	9	35
1	0.186	J Korean Med Sci	26	3	4	8	4	1	1	1	1	1	0	2
2	0.137	Korean J Asthma Allergy Clin Immunol	18	0	0	2	1	1	4	6	0	2	0	2
3	0.164	Korean J Pediatr	13	0	2	1	3	1	0	1	0	2	1	2
4	0.185	Korean J Med	12	0	0	4	0	1	1	0	2	0	0	4
5	0.146	Infect Chemother	11	0	2	2	0	0	4	0	0	0	0	3
6	0.215	Korean J Dermatol	8	0	2	0	2	1	1	0	0	0	0	2
7	0.151	J Korean Neurosurg Soc	8	0	1	0	2	0	2	0	1	1	0	1
8	0.145	Yonsei Med J	8	0	0	2	0	1	2	2	0	0	0	1
9	0.254	Korean J Intern Med	7	0	1	0	2	2	1	0	0	0	0	1
10	0.193	Korean J Obstet Gynecol	7	0	0	2	1	3	0	0	0	1	0	0

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 Citation Information ZIF: Z Impact Factor

IF: Impact Factor
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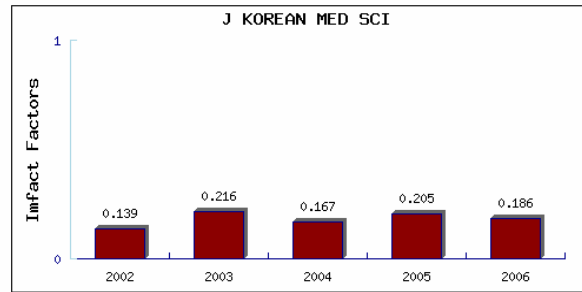
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 2004 = 34
 Sum = 71

Calculation: $\frac{\text{Cites to recent articles}}{\text{Number of recent articles}} = \frac{71}{382} = 0.186$

ZIF: Z Impact Factor
 ZIF TREND

Excluding Journal self-citation 2005 = 33
 Cites in 2006 to articles published in: 2004 = 26
 Sum = 59

Calculation: $\frac{\text{Cites to recent articles}}{\text{Number of recent articles}} = \frac{59}{382} = 0.154$



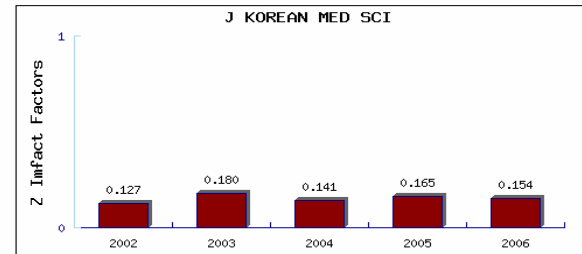
2006 IF: Impact Factor

Cites in 2006 to articles published in: 2005 = 37 Number of articles published in: 2005 = 207
 2004 = 34 2004 = 175
 Sum = 71 Sum = 382

Calculation: $\frac{\text{Cites to recent articles}}{\text{Number of recent articles}} = \frac{71}{382} = 0.186$

2005 IF: Impact Factor

Cites in 2005 to articles published in: 2004 = 37 Number of articles published in: 2004 = 175
 2003 = 35 2003 = 176
 Sum = 72 Sum = 351



2006 ZIF: Z Impact Factor

Excluding Journal self-citation
 Cites in 2006 to articles published in: 2005 = 33 Number of articles published in: 2005 = 207
 2004 = 26 2004 = 175
 Sum = 59 Sum = 382

Calculation: $\frac{\text{Cites to recent articles}}{\text{Number of recent articles}} = \frac{59}{382} = 0.154$

2005 ZIF: Z Impact Factor

Excluding Journal self-citation
 Cites in 2005 to articles published in: 2004 = 28 Number of articles published in: 2004 = 175
 2003 = 30 2003 = 176
 Sum = 58 Sum = 351

KoMCI 2006

Bibliographic Data

IF: Impact Factor

Journal Cited Half Life

Journal Citing Half-Life

Citation Information

ZIF: Z Impact Factor

Cited Journal Graph

Citing Journal Graph

Journal Cited Half-Life

The cited half-life for *J Korean Med Sci* is the median age of its articles cited in the KoMCI 2006. Half of the citations to *J Korean Med Sci* are to articles published within the cited half-life.

Cited Half-Life: 4.94 years

Cited Year	2006	2005	2004	2003	2002	2001	2000	1999	1998	1997	1996-all
No. of Cites from 2006	10	37	34	31	25	17	27	11	13	8	58
Cumulative %	3.69	17.34	29.89	41.33	50.55	56.83	66.79	70.85	75.65	78.6	100

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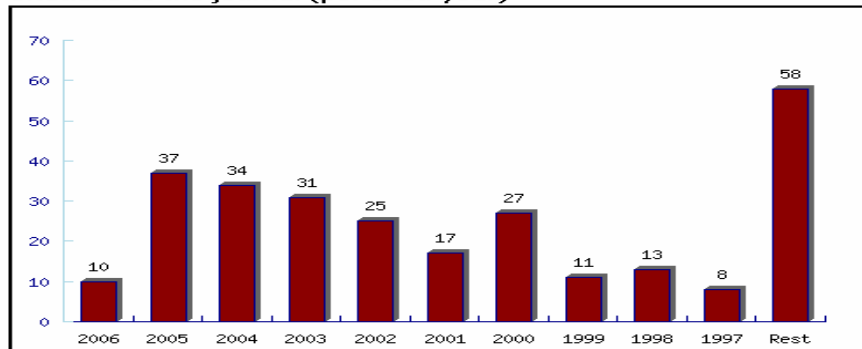
The cited half-life calculation finds the number of publication years from the KoMCI 2006 that account for 50% of citations received by *J Korean Med Sci*.

Cited Journal Graph

CITED JOURNAL DATA

This graph shows the distribution by cited year of citations to articles published in *J Korean Med Sci*.

Citations to the journal (per cited year)



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KoMCI	학술지 종수	논문수	총 참고문헌수 [A]	한국학술지 문헌 인용수 [B]	비율 [B/Ax100]
2000	69종	7,856	170,254	14,511	8.52 %
2001	75종	7,236	156,737	13,739	8.77%
2002	81종	7,301	159,581	16,719	10.48%
2003	92종	7,921	173,245	19,930	11.50%
2004	105종	8,092	180,925	22,691	12.54%
2005	113종	8,728	195,537	25,119	12.85%
2006	119종	8,932	197,880	26,492	13.39%

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- [Immune Network](#) (Immune Netw, 1598-2629)
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Intima-media Thickness and Arterial Stiffness of Carotid Artery in Korean Patients with Behcet's Disease

J Korean Med Sci. 2007 Jun;22(3):387-392

Moo Yong Rhee, Hyun Kyu Chang, Seong Kyu Kim

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Clinical Features Reflect Exon Sites of EGFR Mutations in Patients with Resected Non-Small-Cell Lung Cancer

J Korean Med Sci. 2007 Jun;22(3):393-399

Im II Na, Jin Kyung Rho, Yun Jung Choi, Cheol Hyeon Kim, Jae Soo Koh, Baek Yeol Ryoo, Sung Hyun Yang, Jae Cheol Lee

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Outpatient-basis Chemotherapy of Oxaliplatin, 5-fluorouracil, and Leucovorin as First-line Treatment for Patients with Metastatic or Recurrent Colorectal Cancer

J Korean Med Sci. 2007 Jun;22(3):400-404

Joon Ho Moon, Jong Gwang Kim, Sang Kyun Sohn, Jin Ho Baek, Yoon Young Cho, Yee Soo Chae, Byung Min Ahn, Shi Nae Kim, Soo Jung Lee, In Taek Lee, Gyu Seog Choi, Soo Han Jun

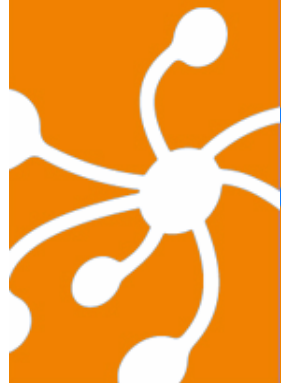
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Folate Receptor Targeted Imaging Using Poly (ethylene glycol)-folate: In Vitro and In Vivo Studies

J Korean Med Sci. 2007 Jun;22(3):405-411

Se Lim Kim, Hwan Jeong Jeong, Eun Mi Kim, Chang Moon Lee, Tae Hyoung Kwon, Myung Hee Sohn

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J Korean Med Sci. 2007 Jun;22(3):387-392.

Intima-media Thickness and Arterial Stiffness of Carotid Artery in Korean Patients with Behcet's Disease

Moo Yong Rhee, Hyun Kyu Chang, Seong Kyu Kim

Division of Cardiology, Department of Internal Medicine, College of Medicine, Dongguk University, Gyeongju, Korea., Chang Hyun Kyu Rheumatism Clinic, Cheonan, Korea., Division of Rheumatology, Department of Internal Medicine, College of Medicine, Dankook University, Cheonan, Korea. ✉ kimsk714@cu.ac.kr

Behcet's disease (BD) is a systemic vasculitis involving diverse sizes of arteries and veins. We performed this study to evaluate the vascular changes by assessment of the arterial stiffness and intima-media thickness (IMT) of carotid artery in Korean patients with BD. Forty-one patients with BD and age-, and sex-matched 53 healthy subjects were recruited in this study. Carotid arterial stiffness and IMT were assessed by using high-resolution B-mode ultrasonography. Arterial stiffness parameters such as carotid arterial distensibility coefficient, stiffness index, and incremental elastic modulus (E(inc)) were significantly increased in BD patients compared with those in healthy subjects, but not in IMT. Positive relationship was noted between age and IMT, whereas age of onset was significantly associated with arterial stiffness in BD. This finding suggests impaired endothelial function before visible structural changes of arterial wall in BD. Age and age of onset may be an independent risk factor for carotid IMT and arterial stiffness, respectively. Further studies in more large populations are required to confirm our results.

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These results suggest that interaction between LMP1 and IRF7 may be necessary for the signal transduction pathway of LMP1 and/or IRF7. Further experiments will be needed in order to provide additional details about the relationship between LMP1 and IRF7 in the signaling pathway of Epstein-Barr virus infected B-lymphocytes.

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Sugimoto Y, Tsukahara S, Sato S, et al. *Journal of Gene Medicine*
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^a Analyses of choice trials based on 22 younger and 14 older adults, as not all participants used both strategies in choice trials.
* $p < .05$.
** $p < .01$.
*** $p < .001$.

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Japanese Journal of Clinical Immunology, Volume 30, Issue 1, 2007, First Page 1
NUNOI, Hiroyuki

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Pharmaceutical Research, 2007,
Yoshioka, Sho; Katayama, Kazuhiro; Okawa, Chikako; Takahashi, Sachiko; Tsukahara, Satomi; Mitsunashi, Junko; Sugimoto, Yoshikazu

Gene therapy with drug resistance genes

Cancer Gene Therapy, Volume 13, Issue 4, 2006, First Page 335
Zabnicki M; Srinivasakumar N; Schuening F

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WormBase, <http://www.wormbase.org/> (for *C. elegans*).

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Am J Hum Genet 2007 January, 80(1): 1–11.
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Identification of a Novel BBS Gene (BBS12) Highlights the Major Role of a Vertebrate-Specific Branch of Chaperonin-Related Proteins in Bardet-Biedl Syndrome

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genes, on the basis of the assumption that these genes are conserved exclusively in ciliated organisms, such as *Chlamydomonas*, but are absent from nonciliated ones, such as *Arabidopsis*.^{12,16,17} However, this approach is insufficient to identify all genes mutated in this phenotype, as highlighted by the recent identification of the vertebrate-specific *BBS10* locus.¹⁸ In fact, the first BBS gene identified (*BBS6*, or *MKKK5*) and *BBS10* are absent from all ciliary data sets, including the recently constructed, manually curated *Ciliary Proteome Database*.⁴² Interestingly, *BBS6* and *BBS10* are both atypical members of the superfamily of type II chaperonins, on the basis of a distant sequence homology.^{18,28}

The extensive genetic heterogeneity of BBS has profound implications for diagnostic and genetics-counseling applications. Two major genes, *BBS1* and *BBS10*, each account for ~20% of the mutational load in families of European descent, whereas each of the other nine genes accounts for <5%, and some of these were found mutated in only a few families or even a single family (the latter in the case of *BBS11*).^{18,19,25} Taken together, the 11 known BBS genes account for ~70% of affected families,¹⁸ suggesting that additional BBS genes remain to be identified. A further complication is the finding that, in some cases, inheritance departs from classic autosomal recessive inheritance and involves three mutated alleles in two genes, defining oligogenic inheritance, and severity can be modulated by an allele of a modifier gene.^{25,26}

Here, we report the identification of *BBS12* on chromosome 4q27, starting with homozygosity mapping in two consanguineous Gypsy families with BBS. *In silico* studies pointed to a candidate gene, *PLJ35630*, which encodes a predicted chaperonin-like protein. We found mutations in both Gypsy families as well as in other BBS-affected families, suggesting that this novel locus accounts for some 5% of the overall mutational load of families with BBS.

BBS12, together with *BBS6* and *BBS10*, defines a new vertebrate-specific and divergent branch of chaperonin-like proteins with particular sequence insertions, with respect to the typical group II chaperonins. Suppression of each member of the family in zebrafish recapitulated previously reported *bbs* morphant phenotypes,²⁶ whereas combinatorial suppression of *bbs6*, *bbs10*, and *bbs12* revealed strong genetic interaction, with the embryos exhibiting markedly more-severe phenotypes when all three chaperonin-like proteins were suppressed simultaneously. This branch of atypical chaperonins is of major importance for BBS, since the three genes together account for about one-third of the mutational load.

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obtained from the [UCSC Genome Browser](http://genome.ucsc.edu/).

DNA Sequencing and Mutation Screening

PCR amplifications of exon 2 of *PLJ35630* were performed with 50 ng of genomic DNA template per amplicon (see [table 1](#) for oligonucleotide primers). The noncoding exon 1 was screened for all the US families and for one family in the French cohort (family III.10). Bidirectional sequencing of the purified PCR products was performed as described elsewhere.¹⁸

Table 1. Primer Sequences for <i>BBS12</i>	
[Table content]	

Database Searches

Systematic exploration in [UniProt](#) was performed with BLASTP on each candidate gene in the interval of interest. For each gene, the sequences detected with expect value <10⁻³ were aligned with the [PipeAlign](#) program.

For *BBS12*, a PSI-BLAST in [UniProt](#) and [RefSeq](#) was performed using a protein sequence from [UniProt](#), accession number Q6ZW61_HUMAN, as the query and was used to obtain a multiple alignment as described elsewhere.¹⁸ The multiple alignment was manually adjusted, with attention paid to secondary structures, leading to nearly 300 sequences corresponding to α , β , γ , δ , ϵ , ζ , and θ chaperonin-containing

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UCSC Genome Browser, <http://genome.ucsc.edu/>

UniProt, <http://www.ebi.ac.uk/uni/prot/> (for *BBS12* [accession numbers Q6ZW61_HUMAN, Q72482_HUMAN, and Q72342_HUMAN]).

WormBase, <http://www.wormbase.org/> (for *C. elegans*).

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“It’s been as much about
process and system as about
technology”

“A virtual library” in the google story.

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