

**Supplementary Table 1.** The need for an open-minded reader

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All humans are biased

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- There is no doubt that cognitive scientists agree that human beings are biased and go through life verifying their own beliefs.<sup>10</sup> Thus, if the reader is a psychiatrist who thinks of himself/herself as a sophisticated scientist and that the leading psychiatrists in the US are sophisticated scientists, there is not much point in continuing to read an article which defends ideas ignored by US clozapine references.<sup>6-9</sup>
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Psychiatrists are practitioners and not scientists

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- In a prior article in this journal,<sup>11</sup> the author has defended the idea that psychiatrists are practitioners and not scientists and there are major problems with current psychiatric nosology. If, in effect, psychiatrists are fundamentally practitioners, their role as practitioners should be to use clozapine safely and effectively and any clozapine science should target that main goal. Unfortunately, this is not true; psychiatric science focuses on the unrealized fantasy that psychiatry is a clinical neuroscience, similar to neurology.<sup>12</sup>
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Limitations of psychopharmacology as science

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- The humbling reality is that the discovery of most important psychiatric drugs, including clozapine, is better described as serendipitous.<sup>13</sup>
  - Problems occur when psychiatrists tried to focus on advances in scientific knowledge in general or focus on brain mechanisms, which pharmacologists call pharmacodynamic mechanisms.<sup>14</sup> Unfortunately, all the arguments proposed in this article are based on the development of the science of pharmacokinetics.
  - Pharmacokinetics, usually defined in a simplified way as what the body does to the drug, is almost completely ignored by psychiatric textbooks and most leading psychiatric journals.<sup>14</sup>
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Limitations of clozapine science in typical psychiatric articles

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- Thus, scientific discussion in psychiatry tends to focus on brain mechanisms. A recent comprehensive review of clozapine cellular mechanisms for the last 50 years has 755 references<sup>15</sup> but ignores most of the pharmacokinetic advances proposed in this article.
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