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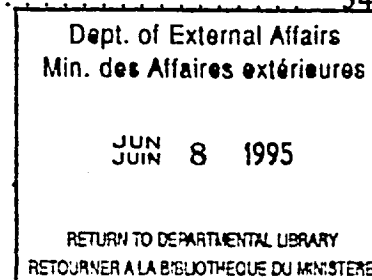
**COLLATERAL ANALYSIS
AND VERIFICATION OF
BIOLOGICAL AND TOXIN RESEARCH:
THE FINAL CASE STUDY**



MAY 1993

TABLE OF CONTENTS

PREFACE	iii
ACKNOWLEDGEMENT	v
PRINCIPAL FINDINGS AND CONCLUSIONS	vi
1.0 INTRODUCTION	1
2.0 STUDY METHODOLOGY	3
2.1 BIOSIS Previews	4
2.2 Embase	5
2.3 CA Search	5
2.4 CAB Abstracts	6
2.5 Medline	7
2.6 Selection Criteria for Canadian Publications	8
3.0 RESULTS OF THE STUDY	11
3.1 Canadian Publications	11
3.2 Sampled Canadian Publications	13
3.3 Research Centres, Agriculture Canada	15
3.4 Institut Armand-Frappier, Laval-des-Rapides, Quebec	16
3.5 The Hospital for Sick Children, Toronto	18
3.6 National Research Council, Ottawa	19
3.7 Health Protection Branch, Ottawa	20
3.8 Clinical Research Institute, Montreal	21
3.9 Faculty of Medicine, University of Calgary, Calgary	22
3.10 Faculty of Medicine, University of British Columbia	23
3.11 Biotechnology Research Institute, National Research Council	24
3.12 Department of Microbiology, University of Alberta	25
3.13 Department of Biochemistry, University of Alberta	26
3.14 Department of Medicine, McGill University	27
3.15 Department Applied Microbiology And Food Sciences, University of Saskatchewan	28
3.16 Department Veterinary Microbiology, University of Saskatchewan	29
3.17 Department of Medical Genetics, University of Toronto	30
3.18 Department of Medical Microbiology And Infectious Diseases, University of Alberta	31
3.19 Defence Research Establishment Suffield, Medicine Hat, Alberta	32
3.20 Other Canadian Publications	33
4.0 CONCLUDING REMARKS	34



43 272 581

LIST OF ANNEXES

- Annex A: Canadian Research Centres
- Annex B: Research Centres, Agriculture Canada, Ottawa
- Annex C: Institut Armand-Frappier, Laval-des-Rapides, Quebec
- Annex D: The Research Institute of the Hospital for Sick Children, Toronto
- Annex E: Division of Biological Sciences, National Research Council of Canada, Ottawa
- Annex F: Health Protection Branch, Tunney's Pasture, Ottawa
- Annex G: Clinical Research Institute of Montreal, Montreal
- Annex H: Faculty of Medicine, University of Calgary, Calgary
- Annex I: Faculty of Medicine, University of British Columbia, Vancouver
- Annex J: Biotechnology Research Institute, National Research Council Canada, Montreal
- Annex K: Department of Microbiology University of Alberta, Edmonton
- Annex L: Department of Biochemistry, University of Alberta, Edmonton
- Annex M: Department of Medicine, McGill University, Montreal
- Annex N: Department Applied Microbiology And Food Sciences., Toxicology Group,
University of Saskatchewan, Saskatoon
- Annex O: Department Veterinary Microbiology, Western College Veterinary Medicine,
University of Saskatchewan, Saskatoon
- Annex P: Department of Medical Genetics, University of Toronto, Toronto
- Annex Q: Department of Medical Microbiology And Infectious Diseases,
University of Alberta, Edmonton
- Annex R: Defence Research Establishment, Suffield, Medicine Hat, Alberta
- Annex S: Other Canadian Publications

PREFACE

This is the fourth and final report in a series which examines the potential contribution of collateral analysis to the consideration of issues related to the Biological and Toxin Weapons Convention (BTWC).

Canada is a Party to the BTWC and, furthermore, has participated in every exchange of information pursuant to the confidence-building measures (CBMs) agreed by the States Parties in 1986 and subsequently revised and extended in 1991. It is clear, however, that more can be done to enhance transparency in relation to activities involving biological materials and toxins.

Since so much of the concern about biological and toxin weapons falls in the realm of dual-use technology, any detailed discussion of such matters must inevitably comprise an assessment -- a "picture" -- of States' technological capabilities. Only on such a basis can an informed discussion take place. It was with this in mind that Canada embarked upon the current series of studies of three States Parties (Iraq, Iran and Canada) and one non-party (Israel) to the BTWC. These countries were chosen because their levels of technological capability were considered to be demonstrably different as was indicated by a preliminary examination of scientific publication databases. Although the studies do not purport to be exhaustive, they can serve as useful reference points in undertaking a comparison of what can be accomplished through the use of collateral analysis. Collateral analysis is a low-technology means of verification which is available to all States Parties to the BWTC.

It must be absolutely clear from the outset that this paper does not address the question of biological and toxin warfare. It is about technology, research and the free flow of information. This case study uses "collateral analysis" to identify from the public literature the types of

biological and toxin research that have been conducted in Canada and published. This can accomplish the following:

- identify specific areas of published research activity;
- identify institutions and scientists associated with such published research activity;
- identify the absence of published research activity in specific areas of endeavour.

Such collateral analysis can give indications of a country's technological base. It bears repeating that this report does not suggest that the work of the institutes and authors cited is in any way associated with biological warfare programs. This paper simply collates publicly-available documentation as published by the institutes and authors concerned, and it draws no conclusions from that information.

The purpose of this document is to promote discussion with regard to transparency and the utility of collateral analysis. This, and other case studies like it, are intended to contribute to the work of the Ad Hoc Group of Experts examining potential verification measures in relation to the BTWC.

This is a report of a research project. The views expressed herein do not necessarily reflect those of the Canadian Government.

ACKNOWLEDGEMENT

External Affairs and International Trade Canada wishes to acknowledge the work performed under contract by Brac Scientific Consulting, in collaboration with the Verification Research Unit of this Department.

PRINCIPAL FINDINGS AND CONCLUSIONS

- Various computerized databases, containing over 33 million records of scientific research from over 130 countries during the period 1966-1993, were accessed using key-words for this study.
- A total of 433,334 publications were identified from Canada during the period 1970 to 1992.
- 41,331 publications were selected, by the use of key-words related to biological and toxin research, from the identified Canadian research.
- 1244 unique publications (3%) were sampled from the selected Canadian publications. This sample formed the basis of the subsequent analysis. The titles of the sampled publications appear in the Annexes to this paper.
- The records of the published research were analyzed in terms of:
 - sites of research;
 - patterns of the publications over time;
 - authors; and
 - apparent nature of the research.

- The following were identified as the main Canadian laboratories and institutes publishing on biological and toxin matters:

- Research Centres, Agriculture Canada, Ottawa;
- Institut Armand-Frappier, Laval-des-Rapides, Quebec;
- The Research Institute of the Hospital for Sick Children, Toronto;
- Division of Biological Sciences, National Research Council of Canada, Ottawa;
- Health Protection Branch, Tunney's Pasture, Ottawa;
- Clinical Research Institute of Montreal, Montreal;
- Faculty of Medicine, University of Calgary, Calgary;
- Faculty of Medicine, University of British Columbia, Vancouver;
- Biotechnology Research Institute, National Research Council, Montreal;
- Department of Microbiology University of Alberta, Edmonton;
- Department of Biochemistry, University of Alberta, Edmonton;
- Department of Medicine, McGill University, Montreal;
- Department Applied Microbiology And Food Sciences., Toxicology Group, University of Saskatchewan, Saskatoon;
- Department Veterinary Microbiology, Western College Veterinary Medicine University of Saskatchewan, Saskatoon;

- Department of Medical Genetics, University of Toronto, Toronto;
 - Department of Medical Microbiology And Infectious Diseases
University of Alberta, Edmonton; and,
 - Defence Research Establishment Suffield, Medicine Hat, Alberta.
- The Research Centres of Agriculture Canada, located in Ottawa, accounted for 2.3% of the sampled publications. This was the leading Canadian centre for biological and toxin research as determined by this study.
 - The five top research centres, as determined by the number of publications, are listed below. Together, they accounted for 9.7% of the sampled biological and toxin publications. These centres are:
 - Research Centres, Agriculture Canada, Ottawa;
 - Institut Armand-Frappier, Laval-des-Rapides, Quebec;
 - The Research Institute of the Hospital for Sick Children, Toronto;
 - Division of Biological Sciences, National Research Council of Canada,
Ottawa; and,
 - Health Protection Branch, Tunney's Pasture, Ottawa.
 - Biological and toxin research in Canada, as sampled by this study, is spread out over many research centres and not concentrated in a few locations.

1.0 INTRODUCTION

In the aftermath of the Gulf War, there has been heightened concern about the potential for proliferation of biological and toxin weapons. This paper does not directly address the issue of biological and toxin weapons per se.

The Biological and Toxin Weapons Convention (BTWC) forbids the development, production and stockpiling of such weapons, as is shown from the enclosed box (see below). The Third Review Conference of the BTWC, held in September 1991, sought to address certain concerns through agreement on a refined and expanded set of confidence-building measures (CBMs) designed to provide greater transparency among States.

In September 1991, the Third Review Conference of the Parties to the BTWC also decided to establish an Ad Hoc Group of Governmental Experts open to all States Parties to identify and examine potential verification measures from a scientific and technical standpoint. One category of measures agreed by the Group for evaluation is entitled "Information Monitoring" which includes "Surveillance of publications" as a separate measure.

Article I

Each State Party to this Convention undertakes never in any circumstances to develop, produce, stockpile or otherwise acquire or retain:

(1) Microbial or other biological agents, or toxins whatever their origin or method of production, of types and in quantities that have no justification for prophylactic, protective or other peaceful purposes;

(2) Weapons, equipment or means of delivery designed to use such agents or toxins for hostile purposes or in armed conflict.

This is the recent background against which this case study was conducted of Canadian publications on the subject of biological and toxin research. The term "collateral analysis" has been used to describe a method of information gathering that involves the collection and systematic examination of publicly-available scientific, technical and other information. It makes no judgement as to the purpose of the research. The analysis may provide an initial picture of a country's technological base and of its capacity to move in a variety of directions, some of which may be of concern and, therefore, a potential area for directing CBM or verification effort to enhance transparency. Similarly, a lack of publication in certain areas of endeavour, when coupled with other sources of information, could raise certain questions. It must be added, however, that collateral analysis cannot be expected to reveal activities that are deliberately concealed.

This study had three objectives:

- 1) to identify specific areas of Canadian published research activity;
- 2) to identify institutions and scientists associated with such published research activity; and
- 3) to identify the absence of published research activity in specific areas of endeavour.

2.0 STUDY METHODOLOGY

A major part of this study depended upon the gathering of information available in the open scientific literature, specifically computerized databases. Therefore, the sources and quality of the information played an important role in determining the usefulness of collateral analysis. Table 1 lists five databases that have been found to provide particularly useful coverage of biological, biochemical and medical research. Together, these databases contain over 33 million records of scientific research from over 130 countries. Accordingly, they were the primary sources of information used in this study of Canadian research. The databases provide consistent coverage during the time period 1970-1992. The following sections will describe the databases and their breadth of coverage.

Table 1. Major Scientific Databases

DATABASE	COVERAGE	RECORDS
BIOSIS Previews	1969-present	7,813,897
Embase	1974-present	4,815,654
Medline	1966-present	6,958,530
CAB Abstracts	1972-present	2,979,945
CS Search	1969-present	10,870,743

2.1 BIOSIS Previews

BIOSIS Previews contains citations from Biological Abstracts (BA); Biological Abstracts/RRM (Reports, Reviews, Meetings [BA/RRM]); and BioResearch Index (BioI), which are the major publications of BIOSIS. Together, these publications constitute the major English-language service providing comprehensive worldwide coverage of research in the biological and biomedical sciences. Biological Abstracts includes approximately 280,000 accounts of original research yearly from nearly 9,000 primary journal and monograph titles. Biological Abstracts/RRM includes an additional 260,000 citations a year drawn from meeting abstracts, reviews, books, notes, letters, selected institutional and government reports, and research communications. U.S. patents are included from 1986 through 1989. The time period covered by BIOSIS Previews is 1969 to the present. The subject coverage in this database includes all life science subjects, including but not limited to the following: Agriculture, Anatomy, Bacteriology, Behavioral Sciences, Biochemistry, Bioengineering, Biophysics, Biotechnology, Botany, Cell Biology, Clinical Medicine, Environmental Biology, Experimental Medicine, Genetics, Immunology, Microbiology, Occupational Health, Parasitology, Pathology, Pharmacology, Physiology, Public Health, Radiation Biology, Systematic Biology, Toxicology, Toxin Research, Veterinary Science, Virology, Zoology. The material scanned for BIOSIS Previews includes the following: approximately 9,000 primary journal and monograph titles, books, reviews, technical reports, meetings and meeting abstracts, notes, letters, annual reviews, U.S. patents from 1986 through 1989, bibliographies, guides, and research communications.

2.2 Embase

Embase (Excerpta Medica) is a comprehensive index of the world's literature on human medicine and related disciplines. About 350,000 records are added annually, over 60% of which contain abstracts. All journal articles are added to the database within 30 days after receipt of the journal, and all records appear on-line with complete indexing.

Subject coverage includes Adverse Drug Reactions, Anaesthesiology, Bacteriology, Biochemistry, Bioengineering, Cancer, Cardiovascular Disease, Endocrinology, Environmental Health, Forensic Science, Genetics, Gene Technology, Health Economics, Hematology, Immunology, Industrial Medicine, Infectious Diseases, Internal Medicine, Microbiology, Neurology, Nuclear Medicine, Occupational Health, Parasitology, Pathology, Pharmacology, Physiology, Physiotherapy, Pollution Control, Psychiatry, Public Health, Radiology, Toxicology, Virology.

Embase provides access to periodical articles from more than 2,900 primary journals from over 110 countries. An additional 600 journals are screened for drug articles.

This database provides coverage of research from 1974 to the present.

2.3 CA Search

The CA Search database includes citations to the literature of chemistry and its applications. A significant part of its coverage includes biological chemistry, biotechnology, peptides, toxins and genetic engineering. CA Search is an expanded database which contains the basic bibliographic information appearing in the printed Chemical Abstracts. The subject coverage

includes Applied Chemistry, Biochemistry and Biology, Chemical Engineering, Classes of Substances, Macromolecular Chemistry, Organic and Inorganic Chemistry, Physical and Analytical Chemistry, Properties and Reactions.

The following sources are included in CA Search: journal articles, patents, reviews, technical reports, monographs, conference and symposium proceedings, dissertations, and books. Over 10,000 primary journals are abstracted. This includes coverage from over 120 countries. The database is updated two times a week with approximately 17,000 records per update. CA Search covers the time period 1967 to the present.

2.4 CAB Abstracts

CAB Abstracts is a comprehensive file of agricultural and biological information containing all records in the 26 main abstract journals published by Commonwealth Agricultural Bureaux. Over 8,500 journals in 37 different languages are scanned for inclusion, as well as books, reports, theses, conference proceedings, patents, annual reports, and guides. In some instances less accessible literature is abstracted by scientists working in other countries. About 130,000 items are indexed each year. Significant papers are abstracted, while less important works are reported with bibliographic details only. The journals included in CAB Abstracts cover the following subjects: agricultural engineering, animal disease, biotechnology, horticulture, nutrition, veterinary science, entomology, plant pathology, pesticides, fertilizers, weeds, and world agricultural economics.

The material scanned for CAB Abstracts includes the following: scientific journals, books, monographs in series, textbooks, technical reports, published theses, symposia, conference proceedings, review journals, patents, annual reports, bibliographies and guides, and translated journals. The dates covered are 1972 to the present. The database is updated monthly.

2.5 Medline

Medline (Medlars on-line), produced by the U.S. National Library of Medicine, is one of the major sources for biomedical literature materials. Additional materials not published in Index Medicus are included in the Medline database in the areas of population and reproductive biology. Abstracts, which are taken directly from the published articles, are included for over 47% of the records added from 1975 forward. Records added before 1975 do not contain abstracts; records added from 1984 to the present have abstracts for about 59% of the records. Approximately 300,000 records are added per year, of which over 75% represent publications written in the English language.

Subject coverage includes Clinical Medicine, Experimental Medicine, Hospital Literature, Population and Reproductive Biology, Pharmacology, Psychiatry, Environment and Public Health, Veterinary Medicine, Pathology, Anatomy and Physiology, Microbiology and Parasitology, Toxicology. Medline indexes articles from approximately 3,300 journals published in over 70 countries. Dates covered include 1966 to the present.

2.6 Selection Criteria for Canadian Publications

After Canadian publications were identified in the major databases, the next step was to use key words to identify certain types of research. These key words are some of the indicators sometimes suggested to have a potential relationship to biological warfare research. It should be clearly noted that the presence of these key words described below does not imply any direct connection between the research and biological weapons programs.

Key words are tools with which to identify particular types of research.

Search strategies were used to identify Canadian research associated with the following topics. The following are key words:

- 1) Microbiology, virology, bacteriology, infectious diseases;
- 2) toxins, neurotoxins, (see Box 1 for specific key words);
- 3) recombinant DNA, gene-cloning, biotechnology;
- 4) large-scale production, fermentation, bioreactors;
- 5) vaccine technology, immunology, immunization;
- 6) aerosol, lyophilization;
- 7) specific biological agents, such as bacteria, viruses, rickettsia (see Box 2 for specific key-words); and,
- 8) bioregulators (see Box 3 for specific key words).

abrin
apamin
brevetoxin
batrachotoxin
botulinum toxin
bungarotoxin
conotoxin
curare
diamphotoxin
latrotoxin
mycotoxin
palytoxin
phospholipase
ricin
sarafotoxin
saxitoxin
tetanus toxin
tetrodotoxin
tubocurare

1. Toxin Key Words

The approach was taken to use broad terms such as microorganism, virus, bacteria and toxin as well as specific key words. This was done to ensure coverage and not miss any significant area of Canadian research.

Truncation was used so that minor differences in the ending of a key word would still allow its retrieval. For example, Box 2 gives *tulare?* as a key word. In this case, the question mark allows tularemia or *tularensis* to be selected. *Francisella tularensis* is the bacterium that causes the disease tularemia.

tulare?
anthrax
anthracis
brucell?
glanders
pseudomonas
cholera
salmonella
plague
typhoid
typhi
q-fever
influenza
ebola
marburg
lassa
west-nile
congo-crimean
dengue
yellow fever
smallpox

2. Key Words for Biological Agents

angiotensin
atrial natriuretic
peptide
bombesin
bradykinin
cholecystokinin
delta sleep-
inducing peptide
dynorphin
endorphin
endothelin
enkephalin
gastrin
gonadoliberin
neurotensin
neuropeptide Y
somatostatin
substance P
thyroliberin
vasopressin

3. Bioregulator Key Words

Canadian publications were selected on the basis of the presence of key-words described above. Publications from all five databases were selected containing these key-words. It was estimated at the outset that the number of potential publications would be much greater than this study could directly examine. Therefore, it was decided to sample an appropriate number of key-word selected Canadian publications during the years 1970 to 1992. Preliminary surveys found approximately 16,700 publications from Canadian centres of research in all databases containing the key-words described above. To achieve a sample size of approximately one thousand unique publications for analysis, three percent of the publications were sampled. The box below contains an explanation of the terms used throughout this study.

The unique set of references were stored in a Pro-Cite bibliographic database. The records of the published research were analyzed in terms of:

- 1) sites of research;
- 2) patterns of publications over time;
- 3) authors of the publications; and
- 4) apparent nature of the research.

In the case of some research centres, further searches were done

with the on-line scientific databases. This was done to compare the results obtained from the sample of Canadian research with the overall picture contained in the major scientific databases.

Explanation of Terms

Identification is the process of finding all the publications from the major scientific databases for a particular country. In the case of Canada, there were 433,334 publications.

Selection is the process of using the key-words to find research on particular topics. In the case of Canada, approximately 100 key-words selected 41,331 publications.

Sampling is the process of taking a fixed percentage of the selected publications. In the case of Canada, 3% of 41,331 publications or 1244 publications were sampled, and each abstract was examined in detail.

3.0 RESULTS OF THE STUDY

3.1 Canadian Publications

One of the prime objectives of this study was to develop a baseline of information in order to analyze the trends of published Canadian biological research. Once a baseline of research in Canada had been established, questions could be asked about the possible scope of the described research. This report uses the number of scientific publications as a measure of research done. Table 2 gives the total number of publications identified as having research carried out in Canada. In the time period under consideration, there were **433,334 publications from Canada**. Clearly, this number of publications is indicative of a significant level of research, but it is important to ask when and where this research was carried out. Of course, the nature of the research is of prime importance. As well, a unique set of publications must be selected on the pertinent topics as described in section 2.6, since the multiple databases may report the same publication.

Table 2. Canadian Publications in Major Scientific Databases

DATABASE	COVERAGE	PUBLICATIONS
Biosis Previews	1970-1992	95,864
Embase	1974-1992	174,421
Medline	1970-1992	49,902
CAB Abstracts	1972-1992	80,750
CA Search	1970-1992	32,397

This section also describes the time course of the total publications from Canada. Analysis of publications from Canada in scientific databases showed marked changes in the number of publications during the period January 1, 1970 to the end of 1992. Figure 1 shows the pattern of publications in all the major scientific databases identified in this study. **It should be noted that this figure gives the total annual Canadian publications in each respective database.** Thus, there were under 2,000 publications in 1970. This increased to over 10,000 publications by 1974. All databases showed an increasing number of publications until 1982. After a small dip, the total (annual) number of publications reached a peak in 1990 at over 39,000 publications. After the peak, the number of publications declined both in 1991 and 1992. Caution should be used in interpreting these numbers since they were obtained from 5 different databases. **There is a significant amount of overlap of coverage in the databases.**

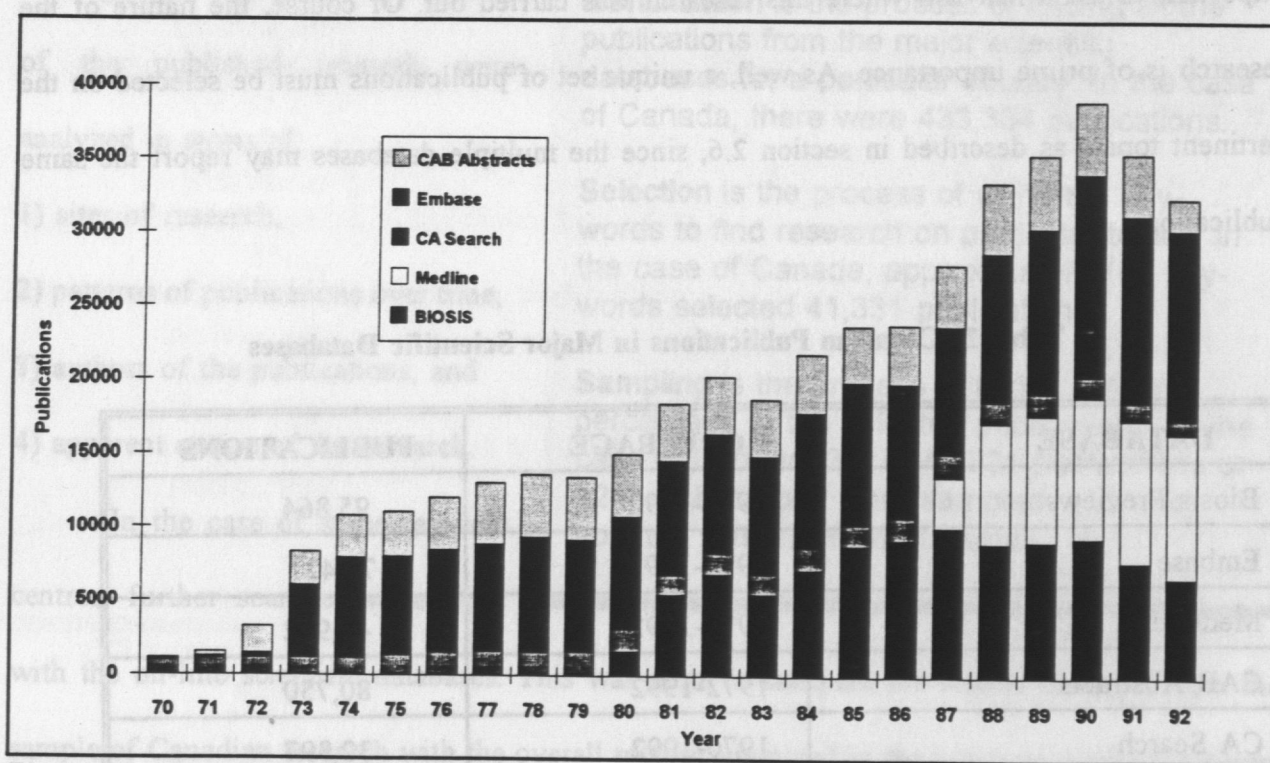


Figure 1. Total Canadian Publications in Scientific Databases

These results give an overview of all Canadian publications in major scientific databases used in this study. Clearly, there will be research included in these databases that is not at all pertinent to questions that this study is examining. Therefore, it was important to sample publications that have been selected by the key-words that establish more direct relevance to biological and toxin research programs.

3.2 Sampled Canadian Publications

There were 41,331 Canadian publications selected on the basis of key-words described in section 2.6. The objective was to select research containing key words describing research in key subject areas dealing with recombinant DNA, viruses, bacteria, toxins, peptides, bioregulators and other key areas of biological and toxin research. However, it was beyond the scope of this study to examine all 41,331 publications. Therefore, from these selections, 3% of the publications were sampled from each year during 1970 to 1992. Figure 2 shows time course of the sampled publications.

In order to provide more detailed analysis, the research contained in the 1244 sampled publications was further broken down to describe the major research centres publishing this research. Annex A lists these centres as determined by the institutional affiliation that the publications gave. The following sections describe the main laboratories and institutes that publish in the scientific literature from Canada.

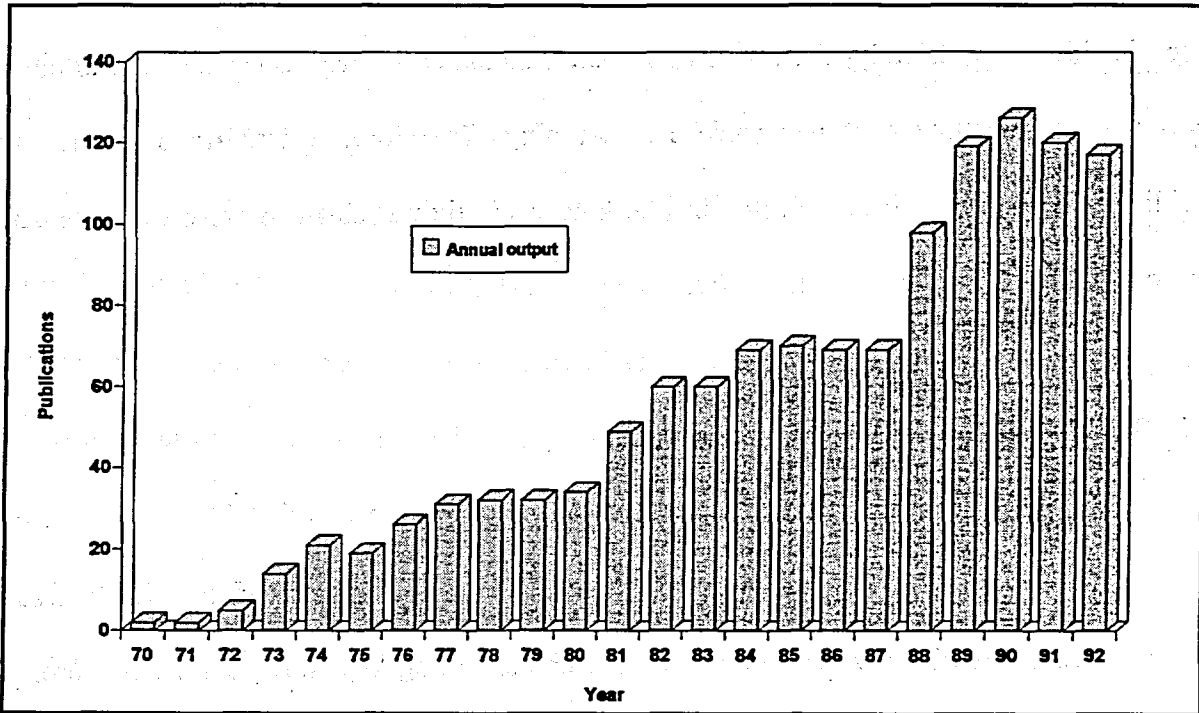


Figure 2. Sampled Canadian Biological and Toxin Publications

The following sections give a brief description of the major Canadian centres publishing biological and toxin research.

3.3 Research Centres, Agriculture Canada

The Research Centres of Agriculture Canada, located in Ottawa Canada published 29 papers sampled by this study. This represents 2.3% (29/1244) of the sample. Subsequent literature searches found 9,277 publications from the Research Centres in Ottawa Agriculture Canada out of 433,334 publications. This corresponds to 2.1% of total Canadian publications identified. Figure 3 shows the time course of sampled publications.

The main laboratories conducting research for Agriculture Canada were:

- Animal Disease Research Institute, Nepean, Ontario K2H 8P9
- Food Research Institute, Ottawa, Ontario K1A 0C6
- Plant Research Centre, Ottawa, Ontario K1A 0C6.

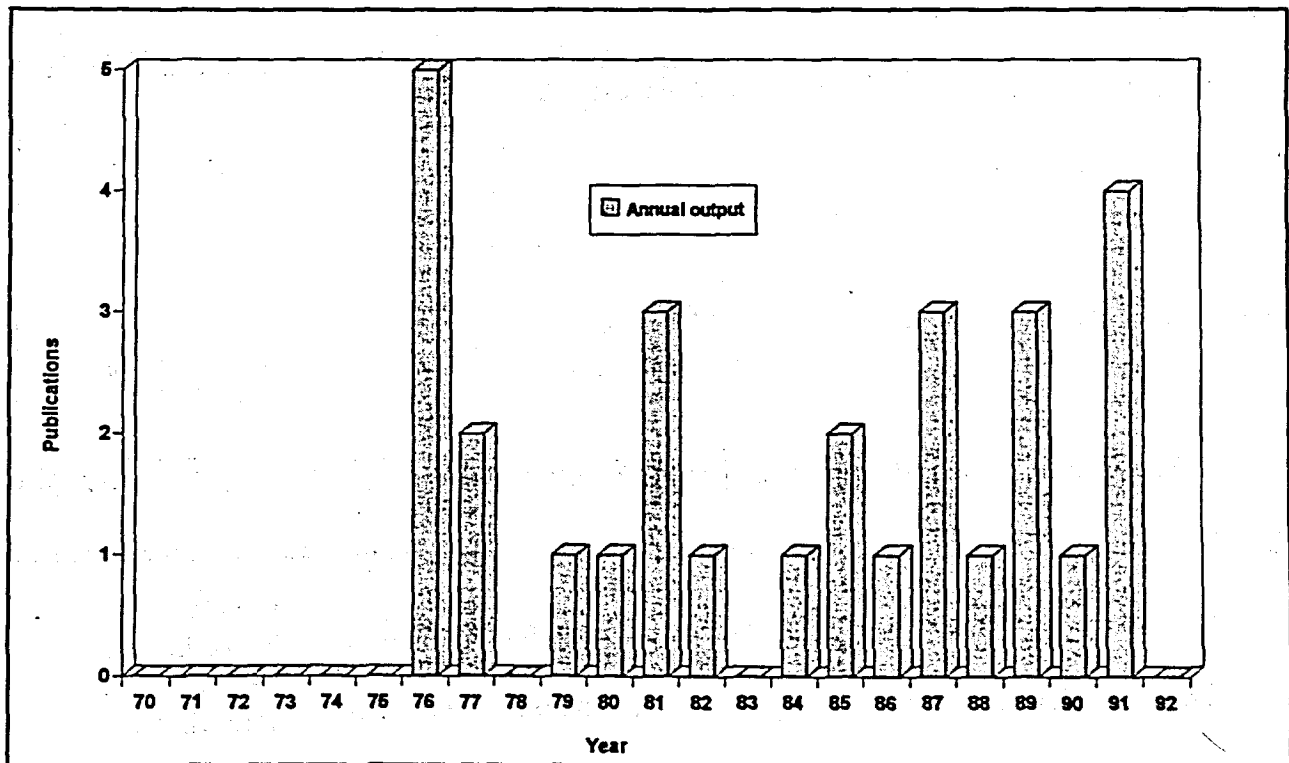


Figure 3. Sampled Publications of Agriculture Canada, Ottawa

Annex B of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 4 lists the authors who have appeared on 5 or more publications from the during the time period under consideration. Box 5 lists the apparent research priorities as suggested by the emphasis of sampled publications.

Chander, S.
Hamilton, R. M. G.
Hugh, Jones M. E.
Morley, R. S.
Prelusky, D. B.
Sauer, F. D.
Trenholm, H. L.

4. Main Authors

There is a wide range of research published by the laboratories associated with Agriculture Canada. This

ranges from applied studies dealing with agricultural problems such as disease prevention, to

Bovine leukemia virus
Disease epidemiology and prevention
Foot mouth disease virus
Fusarium-spp
Mycotoxins (Aflatoxin B-1, Deoxynivalenol, T-2 Toxin)
Rabies virus

basic research on toxin

5. Research Priorities

action and bacterial metabolism.

3.4 Institut Armand-Frappier, Laval-des-Rapides, Quebec

The Institut Armand-Frappier, located at 531 Blvd. Des Prairies, Laval-des-Rapides, Quebec published 27 papers sampled by this study. The topics covered in the sampled papers covered a wide range topics, including toxin detection and vaccine production. Box 6 lists the main authors.

Ishaque, M.
Kato, L.
Payment, P.
Trudel, M.

6. Main Authors

Box 7 lists the apparent research priorities as suggested by the emphasis of sampled publications. Annex C of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited

- Powassan virus
- St. Louis encephalitis virus
- toxin detection
- vaccine production
- Western Equine encephalitis virus

7. Research Priorities

as co-authors. Figure 4 shows the time course of the sampled publications.

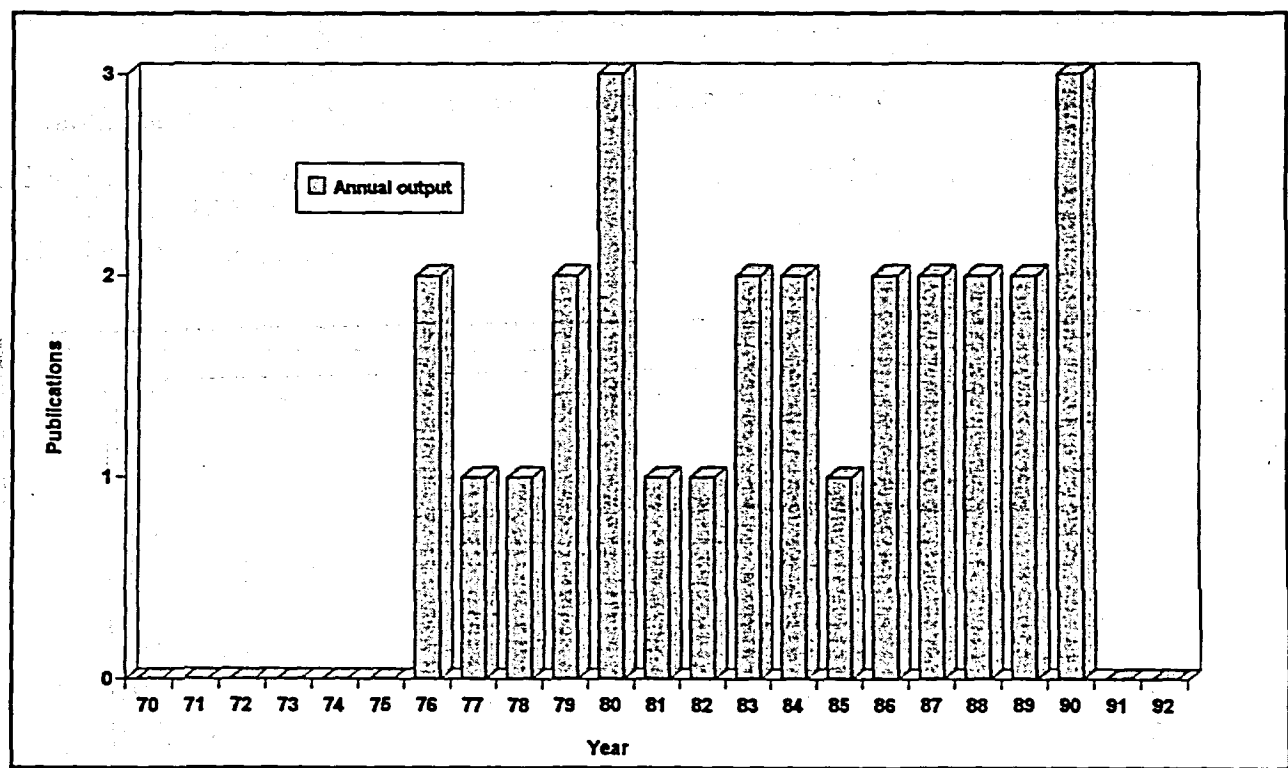


Figure 4. Sampled Publications of the Institut Armand-Frappier

3.5 The Hospital for Sick Children, Toronto

The Research Institute of the Hospital for Sick Children, located at 555 University Avenue, Toronto, Ontario published 24 papers sampled by this study. Annex D of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 8 lists the main authors Box 9 lists the apparent research priorities as suggested by the emphasis of sampled publications. Figure 5 shows the time course of the sampled publications.

Cohen, A.
Dosch, H. M.
King, S. M.
Lingwood, C. A.

8. Main Authors

Interferon
Interleukin
Tumor necrosis factor

9. Research Priorities

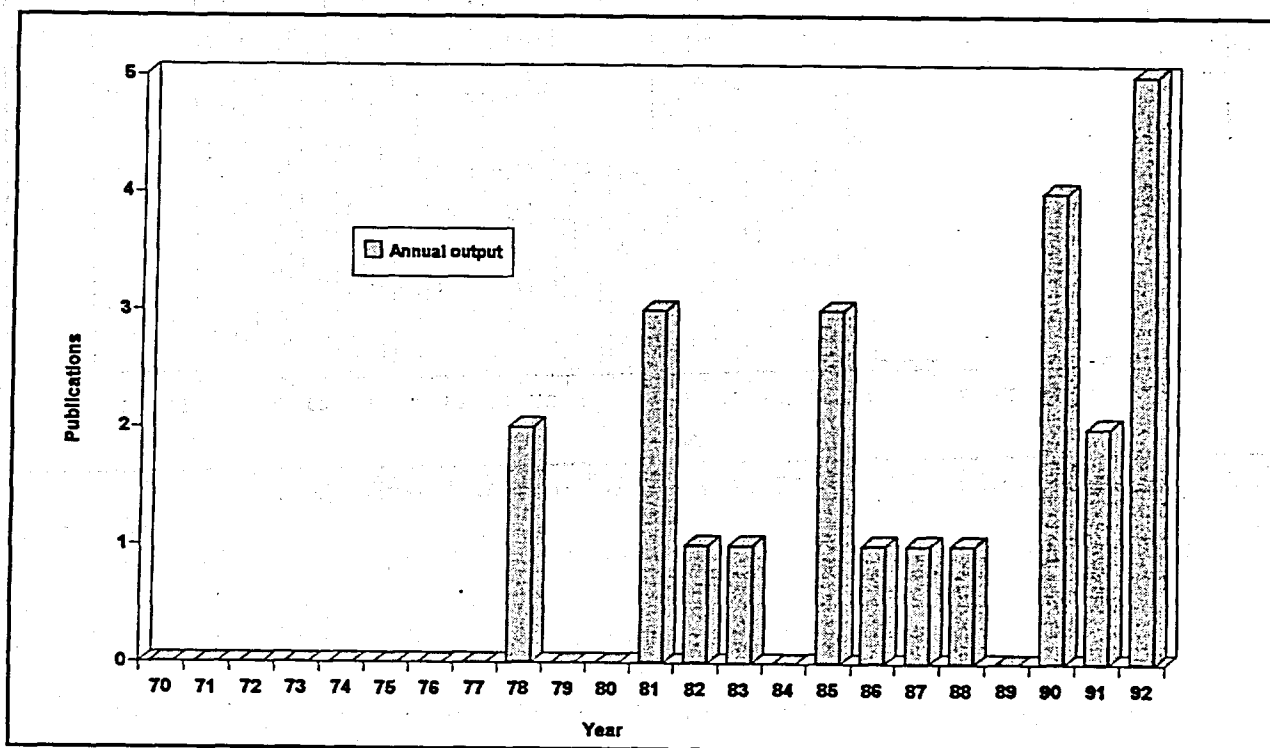


Figure 5. Sampled publications from the Hospital for Sick Children, Toronto

3.6 National Research Council, Ottawa

The Division of Biological Sciences, National Research Council of Canada, located in Ottawa, Ontario published 21 papers sampled by this study. Annex E of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 10 lists the main authors. Box 11 lists the apparent research priorities as suggested by the emphasis of sampled publications. Figure 6 shows the time course of sampled publications from the Division of Biological Sciences.

Bundle, D. R.
Durkin, J. P.
Khan, A. W.
Perry, M. B.
Whitfield, J. F.

10. Main Authors

cell proliferation
DNA Sequence Analysis
NMR Spectroscopy
Vibrio cholerae

11. Research Priorities

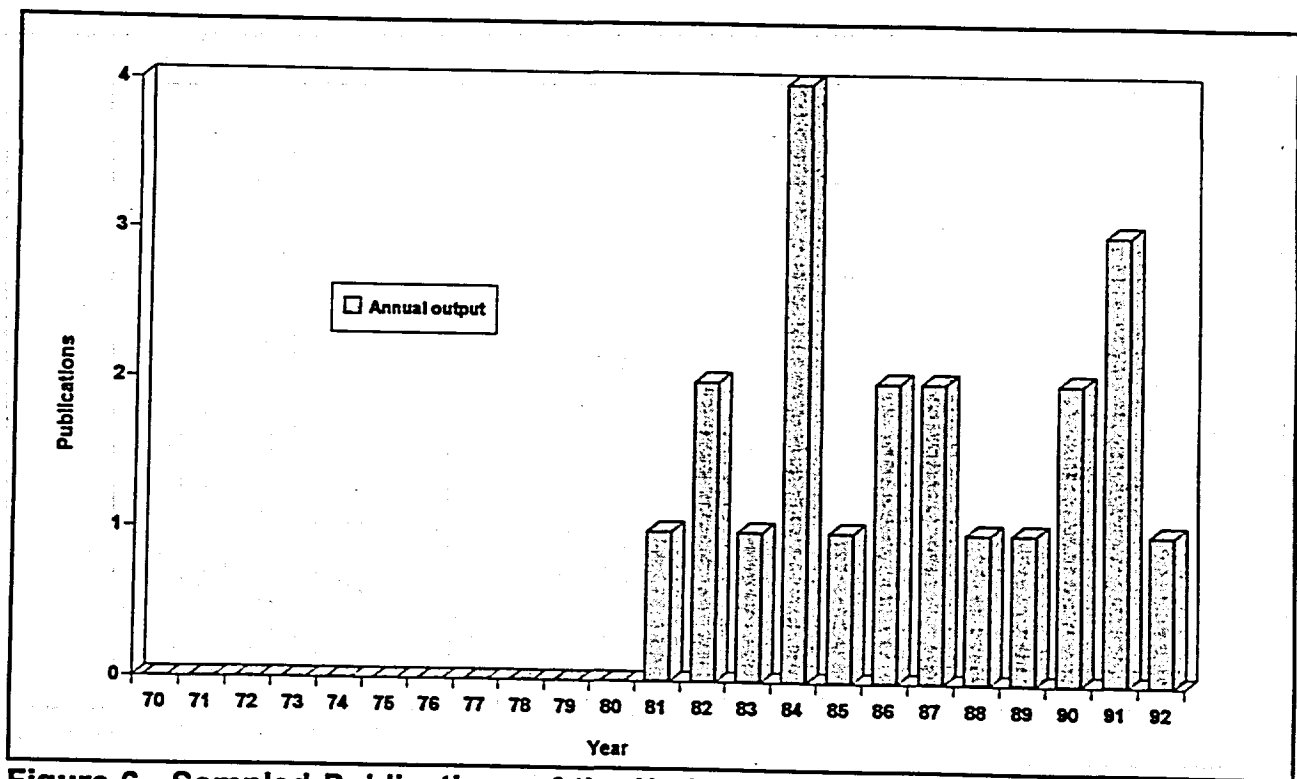


Figure 6. Sampled Publications of the National Research Council, Ottawa

3.7 Health Protection Branch, Ottawa

The Health Protection Branch, located at Tunney's Pasture, Ottawa, Ontario published 20 papers sampled by this study. Annex F of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 12 lists the main authors. Box 13 lists the apparent research priorities as suggested by the emphasis of sampled publications.

Figure 7 shows the time course of the sampled papers.

Blanchfield, B.
Daoust, J. Y.
Farber, J. M.
Sharpe, A. N.

12. Main Authors

arboviruses
mycotoxins
Pseudomonas aeruginosa
Salmonella typhimurium
toxins

13. Research Priorities

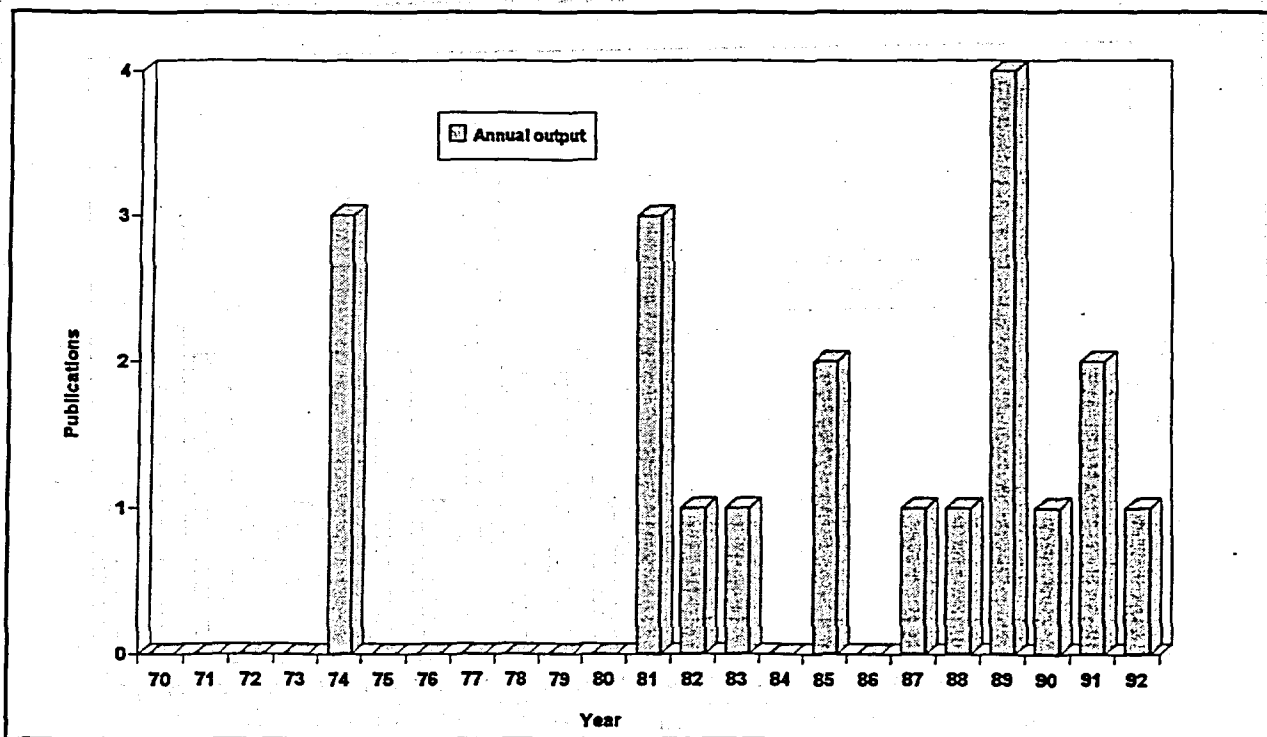


Figure 7. Sampled Publications of the Health Protection Branch, Ottawa

3.8 Clinical Research Institute, Montreal

The Clinical Research Institute of Montreal, located at 110 West Pine Ave., Montreal, Quebec, published 19 papers sampled by this study. Annex G of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 14 lists the main authors. Box 15 lists the apparent research priorities as suggested by the emphasis of sampled publications. Figure 8 shows the time course of the sampled publications.

Brousseau, R.
Greer, C. W.
Lau, P. C. K.
Luong, J. H. T.
Prefontaine, G.

14. Main Authors

Bioregulatory peptides
Endorphin
Substance P
Vasopressin

15. Research Priorities

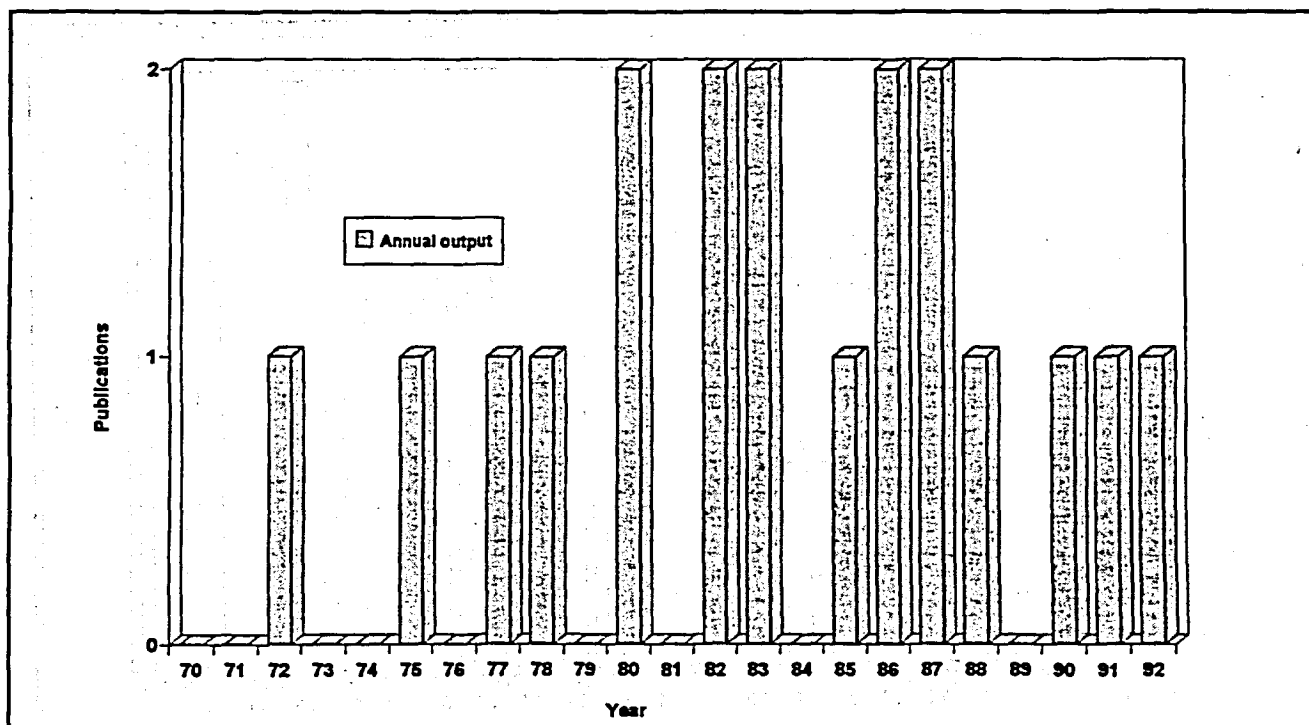


Figure 8. Sampled Publications from the Clinical Research Institute of Montreal

3.9 Faculty of Medicine, University of Calgary, Calgary

The Faculty of Medicine, University of Calgary, located in Calgary, Alberta, published 19 papers sampled by this study. Annex H of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 16 lists the main authors. Box 17 lists the apparent research priorities as suggested by the emphasis of sampled publications. Figure 9 shows the time course of sampled publications.

Giles, W. R.
Lederis, K.
Yoon, J. W.

16. Main Authors

alpha bungarotoxin
Neuropeptide Y
Tetrodotoxin

17. Research Priorities

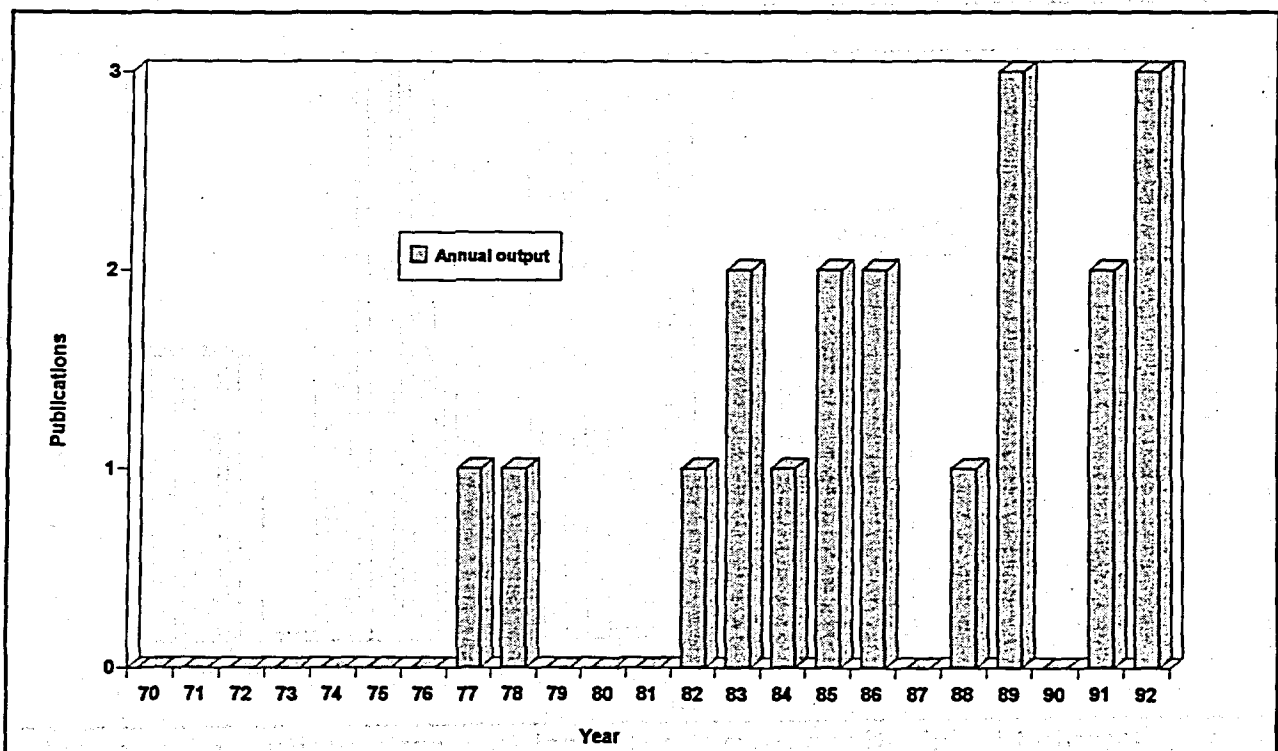


Figure 9. Sampled Publications of the Faculty of Medicine, University of Calgary

3.10 Faculty of Medicine, University of British Columbia

The Faculty of Medicine, University of British Columbia, located in Vancouver, published 16 papers sampled by this study. Annex I of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 18 lists the main authors. Box 19 lists the apparent research priorities as suggested by the emphasis of sampled publications. Figure 10 shows the time course of the sampled publications.

arboviruses
tetradotoxin

18. Research Priorities

Mclean, D.M.
Puil, E.

19. Main Authors

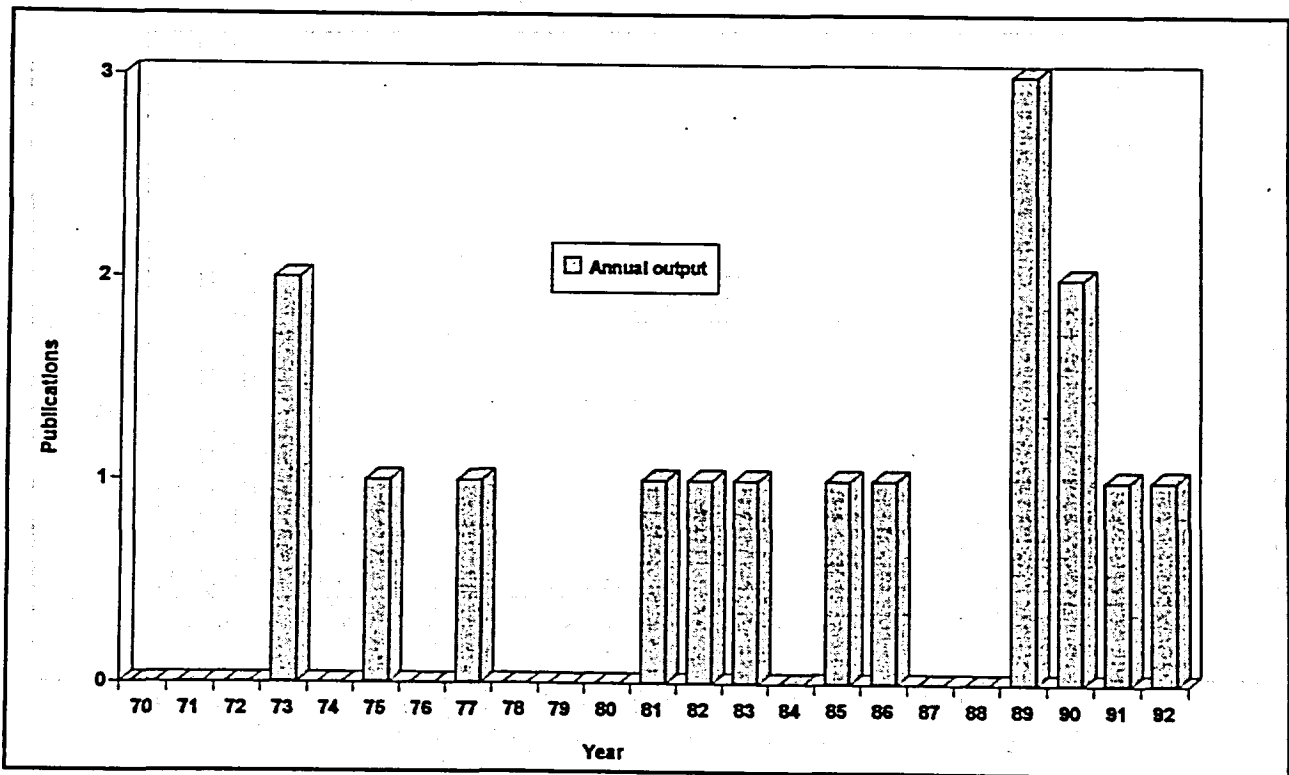


Figure 10. Sampled Publications of the Faculty of Medicine, University of British Columbia

3.11 Biotechnology Research Institute, National Research Council

The Biotechnology Research Institute, National Research Council located at 6100 Royalmount Ave., Montreal, published 13 papers sampled by this study. Annex J of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 20 lists the main authors. Box 21 lists the apparent research priorities as suggested by the emphasis of sampled publications. Figure 11 shows the time course of the sampled publications.

Brousseau, R.
Greer, C. W.
Prefontaine, G.

20. Main Authors

Biosensors
Large-scale -production

21. Research Priorities

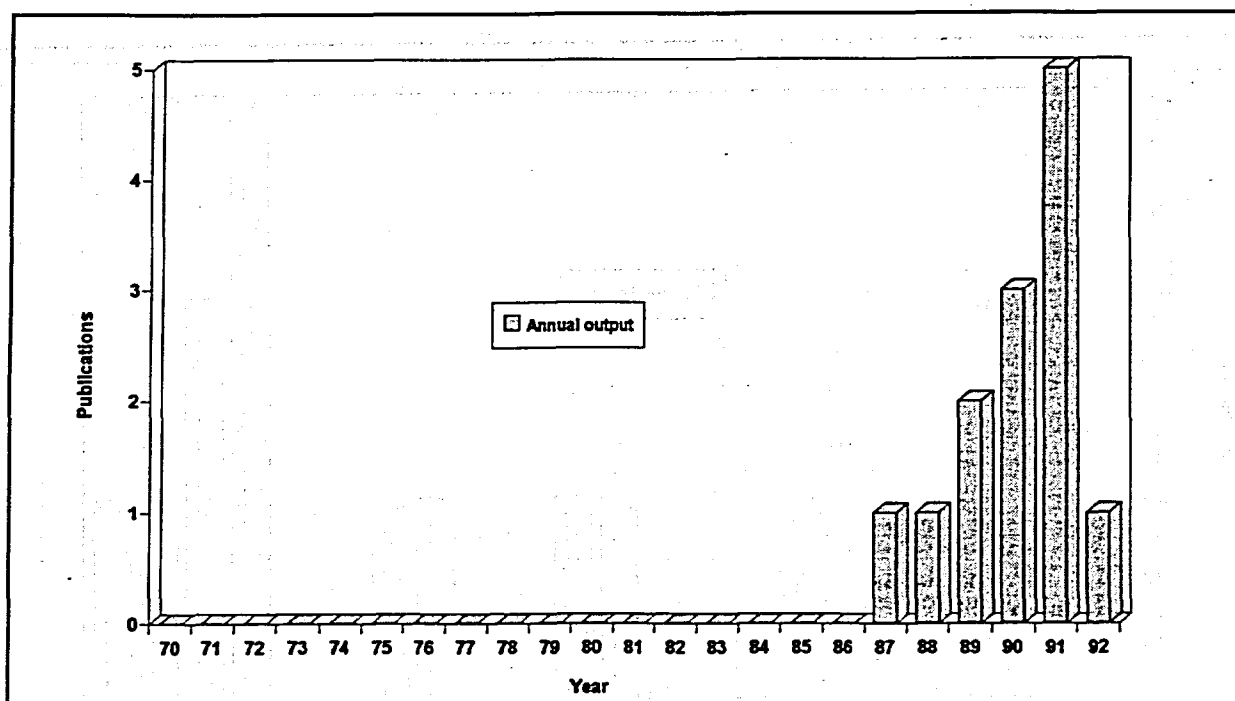


Figure 11. Sampled Publications from Biotechnology Research Institute, Montreal

3.12 Department of Microbiology, University of Alberta

The Department of Microbiology, University of Alberta, located in Edmonton, Alberta, published 13 papers sampled by this study. Annex K of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 22 lists the main authors. Box 23 lists the apparent research priorities as suggested by the emphasis of sampled publications. Figure 12 shows the time course of the sampled publications.

Jensen, S. E.
Page, W. J.
Westlake, D. W.

22. Main Authors

Antibiotics
Fermentation

23. Research Priorities

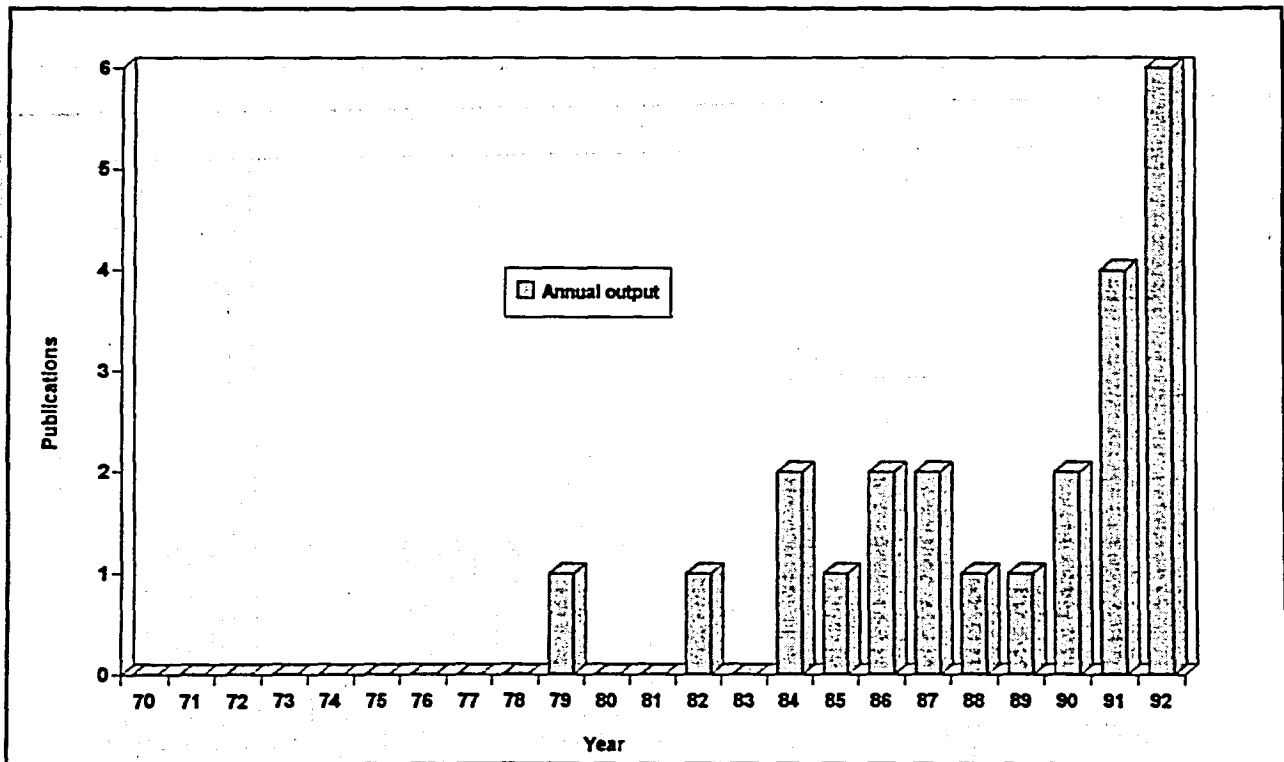


Figure 12. Sampled Publications from the Department of Microbiology, University of Alberta

3.13 Department of Biochemistry, University of Alberta

The Department of Biochemistry, University of Alberta, located in Edmonton, Alberta published 12 papers sampled by this study. Annex L of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 24 lists the main authors. Box 25 lists the apparent research priorities as suggested by the emphasis of sampled publications. Figure 13 shows the time course of selected publications.

Finlay, B. B.
Frost, L. S.
Paranchych, W.

24. Main Authors

Foot-and-mouth disease virus
Protein crystallography

25. Research Priorities

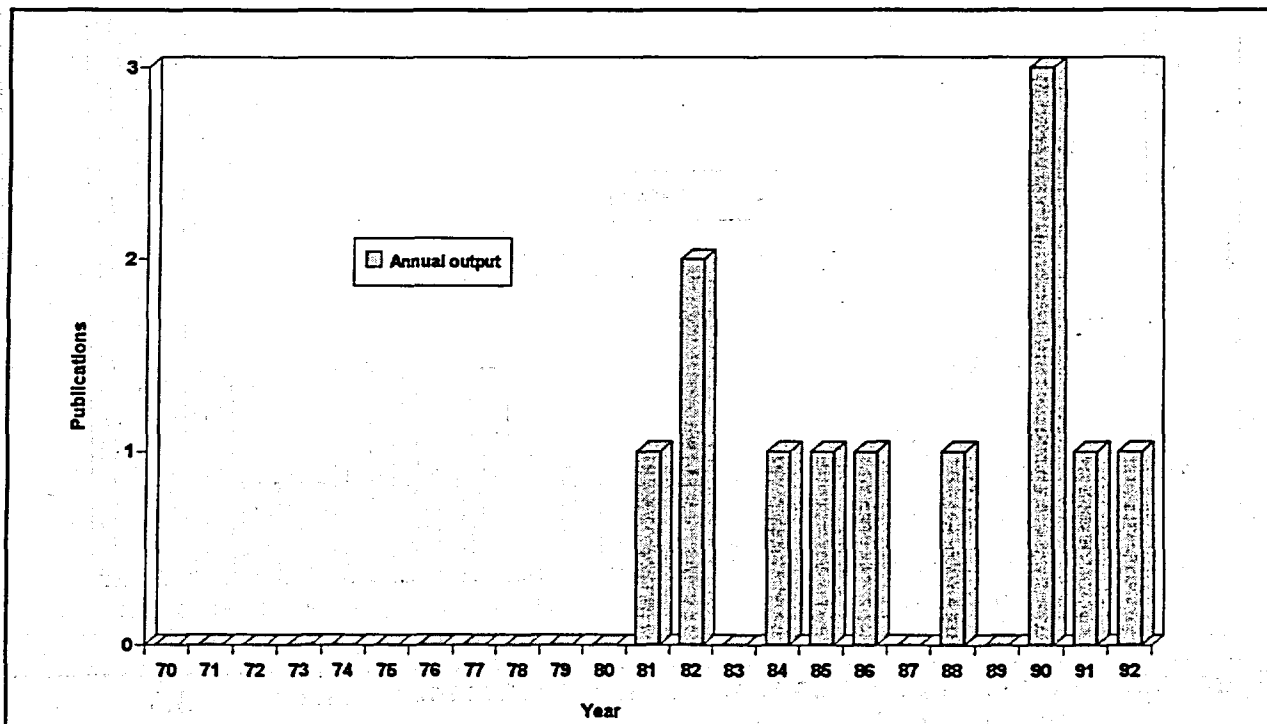


Figure 13. Sampled Publications from the Department of Biochemistry, University of Alberta

3.14 Department of Medicine, McGill University

The Department of Medicine, McGill University located in Montreal, Quebec published 11 papers sampled by this study. Annex M of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 26 lists the main authors. Box 27 lists the apparent research priorities as suggested by the emphasis of sampled publications.

Chang, T. M. S.
Patel, Y. C.

26. Main Authors

Bioregulatory peptides
Hepatitis A virus

27. Research Priorities

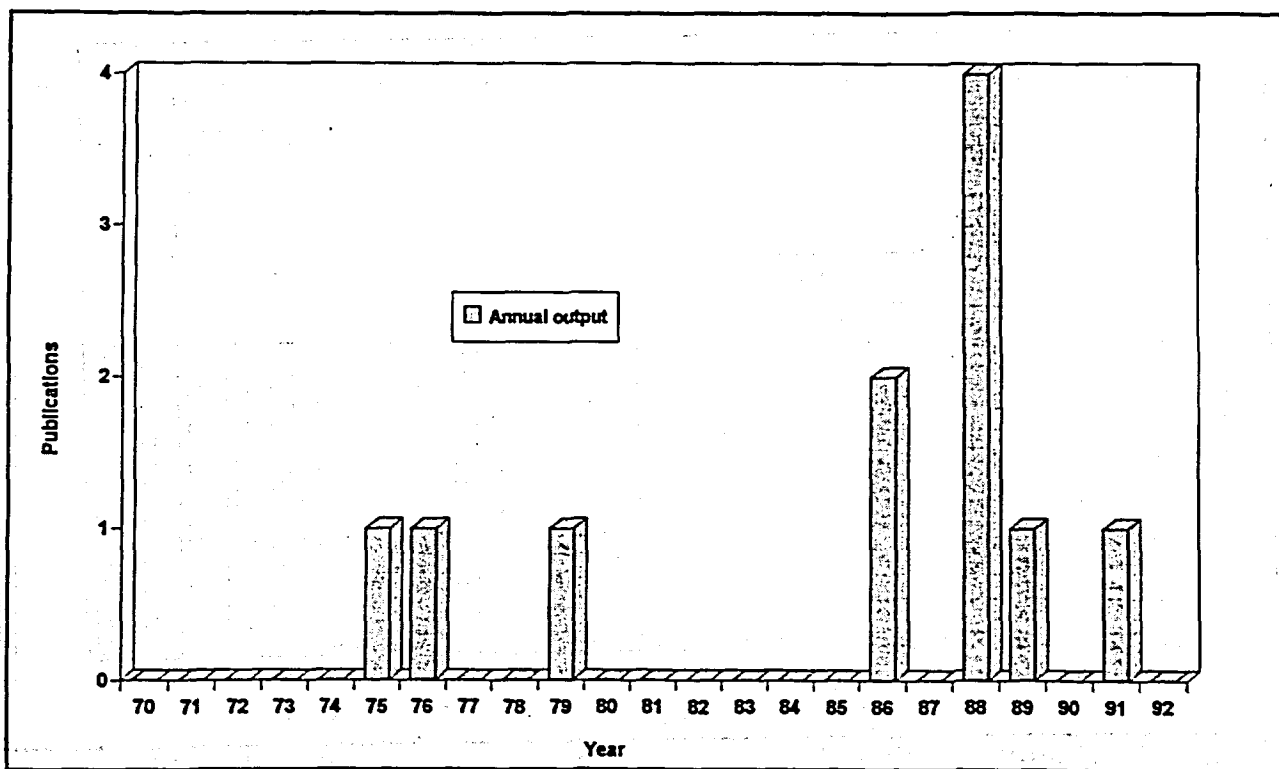


Figure 14. Sampled Publications from the Department of Medicine, McGill University

3.15 Department Applied Microbiology And Food Sciences,
University of Saskatchewan

Hayes, S.
Khachatourians, G.

The Department Applied Microbiology And Food Sciences,
Toxicology Group, University of Saskatchewan, Saskatchewan

28. Main Authors

published 9 papers sampled by this study. Annex N of this
report contains the bibliographic citations to the publications

Hemophilus influenzae
Mycotoxins

as well an alphabetical listing of all personnel cited as co-

29. Research Priorities

authors. Box 28 lists the main authors. Box 29 lists the apparent research priorities as by the
emphasis of sampled publications. Figure 15 shows the time course of sampled publications.

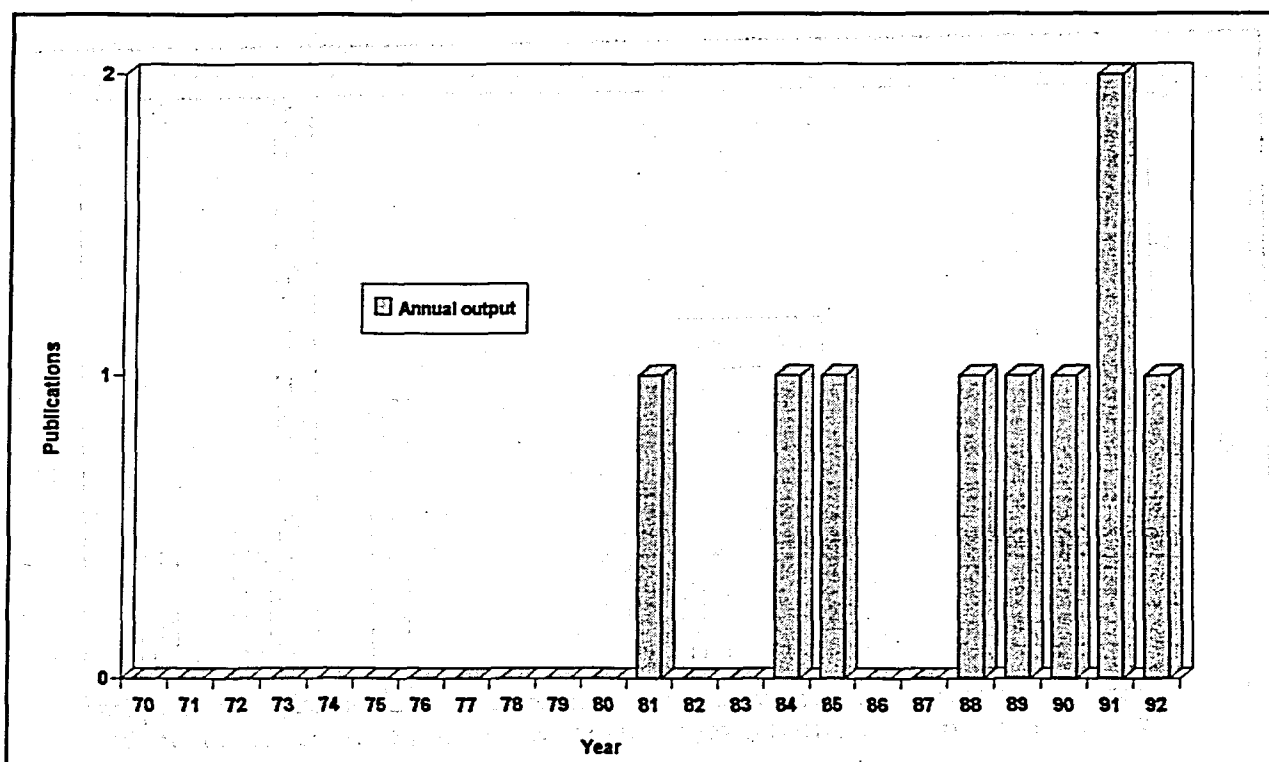


Figure 15. Sampled Publications of the Department of Applied Microbiology and Food Sciences, University of Saskatchewan

3.16 Department Veterinary Microbiology,

University of Saskatchewan

The Department Veterinary Microbiology, Western College

Veterinary Medicine, University of Saskatchewan, located in Saskatoon, Saskatchewan published 8 papers sampled by this study. Annex O of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 30 lists the main authors.

Box 31 lists the apparent research priorities as suggested by the emphasis of sampled publications. Figure 16 shows the time course of the sampled publications.

Babiuk, L. A.
Misra, V.
Sebunya, T. N.

30. Main Authors

Antiviral drugs
Pasteurella multocida

31. Research Priorities

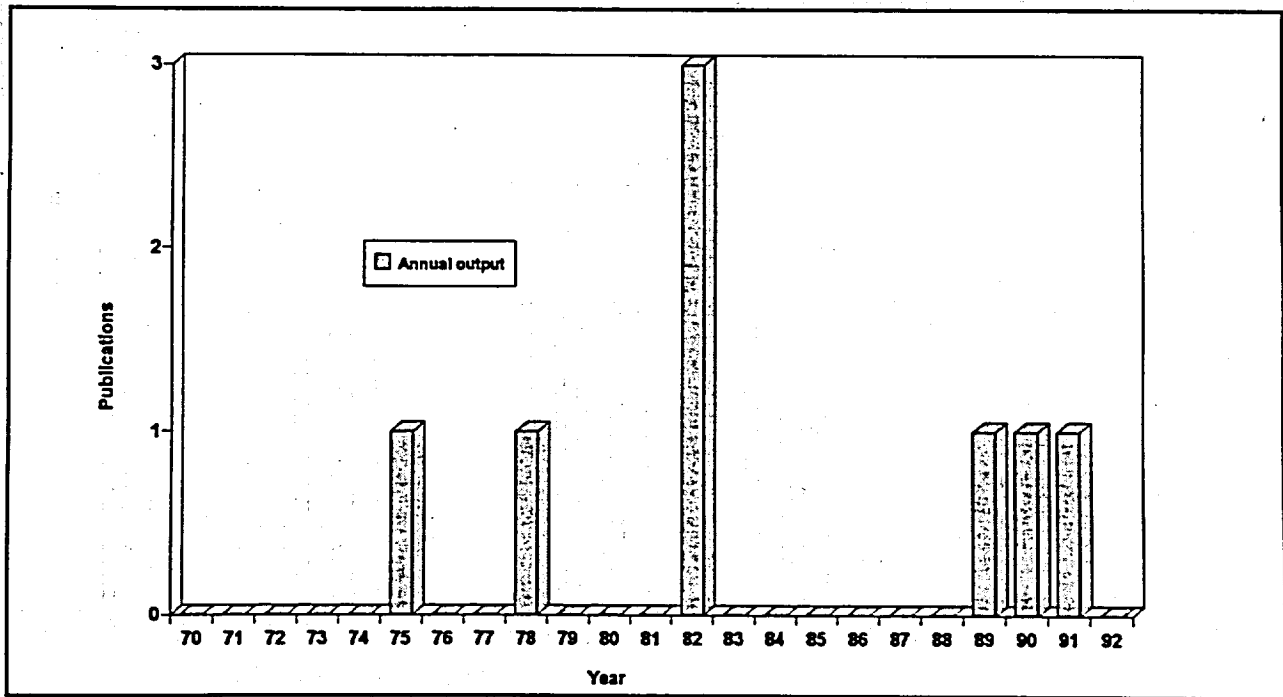


Figure 16. Sampled Publications of the Department of Veterinary Microbiology, University of Saskatchewan

3.17 Department of Medical Genetics, University of Toronto

The Department of Medical Genetics, University of Toronto, located in Toronto published 8 papers sampled by this study. Annex P of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 32 lists the main authors. Box 33 lists the apparent research priorities as suggested by the emphasis of sampled publications. Figure 17 shows the time course of selected publications.

Friesen, J. D.
Murialdo, H.

32. Main Authors

Bacteriophages
Retroviruses

33. Research Priorities

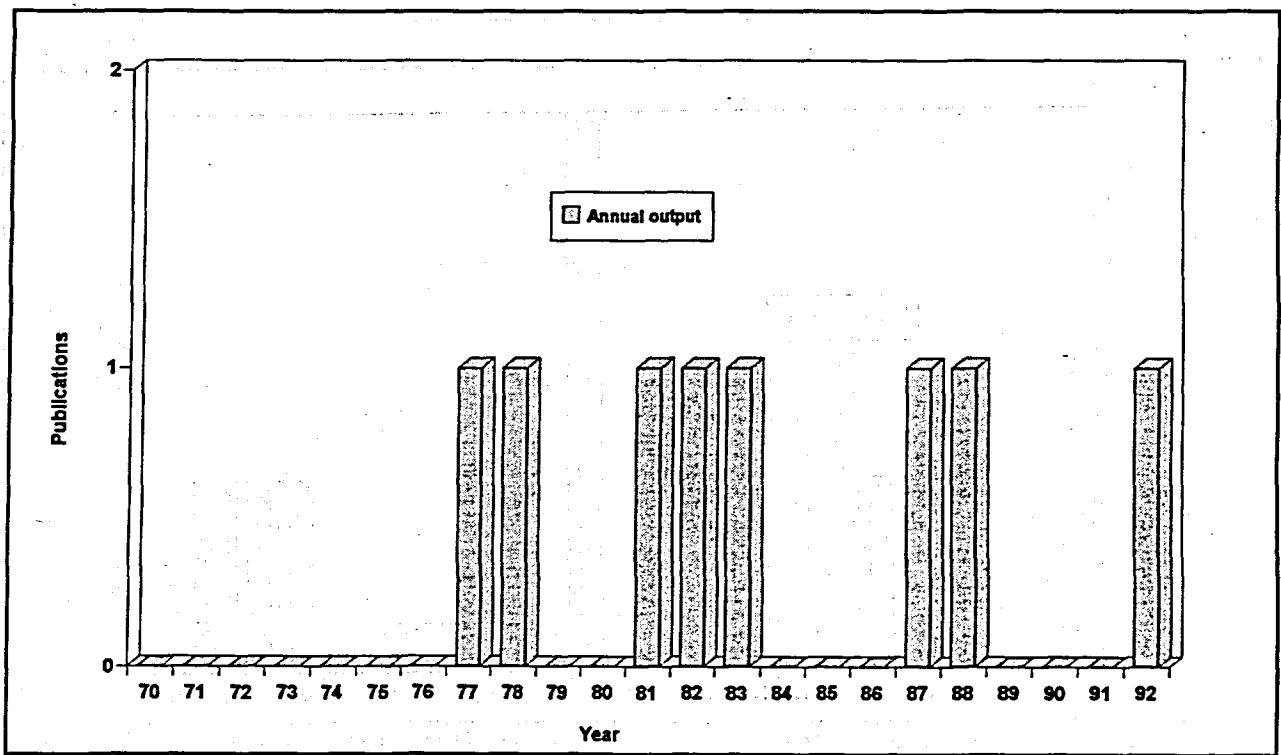


Figure 17. Sampled Publications of the Department of Medical Genetics, University of Toronto

3.18 Department of Medical Microbiology And Infectious Diseases, University of Alberta

The Department of Medical Microbiology And Infectious Diseases, located at the University of Alberta, Edmonton, published 6 papers sampled by this study. Annex Q of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 34 lists the main authors. Box 35 lists the apparent research priorities as suggested by the emphasis of sampled publications. Figure 18 shows the time course of the sampled publications.

Armstrong, G. D.
Irvin, R. T.

34. Main Authors

Bordetella pertussis toxin
Hepatitis b virus

35. Research Priorities

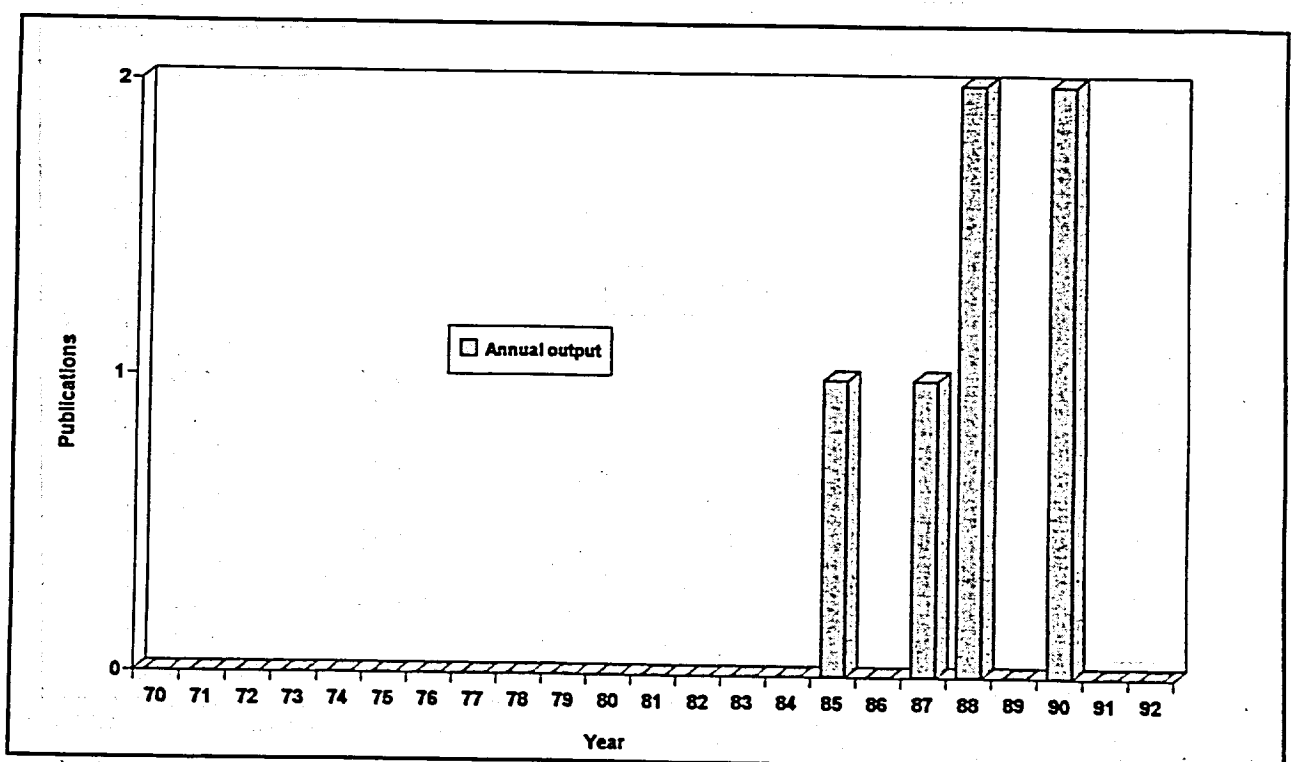


Figure 18. Sampled Publications from the Department of Medical Microbiology and Infectious Diseases, University of Alberta

3.19 Defence Research Establishment Suffield, Medicine Hat, Alberta

The Defence Research Establishment Suffield, located in Medicine Hat, Alberta published 3 papers sampled by this study. Annex R of this report contains the bibliographic citations to the publications as well an alphabetical listing of all personnel cited as co-authors. Box 36 gives the main authors. Box 37 lists the apparent research priorities as suggested by the emphasis of sampled publications. Figure 19 shows the time course of selected publications.

Frew, R.
Hamilton, M. G.

36. Main Authors

Conotoxin
Cytomegalovirus

37. Research Priorities

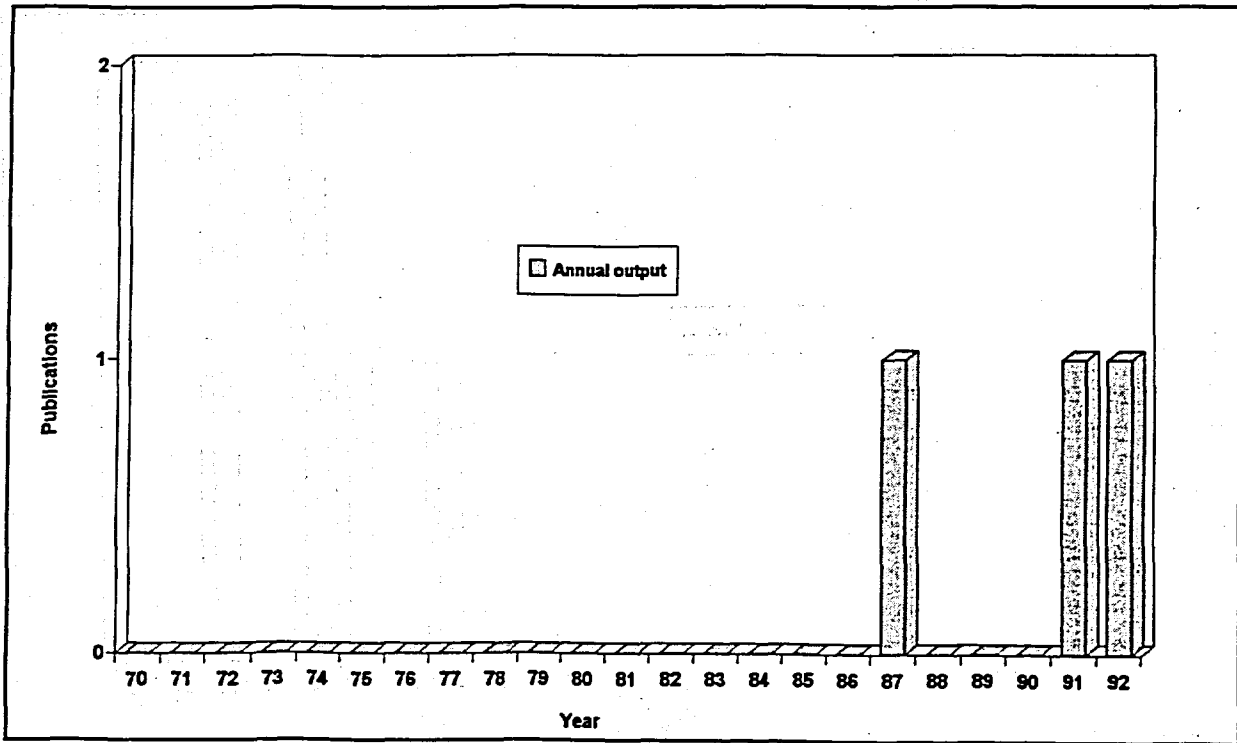


Figure 19. Sampled Publications of the Defence Research Establishment Suffield, Medicine Hat, Alberta

3.20 Other Canadian Publications

This category contains 986 sampled Canadian publications which did not originate from the major centres of research. Annex S of this report contains the bibliographic citations to these publications as well an alphabetical listing of all personnel cited as co-authors. Figure 20 shows the time course of sampled publications.

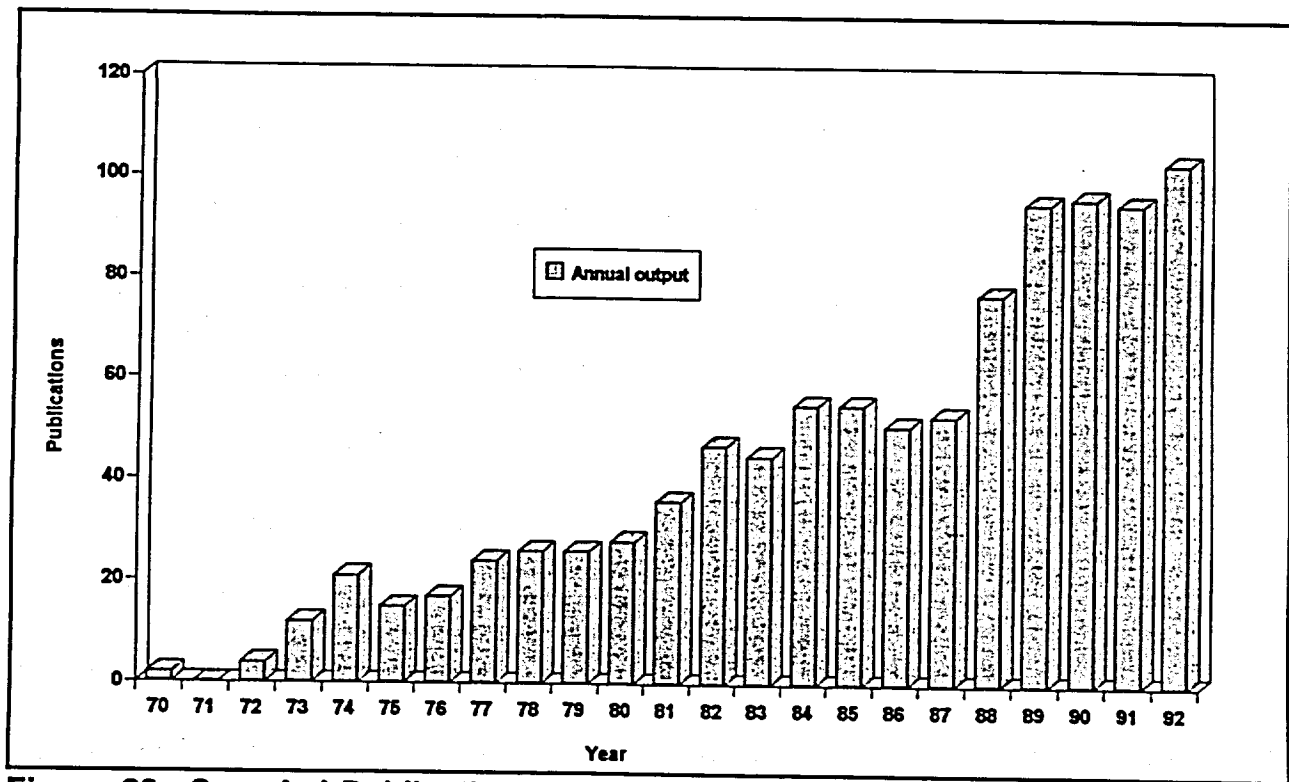


Figure 20. Sampled Publications from other Canadian research centres

4.0 CONCLUDING REMARKS

This study has surveyed Canadian scientific publications dealing with biological and toxin research. In all databases examined, there were approximately 433,000 publications, including duplicate reporting, from Canada between 1970 and 1992. Using over 100 key-words related to biological and toxin research, 41,331 publications were selected from Canada. Since it was beyond the scope of this study to examine all these publications, a unique set of 1244 publications (3% of the key-word selected publications) were sampled to survey Canadian research.

The 5 main centres for biological and toxin research are:

- Research Centres, Agriculture Canada, Ottawa;
- Institut Armand-Frappier, Laval-des-Rapides, Quebec;
- Research Institute of the Hospital for Sick Children, Toronto;
- Division of Biological Sciences, National Research Council of Canada, Ottawa;
- Health Protection Branch, Tunney's Pasture, Ottawa.

These centres carried out research using advanced biological and chemical techniques. The Research Centres, Agriculture Canada, located in Ottawa accounted for 2.3% of the sampled publications. The five centres listed above accounted for 9.7% of the sampled Canadian publications. The Canadian research sampled by this study, was spread noticeably out over many research centres and not concentrated in a few locations.

Annex A
Canadian Research Centres Publishing
Biological and Toxin Research

Research Centres, Agriculture Canada, Ottawa
Canada K1A 0C6

Institut Armand-Frappier, 531 Blvd. Des Prairies,
Laval-des-Rapides, Quebec, Canada, H7V1B7

The Hospital for Sick Children, 555 University Avenue, Toronto, Ontario M5G 1X8

Division of Biological Sciences, National Research Council of Canada,
Ottawa, Ontario K1A 0R6

Health Protection Branch, Tunney's
Pasture, Ottawa, Ontario K1A 0L2, Canada

Clinical Research Institute of Montreal, 110 West Pine Ave.,
Montreal, Quebec, Canada, H2W 1R7

Faculty of Medicine, University of Calgary, Calgary Alberta, T2N 4N1

Faculty of Medicine, University of British Columbia,
Vancouver V6T 1W5 Canada

**Biotechnology Research Institute, National Research Council
Canada, 6100 Royalmount Ave., Montreal, Can. H4P 2R2**

**Department of Microbiology University of Alberta, Edmonton, Alberta,
Canada T6G 2E9**

**Department of Biochemistry, University of Alberta, Edmonton, Alberta,
Canada T6G 2H7**

**Department of Medicine, McGill University
Montreal, Quebec, Canada H3G 1Y6**

**Department Applied Microbiology And Food Sci., Toxicology Group,
University of Saskatchewan, Saskatchewan S7N 0W0, Canada**

**Department Veterinary Microbiol., Western College Veterinary Medicine,
University of Saskatchewan, Saskatoon, Saskatchewan S7N 0W0, Canada**

Department of Medical Genetics, University of Toronto, Toronto M5S 1A8

**Department of Medical Microbiology And Infectious Diseases,
University of Alberta, Edmonton, Alta., Canada T6G 2G3**

**Defence Research Establishment, Suffield, Box 4000, Medicine Hat,
Alberta T1A 8K6, CANADA.**

ANNEX B

Research Centres, Agriculture Canada, Ottawa

This Annex contains two parts. The first part lists the sampled publications (in alphabetical order of the authors). The second part lists all the authors, in alphabetical order, of the publications listed in part one.

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12. Lewis, A. E. Significance of animal products in the spread of disease. Bull. Off. Int. Epizoot.; 87/9-10 (855-858). 1977;
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15. MCGowan, J. E. The control of newcastle disease in canada. Bull. Off. Int. Epizoot.; 85/7-8 (819-824). 1976;
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20. Morley, R. S.; Hugh, Jones M. E. Seroprevalence of anaplasmosis in the red river plains and south-east areas of louisiana usa. *Vet res commun.* 1989; 13(4): 287-296.
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Mckellar, R. C.
Miller, J. D.
Modler, H. W.
Morley, R. S.
Neish, G. A.
Pearson, J.
Prelusky, D. B.
Pringle, Ross B.
Sauer, F. D.
Sauer, F.D
Scott, P. M.
Southam, G. G.
Teather, R. M.
Teather, R.M.
Thompson, B. K.
Trenholm, H. L.
Yang, A. F.

ANNEX C

**Institut Armand-Frappier,
Laval-des-rapides, Quebec**

This Annex contains two parts. The first part lists the sampled publications (in alphabetical order of the authors). The second part lists all the authors, in alphabetical order, of the publications listed in part one.

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Collet, H.
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ANNEX D

**Research Institute
The Hospital for Sick Children,
Toronto, Ontario**

This Annex contains two parts. The first part lists the sampled publications (in alphabetical order of the authors). The second part lists all the authors, in alphabetical order, of the publications listed in part one.

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Wang, E.
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ANNEX E

**Division of Biological Sciences,
National Research Council of Canada,
Ottawa, Ontario**

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ANNEX F

**Health Protection Branch,
Tunney's Pasture, Ottawa, Ontario**

This Annex contains two parts. The first part lists the sampled publications (in alphabetical order of the authors). The second part lists all the authors, in alphabetical order, of the publications listed in part one.

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ANNEX G

**Clinical Research Institute of Montreal,
Montreal, Quebec**

This Annex contains two parts. The first part lists the sampled publications (in alphabetical order of the authors). The second part lists all the authors, in alphabetical order, of the publications listed in part one.

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ANNEX H

Faculty of Medicine, University of Calgary, Calgary Alberta

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ANNEX I

Faculty of Medicine, University of British Columbia, Vancouver, British Columbia

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ANNEX J

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ANNEX K

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ANNEX L

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ANNEX M

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ANNEX N

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This Annex contains two parts. The first part lists the sampled publications (in alphabetical order of the authors). The second part lists all the authors, in alphabetical order, of the publications listed in part one.

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ANNEX O

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ANNEX P

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ANNEX Q

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ANNEX R

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ANNEX S

Other Canadian Publications

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Denis, M.
Deniz, G.
Dent, D.
Dent, P. B.
Derbyshire, J.B
Deregt, D.
Dery, C. V.

Desgroseillers, L.
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Desrochers, M.
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Desser, S. S.
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Dimopoulos, M. A.
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Dion, P.
Djeu, J. Y.
Djoneidi, M.
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Elkashab, M.
Ellen, R. P.
Elliott, B. E.
Ellis, C.
Embil, J. A.
Enarson, D. A.
Entis, P.
Er, S. S.
Erlandson, M. A.
Ernst, P. B.
Escher, A.
Escher, E.
Eskinazi, D.
Etches, R.J
Evelegh, M.
Evelyn, T. P. T.
Evelyn, T.P.T
Everett, R.
Evers, M. C. J. M.
Ewan, P.
Ewen, M.
Fabricant, J.
Fahmy, N. W.
Fairbrother, J. M.
Fairfull, R. W.
Falbo, M.
Falk, J.
Falk, J. A.
Falk, R.
Falk, R. E.
Falkenberg, Anderson K.
Falkow, S.
Fang, R.
Fanning, A.
Farewell, V.
Farkas-Himsley, H.
Farmer, C.
Farrell, M. A.
Fast, P.
Fathi, Afshar R.
Faulkner, G.
Fedida, D.
Fedorak, P. M.
Feld, R.
Feller, T.
Fenje, P.
Fenton, T. W.
Ferin, J.
Ferket, P.
Fernandez, L. A.

Fettes, I.
Fibiger, H. C.
Fidai, S.
Figueroa, J.
Filep, J. G.
Filteau, S. M.
Finch, G. R.
Findlay, C. J.
Findlay, J. A.
Finkelstein, J. A.
Finlayson, P. G.
Finnegan, P. M.
Fisher, B. K.
Fisher, L. J.
Fitzgerald, J.D
Fitzpatrick, D. R.
Fitzsimmons, B.
Fix, D. F.
Flaschel, E.
Fleming, C. A.
Flintoff, W.
Flores, D. A.
Flumerfelt, B. A.
Fogal, W. H.
Foldes, Filep E.
Fontaine, F.
Foran, S.
Forbes, L. B.
Ford, Hutchinson A. W.
Forest, J.C
Forest, M. G.
Forget, A.
Forget, C.
Forgie, M. L.
Formosa, P. J.
Fornasier, V.
Forsberg, C. W.
Forsberg, C.W.
Forster, C.
Fortier, J.
Fortier, M.
Fortin, J. A.
Fortin, N.
Foskett, J. K.
Foster, A.
Fournier, A.
Fox, G. R.
Frackowiak, D.
Fraher, L. J.
Frappier, A.
Fraser, A. D. E.
Fraser, C. C.
Fraser, D.J
Fredericksen, T.L.

Freedman, J.
French, C. J.
Fretz, P.
Friesen, H. G.
Friesen, J. D.
Froese, A.
Frosch, M.
Fryer, J. N.
Fulop, G.
Fulthorpe, R. R.
Fung, L. S.
Furr, P.
Fyfe, C.
Gaboury, L.
Gadsden, M. H.
Gaese, C.
Gagne, C.
Gagnon, D.J.
Gagnon, P.
Gagnon, P. M.
Gagnon, R. F.
Gaisano, H. Y.
Galanopoulou, A. S.
Gall, D. E. J.
Gall, D. G.
Gall, D.G.
Gallinger, L. A.
Gallo, Payet N.
Galloway, P.
Gamache, B.
Gamble, C. L.
Gamble, C.L.
Gandhi, A.
Ganguly, P. K.
Gannon, V. P. J.
Ganz, P. R.
Garber, G. E.
Garber, Goldsman C. B.
Garcia, M. M.
Gardiner, P. F.
Gardner, N.
Gareau, A. B.
Gareau, R.
Gariepy, J.
Garland, J. A.
Garvey, M. B.
Garzon, S.
Gascoyne, R. D.
Gaudin, D.
Gaudreau, C. L.
Gaudreau, L. R.
Gauldie, J.
Gauthier, J. Y.
Gauthier, L.

Gauthier, S.
Gavora, J. S.
Gawel, M.
Ge, A. Z.
Gedamu, L.
Gentner, N.E
Gentry, P. A.
Gerhardt, D.
Gershey, R. M.
Gervais, F.
Geyer, R.
Ghadirian, E.
Ghadirian, G.
Gharapetian, H.
Gianoulakis, C.
Gibb, W.
Gibbins, A. M. V.
Gibbon, K.
Gibson, L.
Gidney, M. A. J.
Gierasch, L. M.
Gilbert, C.
Gilbert, J. J.
Gilbride, K. A.
Gilkes, M. R.
Gilkes, N. R.
Gill, C. O.
Gill, F.
Gill, S.
Gingras, G.
Gingras, M. C.
Girard, F.
Girard, M.
Giroux, J. M.
Giroux, S.
Gladu, J. M.
Glaeske, D.
Glass, K. A.
Glass, N. L.
Glickman, B. W.
Globus, M.
Glode, L. M.
Gnanmanickam, S. S.
Goda, Y.
Godbout, R.
Godden, D. J.
Godkin, S. E.
Godson, D. L.
Goerzen, D. W.
Gold, L.
Goldberg, H.
Goldie, H.
Golomb, M.
Golsteyn, Thomas E. J.

Gong, J.
Gonzalez, A.
Good, P. W.
Goodman, D. M.
Goodyer, C. G.
Gopaul, D.
Goplen, B. P.
Gorzynski, R. M.
Gordon, A. J. E.
Gordon, S.
Goring, D. R.
Gornitsky, M.
Goubau, S.
Goulet, J.
Gourdeau, M.
Graham, A. F.
Graham, A. M.
Graham, F. L.
Graham, F.L
Granger, Julien M.
Grant, E.
Gratton, A.
Green, M.
Green, M. J.
Greenberg, L.
Greenblatt, J.
Greenfield, J.
Greenway, C.V.
Greer, C. N.
Greer, G. G.
Greer, P. A.
Gregor, L.
Gregor, L. L.
Gregor, M.
Greig, P.
Gribble, M.
Grimsrud, K.
Grinstein, S.
Grondin, G.
Gros, P.
Grose, J.H
Grosovsky, A. J.
Groulx, C.
Groves, D.
Groves, D.J
Grofum, W. L.
Growe, G. H.
Gruda, J.
Gruia, Gray J.
Grupp, L. A.
Grzybowski, S.
Gu, M.
Guay, J. M.
Guertin, Carignan M.

Guidoin, R.
Guidoin, R. G.
Guidos, C.
Guilbert, L. J.
Guillemette, G.
Guillon, G.
Guilmette, H.
Gurd, W.
Gurevich, N.
Gutkowska, J.
Guttman, R. D.
Guttman, R.D
Guy, D.
Guyda, H. J.
Guzman, R.
Gyles, C.
Gyles, C.L.
Haber, S.
Hackett, N. R.
Hackmann, T.
Haddad, G.
Haines, D.
Hale, E.J.
Hall, J. W.
Hall, R.
Hall, R. A.
Halloran, P.
Halloran, P. F.
Hamalian, T.
Hamilton, J. T.
Hamilton, J.R.
Hammerberg, O.
Hammerlindl, J. K.
Hammond, G. W.
Hancock, E. J.
Hancock, R. E. W.
Hand, R.
Hanly, P.
Haour, F.
Haraphongse, M.
Hardin, J. A.
Hardy, G. M.
Hare, W. C. D.
Harel, J.
Harkness, R. E.
Harnett, J. D.
Harrington, M. E.
Harris, M.
Harrison, J. C. L.
Harrison, P. G.
Hasel, K. W.
Hasnain, S. E.
Hassell, J. A.
Hatton, M. W. C.

Havlicek, V.
Hay, G. W.
Hayashi, Yoshiyuki
Hayes, C.
Hayes, L.
Hayes, S.
Haynes, R. H.
Hayward, G.
Heath, R.B
Heathcote, J.
Hebert, J.
Heidemann, S. R.
Helbecque, D.M.
Henderson, H. M.
Henderson, M.
Hendrickson, O. Q.
Hendy, G. N.
Hengartner, H.
Hennessy, J. N.
Henni, H.
Henning, M. D.
Henry, E.
Henry, J. G.
Henry, J. L.
Henry, M.
Hicks, K. A.
Hiebert, L.M.
Higashitani, F.
Higgins, R.
Hildebrand, P. D.
Hill, G. A.
Hill, R. D.
Hipkiss, A. R.
Hirata, F.
Hiruki, C.
Hiscott, J.
Hitt, M. M.
Ho, D.
Hobbs, S. L. A.
Hodge, D. S.
Hoffman, A.G.D
Hoffman, D.
Hofman, J.D.
Hogg, G.R.
Holdeman, L. V.
Holder, M. D.
Holl, F. B.
Holland, G. R.
Holland, H. L.
Hollenberg, M.J.
Holme, R. H. G.
Holton, D.
Holy, S.
Homonylo, M. K.

Honore, L. H.
Honrath, U.
Hope, B. T.
Horgen, P.A.
Horne, C.
Horsewood, P.
Horvath, E.
Horwitz, R.
Hosseinzadeh, M.
Howard, R.
Howley, T. P.
Howse, C.W.
Hoyle, B. D.
Hoyt, D. W.
Hrdina, P.D
Hsia, J. C.
Hu, B.
Hu, D. C.
Huan, S. D.
Huang, D.
Huang, H. C. K.
Huard, G.
Hubler, S.
Hubschman, S.
Huchzermeyer, F. W.
Hudon, C.
Hudson, A. J.
Hudson, J.B.
Hughes, H. P. A.
Huizinga, J. D.
Humphries, K.
Humphries, R. K.
Hung, S.
Hunt, R. H.
Hunter, B.
Hunter, E.
Huppe, H.
Hurlbert, R. E.
Hurst, R. D.
Hurst, S.
Hussain, Z.
Hutchison, D. M.
Hylemon, P. B.
Hyodo, A.
Hyslop, D. B.
Hyslop, N.St.G
Ibrahim, R. K.
Idziak, E. S.
Ijaz, M. K.
Ingledow, W. M.
Ingraham, J.L
Inman, R. D.
Innes, I.R.
Inniss, W.E

Inoue, M.
Inouye, N.
Irvin, R. T.
Irwin, D.
Ishac, E. J. N.
Ishaque, M.
Ishida, K.
Ishiguro, E. E.
Ishiguro, E.E.
Ismail, A. A.
Ismail, A. I.
Israel, Y.
Issekutz, A. C.
Issekutz, T. B.
Itakura, C.
Ito, E.
Iwaya, M.
Iyer, V. N.
Iyer, V.N
Jabaji, Hare S. H.
Jack, T.R.
Jacobs, H.
Jacobsen, K.
Jacques, L.
Jacques, M.
Jakubovic, H. R.
Jalowiec, J.E.
James, D.
James, H.
Jamieson, W. R. E.
Jansz, G. F.
Janzen, R. A.
Jao, King C.
Jaques, L.B
Jarema, A. J.
Jarrell, K.
Jarrell, K. F.
Jasmin, G.
Jaswal, A. S.
Javorsky, P.
Jayatileka, T. N.
Jeffrey, V. M.
Jellinck, P. H.
Jenkins, D.
Jenkins, D. J. A.
Jenkins, H. J.
Jensen, S. E.
Jericho, K. W. F.
Jessop, J.
Jiang, M. Y.
Jobin, M.
Johns, A.
Johnson, D.
Johnson, D. C.

Johnson, P. M.
Johnson, R.
Johnson, S.E.
Johnson, W. H.
Johnson, W. M.
Jol, D.
Jolicoeur, F.
Jolicoeur, F. B.
Joncas, J. H.
Joncas, J.H.
Jones, G. A.
Jones, J. B.
Jones, M. W.
Jones, S. R. M.
Jones, T.
Jones, T. R.
Jones, W.
Jongstra, J.
Joseph, M. G.
Joshi, S.
Joyner, A.
Juhasz, J.
Juorio, A. V.
Jurkowska, G.
Kabata, Z.
Kabore, A. F.
Kadam, S. K.
Kakulphimp, J.
Kalnins, R.
Kalnins, V.I
Kamal, V. S.
Kamekura, M.
Kaneda, T.
Kang, C. Y.
Kannan, L.
Kannan, M. S.
Kapasi, K.
Kaplan, H.
Kapoor, S.
Kapusta, M.A.
Karasira, P.
Karpowicz, E.
Karska, Wysocki B.
Kantha, K. K.
Kates, M.
Kato, L.
Kaufert, J. M.
Kaufert, P. L.
Kaufmann, J. C. E.
Kaupp, W. J.
Kavaliers, M.
Kawarabata, T.
Kawchuk, L. M.
Kay, W. W.

Keenan, C. M.
Kelly, R.
Kellen, J.A
Kelley, J. L.
Kelly, J. J.
Kelly, L. J.
Kelly, N.
Kelly, W. A.
Kelso, A.
Kemp, G. A.
Kempton, J. B.
Kennedy, M.
Kennedy, T. G.
Kent, G.
Keough, K. M. W.
Kerbel, R.S
Kershaw, M.
Kerzner, B.
Keystone, E. C.
Khachatourians, G. G.
Khakhria, R.
Khan, A.W
Khan, R. A.
Khanna, V.
Khoury, J.
Kibenge, F. S. B.
Kibsey, P.
Kikcio, S. I.
Kilborn, M. J.
Kilburn, D. G.
Kim, S.J.
Kimpinski, J.
King, G. J.
King, M.
King, R.
Kingma, I.
Kinlough, Rathbone R. L.
Kintz, J.
Kirby, T. J.
Kirby, W.M.M
Kisil, F. T.
Kiso, Y.
Kiyono, H.
Klein, E.
Klein, M.
Klicius, J.
Klisko, B.
Kluge, R.
Kluger, R.
Knapen, H.
Knowles, R.
Koehler, G.
Kollerup, F.
Kongshavn, P. A. L.

Koning, R.
Korber, D. R.
Kostiner, G. B.
Kourounakis, L.
Kovacs, K.
Kowbel, D.J
Kozbor, D.
Kozlowski, M.
Kozub, G. C.
Kraicer, J.
Krantis, A.
Kreiwirth, B. N.
Kremer, S.
Krestynski, F.
Krezolek, M.
Krishnan, C.
Kropinski, A. M. B.
Kropinski, A.M
Krouse, H.R
Krygsman, P.
Krystal, G.
Kudlow, J. E.
Kuhnlein, U.
Kukolj, G.
Kuksis, A.
Kuma, Mintah A.
Kunos, G.
Kurstak, E.
Kushalappa, A. C.
Kushner, D. J.
Kusser, W.
Kuyucak, N.
Kuzniak, S.
Kwan, K. K.
Kwok, Y. N.
Laakso, D. H.
Laarveld, B.
Labella, F. S.
Laberge, C.
Laberge, S.
Labow, R. S.
Labrie, F.
Labzoffsky, N.A
Lachance, J.G.
Lachance, R.
Lacoste, J.
Lacroix, A.
Lacroix, R.
Lacroute, F.
Ladd, T. I.
Ladenius, A. R. C.
Lafleur, M.
Lafond, M.
Lafontaine, P. J.

Lafreniere, C.
Lagace, J.
Lagace, Simard J.
Lagarde, A. E.
Laing, A.C
Laishley, E. J.
Lajoie, G.
Lala, P.
Lalli, C. M.
Lalonde, M.
Lalonde, R. G.
Lam, J. S.
Lam, M. Y. C.
Lam, W. L.
Lamas, S.
Lambert, I. B.
Lammerding, A. M.
Lamothe, F.
Lancaster, J. E.
Landry, M.
Langer, B.
Langleben, D.
Lannigan, R.
Lansdorp, P. M.
Lantos, G.
Lapchak, P. A.
Lapointe, J.
Lapointe, J. R.
Lapointe, M.
Lapp, M. S.
Larabie, M.
Lariviere, Durand C.
Lariviere, L. A.
Lariviere, S.
Laroche, R.
Larose, L.
Larose, P.
Larue, T. A.
Lassam, N.J.
Latour, J.G.
Lau, R. H.
Lau, R.H.
Laughrea, M.
Laurence, G.D.
Laurence, R.A
Lauri, D.
Lauterbach, P.
Lauzon, H.
Lavallee, M.
Lavoie, B.
Lavoie, G. Y.
Lavoie, J. M.
Lavoie, M. C.
Lawford, H. G.

Lawrence, J. R.
Laycock, M. V.
Layzell, D. B.
Lazar, G.
Lazarovits, G.
Le, Blanc A.E
Le, Gal M. F.
Lea, R. G.
Lebel, G.L.
Lebel, M.
Leclerc, D.
Ledbetter, J. A.
Lederis, K.
Lee, B. H.
Lee, C. Y. G.
Lee, H.
Lee, K.
Lee, K. C.
Lee, L.
Lee, P. E.
Lee, S. F.
Lee, S. T.
Lee, S. W.
Lee, S. Y.
Lees, H.
Lefcoe, N. M.
Lefebvre, G. M.
Lefebvre, S.
Legace, Simard J.
Leger, S.
Leger-Gauthier, C.
Lehmann, D. L.
Lehoux, J. G.
Lehoux, J.G.
Leibowitz, J. L.
Leitner, M.
Lejohn, H. B.
Lemaire, I.
Lemaire, S.
Lemaistre, C. F.
Lemay, G.
Lemchuk, Favel L. T.
Lemieux, M.
Lemoine, F. M.
Lemonde, P.
Lenaerts, V.
Leonard, D.
Leong, M. M. L.
Lepine, F.
Lesiak, J.M.
Lessard, C.
Lester, W. M.
Letts, G.
Leung, J. W. C.

Leung, N.
Leung, W. C.
Leung, W. W. S.
Levesque, A.
Levesque, P.
Levesque, R. C.
Levine, M.
Levinsky, H.V.
Levy, G.
Levy, G. A.
Levy, J. G.
Levy, M.
Lewing, P. J.
Lewis, B.J
Lewis, M. E.
Lewis, R. N. A. H.
Li, C.
Li, J.
Li, T. S. C.
Li, X. M.
Li, Z.
Liao, N.
Liao, S. K.
Lien, K. A.
Lieu, H.
Liew, C. C.
Lifshitz, R.
Ligas, M. W.
Light, R. B.
Ligier, S.
Likavcanova, E.
Liljelund, P.
Lillie, L. E.
Lim, Fing R.
Ling, L.
Linn, T.
Linsley, P. S.
Lior, H.
Liss, S. N.
Little, C. H. A.
Little, P.B.
Liu, D.
Lo, J.S.
Lo, R. Y. C.
Lo, T.C.Y
Lobo, A.
Lockwood, C. M.
Loewen, P. C.
Loh, B. A.
Lonergan, M.
Long, D.G.
Longenecker, B.M.
Loo, T. W.
Loosmore, S.

Lopez, A.
Lopez, J.
Lord, A.
Lorencz, B.
Louie, M.
Louie, T. J.
Loutit, J. S.
Low, D. E.
Lowe, T.
Luckevich, M. D.
Lugtig, L.
Lui, L. C.
Lukowiak, K.
Lulsdorf, M. M.
Lundberg, M. S.
Luo, H.
Luong, J. H. T.
Lupien, P. J.
Lussier, G.
Lwin, U.S.
Ma, S. W.
Ma, W.
Macdonald, I. A.
Macdonald, J. F.
Macdonald, J.A
Macdonald, K.R
Macdonald, L. A.
Macdonald, M. E.
Macdonald, M.E.
Macdonald, P. M.
Macdonald, S.
Macfarlane, T. W.
Macgillivray, R. T. A.
Macinnes, C. D.
Macintyre, S.
Mackenzie, C. R.
Mackenzie, S. L.
Mackie, G. A.
Mackie, G. O.
Maclean, A. A.
Maclean, L.D.
Macleod, J. E.
Macleod, K. M.
Macmillan, D. A.
Macnaughton, W. K.
Macphee, P. J.
Macpherson, L. M. D.
Macrae, S.
Macsween, H. M.
Macsween, J. M.
Madduri, K.
Madryga, F. J.
Mager, D.
Maguire, R.J

Mah, D.
Maher, E.
Mahony, J. B.
Maier, T.
Majak, W.
Mak, T. W.
Maki, A Jr
Makowka, L.
Male, K. B.
Malek, L. T.
Malmfors, T.
Malott, D.W.
Malsbury, C. W.
Mamet, Bratley M. D.
Man, M.S. .
Manavathu, E. K. .
Mancino, L.
Mann, V. M.
Manns, J. G.
Manuel, J.
Mapletoft, R. J.
Margolese, R. G.
Margolis, B. L.
Marinier, R.
Marion, D.
Markham, R. J. F.
Markowitz, S.
Marois, M.
Marois, Y.
Maroun, J. A.
Marsden, P. A.
Marshall, J. S.
Marshall, K. C.
Martel, A.
Martel, R.R.
Martell, M.
Martin, De Leon P.A.
Martin, L.
Martin, R. R.
Marusyk, R. G.
Marx, J.
Mason, P.
Mason, S.T
Massicotte, J.
Matheson, A.T.
Mathieu, L.G.
Mathieu, Serra A.
Matsui, T.
Maxie, M. G.
Maxwell, I. H.
Mayfield, C.I.
Mayrand, D.
Mazaheri, R.

Mazumder, A.
Mcallister, T. A.
Mcauley, L.
Mccalla, D. R.
Mccarter, J. A.
Mccoll, S. R.
Mccready, R.G.L.
Mcculloch, P. B.
Mccurdy, H.D
Mcdermid, K. P.
Mcdermott, G. F.
Mcdermott, J. J.
Mcdonald, J. G.
Mcdonald, J.C.
Mcdougall, L.
Mcdowell, J.
Mcduffie, N. G.
Mcelhaney, R. N.
Mcfadden, R. G.
Mcfarlane, C. S.
Mcfarlane, E. S.
Mcgavin, M. J.
Mcgeer, E. G.
Mcgeer, E.G.
Mcgeer, P.L
Mcgee, J. R.
Mcgill, W. B.
Mcgirr, L. G.
Mcgraw, B.M
Mchugh, N.
Mcinnes, A. G.
Mcintosh, A. R.
Mcintosh, C. H. S.
Mcintosh, E. M.
Mciver, D. J. L.
Mckay, T.
Mckenna, P. K.
Mckeown, B. A.
Mckinley, J. B.
Mcknight, G. W.
Mclean, D. M.
Mclean, R. H. C.
Mclellan, B. A.
Mclennan, H.
Mcminn, R. G.
Mneil, R.
Mneill, K.
Mcperson, J.
Mcrae, K.B.
Meaney, M.
Meaney, M. J.
Mecheri, S.
Medina, V.
Meghji, K.

Meiering, A. G.
Meighen, E. A.
Meisels, A.
Melancon, P.
Melnyk, D.
Mendelson, J.
Meneghetti, C. M.
Merand, Y.
Mes, Hartree M.
Mes, Masson A. M.
Messier, S.
Metcalf, A.
Metcalf, D. R.
Miarczynski, D.
Miceli, M. O.
Micetich, R. G.
Michel, T.
Michon, F.
Miginiac, Maslow M.
Milburn, P. H.
Miller, F. W.
Miller, G.G.
Miller, M. A.
Miller, R. C Jr
Miller, R. G.
Milliron, M. L.
Mills, J. T.
Millward, S.
Milne, R.
Milton, G.
Miniats, O. P.
Miniats, O.P
Mirakian, A.
Misener, V.
Misra, S.
Misra, V.
Mistry, G.
Mitchell, C.A
Mittal, K. R.
Mobry, P. M.
Moe, G. W.
Mogenson, G.J
Mohapatra, S. S.
Mohring, J.
Moineau, S.
Mokresh, A. H.
Mokry, J.
Moldoveanu, N.
Molgat, G. F.
Molnar, S.
Molrency, M. J.
Monck, M. A.

Monette, P. L.
Mongeau, C. J.
Montplaisir, J.
Moor, B. C.
Moore, G. J.
Moore, K. C.
Moore, R. A.
Moorjani, S.
Morales, A.
Morales, M. A.
Morali, G. A.
Moran, E. T Jr
Moresoli, C.
Morgan, W. K. C.
Morgante, O.
Morin, C.
Morin, P.
Morin, Y.
Morisset, J.
Morris, V. L.
Morrow, G. W.
Mort, J. S.
Morton, K. D. R.
Moser, B.
Mourad, W.
Movat, H. Z.
Mowery, J.
Mueller, C. R.
Mukherjee, B. B.
Mukkur, T.K.S
Mulvey, M. R.
Murdoch, J. H.
Murphy, B. D.
Murray, G.
Murray, R. G. E.
Murray, W.D.
Mustard, J. F.
Mutt, V.
Myers, D. J.
Myers, Mason N.
Nabi, I. R.
Naccache, P. H.
Nadeau, M.
Nadeau, P.
Nadeau, R.
Nadler, S. P.
Nag, K.
Nagy, E.
Naiman, S. C.
Nakamura, Y.
Nance, D. M.
Nanci, A.
Nano, F. E.

Nasmith, P. E.
Nathanielsz, P.
Nattel, S.
Naus, C. C. G.
Nazar, R. N.
Nelson, L. M.
Nelson, W. R.
Neshat, S.
Nestmann, E.R.
Neugebauer, E.
Newcomb, W.
Newhouse, M.
Ng, L. K.
Ngeleka, M.
Nguyen, T.M.D.
Nicholson, R.
Nicholson, J. W. G.
Nicholson, R.
Nickel, J. C.
Nicolle, L. E.
Nielsen, K. H.
Niilo, L.
Ninniss, R. W.
Nishi, S.
Niven, D. F.
Nodwell, J. R.
Noel, C.
Northover, J.
Northrop, D. B.
Nossal, N.
Notenboom, R. H.
Nowicki, T. W.
Nsanze, H.
Nurse, C. A.
Obrodovich, H. M.
Odense, P. H.
Ogilvie, K. K.
Ohara, C. J.
Ohashi, P. S.
Okane, D. J.
Okazawa, M.
Oku, H.
Oldham, W. K.
Olivier, C.
Olivier, M.
Oloughlin, E. V.
Olsen, E.
Olson, M. E.
Olsson, O.
Ong, H.
Oreilly, T.
Orr, F. W.
Orr, P.
Osmond, D. G.

Ossenkopp, K. P.
Ottaway, C. A.
Owens, J. N.
Owens, T.
Owttrim, G. W.
Oyelese, A. O.
Ozanne, G.
Pabst, H. F.
Pacepavicius, G.
Packham, M. A.
Page, M.
Pagel, J. E.
Pagtakhan, R.D.
Painson, J. C.
Panagia, V.
Pandian, S.
Pantaloni, C.
Pap, K.
Papp, Vid G.
Pappu, H. R.
Paradis, R.
Paraskevas, F.
Pare, P. D.
Parekh, S. R.
Parent, A.
Parfrey, P. S.
Park, S. J.
Park, W.K
Parker, M. D.
Parker, S.
Parkin, K. L.
Parkinson, D.
Parnell, T.
Parniak, M.
Parry, L. J.
Parsons, R. V.
Parsons, T. R.
Pass, M. A.
Passador, L.
Patel, M.
Patel, P.
Patel, Y. C.
Paterson, M.C.
Paterson, N. A. M.
Patil, A. D.
Paton, D.M.
Patrick, M. K.
Patterson, T. G.
Paul, B.
Paul, L. C.
Paulitz, T. C.
Pausova, Z.
Pavlicik, V.
Payette, M.

Pazderka, F.
Pearce, C. M.
Pearce, M.
Pearson, D.
Pearson, E.W
Pechere, J. C.
Pecquet, S. S.
Pederson, R. A.
Pelcher, L. E.
Pelletier, G.
Peltre, G.
Pena, S. D. J.
Pennells, G. C. L.
Penner, J. L.
Penney, C. L.
Penney, M. T.
Pepler, M. S.
Pepys, M. B.
Pereles, L.
Perez, Gil J.
Perlanski, E.
Perreault, C.
Perreault, T.
Perrone, M. A.
Perry, K. J.
Peter, R. E.
Peterkin, P. I.
Peterson, J.
Peterson, J. B.
Petit, A.
Petitclerc, D.
Petric, M.
Pettigrew, M.
Pettit, J. R.
Pharoah, G.
Phillip, L. E.
Phillips, A.
Phillips, D. A.
Phillips, D. M.
Phillips, M. J.
Phillips, R.
Picard, D.
Picard, V.
Picaud, A.
Piche, Y.
Pichon, F.
Picorel, R.
Piechuta, H.
Pier, G. B.
Pierce, C.H.
Pierce, G. N.
Pim, C.
Pinski, J.
Pinto, B. M.

Pittman, Q. J.
Plante, G. E.
Plisetskaya, E. M.
Plourde, G.
Plummer, F. A.
Poffenroth, L.
Poirier, G.
Polc, P.
Polidoulis, I.
Pollak, M. N.
Polosa, C.
Pomerantz, D. K.
Pomeranz, I. S.
Pontarollo, R.
Poole, A. R.
Poole, K.
Poon, K.
Pope, B. L.
Pope, C.
Poplonski, L.
Porter, K. G.
Portnoy, J. D.
Posner, B. I.
Pot, J.
Potter, A. A.
Pouyssegur, J.
Povey, R.C
Power, M. E.
Prasad, D.
Prasad, E.
Prentis, J.
Prevost, D.
Price, G. B.
Prior, M. G.
Procter, B.G.
Pross, H. F.
Proulx, M.
Pruzanski, W.
Purchase, E. C. R.
Qi, B. J.
Quail, J. A.
Quantz, M. A.
Quarrington, C.
Quee, T. C.
Quessy, S.
Quinn, T.
Quirion, R.
Qureshi, J.
Rabinovitch, N.
Rabinowitz, B.
Rabkin, S. W.
Radika, K.
Rae, K.
Rainbow, A. J.

Rajakumar, N.
Rajaraman, R.
Rajaraman, S.
Ralph, E.D.
Ramia, S.
Ramotar, K.
Ramsden, D. A.
Randle, J. C. R.
Ransom, T. P. P.
Rao, A. V.
Rao, L. G.
Raper, C. D Jr
Rapin, N.
Rashid, A.
Rawls, J. A.
Rawls, W. E.
Rawls, W.E.
Read, S.
Rector, E. S.
Reece, D. E.
Reeves, W. C.
Regoeczi, E.
Regoli, D.
Regoli, D.C
Reimer, K. B.
Reithmeier, R.A.F.
Remer, Z.
Renaud, L. P.
Renken, A.
Rennie, R. J.
Renton, K. W.
Renzo, A.
Retnakaran, A.
Rhaleb, N. E.
Rho, D.
Ribeiro, Da Silva A.
Ribuot, C.
Richards, G. K.
Richards, J. E.
Richards, W. R.
Richardson, C. D.
Richardson, P. M.
Richardson, S.
Richter, M.
Riddell, R. H.
Riddle, D. L.
Riendeau, D.
Rigby, C. E.
Ring, R. A.
Riopelle, R. J.
Rioux, F.
Rivard, S. R.
Rivers, D.
Riviere, C. J.

Rivington, R. N.
Roberts, F. J.
Roberts, J.R
Robertson, F. J.
Robertson, J. A.
Robey, W. G.
Robin, J.
Robinette, M.
Robinson, C. W.
Robson, D. A.
Robson, H. G.
Rochette, L.
Rochon, D. M.
Rochon, Y. P.
Rock, G.
Rocourt, J.
Rodaros, D.
Rode, L. M.
Roder, J. C.
Rodrigue, L.
Rodriguez, A.
Roeder, R. G.
Rogalski, T. M.
Rogers, R. E.
Rokach, J.
Rola, Pleszczynski M.
Rollins, L.
Romanowski, B.
Rompre, P. P.
Ronald, A. R.
Ronald, W.P
Rondeau, J. J.
Rood, S. B.
Rosendal, S.
Roslycky, E. B.
Ross, A. D.
Ross, Chouinard A.
Rossant, J.
Rossignol, B.
Rotstein, L. E.
Roughley, P. J.
Rouse, B.T
Rousseau, A.
Rowley, R. B.
Roy, D. C.
Roy, M.
Roy, M. C.
Roy, P. E.
Roy, R.
Rubin, B.
Rubin, P.
Rupitz, K.
Rusak, B.
Ruseska, I.

Rushlow, W. J.
Russell, I.
Ryan, J. A.
Rydall, J. R.
Saad, W. P.
Sabara, M.
Sabour, M.
Sacks, S. L.
Sackston, W. E.
Saddler, J. N.
Saddler, J.N.
Saenger, P.
Saez, J. M.
Saftic, S.
Sahai, A.
Saito, S.
Salari, H.
Salomons, M. O.
Samagh, B. S.
Sambhara, S. R.
Sandberg, G.
Sanderson, K. E.
Sanfacon, H.
Sangalang, V. E.
Sankaranarayanan, A.
Sanni, A.
Sargent, M.D.
Sarmiento, U. M.
Sasakura, Y.
Sato, H.
Sattar, S. A.
Sattar, S.A.
Sauder, D. N.
Sauer, W. C.
Saunders, J. M.
Saunders, J. R.
Saunders, W. S.
Savard, P.
Savoie, S.
Sawynok, J.
Scalbert, E.
Schaaper, R. M.
Schappert, K. T.
Schechter, M. T.
Scheibling, R. E.
Schellenberg, R. R.
Scher, F. M.
Schiefer, H.B
Schieven, B. C.
Schmiel, D.
Schneider, T. D.
Schoun, J.
Schrader, J. W.
Schurch, S.

Schwartz, C. J.
Science, Russel A.
Scott, G. W.
Scott, R. B.
Scott, Taylor T.
Seagull, R. W.
Sebunya, T. N. K.
See, R. H.
Seech, A. G.
Segall, J.
Seguin, A.
Sehon, A. H.
Seip, A. E.
Sekla, L.
Selvaraj, G.
Selzler, L.
Semba, K.
Semple, J. W.
Sengar, D.P.
Senikas, V. M.
Setterfield, G.
Seuwen, K.
Seyfried, P.L.
Shaffer, E. A.
Shaffer, Tannenbaum G.
Shah, A.
Shah, P. P.
Shaikh, N.A.
Shapiro, S.
Sharma, A.
Sharpe, A. N.
Shattuck, V.
Shaw, D. H.
Shek, P. N.
Shelford, J. A.
Shelp, B. J.
Shenouda, N.
Sheppard, M. S.
Sheps, S. B.
Sheu, L.
Shewen, P. E.
Shibuya, T.
Shieh, H. S.
Shieh, H.S
Shiffman, N.
Shimo, Takahara S.
Shiu, R. P. C.
Shizgal, H.M
Sia, C.
Siebenmann, C.O.
Silim, A.
Silverman, M.
Silversides, D. W.
Simard, C. L.

Simard, J.
Simatos, G.
Siminovitch, K. A.
Simon, M.
Simon, P. C.
Simor, A. E.
Simpelkamp, J.
Simpson, S.
Sinclair, P.
Singal, D.
Singal, D. P.
Singh, A.P.
Singh, E. L.
Singh, G.
Singh, H.
Singh, I. P.
Singh, M. P.
Singh, R.
Singh, R. P.
Singla, C. L.
Sinha, R. N.
Sinnott, N. M.
Sircar, R.
Sirois, P.
Sit, T. L.
Sitbon, F.
Skamene, E.
Skorecki, K.
Skorecki, K. L.
Skup, D.
Slapsys, R. M.
Slocombe, J.O.D.
Smirnoff, W. A.
Smith, B.P.
Smith, D. W.
Smith, E.A.
Smith, E.K.M.
Smith, J. A.
Smith, J. W.
Snajdarova, I.
Snelling, C. F. T.
Sniderman, K. W.
Snow, K. M.
Snutch, T. P.
Soll, A. H.
Somerville, T. H.
Sonenberg, N.
Song, J.
Sonnenberg, H.
Sorensen, O.
Souhami, L.
Spady, D. W.
Speert, D.
Speert, D. P.

Spence, L.
Spence, L.P
Spencer, J. L.
Spencer, V.
Spero, L.
Spiegelman, G. B.
Spira, B.
Spitzer, G.
Sprague, E. A.
Springthorpe, V. S.
Sprott, G. D.
Spruston, N.
Srikant, C. B.
St, Amand J.
St, Germain J.
St, Louis J.
St, Pierre C.
St, Pierre D. M. M.
St, Pierre S.
Stace, Smith R.
Stanacev, N. Z.
Stanisz, A.
Stanisz, A. M.
Stankova, J.
Stanners, C. P.
Starratt, A. N.
Starreveld, E.
Stauffer, G. V.
Stedronska, Clark J.
Steinmann, T. I.
Stemke, G. W.
Stemshorn, B. W.
Stephen, K. W.
Stevenson, Ian L.
Stevenson, M. M.
Stevenson, R. M. W.
Stevnsborg, N.
Stewart, D. J.
Stewart, G. G.
Stewart, J.
Stewart, R. B.
Stewart, T. H. M.
Stickney, H.
Stiles, M. E.
Stillman, M. J.
Stoakes, L.
Stobbs, L. W.
Stockner, J. G.
Stoessl, A.
Stone, B. R.
Storey, D. G.
Storey, K. B.
Stormo, G. D.
Strasberg, S.

Strathdee, C. A.
Street, I. P.
Streit, W.
Strejan, G. H.
Stricker, E.M
Strnad, C. F.
Stuhne, Sekalec L.
Sturges, W. T.
Stuttard, C.
Subden, R. E.
Subrahmanyam, T.P.
Sullivan, A. K.
Sullivan, S. N.
Sulochana, S.
Summerlee, A. J. S.
Sun, A. M.
Sun, S. K.
Sung, J. Y.
Superina, R.
Susan, M.
Suzuki, I.
Swartzman, E.
Sweeney, G. D.
Sweeney, M. I.
Sweeting, R. M.
Switala, J.
Sylla, B. S.
Szalay, A. A.
Szarek, W. A.
Szewczuk, M. R.
Szilvassy, S. J.
Szpirer, C.
Szpirer, J.
Tabel, H.
Tackaberry, E. S.
Tai, G. C. C.
Takahashi, I.
Takahashi, Y.
Talor, E.
Tan, B.
Tan, D. T. M.
Tan, R.
Taneva, S.
Tannenbaum, G. S.
Tarrand, J. J.
Taubchi, K.
Taurus, T. E.
Tawar, V.
Taylor, A.
Taylor, B.
Taylor, E.
Taylor, F. J. R.
Taylor, L.
Taylor, Robinson D.

Tchakarov, L.
Teh, C. Z.
Teh, H. S.
Teh, S. J.
Teitel, J.
Telemaque, S.
Temesvari, P.
Tence, M.
Thatte, V.
Thavundayil, J.
Thibault, A.
Thibault, L.
Thibault, P.
Thiers, H.D
Thiessen, J.J.
Thng, C.
Thode, H. G.
Thomas, D. Y.
Thomas, F. C.
Thomas, G.
Thomas, K. C.
Thompson, J.
Thompson, J. S.
Thomson, R. J.
Thornhill, J. A.
Thorsen, J.
Tian, S.
Tilley, P.
Tinline, R. R.
Tisa, L.S.
Tiwari, K. N.
To, M.
Tobe, S.
Tobey, J. C.
Tobin, R. S.
Todesco, T.
Tolias, P. P.
Tomar, R. S.
Tomita, F.
Too, C. K. L.
Toogood, J. H.
Tostowaryk, W.
Tousignant, C.
Towell, M. E.
Towers, G. H. N.
Townsend, G. E.
Tremaine, J. H.
Tremaine, J.H.
Tremblay, C.
Tremblay, F. M.
Tremblay, J.P.
Treves, S.
Trevors, J. T.
Trew, J. A.


Triggs, B. L.
Trimble, R. M.
Trimbur, D.
Trip, E.
Tron, V. A.
Trottier, S.
Trudel, I.
Trudel, L.
Trust, T. J.
Tse, K. S.
Tse, K.S
Tsui, F. W. L.
Tsui, L. C.
Tsui, W. C.
Tucker, J.
Tuffnell, P.G.
Tulloch, A. P.
Turcotte, A.
Turcotte, B.
Turgeon, F. F.
Turgeon, P.
Turgeon, P.L
Turkington, R.
Turley, W.
Twiddy, R.R.
Tyler, K. L.
Urtasun, R.C.
Usman, N.
Utkhede, R.
Utkhede, R. S.
Vadas, P.
Vadeboncoeur, C.
Valente, A. J.
Valera, V.
Valli, V. E.
Valli, V. E. O.
Valvano, M. A.
Van, De Kar L. D.
Van, Den Hurk J. V.
Van, Drunnen Littel Van Den Hurk S.
Van, Dyke M. I.
Van, Everdingen R. O.
Van, Loon G. R.
Vanderwel, D.
Varughese, P.
Vassos, T. D.
Vats, S.
Vaudry, H.
Veldhuyzen, Van Zanten S. J. O.
Venkateswarlu, K.
Venne, D.
Verge, V. M. K.
Verma, D. P. S.
Verreault, A.

Verreault, H.
Vessey, J. K.
Vethamany, Globus S.
Vezina, G.
Vezina, W.
Viau, V.
Villanueba, K.
Vincent, S. R.
Vining, L. C.
Vinitski, S.
Viol, G.W.
Virdi, K.
Visentin, L.P
Viswanatha, T.
Vogel, H. J.
Voigt, D. R.
Volesky, B.
Von, Rudloff E.
Vydelingum, S.
Wade, A. W.
Wainberg, M. A.
Walczyk, E.
Walker, M.
Walker, P.
Wall, R.
Wallace, C. J. A.
Wallace, J. L.
Wallace, R.
Wallace, V. A.
Wallbank, A.M.
Wallerstein, R. O.
Wallis, P. M.
Walsh, A.
Walsh, K.
Walter, J. A.
Walton, R.
Wan, K. K.
Wang, H. Y.
Wang, N. S.
Wang, P.Y.
Wang, W. S.
Wanklin, R. J.
Ward, E. W. B.
Ward, O. P.
Warner, H. A.
Warren, C. P. W.
Warren, K. G.
Warren, R. A. J.
Warren, W.P
Warrington, R.J.
Wat, L. L.
Waterer, J. G.
Watson, W.
Waye, M. M. Y.

Wayman, M.
Weaver, L. C.
Webster, J. M.
Webster, W. A.
Wei, C.
Weld, M. M.
Welham, M. J.
Wellington, C. L.
Wellington, W. G.
Wells, G. A.
Werner, D.
Werstuck, G.
Wesley, I. V.
West, M.
Westlake, D. W. S.
Wheeler, M. B.
White, B. A.
White, B. N.
White, J. L.
White, K. A.
White, M. J.
White, R. L.
Whiteside, C. I.
Whitfield, C.
Whitney, F. A.
Wieczorek, A.
Wieler, J. S.
Wight, C. P.
Wildeman, A. G.
Wilfong, A. A.
Wilkie, J.S.
Wilkinson, R. D.
Willetts, N.
Willey, B. M.
Williams, D. R.
Williams, D.T.
Williams, Fulton N. R.
Williams, G.
Williams, M.C.
Williams, R.
Williams, T.
Wilmot, S. J.
Wilson, D. R.
Wilson, N.
Wise, R. A.
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Withers, S. G.
Withler, R. E.
Witt, Sullivan H.
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Wojtczak, G.
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Wong, C. S.

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Woo, P. T. K.
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Wright, J. B.
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Wu, J.
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Yacoob, R.
Yada, R. Y.
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Yamaguchi, N.
Yamamoto, Y.
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Yang, C. R.
Yang, W. H.
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Youson, J. H.
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Yu, S.
Yulis, C. R.
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Zamb, T. J.
Zamboni, R.
Zealey, G.
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Zeng, W.

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Zhang, L.
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