Cancer Dermatology

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Foreword

This collective effort was undertaken by members and former members of the medical staff of the Roswell Park Memorial Institute to make available a clinical guide to aid in the diagnosis and treatment of the malignant diseases afflicting the human integument. This is not intended to be a textbook, nor an encyclopedic work restricted to the reference library, but rather a concise manual to be readily available and easily used when needed.

However, even on the most pragmatic level, merely didactic methodology is not enough. Our concern must include concepts, no matter how briefly stated. Thus condensed discussions of the various facets of such large complex fields of inquiry as carcinogenesis, brief descriptive profiles of the more important cutaneous neoplasms, and consideration of neoplasms of other organs that invade or influence the skin are provided to help the therapist attain understanding as well as skill. To extend the horizon further. most of the chapters present carefully selected references for the reader who wishes to probe deeper and more broadly.

Part III is essentially a descriptive summary of the actual techniques used by cach member of a team composed of difterent specialties working in our busy cancer clinic. The unusual opportunity for cooperative and easy exchange of consultations among the various specialists, dermatologists, radiologists, surgeons, chemotherapists. pathologists, cytologists, and others have been a catalyst of incalculable value in potentiating the very rich personal experience of the individual therapist. This is transmitted not only into better patient care, but also into better research.

In any undertaking of multiple authorship, some overlap of discussion is inevitable. On the other hand, this very feature has the advantage of bringing into focus differences of emphasis from the perspective of more than one area of interest.

It is hoped that this book will fulfill the expectation of the reader by expanding his awareness of diagnostic possibilities and helping him weigh the advantages and disadvantages of available therapeutic options, as well as provide some guidance in the employment of the modality he has chosen.

The editor is to be commended on the judicious selection of topics, which for the most part encompass the whole spectrum of cancer dermatology.

GERALD P. MURPHY, M.D., D.S.C. Director Roswell Park Memorial Institute

Preface

More malignant neoplasms occur on the skin of man than on any other anatomic site. It is estimated that from 300,000 to 600,000 skin cancers develop each year in the United States alone. The skin is a site most amenable to therapy; however, an accurate diagnosis and a proper assessment of the circumstances are necessary. In addition, neoplasms on the skin can serve as a model system because of their accessibility and thereby increase our knowledge about the broader aspects of tumor biology.

This book stresses a team approach to the management of skin cancer. As practiced at Roswell Park Memorial Institute in Buffalo, the concept has worked well for both the patient and the physician. Access to immediate consultation with all specialties concerned in the treatment of cancer (dermatology, pathology, plastic surgery, radiology, and oncology) not only helps to quickly establish a diagnosis, but also to outline a plan of early treatment. The dermatologist has a great advantage, in that he is not only trained to make a correct diagnosis based on the clinical appearance and histologic examination, but he is also acquainted with all treatment modalities, without necessarily using them himself. Therefore without bias, he is better qualified to refer patients for the appropriate therapy.

Buffalo, New York Frederick Helm, M.D.

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I also wish to express my appreciation for the technical assistance and secretarial efforts of many persons.

All pictures, when not otherwise marked, are from the collection of the Department of Dermatology at Roswell Park Memorial Institute, Buffalo, New York.

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Contents

| PART | T | ETIOLOGY | OF | NEOPI. | ASMS | OF | SKIN |
|------|---|----------|----|--------|------|----|------|

| | 1. | Epidemiology | 3 |
|---------------|---|--|------------------------|
| | | Carcinogenesis | 17 |
| | | 2. Experimental Carcinogenesis Fred G. Bock | 20 |
| | 3. | Epithelioma from Single Trauma Howard L. Stoll, Jr., and John T. Crissey | ; 5 |
| | 4. | Late Radiation Injury and Cutaneous Neoplasia Herbert L. Traenkle† | 31 |
| | 5. | Genodermatoses Associated with Malignancies Bernard F. McEvoy | 39 |
| | | OF THE CLUB | |
| PART II. NEOP | LASMS | OF THE SKIN | |
| PART II. NEOP | | OF THE SKIN Precancerous Lesions | 59 |
| PART II. NEOP | 6. | Precancerous Lesions | 59 81 |
| PART II. NEOP | 6.7. | Precancerous Lesions | |
| PART II. NEOP | 6.7.8. | Precancerous Lesions | 81 91 |
| PART II. NEOP | 6. 7. 8. | Precancerous Lesions John C. Maize and James E. Rasmussen Intraepidermal Epithelioma of Borst and Jadassohn Karl Holubar Basal Cell Epithelioma Gordon H. Burgess and Blair V. Jager Malignant Tumors of Skin Appendages | 81 91 103 |
| PART II. NEOP | 6.7.8.9.10. | Precancerous Lesions John C. Maize and James E. Rasmussen Intraepidermal Epithelioma of Borst and Jadassohn Karl Holubar Basal Cell Epithelioma Gordon H. Burgess and Blair V. Jager Malignant Tumors of Skin Appendages Kenneth Hashimoto Squamous Cell Carcinoma | 81 91 103 113 |

| | 12. | Malignant Melanoma | 141 |
|------------------|-----|---|-----|
| | 13. | Soft Tissue Sarcomas | |
| | 14. | Kaposi's Hemorrhagic Sarcoma | 177 |
| | 15. | Malignant Tumors of Blood and Lymph Vessels Frederick Helm | 185 |
| | 16. | Lymphoma Cutis | 191 |
| | 17. | Mycosis Fungoides | 205 |
| | 18. | The Sézary Syndrome | 225 |
| | | Cutaneous Lesions in Multiple Myeloma Peter J. Lynch | 229 |
| | 20. | Malignant Tumors of the Oral Cavity | 235 |
| | 21. | Cutaneous Markers of Internal Malignancies Frederick Helm and Juta Helm | 247 |
| | 22. | Pseudomalignancies | 285 |
| PART III. MANAGE | EME | ENT OF SKIN CANCER | |
| | 23. | Prevention of Skin Malignancies | 313 |
| | 24. | Diagnostic Procedures | |
| | | Maria C. Gamarra 2. Skin Biopsy John F. Gaeta | |
| | 25. | Surgery | 325 |
| | 26. | Ionizing Radiation 1. X-ray Therapy Walter Murphy 2. Electron Beam Therapy Moshe Friedman and John I. Pearce | 365 |
| | 27. | Electrosurgery | 411 |

| CONTENTS | xvii |
|----------|------|
| | |

| 28. | Cryosurgery 421 |
|------------|---|
| 201 | Gordon H. Burgess |
| 29. | Mohs' Chemosurgical Technique 425 John T. Phelan and Halina Milgrom |
| 30. | Chemotherapy |
| | 2. Systemic Chemotherapy |
| | 3. Chemotherapeutic Agents |
| 31. | Introduction to Tumor Immunotherapy 471 Pierluigi E. Bigazzi, Edmund Klein, and Frederick Helm |
| 32. | Comparison of Different Methods of Treatment 481 Frederick Helm |
| APPENDIXES | |
| | Living with Lymphoma |
| | Counseling the Patient |
| INDEX | |

| | | | | _ |
|-----|-------|-------|-----|---|
| 00 | JA II | אישים | THE | 2 |
| 1.1 | ON' | | NIC | 3 |

xvii

| 28. | Cryosurgery |
|------------|--|
| 29. | Mohs' Chemosurgical Technique 425 John T. Phelan and Halina Milgrom |
| 30. | Chemotherapy |
| | Systemic Chemotherapy |
| 31 | Introduction to Tumor Immunotherapy 471 Pierluigi E. Bigazzi, Edmund Klein, and Frederick Helm |
| 32 | Comparison of Different Methods of Treatment 481 Frederick Helm |
| APPENDIXES | |
| | Living with Lymphoma |
| | Counseling the Patient |
| INDEX | * 495 |

Part I Etiology of Neoplasms of Skin

1 Epidemiology

Josef Vana

Cancers of the skin are the most common malignant tumors of man, at least of white man. It is surprising, therefore, that they have received so little attention from the epidemiologist. Their high frequency, relatively easy diagnosis, and suitability for detection by screening techniques should render these tumors highly relevant for epidemiologic study. There are, however, significant obstacles in data collection. It is difficult to obtain complete and reliable information on all skin cancers occurring in a defined population during a specified period of time. Most skin cancer patients are treated in physicians' offices or clinics, and histologic diagnosis is not always made. Data on such patients are not systematically recorded in registries, hospital charts, or records of pathology laboratories or outpatient facilities. These difficulties are aggravated both in countries with a large sector of private medical practice and in countries with underdeveloped health services.

Epidemiologic cancer research has been greatly facilitated in the past 20 years by the growing network of regional, national, and international cancer registries. Al-

though useful in providing information on many neoplasms, cancer registries are notably unsuccessful in compiling data on malignant skin tumors. Many registries that began by including all malignant tumors have dropped the obligatory reporting of skin cancer, except melanoma, because of the difficulties in obtaining complete data. In some instances ad hoc population surveys in registry populations have confirmed how selective and fragmentary the registry data on skin cancer were.

In recognition of these obstacles in data collection, research workers have turned away from nationwide efforts to ad hoc surveys in selected areas to obtain the true prevalence or incidence of skin cancer in the population. Recent studies of this type have shown that the real extent of skin cancer is much higher than that indicated by national statistical estimates or registry data. 49.68,80 In one such study in Victoria, Australia, for every four known and treated cases, roughly three untreated cases were detected.73 In the Third National Cancer Survey conducted by the U.S. National Cancer Institute in 1969-71. superficial skin cancers were found to be

so underreported that incidence may vary from 300,000 to 600,000 new cases annually.

A further constraint that slows the progress of epidemiologic research in skin cancer is the variation in reliability of diagnostic procedures. Diagnoses should be verifiable, and histologic evidence should be available for confirmation. At present, histologic verification is not recorded for all reported cases even in areas where laboratory facilities are available. Moreover, pathologists are still likely to differ considerably in their microscopic typing of the common types of skin cancer.

Malignant melanoma differs significantly from the superficial skin cancers: it has a much lower incidence but a much higher mortality. In contrast to the other skin cancers, it is unlikely to be treated on an outpatient basis, far more likely to reach hospital diagnosis, treatment, and therefore records. Many registries that have discontinued notification of the superficial skin cancers retain registry of malignant melanoma. Moreover. melanoma is classified separately from other skin cancers in the international codes for certification of death, and in some countries site of primary lesion is also routinely recorded on the death certificate. The greatest constraint to epidemiologic study of malignant melanoma is its low incidence: large population data bases or pooled uniform data are necessary to provide a sufficient number of cases for epidemiologic study. The improved regional and national health statistics developed over the past years are beginning to provide such a data base. Moreover, improved diagnostic and therapeutic techniques with consequent improved survival rates are prompting a new focus on etiology and prevention of malignant melanoma.

Squamous cell and basal cell carcinomas and melanoma together comprise by far the largest group of malignant skin neoplasms. For these tumors some data on frequency, site, and type are available for epidemiologic review.

MORTALITY

Unlike statistics for cancers of high malignancy, mortality statistics for superficial skin cancer cannot be used for even rough estimates of the extent of morbidity. Death rates for squamous cell and basal cell carcinoma reflect not only incidence but also the fatality rates, the level of health services provided, and the comprehensiveness of the statistical reporting systems. Furthermore, as deaths from malignant melanoma comprise up to two thirds of all deaths from primary malignant tumors of the skin,40,42 mortality figures for skin cancer are inflated with respect to superficial skin cancers and therefore are misleading.

Death rates for malignant melanoma alone are more useful. Comparative studies have shown that in areas where incidence data are not available, estimates of incidence may cautiously be drawn from mortality data. 45 Since 1957 deaths from melanoma have been classified and reported separately from other skin cancers in the International Statistical Classification of Diseases, Injuries, and Causes of Death (7th Revision), and this has facilitated analysis of mortality data on an international level. Death rates for malignant melanoma are reported to range from 3.2/100,000 population in Australia⁶ to 0.2/100,000 population among Asian females.65 In the United States the mortality rate in 1967 was 1.6/100,000 among both sexes of the white population.41

Considerable variation in melanoma death rates exists within, as well as between, countries. In Australia, an increase in death rates has been observed as one proceeds toward the Equator from Victoria through New South Wales to Queensland, and this increase in mortality by latitude has been observed every year since 1950. 16 Recent analysis of age-

standardized malignant melanoma mortality rates in the United States and Canada by state and province shows a strong negative association of mortality with latitude of the largest city. ²¹ Moreover, extensive variations in mortality are also found even within certain states. For example, in Nebraska death rates by county among males range from 0.4 to 17.8/100,000, and in Georgia the rates for females range from 0.4 to 11.4. ⁵³

Although of limited value for epidemiologic study of morbidity, skin cancer mortality data did provide the first indications of the large geographical differences in the distribution of skin cancer. thus suggesting the etiologic importance of environmental factors. These large differences were later confirmed by incidence studies. Reported age-adjusted death rates for total skin cancers in 24 countries for 1958-59 ranged from 3.8/ 100,000 population among Australian males to 0.7/100,000 in Japanese females.69 In the United States ageadjusted skin cancer mortality rates in 1969-71 were 2.4/100,000 for males and 1.5/100,000 among females.1

INCIDENCE

Published studies of incidence are based on large clinical series compiled over a long period of time, on registry data, and on population surveys. The clinical series from hospitals provide useful information, but they do not necessarily include all cases in a defined population and cannot therefore be used to derive true incidence rates. The varying quality of registry data has already been mentioned; it should be pointed out, however, that registries in countries with comprehensive national health insurance that include notification from all types of services are often able to provide reliable incidence data on skin cancers. The population surveys provide the most reliable information on prevalence.

One striking feature of a review of the

literature on frequency and distribution of skin cancers is that the same pattern of geographic variation emerges. The highest incidence of skin cancers of all types is reported from geographic areas where fair-skinned people are exposed to large amounts of ultraviolet radiation. Reported frequencies are lower among darkskinned and Asian people and lower in areas where populations are exposed to less ultraviolet radiation. 18,72

These geographic differences span a wide range. Age-standardized incidence rates for skin cancer (excluding melanoma) in various countries range from 0.5/ 100,000 population among Bantu males in South Africa to as high as 239.7 new cases per 100,000 among non-Latin males in El Paso, Texas—a 480-fold span. Rates among Europeans are, roughly, in the range of 20 to 40/100,000 for both sexes, though the incidence figures for females lie generally in the lower range. Incidence rates for skin cancer (excluding melanoma) in various countries range from 0.5/ the exception of colored people in Colombia, Jamaica, and Puerto Rico, where the incidence is reported to approach European levels. Similar large variations in incidence have also been observed within. as well as between, countries, 18,31 Selected incidence rates for skin cancer as well as data on skin cancer as a percentage of all cancers in selected countries are given in Table 1-1.

Incidence of malignant melanoma is much lower and spans a more narrow range but follows the same geographic pattern. Reported rates range from 0.1/100,000 in Japanese males to 16/100,000 in Queensland, Australia. 17.31 In tropical and subtropical Queensland, the incidence is five times that of Tasmania in the South. 16 Within Queensland a higher incidence is reported from the coastal areas than from the interior. 16.30 Similar geographic variations that could reflect gradients in intensity and duration of sun exposure have also been described in the