# DETERMINATION OF THE REASONABLE OR PERMISSIBLE MARGIN OF ERROR IN DISPENSING. V. LIQUIDS.* 

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## INTRODUCTION.

In the first paper of this series, it was stated that the different types of prescriptions which the pharmacist is ordinarily called upon to fill may be divided, roughly, into two groups, namely, liquids and solids. The four papers presented to date deal only with the more frequently encountered types of the latter, viz.: Powders and Capsules (1); Ointments (2); Suppositories (3); and Pills (4). This, the fifth paper of the series, deals with Liquids.

Liquids called for on prescriptions are usually measured, the volume in most cases, being determined by using either a cylindrical or conical graduate. The possibilities for error in measuring a definite volume of a liquid are greater in number than is commonly held. Fortunately, most of them may be ignored as the error involved is too small to be of practical significance. In fact, only three need be considered for the purpose of this study. They are believed to be, in the order of their importance: (1) the nature of the liquid to be measured, (2) the shape and size of the graduate used, and (3) the personal equation. To determine to what extent each of these factors is responsible for the total deviation from the standard, the studies reported in this paper were undertaken.

## EXPERIMENTAL PART.

For the purpose of the study reported in this paper, two series of tests were made. The object of the first was to determine the relationship, if any, between the size and shape of the graduate used and the magnitude of the observed error in the measurement of definite volumes. The object of the second was to determine to what extent the magnitude of error was effected by certain physical properties of liquids, such as color, viscosity, etc.

In the actual performance of these tests, the liquids were measured in both cylindrical and conical graduates by 100 members of the senior class in dispensing pharmacy at the School of Pharmacy of the University of Maryland. In each case the liquid was poured from a quart bottle into the graduate, held in the hand of the dispenser, then transferred to a prescription bottle. The contents of the bottle were again transferred to a tared container and accurately weighed on a chainomatic balance. The temperature of the liquids in both series of tests ranged from $22^{\circ} \mathrm{C}$. to $25^{\circ} \mathrm{C}$.

Each dispenser was assigned a definite number so the variation in the work of any individual could be followed throughout both series. The weight of all measurements is reported.

## SERIES I.

In the first series of tests definite volumes of distilled water were measured by the dispenser in $10-, 25-, 50$ - and $100-\mathrm{cc}$. cylindrical and conical graduates.

[^0]The results of the first series of tests are presented in Table I.
Table I.-Effect of Size and Shape of Graduate on Measurement of Definite Volumes of Ligutd.

| Dispen |  |  |  |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Numbe | Cyl. | Con. | yl. | Con. | Cyl. | Co | Cyl. | Con. |
| 1. | 9.579 | 9.555 | 24.653 | 24.510 | 49.457 | 48.285 | 98.283 | 98.052 |
| 2. | 9.647 | 9.860 | 24.615 | 23.957 | 49.801 | 48.653 | 98.048 | 99.242 |
| 3. | 9.855 | 9.718 | 24.812 | 24.474 | 49.593 | 48.890 | 98.439 | 98.917 |
| 4. | 9.653 | 9.410 | 23.622 | 25.231 | 49.417 | 47.836 | 96.715 | 96.637 |
| 5. | 9.528 | 9.523 | 24.550 | 24.219 | 50.208 | 46.385 | 99.313 | 98.319 |
| 6. | 9.729 | 9.798 | 24.742 | 24.354 | 49.659 | 46.385 | 98.902 | 97.343 |
| 7. | 9.343 | 9.948 | 24.961 | 24.856 | 49.835 | 48.689 | 98.002 | 101.368 |
| 8. | 9.373 | 9.529 | 23.444 | 24.745 | 49.140 | 47.379 | 97.815 | 93.970 |
| 9. | 9.691 | 10.004 | 24.995 | 24.501 | 49.789 | 50.147 | 99.046 | 100.590 |
| 10. | 9.227 | 9.182 | 24.287 | 22.919 | 48.841 | 47.253 | 97.673 | 98.600 |
| 11. | 9.718 | 9.778 | 24.716 | 23.710 | 50.022 | 46.766 | 98.745 | 95. 200 |
| 12. | 9.455 | 9.586 | 24.383 | 23.721 | 48.842 | 48.948 | 97.804 | 96.911 |
| 13. | 9.517 | 9. 584 | 25.626 | 24.328 | 50.023 | 49.023 | 98.705 | 96.561 |
| 14. | 9.700 | 9.459 | 24.704 | 22.910 | 49.400 | 48.177 | 98.942 | 93.812 |
| 15. | 9.401 | 9.965 | 24.170 | 24.462 | 50.081 | 49.300 | 98.773 | 98.995 |
| 16. | 9.717 | 9.632 | 24.336 | 23.848 | 48.964 | 48.553 | 9.8.478 | 99.201 |
| 17. | 9.487 | 9.620 | 24.340 | 23.339 | 48.787 | 47.335 | 97.637 | 100.009 |
| 18. | 9.674 | 9.792 | 24.535 | 24.669 | 49.484 | 50.331 | 98.548 | 99.264 |
| 19. | 9.660 | 9.533 | 24.829 | 25.110 | 49.333 | 49.842 | 99.523 | 100.424 |
| 20. | 9.561 | 9.944 | 24.170 | 25.135 | 50.170 | 50.036 | 98.753 | 99.740 |
| 21. | 9.475 | 9.768 | 23.844 | 24.729 | 49.047 | 49.475 | 98.534 | 98.856 |
| 22. | 9.130 | 9.484 | 24.500 | 24.481 | 49.312 | 49.553 | 97.866 | 96.409 |
| 23. | 9.617 | 9.838 | 24.765 | 24.610 | 50.024 | 49.784 | 98.741 | 98.443 |
| 24. | 9.617 | 9.951 | 24.905 | 24.604 | 49.131 | 49.362 | 98.400 | 98.777 |
| 25. | 9.844 | 9.600 | 24.177 | 23.843 | 49.152 | 47.778 | 97.963 | 97.034 |
| 26. | 8.985 | 8.748 | 24.418 | 23.986 | 49.015 | 47.681 | 98.212 | 94.765 |
| 27. | 9.620 | 10.000 | 24.935 | 23.584 | 47.962 | 47.351 | 98.311 | 95.958 |
| 28. | 9.231 | 9.145 | 24.060 | 23.642 | 47.910 | 48.739 | 97.456 | 95.683 |
| 29. | 9.515 | 9.664 | 24.830 | 23. 274 | 49.927 | 50.154 | 97.603 | 98.490 |
| 30. | 9.600 | 9.810 | 24.775 | 24.476 | 49.727 | 49.469 | 98.353 | 100.326 |
| 31. | 9.258 | 9.081 | 21.951 | 24.130 | 49.154 | 49.673 | 98.389 | 99.169 |
| 32. | 9.572 | 9.354 | 24.655 | 23.406 | 49.746 | 48.444 | 95.119 | 95.420 |
| 33. | 9.661 | 9.779 | 24.686 | 24.867 | 49.978 | 50.067 | 98.871 | 100.131 |
| 34. | 9.522 | 9.843 | 23.980 | 23.851 | 49.537 | 48.623 | 98.605 | 99.491 |
| 35. | 9.355 | 9.681 | 24.880 | 24.296 | 48.877 | 50.013 | 97.450 | 96.529 |
| 36. | 9. 284 | 9.377 | 24.556 | 24.252 | 48.741 | 48.459 | 98.537 | 96.729 |
| 37. | 9.636 | 9.417 | 24.302 | 23.104 | 49.365 | 47.971 | 100.021 | 97.117 |
| 38. | 9.654 | 9.568 | 24.445 | 24.829 | 50.472 | 49.367 | 98.440 | 98.608 |
| 39. | 9. 385 | 9.577 | 24.246 | 24.476 | 49.500 | 51.393 | 98.250 | 97.262 |
| 40. | 9.540 | 9.601 | 23.355 | 24.021 | 46.653 | 49.337 | 97.403 | 98.365 |
| 41. | 9.505 | 9.250 | 24.542 | 24.161 | 48.670 | 49.170 | 98.262 | 96.100 |
| 42. | 9.476 | 9.408 | 23.986 | 23.359 | 48.353 | 47.907 | 97.661 | 97.286 |
| 43. | 9.537 | 9.454 | 24.441 | 23.058 | 49.400 | 49.000 | 98.444 | 95.646 |
| 44. | 9.832 | 9.705 | 24.920 | 24.470 | 49.521 | 49.221 | 98.540 | 96.049 |
| 45. | 9.349 | 9.753 | 24.674 | 24. 247 | 49.751 | 49.911 | 98.623 | 98.312 |
| 46. | 9.337 | 9.337 | 24.500 | 21.000 | 49.298 | 45.857 | 97.865 | 98.800 |
| 47. | 9.208 | 9. 263 | 24.620 | 23.809 | 47.842 | 49.876 | 98.601 | 93.335 |
| 48. | 9.567 | 9.202 | 24.362 | 24.107 | 48.875 | 49. 205 | 97.904 | 96.937 |
| 49. | 9.588 | 9.571 | 24.304 | 23.119 | 49.344 | 49.308 | 98.341 | 94.980 |
| 50. | 9.235 | 9.772 | 22.684 | 25.102 | 49.928 | 49.272 | 97.600 | 96.069 |


| 51. | 9.604 | 9.004 | 24.352 | 22.985 | 49.055 | 47. 230 | 98.974 | 96.120 |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 52. | 9.515 | 10. 209 | 25.003 | 24.107 | 49.653 | 48.348 | 98. 210 | 99.056 |
| 53. | 9.476 | 9.606 | 24.448 | 23. 504 | 49.764 | 49.676 | 98.479 | 98.191 |
| 54. | 9.510 | 9.842 | 24.608 | 24.445 | 49.744 | 49.228 | 98.461 | 97.565 |
| 55. | 9.817 | 10.075 | 25.416 | 23.201 | 49.194 | 49.006 | 97.981 | 97.134 |
| 56. | 9.601 | 9.348 | 24.376 | 23.210 | 49.341 | 48. 258 | 98.003 | 99.001 |
| 57. | 9.556 | 9.406 | 24.431 | 23.613 | 48.353 | 47.633 | 98. 242 | 99.705 |
| 58. | 8.943 | 9.468 | 24.347 | 24.782 | 49.971 | 47.843 | 97.970 | 98.831 |
| 59. | 9.638 | 9.550 | 24.391 | 23.562 | 47.342 | 48.668 | 98.003 | 96.674 |
| 60. | 9.660 | 10.301 | 24.239 | 23.504 | 49.396 | 47.962 | 98.966 | 97.748 |
| 61. | 9.949 | 9.671 | 23.998 | 24.200 | 47.902 | 49.941 | 98.827 | 97.990 |
| 62. | 9.777 | 9.864 | 24.539 | 23.121 | 50.155 | 48. 261 | 97.050 | 97.957 |
| 63. | 9.682 | 9.038 | 24.952 | 24.452 | 50.065 | 49.724 | 98.937 | 100.421 |
| 64. | 9.452 | 9.262 | 23.641 | 23.490 | 49.078 | 48.531 | 97.730 | 100.348 |
| 65. | 9.708 | 9.702 | 24.601 | 23.556 | 49.762 | 49.783 | 98.551 | 95.742 |
| 66. | 9.678 | 9.910 | 24.640 | 24.228 | 48.741 | 49.190 | 98.320 | 99.311 |
| 67. | $\bigcirc .523$ | 9.790 | 24.503 | 23.727 | 49.622 | 50.169 | 98.882 | 98.009 |
| 68. | 9.501 | 9.164 | 24.107 | 22.746 | 49.104 | 47.386 | 98.816 | 98.612 |
| 69. | 9.938 | 9.726 | 24.955 | 24.945 | 49.940 | 50.860 | 99.830 | 97.723 |
| 70. | 9.573 | 9.587 | 24.872 | 24.900 | 49.590 | 48.976 | 97.631 | 96.867 |
| 71. | 9.912 | 9.716 | 24.182 | 22.911 | 47.654 | 49.829 | 98.506 | 99.004 |
| 72. | 9.721 | 9.543 | 24.748 | 22.733 | 47.820 | 46.303 | 98. 209 | 94.402 |
| 73. | 9.357 | 9.543 | 23.126 | 24. 283 | 47.124 | 49.100 | 94.564 | 98.696 |
| 74. | 9.453 | 9.712 | 24.389 | 21.795 | 49.281 | 48.472 | 98.872 | 97.838 |
| 75. | 9.588 | 9.739 | 24.941 | 24.481 | 49.502 | 49.777 | 98.768 | 97.594 |
| 76. | 9.485 | 9.229 | 24.633 | 23. 268 | 49.239 | 50.036 | 98.002 | 98.753 |
| 77. | 9.483 | 10.035 | 24.913 | 24.076 | 49.046 | 48.921 | 98.470 | 99.126 |
| 78. | 9.732 | 9.451 | 24.105 | 24.929 | 49.835 | 46.891 | 98.422 | 96.770 |
| 79. | 9.462 | 9.383 | 24.314 | 23.709 | 49.876 | 49.934 | 98.104 | 98.163 |
| 80. | 9.589 | 9.678 | 24.483 | 23.771 | 49.186 | 49.260 | 98.583 | 96.768 |
| 81. | 9.733 | 10.028 | 25.007 | 23.928 | 49.644 | 48.386 | 99.172 | 97.804 |
| 82. | 9.740 | 9.681 | 24.510 | 23.674 | 48.765 | 50.984 | 98.644 | 99.433 |
| 83. | 9.667 | 9.760 | 24.590 | 24.184 | 49.776 | 47.566 | 98. 238 | 95.723 |
| 84. | 9.629 | 9.630 | 24.710 | 24.740 | 49.386 | 49.937 | 98.636 | 100.388 |
| 85. | 9.819 | 10.151 | 24.814 | 24.281 | 49.653 | 48.447 | 98.907 | 100. 233 |
| 86. | 9.637 | 9.469 | 24.933 | 24.912 | 49.763 | 48.885 | 96.876 | 99.423 |
| 87. | 9.369 | 9.365 | 24.799 | 24.043 | 48.065 | 50.226 | 97.986 | 98.149 |
| 88. | 9.637 | 9.722 | 24.988 | 24.341 | 48.382 | 47.635 | 97.380 | 97.932 |
| 89. | 9.466 | 9.691 | 24.905 | 24.307 | 48.363 | 45. 516 | 98.780 | 101.177 |
| 90. | 9.506 | 9.512 | 24. 292 | 24.178 | 49.374 | 49.783 | 98.389 | 99.905 |
| 91. | 9.650 | 9.108 | 24.777 | 23.876 | 48.350 | 47.031 | 98.704 | 94. 353 |
| 92. | 9.991 | 9.104 | 24.825 | 24.202 | 49.601 | 47.561 | 96.550 | 99.961 |
| 93. | 9.596 | 10.228 | 24.669 | 24.525 | 49.128 | 50.146 | 98.722 | 99.547 |
| 94. | 9.623 | 9.349 | 24.763 | 24.892 | 49.104 | 49.582 | 98.315 | 97.081 |
| 95. | 10.651 | 9.876 | 24.339 | 23.653 | 50.102 | 46.1783 | 97.014 | 101.900 |
| 96. | 9.484 | 9.806 | 24.341 | 24.256 | 50.653 | 47.343 | 98.333 | 99.502 |
| 97. | 9.292 | 9.402 | 24.349 | 23.406 | 49.876 | 51.437 | 98.136 | 96.332 |
| 98. | 9.154 | 9.135 | 24.687 | 23.778 | 49.783 | 48.106 | 98.268 | 98.030 |
| 99. | 9.724 | 9.630 | 24.678 | 24.040 | 49.386 | 48.462 | 98.336 | 96.668 |
| 100. | 9.203 | 9.973 | 25.067 | 24.103 | 49.336 | 47.286 | 97.643 | 94.192 |
| Av. Wt. | 9.561 | 9.606 | 24.481 | 23.994 | 49.258 | 48.758 | 98.240 | 97.881 |
| S. D.* | 0.224 | 0.290 | 0.503 | 0.708 | 0.727 | 1.203 | 0.767 | 1.802 |
| $\% \mathrm{D}^{1}$ | 2.34\% | 3.11\% | 2.05\% | 2.95\% | 1.48\% | 2.47\% | 0.78\% | 1.84\% |

[^1]The tabulated data given in Table I shows there is an error due to the size and shape of the graduate. The magnitude of error is considerably greater when a conical graduate is used than when a cylindrical graduate is used. With respect to size just the opposite is true, the larger the graduate and the larger the volume measured the smaller the per cent of error. For instance in the measurement of 10 cc. of distilled water in a cylindrical graduate the average error based on the standard deviation amounted to $2.34 \%$. When a $10-\mathrm{cc}$. conical graduate was used it amounted to $3.11 \%$. That the magnitude of the error is greater when the smaller graduate was used than when the larger graduate was used is shown when the volume measured was 100 cc ., and a $100-\mathrm{cc}$. graduate was used in making the measurements, the average error amounted to only $0.78 \%$ in the case of the cylindrical graduate and $1.84 \%$ in the case of the conical graduate.

The error due to the personal equation is naturally indefinite. In fact, it was found to be impractical to attempt to measure it separately. The error is revealed by a definite trend in a series of measurements made by an individual rather than by the observation made on a single measurement. For example, in the series of measurements made by dispenser number 1 it will be observed that a majority of the measurements here were high when compared with the average, while those of dispenser number 10 were below the average.

No effort was made to determine what the personal equation was in these cases so that an accurate statement cannot be made concerning its nature. In some cases it may have been the result of defective vision, in others it may have been due to natural carelessness, in other instances it may have been due to using the upper meniscus at times and at other times the lower meniscus for making measurements, or to some other trait of the individual making the measurements.

The results presented in Table I are summarized in Table II, which follows. This table shows the actual number of measurements falling within the standard deviation and multiples thereof. Furthermore, the table shows the per cent of measurements falling within any one group, since the total number of measurements made in each case was exactly 100 .

| Liquid to Be Measured. | Table II.-Summary of Resulis Presented in Table I. |  |  |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | Volume | Shape of | Average | Standard Number of Measurements Falling within |  |  |  |  |
|  | Measured in Cc . | Graduate Used. | Weight in Gm . | Deviation in Gm. | $\begin{array}{cc} 1 \times \\ \text { S. } & \times \\ \hline \end{array}$ | $\begin{gathered} 2 \times \\ \text { S. } \\ \hline \end{gathered}$ | $\begin{gathered} 3 \times \\ \text { S. } \\ \hline \end{gathered}$ | $4 \times S .0$ or Over. |
| Distilled Water | 10 | Cyl. | 9.561 | 0.224 | 78 | 19 | 2 | 1 |
| Distilled Water | 10 | Con. | 9.606 | 0.290 | 70 | 25 | 5 | 0 |
| Distilled Water | 25 | Cyl. | 24.481 | $0.50: 3$ | 85 | 9 | 4 | 2 |
| Distilled Water | 25 | Con. | 23.994 | 0.708 | 68 | 30 | 0 | 2 |
| Distilled Water | 50 | Cyl. | 49.258 | 0.727 | 74 | 22 | 3 | 1 |
| Distilled Water | 50 | Con. | 48.758 | 1. 203 | 69 | 26 | 5 | 0 |
| Distilled Water | 100 | Cyl. | 98.240 | 0.767 | 83 | 12 | 3 | 2 |
| Distilled Water | 100 | Con. | 97.881 | 1.802 | 67 | 28 | 5 | 0 |

The second series of tests was conducted to determine the effect of certain physical properties of liquids on the measurement of a definite volume. With this object in view the following liquids were selected:
(1) Elixir of Iron, Quinine and Strychnine, N. F., as a green colored liquid; (2) Syrup, U. S. P., as a viscous clear liquid; (3) Milk of Magnesia, U. S. P., as an opaque liquid; and (4) Castor Oil, U. S. P., as an oily liquid.

To obtain comparative data, the same cylindrical and conical graduates used in measuring the 100 cc . of Distilled Water in the first series of tests, were used in this series. The results of the second series of tests are given in Table III.

Table III.-Effect of Certain Physical Properties of Liquids on the Magnitude of Error.

| $\begin{aligned} & \text { Elixer I. Q.S. } \\ & \text { Cyl. } \end{aligned}$ |  | Syrup. |  | Milk of Magnesia. |  | $\begin{aligned} & \text { Castor Oil. } \\ & \text { Cyl. } \end{aligned}$ |  | Dispenser Num ber. |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 103.912 | 105.721 | 126. 239 | 130.634 | 94.277 | 95.533 | 81.218 | 94.358 | 1. |
| 104.616 | 105.169 | 125.316 | 128.371 | 101.501 | 93.585 | 90.491 | 87. 204 | , |
| 104.352 | 103.355 | 118.564 | 129.030 | 93.607 | 92.510 | 89.362 | 84.856 | 3. |
| 104.251 | 104.403 | 123.108 | 126.035 | 94.705 | 101.491 | 89.300 | 82.680 | 4. |
| 104.318 | 106.825 | 125.292 | 126.832 | 97.774 | 99. 214 | 85.930 | 90.081 | 5 |
| 103.256 | 101.860 | 122.514 | 129.866 | 94.000 | 93.933 | 84.333 | 84.352 | 6. |
| 104.318 | 100.819 | 126.471 | 122.857 | 98.027 | 89.337 | 86. 222 | 93.861 | 7. |
| 101.712 | 101.838 | 124.300 | 124.772 | 92.631 | 88.506 | 89.515 | 85.641 | 8. |
| 104.543 | 103.834 | 125.084 | 129.579 | 93.107 | 99.102 | 89.801 | 87.405 | 9. |
| 103.953 | 100.100 | 126.656 | 120.159 | 93.239 | 88.941 | 89.086 | 80.474 | 10. |
| 104.229 | 101. 246 | 124.049 | 123.073 | 95. 297 | 100.287 | 92.100 | 87.673 | 11. |
| 104.054 | 102.125 | 125.450 | 122. 511 | 90.924 | 90.785 | 91.311 | 84.130 | 12. |
| 104.375 | 105. 640 | 127.315 | 128.417 | 99.762 | 100.747 | 86.416 | 87.381 | 13. |
| 103.088 | 100.714 | 125.833 | 127.586 | 92.725 | 93.730 | 84.803 | 86.960 | 14. |
| 104.122 | 103.030 | 124.522 | 130.190 | 91.873 | 93. 310 | 88.875 | 82.388 | 15. |
| 104.451 | 105.937 | 125.141 | 130.632 | 91.260 | 92.836 | 91. 340 | 92. 253 | 16. |
| 103.743 | 102.488 | 123.965 | 123.376 | 93.453 | 94.687 | 87.587 | 90.040 | 17. |
| 104.442 | 104.002 | 121. 520 | 126.646 | 97.266 | 96. 361 | 86.889 | 91.420 | 18. |
| 104.882 | 107.639 | 124.420 | 127.288 | 99.424 | 94.666 | 85.829 | 86. 264 | 19. |
| 104.455 | 107.165 | 124.510 | 133.900 | 93.595 | 101.015 | 90.888 | 88. 206 | 20. |
| 103. 505 | 106.843 | 121.258 | 123.896 | 92.909 | 91.051 | 89.712 | 89.914 | 21. |
| 104.383 | 103. 567 | 127.533 | 129.635 | 95.747 | 96.066 | 91.535 | 90.416 | 22. |
| 104.417 | 107.950 | 128.727 | 132.664 | 98. 576 | 96.087 | 89.151 | 89.097 | 23. |
| 103.758 | 103.323 | 124.188 | 132.217 | 92.377 | 93.932 | 95.985 | 97.780 | 24. |
| 103.455 | 98.961 | 125. 296 | 121.748 | 94.776 | 94.764 | 91.856 | 89.042 | 25. |
| 101.128 | 103.885 | 124.703 | 128.611 | 92.271 | 96.410 | 81.262 | 84 | 26. |
| 104.200 | 105.251 | 124.510 | 129.791 | 93.438 | 97.810 | 80.374 | 89.891 | 27. |
| 103.290 | 101.965 | 125.746 | 123.473 | 93.091 | 91.271 | 87.200 | 85.853 | 28. |
| 102.850 | 103.075 | 124.602 | 125. 312 | 94.425 | 95.168 | 83.950 | 82.057 | 29. |
| 104.991 | 103. 581 | 122.672 | 132.161 | 98.744 | 100.002 | 84.442 | 85.214 | 30. |
| 102.492 | 104.034 | 120.040 | 125. 252 | 92.776 | 91.035 | 85.150 | 83.727 | 31. |
| 103.902 | 102.730 | 123.021 | 125.508 | 90.431 | 93.059 | 82.224 | 79. 224 | 32. |
| 104.053 | 101.387 | 127.582 | 134.551 | 93.784 | 93.823 | 88.900 | 88.389 | 33. |
| 104.392 | 102.056 | 127.613 | 126.704 | 97.487 | 98.066 | 93.470 | 84.637 | , |
| 103.792 | 103. 585 | 124.763 | 125.242 | 93.981 | 91.906 | 87.990 | 88.673 | 35. |
| 104.267 | 108.794 | 123.323 | 127.172 | 93.381 | 92.345 | 77.756 | 78.931 | 36. |
| 103.580 | 103.600 | 122.779 | 124.210 | 92. 234 | 96.937 | 78.873 | 74.778 | 37. |
| 106.219 | 103.783 | 124.039 | 129.918 | 94.546 | 98.823 | 82.670 | 84.768 | 38. |
| 104.002 | 104.135 | 121.910 | 127.921 | 96. 919 | 93.900 | 81.307 | 86. 300 | 39. |
| 104.379 | 105.932 | 123.700 | 125.678 | 94.222 | 98.302 | 91.318 | 81.351 | 40. |
| 103.973 | 104.077 | 120.211 | 120.660 | 95.040 | 90.997 | 90.588 | 83.593 | 1 |
| 103.472 | 104.082 | 124.862 | 123.270 | 97.088 | 92.449 | 78.413 | 99.688 | 42. |
| 103.874 | 98.793 | 123.566 | 126.066 | 94.700 | 94.738 | 87.579 | 87.730 | 43. |
| 104.048 | 103.592 | 124.363 | 128.529 | 92. 859 | 93.882 | 88.885 | 81.158 | 44. |
| 103.728 | 103.366 | 123.234 | 127.907 | 93.537 | 100.837 | 83.584 | 82. 365 | 45. |
| 103.920 | 102.187 | 121.090 | 122.183 | 92.327 | 90.486 | 86.873 | 83. 277 | 746. |
| 104.056 | 102.192 | 127.164 | 125.773 | 91. 200 | 91.680 | 88.609 | 83.551 | 47. |
| 103.881 | 105.031 | 124.634 | 127.411 | 93.643 | 94.768 | 87.549 | 79.439 | 48. |
| 101.942 | 105.649 | 123.814 | 127.059 | 94. 200 | 94.830 | 86.533 | 81.672 | 49。 |
| 100.130 | 104.144 | 125.064 | 125.902 | 97.102 | 91.874 | 87.589 | 97.041 | 50. |


| 104.220 | 100.849 | 127.508 |  |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 104.221 | 105.980 | 126.184 | 131.553 | 94.949 | 96. 272 | 88.733 | 86.821 | 2. |
| 104.251 | 98.261 | 125.672 | 134.053 | 96.126 | 99.5 | 87.221 | 94,999 | 53. |
| 105. 263 | 106.625 | 123.988 | 127.516 | 101. 291 | 93. 27 | 88.73 | 95.194 |  |
| 104. 257 | 103.140 | 120.802 | 123.582 | 96.449 | 95.500 | 86.8 | 86. | 5. |
| 104.013 | 106. 221 | 121.343 | 125.064 | 94. 508 | 97.65 | 88.382 | 80.959 | 56. |
| 106.003 | 104.612 | 122.952 | 129.540 | 89. 383 | 94.97 | 87.486 | 91.723 | 57. |
| 104. 583 | 104.248 | 127.944 | 125.593 | 98.427 | 96. 22 | 90.49 | 88.45 | 58. |
| 103.788 | 107.811 | 125. 285 | 125.868 | 92.978 | 94.71 | 94.942 | 89.89 | 59 |
| 102.384 | 104,025 | 117.325 | 124.333 | 91.980 | 94.233 | 84.36 | 87. 27 | 60 |
| 102. 681 | 99.258 | 123.321 | 125.324 | 88.832 | 89.121 | 91.836 | 84.65 | 61 |
| 103.377 | 107.395 | 126.056 | 132.802 | 94. 200 | 97.115 | 87.321 | 89. 336 | 62. |
| 106.615 | 104.880 | 126. 326 | 130.639 | 95. 236 | 94.993 | 87.749 | 93.265 | 63. |
| 105.527 | 104.623 | 120.935 | 125.761 | 100. 304 | 94.160 | 83.647 | 88.772 | 64 |
| 103.265 | 101.985 | 122.343 | 122.693 | 96. 382 | 99.416 | 89. 280 | 85.191 | 65. |
| 103.527 | 99.563 | 124.384 | 122.876 | 91.104 | 96.403 | 91.630 | 84.180 | 66. |
| 104.051 | 106.786 | 124.678 | 130.639 | 94. 507 | 93.300 | 85. 676 | 81.684 | 67. |
| 105.357 | 106.001 | 125.565 | 123.774 | 99.909 | 94.948 | 96.776 | 87.603 | 68. |
| 103.902 | 103.663 | 124.621 | 126.795 | 93.430 | 93.915 | 85. 690 | 87.116 | 69 |
| 104.77 | 105.370 | 125.859 | 129. 509 | 99.742 | 95.432 | 83. 529 | 84.965 | 70 |
| 103.690 | 102.844 | 125.231 | 122. 525 | 95.461 | 94.111 | 87.325 | 79.603 | 71 |
| 103.672 | 100.578 | 118.993 | 121.108 | 93.729 | 90.062 | 96.146 | 97.798 | 72 |
| 104.872 | 104.530 | 124.484 | 125.028 | 93.606 | 96.050 | 84.382 | 88.743 | 73 |
| 105.477 | 103.923 | 123.120 | 123. 579 | 93.428 | 94.127 | 87.175 | 86. 262 | 74. |
| 104.452 | 104.038 | 122.561 | 128.706 | 96.711 | 96.045 | 84.589 | 91.220 | 75. |
| 104.005 | 105.483 | 126.936 | 122. 363 | 95.369 | 98.710 | 91.365 | 94.323 | 76 |
| 103.714 | 104.935 | 125.758 | 133.774 | 94.805 | 94.350 | 92.78 | 81.604 | 7 |
| 104.162 | 105.057 | 125.101 | 127.424 | 98.728 | 94.779 | 86.487 | 89.463 | 78. |
| 103.610 | 101.683 | 126.802 | 123.945 | 93.347 | 97.996 | 86.170 | 81.147 | 79 |
| 103.334 | 101.810 | 126.740 | 123.543 | 95. 250 | 95.584 | 91.765 | 88.502 | 80 |
| 104.53 | 104.403 | 124.763 | 129.183 | 98.730 | 94.618 | 96.881 | 81.361 | 81 |
| 105.631 | 103.457 | 120.999 | 125.440 | 95.672 | 97.33 | 88.461 | 87.732 | 82 |
| 104.72 | 100.157 | 123. 364 | 127.942 | 95.661 | 92.30 | 85.395 | 95.156 | 83 |
| 105.006 | 104.987 | 127.376 | 129.519 | 96.100 | 94.58 | 95.601 | 82.397 | 84 |
| 103.679 | 105. 343 | 124.122 | 124.110 | 100.375 | 101.652 | 88.649 | 88.760 | 85. |
| 105.104 | 106.424 | 123.428 | 124.300 | 94.899 | 99.031 | 85.031 | 87.515 | 86 |
| 105.643 | 108.707 | 121.417 | 127.986 | 93.390 | 98.470 | 88.463 | 74.383 | 87. |
| 104. 325 | 100.955 | 122.765 | 125.045 | 95.58 | 94.613 | 87.637 | 91.687 | 88. |
| 103. 762 | 100.040 | 125. 502 | 127.800 | 95. 347 | 92.32 | 88.4 | 89.579 | 89. |
| 103.624 | 105.79 | 118. 640 | 121.091 | 93.493 | 97.00 | 85.36 | 84.652 | . |
| 104.13 | 106.505 | 124.335 | 130.632 | 94. 240 | 101.964 | 88,912 | 89.571 | 91. |
| 104.8 | 105.122 | 126. 255 | 129.312 | 96. 400 | 101. 782 | 87.25 | 90.784 | 92. |
| 103. 27 | 103.505 | 123.035 | 129.669 | 96.662 | 100.734 | 91.3 | 96.646 | 93. |
| 104,018 | 106.061 | 128.643 | 126.38 | 97.483 | 95.60 | 88.1 | 99.648 | 94. |
| 104,051 | 99.018 | 125.340 | 128.006 | 94.30 | 94.32 | 85. 337 | 87.934 | 5. |
| 104.688 | 104.350 | 129.288 | 125,665 | 95.670 | 99, 775 | 81.565 | 84.572 | 96. |
| 103.672 | 104.531 | 129.780 | 129.822 | 92.445 | 91.839 | 86.861 | 75.121 | 97. |
| 103. 676 | 98.521 | 122.793 | 120.471 | 90. 347 | 90.370 | 80.845 | 85.813 | 98. |
| 103.395 | 103.320 | 122.176 | 127.747 | 93. 52' | 95.846 | 94.033 | 86.527 | 9. |
| 104.783 | 107.346 | 120.760 | 121. 229 | 100.674 | 95. 201 | 86.130 | 86.941 | 100 |
| 104.049 | 103.752 | 124.298 | 126.75 | 94.895 | 95.254 | 87.742 | 87.017 |  |
| 0.9 | 2. | 2 | .3 | 2. | 3.15 | 3.938 | 5.087 |  |
| . $89 \%$ | 2.24\% | .91 | 2.63 | 2.79 | .31\% | $4.49 \%$ | 5.85 |  |

[^2]The foregoing tabulation shows that errors are made in measurement and therefore the deviation from the standard is affected by the nature of the liquid measured. The magnitude of the error observed was in the following order: Distilled Water, Elixir of Iron, Quinine and Strychnine, Syrup, Milk of Magnesia and Castor Oil. The physical properties responsible for the great part were found to be color and viscosity.

The data presented in Table III reveal that color in a liquid has a tendency to increase the magnitude in error made in measurement, for example: the average error for 100 cc . of Distilled Water measured in a cylindrical graduate is $0.78 \%$, while that of the green-colored liquid, Elixir of Iron, Quinine and Strychnine is $0.89 \%$.

A similar effect was observed with respect to viscosity. In the case of Distilled Water the average error was $0.78 \%$ as previously stated, whereas the average error found in the measurement of Syrup was $1.91 \%$, and for Castor Oil was $4.49 \%$.

The large error in the case of Castor Oil is no doubt due to the fact that the refractive index of Castor Oil is so near that of glass that the adherence of the oil to the sides of the graduate is not detected and not sufficient time is allowed by the dispenser for complete drainage.

Milk of Magnesia while not a liquid in the true sense of the word is nevertheless generally dispensed by volume rather than by weight, hence it must be measured. It was therefore included in this series of experiments. The comparatively large error found in this instance was no doubt due to the adherence of a considerable amount of the magnesium hydroxide to the inside of the glass graduate from which it was impossible to drain it, but could be readily seen. The average error amounted to $2.79 \%$ as compared with that of Distilled Water which was $0.78 \%$.

The results of Table III are best summarized in Table IV.

Table IV.-Summary of Results Presented in Table III.

| Liquid to Be Measured. | Volume Measured in Cc. | Shape of Graduate Used. | Average Weight in $\mathbf{G m}$. | Standard Deviation in Gm . | Number of Measurements Falling within |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  |  |  |  | $\begin{aligned} & 1 \times \\ & \text { s. } \times \mathbf{D} . \end{aligned}$ | $\begin{gathered} 2 \times \\ \text { s. } \\ \text { D. } \end{gathered}$ | S. ${ }^{3} \times$ |  |
| Distilled Water | 100 | Cyl. | 98.240 | 0.767 | 83 | 12 | 3 | 2 |
| Elix. I. Q. \& S. | 100 | Cyl. | 104.049 | 0.926 | 79 | 14 | 5 | 2 |
| Syrup | 100 | Cyl. | 124.298 | 2.368 | 69 | 25 | 6 | 0 |
| Milk of Magnesia | 100 | Cyl. | 94.895 | 2.644 | 73 | 20 | 7 | 0 |
| Castor Oil | 100 | Cyl. | 87.742 | 3.938 | 72 | 21 | 7 | 0 |
| Distilled Water | 100 | Con. | 97.881 | 1.802 | 67 | 28 | 5 | 0 |
| Elix. I. Q. \& S. | 100 | Con. | 103.752 | 2.329 | 70 | 25 | 5 | 0 |
| Syrup | 100 | Con. | 126.753 | 3.333 | 68 | 28 | 4 | 0 |
| Milk of Magnesia | 100 | Con. | 95.254 | 3.151 | 65 | 30 | 5 | 0 |
| Castor Oil | 100 | Con. | 87.017 | 5.087 | 70 | 24 | 6 | 0 |

To make it possible to compare the results given in Tables I and III with similar data that may have been published, but which have not been expressed in terms of the standard deviation, the per cent deviation from the average weight has been calculated and is given in Table V.

Table V.-Percentage of Error Computed from Data in Tables I and III.

| Liquid to Be Measured. | Volume Measured in Ce. | Shape uate Used. | Average Weight in Gm. | Number of Measurements Falling within $1 \% .2 \% .3 \% .4 \%$. $5 \% .6 \%$. $7 \% .8 \% .9 \%$. |  |  |  |  |  |  |  |  | $\begin{aligned} & 10 \% \\ & \text { More } \end{aligned}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Distilled Water | 10 | Cyl. | 9.561 | 4 | 5 | 11 | 24 | 22 | 13 | 9 | 8 | 2 | 2 |
| Distilled Water | 10 | Con. | 9.606 | 11 | 9 | 19 | 14 | 14 | 11 | 7 | 5 | 6 | 4 |
| Distilled Water | 25 | Cyl. | 24.481 | 27 | 29 | 24 | 9 | 4 | 2 | 2 | 1 | 0 | 2 |
| Distilled Water | 25 | Con. | 23.994 | 13 | 9 | 18 | 16 | 10 | 13 | 7 | 6 | 4 | 4 |
| Distilled Water | 50 | Cyl. | 49.258 | 55 | 33 | 8 | 2 | 1 | 1 |  |  |  |  |
| Distilled Water | 50 | Con. | 48.758 | 29 | 23 | 30 | 7 | 6 | 4 | 1 |  |  |  |
| Distilled Water | 100 | Cyl. | 98.240 | 89 | 9 | 0 | 2 |  |  |  |  |  |  |
| Distilled Water | 100 | Con. | 97.881 | 38 | 34 | 18 | 7 | 3 |  |  |  |  |  |
| Elix. I. Q. \& S. | 100 | Cyl. | 104.049 | 82 | 12 | 5 | 1 |  |  |  |  |  |  |
| Elix. I. Q. \& S. | 100 | Con. | 103.752 | 38 | 26 | 18 | 9 | 7 | 2 |  |  |  |  |
| Syrup | 100 | Cyl. | 124.298 | 45 | 27 | 18 | 4 | 5 | 1 |  |  |  |  |
| Syrup | 100 | Con. | 126.753 | 30 | 18 | 24 | 14 | 9 | 4 | 1 |  |  |  |
| Milk of Magnesia | 100 | Cyl. | 94.895 | 30 | 26 | 18 | 9 | 7 | 6 | 4 |  |  |  |
| Milk of Magnesia | 100 | Con. | 95.254 | 29 | 20 | 12 | 12 | 11 | 7 | 7 | 2 |  |  |
| Castor Oil | 100 | Cyl. | 87.742 | 22 | 20 | 11 | 11 | 15 | 3 | 2 | 6 | 3 | 7 |
| Castor Oil | 100 | Con. | 87.017 | 19 | 10 | 17 | 13 | 4 | 7 | 9 | 3 | 5 | 13 |

Note: All percentages are calculated from the average weight. $1 \%=1 \%$ or less; $2 \%=$ from $1 \%$ plus to $2 \%$; etc.

For the purpose of comparison with previously published data it is also desirable to have information showing the percentage of the total measurements in which the error falls below certain magnitudes, the latter being expressed in terms of per cent. The following table is intended to accomplish this purpose.

Table VI.-Table Showing the Maximum Per Cent of Error in $90 \%$ of the Measurements Recorded in Table V.

| Shape of Graduate. | 10 Cc . | $\begin{aligned} & \text { Distilled } \\ & 25 \mathrm{Cc} . \end{aligned}$ | Water. <br> 50 Cc . | 100 Cc . | Elix. <br> I. $Q$. <br> 100 Cc . | $\underset{100}{\text { Syrup }}$ | Milk of Magnesia 100 Cc . | $\begin{gathered} \text { Castor Oil } \\ 100 \mathrm{Cc} . \end{gathered}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Cyl. | 8\% | $5 \%$ | $3 \%$ | 2\% | $2 \%$ | $3 \%$ | $5 \%$ | 8\% |
| Con. | 8\% | 8\% | 5\% | $3 \%$ | $4 \%$ | 5\% | 6\% | 10\% |

## CONCLUSIONS.

1. The factors largely responsible for the errors made by pharmacists in the measurement of specified volumes were found to be three in number. These are in the order of their importance: (1) The nature of the liquid to be measured, (2) the shape and size of the graduate used, and (3) the personal equation.

The error due to the personal equation naturally cannot be predicted with any degree of accuracy as it depends entirely upon the idiosyncrasy of the individuals making the measurements. In some instances it may far exceed one or both of the other two factors but in all of the measurements made in the foregoing series of tests it exceeded twice the standard deviation in less than 7 per cent of all cases.
2. From the data obtained in the tests made it would seem that twice the standard deviation is a reasonable margin of error for the measurement of the volume of liquids. A margin of this magnitude will permit the acceptance of the following:

| Shape of Graduate. | 10 Cc . | ${ }_{25}$ Distilled 25 Cc . | Water. <br> 50 Cc . | 100 Cc . |  | $\begin{aligned} & \text { Syrup } \\ & 100 \text { Cc. } \end{aligned}$ | Milk of Magnesia 100 Cc . | $\begin{aligned} & \text { Castor Oii } \\ & 100 \mathrm{Cc} . \end{aligned}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Cyl. | $97 \%$ | 94\% | 96\% | 95\% | 93\% | 94\% | 93\% | 93\% |
| Con. | 95\% | 98\% | 95\% | 95\% | 95\% | 96\% | 95\% | 94\% |
| REFERENCES. |  |  |  |  |  |  |  |  |
| (1) Andrews, Marvin J., Jour. A. Ph. A., 22, 755 and 838 (1933). |  |  |  |  |  |  |  |  |
| (2) Andrews, Marvin J., Ibid., 23, 350 and 421 (1934). |  |  |  |  |  |  |  |  |
| (3) Andrews, Marvin J., Ibid., 23, 1003 (1934). |  |  |  |  |  |  |  |  |
| (4) Andrews, Marvin J., Ibid., 23, 1117 and 1210 (1934). |  |  |  |  |  |  |  |  |

## UNITED STATES PATENTS GRANTED FOR MEDICINES DURING THE PIONEER YEARS OF THE PATENT OFFICE.*

## BY LYMAN F. KEBLER. ${ }^{1}$

The word "patent" means open, not secret. A patent cannot be granted for a medicine of secret composition. The term "Patent Medicine" applied to a medicine of secret composition is a misnomer. The term in general conveys an erroneous impression. Many think that all proprietaries, foods and drugs, and medicines in package form, are of secret composition and consequently plain frauds. It is true that some outright medical frauds have been and still are perpetrated on the suffering sick. And this is true even in the case of a goodly number of medicines for which patents have been granted. Patenting a product does not preclude telling fairy tales about it. In fact, the therapeutic claims contained in the description of some of the patents for medicines are grossly false and fraudulent, as will be pointed out later.

## SOME INTERESTING PHASES IN PATENTING MEDICINES.

Secret medicines with their air of mysticism have held sway for the ages in all lands. The alchemistic era produced some of the most phony ideas in the matter of the philosopher's stone being a universal medicine and panacea. Secret medicines made marked advances during the time of iatrochemistry, when Paracelsus (14931541), with his lapis infernales, held sway, and John R. Glauber (1610-1770), the distinguished physician-chemist and discoverer of Glauber's salt, played such prominent parts. Glauber not only discovered the salt named after him but ascertained its medicinal virtues and sold it at a handsome profit under the name sal mirabile, for many years. It is claimed that he made a living selling secret medicines.

England set a precedent in granting patents to medicines. Among the earliest may be mentioned "Dr. Bateman's Pectoral Drops" (1726), "Dr. James' Fever Powder" (1747), "Ann Pike's Ointment for the Cure of Cutaneous Eruptions" (1760), and "Gale's Spa Elixir" (1782). The Ann Pike Ointment is probably one of the most glaring of frauds. It is a mixture of pomatum, lard, deer suet, calomel, Jesuits' bark, quicksilver, turpeth mineral, tutty powder, flowers of brimstone and "wood sut." The patent alleges that it is a "Grand Antidote for the Itch and All

[^3]
[^0]:    * Joint Session, Scientific Section and Section on Practical Pharmacy and Dispensing, A. Рн. A., Washington meeting, 1934.
    ${ }^{1}$ In collaboration with A. G. DuMez, Professor of Pharmacy, School of Pharmacy, University of Maryland.

[^1]:    *Standard Deviation expressed in grams.
    ${ }^{1}$ Percentage Deviation based on average weight.

[^2]:    * Standard Deviation expressed in grams.
    ${ }^{1}$ Percentage Deviation based on average weight.

[^3]:    * Section on Historical Pharmacy, A. Ph. A., Washington meeting, 1934.
    ${ }^{1}$ Former Chief of Drug Division, Bureau of Chemistry, United States Department of Agriculture.

