MEDICAL GRAND ROUNDS October 10, 1957

Protein Abnormalities in Multiple Myeloma and Allied States

This 59 year old colored male was admitted to

-57. He had had low back pain for 9 months, and 5 months before admission he had lower thoracic cage soreness and "rheumatic" aching of shoulders. Two months before admission he noted a "caving in" of his upper chest. Physical examination revealed cachexia, marked stiffness of spine, deep depression of the upper sternum, tenderness over ribs. with some crepitus over right anterior chest wall. Laboratory: Hgb - 8.1 gms%, WBC - 9.100 with normal differential, Urine showed 3+ albuminuria. Agglutii nations for Typh. H & 0, Brucella, Tularemia, Paratyphoid A & B, Proteus OX-19 were Negative, Tuberculin I/100 positive, BUN - 26. Total protein - 7 gms, Albumin - 4.3, Globulin - 2.7, Calcium - 11.7, Phosphorus - 6.0, Alkaline phosphotase - 5.5, Bence-Jones - Neg., Blood type - A, Antibodies - 1:128. Electrophoresis: abnormal protein in B-globulin zone. Bone marrow loaded with plasma cells. X-rays - disseminated osteolytic lesions. Treated with P-32 (5 millicuries).

Case #2 This 44 year old colored female was admitted —57. Two months prior to admission she began to have shortness of breath, palpitation, easy fatiguability. Physical examination revealed a temperature of 101.8°F., findings suggestive of congestive heart failure, and a liver edge palpable 4 finger breadths below the costal margin. Laboratory data: Hgb - 5.6 gm%, ESR - 30 mm/hr, WBC - 7,400 with normal differential, Urinalysis negative, Bence-Jones negative, Agglutinations for Typhoid, Brucella, Tularemia negative. BUN - 22; Total protein - 8.5 gm% with 3.0 gm% Albumin and 5.5 gm% Globulin. Calcium - 10.6 mgm%, Phorsphorus - 3.9 mgm%, Blood type-A. Electrophoresis showed abnormal B-globulin peak (62%). Bone marrow revealed plasma cytosis. Bone x-ray survey negative.

This 66 year old colored female was admitted to compared to compare the colored female was admitted to compared to compare the colored female was admitted to compared to compare the colored female was admitted to compared to compared the colored female was admitted to compared to compared the colored female was admitted to compared to compared female was admitted to compared to compared female was admitted female was admitted to compared female was admitted female was admitted female was admitted for compared female was admitted female was admitted female was admitted female was admitted for compared female was admitted for compared female was admitted female was admitted female was admitted female was admitted for compared female was admitted female was admitted female was admitted fema

BIBLIOGRAPHY

History

(I) Bence-Jones, H. On a new substance occurring in the urine of a patient with "mollities ossium". Phil. Tr. Royal Soc., London, 1848, p. 55.

A report of studies on an abnormal urinary protein first observed by

Drs. MacIntyre and Watson.

- (2) MacIntyre, W. Case of mollities and fragilitas ossium accompanied with urine strongly charged with animal matter. Med. Clin. Soc. Tr. 1850, 33, 211. Clinical description of first case.
- (3) Dalrymple, J. On the microscopic character of mollities ossium. Dublin Quart. J. M. Sci. 1846, 2, 85. Autopsy findings of first case. Clinical Description
- (4) Snapper, I., Turner, L.B. and Moscovitz, H.L. Multiple Myeloma, Greene and Strather, N.Y., 1953.

Chemical Considerations

(5) Multiple Myeloma. Current Clincal and Chemical Concepts. Am. J. Med., 1957, 23, 283.

An excellent review of chemical and immunological studies of myeloma and Bence-Jones proteins.

Lawson, H. A., Stuart, C.A., Paull, Wm., Phillips, Wm., and Phillips, R.N. Observations on the antibody content of the blood in patients with multiple myeloma. N.E.J.Med. 252, 13, 1955.

Demonstration of the paucity of antibodies in myeloma serum.

(7) Laurell, C.B., Laurell, H., and Waldenstrom, J. Glycoproteins in Serum from Patient with Myeloma, Macroglobulinemia and Related Conditions. Am. J. Med., 1957, 23, 24.

Allied States

(8) Azar, H.A., Hill, W.T., and Osserman, E.F. Malignant Lymphoma and Lymphatic Leukemia Associated with Myeloma-Type Serum Protein. Am. J. Med., 1957, 23, 239.

13 cases of lymphatic leukemia and malignant lymphoma, including one of Hodgkin's disease associated with myeloma-type proteins.

(9) Buffa, F. and Rappaport, H. Chronic Lymphatic Leukemia Associated with

- Dysproteinemia and Acquired Hemolytic anemia Am. J. Med., 1957, 23, 504. (10) Jencks, W.P., Smith, E.R.B., Durrum, E.L. The Clinical Significance of the Analysis of Serum Protein Distribution by Filter Paper Electrophoresis. Am. J.
- Med. 1956, 21, 387. An excellent review of electrophoretic abnormalities.
 (11) Henstell, H.H. and Feinstein, M. Interference of Abnormal Plasma Proteins with the Clotting Mechanism. Am. J. Med. 1957, 23, 381.
- (12) Stidworthy, G., Payne, R.W., Shetlar, C.L., and Shetlar, M.R. Objective evaluation of patient with Rheumatic Disease, II. Paper electrophoretic studies of serum glycoproteins and protein from patients with Rheumatoid Arthritis. J.C.1. 36, 309, 1957.
- (13) Weissman, N. and Perlmutter, M. An electrophoretic study of serum proteins in Thyroiditis. J.C.I. 1957, 36, 780.

Nonspecific elevated al, a2, and gamma globulin.

(14) Engle, R.L. and Wallis, L. Multiple Myeloma and the Adult Fanconi Syndrome 1. Am. J. Med. 1957, 23, 5.

Report of an adult with multiple myeloma and adult Fanconi syndrome. Defective tubular reabsorption may result from Myelomatosis.

Cryo and Macroglobulinemia

(15) MacKay, I.R., Eriksen, N., Motulsky, A.G., and Volwiler, W. Cryo- and Macroglobulinemia; Electrophoretic, Ultracentrifugal and Clinical Studies. Am. J. Med. 20, 564, 1956.

An excellent study of 5 patients with one or the other or both abnormalities. Analysis of literature on the subject.

(16) Wirtschafter, Z.T., Williams, D.W., and Gaulden, E.C. Cryoproteinemia: An Immunologic Phenomenon. Am. J. Med., 20, 624, 1956.

Describes the beneficial effects of adrenal cortical hormone in suppressing

the cryoglobulin and symptoms therefrom.

(17) Sehon, A.H., Gyenes, L., Gordon, L., Richter, M. and Rose, B. Phsyico-Chemical and Immunologic Studies on Macroglobulins. J.C.I. 1957, 36,456.

(18) Waldenstrom, J. Incipient Myelometose or "essential" hyperglobulinemia with fibrinogenopenia - a new syndrome." Acta Med. Scand., 117,216, 1944.

First description of syndrome due to macroglobulinemia

(19) Waldenstrom, J. "Abnormal proteins in myeloma" Adv. Internal Med.,

<u>5,398, 1952.</u>

(20) Lerner, A.B. and Watson, C.J. "Studies of Cryoglobulines I. Unusual purpura associated with the presence of a high concentration of cryoglobulin (cold precipitable serum globulin) Am. J. Med. Sci., <u>214</u>,410 (1947).

(21) Lerner, A.B., Barnum, C.P. and Watson, C.J. "Study of Cryoglobulin II. The Spontaneous precipitation of protein for Serum at 5°C in various disease States. Am. J. Med.Sci., <u>214</u>,416 (1947).