18062 - Serum IgE level with asthma severity in children

by Athar Somro

Submission date: 15-Dec-2017 09:36PM (UTC+0500) Submission ID: 896621446 File name: 18062-71336-1-RV.docx (60.4K) Word count: 3002 Character count: 16386

ABSTRACT

Objectives: To determine the correlation of serum IgE levels with asthma severity in children (2 to 12 years old) presenting to Pediatric Out Door Department, Liaquat National Hospital Karachi.

2 Study Design: Descriptive Cross-Sectional Study

Place and duration of Study: Department of Pediatrics Medicine, Liaquat National Hospital Karachi from June 2016 to December 2016

Methodology: A total of 106 children with symptoms of airflow obstruction characterized by recurrent cough, difficulty in breathing and wheeze were included in this study. All the study subjects were asked about clinical symptoms and on the basis of that severity of asthma were classified as intermittent, mild, moderate or severe persistant asthma. Blood samples for Serum IgE levels were taken.

Results: - The average age of the children was 6.38 ± 6.0 years. In this study, intermittent asthma was observed in 24(22.64%) cases. There were 30(28.6%) children with mild asthma, 40(37.74%) children with moderate asthma and severe asthma was found in 12(11.32%) children. The mean IgE concentration was lowest in children with intermittent asthma 277.92±45.87 IU/ml, higher in children with mild persistent asthma 487.47±58.47 IU/ml) and highest in children with moderate and severe persistent asthma 688.75±52.08 and 1078.83±64.10IU/ml.

Conclusion: Our results support the hypothesis that severity of asthma is reflected by serum IgE levels in children. We found that the mean IgE concentration was highest in children with moderate persistent and severe persistent asthma hence the patients with raised serum IgE levels and their attendants should be counseled and advised to avoid/minimize allergen exposure for better asthma control.

Key Words: Asthma, IgE concentration, bronchoconstriction

Correlation of Serum IgE level with asthma severity in children (2 to 12 years old), presenting to a tertiary care hospital

Athar Soomro, Muhammad Nadeem Chohan, Samina Shamim, Kashif Abbas

Department of Pediatrics, Liaquat National Hospital Karachi

Corresponding Author: Muhammad Nadeem Chohan

Introduction:

Atopy is the most important risk factor for asthma in children, in this condition allergen exposure causes excessive production of IgE antibodies.¹ IgE causes bronchoconstriction by releasing various inflammatory mediators.²

High serum IgE level is an indicator of allergen exposure in genetically predisposed individuals.³ There is a strong relationship between total IgE and asthma prevalence.⁴ It is also used to determine the severity and persistence of asthma in later adulthood^{5, 6} Serum IgE level can

provide useful information regarding the severity of asthma and persistence of bronchial hyper responsiveness.

Pulmonary function test is the most accurate method of determining air flow obstruction⁷, but it is difficult to perform in children. There is an association between Serum IgE levels and the degree of airflow obstruction, ¹ hence the severity of asthma can be determined by it. There is no consensus on level of serum IgE in relation to asthma severity; different studies have reported varied levels of IgE. The usually accepted range is from 1.5-100 IU/ml.⁹ A strong relationship found between total serum IgE and asthma severity.^{10, 11, 12}

Our hypothesis is that the severity of asthma in children can be reflected by serum IgE levels. The study is intended to estimate the current magnitude of asthma severity and IgE level in our local population. The rationale behind it is that there is no consensus on mean IgE level and it varies from study to study.⁹⁻¹¹ The outcome of this study can be used for risk stratification and will help in counselling of patients and attendants. The patients with raised serum IgE levels can then be advised to avoid/minimize allergen exposure for better asthma control.

Objective:

To determine the correlation of serum IgE levels with asthma severity in children (2 to 12 years old) presenting to a tertiary care hospital.

Methodology:

This Cross sectional study was done at Department of Pediatric Medicine, Liaquat National Hospital Karachi from June 2016 to December 2016. Total 106 study subjects were included in the study with Non probability consecutive sampling technique. WHO software for sample size determination was used considering $P=13.33\%^{12}$, d=6.5%, with 95% confidence level. Inclusion Criteria were age between 2 to 12 years of either Sex with symptoms of airflow obstruction characterized by recurrent cough, difficulty in breathing and wheeze.

Children having Known or suspected immunodeficiency, congenital heart disease, Neurologically impaired children (as they are at risk for recurrent aspiration), Lung diseases like Tuberculosis, bronchiectasis, cystic adenomatoid malformation of lung and cystic fibrosis were excluded from study.

Children fulfilling the inclusion criteria were included in the study after taking informed consent from the parents/care takers. All the study subjects were asked about clinical symptoms and on the basis of their responses, severity of asthma (as per operation definition) was classified as intermittent, mild, moderate or severe persistent asthma. Blood samples for Serum IgE levels was drawn by a trained pediatric nurse and sent to the laboratory of Liaquat National Hospital. Serum IgE levels were estimated by automated Chemiluminescence system (The Chiron Diagnostics ACS: 180). The results of Serum IgE were reported by the consultant microbiologist of experience more than 5 years. All data about patients' demographics, history, physical examination, severity of asthma, and Serum IgE level was recorded by principal investigator on a predesigned Performa.

Operational Definitions:

4

Asthma: Presence of any two or more of the following symptoms of airflow obstruction more than 4 times in the previous year characterized by: cough, shortness of breath/fast breathing or presence of wheeze

Severe persistent asthma: patient with continuous day time symptoms (as stated above) and frequent night symptoms (>3 times per week).⁸

Moderate persistent asthma: patient with daily day time symptoms and night symptoms >1 per week.

Mild persistent asthma: patient with day time symptoms >2 times per week and night symptoms >2 times per month.

Intermittent asthma: patient with day time symptoms ≤ 2 times per week and night time symptoms ≤ 2 times per month.

Data was entered and analyzed in SPSS version 19. Mean \pm SD was calculated for quantitative variables i.e. Age, weight, and height, duration of disease and serum IgE level. Frequency and percentage was calculated for qualitative variables i.e. gender and severity of asthma. Stratification was done for gender, age, weight, height and duration of asthma to see the effect of these on severity of asthma and IgE levels, chi square test and one way ANOVA were applied respectively, at 95% CI and p value ≤ 0.05 was considered as significant.

RESULTS

There were total 106 children in this study. Most of the children were below 10 years of age as presented in Table 1. The average age of the children was 6.38± 6.0 years similarly average weight, height and duration of asthma is also presented in table 2. Out of 106 patients, 59.43% were male and 40.57% were female Table 1. Asthma severity is presented in figure 1. The mean IgE concentration was lowest in children with intermittent asthma 277.92±45.87 IU/ml, higher in children with mild persistent asthma 487.47±58.47 IU/ml) and highest in children with moderate and severe persistent asthma 688.75±52.08 and 1078.83±64.10IU/ml. 95% confidence intervals were also given in table 2. Comparison of mean serum IgE levels among asthma severity with respect to age groups, gender, height, weight and duration of asthma is shown in table 3 to 7. Significant difference of mean serum IgE levels was observed among severity of asthma.

AGE AND GENDER DISTRIBUTION OF THE PATIENTS

n=106

| Age | Number (Percentage) |
|------------|---------------------|
| 2-5 Years | 47 (44.34%) |
| 6-10 Years | 44 (41.51%) |
| >10 Years | 15 (14.15%) |
| Gender | |
| Male | 63 (59.43%) |
| Female | 43 (40.57%) |

TABLE 2

MEAN SERUM IGE LEVELS IN ASTHMA SEVERITY IN ASTHMATIC CHILDREN

| IgE Level (IU/ml) | | Intermittent Asthma | Mild Persistent Asthma | Moderate Persistent Asthma | Severe Persistent Asthma |
|-------------------|----------------|------------------------|------------------------------|----------------------------------|--------------------------------|
| Mean | | 277.92 | 487.47 | 688.75 | 1078.83 |
| Std. Deviation | Std. Deviation | | 58.47 | 52.08 | 64.10 |
| 95% Confidence | Lower Bound | 258.55 | 465.63 | 672.09 | 1038.10 |
| Interval for Mean | Upper Bound | 297.29 | 509.30 | 705.41 | 1119.56 |

COMPARISON OF MEAN SERUM IGE LEVELS IN ASTHMA SEVERITY WITH

RESPECT TO AGE GROUPS

| Age Grou | Age Groups (Years) and severity of asthma | | IGE LE | P-Values | |
|----------|---|----|--------|-------------------|--------|
| | | | Mean | Std. Deviation | |
| 2 to 5 | Intermittent Asthma | 11 | 280.91 | 33 | 0.0005 |

| | Mild Persistent Asthma | 22 | 481.05 | 50.15 | |
|---------|----------------------------|----|---------|--------|--------|
| | Moderate Persistent Asthma | 11 | 668.27 | 39.64 | |
| | Severe Persistent Asthma | 3 | 1094 | 102.67 | |
| | Intermittent Asthma | 13 | 275.38 | 55.77 | 0.0005 |
| 6 to 10 | Mild Persistent Asthma | 7 | 516.57 | 77.01 | |
| | Moderate Persistent Asthma | 22 | 681.59 | 51.09 | |
| | Severe Persistent Asthma | 2 | 1015.5 | 14.84 | |
| | Mild Persistent Asthma | 1 | 425 | | 0.0005 |
| >10 | Moderate Persistent Asthma | 7 | 743.43 | 37.85 | |
| | Severe Persistent Asthma | 7 | 1090.43 | 48.72 | |

One Way ANOVA applied for each age stratification



COMPARISON OF MEAN SERUM IGE LEVELS IN ASTHMA SEVERITY WITH

RESPECT TO GENDER

| Gender with severity of asthma | n | IGE LEVELS | P-Value | |
|--------------------------------|---|------------|---------|--|
| | | | | |

| | | | Mean | Std. Deviation | |
|--------|-------------------------------|----|---------|-------------------|--------|
| | Intermittent Asthma | 13 | 273.85 | 48.741 | |
| | Mild Persistent Asthma | 18 | 490.17 | 55.730 | |
| Male | Moderate Persistent Asthma | 26 | 688.46 | 51.603 | 0.0005 |
| | Severe Persistent Asthma | 6 | 1082.00 | 70.529 | |
| | Intermittent Asthma | 11 | 282.73 | 44.066 | |
| | Mild Persistent Asthma | 12 | 483.42 | 64.679 | |
| Female | Moderate Persistent Asthma | 14 | 689.29 | 54.937 | 0.0005 |
| | Severe Persistent Asthma | 6 | 1075.67 | 63.579 | |

COMPARISON OF MEAN SERUM IGE LEVELS IN ASTHMA SEVERITY WITH

RESPECT TO WEIGHT

| | | | IgE Level (IU/ml) | | P-Value |
|---------------|--------------------------------|----|-------------------|-----------|---------|
| | | | | Std. | |
| Weight with s | Weight with severity of asthma | | Mean | Deviation | |
| | Intermittent Asthma | 14 | 284.29 | 35.020 | 0.0005 |
| | Mild Persistent Asthma | 21 | 490.62 | 57.630 | |
| <= 16 kg | Moderate Persistent | 15 | 661.53 | 37.960 | |
| | Asthma | | | | |
| | Severe Persistent | 3 | 1094.00 | 102.679 | |
| | Asthma | | | | |
| | Intermittent Asthma | 10 | 269.00 | 58.775 | 0.0005 |
| | Mild Persistent Asthma | 9 | 480.11 | 63.268 | |
| >16 kg | Moderate Persistent | 25 | 705.08 | 53.175 | |
| | Asthma | | | | |
| | Severe Persistent | 9 | 1073.78 | 53.849 | |
| | Asthma | | | | |

COMPARISON OF MEAN SERUM IGE LEVELS IN ASTHMA SEVERITY WITH RESPECT TO HEIGHT

| | | | IgE Level (IU/ml) | | P-Value |
|--------------------------------|-------------------------------|----|-------------------|-----------|---------|
| | | | | Std. | |
| Height with severity of asthma | | n | Mean | Deviation | |
| | Intermittent Asthma | 13 | 284.62 | 36.427 | 0.0005 |
| | Mild Persistent Asthma | 24 | 489.13 | 55.251 | |
| <= 116 cm | Moderate Persistent Asthma | 12 | 667.42 | 37.917 | |
| | Severe Persistent Asthma | 3 | 1094.00 | 102.679 | |
| >116 cm | Intermittent Asthma | 11 | 270.00 | 55.857 | 0.0005 |

| Mild Persistent Asthma | 6 | 480.83 | 75.634 | |
|------------------------|----|---------|--------|--|
| Moderate Persistent | 28 | 697.89 | 55.175 | |
| Asthma | | | | |
| Severe Persistent | 9 | 1073.78 | 53.849 | |
| Asthma | | | | |

COMPARISON OF MEAN SERUM IGE LEVELS IN ASTHMA SEVERITY WITH

RESPECT TO DURATION OF ASTHMA

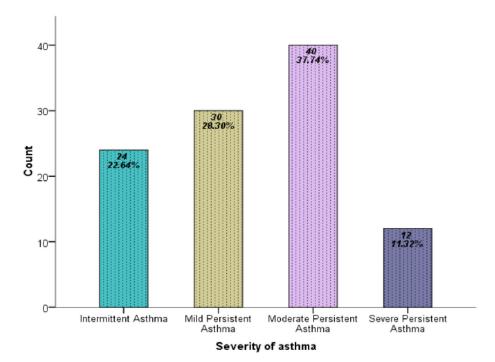
| | | IgE Level (IU/ml) | | P-Value |
|----------------------------------|---|-------------------|-----------|---------|
| | | | Std. | |
| Duration and severity of asthma | n | Mean | Deviation | |
| <= 24 months Intermittent Asthma | 8 | 282.50 | 36.93 | 0.0005 |

| | Mild Persistent Asthma | 16 | 498.19 | 57.05 | |
|------------|-------------------------------|----|---------|-------|--------|
| | Moderate Persistent Asthma | 19 | 671.89 | 46.60 | |
| | Severe Persistent Asthma | 6 | 1076 | 73.72 | |
| | Intermittent Asthma | 16 | 275.63 | 50.72 | |
| | Mild Persistent Asthma | 14 | 475.21 | 59.73 | |
| >24 months | Moderate Persistent Asthma | 21 | 704 | 53.12 | 0.0005 |
| | Severe Persistent Asthma | 6 | 1081.67 | 59.88 | |

FIGURE 1

FREQUENCY OF SEVERITY OF ASTHMA IN CHILDREN

n=106





DISCUSSION:

Acute exacerbation of asthma is precipitated by environmental factors, like inhalant allergens.¹³ IgE is responsible for inflammation of airways and various allergic reactions. Severity of asthma may be modulated by IgE. Ig-E levels are associated with asthma. The combination of reported respiratory symptoms and nonspecific bronchial hyper responsiveness has been shown to correctly identify patients with severe asthma ¹⁴. Serum IgE levels is also related to airway hyperresponsiveness in asymptomatic children ²⁰. Higher IgE levels may be correlated with lower lung function and more severe asthma attacks ²⁴.

To our knowledge no such kind of study did previously locally or internationally, that correlate serum IgE level with asthma severity in children. In our study the mean IgE concentration was lowest in children with intermittent asthma higher in children with mild persistent asthma and highest in children with moderate and severe persistent asthma, while in an unsimilar international study there was no correlation between serum IgE level and mild to moderate and severe asthma ²⁵, this may be due to different race and allergens in different countries. Another similar international study showed mean values of FEV1, FVC, FEV1/FVC% were significantly lower in high Ig E level group as compared to low Ig E group ²⁶.

In a unsimilar local study there was no significant association of serum total IgE levels with allergy severity ¹⁵, these results are different from our study because in this study they checked overall allergic symptoms, not the severity of asthma as related to Serum IgE level. Other local study also revealed positive correlation between serum IgE level and asthma but they did not compared asthma severity ^{16, 18}. A similar local study showed significant high level of Serum IgE level in poorly controlled asthma children ¹⁷ but asthma severity categorization was lacking in

their study. Other similar international study showed correlation between higher asthma severity and higher serum concentration of total IgE ²².

In our study there was no association of Mean Serum IgE level with age, gender, weight, height and duration of asthma. In a similar international study there was no association of age with Serum IgE level ¹⁹. In an unsimilar international study mean Serum IgE levels more in adolescent children ²¹. While another unsimilar study showed higher mean Serum IgE levels in males as compared to females ²³.

Our study had several limitations. It was a tertiary-care based study, so the most of the children were referred from primary care physicians; there may be bias regarding asthma severity. Socioeconomic characteristics may be different than the general asthma population. Furthermore 4 we did not assess the presence of asthma co-morbidities such as allergic rhinitis that may also affect the severity of asthma.

CONCLUSION:

Our results support the hypothesis that serum IgE levels reflect the severity of asthma in children. We found that the mean IgE concentration was highest in children with moderate and severe persistent asthma hence the patients with raised serum IgE levels and their attendants should be counseled and advised to avoid/minimize allergen exposure for better asthma control.

REFERENCES:

- Anupama N, Sharma MV, Nagaraja HS, Bhat MR. The serum immunoglobulin E level reflects the severity of bronchial asthma. Thai J Physiological Sci. 2005; 03:35-40.
- Bousquet J, Peter K, Jeffery S, William W, Busse A, Johnson M, et al. Asthma from bronchoconstriction to airways inflammation and remodeling. Am J Respir Crit Care Med. 2000; 161:1720-45.
- Blumenthal MN. The role of genetics in the development of asthma and atopy. Curr Opin Allergy Clin Immunol. 2005; 5:141-5.
- Sears MR, Burrows B, Flannery EM, Herbison GP, Hewitt CJ. Holdway MD. Relation between airway responsiveness and serum IgE in children with asthma and in apparently normal children. N Eng J Med. 1991; 325:1067-71.
- Burrows B, Sears MR, Flannery EM, Herbison GP, Holdway MD, Silva PA. Relation of the cource of bronchial responsiveness from age 9 to age 15 to allergy. Am J Respir Crit Care Med. 1995; 152:1302-8.
- Sherrill DL, Stein R, Halonen M, Holberg CJ, Wright A, Martinez FD. Total serum IgE and its associatiom with asthma symptoms and allergic sensitization among children. J Allergy Clin Immunol. 1999 Jul; 104(1):28-36.
- Hayes D Jr, Kraman SS. The Physiologic Basis of Spirometry. Respir Care. 2009; 54(12):1717–26.
- Jamalvi SW, Raza SJ, Naz F, Shamim S, Jamalvi SM. Management of acute asthma in children using metered dose inhaler and small volume nebulizer. J Pak Med Assoc. 2006; 56:595-9.

- Dati F, Ringelkp, reference values for serum IgE in healthy non atopic children and adult. Clin Chem. 1982; 28(7):1556.
- 10. Siroux V, Oryszczyn MP, Paty E, Kauffmann F, Pison C, Vervloet D, et al. Relationships of allergic sensitization, total immunoglobulin E and blood eosinophils to asthma severity in children of the EGEA Study. Clin Exp Allergy. 2003 Jun; 33(6):746-51.
- Sandeep T, Roopakala S, Rayappa C, Delphin W, Chandrashekara S, Rao M. Evaluation of serum immunoglobulin E levels in bronchial asthma. Lung India. 2010; 27:138-40.
- 12. Kovac K, Dodig S, Tjesić-Drinković D, Raos M. Correlation between asthma severity and serum IgE in asthmatic children sensitized to Dermatophagoidespteronyssinus. Arch Med Res. 2007 Jan; 38(1):99-105.
- Toelle BG, Peat JK, Salome CM, Mellis CM, Woolcock AJ. Toward a definition of asthma for epidemiology. Am Rev Respir Dis. 1992; 146:633-7.
- 14. Peat JK, Toelle BG, Salome CM, Woolcock AJ. Predictive nature of bronchial responsiveness and respiratory symptoms in a one year cohort study of Sydney school children. Eur Respir J. 1993; 6:662-9.
- 15. Hema Satwani, Arshalooz Rehman, Sohail Ashraf, Anwar Hassan. Is a Serum Total IgE level a good predictor of Allergies in Children? JPMA; 2009
- 16. Muhammad Afzal, Shaukat Mahmood Qureshi, Shabbir Hussain. Risk factors associated with childhood asthma among children aged 1-12 years in Rawalpindi. PAFMJ; 2011 (61) 3
- Javed Iqbal, Munir Ahmad Azhar, Arshad Gardezi. IgE Mediated Asthma- Effect of Gender on Management. Ann. Pak. Inst. Med. Sci. 2014; 10(4): 198-202

- M. Lama, M. Chatterjee, T. K. Chaudhuri. Total Serum Immunoglobulin E in Children with Asthma. Indian J Clin Biochem. 2013; 28(2): 197–200
- 19. Sherrill DL1, Stein R, Halonen M, Holberg CJ, Wright A, Martinez FD. Total serum IgE and its association with asthma symptoms and allergic sensitization among children. J Allergy Clin Immunol. 1999 Jul; 104(1):28-36.
- 20. M.R. Sears, B. Burrows, E.M. Flannery, G.P. Herbison, C.J. Hewitt, and M.D. Holdaway. Relation between Airway Responsiveness and Serum IgE in Children with Asthma and in Apparently Normal Children. N Engl J Med 1991; 325:1067-71
- 21. Haselkorn T, Szefler SJ, Simons FER, Zeiger RS, Mink DR, Chipps BE, Borish L, Wong DA, for the TENOR Study Group. Allergy, total serum immunoglobulin E, and airflow in children and adolescents in TENOR. Pediatr Allergy Immunol 2010: 21: 1157–65.
- 22. Kovac K1, Dodig S, Tjesić-Drinković D, Raos M. Correlation between asthma severity and serum IgE in asthmatic children sensitized to Dermatophagoides pteronyssinus. Arch Med Res. 2007 Jan; 38(1):99-105.
- 23. Borish L1, Chipps B, Deniz Y, Gujrathi S, Zheng B, Dolan CM; TENOR Study Group. Total serum IgE levels in a large cohort of patients with severe or difficult-to-treat asthma. Ann Allergy Asthma Immunol. 2005 Sep; 95(3):247-53.
- 24. Mariam Naqvi, BA, Shweta Choudhry, PhD, MSc, Hui-Ju Tsai, PhD. Association between IgE levels and asthma severity among African American, Mexican, and Puerto Rican patients with asthma. J Allergy Clin Immunol 2007; 120:137-43.
- 25. Cristine S. Rosario, MD, Nelson A. Rosario, MD, PhD, FAAAAI, Herberto J. Chong Neto, MD, PhD, FAAAA. Total Serum IgE Levels in Asthmatic Children. J ALLERGY CLIN IMMUNOL; (137) 2

26. Begum JA, Islam MI, Hoque SKA, Islam MT, Hossain MZ, Matin. Relationship between IgE levels and lung function tests in children with Asthma. J Dhaka Med Coll. 2010; 19(1): 3-6.



18062 - Serum IgE level with asthma severity in children

| | ALITY REPORT | n ige level with as | sthma severity | in children |
|--------|---|--|--|-------------------------------------|
| 1 | 1% RITY INDEX | 9% INTERNET SOURCES | 10% PUBLICATIONS | % STUDENT PAPERS |
| PRIMAR | Y SOURCES | | | |
| 1 | Claudino Junquei 6 month phagocy | -Martins, Carmer o Couto, and Mar ra. "Inhaled cortic s was not sufficie tosis in asthmati nslational Allergy | ia Imaculada costeroid treat ent to normali c children", Cl | Muniz- ment for ze |
| 2 | ompeter | | | 2% |
| 3 | Ayşenur Gümüşa asthma with astl | avuzyılmaz, Şeb Kaya, Pınar Kara y Topkara. "The severity and asth nma", SiSli Etfal dical Bulletin of S | adeniz, Melter effect of atop ima control in Hastanesi Tip | m oy on children Bulteni / |
| 4 | Children Medicat | charier. "Classifyi : Mismatch Betw on Use, and Lun | een Symptom g Function", A | ns, American |

Journal of Respiratory and Critical Care

| | Medicine, 08/15/2004 Publication | |
|----|---|----|
| 5 | www.jcpsp.pk Internet Source | 1% |
| 6 | www.j-pbs.org Internet Source | 1% |
| 7 | banglajol.info Internet Source | 1% |
| 8 | Dolan, Chantal, James Reimann, and Sharon Safrin. "Serum IgE Distributions in Normal and Asthmatic Subjects", Lung Biology in Health and Disease, 2002. Publication | 1% |
| 9 | www.cefir.com.br Internet Source | 1% |
| 10 | ijaai.tums.ac.ir Internet Source | 1% |
| 11 | V. Siroux. "Relationships of allergic sensitization, total immunoglobulin E and blood eosinophils to asthma severity in children of the EGEA Study", Clinical & Experimental Allergy, 6/2003 Publication | 1% |

| Exclude | quotes | On | ١ |
|---------|--------|----|---|
|---------|--------|----|---|

Exclude bibliography On

Exclude matches < 1%